Case 22617

Heterocyclic Antiviral Compounds

This invention relates to octahydro-pyrrolo[3,4-c]pyrrole derivatives useful in the treatment of a variety of disorders, including those in which the modulation of CCR5 receptors is implicated. More particularly, the present invention relates to 3-(hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propylamine and [3-(hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-propyl]-phenyl-amine compounds and related derivatives, to compositions containing, to uses of such derivatives and to processes for preparing said compoundsz. Disorders that may be treated or prevented by the present derivatives include HIV and genetically related retroviral infections (and the resulting acquired immune deficiency syndrome, AIDS), diseases of the immune system and inflammatory diseases.

A-M. Vandamme *et al.* (*Antiviral Chemistry & Chemotherapy*, 1998 9:187-203) disclose current HAART clinical treatments of HIV-1 infections in man including at least triple drug combinations. Highly active anti-retroviral therapy (HAART) has traditionally consisted of combination therapy with nucleoside reverse transcriptase inhibitors (NRTI), non-nucleoside reverse transcriptase inhibitors (NNRTI) and protease inhibitors (PI). These compounds inhibit biochemical processes required for viral replication. In compliant drug-naive patients, HAART is effective in reducing mortality and progression of HIV-1 to AIDS. While HAART has dramatically altered the prognosis for HIV infected persons, there remain many drawbacks to the current therapy including highly complex dosing regimes and side effects which can be very severe (A. Carr and D. A. Cooper, *Lancet* 2000 356(9239):1423-1430). Moreover, these multidrug therapies do not eliminate HIV-1 and long-term treatment usually results in multidrug resistance, thus limiting their utility in long term therapy. Development of new drug therapies to provide better HIV-1 treatment remains a priority.

Compounds of the present invention modulate the activity of the chemokine CCR5 receptors. The chemokines are a large family of pro-inflammatory peptides that exert their pharmacological effect through G-protein-coupled receptors. The name "chemokine", is a contraction of "chemotactic cytokines". The chemokines are a family of leukocyte chemotactic proteins capable of attracting leukocytes to various tissues, which is an essential response to inflammation and infection. Human chemokines include

approximately 50 small proteins of 50-120 amino acids that are structurally homologous. (M. Baggiolini *et al.*, *Annu. Rev. Immunol.* 1997 15:675-705)

Modulators of the CCR5 receptor may be useful in the treatment of various inflammatory diseases and conditions, and in the treatment of infection by HIV-1 and genetically related retroviruses. As leukocyte chemotactic factors, chemokines play an indispensable role in the attraction of leukocytes to various tissues of the body, a process which is essential for both inflammation and the body's response to infection. Because chemokines and their receptors are central to the pathophysiology of inflammatory and infectious diseases, agents which are active in modulating, preferably antagonizing, the activity of chemokines and their receptors, are useful in the therapeutic treatment of such inflammatory and infectious diseases. The chemokine receptor CCR5 is of particular importance in the context of treating inflammatory and infectious diseases. CCR5 is a receptor for chemokines, especially for the macrophage inflammatory proteins (MIP) designated MIP-1a and MIP-1b, and for a protein which is regulated upon activation and is normal T-cell expressed and secreted (RANTES).

HIV-1 infects cells of the monocyte-macrophage lineage and helper T-cell lymphocytes by exploiting a high affinity interaction of the viral enveloped glycoprotein (Env) with the CD-4 antigen. The CD-4 antigen, however appeared to be a necessary, but not sufficient requirement for cell entry and at least one other surface protein was required to infect the cells (E. A. Berger *et al.*, *Ann. Rev. Immunol.* 1999 17:657-700). Two chemokine receptors, either the CCR5 or the CXCR4 receptor were subsequently found to be co-receptors along with CD4 which are required for infection of cells by the human immunodeficiency virus (HIV). The central role of CCR5 in the pathogenesis of HIV was inferred by epidemiological identification of powerful disease modifying effects of the naturally occurring null allele CCR5 Δ32. The Δ32 mutation has a 32-basepair deletion in the CCR5 gene resulting in a truncated protein designated Δ32. Relative to the general population, Δ32/Δ32 homozygotes are significantly common in exposed/uninfected individuals suggesting the role of CCR5 in HIV cell entry (R. Liu *et al.*, *Cell* 1996 86(3):367-377; M. Samson *et al.*, *Nature* 1996 382(6593):722-725).

The HIV-1 envelope protein is comprised of two subunits: gp120, the surface subunit and gp41, the transmembrane subunit. The two subunits are non-covalently associated and form homotrimers which compose the HIV envelope. Each gp41 subunit contains two helical heptad repeat regions, HR1 and HR2 and a hydrophobic fusion region on the C-terminus.

The CD-4 binding site on the gp120 of HIV appears to interact with the CD4 molecule on the cell surface that induces a conformation change in gp120 which creates

or exposes a cryptic CCR5 (or CXCR-4) binding site, and undergoes conformational changes which permits binding of gp120 to the CCR5 and/or CXCR-4 cell-surface receptor. The bivalent interaction brings the virus membrane into close proximity with the target cell membrane and the hydrophobic fusion region can insert into the target cell membrane. A conformation change in gp41 allows the contact between the outer leaflet of the target cell membrane and the viral membrane which produces a fusion pore whereby the virus RNA is injected into the cytoplasm. Accordingly, an agent which could inhibit binding between gp120 and chemokine receptors should prevent or moderate infection in healthy individuals and slow or halt viral progression in infected patients. (B. Tomkowicz and R. G. Collman, *Expert Opin. Ther. Targets* 2004 8(2):65-78; J. P. Moore and R. W. Doms, *Proc. Nat. Acad. Sci. USA*, 2003 100(19):10598-10602)

Viral fusion and cell entry is a complex multi-step process and each step affords the potential for therapeutic intervention. These steps include (*i*) CD-40-gp120 interactions, (*ii*) CCR5 and/or CXCR-4 interactions and (*iii*) gp41 mediated membrane fusion. Each of these steps affords an opportunity for therapeutic intervention in preventing or slowing HIV infection

RANTES, a natural ligand for the CCR5 receptor, and an analog chemically modified on the N-terminus, aminooxypentane RANTES, were found to block HIV entry into the cells. (G. Simmons *et al.*, *Science* 1997 276:276-279). Other compounds have been demonstrated to inhibit the replication of HIV, including soluble CD4 protein and synthetic derivatives (Smith, *et al.*, *Science* 1987 238:1704-1707), dextran sulfate, the dyes Direct Yellow 50, Evans Blue, and certain azo dyes (U.S. Pat. No. 5,468,469). Some of these antiviral agents have been shown to act by blocking the binding of gp120, the coat protein of HIV, to its target, the CD4 glycoprotein of the cell.

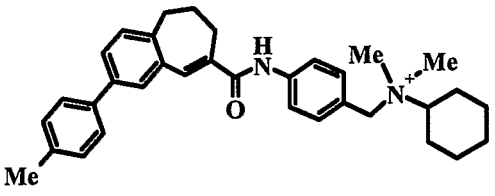
T20 (Entuvirtide) is a 36 amino acid peptide corresponding to residues 643-678 in the HR2 domain of gp41. T-20 binds a transient intermediate formed interaction after interaction of gp120 and the target cell which inhibits viral fusion thereby suppressing viral replication. (J. M. Kilby *et al.*, *New Eng. J. med.* 1998 4(11):1302-1307). T-20 has been approved for clinical use.

In addition to the potential for CCR5 modulators in the management of HIV infections, the CCR5 receptor is an important regulator of immune function and compounds of the present invention may prove valuable in the treatment of disorders of the immune system. Treatment of solid organ transplant rejection, graft v. host disease, arthritis, rheumatoid arthritis, inflammatory bowel disease, atopic dermatitis, psoriasis, asthma, allergies or multiple sclerosis by administering to a human in need of such

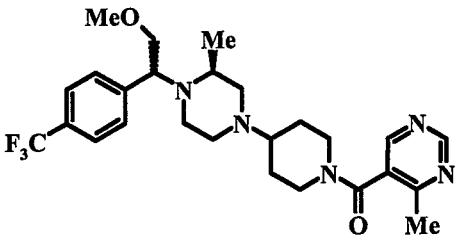
treatment an effective amount of a CCR5 antagonist compound of the present invention is also possible.

The pharmacokinetic challenges associated with large molecules, proteins and peptides resulted in the establishment of programs to identify low molecular weight antagonists of CCR5. The efforts to identify chemokine modulators have been reviewed (W. Kazmierski *et al. Biorg Med. Chem.* 2003 11:2663-76; L. Agrawal and G. Alkhatib, *Expert Opin. Ther. Targets* 2001 5(3):303-326; *Chemokine CCR5 antagonists incorporating 4-aminopiperidine scaffold, Expert Opin. Ther. Patents* 2003 13(9):1469-1473; M. A. Cascieri and M. S. Springer, *Curr. Opin. Chem. Biol.* 2000 4:420-426, and references cited therein)

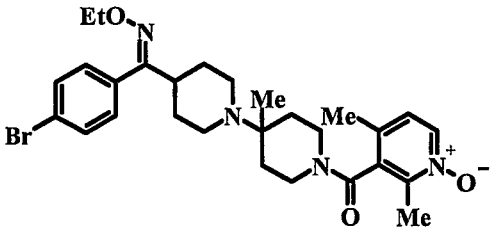
Takeda's program was the first to lead to fruition with the identification of TAK-779 (M. Shiraishi *et al., J. Med. Chem.* 2000 43(10):2049-2063). Schering has advanced Sch-351125 into Phase I/II clinical studies and reported the advance of a more potent follow-up compound, Sch-417690 into Phase I studies. (S. W. McCrombie *et al.*, WO00066559; B. M. Baroudy *et al.* WO00066558; A. Palani *et al., J. Med Chem.* 2001 44(21):3339-3342; J. R. Tagat *et al., J. Med Chem.* 2001 44(21):3343-3346; J. A. Esté, *Cur. Opin. Invest. Drugs* 2002 3(3):379-383).



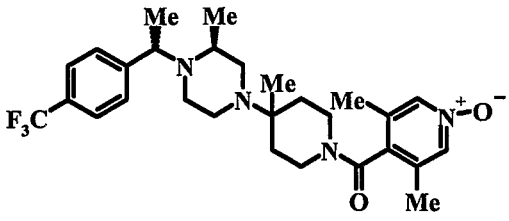
TAK-779



Sch-417690



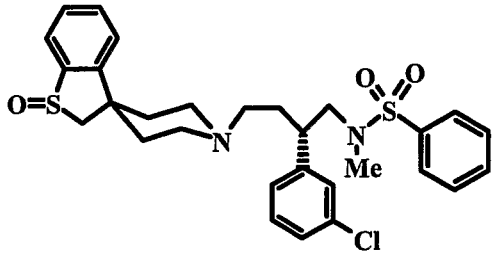
Sch-351125



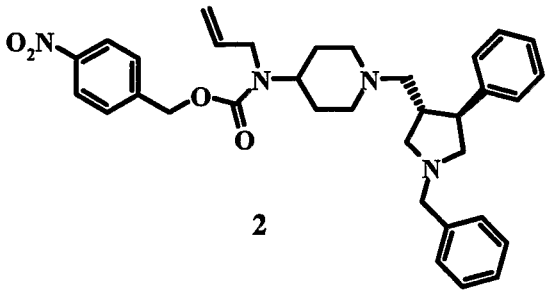
Sch-350634

Merck has disclosed the preparation of (2S)-2-(3-chlorophenyl)-1-N-(methyl)-N-(phenylsulfonyl)amino]-4-[spiro(2,3-dihydrobenzothiophene-3,4'-piperidin-1'-yl)butane S-oxide (1) and related derivatives, trisubstituted pyrrolidines 2 and substituted piperidines 3 with good affinity for the CCR5 receptor and potent-HIV activity. (P. E. Finke *et al., Bioorg. Med. Chem. Lett.*, 2001 11:265-270; P. E. Finke *et al., Bioorg. Med. Chem. Lett.*, 2001 11:2469-2475; P. E. Finke *et al., Bioorg. Med. Chem. Lett.*, 2001 11:2475-

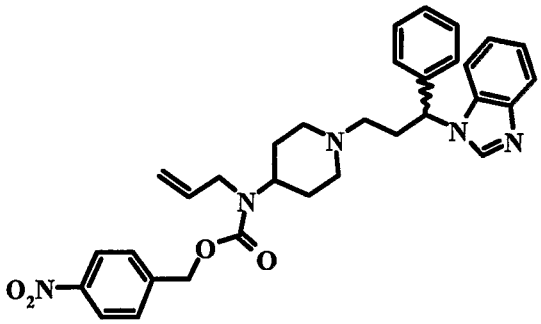
2479; J. J. Hale *et al.*, *Bioorg. Med. Chem. Lett.*, 2001 11:2741-22745; D. Kim *et al.*, *Bioorg. Med. Chem. Lett.*, 2001 11:3099-3102)



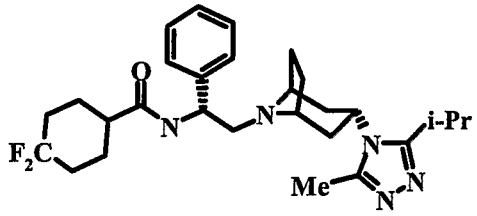
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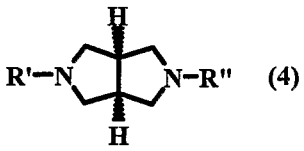


3



UK-427857

WO0039125 (D. R. Armour *et al.*) and WO0190106 (M. Perros *et al.*) disclose heterocyclic compounds that are potent and selective CCR5 antagonists. UK-427857 has advanced to clinical trials and show activity against HIV-1 isolates and laboratory strains (M. J. Macartney *et al.*, *43rd Intersci. Conf. Antimicrob. Agents Chemother.* (September 14-17, 2003, Abstract H-875).



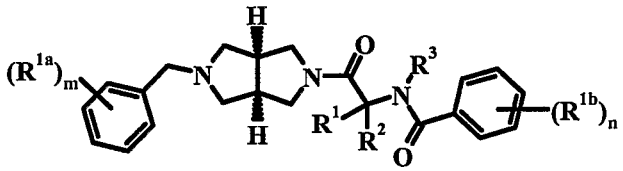
(4)

Due to their rigidity and the ease of functionalization, bicyclic and tricyclic amines have proven to be useful scaffolds in drug design. Compounds based upon the 3,7-diazabiyclo[3.3.0]octane (4: R' = R" = H) are known for use in a variety of medical applications, including *inter alia* as migrane (as described in WO 98/06725 and WO97/11945); antibiotics (as described in WO 97/10223 and WO 96/25691, neuroleptics (as described in WO 95/15327 and WO95/13279); serotonin reuptake inhibitors (as described in WO 96/07656); thrombin inhibitors (as described in *Helv. Chim. Acta* 2000 83:855, *Chem. & Biol.* 1997 4:287 and *Angew. Chem. Int. Ed. Eng.*1995 34:1739); and, anxiolytic agents (*J. Med. Chem.* 1989 32:1024). In addition, compounds based upon 3,7-diazabiyclo[3.3.0]octane have been used in the treatment of gastrointestinal disorders (DE 39 30 266 A1) disorders of the glutaminergic system. WO 00/55143 discloses

diamine compounds, including 3,7-diazabicyclo[3.3.0]octane compounds linked to an oxazolone which compounds are α-1 adenoreceptor modulators. EP 1 122 257 (F. Ito *et al.*) disclose benzimidazole compounds linked to a variety bicyclic diamine compounds including diazabicyclooctanes which compounds are ORL-1 receptor agonists. The ORL-1 receptor is an opioid receptor subtype.

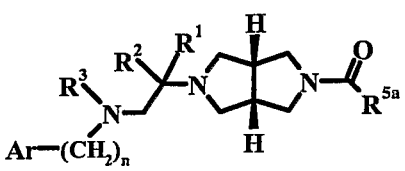
WO 96/07656 (J. M. Schaus and R.D. Titus) disclosed aralkyl substituted 3,7-diazabicyclo[3.3.0]octane compounds with selective reuptake inhibitory activity. WO 97/11945 (A. Madin) disclose 3-substituted 3,7-diazabicyclo[3.3.0]octane compounds, which may be further substituted at the 7-position, with selective 5-HT_{1D} activity. WO 01/44243 (D. Peters *et al.*) disclose novel heteroaryl 3-substituted-diazabicycloalkane compounds, which may be optionally substituted at the 7-position, which are muscarinic and nicotinic receptor modulators which are useful in the treating diseases associated with degeneration of the cholinergic system. The preferred embodiments include 3,7-diazabicyclo[3.3.0]octane compounds 4 wherein R' is a heteroaryl group and R" is hydrogen, alkyl, aryl, aralkyl or a fluorescent group.

WO 02/07523 A1 (R. Colon-Cruz *et al.*) discloses 3,7-diazabicyclo[3.3.0]octane compounds which are anatagonists of the chemokine CCR2 and CCR3 receptors. MCP-1 is believed the natural ligand for the CCR2 receptor and interaction of the ligand with the receptor increases histamine release, calcium influx, cAMP activation, increases integrin expression and acts as a chemotactic factor for monocytes and macrophages. Compounds disclosed in the invention were claimed useful for the treatment of diseases of monocyte, lymphocyte and leukocyte accumulation and more specifically atherosclerosis, restenosis, gingivitis, psoriasis, rheumatoid arthritis, glomerulonephritis, Crohn's disease, encephalomyelitis and transplant rejection. The preferred embodiments include compounds with the generic formula 5.



(5)

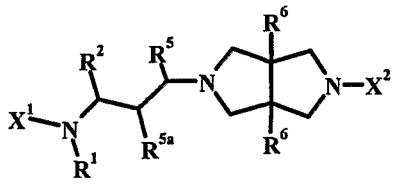
WO 02/060902 A1 (M. Björsne *et al.*) disclose 3,7-diazabicyclo[3.3.0]octane compounds useful in the treatment of cardiac arrhythmias. The preferred embodiments include compounds with the generic formula 6.



(6)

The present invention relates to a compounds according to formula I, methods for treating diseases alleviated by administration of a compound according to formula I that is a CCR5 antagonists and pharmaceutical compositions for treating diseases containing a compound according to formula I that is a CCR5 antagonist admixed with at least one carrier, diluent or excipient.

One object of the present invention is (i) a compound of formula (I)



(I)

wherein:

X^1 is selected from the group consisting $-C(=O)R^3$, $-S(=O)_2R^3$ and $-C(=O)OR^3$;

X^2 is selected from the group consisting $-C(=O)R^4$, $-S(=O)_2R^4$, benzyl or $-CH_2$(pyridin-3-yl) said benzyl optionally substituted with halogen or C_{1-3} alkyl;

one of R^1 and R^2 is:

(i) phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of halogen, C_{1-6} alkyl and C_{1-6} alkoxy;

(ii)heteroaryl selected from the group consisting of pyridinyl, pyridinyl-N-oxide, pyrimidinyl and thiazolyl said heteroaryl optionally substituted with one to three substituents independently selected from the group consisting of halogen, C_{1-6} alkyl, C_{1-6} haloalkyl and C_{1-6} alkoxy; and,

the other of R^1 and R^2 is hydrogen;

R^3 is selected from the group consisting of:

(i) phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of:

(a) C_{1-10} alkyl,

(b) C_{1-10} heteroalkyl,

(c) C_{1-6} haloalkyl,

(d) C_{1-6} alkoxy,

(e) C_{1-6} thioalkyl,

(f) amino,

(g) C_{1-6} alkyl amino,

(h) C_{1-6} dialkylamino,

(g) C_{1-6} acylamino,

(i) carbamoyl, N-C_{1-6} alkylcarbamoyl or N,N-C_{1-6} dialkylcarbamoyl,

(j) ureido,

(k) nitro,

(l) cyano,

(m) halogen,

(n) C_{1-6} alkylsulfonyl,

(o) sulfamoyl, N-C_{1-6} alkylsulfamoyl or N, N-C_{1-6} dialkylsulfamoyl,

(p) C_{1-6} alkylsulfonamido or optionally substituted phenylsulfonamido,

(q) optionally substituted phenoxy,

(r) optionally substituted heteroaryloxy, and,

(s) $-Y(CH_2)_nR^{11}$ wherein R^{11} is selected from the group consisting of cyano, $-CO_2R^{12}$, $-CONR^{12}R^{13}$, $-SO_2N^{12}R^{13}$, $-NHSO_2R^{12}$ and $-NHSO_2NR^{12}R^{13}$,

(t) CO_2R^{12},

(u) C_{1-6} acyloxy,

(v) C_{1-6} alkylcarbonyl, and,

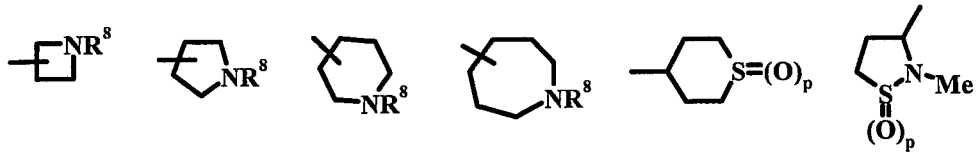
(w) C_{1-6} haloalkoxy;

(ii) phenyl C_{1-6} alkyl wherein phenyl as described in (i) above;

(iii) heteroaryl selected from the group consisting of pyridinyl, pyridinyl-N-oxide, pyrimidinyl, pyrazinyl, pyridazinyl, isoxazolyl, isothiazolyl, pyrrolyl, furyl, thiazolyl, oxazolyl, pyrazolyl, oxadiazolyl, thiadiazolyl, imidazolyl, triazolyl, tetrazolyl, indolyl, quinolinyl, isoquinolinyl, 2,4-dimethyl-6-oxo-6H-pyranyl and thienyl said heteroaryl optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, phenyl, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, C_{1-6} alkoxycarbonyl, carbamoyl, acetyl, nitro, cyano, amino, C_{1-3} alkylamino, C_{1-3} dialkylamino, N-morpholino, sulfamoyl and halogen,

(iv) heteroaryl C_{1-6} alkyl wherein heteroaryl is as described above,

(v) heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is selected from the group consisting of:

(i) C_{1-6} alkyl optionally substituted with one or two groups independently selected from the group consisting of C_{1-3} alkoxy, C_{1-3} acyloxy, hydroxyl, phenyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino and, aminocarbonylpyridyl wherein said aminocarbonylpyridyl can optionally be the N-oxide and aminobenzoyl wherein said phenyl or said benzoyl radical is optionally substituted with one to three groups independently selected from the group consisting of amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, hydroxyl, halogen and C_{1-3} alkoxy,

(ii) phenyl optionally substituted with one or two groups independently selected from the group consisting of sulfamoyl, acetylamino, halogen, C_{1-6} alkyl, C_{1-6} haloalkyl, and C_{1-6} alkoxy,

(iii) NH_2, C_{1-6} alkylamino, C_{1-6} dialkylamino,

(iv) $C(=O)NH_2$, $C(=O)OC_{1-6}$ alkyl, or $C(=O)OH$;

(v) 1H indole-3-carbonyl,

(vi) furyl, pyridinyl, or pyridinyl N-oxide,

(vii) N-acetylpiperidin-4-yl,

(viii) furfuryl,

(ix) C_{3-8} cycloalkyl,

(x) C_{1-6} haloalkyl,

(xi) phenyl C_{1-3} alkyl, and,

(xii) C_{1-6} alkoxy,

R^{10} is selected from the group consisting of:

(i) C_{1-6} alkyl,

(ii) C_{1-3} haloalkyl,

(iii) C_{3-8} cycloalkyl, and

(iv) NH_2, C_{1-3} alkylamino, C_{1-3} dialkylamino;

(vi) heterocycle C_{1-6} alkyl wherein heterocycle is as defined above;

(vii) hydrogen;

(viii) C_{1-10} alkyl optionally independently substituted in each occurrence with one to three substituents independently selected at each occurrence from the group consisting of hydroxy, halogen, C_{1-6} alkoxy, C_{1-6} thioalkyl, C_{1-6} acyloxy, C_{1-6} alkoxycarbonyl, C_{1-6} acylamino, $NR^{14a}R^{14b}$, cyano, C_{1-6} alkylsulfonyl, phenyl, N-methyl-methyl-sulfonamido, N-piperidinyl, N-pyrrolidinyl, imidazolyl, 2-aza-bicyclo[2.2.1]hept-2-yl, carbamoyl, C_{1-6} alkoxycarbonyl, and optionally substituted phenyl as defined in R^3 (*i*) above;

(ix) C_{3-7} cycloalkyl or [3.1.0]bicyclohexyl, 4-oxo-cyclohexyl or 3-oxo-cyclobutyl said cycloalkyl optionally substituted with 1 to 4 fluorine, cyano, hydroxyl, C_{1-3} alkyl or phenyl;

(x) C_{1-3} alkyl-C_{3-7} cycloalkyl said cycloalkyl optionally substituted with 1 to 4 fluorine, C_{1-3} alkyl or phenyl;

(xi) $NR^{14a}R^{14b}$; and,

(xii) $C(=O)OR^{14c}$;

R^4 is selected from the group consisting of:

(i) aryl selected from the group consisting of:

(a) phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of a phenyl substituent (i)(a) to (i)(w) as described in R^3, C_{1-6} haloalkoxy, pyridinyl, 2-oxo-pyrrolidin-1-yl, 2,5-dimethyl-pyrrolidin-1-yl, 3-methyl-5-oxo-4,5-dihydro-pyrazol-1-yl and $NR^{14a}R^{14b}$ wherein R^{14a} and R^{14b} are taken together along with the nitrogen to which they are attached are $(CH_2)_r$ or $[(CH_2)_2X^3(CH_2)_2]$, $-NMe(CH_2)_2OH$;

(b) 2,3-dihydrobenzofuran-4-yl,

(c) benzofuran-4-yl,

(d) 2,3-dihydrobenzofuran-7-yl,

(e) quinolin-8-yl,

(f) 1,2,3,4-tetrahydro-isoquinolin-5-yl,

(g) N-acetyl-1,2,3,4-tetrahydro-isoquinolin-5-yl,

(h) N-Boc-1,2,3,4-tetrahydro-isoquinolin-5-yl,

(i) 1,2,3,4-tetrahydro-isoquinolin-8-yl,

(j) N-Boc-1,2,3,4-tetrahydro-isoquinolin-8-yl,

(k) isoquinolin-7-yl,

(l) 1H-indol-5-yl,

(m) 2-acetyl-1,2,3,4-tetrahydro-isoquinolin-5-yl, and

(n) 6-fluoro-benzo[1,3]dioxin-8-yl, and,

(o) quinolin-6-yl;

(ii) phenyl C_{1-6} alkyl wherein phenyl as described above;

(iii)heteroaryl selected from the group consisting of pyridinyl, 2-azetidin-1-yl, pyridinyl N-oxide, pyrimidinyl, pyrazinyl, pyridazinyl, isoxazolyl, isothiazolyl, pyrrolyl, furyl, thiazolyl, oxazolyl, pyrazolyl, oxadiazolyl, thiadiazolyl, imidazolyl, triazolyl, tetrazolyl, indolyl, quinolyl, isoquinolyl, quinoxalin-2-yl, indazole, benzofuranyl, 4,5,6,7-tetrahydro-benzofuranyl, 2,4-dimethyl-6-oxo-6H-pyranyl, benzo[b]thiophenyl, 4,5,6,7-tetrahydroindazole, 1,4,5,6-tetrahydrocyclopentylpyrazolyl, imidazo[2,1-b]thiazolyl, 6-fluoro-4H-benzo[1,3]dioxin-8-yl, cinnolin-4-yl, and thienyl said heteroaryl optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of substituents (i)(a) to (i)(w) as described for R^3 above, benzyl, pyridinyl, 2-methyl-thiazolyl, acetyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, aminobenzyl, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, C_{1-6} alkylsulfonyl, C_{1-6} alkoxycarbonyl, carboxyl, carbamoyl, nitro, cyano, sulfamoyl, $-OCH_2CO_2R^{14c}$, $-SCH_2CO_2R^{14c}$ and halogen;

(iv) heteroaryl C_{1-6} alkyl wherein heteroaryl is as described above;

(v) heterocycle selected from the group consisting of IIa-f as described in (v) of R^3 above, furyl, oxetanyl, tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, piperazinyl, imidazolinyl, N-benzyl-morpholine, N-methyl-1,2,3,4-tetrahydroquinolin-2-yl, 4,5-dihydro-pyrazolyl, and 1,2,3,4-tetrahydroisoquinolinyl said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, acetyl, hydroxy, C_{1-6} alkylcarbonyl, C_{1-6} alkoxycarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo);

(vi) heterocycle C_{1-6} alkyl wherein heterocycle is as defined above;

(vii) hydrogen;

(viii)C_{1-10} alkyl substituted with one to three substituents independently selected at each occurrence from the group consisting of hydroxy, halogen, C_{1-6} alkoxy, C_{1-6} thioalkyl, acyloxy, C_{1-6} acylamino, $NR^{14a}R^{14b}$, cyano, C_{1-6} alkylsulfonyl, N-methyl-methyl-sulfonamido, phenyl, N-piperidinyl, N-pyrrolidinyl, imidazolyl, carbamoyl and C_{1-6} alkoxycarbonyl,

(ix) C_{3-7} cycloalkyl optionally substituted with 1 to 4 fluorine, C_{1-3} alkyl or phenyl,

(x) C_{1-3} alkyl-C_{3-7} cycloalkyl optionally substituted with 1 to 4 fluorine, C_{1-3} alkyl or phenyl; and,

(xi) $NR^{14a}R^{14b}$;

R^5 is hydrogen, C_{1-6} alkyl, C_{1-6} haloalkyl, C_{1-6} hydroxyalkyl, methoxymethyl, -CO_2R^{12} or $CONR^{12}R^{13}$;

R^{5a} is hydrogen or C_{1-6} alkyl;

R^6 is independently selected from the group consisting of hydrogen, C_{1-10} alkyl, hydroxy, C_{1-6} alkoxy, C_{1-3} hydroxyalkyl and C_{1-3} alkoxy-C_{1-3} alkyl;

R^7 is hydrogen or Boc;

R^{12} and R^{13} (i) are independently hydrogen, C_{1-6} alkyl or C_{1-6} heteroalkyl, or (*ii*) R^{12} and R^{13}, when both attached to a nitrogen atom, together can be C_{2-5} alkylene;

R^{14a} and R^{14b} (i) taken independently are R^3, or (ii) taken together along with the nitrogen to which they are attached are $(CH_2)_r$ or $[(CH_2)_2X^3(CH_2)_2]$;

R^{14c} is hydrogen or C_{1-6} alkyl;

R^{15} is hydrogen, methyl, *iso*-propyl, *iso*-butyl, *sec*-butyl, -CH_2OH, -$CH(OH)CH_3$, -CH_2SH, -CH_2CH_2SMe, -$(CH_2)_sCOR^{16}$ wherein R^{16} is -OH or -NH_2 and s is 1 or 2, -$(CH_2)_t$-NH_2 where t is 3 or 4, -$(CH_2)_3$-$NHC(=NH)NH_2$, -$CH_2C_6H_5$, -CH_2-*p*-C_6H_4-OH, (3-indolinyl)methylene or (4-imidazolyl)methylene;

X^3 is -O-, -$S(O)_p$-, NR^{14c};

Y is a direct bond, -O-, -S- or -NR^{12}-;

n is an integer from 1 to 6;

p is an integer from 0 to 2;

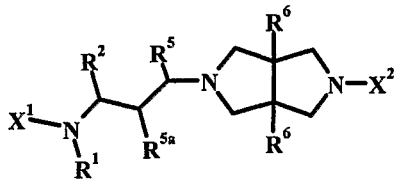
r is an integer from 3 to 6; and,

pharmaceutically acceptable salts, hydrates, solvates and stereoisomers thereof.

Compounds and compositions of the present invention are useful for treating diseases mediated by human immunodeficiency virus in humans. Compounds and

compositions of the present invention also may be used for treatment of disorders alleviated by a CCR5 antagonist including respiratory disorders, including adult respiratory distress syndrome (ARDS), bronchitis, chronic bronchitis, chronic obstructive pulmonary disease, cystic fibrosis, asthma, emphysema, rhinitis and chronic sinusitis. Conditions triggered, affected or are in any other way correlated with T-cell trafficking in different organs may be treated with compounds of the invention. Compounds of the present invention may be useful for the treatment of such conditions and in particular, but not limited to the following for which a correlation with CCR5 or CCR5 chemokines has been established: inflammatory bowel disease, including Crohn's disease and ulcerative colitis, multiple sclerosis, rheumatoid arthritis, graft rejection, in particular but not limited to kidney and lung allografts, endometriosis, type I diabetes, renal diseases, chronic pancreatitis, inflammatory lung conditions or chronic heart failure. For recent reviews of possible applications of chemokines and chemokine receptor blockers see: Cascieri, M.A., and Springer, M.S., *The chemokine/chemokine receptor family: potential and progress for therapeutic intervention, Curr. Opin. Chem. Biol.* 2000 4(4):420-7; A. E. I. Proudfoot *The Strategy of Blocking the Chemokine System to Combat Disease, Immunol. Rev.* 2000 177:246-256.

Further objects of the present invention are (ii) A compound according to formula I



(I)

wherein:

one of R^1 and R^2 is:

phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of halogen and C_{1-6} alkyl;

heteroaryl selected from the group consisting of pyridinyl, pyrimidinyl and thiazolyl said heteroaryl optionally substituted with one to three substituents independently selected from the group comprising halogen, C_{1-6} alkyl and C_{1-6} haloalkyl; and,

the other of R^1 and R^2 is hydrogen;

R^5 is hydrogen, C_{1-6} alkyl methoxymethyl or C_{1-3} hydroxyalkyl;

R^{5a} is hydrogen, hydroxyl or C_{1-6} alkyl;

R^6 is hydrogen, C_{1-6} alkyl or C_{1-3} hydroxyalkyl;

X^1 is selected from the group consisting of $-C(=O)R^3$, $-S(=O)_2R^3$ and $-C(=O)OR^3$;

X^2 is selected from the group consisting of $-C(=O)R^4$, $-S(=O)_2R^4$ or benzyl, said benzyl optionally substituted with halogen or C_{1-3} alkyl;

R^3 is selected from the group consisting of:

hydrogen, C_{1-6} alkyl, $-CH_2NH2$, $-CH_2CH_2NH_2$, $-C(CH_3)NH_2$, $-CH_2CH_2N(ethyl)_2$, $-N(CH_3)_2$, $-N(CH_2CH_3)_2$, -N(iso-propyl)$_2$, $-NCH_3$-benzyl, $-NHCH_2COOCH_2CH_3$, $-CH_2NHCOCH_3$, $CH_2CH_2CONH_2$, $-C(NH_2)CH_2C(CH_3)_2$, $-CH(NH_2)CH(CH_3)_2$, CH_2CN,
$-C_{1-6}$ alkyl-OH, $-CH_2CH_2COOCH_3$, $-C(CH_3)_2OCOCH_3$, $-CH_2OCOCH_3$, $-CH_2SO_2CH_3$,
$-CH_2(CH_3)SO_2CH_3$,

heteroalkyl, said heteroalkyl is optionally substituted with $-NHCOCH_3$,

$-(CH_2)_t$-C_{3-8} cycloalkyl, wherein t is 0 to 2, and

said $-C_{3-8}$ cycloalkyl is optionally substituted with one to three substituents selected independently from the group comprising of C_{1-6} alkyl, halogen, hydroxyl, cyano, oxo, phenyl,

$-(CHR^m)_u$-phenyl, wherein R^m is H, hydroxyl or C_{1-6} alkyl, u is 0 to 3, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of:

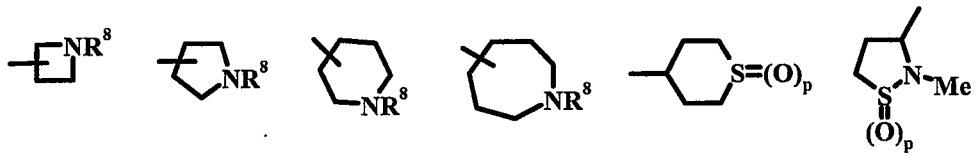
C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, $-SO_2NH_2$,
$-NHCOCH_3$ or $-COOCH_3$,

$-(CH_2)_v$-heteroaryl selected from the group consisting of pyridinyl, pyridinyl-N-oxide, pyrimidinyl, pyrazinyl, pyridazinyl, isoxazolyl, isothiazolyl, pyrrolyl, furyl, thiazolyl, oxazolyl, pyrazolyl, oxadiazolyl, thiadiazolyl, imidazolyl, triazolyl, tetrazolyl, indolyl, quinolinyl, isoquinolinyl, 2,4-dimethyl-6-oxo-6H-pyranyl and thienyl,

wherein v 0 to 3, and

said heteroaryl is optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, phenyl, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, C_{1-6} alkoxycarbonyl, carbamoyl, acetyl, nitro, cyano, amino, C_{1-3} alkylamino, C_{1-3} dialkylamino, N-morpholino, sulfamoyl and halogen,

$-(CHR^n)_w$-heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl,

wherein R^n is hydrogen or C_{1-6} alkyl, w is 0 to 3, and

said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is C_{1-6} alkyl, C_{1-6} haloalkyl, furanyl, C_{1-6} alkoxy, C_{3-5} cycloalkyl, C_{1-6} alkylamino, -COO- C_{1-6} alkyl, -COOH, $-CH_2OCH_2OOH$, $-CH_2CH_2COOH$;

R^{10} is C_{1-6} alkyl, C_{1-6} dialkylamino, C_{3-8} cycloalkyl;

R^7 is hydrogen or Boc;

R^{15} is hydrogen or C_{1-6} alkyl;

$-NH-(CH_2)_x$-phenyl, wherein x is 0 or 1, and

said phenyl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, $-OCF_3$, heteroalkyl, haloalkyl, acetyl, dialkylamin, cyano, -COOH, $COOCH_2CH_3$,

$-NH(CH_2)_y$-heteroaryl, wherein y is 0 or 1, and

said heteroaryl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, haloalkyl,

-NH-heterocycle, wherein said heteroaryl is optionally substituted with 1 to 2 substituents independently selected in each occurrence from the group consisting of $-COCF_3$, $COOCH_2$-phenyl,

$-NH(CH_2)_z-C_{3-8}$ cycloalkyl, wherein z is 0 or 1;

R^4 is selected from the group consisting of:

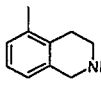
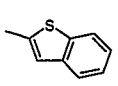
C_{1-6} alkyl, $-CH_2CH_2CONH_2$, $-CH_2NCH_3SO_2CH_3$, $-CH(CH(CH_3)_2)NHCOCH_3$,

$-(CH_2)_eC_{3-8}$ cycloalkyl, wherein e is 0 to 2, and said cycloalkyl is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl,

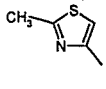
C_{3-8} heterocycle, wherein said heterocyle is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxyl, -CO- C_{1-6} alkyl, oxo, benzyl, , or ,

$-(CHR^f)_f$-phenyl, wherein R^f is hydrogen or –NHCOCH3, f is 0 or 1, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, NO_2, $-CO-C_{1-6}$ alkyl, $-COOCH_3$, $COOC(CH_3)_3$, $-NHSO_2CH_3$, NH_2, $-N(CH_3)_2$, $-N(CH_3)CH_2CH_2OH$, $-NHCOCH_3$,$-OCF_3$ $-OCH_2COOH$, pyridine-4-yl, , , , ;

$(CH_2)_g$-heteroaryl, wherein g is 0 or 1, sais heteroary selected from the group consisting of furanyl, thiophenyl, isoxazolyl, thiazolyl, oxazolyl, pyrroly, pyrazolyl, imidazolyl, indolyl, benzofuranyl, pyranyl, pyridinyl, pyrimidinyl, chinoxalinyl, chinolinyl, , , , , and ,

said heteroaryl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-6} alkyl, oxo, oxido,

heteroalkyl, halogen, haloalkyl, methoxy-phenyl, haloalkyl, -NH_2, -$N(CH_3)_2$, -NHCH2-phenyl, -NHCOCH3, cyano, -SO_2NH_2, -SO_2CH_3, -OCH_2COOCH_3, -OCH_2COOH, -SCH_2COOH, -$COCH_3$, -$COOC(CH_3)_3$, -$COOC(CH_3)_3$, -$COOCH_2CH_3$,- COOH, pyridine-4-yl, , , benzyl, phenyl, said phenyl is optionally substituted with one to three substitutuents consisting of C_{1-6} alkyl , halogen, C_{1-6} alkoxyl, -$COOCH_3$;

-(NR^h)-$(CH_2)_h$-phenyl, wherein R^h is hydrogen or C_{1-6} alkyl, h is 0 or 1, and said phenyl is optionally substituted with one to three substituents selected from C_{1-6} alkyl, and

pharmaceutically acceptable salts, hydrates, solvates and stereoisomers thereof.

(iii). A compound according to (ii), wherein:

R^2 is phenyl optionally substituted with one to four halogen, or pyridinyl;

R^1 is hydrogen;

R^5 is hydrogen, C_{1-6} alkyl or methoxymethyl, C_{1-3} hydroxyalkyl;

R^{5a} is hydrogen hydroxyl or C_{1-6} alkyl;

R^6 is hydrogen, C_{1-6} alkyl or C_{1-3} hydroxyalkyl;

X^1 is selected from the group consisting of -$C(=O)R^3$, -$S(=O)_2R^3$ and -$C(=O)OR^3$;

X^2 is selected from the group consisting of -$C(=O)R^4$, -$S(=O)_2R^4$ or benzyl, said benzyl optionally substituted with halogen or C_{1-3} alkyl;

R^3 is selected from the group consisting of:

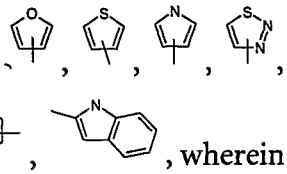
hydrogen, C_{1-6} alkyl, -CH_2NH2, -$CH_2CH_2NH_2$, -$C(CH_3)NH_2$, -$CH_2CH_2N(ethyl)_2$,, -$N(CH_3)_2$, -$N(CH_2CH_3)_2$, -N(iso-propyl)$_2$, -NCH_3-benzyl, -$NHCH_2COOCH_2CH_3$, -NH-phenyl, -$CH_2NHCOCH_3$, $CH_2CH_2CONH_2$, -$C(NH_2)CH_2C(CH_3)_2$, -$CH(NH_2)CH(CH_3)_2$,-CH2CN, -C_{1-6} alkyl-OH, -$CH_2CH_2COOCH_3$, -$C(CH_3)_2OCOCH_3$, -CH_2OCOCH_3, -CH2SO2CH3, -CH2(CH3)SO2CH3, heteroalkyl,

-$(CH_2)_t$-C_{3-8} cycloalkyl selected from cyclopropanyl, cyclobutanyl, cyclopentanyl, cyclohexanyl, cycloheptanyl and , wherein t is 0 to 2, and

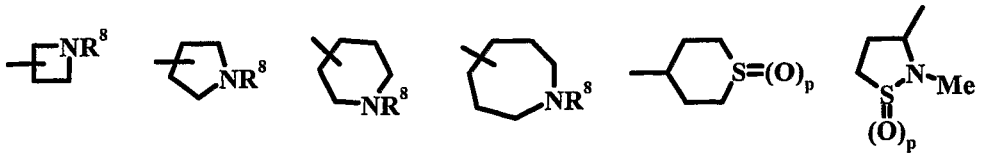
said -C_{3-8} cycloalkyl is optionally substituted with one to three substituents selected independently from the group comprising of C_{1-6} alkyl, halogen, hydroxyl, cyano, oxo, phenyl,

-$(CHR^m)_u$-phenyl, wherein R^m is H, hydroxyl or C_{1-6} alkyl, u is 0 to 2, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, -SO_2NH_2, -$NHCOCH_3$ or -$COOCH_3$,

-$(CH_2)_v$-heteroaryl selected from the group consisting of , wherein v is 0 to 3, and said heteroaryl is optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, halogen, phenyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, -NH_2, SO2NH2, -COCH3, oxo, oxido, ,

-$(CHR^n)_w$-heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl,

wherein R^n is hydrogen or C_{1-6} alkyl, w is 0 to 3, and

said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is C_{1-6} alkyl, C_{1-6} haloalkyl, furanyl, C_{1-6} alkoxy, C_{3-5} cycloalkyl, C_{1-6} alkylamino, -COO- C_{1-6} alkyl, -COOH, $-CH_2OCH_2OOH$, $-CH_2CH_2COOH$;

R^{10} is C_{1-6} alkyl, C_{1-6} dialkylamino, C_{3-8} cycloalkyl;

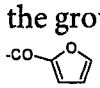
R^7 is hydrogen or Boc;

R^{15} is hydrogen or C_{1-6} alkyl;

R^4 is selected from the group consisting of:

C_{1-6} alkyl, $-CH_2CH_2CONH_2$, $-CH_2NCH_3SO_2CH_3$, $-CH(CH(CH_3)_2)NHCOCH_3$,

$-(CH_2)_eC_{3-8}$ cycloalkyl selected from cyclopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, wherein e is 0 to 2, and said cycloalkyl is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl,

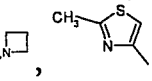
C_{3-8} heterocycle, wherein said heterocyle is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxyl, -CO- C_{1-6} alkyl, oxo, benzyl, , or ,

$-(CHR^f)_f$-phenyl, wherein R^f is hydrogen or –NHCOCH3, f is 0 or 1, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, NO_2, -CO-C_{1-6} alkyl, $-COOCH_3$, $COOC(CH_3)_3$, $-NHSO_2CH_3$, NH_2, $-N(CH_3)_2$, $-N(CH_3)CH_2CH_2OH$, $-NHCOCH_3$,$-OCF_3$ $-OCH_2COOH$, pyridine-4-yl, ;

$(CH_2)_g$-heteroaryl, wherein g is 0 or 1, sais heteroary selected from the group consisting of furanyl, thiophenyl, isoxazolyl, thiazolyl, oxazolyl, pyrroly, pyrazolyl, imidazolyl, indolyl, benzofuranyl, pyranyl, pyridinyl, pyrimidinyl, chinoxalinyl, chinolinyl, , , and ,

said heteroaryl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-6} alkyl, oxo, oxido, heteroalkyl, halogen, haloalkyl, methoxy-phenyl, haloalkyl, $-NH_2$, $-N(CH_3)_2$, -NHCH2-phenyl, -NHCOCH3, cyano, $-SO_2NH_2$, $-SO_2CH_3$,

$-OCH_2COOCH_3$, $-OCH_2COOH$, $-SCH_2COOH$, $-COCH_3$, $-COOC(CH_3)_3$, $-COOC(CH_3)_3$, $-COOCH_2CH_3$,- COOH, pyridine-4-yl, , , benzyl, phenyl, said phenyl is optionally substituted with one to three substitutuents consisting of C_{1-6} alkyl , halogen, C_{1-6} alkoxyl, $-COOCH_3$;

-NH-phenyl, wherein said phenyl is optionally substituted with one to three substituents selected from C_{1-6} alkyl, and

pharmaceutically acceptable salts, hydrates, solvates and stereoisomers thereof.

(iv). A compound according to (iii), wherein

X^1 is $C(=O)R^3$, wherein

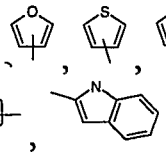
R^3 is selected from the group consisting of:

hydrogen, C_{1-6} alkyl, $-CH_2NH2$, $-CH_2CH_2NH_2$, $-C(CH_3)NH_2$, $-CH_2CH_2N(ethyl)_2$,, $-N(CH_3)_2$, $-N(CH_2CH_3)_2$, -N(iso-propyl)$_2$, $-NCH_3$-benzyl, $-NHCH_2COOCH_2CH_3$, -NH-phenyl, $-CH_2NHCOCH_3$, $CH_2CH_2CONH_2$, $-C(NH_2)CH_2C(CH_3)_2$, $-CH(NH_2)CH(CH_3)_2$,-CH2CN, $-C_{1-6}$ alkyl-OH, $-CH_2CH_2COOCH_3$, $-C(CH_3)_2OCOCH_3$, $-CH_2OCOCH_3$, -CH2SO2CH3, -CH2(CH3)SO2CH3, heteroalkyl,

$-(CH_2)_t$-C_{3-8} cycloalkyl selected from cyclopropanyl, cyclobutanyl, cyclopentanyl, cyclohexanyl, cycloheptanyl or , wherein t is 0 to 2, and
said $-C_{3-8}$ cycloalkyl is optionally substituted with one to three substituents selected independently from the group comprising: C_{1-6} alkyl, halogen, hydroxyl, cyano, oxo, phenyl,

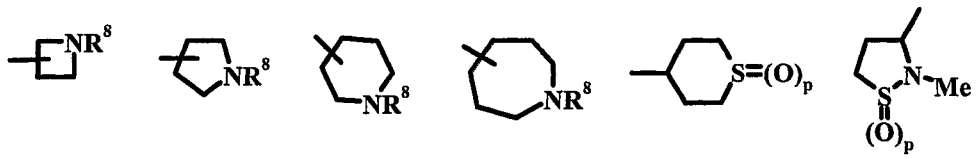
$-(CHR^m)_u$-phenyl, wherein R^m is H, hydroxyl or C_{1-6} alkyl, u is 0 to 2, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, $-SO_2NH_2$, $-NHCOCH_3$ or $-COOCH_3$,

$-(CH_2)_v$-heteroaryl selected from the group consisting of , wherein v 0 to 3, and

said heteroaryl is optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, halogen, phenyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, $-NH_2$, SO2NH2, -COCH3, oxo, oxido, ,

$-(CHR^n)_w$-heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl,

wherein R^n is hydrogen or C_{1-6} alkyl, w is 0 to 3, and

said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is C_{1-6} alkyl, C_{1-6} haloalkyl, furanyl, C_{1-6} alkoxy, C_{3-5} cycloalkyl, C_{1-6} alkylamino, -COO- C_{1-6} alkyl, -COOH, $-CH_2OCH_2OOH$, $-CH_2CH_2COOH$;

R^{10} is C_{1-6} alkyl, C_{1-6} dialkylamino, C_{3-8} cycloalkyl;

R^7 is hydrogen or Boc;

R^{15} is hydrogen or C_{1-6} alkyl.

(v). A compound according to (iii), wherein

X^1 is $S(=O)_2R^3$, wherein $R^3$is phenyl optionally substituted with one to three halogen.

(vi). A compound according to (iii), wherein

X^1 is $C(=O)OR^3$, wherein R^3 is C_{1-6} alkyl.

(vii). A compound according to (iii), wherein

X^2 is $C(=O)R^4$, wherein

R^4 is selected from the group consisting of:

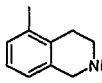
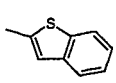
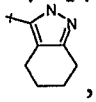
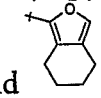
C_{1-6} alkyl, $-CH_2CH_2CONH_2$, $-CH_2NCH_3SO_2CH_3$, $-CH(CH(CH_3)_2)NHCOCH_3$,

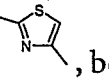
$-(CH_2)_eC_{3-8}$ cycloalkyl, wherein e is 0 to 2, and said cycloalkyl is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl,

C_{3-8} heterocycle, wherein said heterocyle is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxyl, -CO- C_{1-6} alkyl, oxo, benzyl, , or ,

$-(CHR^f)_f$-phenyl, wherein R^f is hydrogen or –NHCOCH3, f is 0 or 1, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, NO_2, -CO-C_{1-6} alkyl, $-COOCH_3$, $COOC(CH_3)_3$, $-NHSO_2CH_3$, NH_2, $-N(CH_3)_2$, $-N(CH_3)CH_2CH_2OH$, $-NHCOCH_3$,$-OCF_3$ $-OCH_2COOH$, pyridine-4-yl, ;

$(CH_2)_g$-heteroaryl, wherein g is 0 or 1, sais heteroary selected from the group consisting of furanyl, thiophenyl, isoxazolyl, thiazolyl, oxazolyl, pyrroly, pyrazolyl, imidazolyl, indolyl, benzofuranyl, pyranyl, pyridinyl, pyrimidinyl, chinoxalinyl, chinolinyl, , , , , and ,

said heteroaryl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-6} alkyl, oxo, oxido, heteroalkyl, halogen, haloalkyl, methoxy-phenyl, haloalkyl, $-NH_2$, $-N(CH_3)_2$, -NHCH2-phenyl, -NHCOCH3, cyano, $-SO_2NH_2$, $-SO_2CH_3$, $-OCH_2COOCH_3$, $-OCH_2COOH$, $-SCH_2COOH$, $-COCH_3$, $-COOC(CH_3)_3$, $-COOC(CH_3)_3$, $-COOCH_2CH_3$,- COOH, pyridine-4-yl, , , benzyl, phenyl, said phenyl is optionally substituted with one to three substitutuents consisting of C_{1-6} alkyl , halogen, C_{1-6} alkoxyl, $-COOCH_3$;

-NH-phenyl, wherein said phenyl is optionally substituted with one to three substituents selected from C_{1-6} alkyl.

(viii). A compound according to (iii), wherein

X^2 is $S(=O)_2R^4$, wherein

R^4 is phenyl optionally substituted with one to three C_{1-6} alkyl, or

heteroaryl selected from  , and said heteroaryl is optionally substituted with one to four substitutuents consisting C_{1-6} alkyl, halogen, $-COOCH_3$ or phenyl.

(ix). A compound according to (ii), wherein:

R^1 is phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of halogen and C_{1-6} alkyl, or

heteroaryl selected from the group consisting of pyridinyl, pyrimidinyl and thiazolyl said heteroaryl optionally substituted with one to three substituents independently selected from the group consisting of halogen, C_{1-6} alkyl and C_{1-6} haloalkyl;

R^2 is hydrogen;

R^5 is hydrogen or C_{1-6} alkyl;

R^{5a} is hydrogen, hydroxyl or C_{1-6} alkyl;

R^6 is hydrogen or C_{1-3} hydroxyalkyl;

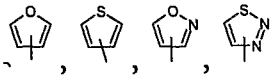
X^1 is selected from the group consisting $-C(=O)R^3$, $-S(=O)_2R^3$;

X^2 is selected from the group consisting $-C(=O)R^4$, $-S(=O)_2R^4$ or benzyl;

R^3 is selected from the group consisting of:

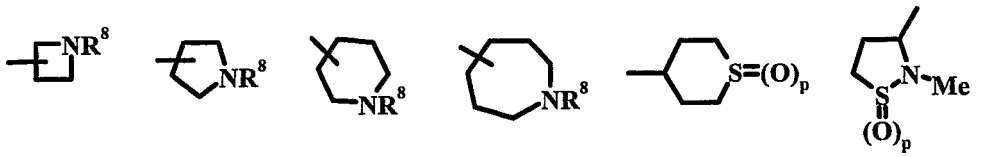
C_{1-6} alkyl, heteroalkyl, -phenyl-SO_2CH_3,

$-(CH_2)_t$-C_{3-8} cycloalkyl selected from the group consisting of cycopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, wherein t is 0 or 2, and said $-C_{3-8}$ cycloalkyl is optionally substituted with one to three substituents selected independently from the group consisting of halogen, hydroxyl and oxo,

heteroaryl selected from the group consisting of 

or ,wherein v 0 to 3, and said heteroaryl is optionally substituted with one to three C_{1-10} alkyl,

heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl, and

said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is C_{1-6} alkyl, C_{1-6} haloalkyl, furanyl, C_{1-6} alkoxy, C_{3-5} cycloalkyl, C_{1-6} alkylamino, -COO- C_{1-6} alkyl, -COOH, $-CH_2OCH_2OOH$, $-CH_2CH_2COOH$;

R^{10} is C_{1-6} alkyl, C_{1-6} dialkylamino, C_{3-8} cycloalkyl;

R^7 is hydrogen or Boc;

R^{15} is hydrogen or C_{1-6} alkyl;

$-NH-(CH_2)_x$-phenyl, wherein x is 0 or 1, and

said phenyl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, $-OCF_3$, heteroalkyl, haloalkyl, acetyl, dialkylamin, cyano, -COOH, $COOCH_2CH_3$,

$-NH(CH_2)_y$-heteroaryl, wherein y is 0 or 1, and

said heteroaryl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, haloalkyl,

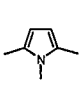
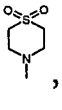
-NH-heterocycle, wherein said heteroaryl is optionally substituted with 1 to 2 substituents independently selected in each occurrence from the group consisting of $-COCF_3$, $COOCH_2$-phenyl,

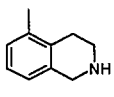
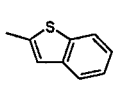
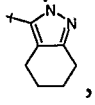
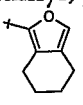
$-NH(CH_2)_z-C_{3-8}$ cycloalkyl selected from cycopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, wherein z is 0 or 1;

R^4 is selected from the group consisting of:

C_{1-6} alkyl,

C_{3-8} cycloalkyl selected from selected from cyclopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, and said cycloalkyl is optionally substituted with 1 to 4 substituents consisting of C_{1-6} alkyl,

C_{3-8} heterocycle selected from tetrahydrofuranyl and ,wherein said heterocyle is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl and -CO- C_{1-6} alkyl,
Phenyl, and said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, NO_2, -CO-C_{1-6} alkyl,
$-COOCH_3$, $COOC(CH_3)_3$,$-NHSO_2CH_3$, NH_2, $-N(CH_3)_2$, $-N(CH_3)CH_2CH_2OH$,
$-NHCOCH_3$,$-OCF_3$ $-OCH_2COOH$, pyridine-4-yl, , , ,

heteroaryl selected from the group consisting of furanyl, thiophenyl, isoxazolyl, thiazolyl, oxazolyl, pyrroly, pyrazolyl, imidazolyl, indolyl, benzofuranyl, pyranyl, pyridinyl, pyrimidinyl, chinoxalinyl, chinolinyl, , , , , and ,

said heteroaryl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-6} alkyl, oxo, oxido, heteroalkyl, halogen, haloalkyl, methoxy-phenyl, haloalkyl, $-NH_2$,

$-N(CH_3)_2$, -NHCH2-phenyl, -NHCOCH3, cyano, $-SO_2NH_2$, $-SO_2CH_3$, $-OCH_2COOCH_3$, $-OCH_2COOH$, $-SCH_2COOH$, $-COCH_3$, $-COOC(CH_3)_3$, $-COOC(CH_3)_3$, $-COOCH_2CH_3$,- COOH, pyridine-4-yl, , benzyl, phenyl, said phenyl is optionally substituted with one to three substituents consisting of C_{1-6} alkyl , halogen, C_{1-6} alkoxyl, $-COOCH_3$;

$-(NR^h)-(CH2)_h$-phenyl, wherein R^h is hydrogen or C_{1-6} alkyl, h is 0 or 1, and said phenyl is optionally substituted with one to three substituents selected from C_{1-6} alkyl, and

pharmaceutically acceptable salts, hydrates, solvates and stereoisomers thereof.

(x). A compound according to (ix), wherein

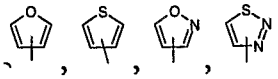
X^1 is $C(=O)R^3$, wherein

R^3 is selected from the group consisting of:

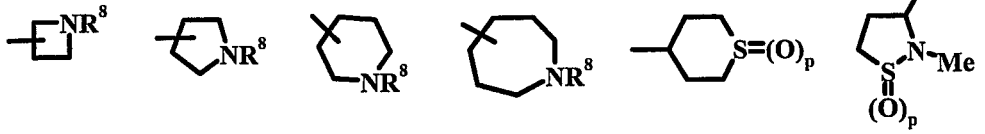
is selected from the group consisting of:

C_{1-6} alkyl, heteroalkyl, $-phenyl-SO_2CH_3$,

$-(CH_2)_t-C_{3-8}$ cycloalkyl selected from the group consisting of cycopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, wherein t is 0 or 2, and said $-C_{3-8}$ cycloalkyl is optionally substituted with one to three substituents selected independently from the group consisting of halogen, hydroxyl and oxo,

heteroaryl selected from the group consisting of , or ,wherein v 0 to 3, and said heteroaryl is optionally substituted with one to three C_{1-10} alkyl,

heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl, and

said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is C_{1-6} alkyl, C_{1-6} haloalkyl, furanyl, C_{1-6} alkoxy, C_{3-5} cycloalkyl, C_{1-6} alkylamino, -COO- C_{1-6} alkyl, -COOH, $-CH_2OCH_2OOH$, $-CH_2CH_2COOH$;

R^{10} is C_{1-6} alkyl, C_{1-6} dialkylamino, C_{3-8} cycloalkyl;

R^7 is hydrogen or Boc;

R^{15} is hydrogen or C_{1-6} alkyl;

$-NH-(CH_2)_x$-phenyl, wherein x is 0 or 1, and

said phenyl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, $-OCF_3$, heteroalkyl, haloalkyl, acetyl, dialkylamin, cyano, -COOH, $COOCH_2CH_3$,

$-NH(CH_2)_y$-heteroaryl, wherein y is 0 or 1, and

said heteroaryl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, haloalkyl,

-NH-heterocycle, wherein said heteroaryl is optionally substituted with 1 to 2 substituents independently selected in each occurrence from the group consisting of $-COCF_3$, $COOCH_2$-phenyl,

$-NH(CH_2)_z$-C_{3-8} cycloalkyl selected from cycopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, wherein z is 0 or 1.

(xi). A compound according to (ix), wherein

X^1 is $-S(=O)_2R^3$, wherein

R^3 phenyl.

(xii). A compound according to (ix), wherein

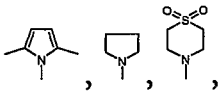
X^2 is $-C(=O)R^4$,

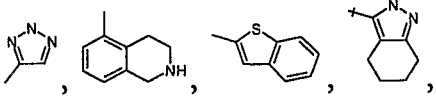
R^4 is selected from the group consisting of:

C_{1-6} alkyl,

C_{3-8} cycloalkyl selected from selected from cyclopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, and said cycloalkyl is optionally substituted with 1 to 4 substituents consisting of C_{1-6} alkyl,

C_{3-8} heterocycle selected from tetrahydrofuranyl and ,wherein said heterocyle is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl and -CO- C_{1-6} alkyl,

Phenyl, and said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, NO_2, -CO-C_{1-6} alkyl,
$-COOCH_3$, $COOC(CH_3)_3$,$-NHSO_2CH_3$, NH_2, $-N(CH_3)_2$, $-N(CH_3)CH_2CH_2OH$,
$-NHCOCH_3$,$-OCF_3$ $-OCH_2COOH$, pyridine-4-yl, ,

heteroaryl selected from the group consisting of furanyl, thiophenyl, isoxazolyl, thiazolyl, oxazolyl, pyrroly, pyrazolyl, imidazolyl, indolyl, benzofuranyl, pyranyl, pyridinyl, pyrimidinyl, chinoxalinyl, chinolinyl, ,
and ,

said heteroaryl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-6} alkyl, oxo, oxido, heteroalkyl, halogen, haloalkyl, methoxy-phenyl, haloalkyl, $-NH_2$, $-N(CH_3)_2$, -NHCH2-phenyl, -NHCOCH3, cyano, $-SO_2NH_2$, $-SO_2CH_3$, $-OCH_2COOCH_3$, $-OCH_2COOH$, $-SCH_2COOH$, $-COCH_3$, $-COOC(CH_3)_3$, $-COOC(CH_3)_3$, $-COOCH_2CH_3$,- COOH, pyridine-4-yl, , benzyl, phenyl, said phenyl is optionally substituted with one to three substituents consisting of C_{1-6} alkyl , halogen, C_{1-6} alkoxyl, $-COOCH_3$;

$-(NR^h)-(CH2)_h$-phenyl, wherein R^h is hydrogen or C_{1-6} alkyl, h is 0 or 1, and said phenyl is optionally substituted with one to three substituents selected from C_{1-6} alkyl.

(xiii). A compound according to (ix), wherein

X^2 is $-S(=O)_2R^4$,

R^4 is phenyl optionally substituted with one to four substituents consiting of C_{1-6} alkyl, halogen, cyano, $-COOCH_3$ and COOH, or

heteroaryl selected from isoxazolyl, pyrazolyl, furanyl, pyrrolyl and pyridinyl, and said heteroaryl is optionally substituted with one to four substituents consisting of C_{1-6} alkyl, halogen, phenyl, $-N(CH_3)_2$, -COOH, $-COOCH_3$ and $-COOCH_2CH_3$.

In one embodiment of the present invention there is provided a compound according to formula I wherein R^1, R^2, R^3, R^4, R^5, R^{5a}, R^6, R^7, R^8, R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14a}, R^{14b}, R^{14c}, R^{15}, R^{16}, X^1, X^2, X^3, Y, r, s, n and p are as defined hereinabove

In the following embodiments of the present invention all substituent definitions are intended to take the broadest form as disclosed in the Summary of the invention unless limited in the description of the following embodiments.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^1 is optionally substituted phenyl; X^1 is $-C(=O)R^3$ or $-SO_2R^3$; X^2 is $-C(=O)R^4$; and R^2, R^5, R^{5a} and R^6 are hydrogen.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^1 is optionally substituted phenyl; X^1 is $-C(=O)R^3$; X^2 is $-C(=O)R^4$; R^3 is a heterocycle as defined in section (v) of R^3; and R^2, R^5, R^{5a} and R^6 are hydrogen.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^1 is optionally substituted phenyl; X^1 is $-C(=O)R^3$; X^2 is $-C(=O)R^4$; R^3 is a heterocycle according to formulae IIa-IId as defined in section (v) of R^3; R^4 is a phenyl or heteroaryl group wherein at least one, and preferably both, atom(s) adjacent to the atom linked to X^2 are substituted carbon atoms and R^2, R^5, R^{5a} and R^6 are hydrogen. A substituted carbon as used herein refers to a carbon bearing a non-hydrogen substituent.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^1 is 4-methyl-3-chloro-phenyl; X^1 is $-C(=O)R^3$; X^2 is $-C(=O)R^4$; R^3 is a heterocycle according to formula IIb or IIc defined in section (v) of R^3; R^4 is 2,6-dimethyl-pyrimidin-5-yl, 2,6-dimethyl-pyridine-3-yl or 2,6-dimethyl-phenyl; and R^2, R^5, R^{5a} and R^6 are hydrogen.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^1 is 4-methyl-3-chloro-phenyl; X^1 is $-C(=O)R^3$; X^2 is $-C(=O)R^4$; R^3 is a heterocycle according to formula IIb or IIc defined in section (v) wherein the piperidinyl or pyrrolidinyl nitrogen atom is substituted by $-C(=O)CH_3$, $-C(=O)C(=O)O-C_{1-6}$ alkyl or $-C(=O)C(=O)OH$; R^4 is 2,6-dimethyl-pyrimidin-5-yl, 2,6-dimethyl-pyridine-3-yl or 2,6-dimethyl-phenyl; and R^2, R^5, R^{5a} and R^6 are hydrogen.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^1 is optionally substituted phenyl; X^1 is $-C(=O)NR^{14a}R^{14b}$ or $-S(=O)_2NR^{14a}R^{14b}$; X^2 is $-C(=O)R^4$; and R^2, R^5, R^{5a} and R^6 are hydrogen.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^1 is optionally substituted phenyl; X^1 is $-C(=O)NR^{14a}R^{14b}$ or $-S(=O)_2NR^{14a}R^{14b}$ wherein R^{14a} is optionally substituted phenyl and R^{14b} is hydrogen; X^2 is $-C(=O)R^4$; and R^2, R^5, R^{5a} and R^6 are hydrogen. Optionally substituted phenyl refers to the substituents listed in the definition in section (i) of R^3.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^1 is 4-methyl-3-chloro-phenyl; X^1 is $-C(=O)NR^{14a}R^{14b}$ or $-S(=O)_2NR^{14a}R^{14b}$ wherein R^{14a} is phenyl substituted by a carboxyl or C_{1-6} alkoxycarbonyl and R^{14b} is hydrogen; X^2 is $-C(=O)R^4$; and R^2, R^5, R^{5a} and R^6 are hydrogen. Optionally substituted phenyl refers to the substituents listed in the definition in section (i) of R^3.

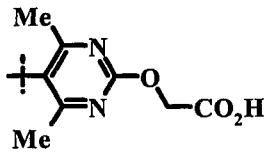
In another embodiment of the present invention there is provided a compound according to formula I wherein R^2 is optionally substituted phenyl; X^1 is $-C(=O)R^3$ or $-SO_2R^3$; X^2 is $-C(=O)R^4$; and R^1, R^5, R^{5a} and R^6 are hydrogen.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^2 is optionally substituted phenyl; X^1 is $-C(=O)R^3$; X^2 is $-C(=O)R^4$; and R^1, R^5, R^{5a} and R^6 are hydrogen. In this embodiment R^3 is optionally substituted aryl, optionally substituted cycloalkyl, 4-oxo-cyclohexyl, 3-oxo-cyclobutyl or tetrahydrofuranyl and R^4 is a phenyl or heteroaryl group wherein at least one, and preferably both, atom(s) adjacent to the atom linked to X^2 are substituted carbon atoms.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^2 is optionally substituted phenyl; X^1 is $-C(=O)R^3$; X^2 is $-C(=O)R^4$; and R^1, R^5, R^{5a} and R^6 are hydrogen. In this embodiment R^3 is cycloalkyl optionally substituted with 1 to 4 fluorine atoms and R^4 is a phenyl or heteroaryl group

wherein at least one, and preferably two, atom(s) adjacent to the atom linked to X^2 are substituted carbon atoms.

In another embodiment of the present invention there is provided a compound according to formula I wherein R^2 is optionally substituted phenyl; X^1 is $-C(=O)R^3$; X^2 is $-C(=O)R^4$; and R^1, R^5, R^{5a} and R^6 are hydrogen. In this embodiment R^3 is cyclopentyl, 4,4-difluorocyclohexyl or 3,3-difluorocyclobutyl and R^4 is 2,6-dimethyl-pyrimidin-5-yl, 2,6-dimethyl-pyridine-3-yl, 2,6-dimethyl-phenyl or III.



(III)

In another embodiment of the present invention there is provided a compound according to formula I wherein R^2 is phenyl; X^1 is $-C(=O)R^3$; X^2 is $-C(=O)R^4$; and R^1, R^5, R^{5a} and R^6 are hydrogen. In this embodiment R^3 is cyclopentyl, 4,4-difluorocyclohexyl or 3,3-difluorocyclobutyl and R^4 is 2,6-dimethyl-pyrimidin-5-yl, 2,6-dimethyl-pyridine-3-yl, 2,6-dimethyl-phenyl or III.

In another embodiment of the present invention there is provided a compound according to formula I selected from the following:

1-acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide,

1-acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide,

[4-((3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-piperidin-1-yl]-oxo-acetic acid; compound with trifluoro-acetic acid,

cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

cyclopentanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide,

cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide,

3-(3-(3-Chloro-4-methyl-phenyl)-3-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-ureido)-benzoic acid; compound with trifluoro-acetic acid

4,4-difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-butyl}-amide, and,

(5-{5-[(S)-3-(cyclopentanecarbonyl-amino)-3-phenyl-propyl]-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl}-4,6-dimethyl-pyrimidin-2-yloxy)-acetic acid; compound with trifluoro-acetic acid.

In another embodiment of the present invention there is provided a method for treating or preventing an human immunodeficiency virus (HIV) infection, or treating AIDS or ARC, in a patient in need thereof which comprises administering to the patient in need thereof a therapeutically effective amount of a compound of according to formula I wherein R^1, R^2, R^3, R^4, R^5, R^{5a}, R^6, R^7, R^8 , R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14a}, R^{14b}, R^{14c}, R^{15}, R^{16}, X^1, X^2, X^3, Y, r, s, n and p are as defined hereinabove.

In another embodiment of the present invention there is provided a method for treating or preventing an human immunodeficiency virus (HIV) infection, or treating AIDS or ARC, in a patient in need thereof which comprises co-administering to a patient in need thereof a therapeutically effective amount of a compound of according to formula I wherein R^1, R^2, R^3, R^4, R^5, R^{5a}, R^6, R^7, R^8 , R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14a}, R^{14b}, R^{14c}, R^{15}, R^{16}, X^1, X^2, X^3, Y, r, s, n and p are as defined hereinabove and at least one of an HIV nucleoside reverse transcriptase inhibitors, HIV non-nucleoside reverse transcriptase inhibitors, HIV protease inhibitors or viral fusion inhibitors.

In another embodiment of the present invention there is provided a method for treating or preventing an human immunodeficiency virus (HIV) infection, or treating AIDS or ARC, in a patient in need thereof which comprises co-administering to a patient in need thereof a therapeutically effective amount of a compound of according to formula I wherein R^1, R^2, R^3, R^4, R^5, R^{5a}, R^6, R^7, R^8 , R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14a}, R^{14b}, R^{14c}, R^{15}, R^{16}, X^1, X^2, X^3, Y, r, s, n and p are as defined hereinabove and at least one of efavirenz, nevirapine or delavirdine, zidovudine, didanosin, zalcitabine, stavudine.; lamivudine, abacavir, adefovir and dipivoxil, saquinavir, ritonavir, nelfinavir, indinavir, amprenavir and lopinavir or T-20.

In another embodiment of the present invention there is provided a method of treating a mammal with a disease state that is alleviated by a CCR5 receptor antagonist wherein said disease state that is solid organ transplant rejection, graft v. host disease, arthritis, rheumatoid arthritis, inflammatory bowel disease, atopic dermatitis, psoriasis,

asthma, allergies or multiple sclerosis comprising administering to a mammal in need thereof a therapeutically effective amount of a compound of according to formula I wherein R^1, R^2, R^3, R^4, R^5, R^{5a}, R^6, R^7, R^8 , R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14a}, R^{14b}, R^{14c}, R^{15}, R^{16}, X^1, X^2, X^3, Y, r, s, n and p are as defined hereinabove.

In another embodiment of the present invention there is provided a method of treating a mammal with a disease state that is alleviated by a CCR5 receptor antagonist wherein said disease state that is solid organ transplant rejection, graft v. host disease, arthritis, rheumatoid arthritis, inflammatory bowel disease, atopic dermatitis, psoriasis, asthma, allergies or multiple sclerosis comprising co-administering to a mammal in need thereof a therapeutically effective amount of a compound of according to formula I wherein R^1, R^2, R^3, R^4, R^5, R^{5a}, R^6, R^7, R^8 , R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14a}, R^{14b}, R^{14c}, R^{15}, R^{16}, X^1, X^2, X^3, Y, r, s, n and p are as defined hereinabove and at least one other immune system modulator.

In another embodiment of the present invention there is provided a method of treating a human with a disease state that is alleviated by a CCR5 receptor antagonist wherein said disease state that is solid organ transplant rejection, graft v. host disease, arthritis, rheumatoid arthritis, inflammatory bowel disease, atopic dermatitis, psoriasis, asthma, allergies or multiple sclerosis comprising co-administering to a mammal in need thereof a therapeutically effective amount of a compound of according to formula I wherein R^1, R^2, R^3, R^4, R^5, R^{5a}, R^6, R^7, R^8 , R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14a}, R^{14b}, R^{14c}, R^{15}, R^{16}, X^1, X^2, X^3, Y, r, s, n and p are as defined hereinabove and at least one other immune system modulator.

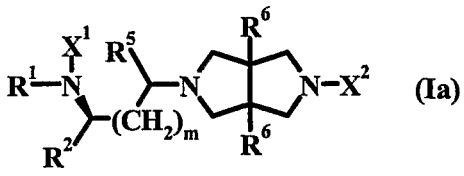
In another embodiment of the present invention there is provided a pharmaceutical composition for treating or preventing an human immunodeficiency virus (HIV) infection, or treating AIDS or ARC comprising a compound according to formula I wherein R^1, R^2, R^3, R^4, R^5, R^{5a}, R^6, R^7, R^8 , R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14a}, R^{14b}, R^{14c}, R^{15}, R^{16}, X^1, X^2, X^3, Y, r, s, n and p are as defined hereinabove admixed with at least one pharmaceutical acceptable carrier, diluent or excipient.

In another embodiment of the present invention there is provided a pharmaceutical composition for treating a mammal with a disease state that is alleviated by a CCR5 receptor antagonist wherein said disease is solid organ transplant rejection, graft v. host disease, arthritis, rheumatoid arthritis, inflammatory bowel disease, atopic dermatitis, psoriasis, asthma, allergies or multiple sclerosis comprising a compound according to formula I wherein R^1, R^2, R^3, R^4, R^5, R^{5a}, R^6, R^7, R^8 , R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14a}, R^{14b}, R^{14c}, R^{15}, R^{16}, X^1, X^2, X^3, Y, r, s, n and p are as defined hereinabove admixed with at least one pharmaceutical acceptable carrier, diluent or excipient.

In another embodiment of the present invention there is provided a process for the preparation of a compound according to formula wherein R^1 is hydrogen and R^2 is optionally substituted phenyl or optionally substituted heteroaryl comprising a reductive amination step, removal of protecting if present and acylation with an acylation agent selected to provide compounds disclosed in the present invention.

In another embodiment of the present invention there is provided a process for the preparation of a compound according to formula wherein R^2 is hydrogen and R^1 is optionally substituted phenyl or optionally substituted heteroaryl comprising an alkylation step, removal of protecting if present and acylation with an acylation agent selected to provide compounds disclosed in the present invention.

In another embodiment of the present invention there is provided a compound according to formula **Ia** wherein:



(Ia)

X^1 is selected from the group consisting $-C(=O)R^3$, $-S(=O)_2R^3$, $-C(=O)OR^3$ and $-C(=O)NHR^3$; X^2 is selected from the group consisting $-C(=O)R^4$, $-S(=O)_2R^4$ or $-CH_2$-Aryl; one of R^1 and R^2 is:

(i) aryl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of: (a) C_{1-10} alkyl,(b) C_{1-10} heteroalkyl, (c) C_{1-6} haloalkyl, (d) C_{1-6} alkoxy, (e) C_{1-6} thioalkyl, (f) amino, (g) C_{1-6} alkyl amino, (h) C_{1-6} dialkylamino, (g) C_{1-6} acylamino, (i) carbamoyl, N-alkylcarbamoyl, N,N-dialkylcarbamoyl, (j) ureido, (k) nitro, (l) cyano, (m) halogen, (n) C_{1-6} alkylsulfonyl, (o) sulfamoyl, N-alkylsulfamoyl or N, N-dialkylsulfamoyl, (p) alkylsulfonamido or optionally substituted arylsulfonamido, (q) optionally substituted aryloxy, (r) optionally substituted heteroaryloxy, and, (s) $-Y(CH_2)_nR^{11}$ wherein R^{11} is selected from the group consisting of cyano, $-CO_2R^{12}$, $-CONR^{12}R^{13}$, $-SO_2N^{12}R^{13}$, $-NHSO_2R^{12}$ and $-NHSO_2NR^{12}R^{13}$ R^{12} and R^{13} (i) are independently hydrogen, C_{1-6} alkyl or C_{1-6} heteroalkyl, or (*ii*) R^{12} and R^{13}, when both attached to a nitrogen atom, together can be C_{2-5} alkylene, and Y is a direct bond, -O-, -S- or $-NR^{12}$-; or, (ii) one of R^1 and R^2 are heteroaryl selected from the group consisting of pyridinyl, pyridinyl-N-oxide, pyrimidinyl, pyrazinyl, pyridazinyl, isoxazolyl, isothiazolyl, pyrrolyl, thiazolyl and thienyl said heteroaryl optionally substituted with one to three substituents independently selected from the group of optional substituents in (i) above; and, the other of R^1 and R^2 is hydrogen; R^3 is selected from the group consisting of: (i) aryl optionally substituted with one to four substituents selected independently in

each incidence from the group consisting of (i)(a) to (i)(s) described for R^1 and R^2 above, (ii) aryl C_{1-6} alkyl wherein aryl is as described in (i) above, (iii) heteroaryl selected from the group consisting of pyridinyl, pyridinyl-N-oxide, pyrimidinyl, pyrazinyl, pyridazinyl, isoxazolyl, isothiazolyl, pyrrolyl, thiazolyl, oxazolinyl, pyrazolinyl, oxadiazolinyl, thiadiazolinyl, pyrazolyl, tetrazole, imidazolyl, triazolyl, triazolyl, indolyl, quinolinyl, isoquinolinyl and thienyl said heteroaryl optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, aryl, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, carbamoyl, nitro, cyano, amino, C_{1-3} alkylamino, C_{1-3}-dialkylamino, morpholino and halogen, (iv) heteroaryl C_{1-6} alkyl wherein heteroaryl is as described above, (v) heterocycle selected from the group consisting of IIa-f, furfuryl, oxetanyl, tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl, 1,2,3,4-tetrahydroisoquinolinyl and 2-aza-bicyclo[2.2.1]heptanyl said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, alkylamino, dialkylamino and phenylamino, or the two hydrogens on a carbon bonded to a nitrogen are replaced by oxygen (oxo), wherein: R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, COR^9 or SO_2R^{10}; R^9 is selected from the group consisting of: (i) C_{1-6} alkyl optionally substituted with one or two groups independently selected from the group consisting of C_{1-3} alkoxy, C_{1-3} acyloxy, hydroxyl, phenyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, aminocarbonylpyridyl wherein said aminocarbonylpyridyl can optionally be the N-oxide and aminobenzoyl wherein said phenyl or said benzoyl radical is optionally substituted with one to three groups independently selected from the group consisting of amino, alkylamino, dialkylamino, hydroxyl, halogen and C_{1-3} alkoxy, (ii) phenyl optionally substituted with one or two groups independently selected from the group consisting of sulfamoyl, acetylamino, halogen, C_{1-6} alkyl, C_{1-6} haloalkyl, and C_{1-6}alkoxy, (iii) NH_2, C_{1-3} alkylamino, C_{1-3} dialkylamino, (iv) $C(=O)NH_2$, (v) 1H indole-3-carbonyl, (vi) furyl, pyridinyl, or pyridinyl N-oxide, (vii) N-acetylpiperidin-4yl and, (viii) furfuryl;R^{10} is selected from the group consisting of: (i) C_{1-6} alkyl, (ii) C_{1-3} haloalkyl, (iii) C_{1-3} haloalkoxy, (iv) C_{3-8} cycloalkyl, (v) phenyl optionally substituted with C_{1-3} alkoxy, halogen, nitro, acetamido, carboxy and, (vi) aralkyl, (vi) heterocyclyl C_{1-6} alkyl wherein heterocyclyl is as defined above, (vii) hydrogen, (viii) C_{1-10} alkyl substituted with one to three substituents independently selected at each occurrence with hydroxy, halogen, C_{1-6} alkoxy, C_{1-6} thioalkyl, acyloxy, C_{1-6} acylamino, amino, alkylamino, dialkylamino, cyano, C_{1-6} alkylsulfonyl, C_{1-6} sulfinyl, N-piperidinyl, N-pyrrolidinyl, imidazolyl, carbamoyl and alkoxycarbonyl, (ix) C_{3-7} cycloalkyl optionally substituted with 1 to 4 fluorine, C_{1-3} alkyl or phenyl; (x) C_{1-3} alkyl-C_{3-7} cycloalkyl optionally substituted with 1 to 4 fluorine, C_{1-3} alkyl or phenyl; R^4 is selected from the group consisting of: (i) aryl optionally substituted with one to four substituents selected independently in each incidence from the group

consisting of (i)(a) to (i)(s) decribed for R^1 and R^2 above, (ii) aryl C_{1-6} alkyl wherein aryl is as described above, (iii) heteroaryl selected from the group consisting of pyridinyl, pyrimidinyl, pyrazinyl, pyridazinyl, isoxazolyl, isothiazolyl, pyrrolyl, thiazolyl, oxazolinyl, pyrazolinyl, oxadiazolinyl, thiadiazolinyl, pyrazolyl, tetrazole, imidazolyl, triazolyl, triazolyl, indolyl, quinolinyl, isoquinolinyl, indazole, 4,5,6,7-tetrahydroindazole, 1,4,5,6-tetrahydrocyclopentylpyrazolyl, imidazo[2,1-b]thiazolyl and thienyl said heteroaryl optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, aryl, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, carbamoyl, nitro, cyano and halogen, (iv) heteroaryl C_{1-6} alkyl wherein heteroaryl is as described above, (v) heterocycle selected from the group consisting of azetidinyl, azepinyl, furfuryl, oxetanyl, piperidinyl, pyrrolidinyl, tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and 1,2,3,4-tetrahydroisoquinolinyl and said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, alkylamino, dialkylamino and phenylamino, or the two hydrogens on a carbon bonded to a nitrogen are replaced by oxygen (oxo); (vi) heterocyclyl C_{1-6} alkyl wherein heterocyclyl is as defined above, (vii) hydrogen, (viii)C_{1-10} alkyl substituted with one to three substituents independently selected at each occurrence with hydroxy, halogen, C_{1-6} alkoxy, C_{1-6} thioalkyl, acyloxy, C_{1-6} acylamino, amino, alkylamino, dialkylamino, cyano, C_{1-6} alkylsulfonyl, C_{1-6} sulfinyl, N-piperidinyl, N-pyrrolidinyl, imidazolyl, carbamoyl and alkoxycarbonyl, (ix) C_{3-7} cycloalkyl optionally substituted with 1 to 4 fluorine, C_{1-3} alkyl or phenyl, (x) C_{1-3} alkyl-C_{3-7} cycloalkyl optionally substituted with 1 to 4 fluorine, C_{1-3} alkyl or phenyl; R^5 is hydrogen, C_{1-6} alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl,-CO_2R^{12} or $CONR^{12}R^{13}$, R^6 is independently selected from the group consisting of hydrogen, C_{1-10} alkyl, hydroxy, C_{1-6} alkoxy, C_{1-3} hydroxyalkyl and C_{1-3} alkoxy-C_{1-3} alkyl; m is 1 to 2; n is 1 to 6; p is 0 to 2; and, pharmaceutically acceptable salts, hydrates, solvates and stereoisomers thereof.

The phrase “a” or “an” entity as used herein refers to one or more of that entity; for example, a compound refers to one or more compounds or at least one compound. As such, the terms “a” (or “an”), “one or more”, and “at least one” can be used interchangeably herein.

The phrase "as defined hereinabove" refers to the first definition provided in the Summary of the Invention.

The term “optional” or “optionally” as used herein means that a subsequently described event or circumstance may, but need not, occur, and that the description includes instances where the event or circumstance occurs and instances in which it does

not. For example, "optionally substituted" means that the moiety may be hydrogen or a substituent.

It is contemplated that the definitions described herein may be appended to form chemically-relevant combinations, such as "heteroalkylaryl," "haloalkylheteroaryl," "arylalkylheterocyclyl," "alkylcarbonyl," "alkoxyalkyl," and the like. The term -(ar)alkyl refers to either an unsubstituted alkyl or an aralkyl group. The term (hetero)aryl refers to either an aryl or a heteroaryl group.

The term "alkyl" as used herein denotes an unbranched or branched chain, saturated, monovalent hydrocarbon residue containing 1 to 10 carbon atoms. The term "lower alkyl" denotes a straight or branched chain hydrocarbon residue containing 1 to 6 carbon atoms. "C_{1-10} alkyl" as used herein refers to an alkyl composed of 1 to 10 carbons. One or more of the carbon atoms may optionally be replaced by oxygen, sulfur, substituted or unsubstituted nitrogen atom(s). Examples of alkyl groups include, but are not limited to, lower alkyl groups include methyl, ethyl, propyl, *i*-propyl, *n*-butyl, *i*-butyl, *t*-butyl or pentyl, isopentyl, neopentyl, hexyl, heptyl, and octyl. The term (ar)alkyl or (heteroaryl)alkyl indicate the alkyl group is optionally substituted by an aryl or a heteroaryl group respectively.

When the term "alkyl" is used as a suffix following another term, as in "phenylalkyl," or "hydroxyalkyl," this is intended to refer to an alkyl group, as defined above, being substituted with one to two substituents selected from the other specifically-named group. Thus, for example, "phenylalkyl" refers to an alkyl group having one to two phenyl substituents, and thus includes benzyl, phenylethyl, and biphenyl. An "alkylaminoalkyl" is an alkyl group having one to two alkylamino substituents. "Hydroxyalkyl" includes 2-hydroxyethyl, 2-hydroxypropyl, 1-(hydroxymethyl)-2-methylpropyl, 2-hydroxybutyl, 2,3-dihydroxybutyl, 2-(hydroxymethyl), 3-hydroxypropyl, and so forth. Accordingly, as used herein, the term "hydroxyalkyl" is used to define a subset of heteroalkyl groups defined below.

The term "alkylene" as used herein denotes a divalent saturated linear hydrocarbon radical of 1 to 8 carbon atoms or a branched saturated divalent hydrocarbon radical of 3 to 8 carbon atoms, unless otherwise indicated. Examples of alkylene radicals include, but are not limited to, methylene, ethylene, propylene, 2-methyl-propylene, butylene, 2-ethylbutylene.

The term "haloalkyl" as used herein denotes a unbranched or branched chain alkyl group as defined above wherein 1, 2, 3 or more hydrogen atoms are substituted by a halogen. Examples are 1-fluoromethyl, 1-chloromethyl, 1-bromomethyl, 1-iodomethyl,

trifluoromethyl, trichloromethyl, tribromomethyl, triiodomethyl, 1-fluoroethyl, 1-chloroethyl, 1-bromoethyl, 1-iodoethyl, 2-fluoroethyl, 2-chloroethyl, 2-bromoethyl, 2-iodoethyl, 2,2-dichloroethyl, 3-bromopropyl or 2,2,2-trifluoroethyl.

The term "heteroalkyl" as used herein means an alkyl radical as defined herein wherein one, two or three hydrogen atoms have been replaced with a substituent independently selected from the group consisting of $-OR^a$, $-NR^bR^c$, and $-S(O)_nR^d$ (where n is an integer from 0 to 2), with the understanding that the point of attachment of the heteroalkyl radical is through a carbon atom, wherein R^a is hydrogen, C_{1-6} acyl, C_{1-6} alkyl, C_{3-8} cycloalkyl, or C_{3-8} cycloalkyl-C_{1-6} alkyl; R^b and R^c are independently of each other hydrogen, C_{1-6} acyl, C_{1-6} alkyl, C_{3-8} cycloalkyl, or C_{3-8} cycloalkyl- C_{1-6} alkyl; and when n is 0, R^d is hydrogen, C_{1-6} alkyl, C_{3-8} cycloalkyl, or C_{3-8} cycloalkyl-C_{1-6} alkyl, and when n is 1 or 2, R^d is C_{1-6} alkyl, C_{3-8} cycloalkyl, C_{3-8} cycloalkyl- C_{1-6} alkyl, amino, C_{1-6} acylamino, or C_{1-6} alkylamino. Representative examples include, but are not limited to, 2-hydroxyethyl, 3-hydroxypropyl, 2-hydroxy-1-hydroxymethylethyl, 2,3-dihydroxypropyl, 1-hydroxymethylethyl, 3-hydroxybutyl, 2,3-dihydroxybutyl, 2-hydroxy-1-methylpropyl, 2-aminoethyl, 3-aminopropyl, 2-methylsulfonylethyl, aminosulfonylmethyl, aminosulfonylethyl, aminosulfonylpropyl, methylaminosulfonylmethyl, methylaminosulfonylethyl, methylaminosulfonylpropyl, and the like.

The term “acyl” as used herein denotes a group of formula -C(=O)R wherein R is hydrogen or lower alkyl as defined herein. The term or "alkylcarbonyl" as used herein denotes a group of formula C(=O)R wherein R is alkyl as defined herein. The term "arylcarbonyl" as used herein means a group of formula C(=O)R wherein R is an aryl group; the term "benzoyl" as used herein an "arylcarbonyl" group wherein R is phenyl.

The term “acylamino” as used herein denotes a group of formula -NHC(=O)R wherein R is hydrogen or lower alkyl as defined herein

The term "acyloxy" as used herein denotes the radical -OC(O)R, wherein R is a lower alkyl radical as defined herein. Examples of acyloxy radicals include, but are not limited to, acetoxy, propionyloxy.

The term "alkoxy" as used herein means an -O-alkyl group, wherein alkyl is as defined above such as methoxy, ethoxy, *n*-propyloxy, *i*-propyloxy, *n*-butyloxy, *i*-butyloxy, *t*-butyloxy, pentyloxy, hexyloxy, including their isomers. "Lower alkoxy" as used herein denotes an alkoxy group with a "lower alkyl" group as previously defined. "C_{1-10} alkoxy" as used herein refers to an-O-alkyl wherein alkyl is C_{1-10}.

The term "alkylthio" or "thioalkyl" means an -S-alkyl group, wherein alkyl is as defined above such as meththio, ethylthio, *n*-propylthio, *i*-propylthio, *n*-butylthio,

hexylthio, including their isomers. "Lower alkylthio" or "lower thioalkyl" as used herein denotes an alkylthio group with a "lower alkyl" group as previously defined. "C_{1-10} alkylthio" as used herein refers to an-S-alkyl wherein alkyl is C_{1-10}.

The prefix "carbamoyl" as used herein means the radical -$CONH_2$. The prefix "N-alkylcarbamoyl" and "N,N-dialkylcarbamoyl" means a the radical CONHR' or CONR'R" respectively wherein the R' and R" groups are independently alkyl as defined herein. The prefix "N-arylcarbamoyl" denotes the radical CONHR' wherein R' is an aryl radical as defined herein.

The terms "amino", "alkylamino" and "dialkylamino" as used herein refer to -NH_2, -NHR and -NR_2 respectively and R is alkyl as defined above. The two alkyl groups attached to a nitrogen in a dialkyl moiety can be the same or different. The terms "aminoalkyl", "alkylaminoalkyl" and "dialkylaminoalkyl" as used herein refer to $NH_2(CH_2)$n-, $RHN(CH_2)$n-, and $R_2N(CH_2)$n- respectively wherein n is 1 to 6 and R is alkyl as defined above. "C_{1-10} alkylamino" as used herein refers to an-aminoalkyl wherein alkyl is C_{1-10}. The term "phenylamino" as used herein refers to -NHPh wherein Ph represents an optionally substituted phenyl group.

The term "halogen" or "halo" as used herein means fluorine, chlorine, bromine, or iodine.

The term "aryl" as used herein denotes a monovalent aromatic carbocyclic radical containing 5 to 15 carbon atoms consisting of one individual ring, or one or more fused rings in which at least one ring is aromatic in nature, which can optionally be substituted with one or more, preferably one or three substituents independently selected from hydroxy, thio, cyano, alkyl, alkoxy, lower haloalkoxy, alkylthio, halogen , haloalkyl, hydroxyalkyl, nitro, alkoxycarbonyl, amino, alkylamino, dialkylamino, aminoalkyl, alkylaminoalkyl, and dialkylaminoalkyl, alkylsulfonyl, arylsulfinyl, alkylaminosulfonyl, arylaminosulfonyl, alkylsulfonylamino, arylsulfonylamino, carbamoyl, alkylcarbamoyl and dialkylcarbamoyl, arylcarbamoyl, alkylcarbonylamino, arylcarbonylamino, unless otherwise indicated. Alternatively two adjacent atoms of the aryl ring may be substituted with a methylenedioxy or ethylenedioxy group. Thus a bicyclic aryl substituents may be fused to a heterocyclyl or heteroaryl ring; however, the point of attachment of bicyclic aryl substituent is on the carbocyclic aromatic ring. Examples of aryl radicals include, phenyl, naphthyl, indanyl, anthraquinolyl, tetrahydronaphthyl, 3,4-methylenedioxyphenyl, 1,2,3,4-tetrahydroquinolin-7-yl, 1,2,3,4-tetrahydroisoquinoline-7-yl, and the like.

The term "arylalkyl" or "aralkyl" as used herein denotes the radical R'R"-, wherein R' is an aryl radical as defined herein, and R" is an alkylene radical as defined herein with the understanding that the attachment point of the arylalkyl moiety will be on the alkylene radical. R" is an alkylene chain comprising 1 to 6 methylenes. The term "phenyl C_{1-6} alkyl" refers to a radical R'R" wherein R' is a phenyl group and R" is an alkylene chain comprising 1 to 6 methylenes. Examples of arylalkyl radicals include, but are not limited to, benzyl, phenylethyl, 3-phenylpropyl.

The term "aryloxy" as used herein denotes a O-aryl group, wherein aryl is as defined above. An aryloxy group can be unsubstituted or substituted with one or two suitable substituents. The term "phenoxy" refers to an aryloxy group wherein the aryl moiety is a phenyl ring.

The term "heteroaryl" or "heteroaromatic" as used herein means a monocyclic or bicyclic radical of 5 to 12 ring atoms having at least one aromatic ring containing four to eight atoms per ring, incorporating one or more N, O, or S heteroatoms, the remaining ring atoms being carbon, with the understanding that the attachment point of the heteroaryl radical will be on a heteroaryl ring. As well known to those skilled in the art, heteroaryl rings have less aromatic character than their all-carbon counter parts. Thus, for the purposes of the invention, a heteroaryl group need only have some degree of aromatic character. Examples of heteroaryl moieties include monocyclic aromatic heterocycles having 5 to 6 ring atoms and 1 to 3 heteroatoms include, but is not limited to, pyridinyl, pyrimidinyl, pyrazinyl, pyrrolyl, pyrazolyl, imidazolyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, triazolinyl, thiadiazolyl and oxadiaxolinyl which can optionally be substituted with one or more, preferably one or two substituents selected from hydroxy, cyano, alkyl, alkoxy, thio, lower haloalkoxy, alkylthio, halo, haloalkyl, alkylsulfinyl, alkylsulfonyl, halogen, amino, alkylamino, dialkylamino, aminoalkyl, alkylaminoalkyl, and dialkylaminoalkyl, nitro, alkoxycarbonyl and carbamoyl, alkylcarbamoyl, dialkylcarbamoyl, arylcarbamoyl, alkylcarbonylamino and arylcarbonylamino. Examples of bicyclic moieties include, but are not limited to, quinolinyl, isoquinolinyl, benzofuryl, benzothiophenyl, benzoxazole, benzisoxazole, benzothiazole and benzisothiazole. Bicyclic moieties can be optionally substituted on either ring; however the point of attachment is on a ring containing a heteroatom.

The term "heteroaryl alkyl" or "heteroaralkyl" means the radical of the formula R'R", wherein R' is an optionally substituted heteroaryl radical as defined herein, and R" is an alkylene radical as defined herein with the understanding that the attachment point of the heteroaryl radical will be on the alkylene radical. Examples of heteroarylalky radicals include, but are not limited to, 2-imidazolylmethyl, 3-pyrrolylethyl.

The term "heterocyclyl" or "heterocycle" as used herein denotes a monovalent saturated cyclic radical, consisting of one or more rings, preferably one to two rings, of three to eight atoms per ring, incorporating one or more ring heteroatoms (chosen from N,O or S(O)0-2), and which can optionally be independently substituted with one or more, preferably one or two substituents selected from hydroxy, oxo, cyano, lower alkyl, lower alkoxy, lower haloalkoxy, alkylthio, halo, haloalkyl, hydroxyalkyl, nitro, alkoxycarbonyl, amino, alkylamino, alkylsulfonyl, arylsulfonyl, alkylaminosulfonyl, arylaminosulfonyl, alkylsulfonylamino, arylsulfonylamino, alkylaminocarbonyl, arylaminocarbonyl, alkylcarbonylamino, arylcarbonylamino, unless otherwise indicated. A bicyclic heterocycle can be fused to an aryl or heteroaryl ring; however, the point of attachment is on the heterocyclic ring. Examples of heterocyclic radicals include, but are not limited to, azetidinyl, pyrrolidinyl, hexahydroazepinyl, oxetanyl, tetrahydrofuranyl, tetrahydrothiophenyl, oxazolidinyl, thiazolidinyl, isoxazolidinyl, morpholinyl, piperazinyl, piperidinyl, tetrahydropyranyl, thiomorpholinyl, quinuclidinyl and imidazolinyl.

The term "heterocycloalkyl" (or "heterocyclylalkyl") means the radical of the formula R'R", wherein R' is a heterocyclic radical as defined herein, and R" is an alkylene radical as defined herein and the attachment point of the heterocycloalkyl radical will be on the alkylene radical.. Examples of heterocycloalkyl radicals include, but are not limited to, 1-piperazinylmethyl, 2-morpholinomethyl, and the like.

The term "cycloalkyl" as used herein denotes a saturated carbocyclic ring containing 3 to 8 carbon atoms, i.e. cyclopropyl, cyclobutyl, cyclopentyl, cyclohexyl, cycloheptyl or cyclooctyl. "C_{3-7} cycloalkyl" as used herein refers to an cycloalkyl composed of 3 to 7 carbons in the carbocyclic ring.

The term "cycloalkylalkyl" as used herein refers to the radical R'R"-, wherein R' is a cycloalkyl radical as defined herein, and R" is an alkylene radical as defined herein with the understanding that the attachment point of the cycloalkylalkyl moiety will be on the alkylene radical. Examples of cycloalkylalkyl radicals include, but are not limited to, cyclopropylmethyl, cyclohexylmethyl, cyclopentylethyl. C_{3-7} cycloalkyl-C_{1-3} alkyl refers to the radical R'R" where R' is C_{3-7} cyclolalkyl and R" is C_{1-3} alkylene as defined herein.

The term "cycloalkyl" as used herein denotes a saturated carbocyclic ring containing 3 to 8 carbon atoms, i.e. cyclopropyl, cyclobutyl, cyclopentyl, cyclohexyl, cycloheptyl or cyclooctyl. "C_{3-7} cycloalkyl" as used herein refers to an cycloalkyl composed of 3 to 7 carbons in the carbocyclic ring.

The term "cycloalkylalkyl" as used herein refers to the radical R'R"-, wherein R' is a cycloalkyl radical as defined herein, and R" is an alkylene radical as defined herein with the understanding that the attachment point of the cycloalkylalkyl moiety will be on the alkylene radical. Examples of cycloalkylalkyl radicals include, but are not limited to, cyclopropylmethyl, cyclohexylmethyl, cyclopentylethyl. C_{3-7} cycloalkyl-C_{1-3} alkyl refers to the radical R'R" where R' is C_{3-7} cyclolalkyl and R" is C_{1-3} alkylene as defined herein.

The terms "hydroxyalkyl" and "alkoxyalkyl" as used herein denotes the radical R'R" where R' is an hydroxy radical or a alkoxy radical respectively and R" is alkylene as defined herein and the attachment point of the hydroxyalkyl or alkoxyalkyl radical will be on the alkylene radical.

The term "alkylene" as used herein denotes a divalent saturated linear hydrocarbon radical of 1 to 8 carbon atoms or a branched saturated divalent hydrocarbon radical of 3 to 8 carbon atoms, unless otherwise indicated. Examples of alkylene radicals include, but are not limited to, methylene, ethylene, propylene, 2-methyl-propylene, butylene, 2-ethylbutylene.

The terms "alkylsulfinyl" and "arylsulfinyl"as used herein denotes a group of formula -S(=O)R wherein R is alkyl or aryl respectively and alkyl and aryl are as defined herein

The terms "alkylsulfonyl" and "arylsulfonyl"as used herein denotes a group of formula $-S(=O)_2R$ wherein R is alkyl or aryl respectively and alkyl and aryl are as defined herein.

The term "sulfamoyl" as used herein refers to the radical $-S(O)_2NH_2$. The terms "N-alkylsulfamoyl" and "N, N-dialkylsulfamoyl" as used herein refer to the radical $-S(O)_2NR'R"$, wherein R' and R" are hydrogen and lower alkyl and R' and R" are independently lower alkyl respectively. Examples of N-alkylsulfamoyl substituents include, but are not limited to methylaminosulfonyl, *iso*-propylaminosulfonyl. Examples of N,N-dialkylsulfamoyl substituents include, but are not limited to dimethylaminosulfonyl, *iso*-propyl-methylaminosulfonyl. The prefix N-alkyl or N,N-dialkyl can be replaced with aryl, heteroaryl, heterocyclyl or other radical to indicate a case where the amine is substituted with a group other than alkyl.

The term "alkylsulfonamido" refers to the radical $-NH-S(O)_2$-alkyl. The term alkyl can be replaced by other chemically relevant radicals such as aryl or heteroaryl to indicate, e.g. phenylsulfonamido $-NH-S(O)_2$-Ph. "N-alkylalkylsulfonamido" refers to the radical $-NR-S(O)_2$-alkyl where R is a lower alkyl group.

The term "ureido" as used herein means an -N^1RC(=O)N^3R'R" radical where R, R' and R" are independently hydrogen or lower alkyl. If the nitrogen atoms are substituted by other group other than hydrogen or lower alkyl, locants N or N' or 1 and 3 respectively are used to identify the substituted nitrogen atoms. The point of attachment of the ureido radical is denoted N or 1. For example, using this nomenclature N'-phenylureido refers to-NRC(=O)NPhR' where R and R' are as defined previously. The position of specific alkyl substituents can optionally be specifically designated using the same nomenclature.

The term "carbamate" or "urethane" as used herein refers to a ROC(=O)N^3R'R" radical wherein either R or R' is the core structure and the other of R, R' and R" are as defined in the specification and claims. The term "urea" as used herein refers to a group RR'NC(=O)NR"R'" wherein R is the core structure and R', R" and R'" are as defined in the specification and claims.

The term "aminocarbonylpyridyl" as used herein refers to the radical -NHCOR wherein R is 2-pyridinyl (picolinoyl), 3-pyridinyl (nicotinoyl) or 4-pyridinyl (isonicotinoyl) and the N-oxides derived therefrom.

Compounds of formula I exhibit tautomerism. Tautomeric compounds can exist as two or more interconvertable species. Prototropic tautomers result from the migration of a covalently bonded hydrogen atom between two atoms. Tautomers generally exist in equilibrium and attempts to isolate an individual tautomers usually produce a mixture whose chemical and physical properties are consistent with a mixture of compounds. The position of the equilibrium is dependent on chemical features within the molecule. For example, in many aliphatic aldehydes and ketones, such as acetaldehyde, the keto form predominates while; in phenols, the enol form predominates. Common prototropic tautomers include keto/enol (-C(=O)-CH- □ -C(-OH)=CH-), amide/imidic acid (-C(=O)-NH- □ -C(-OH)=N-) and amidine (-C(=NR)-NH- □ -C(-NHR)=N-) tautomers. The latter two are particularly common in heteroaryl and heterocyclic rings and the present invention encompasses all tautomeric forms of the compounds.

It will be appreciated by the skilled artisan that the compounds of formula I may contain one or more chiral centers and therefore exist in two or more stereoisomeric forms. The racemates of these isomers, the individual isomers and mixtures enriched in one enantiomer, as well as diastereomers when there are two chiral centers, and mixtures partially enriched with specific diastereomers are within the scope of the present invention. It will be further appreciated by the skilled artisan that substitution of the tropane ring can be in either *endo*- or *exo*-configuration, and the present invention covers both configurations. The present invention includes all the individual stereoisomers (e.g.

enantiomers), racemic mixtures or partially resolved mixtures of the compounds of formulae I and, where appropriate, the individual tautomeric forms thereof.

The racemates can be used as such or can be resolved into their individual isomers. The resolution can afford stereochemically pure compounds or mixtures enriched in one or more isomers. Methods for separation of isomers are well known (cf. Allinger N. L. and Eliel E. L. in "Topics in Stereochemistry", Vol. 6, Wiley Interscience, 1971) and include physical methods such as chromatography using a chiral adsorbent. Individual isomers can be prepared in chiral form from chiral precursors. Alternatively individual isomers can be separated chemically from a mixture by forming diasteromeric salts with a chiral acid, such as the individual enantiomers of 10-camphorsulfonic acid, camphoric acid, .alpha.-bromocamphoric acid, tartaric acid, diacetyltartaric acid, malic acid, pyrrolidone-5-carboxylic acid, and the like, fractionally crystallizing the salts, and then freeing one or both of the resolved bases, optionally repeating the process, so as obtain either or both substantially free of the other; i.e., in a form having an optical purity of >95%. Alternatively the racemates can be covalently linked to a chiral compound (auxillary) to produce diastereomers which can be separated by chromatography or by fractional crystallization after which time the chiral auxilliary is chemically removed to afford the pure enantiomers.

The compounds of formula I contain at least one basic center and suitable acid addition salts are formed from acids which form non-toxic salts. Examples of salts of inorganic acids include the hydrochloride, hydrobromide, hydroiodide, chloride, bromide, iodide, sulphate, bisulphate, nitrate, phosphate, hydrogen phosphate. Examples of salts of organic acids include acetate, fumarate, pamoate, aspartate, besylate, carbonate, bicarbonate, camsylate, D and L-lactate, D and L-tartrate, esylate, mesylate, malonate, orotate, gluceptate, methylsulphate, stearate, glucuronate, 2-napsylate, tosylate, hibenzate, nicotinate, isethionate, malate, maleate, citrate, gluconate, succinate, saccharate, benzoate, esylate, and pamoate salts. For a review on suitable salts see Berge *et al, J. Pharm. Sci.*, 66, 1-19, 1977.

The term "solvate" as used herein means a compound of the invention or a salt, thereof, that further includes a stoichiometric or non-stoichiometric amount of a solvent bound by non-covalent intermolecular forces. Preferred solvents are volatile, non-toxic, and/or acceptable for administration to humans in trace amounts.

The term "hydrate" as used herein means a compound of the invention or a salt thereof, that further includes a stoichiometric or non-stoichiometric amount of water bound by non-covalent intermolecular forces.

The term "clathrate" as used herein means a compound of the invention or a salt thereof in the form of a crystal lattice that contains spaces (e. g., channels) that have a guest molecule (e. g., a solvent or water) trapped within.

The term "nucleoside and nucleotide reverse transcriptase inhibitors" ("NRTI"s) as used herein means nucleosides and nucleotides and analogues thereof that inhibit the activity of HIV-1 reverse transcriptase, the enzyme which catalyzes the conversion of viral genomic HIV-1 RNA into proviral HIV-1 DNA. Typical suitable NRTIs include zidovudine (AZT) available as RETROVIR® from Glaxo-Wellcome Inc.; didanosine (ddI) available as VIDEX® from Bristol-Myers Squibb Co.; zalcitabine (ddC) available as HIVID® from Roche Pharmaceuticals; stavudine (d4T) available as ZERIT® from Bristol-Myers Squibb Co.; lamivudine (3TC) available as EPIVIR® from Glaxo-Wellcome; abacavir (1592U89) disclosed in WO96/30025 and available ZIAGEN® from Glaxo-Wellcome; adefovir dipivoxil [bis(POM)-PMEA] available as PREVON® from Gilead Sciences; lobucavir (BMS-180194), a nucleoside reverse transcriptase inhibitor disclosed in EP-0358154 and EP-0736533 and under development by Bristol-Myers Squibb; BCH-10652, a reverse transcriptase inhibitor (in the form of a racemic mixture of BCH-10618 and BCH-10619) under development by Biochem Pharma; emitricitabine [(-)-FTC] licensed from Emory University under U.S. Pat. No. 5,814,639 and under development by Triangle Pharmaceuticals; beta-L-FD4 (also called beta-L-D4C and named beta-L-2', 3'-dicleoxy-5-fluoro-cytidene) licensed by Yale University to Vion Pharmaceuticals; DAPD, the purine nucleoside, (-)-b-D-2,6-diamino-purine dioxolane disclosed in EP-0656778 and licensed by Emory University and the University of Georgia to Triangle Pharmaceuticals; and lodenosine (FddA), 9-(2,3-dideoxy-2-fluoro-b-D-threo-pentofuranosyl)adenine, an acid stable purine-based reverse transcriptase inhibitor discovered by the NIH and under development by U.S. Bioscience Inc.

The term "non-nucleoside reverse transcriptase inhibitors" ("NNRTI"s) as used herein means non-nucleosides that inhibit the activity of HIV-1 reverse transcriptase. Typical suitable NNRTIs include nevirapine (BI-RG-587) available as VIRAMUNE® from Roxane Laboratories; delaviradine (BHAP, U-90152) available as RESCRIPTOR® from Pfizer; efavirenz (DMP-266) a benzoxazin-2-one disclosed in WO94/03440 and available as SUSTIVA® from Bristol-Myers Squibb Co.; PNU-142721, a furopyridine-thio-pyrimide under development by Pfizer 08807; AG-1549 (formerly Shionogi # S-1153); 5-(3,5-dichlorophenyl)-thio-4-isopropyl-1-(4-pyridyl)methyl-1H-imidazol-2-ylmethyl carbonate disclosed in WO 96/10019 and under development by Agouron Pharmaceuticals, Inc.; MKC-442 (1-(ethoxy-methyl)-5-(1-methylethyl)-6-(phenylmethyl)-(2,4(1H,3H)-pyrimidinedione) discovered by Mitsubishi Chemical Co. and under development by Triangle Pharmaceuticals; and (+)-calanolide A (NSC-

675451) and B, coumarin derivatives disclosed in NIH U.S. Pat. No. 5,489,697, licensed to Med Chem Research, which is co-developing (+) calanolide A with Vita-invest as an orally administrable product.

The term "protease inhibitor" ("PI") as used herein means inhibitors of the HIV-1 protease, an enzyme required for the proteolytic cleavage of viral polyprotein precursors (e.g., viral GAG and GAG Pol polyproteins), into the individual functional proteins found in infectious HIV-1. HIV protease inhibitors include compounds having a peptidomimetic structure, high molecular weight (7600 daltons) and substantial peptide character. Typical suitable PIs include saquinavir (Ro 31-8959) available in hard gel capsules as INVIRASE® and as soft gel capsules as FORTOVASE® from Roche Pharmaceuticals, Nutley, N.J. 07110-1199; ritonavir (ABT-538) available as NORVIR® from Abbott Laboratories; indinavir (MK-639) available as CRIXIVAN® from Merck & Co., Inc.; nelfnavir (AG-1343) available VIRACEPT® from Agouron Pharmaceuticals, Inc.; amprenavir (141W94), AGENERASE®, a non-peptide protease inhibitor under development by Vertex Pharmaceuticals, Inc. and available from Glaxo-Wellcome, under an expanded access program; lasinavir (BMS-234475) available from Bristol-Myers Squibb; DMP-450, a cyclic urea discovered by Dupont and under development by Triangle Pharmaceuticals; BMS-2322623, an azapeptide under development by Bristol-Myers Squibb as a 2nd-generation HIV-1 PI; ABT-378 under development by Abbott; and AG-1549 an orally active imidazole carbamate discovered by Shionogi and under development by Agouron Pharmaceuticals, Inc.

Other antiviral agents include hydroxyurea, ribavirin, IL-2, IL-12, pentafuside. Hydroyurea (Droxia), a ribonucleoside triphosphate reductase inhibitor, the enzyme involved in the activation of T-cells, was discovered at the NCI and is in preclinical studies, it was shown to have a synergistic effect on the activity of didanosine and has been studied with stavudine. IL-2 is disclosed in Ajinomoto EP-0142268, Takeda EP-0176299, and Chiron U.S. Pat. Nos. RE 33,653, 4,530,787, 4,569,790, 4,604,377, 4,748,234, 4,752,585, and 4,949,314, and is available under the PROLEUKIN® (aldesleukin) as a lyophilized powder for IV infusion or sc administration upon reconstitution and dilution with water; a dose of about 1 to about 20 million 1U/day, sc is preferred; a dose of about 15 million 1 U/day, sc is more preferred. IL-12 is disclosed in WO96/25171 and is administered in a dose of about 0.5 microgram/kg/day to about 10 microgram/kg/day, sc is preferred. Pentafuside (FUZEON®) a 36-amino acid synthetic peptide, disclosed in U.S. Pat. No. 5,464,933 that acts by inhibiting fusion of HIV-1 to target membranes. Pentafuside (3-100 mg/day) is given as a continuous sc infusion or injection together with efavirenz and 2 PI's to HIV-1 positive patients refractory to a triple combination therapy; use of 100 mg/day is preferred. Ribavirin, 1-.beta.-D-

ribofuranosyl-1H-1,2,4-triazole-3-carboxamide, is available from ICN Pharmaceuticals, Inc., Costa Mesa, Calif.; its manufacture and formulation are described in U.S. Pat. No. 4,211,771.

The term "viral fusion inhibitors" as used herein refers to compounds which inhibit fusion of the free virus particle and introduction of the viral RNA into a host cell independent of the molecular locus of inhibitor binding. Viral fusion inhibitors therefore include, but are not limited to T-20; CD-4 binding ligands including BMS-378806, BMS-488043; CCR5 binding ligands including SCH-351125, Sch-350634, Sch-417690 (Schering Plough), UK-4278957 (Pfizer), TAK-779 (Takeda), ONO-4128 (Ono), AK-602 (Ono, GlaxoSmithKline), compounds 1-3 (Merck); CXCR4 binding ligands KRH-1636 (K. Ichiyama *et al. Proc. Nat. Acad. Sci USA* 2003 100(7):4185-4190), T-22 (T. Murakami *et al. J. Virol.* 1999 73(9):7489-7496), T-134 (R. Arakaki *et al. J. Virol.* 1999 73(2):1719-1723). Viral fusion inhibitors as used herein also include peptide and protein soluble receptors, antibodies, chimeric antibodies, humanized antibodies.

Abbreviations used in this application include: acetyl (Ac), acetic acid (HOAc), azo-*bis*-isobutyrylnitrile (AIBN), 1-N-hydroxybenzotriazole (HOBT), atmospheres (Atm), high pressure liquid chromatography (HPLC), 9-borabicyclo[3.3.1]nonane (9-BBN or BBN), methyl (Me), *tert*-butoxycarbonyl (Boc), acetonitrile (MeCN), di-*tert*-butyl pyrocarbonate or boc anhydride (BOC_2O), 1-(3-dimethylaminopropyl)-3-ethylcarbodiimide hydrochloride (EDCI), benzyl (Bn), *m*-chloroperbenzoic acid (MCPBA), butyl (Bu), methanol (MeOH), benzyloxycarbonyl (cbz or Z), melting point (mp), carbonyl diimidazole (CDI), $MeSO_2$- (mesyl or Ms), 1,4-diazabicyclo[2.2.2]octane (DABCO), mass spectrum (ms) diethylaminosulfur trifluoride (DAST), methyl *t*-butyl ether (MTBE), dibenzylideneacetone (Dba), N-carboxyanhydride (NCA), 1,5-diazabicyclo[4.3.0]non-5-ene (DBN), N-bromosuccinimide (NBS), 1,8-diazabicyclo[5.4.0]undec-7-ene (DBU), N-methylpyrrolidone (NMP), 1,2-dichloroethane (DCE), pyridinium chlorochromate (PCC), N,N'-dicyclohexylcarbodiimide (DCC), pyridinium dichromate (PDC), dichloromethane (DCM), propyl (Pr), diethyl azodicarboxylate (DEAD), phenyl (Ph), di-*iso*-propylazodicarboxylate , DIAD, pounds per square inch (psi), di-iso-propyl-ethylamine (DIPEA), pyridine (pyr), di-*iso*-butylaluminumhydride , DIBAL-H, room temperature, rt or RT, N,N-dimethyl acetamide (DMA), *tert*-butyldimethylsilyl or *t*-$BuMe_2Si$, (TBDMS), 4-N,N-dimethylaminopyridine (DMAP), triethylamine (Et_3N or TEA), N,N-dimethylformamide (DMF), triflate or CF_3SO_2- (Tf), dimethyl sulfoxide (DMSO), trifluoroacetic acid (TFA), 1,1'-*bis*-(diphenylphosphino)ethane (dppe), 2,2,6,6-tetramethylheptane-2,6-dione (TMHD), 1,1'-*bis*-(diphenylphosphino)ferrocene (dppf), thin layer chromatography (TLC), ethyl acetate (EtOAc), tetrahydrofuran (THF), diethyl

ether (Et_2O), trimethylsilyl or Me_3Si (TMS), ethyl (Et), *p*-toluenesulfonic acid monohydrate (TsOH or pTsOH), lithium hexamethyl disilazane (LiHMDS), 4-Me-$C_6H_4SO_2$- or tosyl (Ts), *iso*-propyl (*i*-Pr), N-urethane-N-carboxyanhydride (UNCA), ethanol (EtOH). Conventional nomenclature including the prefixes *normal* (*n*), *iso* (*i*-), *secondary* (*sec*-), *tertiary* (*tert*-) and *neo* have their customary meaning when used with an alkyl moiety. (J. Rigaudy and D. P. Klesney, *Nomenclature in Organic Chemistry,* IUPAC 1979 Pergamon Press, Oxford.).

Compounds of the present invention can be made by a variety of methods depicted in the illustrative synthetic reaction schemes shown and described below. The starting materials and reagents used in preparing these compounds generally are either available from commercial suppliers, such as Aldrich Chemical Co., or are prepared by methods known to those skilled in the art following procedures set forth in references such as *Fieser and Fieser's Reagents for Organic Synthesis*; Wiley & Sons: New York, Volumes 1-21; R. C. LaRock, *Comprehensive Organic Transformations,* 2nd edition Wiley-VCH, New York 1999; *Comprehensive Organic Synthesis,* B. Trost and I. Fleming (Eds.) vol. 1-9 Pergamon, Oxford, 1991; *Comprehensive Heterocyclic Chemistry*, A. R. Katritzky and C. W. Rees (Eds) Pergamon, Oxford 1984, vol. 1-9; *Comprehensive Heterocyclic Chemistry II*, A. R. Katritzky and C. W. Rees (Eds) Pergamon, Oxford 1996, vol. 1-11; and *Organic Reactions*, Wiley & Sons: New York, 1991, Volumes 1-40. The following synthetic reaction schemes are merely illustrative of some methods by which the compounds of the present invention can be synthesized, and various modifications to these synthetic reaction schemes can be made and will be suggested to one skilled in the art having referred to the disclosure contained in this Application.

The starting materials and the intermediates of the synthetic reaction schemes can be isolated and purified if desired using conventional techniques, including but not limited to, filtration, distillation, crystallization, chromatography, and the like. Such materials can be characterized using conventional means, including physical constants and spectral data.

Unless specified to the contrary, the reactions described herein preferably are conducted under an inert atmosphere at atmospheric pressure at a reaction temperature range of from about -78 °C to about 150 °C, more preferably from about 0° C to about 125° C, and most preferably and conveniently at about room (or ambient) temperature, e.g., about 20 °C.

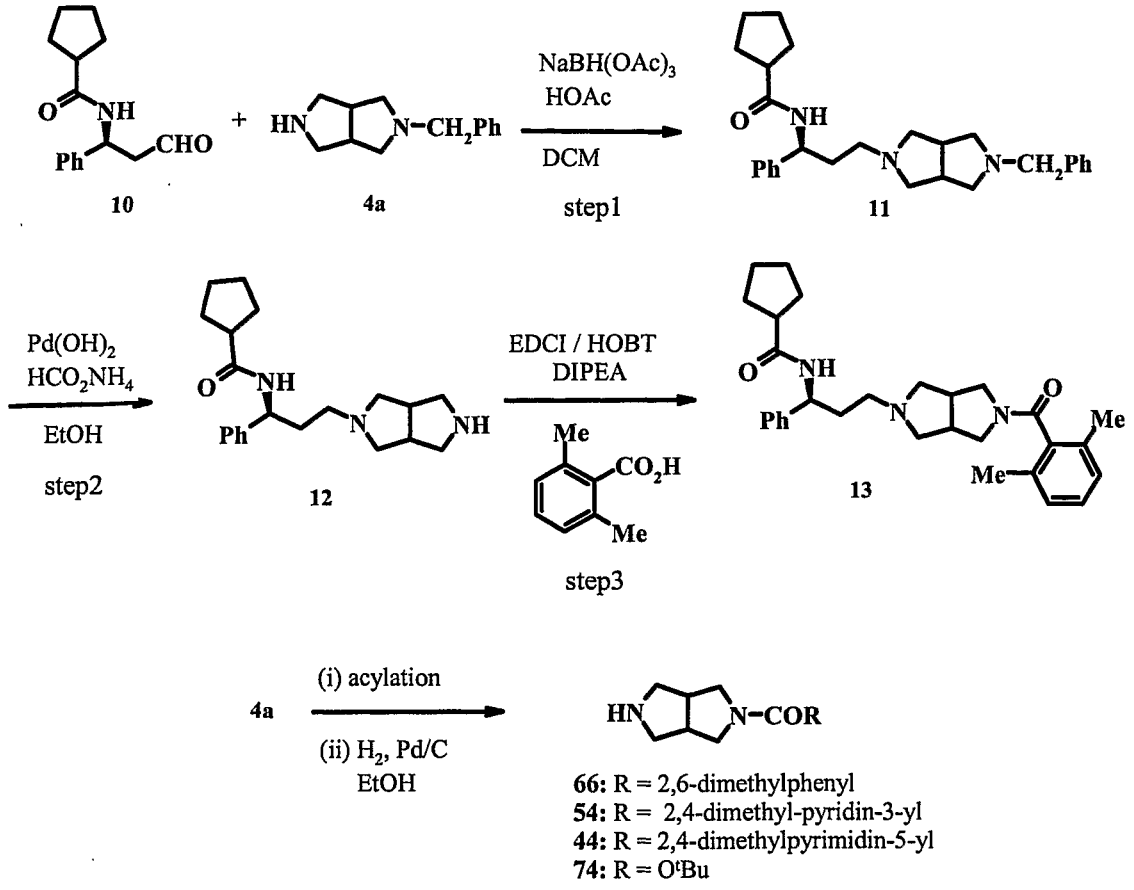
2-Benzyl-octahydro-pyrrolo[3,4-c]pyrrole (4a) was prepared by [2,3]-dipolar cycloaddition of an imine ylide with N-benzylmaleimide as described previously (R.

Colon-Cruz *et al.* WO 02/070523 and M. Björsne *et al.* WO 02/060902). Reduction of the imide, and selective debenzylation are accomplished as described therein.

The preparation of some compounds of the present invention wherein R^2 is phenyl is depicted in Scheme 1. The procedure in Scheme 1 is particularly suited for preparation of a series of compounds in which the amides (or ureas or sulfonamides) linked to the hexahydro-pyrrolo[3,4-c]pyrrol-2-yl] scaffold is varied. The fully elaborated 1-phenyl-1-amino-propyl side chain is introduced before deprotection of the second amine. Debenzylation of the amine allows for the elaboration of the second nitrogen atom.

Alternatively, the amide (or urea or sulfonamide) my be introduced first by acylation of 4a with *e.g.* 2,6-dimethylbenzoyl chloride, 2,4-dimethyl-nicotinoyl chloride or 4,6-dimethyl-pyrimidin-5-carbonyl chloride, and subsequent debenzylation to afford **66**, **54** and **44** respectively. Introduction of a Boc group and subsequent debenzylation affords **74**. The accompanying examples illustrate the utility of these compounds for the preparation of compounds of the invention.

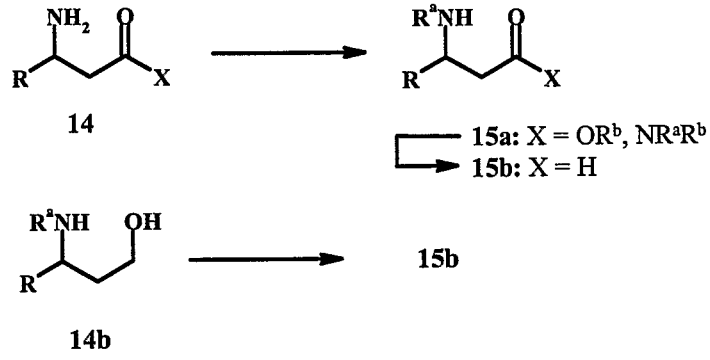
Scheme 1



β-Aminoacylcarboxaldeydes **15b** are convenient synthetic intermediates accessible by reduction of β-acylamino carboxylic acids or carboxylic acid derivatives (15a: X = OH,

OR^b or NR^bR^c wherein R^a is an acyl radical or a protecting group and R^b and R^c are typically lower alkyl), see, *e.g.* Example 15 Acylation of a □-amino ester (14: X is O-alkyl) is conveniently carried out with a corresponding acyl halide, carboxylic acid anhydride or chloroformate in a solvent such as DCM, chloroform, carbon tetrachloride, ether, THF, dioxane, benzene, toluene, MeCN, DMF, sodium hydroxide solution or sulpholane optionally in the presence of an inorganic or organic base at temperatures between -20 and 200° C, but preferably at temperatures between -10 and 100° C.

Scheme 2



Acylation also may be carried out with the carboxylic acid in the presence of an activating agent or a dehydrating agent, *e.g.* in the presence of isobutyl chloroformate, thionyl chloride, trimethylchlorosilane, HCl, H_2SO_4, methanesulphonic acid, p-toluenesulphonic acid, phosphorus trichloride, P_2O_5, N,N'-dicyclohexylcarbodiimide, N,N'-dicyclohexyl-carbodiimide/N-hydroxysuccinimide or HOBt, N,N'-carbonyldiimidazole, O-(benzotriazol-1-yl)-N,N,N',N'-tetramethyl-uronium tetrafluoroborate/N-methylmorpholine, O-(benzotriazol-1-yl)-N,N,N',N'-tetramethyl-uronium tetrafluoroborate/N-ethyldiisopropylamine, N,N'-thionyldiimidazole or triphenylphosphine/CCl_4, at temperatures between -20 and 200° C., but preferably at temperatures between -10 and 100° C.

Aldehydes 15b may be prepared by reduction of the corresponding acid (15a: X = OH), ester (15a: X = OR^b) or amide (15a: X = NR^bR^c), wherein R^b and R^c are lower alkyl, by a hydride reducing agent in a suitable solvent. Alternatively, reduction of an acyl halide may be achieved with a suitable transition metal catalyst, a hydrogen source and in a suitable solvent. Typical hydride reducing agents are aluminum hydrides or boron hydrides such as DIBAL-H, $LiAl(O\text{-}tert\text{-}Bu)_3H$, or $Me_2CHCH(Me)_2BH$. Suitable solvents are inert solvents such as THF, DCM or toluene. An acid chloride (15a: X = Cl) can be reduced with a transition metal catalyst such as Pd/C or $Pd/BaSO_4$, under a hydrogen atmosphere with a modifier such as 2,4-dimethylpyridine and in solvent such as THF or toluene. Another route to □-acylamino-carboxyaldehydes 15b comprises oxidation of a □-acylaminoalcohol 14b which can be carried out by a variety of oxidizing agents, e.g.

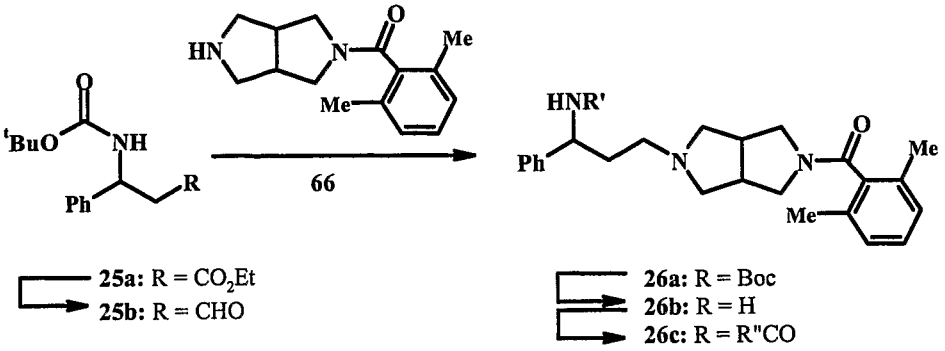
pyridine.SO_3.TEA. Preferably the N-acylated acid or ester is reduced to the aldehydes with DIBAL-H in DCM at -78° C as described by D. R. Armour *et al.* (WO 00/38680).

The aldehyde **15b** (R^a is an acyl radical or a nitrogen protecting group) is incorporated onto the 2-octahydro-pyrrolo[3,4-c]pyrrole scaffold by reductive amination to afford diamine **11**. Reductive amination is preferably carried out by combining an amine and aldehyde in the presence of a complex metal hydride such as sodium borohydride, lithium borohydride, sodium cyanoborohydride, zinc borohydride, sodium triacetoxyborohydride or borane/pyridine conveniently at a pH of 1-7. The reaction mixture optionally includes a dehydrating agent such as molecular sieves or Ti(IV)(O-i-Pr)$_4$ to facilitate formation of the intermediate imine at ambient temperature or with hydrogen in the presence of a hydrogenation catalyst, e.g. in the presence of Pd/C, at a hydrogen pressure of 1 to 5 bar, preferably at temperatures between 20° C. and the boiling temperature of the solvent used. It may also be advantageous during the reaction if reactive groups are protected during the reaction by conventional protecting groups which are cleaved again by conventional methods after the reaction. Reductive amination procedures have been reviewed: R. M. Hutchings and M. K. Hutchings *Reduction of C=N to CHNH by Metal Hydrides* in *Comprehensive Organic Synthesis* col. 8, I. Fleming (Ed) Pergamon, Oxford 1991 pp. 47-54.

Removal of the remaining benzyl protecting group affords amine **12**. Removal of benzyl protecting groups can be readily achieved by catalytic hydrogenolysis using Pd, Pt, Ni or Rh catalysts. Acids are sometimes added to promote hydrogenolysis. Acylation of the free secondary amine affords **13**. Alternatively the nitrogen atom can be alkylated with an aralkyl halide to afford I (X^2 is aralkyl) or sulfonylated to produce sulfonamides I (X^2 is SO_2R^4). Sulfonylation of amines is typically carried out by treating an amine with an alkyl or aryl sulfonyl chloride in the presence of an organic base, *e.g.* pyridine or TEA, in an inert solvent.

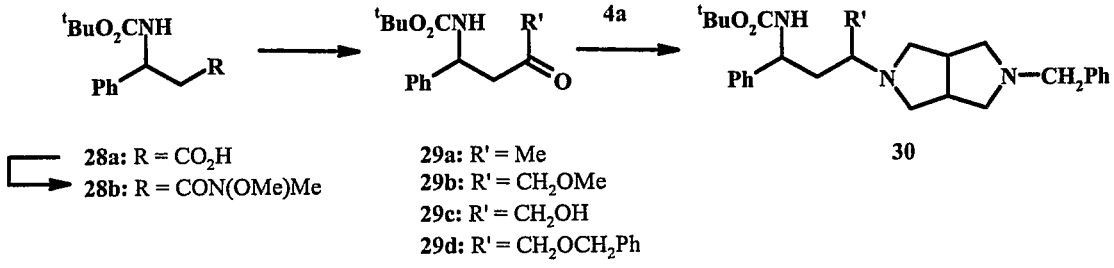
While the preparation of **13** exemplifies the synthesis of a compound of the invention with a cyclopentylcarboxamide, one skilled in the art will appreciate a wide variety of amides, ureas or sulfonamides can be introduced by using an analogous.

Scheme 3



An amino protecting group, such as the Boc derivative **25b**, may be utilized in place of an acyl radical during the reductive amination procedures (Scheme3). Numerous reactions for the formation and removal of such amine protecting groups are described in standard works including, for example, "Protective Groups in Organic Chemistry", Plenum Press, London and New York, 1973; Greene, Th. and Wutts, P. G. M. *Protective Groups in Organic Synthesis*, Wiley, New York, 1999; *The Peptides*, Vol. I, Schroder and Lubke, Academic Press, London and New York, 1965; *Methoden der organischen Chemie, Houben-Weyl*, 4th Edition, Vol 15/I, Georg Thieme Verlag, Stuttgart 1974, the disclosures of which are incorporated herein by reference in their entirety. Protecting groups may be selected which are cleaved under a variety of conditions selected to be compatible with other functional groups in the molecule. Introduction of acyl groups is then accomplished after the reductive amination step by deprotection of the primary amine and acylation with a carboxylic acid as described above. Ureas, carbamates and sulfonamides are accessible by reaction of **26b** with isocyanates, chloroformates and sulfonyl chlorides respectively. Scheme 3 depicts the introduction of a carboxamido group onto the diazabicyclooctane ring prior to the reductive alkylation and elaboration of the aminopropyl side chain after reductive alkylation.

Scheme 4



Embodiments of the invention may have alkyl (**30**: R' = Me), alkoxyalkyl (**30**: R' = CH_2OMe or CH_2OCH_2Ph) or hydroxymethyl (**30**: R' = CH_2OH) substituents on the propylene linker. One route to these compounds is depicted in Scheme 4. The □-acylaminocarboxylic acid was converted to the N-methoxy-N-methyl amide and reacted with methyl lithium to afford butanone **29a** which could be incorporated onto the

pyrrolo[3,4-c]pyrrol-2-yl scaffold by reductive alkylation as described previously. Methoxymethyl- and benzyloxymethyl compounds were prepared by analogous methods starting from stannane organometallics as described in Examples 27 and 28. Debenzylation of 29d affords the corresponding hydroxylmethyl compound 29c, Compounds bearing a methyl or a dimethyl substituent on the linker carbon adjacent to aminomethine carbon were prepared as described in Examples 24 and 26. An example of the preparation of compounds with a hydroxyl substituent on the propylene linker is in Example 35

Examples of representative compounds encompassed by the present invention and within the scope of the claims are provided in the following Tables. These examples and preparations are provided to enable those skilled in the art to more clearly understand and to practice the present invention. They should not be considered as limiting the scope of the invention, but merely as being illustrative and representative thereof. The reagents used to introduce X^1 and X^2 are commercially available or can prepared from commercially available materials by published procedures. An examination of the compounds encompassed in the invention and working examples demonstrate that the generality of the reaction sequence and the nature of the protecting groups which can be optimized for a particular target without deviating from the general procedures discussed herein.

In general, the nomenclature used in this Application is based on AUTONOM™ v.4.0, a Beilstein Institute computerized system for the generation of IUPAC systematic nomenclature. If there is a discrepancy between a depicted structure and a name given that structure, the depicted structure is to be accorded more weight. In addition, if the stereochemistry of a structure or a portion of a structure is not indicated with, for example, bold or dashed lines, the structure or portion of the structure encompasses both isomers. While compounds of the present invention are frequently depicted with the (S) stereochemistry, both stereoisomers are included in the present invention and both can be prepared by identical procedures from the appropriate starting material.

Representative compounds of the present invention in which R^2 is an aryl or heteroaryl group are compiled in Table 1.

Table 1

	Name	MS	Mp
I-1	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoxaline-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	500.3	
I-2	Cyclopentanecarboxylic acid {(S)-3-[5-(3,4-dichloro-benzyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-	500	

	propyl}-amide; compound with trifluoro-acetic acid		
I-3	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	474	60.9-63.9
I-4	Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	447	
I-5	Cyclopentanecarboxylic acid [(S)-1-phenyl-3-(5-phenylacetyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-propyl]-amide; compound with trifluoro-acetic acid	460	
I-6	5-[(S)-3-(Cyclopentanecarbonyl-amino)-3-phenyl-propyl]-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carboxylic acid benzyl ester	476	
I-7	Cyclopentanecarboxylic acid {(R)-2-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-ethyl}-amide; compound with trifluoro-acetic acid	460	
I-8	Cyclopentanecarboxylic acid {(S)-3-[5-(2,5-dimethyl-2*H*-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	464	70.1-72.5
I-9	Cyclopentanecarboxylic acid {(S)-3-[5-(1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	463	62.0-64.3
I-10	Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(1*H*-[1,2,4]triazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	437	107.0-108.0
I-11	Cyclopentanecarboxylic acid {(S)-3-[5-(4-methoxy-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	504.8	
I-12	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dichloro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	514.7	
I-13	Cyclopentanecarboxylic acid {(S)-3-[5-(2-chloro-6-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	494.8	
I-14	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dichloro-4-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	528.7	
I-15	Cyclopentanecarboxylic acid {(S)-3-[5-(4-butoxy-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	546.9	
I-16	Cyclopentanecarboxylic acid {(S)-3-[5-(4-ethoxy-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	518.8	
I-17	Cyclopentanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with hydrochloric acid	498.7	
I-18	Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	488.8	

I-19	Cyclopentanecarboxylic acid {(S)-3-[5-(2-bromo-6-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	540.7	
I-20	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-difluoro-4-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	512.8	
I-21	Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	536.8	
I-22	Cyclopentanecarboxylic acid {(S)-3-[5-(4-chloro-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	510.8	
I-23	Cyclopentanecarboxylic acid {(S)-3-[5-(2,3-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	474.8	
I-24	Cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	474.8	
I-25	Cyclopentanecarboxylic acid {(S)-3-[5-(2-methoxy-4-methylsulfanyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	522.8	
I-26	Cyclopentanecarboxylic acid {(S)-3-[5-(2-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	489.8	
I-27	Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(1*H*-pyrrole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	435.8	
I-28	Cyclopentanecarboxylic acid {(S)-3-[5-(3,5-dimethyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	465.8	
I-29	Cyclopentanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	476	48.0-49.0
I-30	Cyclopentanecarboxylic acid [(S)-3-(5-acetyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	384	
I-31	Cyclopentanecarboxylic acid {(S)-3-[5-(2,2-dimethyl-propionyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	426	
I-32	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-benzenesulfonamide; compound with trifluoro-acetic acid	518.8	
I-33	Cyclopropanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	446.8	
I-34	Furan-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-	472.8	

	amide; compound with trifluoro-acetic acid		
I-35	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-isobutyramide; compound with trifluoro-acetic acid	448.8	
I-36	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-methyl-butyramide; compound with trifluoro-acetic acid	462.8	
I-37	Thiophene-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	488.8	
I-38	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2,2-dimethyl-propionamide; compound with trifluoro-acetic acid	462.8	
I-39	Cyclohexanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	488.9	
I-40	Morpholine-4-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	491.8	
I-41	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-hydroxy-2-methyl-propionamide; compound with trifluoro-acetic acid	464.8	
I-42	Cyclobutanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	460.8	
I-43	Pyrrolidine-1-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	475.8	
I-44	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	420.8	
I-45	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-methoxy-acetamide; compound with trifluoro-acetic acid	450.8	
I-46	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-hydroxy-2-phenyl-acetamide; compound with trifluoro-acetic acid	512.8	
I-47	2-Cyclopentyl-*N*-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	488.8	
I-48	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-methanesulfonyl-benzamide; compound with trifluoro-acetic acid	560.8	
I-49	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid	482.8	
I-50	Furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-	472.8	

	amide; compound with trifluoro-acetic acid		
I-51	1*H*-Pyrrole-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	471.8	
I-52	5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	502.8	
I-53	1-Methyl-1*H*-pyrrole-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	485.8	
I-54	2-Acetylamino-*N*-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	477.9	
I-55	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-sulfamoyl-benzamide; compound with trifluoro-acetic acid	561.8	
I-56	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	476.9	
I-57	2-Oxo-thiazolidine-4-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	507.8	
I-58	Pyrazine-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	484.8	
I-59	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-hydroxy-acetamide; compound with trifluoro-acetic acid	436.8	
I-60	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-methyl-butyramide; compound with trifluoro-acetic acid	462.8	
I-61	Tetrahydro-furan-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	476.8	
I-62	2-Dimethylamino-*N*-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	463.8	
I-63	1-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-phenyl-urea; compound with trifluoro-acetic acid	497.8	
I-64	1-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-isopropyl-urea; compound with trifluoro-acetic acid	463.8	
I-65	1-Acetyl-piperidine-4-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	531.9	
I-66	4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	524.8	

I-67	Cyclopentanecarboxylic acid [(S)-3-(5-isobutyryl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	412	
I-68	Cyclopentanecarboxylic acid {(S)-3-[5-(3-methyl-butyryl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	426	
I-69	Cyclopentanecarboxylic acid [(S)-3-(5-butyryl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	412	
I-70	Cyclopentanecarboxylic acid [(S)-3-(5-cyclopropanecarbonyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	410	
I-71	Cyclopentanecarboxylic acid [(S)-3-(5-cyclobutanecarbonyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	424	
I-72	Cyclopentanecarboxylic acid {(S)-3-[5-(2-methyl-butyryl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	426	
I-73	Cyclopentanecarboxylic acid {(S)-3-[5-(2-ethyl-butyryl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	440	
I-74	{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-pyridin-3-yl-propyl}-carbamic acid *tert*-butyl ester	479	
I-75	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-butyl}-amide	488	
I-76	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-pyridin-3-yl-propyl}-amide	475	
I-77	Cyclopentanecarboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-butyl}-amide	488	
I-78	Cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	475	68.8-73.6
I-79	Cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	491	93.3-95.5
I-80	Cyclopentanecarboxylic acid {(S)-3-[5-(1-acetyl-piperidine-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	495	
I-81	Cyclopentanecarboxylic acid [(S)-3-(5-benzyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	432	
I-82	Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trifluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-	499	

	propyl}-amide; compound with trifluoro-acetic acid		
I-83	Cyclopentanecarboxylic acid {(S)-3-[5-(1-benzyl-5-oxo-pyrrolidine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	543	
I-84	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	460	
I-85	(1S,5R)-Bicyclo[3.1.0]hexane-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	486	
I-86	4,4-Difluoro-cyclohexanecarboxylic acid [(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-amide; compound with trifluoro-acetic acid	542	
I-87	*N*-[(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-acetamide; compound with trifluoro-acetic acid	438	
I-88	(2S,3S)-1-Methyl-5-oxo-2-pyridin-3-yl-pyrrolidine-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	580.8	
I-89	4-Dimethylamino-*N*-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-butyramide; compound with trifluoro-acetic acid	491.9	
I-90	1-Methyl-pyrrolidine-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	489.8	
I-91	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-morpholin-4-yl-propionamide; compound with trifluoro-acetic acid	519.9	
I-92	2-(2-Aza-bicyclo[2.2.1]hept-2-yl)-*N*-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	515.9	
I-93	1-(Furan-2-carbonyl)-4-hydroxy-pyrrolidine-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	585.8	
I-94	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-(5-morpholin-4-yl-tetrazol-2-yl)-acetamide; compound with trifluoro-acetic acid	573.8	
I-95	1-Methyl-piperidine-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	503.8	
I-96	1-Methyl-azepane-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-	517.9	

	phenyl-propyl}-amide; compound with trifluoro-acetic acid		
I-97	2-Methyl-1,2,3,4-tetrahydro-isoquinoline-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	551.8	
I-98	3-(2-Aza-bicyclo[2.2.1]hept-2-yl)-*N*-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-propionamide; compound with trifluoro-acetic acid	529.9	
I-99	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-piperidin-1-yl-acetamide; compound with trifluoro-acetic acid	503.9	
I-100	1-Isopropyl-5-oxo-pyrrolidine-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	531.9	
I-101	1-Ethyl-piperidine-4-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	517.9	
I-102	1-Isobutyl-5-oxo-pyrrolidine-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	545.9	
I-103	*N*-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-pyrrolidin-1-yl-acetamide; compound with trifluoro-acetic acid	489.8	
I-104	1-Methyl-5-sulfamoyl-1*H*-pyrrole-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	564.8	
I-105	(S)-1-Acetyl-pyrrolidine-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	517.8	
I-106	Cyclopentanecarboxylic acid {(S)-3-[5-(2-methyl-1,2,3,4-tetrahydro-isoquinoline-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	515.9	
I-107	Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(1-pyrimidin-2-yl-piperidine-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	531.9	
I-108	Cyclopentanecarboxylic acid {(S)-3-[5-(1-isobutyl-5-oxo-pyrrolidine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	509.9	
I-109	Cyclopentanecarboxylic acid {(S)-3-[5-(1-methyl-5-sulfamoyl-1*H*-pyrrole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	528.8	

I-110	Cyclopentanecarboxylic acid ((S)-3-{5-[1-(furan-2-carbonyl)-4-hydroxy-pyrrolidine-2-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	549.8	
I-111	Cyclopentanecarboxylic acid {(S)-3-[5-(4-benzyl-morpholine-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	545.9	
I-112	Cyclopentanecarboxylic acid {(S)-3-[5-(6-chloro-imidazo[2,1-*b*]thiazole-5-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	562.7	
I-113	Cyclopentanecarboxylic acid {(S)-3-[5-(5-methyl-1-phenyl-1*H*-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	562.8	
I-114	Cyclopentanecarboxylic acid {(S)-3-[5-(5-methyl-2-phenyl-2*H*-[1,2,3]triazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	527.8	
I-115	Cyclopentanecarboxylic acid ((S)-3-{5-[3-(2-chloro-phenyl)-5-methyl-isoxazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	561.8	
I-116	Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(4,5,6,7-tetrahydro-2*H*-indazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	490.8	
I-117	Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(1,4,5,6-tetrahydro-cyclopentapyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	476.8	
I-118	Cyclopentanecarboxylic acid {(S)-3-[5-(5-ethyl-2-methyl-2*H*-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	478.8	
I-119	[(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-carbamic acid *tert*-butyl ester	496	
I-120	Cyclopentanecarboxylic acid {(S)-3-[5-(4-fluoro-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	492	61.6-66.3
I-121	2-Chloro-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-fluoro-benzamide; compound with trifluoro-acetic acid	558.5	
I-122	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-trifluoromethyl-benzamide; compound with trifluoro-acetic acid	574.6	
I-123	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-	536.6	

	pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-methoxy-benzamide; compound with trifluoro-acetic acid		
I-124	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-methoxy-benzamide; compound with trifluoro-acetic acid	536.6	
I-125	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid	506.6	
I-126	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3,3-dimethyl-butyramide; compound with trifluoro-acetic acid	500.7	
I-127	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-succinamic acid methyl ester; compound with trifluoro-acetic acid	516.6	
I-128	2-Chloro-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid	540.6	
I-129	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-cyano-benzamide; compound with trifluoro-acetic acid	531.6	
I-130	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2,4-difluoro-benzamide; compound with trifluoro-acetic acid	542.6	
I-131	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2,6-difluoro-benzamide; compound with trifluoro-acetic acid	542.6	
I-132	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3,4-difluoro-benzamide; compound with trifluoro-acetic acid	542.5	
I-133	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-fluoro-benzamide; compound with trifluoro-acetic acid	524.6	
I-134	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-fluoro-benzamide; compound with trifluoro-acetic acid	524.5	
I-135	Furan-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	496.5	
I-136	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-isobutyramide; compound with trifluoro-acetic acid	472.6	
I-137	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-methyl-butyramide; compound with trifluoro-acetic acid	486.6	
I-138	3-Chloro-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-fluoro-benzamide; compound with trifluoro-acetic acid	558.5	
I-139	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-thiophen-2-	526.5	

	yl-acetamide; compound with trifluoro-acetic acid		
I-140	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-trifluoromethyl-benzamide; compound with trifluoro-acetic acid	574.5	
I-141	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-cyano-benzamide; compound with trifluoro-acetic acid	531.5	
I-142	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-ethoxy-benzamide; compound with trifluoro-acetic acid	550.5	
I-143	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3,5-difluoro-benzamide; compound with trifluoro-acetic acid	542.5	
I-144	Thiophene-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	512.5	
I-145	4-Chloro-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid	540.5	
I-146	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-methoxy-benzamide; compound with trifluoro-acetic acid	536.5	
I-147	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-fluoro-benzamide; compound with trifluoro-acetic acid	524.5	
I-148	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3,4-dimethoxy-benzamide; compound with trifluoro-acetic acid	566.5	
I-149	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2,2-dimethyl-propionamide; compound with trifluoro-acetic acid	586.5	
I-150	Cyclohexanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	512.5	
I-151	Morpholine-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	515.5	
I-152	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-oxalamic acid ethyl ester; compound with trifluoro-acetic acid	502.5	
I-153	Acetic acid 1-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propylcarbamoyl}-1-methyl-ethyl ester; compound with trifluoro-acetic acid	530.5	
I-154	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-phenyl-acetamide; compound with trifluoro-acetic acid	520.5	
I-155	(1R,2R)-2-Phenyl-cyclopropanecarboxylic acid {(S)-3-[5-	546.6	

	(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid		
I-156	Cyclobutanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	484.6	
I-157	Isoxazole-5-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	497.5	
I-158	5-Chloro-4-methoxy-thiophene-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	576.5	
I-159	4-Methyl-[1,2,3]thiadiazole-5-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	528.5	
I-160	3-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-1,1-diethyl-urea; compound with trifluoro-acetic acid	501.6	
I-161	3-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-1,1-diisopropyl-urea; compound with trifluoro-acetic acid	529.6	
I-162	3-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-1,1-dimethyl-urea; compound with trifluoro-acetic acid	473.6	
I-163	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	444.6	
I-164	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-methoxy-acetamide; compound with trifluoro-acetic acid	474.6	
I-165	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-cyclopentyl-propionamide; compound with trifluoro-acetic acid	526.6	
I-166	Acetic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propylcarbamoyl}-methyl ester; compound with trifluoro-acetic acid	502.6	
I-167	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-methoxy-phenyl)-acetamide; compound with trifluoro-acetic acid	550.6	
I-168	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-cyclopentyl-acetamide; compound with trifluoro-acetic acid	512.6	
I-169	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-dimethylamino-benzamide; compound with trifluoro-acetic acid	549.6	

I-170	3,5-Dimethyl-isoxazole-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	525.6	
I-171	3-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-1-methyl-1-phenyl-urea; compound with trifluoro-acetic acid	535.6	
I-172	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-methylsulfanyl-acetamide; compound with trifluoro-acetic acid	490.6	
I-173	3-Chloro-thiophene-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	546.5	
I-174	{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-carbamic acid methyl ester; compound with trifluoro-acetic acid	460.5	
I-175	1-Benzyl-3-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-1-methyl-urea; compound with trifluoro-acetic acid	549.6	
I-176	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dichloro-4-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	592	
I-177	3-Acetylamino-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid	563.6	
I-178	4-Acetylamino-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid	563.6	
I-179	1-Acetyl-piperidine-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	555.6	
I-180	3-Amino-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-propionamide; compound with trifluoro-acetic acid	473.6	
I-181	2-Amino-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	459.6	
I-182	(S)-Pyrrolidine-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	499.6	
I-183	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-cyano-acetamide; compound with trifluoro-acetic acid	469.6	
I-184	Cycloheptanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	526.6	
I-185	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-	526.6	

	pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-cyclohexyl-acetamide; compound with trifluoro-acetic acid		
I-186	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-cyclopropyl-acetamide; compound with trifluoro-acetic acid	484.6	
I-187	Furan-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	496.5	
I-188	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-isonicotinamide; compound with trifluoro-acetic acid	507.6	
I-189	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-terephthalamic acid methyl ester; compound with trifluoro-acetic acid	564.5	
I-190	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-piperidin-1-yl-propionamide; compound with trifluoro-acetic acid	541.6	
I-191	1*H*-Pyrrole-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	495.6	
I-192	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-succinamide; compound with trifluoro-acetic acid	501.6	
I-193	1*H*-Indole-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	545.6	
I-194	5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	526.6	
I-195	1-Methyl-1*H*-pyrrole-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	509.6	
I-196	2-Acetylamino-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	501.6	
I-197	Pyridine-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	507.6	
I-198	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-methylamino-benzamide; compound with trifluoro-acetic acid	535.7	
I-199	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-methylamino-benzamide; compound with trifluoro-acetic acid	535.7	
I-200	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-dimethylamino-benzamide; compound with trifluoro-acetic	549.7	

	acid		
I-201	5-Amino-1-phenyl-1*H*-pyrazole-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	587.7	
I-202	2,2,3,3-Tetramethyl-cyclopropanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	526.7	
I-203	2-Acetylamino-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-methylsulfanyl-butyramide; compound with trifluoro-acetic acid	575.7	
I-204	2-Ethyl-5-methyl-2*H*-pyrazole-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	538.7	
I-205	5-Amino-2*H*-[1,2,4]triazole-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	512.6	
I-206	5-Chloro-thiophene-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	546.5	
I-207	(S)-2-Acetylamino-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-methylsulfanyl-butyramide; compound with trifluoro-acetic acid	575.6	
I-208	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-diethylamino-propionamide; compound with trifluoro-acetic acid	529.7	
I-209	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-methyl-nicotinamide; compound with trifluoro-acetic acid	521.6	
I-210	3-Methyl-furan-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	510.6	
I-211	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-nicotinamide; compound with trifluoro-acetic acid	507.5	
I-212	Pyrazine-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	508.5	
I-213	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-(1-methyl-1*H*-imidazol-4-yl)-acetamide; compound with trifluoro-acetic acid	524.7	
I-214	3*H*-Imidazole-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-	496.6	

	phenyl-propyl}-amide; compound with trifluoro-acetic acid		
I-215	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-methanesulfonyl-acetamide; compound with trifluoro-acetic acid	522.6	
I-216	2-Methyl-thiazole-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	527.6	
I-217	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-imidazol-1-yl-acetamide; compound with trifluoro-acetic acid	510.7	
I-218	1-Methyl-1*H*-pyrazole-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	510.7	
I-219	1-Methyl-1*H*-imidazole-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	510.7	
I-220	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dichloro-4-methylsulfanyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	MH+ 560	67.2-72.6
I-221	5-{(S)-3-Phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carboxylic acid phenylamide; compound with trifluoro-acetic acid	463.3	
I-222	5-{(S)-3-Phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carboxylic acid (2,6-dimethyl-phenyl)-amide; compound with trifluoro-acetic acid	491.3	
I-223	Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-benzenesulfonyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	484.2	
I-224	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(thiophene-2-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	490.2	
I-225	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3,5-dimethyl-isoxazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	503.2	
I-226	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-chloro-1,3-dimethyl-1*H*-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	536.3	
I-227	2,5-Dimethyl-4-(5-{(S)-3-phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-sulfonyl)-furan-3-carboxylic acid methyl ester; compound with trifluoro-acetic acid	560.3	
I-228	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(1,3,5-trimethyl-1*H*-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with	516.3	

	trifluoro-acetic acid		
I-229	1-Methyl-5-(5-{(S)-3-phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-sulfonyl)-1*H*-pyrrole-2-carboxylic acid methyl ester; compound with trifluoro-acetic acid	545.3	
I-230	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-cyano-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	473.3	
I-231	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(1*H*-pyrrole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	437.4	
I-232	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methoxy-1*H*-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	517.4	
I-233	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(pyridine-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	449.3	
I-234	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	477.4	
I-235	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(pyrazine-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	450.3	
I-236	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	453.3	
I-237	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(tetrahydro-furan-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	442.4	
I-238	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-acetylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	505.4	
I-239	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(isoquinoline-7-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	499.4	
I-240	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	499.4	
I-241	Cyclopentanecarboxylic acid {(S)-3-[5-(3,5-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	474	
I-242	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(1*H*-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	487.4	

I-243	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(benzofuran-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	488.4	
I-244	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(benzo[*b*]thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	504.4	
I-245	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(1-methyl-1*H*-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	501.4	
I-246	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-chloro-6-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	496.4	
I-247	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methoxy-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	492.4	
I-248	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-methylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	477.4	
I-249	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[4-(3-methyl-5-oxo-4,5-dihydro-pyrazol-1-yl)-benzoyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	544.5	
I-250	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(tetrahydro-furan-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	442.4	
I-251	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	454.4	
I-252	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2-thiophen-2-yl-acetyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	468.4	
I-253	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-chloro-thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	488.3	
I-254	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-5-phenyl-furan-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	528.4	
I-255	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-fluoro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	480.4	
I-256	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[3-(2-oxo-pyrrolidin-1-yl)-benzoyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	531.4	

I-257	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(benzo[*b*]thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	504.4	
I-258	4-(5-{(S)-3-Phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-benzoic acid *tert*-butyl ester; compound with trifluoro-acetic acid	548.5	
I-259	3-(5-{(S)-3-Phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-benzoic acid *tert*-butyl ester; compound with trifluoro-acetic acid	548.5	
I-260	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3,5-dimethyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	467.4	
I-261	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(1*H*-indole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	487.4	
I-262	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(benzofuran-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	488.4	
I-263	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,3-dihydro-benzofuran-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	490.4	
I-264	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	478.4	
I-265	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	478.4	
I-266	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3,3-dimethyl-butyryl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	442.4	
I-267	Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-cyclobutanecarbonyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	426.4	
I-268	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-cyclohexyl-acetyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	468.4	
I-269	Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-cyclopentanecarbonyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	440.4	
I-270	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-cyclopentyl-acetyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	454.4	

I-271	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dichloro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	516.3	
I-272	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	491.4	
I-273	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	466.4	
I-274	Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-isobutyryl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	414.4	
I-275	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-methyl-butyryl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	428.4	
I-276	Tetrahydro-furan-3-carboxylic acid [(S)-1-phenyl-3-(5-propionyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-propyl]-amide; compound with trifluoro-acetic acid	400.4	
I-277	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	462.4	
I-278	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	462.4	
I-279	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-cyano-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	473.3	
I-280	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-chloro-2-ethyl-2*H*-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	500.3	
I-281	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-chloro-2-methyl-2*H*-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	486.3	
I-282	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,5-trimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	496.4	
I-283	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-fluoro-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	494.4	
I-284	Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-benzoyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	448.4	
I-285	1-Methyl-cyclohexanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	526.6	

I-286	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-morpholin-4-yl-propionamide; compound with trifluoro-acetic acid	543.7	
I-287	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-(4-methyl-piperazin-1-yl)-propionamide; compound with trifluoro-acetic acid	556.7	
I-288	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-(1*H*-[1,2,4]triazol-3-yl)-propionamide; compound with trifluoro-acetic acid	525.7	
I-289	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-(1*H*-tetrazol-5-yl)-acetamide; compound with trifluoro-acetic acid	512.7	
I-290	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-furan-2-yl-propionamide; compound with trifluoro-acetic acid	524.7	
I-291	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-thiophen-2-yl-propionamide; compound with trifluoro-acetic acid	540.7	
I-292	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-pyridin-3-yl-propionamide; compound with trifluoro-acetic acid	535.7	
I-293	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-(3*H*-imidazol-4-yl)-propionamide; compound with trifluoro-acetic acid	524.7	
I-294	Tetrahydro-furan-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	500.7	
I-295	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-(methanesulfonyl-methyl-amino)-acetamide; compound with trifluoro-acetic acid	551.7	
I-296	4-Methoxy-thiophene-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	542.6	
I-297	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3,3,3-trifluoro-propionamide; compound with trifluoro-acetic acid	512.6	
I-298	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-dimethylamino-acetamide; compound with trifluoro-acetic acid	487.7	
I-299	2-Methyl-cyclopropanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	484.6	
I-300	1-Cyano-cyclopropanecarboxylic acid {(S)-3-[5-(2-chloro-	495.6	

	6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid		
I-301	5-Acetyl-2,4-dimethyl-1*H*-pyrrole-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	565.6	
I-302	2,4-Dimethyl-6-oxo-6*H*-pyran-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	552.6	
I-303	4,6-Dimethyl-pyrimidine-5-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	536.6	
I-304	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2,4-dimethyl-nicotinamide; compound with trifluoro-acetic acid	535.6	
I-305	(S)-2-Amino-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-propionamide; compound with trifluoro-acetic acid	473.6	
I-306	(S)-2-Amino-4-methyl-pentanoic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	515.6	
I-307	(S)-2-Amino-*N*-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-methyl-butyramide; compound with trifluoro-acetic acid	501.6	
I-308	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-butyramide; compound with trifluoro-acetic acid	472.5	
I-309	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2,3-dimethyl-benzamide; compound with trifluoro-acetic acid	534.5	
I-310	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2,6-dimethyl-benzamide; compound with trifluoro-acetic acid	534.5	
I-311	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-fluoro-phenyl)-acetamide; compound with trifluoro-acetic acid	538.6	
I-312	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-phenyl-propionamide; compound with trifluoro-acetic acid	534.6	
I-313	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-methanesulfonyl-benzamide; compound with trifluoro-acetic acid	584.6	
I-314	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-methyl-benzamide; compound with trifluoro-acetic acid	520.6	
I-315	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-	520.7	

	pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-methyl-benzamide; compound with trifluoro-acetic acid		
I-316	*N*-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-4-methyl-benzamide; compound with trifluoro-acetic acid	520.7	
I-317	Cyclopentanecarboxylic acid {(S)-3-[5-(3,4-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	474	
I-318	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-amino-2,6-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	499.3	
I-319	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-6-oxo-6*H*-pyran-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	494.3	
I-320	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	492.4	
I-321	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(1-acetyl-piperidine-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	497.3	
I-322	Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-cyclohexanecarbonyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	454.4	
I-323	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(furan-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	438.3	
I-324	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(furan-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	438.3	
I-325	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	526.3	
I-326	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	462.3	
I-327	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	516.3	
I-328	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	454.2	
I-329	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic	491.3	

	acid		
I-330	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-ethyl-5-methyl-2*H*-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	480.3	
I-331	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	463.3	
I-332	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-methyl-furan-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	452.3	
I-333	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-fluoro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	480.3	
I-334	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	484.3	
I-335	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-chloro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	482.3	
I-336	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-thiazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	469.3	
I-337	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(1-methyl-1*H*-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	452.3	
I-338	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-methoxy-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	484.2	
I-339	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-acetylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	505.3	
I-340	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-acetylamino-2-phenyl-acetyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	519.3	
I-341	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-((S)-pyrrolidine-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	441.3	
I-342	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-chloro-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	512.2	
I-343	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(isoquinoline-1-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-	499.3	

	propyl}-amide; compound with trifluoro-acetic acid		
I-344	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	499.3	
I-345	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	499.3	
I-346	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-6-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	499.3	
I-347	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-8-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	499.3	
I-348	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-carbamoyl-propionyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	443.3	
I-349	Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-cyclopropanecarbonyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid	412.3	
I-350	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-butyryl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	428.4	
I-351	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[2-(methanesulfonyl-methyl-amino)-acetyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	493.4	
I-352	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-difluoro-4-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	514.4	
I-353	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trifluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	502.4	
I-354	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-4-methoxy-1-phenyl-butyl}-amide	518	61.1-84.5
I-355	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-((R)-2-acetylamino-3-methyl-butyryl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	485.3	
I-356	Cyclopentanecarboxylic acid {(S)-3-[5-(1-methyl-5-oxo-pyrrolidine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	467	
I-357	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with	478.2	

	trifluoro-acetic acid		
I-358	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid;	493.2	
I-359	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-2-methylsulfanyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	524.3	
I-360	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-4-hydroxy-1-phenyl-butyl}-amide	504	78.9-84.9
I-361	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	538.5	
I-362	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,3,6-trichloro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	550.3	
I-363	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methyl-2-phenyl-2*H*-[1,2,3]triazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	529.5	
I-364	2-Oxo-imidazolidine-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	514.4	
I-365	5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid;	519.5	
I-366	2-Cyclohexyl-*N*-{(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamidetrifluoro-acetic acid;	519.5	
I-367	*N*-{(S)-3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-fluoro-phenyl)-acetamidetrifluoro-acetic acid;	531.5	
I-368	*N*-{(S)-3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-methoxy-phenyl)-acetamidetrifluoro-acetic acid;	543.5	
I-369	1-{(S)-3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-isopropyl-ureatrifluoro-acetic acid;	480.5	
I-370	2-Oxo-thiazolidine-4-carboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid;	524.4	
I-371	Furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid;	489.4	
I-372	*N*-{(S)-3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-	535.4	

	hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3,4-difluoro-benzamidetrifluoro-acetic acid;		
I-373	1-{(S)-3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-phenyl-ureatrifluoro-acetic acid;	514.5	
I-374	4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid;	541.5	
I-375	Tetrahydro-pyran-4-carboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid;	507.5	
I-376	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	477.5	
I-377	5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	503.4	
I-378	2-Cyclohexyl-*N*-{(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	503.5	
I-379	*N*-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-fluoro-phenyl)-acetamide; compound with trifluoro-acetic acid	515.5	
I-380	*N*-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-methoxy-phenyl)-acetamide; compound with trifluoro-acetic acid	527.5	
I-381	Cyclohexanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	489.5	
I-382	Furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	473.4	
I-383	*N*-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	421.4	
I-384	*N*-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3,4-difluoro-benzamide; compound with trifluoro-acetic acid	519.4	
I-385	1-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3-phenyl-urea; compound with trifluoro-acetic acid	498.4	
I-386	4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-	525.5	

	c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid		
I-387	Tetrahydro-pyran-4-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	491.4	
I-388	Tetrahydro-pyran-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	514.4	
I-389	(R)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	476.5	
I-390	(R)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	478.4	
I-391	(R)-Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,5-trimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	496.4	
I-392	(S)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	476.5	
I-393	(S)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	477.5	
I-394	(S)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	478.5	
I-395	(S)-Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,5-trimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	496.4	
I-396	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methyl-3-phenyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	529.4	
I-397	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	499.4	
I-398	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	499.4	
I-399	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-6-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	499.4	
I-400	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-8-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-	499.5	

	yl]-propyl}-amide; compound with trifluoro-acetic acid		
I-401	(R)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	477.7	
I-402	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-4-pyridin-4-yl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	551.3	
I-403	Cyclopentanecarboxylic acid {(S)-3-[5-(4-cyano-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	499.2	
I-404	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid	475	
I-405	Cyclopropanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid	447	
I-406	5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid	503	
I-407	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid	477	
I-408	4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid	525	
I-409	1,1-Dioxo-hexahydro-1λ^6-thiopyran-4-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	538	
I-410	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[5-amino-1-(4-methoxy-phenyl)-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	559.4	
I-411	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(1-phenyl-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	582.3	
I-412	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[1-(4-methoxy-phenyl)-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	612.3	
I-413	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[1-(2-methoxy-phenyl)-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	612.3	

I-414	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[1-(4-chloro-phenyl)-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	616.3	
I-415	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(1-*p*-tolyl-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	596.3	
I-416	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[1-(4-fluoro-phenyl)-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	600.3	
I-417	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methyl-1-phenyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	528.3	
I-418	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methyl-1-*p*-tolyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	542.4	
I-419	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[2-(4-methoxy-phenyl)-5-methyl-2*H*-pyrazole-3-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	558.4	
I-420	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-2-phenyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	554.4	
I-421	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[4,6-dimethyl-2-(2-methyl-thiazol-4-yl)-pyrimidine-5-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	575.3	
I-422	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-2-pyridin-4-yl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	555.4	
I-423	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-bromo-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	529.2	
I-424	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-fluoro-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	467.3	
I-425	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-chloro-pyridine-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	483.3	
I-426	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methoxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-	479.3	

	1-phenyl-propyl}-amide; compound with trifluoro-acetic acid		
I-427	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	526.3	
I-428	Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2-trifluoromethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	532.3	
I-429	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-amino-1-phenyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	529.2	
I-430	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[5-amino-1-(4-fluoro-phenyl)-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	547.2	
I-431	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[5-amino-1-(2-methoxy-phenyl)-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	559.2	
I-432	Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[1-(4-chloro-phenyl)-5-methyl-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid	562.2	
I-433	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-bromo-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	556.2	
I-434	Cyclopentanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-butyl}-amide	540	71.3-72.9
I-435	4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	526	131.8-133.2
I-436	Pentanoic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	463.3	
I-437	Cyclobutanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	461.3	
I-438	2-Cyclopentyl-*N*-{(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	489.3	
I-439	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-5-trifluoromethyl-oxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	521.2	

I-440	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-amino-6-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	531.3	
I-441	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-4-nitro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid;	521.3	
I-442	*N*-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-propionamide; compound with trifluoro-acetic acid	435.3	
I-443	*N*-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-butyramide; compound with trifluoro-acetic acid	449.3	
I-444	Cyclopropanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	447.3	
I-445	{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-carbamic acid *tert*-butyl ester	493	
I-446	Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-3a,6a-dimethyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	502 (M+Na) 524	
I-447	5-Oxo-1-(2,2,2-trifluoro-ethyl)-pyrrolidine-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	571	93.3-94.5
I-448	4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-butyl}-amide	540	71.3-72.9
I-449	4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-butyl}-amide	540	86.4-87.0
I-450	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(6-chloro-2-dimethylamino-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	526.1	
I-451	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-cyano-2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	507.1	
I-452	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-iodo-2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	608	
I-453	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	482.1	
I-454	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,5-di-*tert*-	591.2	

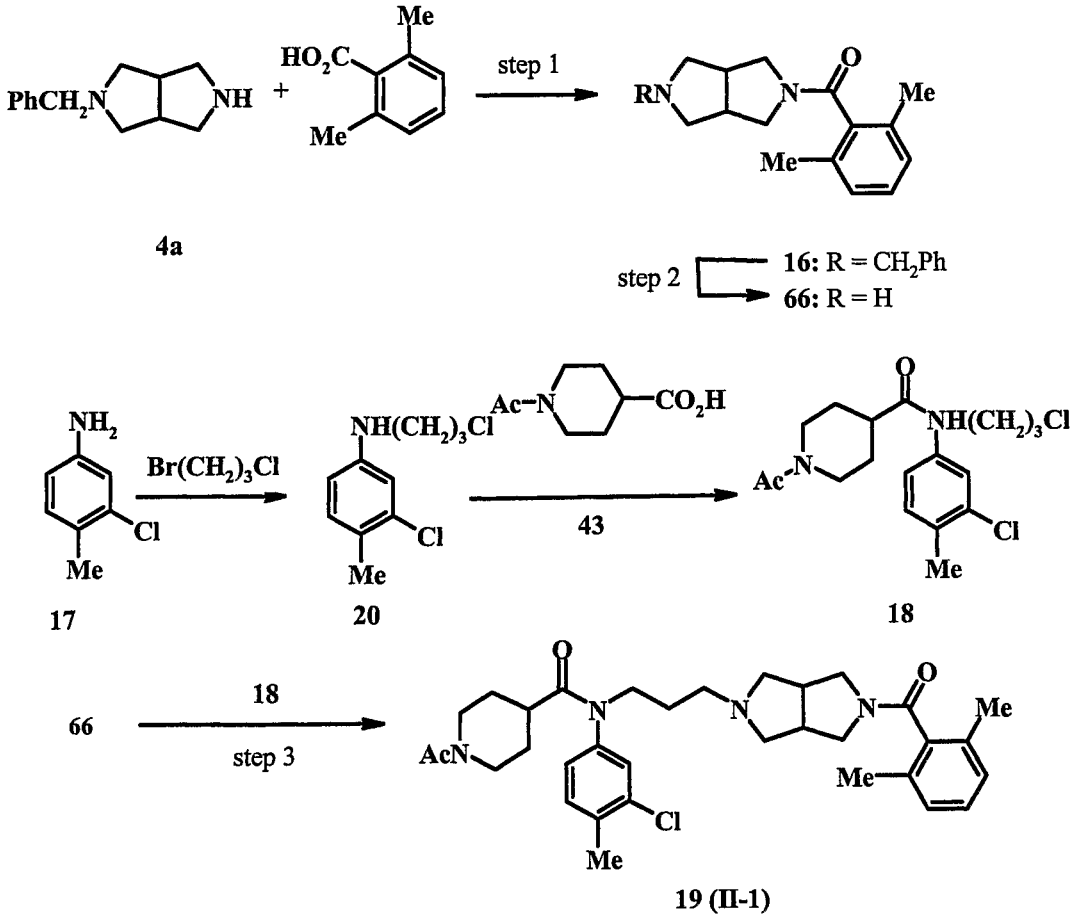
	butyl-4-cyano-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid		
I-455	4-Oxo-cyclohexanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	502	94.3-95.7
I-456	Cyclopentanecarboxylic acid {(S)-3-[5-(2-ethoxy-4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	520	
I-457	Cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	489	
I-458	Cyclopropanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	461	
I-459	5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	517	
I-460	4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	539	
I-461	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-bromo-2,5-di-*tert*-butyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	646.1	
I-462	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3,5-diethyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	495.2	
I-463	Cyclopentanecarboxylic acid {(S)-3-[5-(2-methanesulfonyl-4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	554	
I-464	(5-{5-[(S)-3-(Cyclopentanecarbonyl-amino)-3-phenyl-propyl]-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl}-4,6-dimethyl-pyrimidin-2-yloxy)-acetic acid methyl ester	564	
I-465	Cyclopentanecarboxylic acid {(S)-3-[5-(2-benzylamino-4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	581	
I-466	(5-{5-[(S)-3-(Cyclopentanecarbonyl-amino)-3-phenyl-propyl]-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl}-4,6-dimethyl-pyrimidin-2-yloxy)-acetic acid; compound with trifluoro-acetic acid	550	
I-467	Cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-6-oxo-6*H*-pyran-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-	492	113.0-113.7

	2-yl]-1-phenyl-propyl}-amide		
I-468	Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trimethyl-benzenesulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	524.6	
I-469	Cyclopentanecarboxylic acid {(R)-3-[5-(2-dimethylamino-4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	519	
I-470	*N*-{(S)-3-[5-(4,6-Dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3,4-difluoro-benzamide; compound with trifluoro-acetic acid	520	
I-471	Cyclobutanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	462	
I-472	2-Cyclopentyl-*N*-{(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	490	
I-473	5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	504	
I-474	2-Oxo-thiazolidine-4-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	509	
I-475	2-Cyclohexyl-*N*-{(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid	504	
I-476	3,3-Difluoro-cyclobutanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	498	
I-477	3-Oxo-cyclobutanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	476	
I-478	*N*-{(S)-3-[5-(4,6-Dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-3,4-difluoro-benzenesulfonamide; compound with trifluoro-acetic acid	556	
I-479	Cyclopropanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	448	
I-480	Thiophene-2-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	490	

I-481	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-methyl-benzofuran-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	502	
I-482	Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(7-methoxy-3-methyl-1*H*-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	531	
I-483	Tetrahydro-pyran-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid	492	
I-484	Cyclopentanecarboxylic acid [(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-amide	494	
I-485	4,4-Difluoro-cyclohexanecarboxylic acid [(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-amide	544	
I-486	4-Hydroxy-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	506.3	
I-487	4-Hydroxy-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	506.3	
I-488	3,3-Difluoro-cyclobutanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide	498	

Compounds of formula I wherein R^1 is aryl are conveniently prepared by an alkylation sequence depicted in Scheme 5. 2-Benzyl-octahydro-pyrrolo[3,4-c]pyrrole (4a) is acylated prior to debenzylation as depicted in Scheme 1. The acylation can be carried out by standard techniques described above. Procedures for hydrogenolysis of the benzyl protecting group (step 2) to afford 66 are well known in the art. In other embodiments of the invention 4a may be alkylated or sulfonylated prior to hydrogenolysis of the benzyl group as described above.

Scheme 5



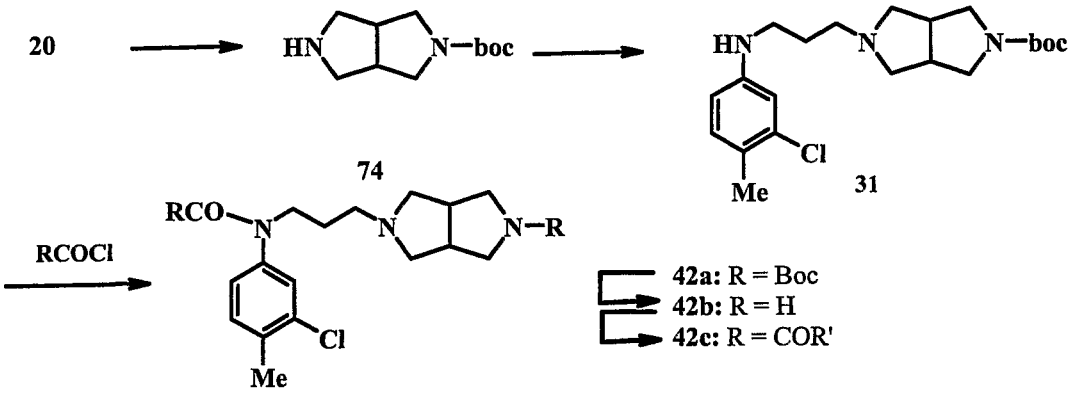
1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-chloro-propyl)-amide (18) was prepared by the general methodology disclosed by S. Imamura *et al.* (WO 01/25200). (Scheme 5) One skilled in the art will immediately recognize that 18 consists of a fragment derived from an aryl amine, a carboxylic acid and a difunctional alkylene chain. Substitution of other aryl or heteroaryl amines and other carboxylic acids readily affords analogs of 18 with other substitution on the aryl ring, with heteroaryl amines and with other acyl radicals.

Alkylation of an aryl amine 17 (or optionally a heteroaryl amine) with 1-bromo-3-chloropropane affords 20. The reaction may typically be carried out in the presence of a base such as TEA, DIPEA or DBU or an inorganic base such as Na_2CO_3, $NaHCO_3$, K_2CO_3 or Cs_2CO_3: optionally in the presence of a phase transfer catalyst, and in a solvent such as acetonitrile, DMF, DMSO, 1,4-dioxane, THF or toluene. While exemplified here with bromo-chloro-propane, other difunctional propylene compounds can also be employed. Alternatively, a metal salt of the amine (i.e. a deprotonated form) may be the nucleophile in a suitable solvent such as THF, DMF or 1,4-dioxane. Preferably the reaction is carried out by reacting the amine 17 and bromo-chloro-propane with either K_2CO_3 or Cs_2CO_3 dissolved in DMF or MeCN, DBU in MeCN or K_2CO_3 and 18-crown-6 (1,4,7,10,13,16-hexaoxacyclooctadecane) in THF. Potassium iodide is frequently added to allow for the

in situ generation of an alkyl iodide. Optimal conditions for the alkylation may vary depending upon the nature of the reactants and one skilled in the art can optimize the specific reaction conditions without undue experimentation.

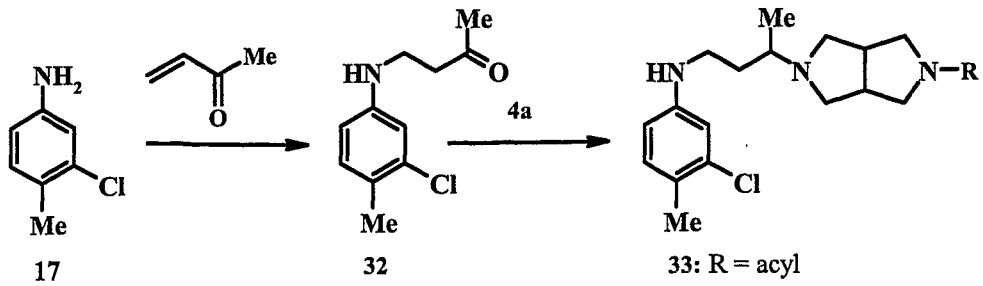
Acylation of the resulting secondary amine 20 by 1-acetyl-piperidine-4-carboxylic acid or the corresponding carboxylic acid chloride, or other carboxylic acids, is carried out by standard protocols as described previously. Urea and sulfonamide analogs are prepared by substituting an isocyanate or sulfonyl chloride for the carboxylic acid.

Scheme 6



Alternative the aryl amine is sufficiently deactivated that alkylation of 74 or other octahydro-pyrrolo[3,4-c]pyrrole derivative can carried out directly with 20 without further modification of the aryl amine functionality to afford 31 which can further acylated. The reaction is exemplified in Scheme 6 utilizing a Boc protecting group which can be removed under acidic conditions for final elaboration of the compounds of the invention (conversion of 42a to 42c)

Scheme 7



Methylation of the propylene linker can be achieved from butanone derivatives which are readily available by reacting an aryl or heteroaryl amine with methyl vinyl ketone (Scheme 7). Reductive alkylation of 32 and 66 (or other amide described herein) and acylation of 33 proceeds as described previously.

The synthesis of compounds wherein R^6 is other then hydrogen is exemplified in Examples 37, 38, 40 and 41.

Representative compounds of the present invention in which R^1 is an aryl or heteroaryl group are compiled in Table 2.

Table 2

II-1	NAME	MS[1]	mp
II-2	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	579	58.1-65.2
II-3	*N*-{3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-*N*-(4-fluoro-phenyl)-benzenesulfonamide; compound with trifluoro-acetic acid	536.6	
II-4	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	580	91.8-96.9
II-5	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	596	67.2-72.6
II-6	4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	596	
II-7	1-Acetyl-piperidine-4-carboxylic acid [3-(5-benzyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-propyl]-(3-chloro-4-methyl-phenyl)-amide	537	
II-8	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,5-trimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	599.3	
II-9	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	581.4	
II-10	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-difluoro-4-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	617.4	
II-11	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	603.3	
II-12	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-6-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	599.4	
II-13	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-	633.3	

	phenyl)-{3-[5-(2,6-dichloro-4-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid		
II-14	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,6-trimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	593.4	
II-15	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	570.4	
II-16	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	594.4	
II-17	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-chloro-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	615.4	
II-18	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-acetyl-2,4-dimethyl-1*H*-pyrrole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	610.5	
II-19	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	589	67.0-74.1
II-20	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with hydrochloric acid	573	152.4 -161.3
II-21	Furan-2-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	521.4	
II-22	*N*-(3-Chloro-4-methyl-phenyl)-*N*-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-isobutyramide; compound with trifluoro-acetic acid	497.5	
II-23	Thiophene-2-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	537.4	
II-24	Cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	537.5	
II-25	Morpholine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide;	540.5	

	compound with trifluoro-acetic acid		
II-26	Isoxazole-5-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	522.4	
II-27	4-Methyl-[1,2,3]thiadiazole-5-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	553.4	
II-28	Pyrrolidine-1-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	524.5	
II-29	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3,3-dimethyl-urea; compound with trifluoro-acetic acid	498.5	
II-30	*N*-(3-Chloro-4-methyl-phenyl)-*N*-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-acetamide; compound with trifluoro-acetic acid	496.4	
II-31	*N*-(3-Chloro-4-methyl-phenyl)-2-cyclopentyl-*N*-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-acetamide; compound with trifluoro-acetic acid	537.5	
II-32	3,5-Dimethyl-isoxazole-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	550.5	
II-33	*N*-(3-Chloro-4-methyl-phenyl)-*N*-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-4-methanesulfonyl-benzamide; compound with trifluoro-acetic acid	609.5	
II-34	Piperidine-1-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	538.5	
II-35	1-Methyl-1*H*-pyrrole-2-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	534.5	
II-36	1-(3-Chloro-4-methyl-phenyl)-3-(4-chloro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	580.4	
II-37	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-butyl}-amide	594	60.9-62.4
II-38	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-	546	60.9-61.1

	yl]-propyl}-*p*-tolyl-amide		
II-39	1-Acetyl-piperidine-4-carboxylic acid (3-{5-[5-chloro-2-(2,5-dimethyl-pyrrol-1-yl)-benzoyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-propyl)-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	678.3	
II-40	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-quinoline-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	616.3	
II-41	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-chloro-4-methyl-6-pyrrolidin-1-yl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	668.4	
II-42	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[2-(1,1-dioxo-1λ^6-thiomorpholin-4-yl)-benzoyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid	684.4	
II-43	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-bromo-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	657.4	
II-44	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-2-pyridin-4-yl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	658.5	
II-45	5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carboxylic acid (2-isopropyl-phenyl)-amide; compound with trifluoro-acetic acid	608.5	
II-46	5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carboxylic acid (2,6-dimethyl-phenyl)-amide; compound with trifluoro-acetic acid	594.5	
II-47	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,6-trimethyl-benzenesulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	629.5	
II-48	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(thiophene-2-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	593.4	
II-49	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-isoxazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	606.5	
II-50	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-chloro-1,3-dimethyl-1*H*-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	639.5	
II-51	4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-	663.6	

	methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-sulfonyl)-2,5-dimethyl-furan-3-carboxylic acid methyl ester; compound with trifluoro-acetic acid		
II-52	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1,3,5-trimethyl-1*H*-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	619.6	
II-53	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(1-acetyl-piperidine-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	600.6	
II-54	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-cyano-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	576.5	
II-55	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	611.6	
II-56	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	579.6	
II-57	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(furan-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	541.5	
II-58	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(furan-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	541.5	
II-59	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(pyridine-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	552.5	
II-60	3-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-benzoic acid methyl ester; compound with trifluoro-acetic acid	609.6	
II-61	4-((3-Chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-piperidine-1-carboxylic acid *tert*-butyl ester	637.6	
II-62	Piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	537.5	
II-63	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-phenyl-amide	532	60.1-61.1
II-64	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methanesulfonyl-benzoyl)-hexahydro-	629.2	

	pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid		
II-65	4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-benzoic acid methyl ester; compound with trifluoro-acetic acid	609.3	
II-66	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1*H*-pyrrole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	540.4	
II-67	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	565.4	
II-68	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	619.5	
II-69	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	571.5	
II-70	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-methyl-1*H*-pyrrole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	554.5	
II-71	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	557.5	
II-72	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(pyridine-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	552.5	
II-73	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	580.6	
II-74	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methoxy-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	609.6	
II-75	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-ethyl-5-methyl-2*H*-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	583.6	
II-76	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	566.5	

II-77	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-methyl-furan-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	555.5	
II-78	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	552.5	
II-79	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-fluoro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	583.6	
II-80	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	587.6	
II-81	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	585.5	
II-82	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3*H*-imidazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	541.5	
II-83	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	556.5	
II-84	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-thiazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	572.5	
II-85	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-methyl-1*H*-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	555.6	
II-86	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(tetrahydro-furan-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	545.5	
II-87	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methoxy-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	587.5	
II-88	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	581.5	
II-89	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-dimethylamino-benzoyl)-hexahydro-	594.6	

	pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid		
II-90	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(isoquinoline-1-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	602.5	
II-91	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-cyclohexanecarbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	571.6	
II-92	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(quinoline-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	602.5	
II-93	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1*H*-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	590.5	
II-94	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(benzofuran-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	591.3	
II-95	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(benzo[*b*]thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	607.4	
II-96	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-methyl-1*H*-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	604.4	
II-97	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methoxy-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	595.5	
II-98	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1*H*-indole-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	590.5	
II-99	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-chloro-thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	591.5	
II-100	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-5-phenyl-furan-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	631.6	
II-101	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-fluoro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	583.6	

II-102	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(1-benzyl-3,5-dimethyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	659.7	
II-103	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,6-trimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	641.6	
II-104	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-chloro-2,6-dimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	645.6	
II-105	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-1-phenyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	645.6	
II-106	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-difluoro-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	601.6	
II-107	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-fluoro-6-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	637.6	
II-108	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(6-chloro-2-fluoro-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	617.6	
II-109	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-6-fluoro-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	617.5	
II-110	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,6-trifluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	605.5	
II-111	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-chloro-2-fluoro-6-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	671.6	
II-112	1-Methanesulfonyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	615.6	
II-113	1-Cyclopropanecarbonyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	605.6	
II-114	Piperidine-1,4-dicarboxylic acid 4-((3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-	608.6	

	pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide) 1-dimethylamide; compound with trifluoro-acetic acid		
II-115	Piperidine-1,4-dicarboxylic acid 4-((3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide) 1-ethylamide; compound with trifluoro-acetic acid	608.6	
II-116	1-Dimethylsulfamoyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	644.6	
II-117	1-(2,2,2-Trifluoro-acetyl)-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	633.5	
II-118	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-pyridin-2-yl-amide	532.5	
II-119	1-Acetyl-piperidine-4-carboxylic acid (3,4-dichloro-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	600	
II-120	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-amino-1-phenyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	632.5	
II-121	1-Acetyl-piperidine-4-carboxylic acid (3-{5-[5-amino-1-(4-methoxy-phenyl)-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-propyl)-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	662.5	
II-122	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-phenyl-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	685.5	
II-123	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[1-(4-methoxy-phenyl)-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid	715.5	
II-124	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[1-(2-methoxy-phenyl)-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid	715.5	
II-125	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[1-(4-chloro-phenyl)-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid	719.5	
II-126	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-*p*-tolyl-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	699.5	
II-127	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-	703.5	

	phenyl)-(3-{5-[1-(4-fluoro-phenyl)-5-trifluoromethyl-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid		
II-128	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-amino-1-*p*-tolyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	646.5	
II-129	1-Acetyl-piperidine-4-carboxylic acid (3-{5-[5-amino-1-(4-fluoro-phenyl)-1*H*-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-propyl)-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	650.5	
II-130	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-1-phenyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	631.5	
II-131	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-1-*p*-tolyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	645.5	
II-132	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-2-*p*-tolyl-2*H*-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	645.5	
II-133	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[2-(4-methoxy-phenyl)-5-methyl-2*H*-pyrazole-3-carbonyl]-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid	661.5	
II-134	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	609.5	
II-135	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-phenyl-amide	531	
II-136	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	632.3	
II-137	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-acetylamino-4-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	626.4	
II-138	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-fluoro-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	570.3	
II-139	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-chloro-pyridine-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	586.3	

II-140	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methoxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	582.3	
II-141	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	629.3	
II-142	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methanesulfonylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	644.3	
II-143	1-(3-Chloro-4-methyl-phenyl)-3-(2,6-dichloro-pyridin-4-yl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	617.2	
II-144	1-(3-Chloro-4-methyl-phenyl)-3-(3,5-dimethyl-isoxazol-4-yl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	565.3	
II-145	1-(3-Chloro-4-methyl-phenyl)-3-cyclopentyl-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	538.3	
II-146	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-thiophen-3-yl-urea; compound with trifluoro-acetic acid	552.3	
II-147	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-furan-2-ylmethyl-urea; compound with trifluoro-acetic acid	550.3	
II-148	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-[1-(2,2,2-trifluoro-acetyl)-piperidin-3-yl]-urea; compound with trifluoro-acetic acid	650.3	
II-149	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-[1-(2,2,2-trifluoro-acetyl)-piperidin-4-yl]-urea; compound with trifluoro-acetic acid	650.3	
II-150	1-(3-Chloro-4-methyl-phenyl)-3-cyclohexylmethyl-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	566.4	
II-151	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-(5-methyl-2-trifluoromethyl-furan-3-yl)-urea; compound with trifluoro-acetic acid	618.3	
II-152	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-	546.3	

	yl]-propyl}-3-phenyl-urea; compound with trifluoro-acetic acid		
II-153	1-(3-Chloro-4-methyl-phenyl)-3-(3-chloro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	580.2	
II-154	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-(4-fluoro-phenyl)-urea; compound with trifluoro-acetic acid	564.3	
II-155	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-*p*-tolyl-urea; compound with trifluoro-acetic acid	560.3	
II-156	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-(4-trifluoromethoxy-phenyl)-urea; compound with trifluoro-acetic acid	630.3	
II-157	(3-(3-Chloro-4-methyl-phenyl)-3-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-ureido)-acetic acid ethyl ester; compound with trifluoro-acetic acid	556.3	
II-158	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-(4-fluoro-benzyl)-urea; compound with trifluoro-acetic acid	578.3	
II-159	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-(4-methoxy-benzyl)-urea; compound with trifluoro-acetic acid	590.3	
II-160	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-(4-trifluoromethyl-phenyl)-urea; compound with trifluoro-acetic acid	614.2	
II-161	3-(4-Acetyl-phenyl)-1-(3-chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	588.3	
II-162	1-(3-Chloro-4-methyl-phenyl)-3-(3,4-difluoro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	582.2	
II-163	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-6-oxo-6*H*-pyran-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	598	
II-164	1-(3-Chloro-4-methyl-phenyl)-3-(4-dimethylamino-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	589.3	

II-165	4-(3-(3-Chloro-4-methyl-phenyl)-3-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-ureido)-piperidine-1-carboxylic acid benzyl ester; compound with trifluoro-acetic acid	687.3	
II-166	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-(4-methoxy-phenyl)-urea; compound with trifluoro-acetic acid	576.3	
II-167	1-(3-Chloro-4-methyl-phenyl)-3-cyclohexyl-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	552.4	
II-168	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-(4-methylsulfanyl-phenyl)-urea; compound with trifluoro-acetic acid	592.3	
II-169	1-(3-Chloro-4-methyl-phenyl)-3-(2,4-dichloro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	616.2	
II-170	1-(3-Chloro-4-methyl-phenyl)-3-(4-chloro-2-trifluoromethyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	648.3	
II-171	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-(4-methyl-benzyl)-urea; compound with trifluoro-acetic acid	574.4	
II-172	1-(3-Chloro-4-methyl-phenyl)-3-(3,4-dichloro-benzyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	630.3	
II-173	1-(3-Chloro-4-methyl-phenyl)-3-(4-cyano-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid	571.4	
II-174	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3,4-dimethyl-phenyl)-amide	559	
II-175	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-3a,6a-dimethyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	608, 610	
II-176	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-6-oxo-6*H*-pyran-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	590	
II-177	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-	563	

	amide; compound with trifluoro-acetic acid		
II-178	4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(5-acetyl-2,4-dimethyl-1*H*-pyrrole-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	603.1	
II-179	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	574.1	
II-180	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,5-trimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	592.1	
II-181	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	572.1	
II-182	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-1-phenyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	638.2	
II-183	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dichloro-4-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	692	
II-184	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-2-pyridin-4-yl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	651.2	
II-185	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-2-phenyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	650.2	
II-186	4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-3-phenyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	625.2	
II-187	Cyclobutanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	509.3	
II-188	Cyclopentanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	523.3	
II-189	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-[1-(2,2,2-trifluoro-acetyl)-piperidin-3-yl]-urea; compound with trifluoro-acetic acid	649.3	
II-190	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-	595	107.9

	phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-2-hydroxy-propyl}-amide		-409.8
II-191	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-*bis*-hydroxymethyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	640	
II-192	4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-*p*-tolyl-amide	538	
II-193	4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-phenyl-amide	525	
II-194	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-5-trifluoromethyl-oxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	624.1	
II-195	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-phenyl-5-trifluoromethyl-oxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	686.1	
II-196	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-amino-6-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	634.1	
II-197	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-4-nitro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amidetrifluoro-acetic acid;	624.3	
II-198	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1,3,5-trimethyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	583.4	
II-199	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-1*H*-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	569.3	
II-200	*N*-(3-Chloro-4-methyl-phenyl)-*N*-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-propionamide; compound with trifluoro-acetic acid	483.3	
II-201	*N*-(3-Chloro-4-methyl-phenyl)-*N*-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-butyramide; compound with trifluoro-acetic acid	497.3	
II-202	Pentanoic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	511.3	
II-203	Cyclopropanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-	495.3	

	pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid		
II-204	3-(4-Chloro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-1-*p*-tolyl-urea	546	60.5-62.5
II-205	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-chloro-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	615.1	
II-206	1-Acetyl-piperidine-4-carboxylic acid [3-(5-benzoyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-propyl]-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	551.2	
II-207	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,4-bis-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	687.1	
II-208	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	611.2	
II-209	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3-dihydro-benzofuran-7-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	593.2	
II-210	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3-dimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	611.3	
II-211	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-chloro-5-methylsulfanyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	631.2	
II-212	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-fluoro-3-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	637.2	
II-213	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-chloro-2-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	603.2	
II-214	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	579.3	
II-215	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-ethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	595.3	
II-216	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-	587.2	

	phenyl)-{3-[5-(2,3-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid		
II-217	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-fluoro-5-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	583.3	
II-218	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-4,5-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	621.2	
II-219	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,5-trifluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	605.2	
II-220	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	587.2	
II-221	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dichloro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	621.2	
II-222	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-dichloro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	621.2	
II-223	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3-dichloro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	621.2	
II-224	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	587.2	
II-225	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-4-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	603.2	
II-226	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-chloro-2-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	603.2	
II-227	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	581.3	
II-228	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-trifluoromethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with	635.3	

	trifluoro-acetic acid		
II-229	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	619.3	
II-230	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	581.3	
II-231	Acetic acid 2-(5-{3-[(1-acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-phenyl ester; compound with trifluoro-acetic acid	609.3	
II-232	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	631.2	
II-233	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-bromo-2-chloro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	665.2	
II-234	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-acetyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	593.3	
II-235	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-dimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	611.3	
II-236	2-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-benzoic acid methyl ester; compound with trifluoro-acetic acid	609.3	
II-237	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,5-bis-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	687.3	
II-238	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3,4-trifluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	605.3	
II-239	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-fluoro-3-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	637.3	
II-240	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dichloro-5-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	639.2	
II-241	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-	637.3	

	phenyl)-{3-[5-(4-fluoro-2-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid		
II-242	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-fluoro-5-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	637.3	
II-243	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-acetyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	593.3	
II-244	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-bromo-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	645.2	
II-245	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	645.2	
II-246	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-5-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	645.2	
II-247	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-bromo-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	645.2	
II-248	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3,5-bis-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	687.2	
II-249	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	569.3	
II-250	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-fluoro-4-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	637.3	
II-251	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	579.3	
II-252	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	619.3	
II-253	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-5-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound	661.2	

	with trifluoro-acetic acid		
II-254	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	599.3	
II-255	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	594.3	
II-256	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-iodo-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	677.2	
II-257	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-ethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	579.3	
II-258	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-[3-(5-{4-[(2-hydroxy-ethyl)-methyl-amino]-benzoyl}-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-propyl]-amide; compound with trifluoro-acetic acid	624.3	
II-259	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(6-fluoro-4*H*-benzo[1,3]dioxine-8-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	627.2	
II-260	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-dibromo-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	709	
II-261	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methoxy-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	595.3	
II-262	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-fluoro-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	599.2	
II-263	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-methoxy-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	595.2	
II-264	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-fluoro-2-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	637.2	
II-265	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(cinnoline-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	603.2	
II-266	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-	612.2	

	phenyl)-{3-[5-(2,6-dimethoxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid		
II-267	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-trifluoromethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	620.2	
II-268	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	566.2	
II-269	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-methyl-pyridine-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	566.2	
II-270	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-pyrimidin-2-yl-amide; compound with trifluoro-acetic acid	533	
II-271	5-Oxo-1-(2,2,2-trifluoro-ethyl)-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	619	68.8-72.9
II-272	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(6-chloro-2-dimethylamino-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	629.1	
II-273	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-cyano-2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	610.1	
II-274	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-iodo-2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	711	
II-275	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	585.1	
II-276	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,6-dimethoxy-pyridazine-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	613.1	
II-277	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-bromo-2,5-di-*tert*-butyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	749.1	
II-278	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-di-*tert*-butyl-4-cyano-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	694.2	

II-279	3-(4-Chloro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-1-phenyl-urea; compound with trifluoro-acetic acid	532	
II-280	1,1-Dioxo-hexahydro-1λ^6-thiopyran-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	586	101.3 -102.7
II-281	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-2-hydroxy-propyl}-amide	596	97.3-98.1
II-282	Cyclopentanecarboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-cyclohexyl}-phenyl-amide	515	108.7 -109.3
II-283	Cyclopentanecarboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-phenyl-amide; compound with trifluoroacetic acid	475	130.3 -130.7
II-284	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-chloro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	599.1	
II-285	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-chloro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	599.1	
II-286	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-chloro-3-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	653.1	
II-287	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-bromo-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	661	
II-288	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-chloro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	599.1	
II-289	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-chloro-pyridine-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	586.1	
II-290	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methyl-2-methylsulfanyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	613.1	
II-291	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-4-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	663.1	
II-292	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-3-	603.1	

	fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid		
II-293	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-bromo-2-chloro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	665	
II-294	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-cyano-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	576.1	
II-295	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,2,3,3-tetramethyl-cyclopropanecarbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	571.2	
II-296	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-ethyl-butyryl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	545.2	
II-297	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-4-fluoro-benzenesulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	639	
II-298	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-cyano-benzenesulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	612.1	
II-299	{1-[4-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-piperidine-1-carbonyl]-2-methyl-propyl}-carbamic acid *tert*-butyl ester; compound with trifluoro-acetic acid	738.2	
II-300	[4-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-piperidin-1-yl]-oxo-acetic acid methyl ester; compound with trifluoro-acetic acid	625.1	
II-301	4-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-piperidine-1-carboxylic acid methyl ester; compound with trifluoro-acetic acid	597.1	
II-302	1-(2-Amino-3-methyl-butyryl)-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	638.2	
II-303	1-(2-Methoxy-ethyl)-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	597.2	
II-304	[4-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-	611.1	

	yl]-propyl}-carbamoyl)-piperidin-1-yl]-oxo-acetic acid; compound with trifluoro-acetic acid		
II-305	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(6-chloro-2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	614	oil
II-306	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-6-methylsulfanyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	626	oil
II-307	Cyclopentanecarboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(4-trifluoromethyl-pyrimidin-2-yl)-amide	544	
II-308	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-cyano-2-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	594	
II-309	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-dimethylamino-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	595	
II-310	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-dimethylamino-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	595.1	
II-311	Cyclopentanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-2-methyl-propyl}-amide	536	
II-312	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-3a,6a-dimethyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	609; 611	
II-313	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-3a,6a-dimethyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	624; 626 (M+Na) 646; 648	
II-314	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-dimethylamino-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	569	
II-315	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-dimethylamino-2-methylsulfanyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	642	
II-316	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-dimethylamino-pyridine-3-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	630.8	
II-317	[4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-	658.8	

	methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-3-chloro-phenoxy]-acetic acid; compound with trifluoro-acetic acid		
II-318	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-bromo-2-methylsulfanyl-pyrimidine-4-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid	678	
II-319	1-Dimethylsulfamoyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	646.3	
II-320	1-(2,2,2-Trifluoro-acetyl)-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	635	
II-321	1-Dimethylsulfamoyl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	618	
II-322	1-(2,2,2-Trifluoro-acetyl)-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	607	
II-323	1-Methanesulfonyl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	589	
II-324	Cyclopentanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-2-hydroxy-propyl}-amide; compound with trifluoro-acetic acid	538	
II-325	4-Oxo-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	550	
II-326	[5-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-4,6-dimethyl-pyridin-2-ylsulfanyl]-acetic acid; compound with trifluoro-acetic acid	670	
II-327	Cyclopentanecarboxylic acid {3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-fluoro-phenyl)-amide; compound with trifluoro-acetic acid	494	
II-328	4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-fluoro-phenyl)-amide	544	
II-329	3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-pyrrolidine-1-carboxylic acid *tert*-butyl ester	625	

II-330	1-Acetyl-piperidine-4-carboxylic acid {2-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-ethyl}-(3-fluoro-phenyl)-amide	537	
II-331	2-Cyclopentyl-*N*-{2-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-ethyl}-*N*-(3-fluoro-phenyl)-acetamide	494	
II-332	*N*-{2-[5-(4,6-Dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-ethyl}-*N*-(3-fluoro-phenyl)-propionamide	440	
II-333	*N*-{2-[5-(4,6-Dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-ethyl}-*N*-(3-fluoro-phenyl)-3-methyl-butyramide	468	
II-334	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-*p*-tolyl-amide	547	
II-335	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-*p*-tolyl-amide	562	
II-336	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(6-dimethylamino-2-ethoxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	639	
II-337	1,1-Dioxo-hexahydro-1λ^6-thiopyran-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	587	
II-338	5-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-3,4-dihydro-1*H*-isoquinoline-2-carboxylic acid *tert*-butyl ester	706	
II-339	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1,2,3,4-tetrahydro-isoquinoline-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	606	
II-340	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-acetyl-1,2,3,4-tetrahydro-isoquinoline-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide	648	
II-341	4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-3,5-dimethyl-1*H*-pyrrole-2-carboxylic acid ethyl ester	640	
II-342	[3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-pyrrolidin-1-yl]-oxo-acetic acid methyl ester	611	
II-343	5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carboxylic acid methyl-phenyl-amide; compound with trifluoro-acetic acid	580	

II-344	5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carboxylic acid benzylamide; compound with trifluoro-acetic acid	580	
II-345	1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-3-methyl-3-phenyl-urea; compound with trifluoro-acetic acid	561	
II-346	1-Acetyl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	553	
II-347	1-Propionyl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	567	
II-348	{2-[3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-azetidin-1-yl]-2-oxo-ethoxy}-acetic acid; compound with trifluoro-acetic acid	627	
II-349	4-[3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-azetidin-1-yl]-4-oxo-butyric acid; compound with trifluoro-acetic acid	611	
II-350	3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-azetidine-1-carboxylic acid methyl ester; compound with trifluoro-acetic acid	569	
II-351	[3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-azetidin-1-yl]-oxo-acetic acid methyl ester; compound with trifluoro-acetic acid	597	
II-352	1-Cyclopropanecarbonyl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	579	
II-353	1-Butyryl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	581	
II-354	1-(2-Methoxy-acetyl)-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	583	
II-355	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-2,2-dimethyl-propyl}-amide	609	
II-356	4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-	676	

	c]pyrrole-2-sulfonyl)-3,5-dimethyl-1*H*-pyrrole-2-carboxylic acid ethyl ester		
II-357	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-azetidin-1-yl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide	607	
II-358	4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-1,3,5-trimethyl-1*H*-pyrrole-2-carboxylic acid ethyl ester	654	
II-359	5-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-1,2,4-trimethyl-1*H*-pyrrole-3-carboxylic acid ethyl ester	654	
II-360	4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-3,5-dimethyl-1*H*-pyrrole-2-carboxylic acid	612	
II-361	1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,6-bis-dimethylamino-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide	638	
II-362	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-1-phenyl-1*H*-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	667	
II-363	4-Oxo-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	552	
II-364	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-pyridine-3-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	602	
II-365	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-methyl-benzofuran-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	605	
II-366	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(7-methoxy-3-methyl-1*H*-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	634	
II-367	Tetrahydro-pyran-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	540	
II-368	*N*-(3-Chloro-4-methyl-phenyl)-*N*-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-acetamide; compound with trifluoro-acetic acid	470	
II-369	Cyclopropanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with	496	

	trifluoro-acetic acid		
II-370	1-Acetyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	567	
II-371	1-(2,2,2-Trifluoro-acetyl)-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	621	
II-372	1-Methanesulfonyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	603	
II-373	1-Cyclopropanecarbonyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	593	
II-374	3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-pyrrolidine-1-carboxylic acid methyl ester; compound with trifluoro-acetic acid	583	
II-375	1-Dimethylsulfamoyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	632	
II-376	[3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-pyrrolidin-1-yl]-oxo-acetic acid; compound with trifluoro-acetic acid	597	
II-377	2-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-sulfonyl)-benzoic acid methyl ester; compound with trifluoro-acetic acid	645	
II-378	2-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-sulfonyl)-benzoic acid; compound with trifluoro-acetic acid	631	
II-379	1-Methanesulfonyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	617	
II-380	Pyrrolidine-1,3-dicarboxylic acid 3-((3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide) 1-dimethylamide; compound with trifluoro-acetic acid	596	
II-381	1-Cyclopropanesulfonyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	629	
II-382	1-Isobutyryl-pyrrolidine-3-carboxylic acid (3-chloro-4-	595	

	methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid		
II-383	1-Propionyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	581	
II-384	1-(2,2,2-Trifluoro-ethyl)-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	607	
II-385	4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-sulfonyl)-3,5-dimethyl-1*H*-pyrrole-2-carboxylic acid	648.3	
II-386	8-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-3,4-dihydro-1*H*-isoquinoline-2-carboxylic acid *tert*-butyl ester	706	
II-387	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-methyl-1*H*-imidazole-2-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	555	
II-388	4-Hydroxy-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	554	
II-389	1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1,2,3,4-tetrahydro-isoquinoline-8-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide	606	
II-390	1-Acetyl-piperidine-4-carboxylic acid (5-chloro-thiazol-2-yl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	572	
II-391	3-[4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl)-3,5-dimethyl-pyrazol-1-yl]-benzoic acid methyl ester	703	
II-392	1-Acetyl-piperidine-4-carboxylic acid (5-chloro-thiazol-2-yl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid	572	
II-393	3-(3-(3-Chloro-4-methyl-phenyl)-3-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-ureido)-benzoic acid; compound with trifluoro-acetic acid	591	
II-394	3-(3-(3-Chloro-4-methyl-phenyl)-3-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-ureido)-benzoic acid ethyl ester; compound with trifluoro-acetic acid	619	

II-395	1. The observed mass spectra are (M)+ or (M+H)+ and are consistent with the compound		

DOSAGE AND ADMINISTRATION

The compounds of the present invention may be formulated in a wide variety of oral administration dosage forms and carriers. Oral administration can be in the form of tablets, coated tablets, hard and soft gelatin capsules, solutions, emulsions, syrups, or suspensions. Compounds of the present invention are efficacious when administered by other routes of administration including continuous (intravenous drip) topical parenteral, intramuscular, intravenous, subcutaneous, transdermal (which may include a penetration enhancement agent), buccal, nasal, inhalation and suppository administration, among other routes of administration. The preferred manner of administration is generally oral using a convenient daily dosing regimen which can be adjusted according to the degree of affliction and the patient's response to the active ingredient.

A compound or compounds of the present invention, as well as their pharmaceutically useable salts, together with one or more conventional excipients, carriers, or diluents, may be placed into the form of pharmaceutical compositions and unit dosages. The pharmaceutical compositions and unit dosage forms may be comprised of conventional ingredients in conventional proportions, with or without additional active compounds or principles, and the unit dosage forms may contain any suitable effective amount of the active ingredient commensurate with the intended daily dosage range to be employed. The pharmaceutical compositions may be employed as solids, such as tablets or filled capsules, semisolids, powders, sustained release formulations, or liquids such as solutions, suspensions, emulsions, elixirs, or filled capsules for oral use; or in the form of suppositories for rectal or vaginal administration; or in the form of sterile injectable solutions for parenteral use. A typical preparation will contain from about 5% to about 95% active compound or compounds (w/w). The term "preparation" or "dosage form"is intended to include both solid and liquid formulations of the active compound and one skilled in the art will appreciate that an active ingredient can exist in different preparations depending on the target organ or tissue and on the desired dose and pharmacokinetic parameters.

The term “excipient” as used herein refers to a compound that is useful in preparing a pharmaceutical composition, generally safe, non-toxic and neither biologically nor otherwise undesirable. The term “excipient” as used herein includes both one and more than one such excipient.

The phrase "pharmaceutically acceptable salt" of a compound means a salt that is pharmaceutically acceptable and that possesses the desired pharmacological activity of the parent compound. Such salts include: (1) acid addition salts, formed with inorganic acids such as hydrochloric acid, hydrobromic acid, sulfuric acid, nitric acid, phosphoric acid, and the like; or formed with organic acids such as acetic acid, propionic acid, hexanoic acid, cyclopentanepropionic acid, glycolic acid, pyruvic acid, lactic acid, malonic acid, succinic acid, malic acid, maleic acid, fumaric acid, tartaric acid, citric acid, benzoic acid, 3-(4-hydroxybenzoyl)benzoic acid, cinnamic acid, mandelic acid, methanesulfonic acid, ethanesulfonic acid, 1,2-ethane-disulfonic acid, 2-hydroxyethanesulfonic acid, benzenesulfonic acid, 4-chlorobenzenesulfonic acid, 2-naphthalenesulfonic acid, 4-toluenesulfonic acid, camphorsulfonic acid, 4-methylbicyclo[2.2.2]-oct-2-ene-1-carboxylic acid, glucoheptonic acid, 3-phenylpropionic acid, trimethylacetic acid, tertiary butylacetic acid, lauryl sulfuric acid, gluconic acid, glutamic acid, hydroxynaphthoic acid, salicylic acid, stearic acid, muconic acid, and the like; or (2) salts formed when an acidic proton present in the parent compound either is replaced by a metal ion, *e.g.*, an alkali metal ion, an alkaline earth ion, or an aluminum ion; or coordinates with an organic base such as ethanolamine, diethanolamine, triethanolamine, tromethamine, N-methylglucamine, and the like.

The preferred pharmaceutically acceptable salts are the salts formed from acetic acid, hydrochloric acid, sulphuric acid, methanesulfonic acid, maleic acid, phosphoric acid, tartaric acid, citric acid, sodium, potassium, calcium, zinc, and magnesium. It should be understood that all references to pharmaceutically acceptable salts include solvent addition forms (solvates) or crystal forms (polymorphs) as defined herein, of the same acid addition salt.

Solid form preparations include powders, tablets, pills, capsules, cachets, suppositories, and dispersible granules. A solid carrier may be one or more substances which may also act as diluents, flavoring agents, solubilizers, lubricants, suspending agents, binders, preservatives, tablet disintegrating agents, or an encapsulating material. In powders, the carrier generally is a finely divided solid which is a mixture with the finely divided active component. In tablets, the active component generally is mixed with the carrier having the necessary binding capacity in suitable proportions and compacted in the shape and size desired. Suitable carriers include but are not limited to magnesium carbonate, magnesium stearate, talc, sugar, lactose, pectin, dextrin, starch, gelatin, tragacanth, methylcellulose, sodium carboxymethylcellulose, a low melting wax, cocoa butter, and the like. Solid form preparations may contain, in addition to the active component, colorants, flavors, stabilizers, buffers, artificial and natural sweeteners, dispersants, thickeners, solubilizing agents, and the like.

Liquid formulations also are suitable for oral administration include liquid formulation including emulsions, syrups, elixirs, aqueous solutions, aqueous suspensions. These include solid form preparations which are intended to be converted to liquid form preparations shortly before use. Emulsions may be prepared in solutions, for example, in aqueous propylene glycol solutions or may contain emulsifying agents such as lecithin, sorbitan monooleate, or acacia. Aqueous solutions can be prepared by dissolving the active component in water and adding suitable colorants, flavors, stabilizing, and thickening agents. Aqueous suspensions can be prepared by dispersing the finely divided active component in water with viscous material, such as natural or synthetic gums, resins, methylcellulose, sodium carboxymethylcellulose, and other well known suspending agents.

The compounds of the present invention may be formulated for parenteral administration (*e.g.*, by injection, for example bolus injection or continuous infusion) and may be presented in unit dose form in ampoules, pre-filled syringes, small volume infusion or in multi-dose containers with an added preservative. The compositions may take such forms as suspensions, solutions, or emulsions in oily or aqueous vehicles, for example solutions in aqueous polyethylene glycol. Examples of oily or nonaqueous carriers, diluents, solvents or vehicles include propylene glycol, polyethylene glycol, vegetable oils (*e.g.*, olive oil), and injectable organic esters (*e.g.*, ethyl oleate), and may contain formulatory agents such as preserving, wetting, emulsifying or suspending, stabilizing and/or dispersing agents. Alternatively, the active ingredient may be in powder form, obtained by aseptic isolation of sterile solid or by lyophilisation from solution for constitution before use with a suitable vehicle, *e.g.*, sterile, pyrogen-free water.

The compounds of the present invention may be formulated for topical administration to the epidermis as ointments, creams or lotions, or as a transdermal patch. Ointments and creams may, for example, be formulated with an aqueous or oily base with the addition of suitable thickening and/or gelling agents. Lotions may be formulated with an aqueous or oily base and will in general also containing one or more emulsifying agents, stabilizing agents, dispersing agents, suspending agents, thickening agents, or coloring agents. Formulations suitable for topical administration in the mouth include lozenges comprising active agents in a flavored base, usually sucrose and acacia or tragacanth; pastilles comprising the active ingredient in an inert base such as gelatin and glycerin or sucrose and acacia; and mouthwashes comprising the active ingredient in a suitable liquid carrier.

The compounds of the present invention may be formulated for administration as suppositories. A low melting wax, such as a mixture of fatty acid glycerides or cocoa butter is first melted and the active component is dispersed homogeneously, for example,

by stirring. The molten homogeneous mixture is then poured into convenient sized molds, allowed to cool, and to solidify.

The compounds of the present invention may be formulated for vaginal administration. Pessaries, tampons, creams, gels, pastes, foams or sprays containing in addition to the active ingredient such carriers as are known in the art to be appropriate.

The compounds of the present invention may be formulated for nasal administration. The solutions or suspensions are applied directly to the nasal cavity by conventional means, for example, with a dropper, pipette or spray. The formulations may be provided in a single or multidose form. In the latter case of a dropper or pipette, this may be achieved by the patient administering an appropriate, predetermined volume of the solution or suspension. In the case of a spray, this may be achieved for example by means of a metering atomizing spray pump.

The compounds of the present invention may be formulated for aerosol administration, particularly to the respiratory tract and including intranasal administration. The compound will generally have a small particle size for example of the order of five (5) microns or less. Such a particle size may be obtained by means known in the art, for example by micronization. The active ingredient is provided in a pressurized pack with a suitable propellant such as a chlorofluorocarbon (CFC), for example, dichlorodifluoromethane, trichlorofluoromethane, or dichlorotetrafluoroethane, or carbon dioxide or other suitable gas. The aerosol may conveniently also contain a surfactant such as lecithin. The dose of drug may be controlled by a metered valve. Alternatively the active ingredients may be provided in a form of a dry powder, for example a powder mix of the compound in a suitable powder base such as lactose, starch, starch derivatives such as hydroxypropylmethyl cellulose and polyvinylpyrrolidine (PVP). The powder carrier will form a gel in the nasal cavity. The powder composition may be presented in unit dose form for example in capsules or cartridges of *e.g.*, gelatin or blister packs from which the powder may be administered by means of an inhaler.

When desired, formulations can be prepared with enteric coatings adapted for sustained or controlled release administration of the active ingredient. For example, the compounds of the present invention can be formulated in transdermal or subcutaneous drug delivery devices. These delivery systems are advantageous when sustained release of the compound is necessary and when patient compliance with a treatment regimen is crucial. Compounds in transdermal delivery systems are frequently attached to an skin-adhesive solid support. The compound of interest can also be combined with a penetration enhancer, e.g., Azone (1-dodecylaza-cycloheptan-2-one). Sustained release delivery systems are inserted subcutaneously into to the subdermal layer by surgery or

injection. The subdermal implants encapsulate the compound in a lipid soluble membrane, e.g., silicone rubber, or a biodegradable polymer, e.g., polyactic acid.

Suitable formulations along with pharmaceutical carriers, diluents and expcipients are described in *Remington: The Science and Practice of Pharmacy* 1995, edited by E. W. Martin, Mack Publishing Company, 19th edition, Easton, Pennsylvania. A skilled formulation scientist may modify the formulations within the teachings of the specification to provide numerous formulations for a particular route of administration without rendering the compositions of the present invention unstable or compromising their therapeutic activity.

The modification of the present compounds to render them more soluble in water or other vehicle, for example, may be easily accomplished by minor modifications (salt formulation, esterification, etc.), which are well within the ordinary skill in the art. It is also well within the ordinary skill of the art to modify the route of administration and dosage regimen of a particular compound in order to manage the pharmacokinetics of the present compounds for maximum beneficial effect in patients.

The term "therapeutically effective amount" as used herein means an amount required to reduce symptoms of the disease in an individual. The dose will be adjusted to the individual requirements in each particular case. That dosage can vary within wide limits depending upon numerous factors such as the severity of the disease to be treated, the age and general health condition of the patient, other medicaments with which the patient is being treated, the route and form of administration and the preferences and experience of the medical practitioner involved. For oral administration, a daily dosage of between about 0.01 and about 100 mg/kg body weight per day should be appropriate in monotherapy and/or in combination therapy. A preferred daily dosage is between about 0.1 and about 500 mg/kg body weight, more preferred 0.1 and about 100 mg/kg body weight and most preferred 1.0 and about 10 mg/kg body weight per day. Thus, for administration to a 70 kg person, the dosage range would be about 7 mg to 0.7 g per day. The daily dosage can be administered as a single dosage or in divided dosages, typically between 1 and 5 dosages per day. Generally, treatment is initiated with smaller dosages which are less than the optimum dose of the compound. Thereafter, the dosage is increased by small increments until the optimum effect for the individual patient is reached. One of ordinary skill in treating diseases described herein will be able, without undue experimentation and in reliance on personal knowledge, experience and the disclosures of this application, to ascertain a therapeutically effective amount of the compounds of the present invention for a given disease and patient.

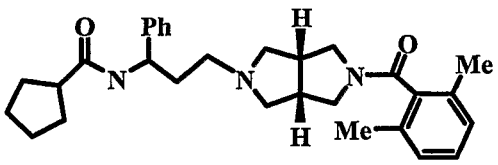
In embodiments of the invention, the active compound or a salt can be administered in combination with another antiviral agent, such as a nucleoside reverse transcriptase inhibitor, another nonnucleoside reverse transcriptase inhibitor, HIV protease inhibitor and/or a viral fusion inhibitor. When the active compound or its derivative or salt are administered in combination with another antiviral agent the activity may be increased over the parent compound. When the treatment is combination therapy, such administration may be concurrent or sequential with respect to that of the nucleoside derivatives. "Concurrent administration" as used herein thus includes administration of the agents at the same time or at different times. Administration of two or more agents at the same time can be achieved by a single formulation containing two or more active ingredients or by substantially simultaneous administration of two or more dosage forms with a single active agent.

Furthermore, treatment of a HIV infection, as used herein, also includes treatment or prophylaxis of a disease or a condition associated with or mediated by HIV infection, or the clinical symptoms thereof.

The pharmaceutical preparations are preferably in unit dosage forms. In such form, the preparation is subdivided into unit doses containing appropriate quantities of the active component. The unit dosage form can be a packaged preparation, the package containing discrete quantities of preparation, such as packeted tablets, capsules, and powders in vials or ampoules. Also, the unit dosage form can be a capsule, tablet, cachet, or lozenge itself, or it can be the appropriate number of any of these in packaged form.

Example 1

Cyclopentanecarboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide (I-3; Scheme 1)



step 1 - A solution of 2-benzyl-octahydro-pyrrolo[3,4-c]pyrrole (4a,1.22g, 4.97 mmol) in DCM (20 mL) was added to a solution of cyclopentanecarboxylic acid (3-oxo-1-phenyl-propyl)-amide (10, 1.0 g, 4.97 mmol) in toluene (8 mL) containing HOAc (0.34 mL). Sodium triacetoxyborohydride (1.26 g, 5.96 mmol) was added in two portion separated by 0.5 h and reaction stirred for 18 h at room temperature. The reaction was quenched by addition a solution 0.5 M of KOH (30 mL) and product extracted with DCM (3x50 mL). The combined organic extracts dried ($MgSO_4$) and concentrated *in*

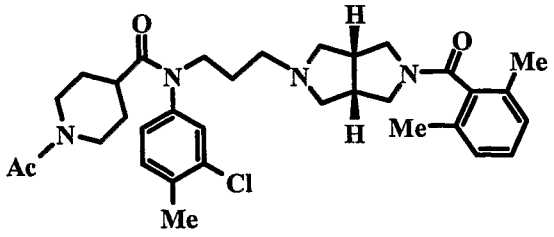
vacuo. The crude product was purified by flash chromatography on silica gel eluting with 4% (10% ammonium hydroxide in methanol) in DCM to afford 11 as a viscous liquid (0.62g, 29% theory):^{1}H NMR ($CDCl_3$) δ 1.46-1.64 (m, 2H), 1.65-1.94 (m, 7H), 1.96-2.11 (m, 1H), 2.23-2.44 (m, 4H), 2.45-2.83 (m, 9H), 3.59 (q, 2H, *J*= 18.2, 12.8 Hz), 5.11(q, 1H, *J*= 6.2, 5.8 Hz), 7.07-7.39 (m, 10H), 7.93 (d, 1H, *J*= 7.1 Hz); MS (ES+) m/z 432 $(M + H)^+$.

step 2 - Palladium hydroxide (0.2 g) and ammonium formate (2.92 g, 0.046 mol) were added to a solution of cyclopentanecarboxylic acid [3-(5-benzyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide (11, 2.0 g, 0.004 mol) in EtOH (50 mL). The solution was heated at reflux for 2 h and filtered through a CELITE® pad. The resulting solution was concentrated *in vacuo* and crude product purified by flash column chromatography on silica gel (CH_2Cl_2: MeOH: NH_4OH=60:10:1) to afford 12: ^{1}H NMR (DMSO-d_6) δ 1.4-1.8 (m, 11H), 2.15-2.42 (m, 6H), 2.55-2.65 (m, 2H), 2.75-2.85 (m, 2H), 4.33 (q, 1H), 7.15-7.35 (m, 5H), 8.2 (dd, 1H) ; ^{13}C NMR (DMSO-d_6) δ 25.96, 26.00, 30.15, 30.45, 35.60, 39.07, 39.35, 39.62, 39.90, 40.18, 40.46, 40.74, 41.37, 43.43, 44.61, 51.02, 52.14, 54.02, 60.20, 60.30, 126.64, 126.82, 128.51, 144.46, 174.81; MS (ES+) m/z 341 $(M + H)^+$;

step 3 - To a solution of cyclopentanecarboxylic acid [3-(hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide (12 0.20 g, 0.585 mmol) in dichloromethane at rt was added 2,6- dimethylbenzoic acid was added to a solution. To the resulting solution was added sequentially EDCI (0.14 g, 0.760 mmol), HOBt (0.10 g, 0.760 mmol) and diisopropylethylamine (0.3 mL, 1.755 mmol) were added to the mixture. The mixture was stirred overnight at rt. The reaction mixture was washed with 5% $NaHCO_3$ solution and dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by flash column chromatography on silica gel (5% MeOH/CH_2Cl_2) to afford 13: mp 60.9-63.9 °C; ^{1}H NMR (DMSO-d_6) δ 1.4-1.9 (m, 10H), 2.1 (s, 3H), 2.2 (s, 3H), 2.3-2.5 (m, 5H), 3.35 (m, 1H), 3.5 (dd, 1H), 3.75 (m, 1H), 4.88 (m, 1H), 7.05 (d, 2H), 7.15-7.35 (m, 6H), 8.0 (d, 1H); ^{13}C NMR (DMSO-d_6) δ 18.85, 18.98, 25.96, 26.00, 30.12, 30.42, 35.73, 39.05, 39.33, 39.60, 39.88, 41.69, 44.56, 50.49, 51.05, 52.37, 53.11, 60.37, 126.64, 126.67, 126.89, 127.53, 127.61, 128.31, 128.52, 133.05, 138.25, 144.39, 167.72, 174.81; MS (ES+) m/z 473 $(M + H)^+$; Anal. ($C_{30}H_{39}N_3O_3$. 0.15M CH_2Cl_2) C; calcd, 74.45; found, 74.55; H; calcd, 8.14; found, 7.74; N; calcd, 8.64; found, 8.95;

Example 2

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl} (II-1)



step 1 - 1-chloro-3-iodopropane (3.0 mL, 28.49 mmol) and cesium carbonate (18.0 g, 56.49 mol) were added to a solution of 2-chloro-4-aminotoluene (4 g, 28.49 mmol) in DMF (8 mL). The mixture was stirred overnight at RT. Water (15 mL) was added to the mixture and the resulting solution was extracted with hexane. The organic layer was washed with water again and dried ($MgSO_4$). The crude product was purified by flash column chromatography on silica gel (5% EtOAc/hexane) to afford **20**: ^1H NMR ($CHCl_3$) δ 1.5 (s, 1H), 2.05 (m, 2H), 2.25 (s, 3H), 3.3 (t, 2H), 3.65 (t, 2H), 6.4 (dd, 1H), 6.65 (d, 1H), 7.0 (d, 1H)

step 2 - A solution of (3-chloro-4-methyl-phenyl)-(3-chloro-propyl)-amine (**20**, 3.20 g, 14.70 mmol) in CH_2Cl_2 (100 mL) was cooled to 0° C. After the mixture was treated with triethylamine (4.9 mL, 35.28 mmol)and 1-acetyl-piperidine-4-carbonyl chloride (5.57 g, 29.40 mmol), it was allowed to stir for 5 h at 0 °C. Saturated $NaHCO_3$ was added to the mixure and the mixture was stirred for 20 min. The solvent was evaporated and the mixture was extracted with EtOAc. The organic layer was washed with 2N HCl and brine, dried ($MgSO_4$) and purified by flash column chromatography on silica gel (5% MeOH/EtOAc) to afford **18**:

^1H NMR ($CHCl_3$) δ 1.55-1.8 (br, 4H), 2.0 (m, 3H), 2.05 (s, 3H), 2.3-2.4 (m, 2H), 2.45 (s, 3H), 3.52 (t, 2H), 3.3 (t, 2H), 6.97 (dd, 1H), 7.18 (t, 1H), 7.32 (d, 1H); ^{13}C NMR ($CHCl_3$) δ 20.19, 21.68, 28.94, 31.17, 39.64, 42.64, 48.03, 126.63, 128.79, 132.47, 135.79, 137.12, 141.27, 169.27, 174.58; ms (ES+) m/z 370 (M + H).

step 3 - To a solution of solution of 2-benzyl-octahydro-pyrrolo[3,4-c]pyrrole (**4b**, 1.03g, 5.11 mmol) in DCM (30 mL) at rt was added 2,6- dimethylbenzoic acid (1.53 g, 10.22 mmol), BOP-Cl (2.60 g, 10.22 mmol), DMAP (1.24 g, 10.22 mmol) and Et_3N (1.5 mL, 10.22 mmol). The reaction mixture was stirred overnight at RT and washed with 5% $NaHCO_3$ and brine. The resulting was dried ($MgSO_4$), filtered and evaporated *in vacuo*. The crude product was purified by flash chromatography on silica gel (CH_2Cl_2: MeOH: NH_4OH / 4000:10:1) to afford **16**: ms (ES+) m/z = 235 $(M + H)^+$;.

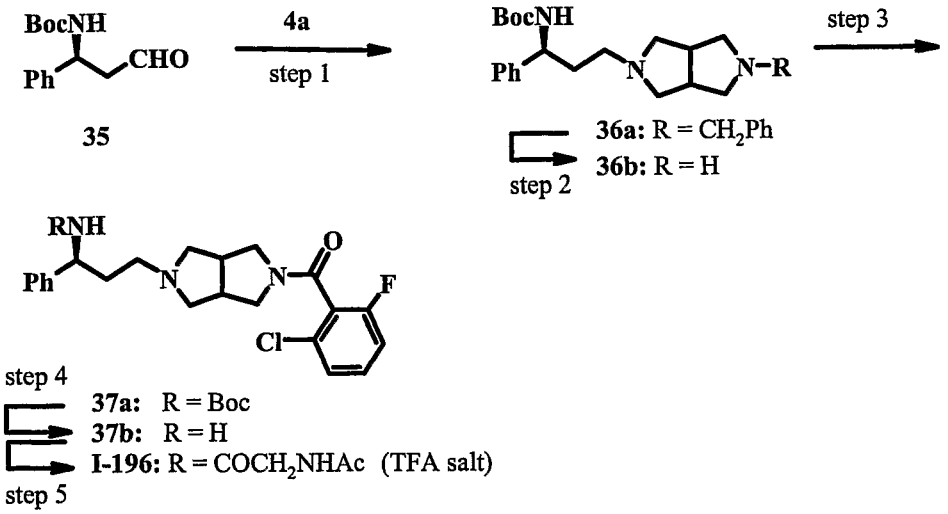
step 4 - Palladium hydroxide (0.13 g) and ammonium formate (2.45 g, 0.038 mol) were added to a solution of (5-benzyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-(2,6-dimethyl-phenyl)-methanone (**16**, 1.30 g, 3.88 mmol) in EtOH (30 mL). The solution

was heated at reflux for 2 h and filtered through a CELITE® pad. The volatile solvents were removed by evaporation *in vacuo* and the crude reaction product was purified by flash column chromatography on silica gel (CH_2Cl_2: MeOH: NH_4OH / 120:10:1) to afford 66: ^1H NMR ($CHCl_3$) δ 2.13 (s, 1H), 2.25 (s, 3H), 2.28 (s, 3H), 2.65 (dd, 1H), 2.2-2.95 (m, 3H), 3.08-3.25 (m, 2H), 3.35 (dd, 1H), 3.65 (dd, 1H), 3.85-3.95 (m, 1H), 7.0 (m, 2H), 7.15 (q, 1H); ms (ES+) m/z 245 (M + H).

step 5 - 1-Acetyl-piperidine-4-carboxylic acid(3-chloro-4-methyl-phenyl)-(3-chloro-propyl)-amide (17, 0.22 g, 0.61 mmol) and (2,6-dimethyl-phenyl)-(hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-methanone (18, 0.15 g, 0.61 mmol) were dissolved in MeCN (15 mL). K_2CO_3 (0.25 g, 1.84 mmol) and potassium iodide (0.11 g, 0.67 mmol) were added to the mixture and the resulting solution was heated at reflux overnight. The mixture was cooled, diluted with water and extracted with EtOAc. The combined EtOAc extracts were washed with brine and dried ($MgSO_4$) and evaporated. The crude product was purified by flash column chromatography on silica gel (5% MeOH/CH_2Cl_2) to afford 19: mp 58.1-65.2 °C; ^1H NMR (DMSO-d_6) δ 1.5-1.65(m, 6H), 1.95 (s, 3H), 2.12 (d, 6H), 2.23-2.52 (m, 12H), 2.68-2.85 (m, 4H), 3.2-3.25 (m, 1H), 3.45 (dd, 1H), 3.6-3.7 (m, 3H), 7.05 (d, 2H), 7.1-7.2 (m, 2H), 7.35 (d, 1H), 7.43 (d, 1H) ; ^{13}C NMR (DMSO-d_6) δ 18.84, 18.93, 19.60, 21.53, 26.64, 28.37, 29.04, 39.05, 39.33, 39.61, 39.88, 41.63, 45.23, 47.46, 50.62, 52.37, 53.13, 60.18, 60.60, 127.42, 127.52, 127.57, 128.29, 128.62, 132.43, 132.96, 133.05, 134.13, 135.64, 138.24, 141.63, 167.65, 168.18, 173.47; ms (ES+) m/z 578 (M + H); Anal. ($C_{33}H_{43}ClN_4O_3$-0.2M CH_2Cl_2) C; calcd, 66.89; found, 66.70; H; calcd, 7.34; found, 7.10; N; calcd, 9.40; found, 9.50.

Example 3

2-Acetylamino-N-{3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide (I-196)



step 1 - A solution of **4a** (6.25g , 31 mmol) in DCM (200 mL) was added to **35** (1.0 g, 4.97 mmol). To the resulting mixture was added $NaBH(OAc)_3$ (9.86 g, 47 mmol) in two portions and the reaction stirred for 18 h at RT. The reaction was washed with a solution 5% $NaHCO_3$ (100 mL). The organic phase was dried (Na_2SO_4) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica gel eluting with DCM/ 3.5% MeOH (containing 10% NH_4OH) to afford 6.8 g (46%) of **36a** as a viscous liquid: MS (ES+) m/z 436 $(M + H)^+$.

step 2 - Palladium hydroxide (0.2 g) and ammonium formate (6.3 g, 0.100 mol) were added to a solution of **36a** (4.36 g, 0.010 mol) in EtOH (200 mL). The solution was heated at reflux for 2 h and filtered through a CELITE® pad. The resulting solution was concentrated *in vacuo* to afford to afford 3.5 g (100%) of**36b**: ms (ES+) m/z 346 $(M + H)^+$.

step 3 - To a solution of **36b** (3.5 g, 10.0 mmol) in DMF (100 ml) at RT was added 2-chloro-6-fluorobenzoic acid (1.75g, 10.0 mmol). To the resulting solution were added sequentially EDCI (2.0 g, 10.0 mmol), HOBt (1.35 g, 10.0 mmol) and $NaHCO_3$ (3.4g, 40.0 mmol) and the mixture was stirred overnight at RT. The solvent was evaporated under reduced pressure and the residue was dissolved in DCM (100 ml). The organics were washed with 2% HCl, water, saturated. $NaHCO_3$ solution, dried (Na_2SO_4) and concentrated *in vacuo*. The crude product was purified by flash column chromatography on silica gel (3.5% MeOH/DCM) to afford 3.3 g (66%) of**37a**: MS (ES+) m/z 502 $(M + H)^+$.

step 4 - To a solution of **37a** (3.3 g, 6.6 mmol) in DCM (50 ml) was added a solution of ethereal HCl (100 mL, 1 N in ether). The reaction stirred 18 h. The solution was evaporated under reduced pressure to afford 3.6 g (110%) of **37b** HCl as a white solid: MS (ES+) m/z 401 $(M + H)^+$.

step 5 - Acetylamino-acetic acid (6.1 mg,52.5 μmol) was weighed into the reaction vessel and resin-bound carbodimide (78 mg, 2.0 equiv) was added. HOBt (60 μmol, 1.7 equiv. in 10% DMF in DCM, 1.0 ml) was added and the reactions was shaken for 1h. A solution of 37b HCl in DCM (35 μmol, 500 μl, with 30 μl DIEA) was added. The reaction was shaken for 72 h. The resin was filtered, washed with DCM (3 x 1.0 mL) and the solvent was evaporated under reduced pressure. The residue was purified by reverse-phase preparative HPLC to afford I-196: ms (ES+) m/z 501 $(M+H)^+$.

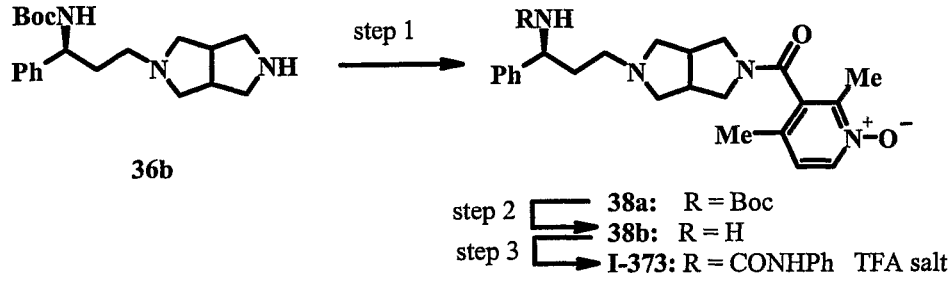
Example 4

N-{3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-phenyl-propionamide (I-312)

I-312 was synthesized in the same manner as example 3 except in step 5, acetylamino-acetic acid was replaced with 2-methylphenyl acetic acid to afford I-312 which was isolated as the TFA salt: ms (ES+) m/z 534 $(M+H)^+$.

Example 5

1-{3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-phenyl-urea (I-373)



step 1 - To a solution of 36b (0.456 g, 1.1 mmol) in DMF (10 ml) at RT was added 2,4-dimethyl-3-pyridine carboxylic acid N-oxide (0.184 g, 1.1 mmol). To the resulting solution were added sequentially EDCI (0.221 g, 1.2 mmol), HOBt (0.149 g, 1.1 mmol) and $NaHCO_3$ (0.370g, 4.4 mmol). The mixture was stirred overnight at RT then evaporated under reduced pressure. The residue was dissolved in DCM (25ml). The organics were washed with 2% HCl, water, saturated. $NaHCO_3$ solution,dried (Na_2SO_4) and concentrated *in vacuo*. The crude product was purified by flash column chromatography on silica gel (3.5% $MeOH/CH_2Cl_2$) to afford 0.360 g (66%) of 38a: ms (ES+) m/z 495 $(M + H)^+$.

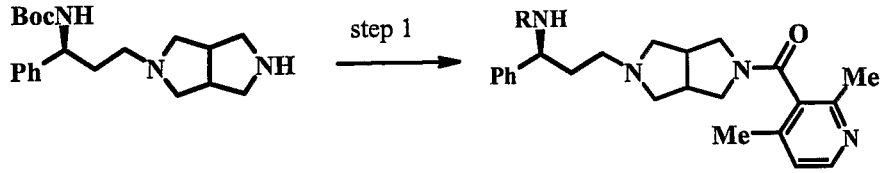
step 2 - To a solution of 38a (0.36 g, 0.73 mmol) in DCM (20 ml) was added a solution of HCl (1 N in ether, 20 ml). The reaction stirred 18 h. The solution was

evaporated under reduced pressure to afford 0.31 g (100%) of 38b HCl as a white solid: ms (ES+) m/z 395 $(M + H)^+$.

step 3 - Phenyl isocyanate (8.1 mg, 75 μmol) was weighed into the reaction vessel and 38b HCl (50 μmol) in DCM (500 μl containing 60 μl DIEA) was added. The reaction was shaken for 18 h. The solvent was evaporated under reduced pressure. The residue was purified by reverse-phase preparative HPLC to afford I-373: ms (ES+) m/z 514 $(M + H)^+$.

Example 6

2-Cyclohexyl-N-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide, TFA salt (I-378)



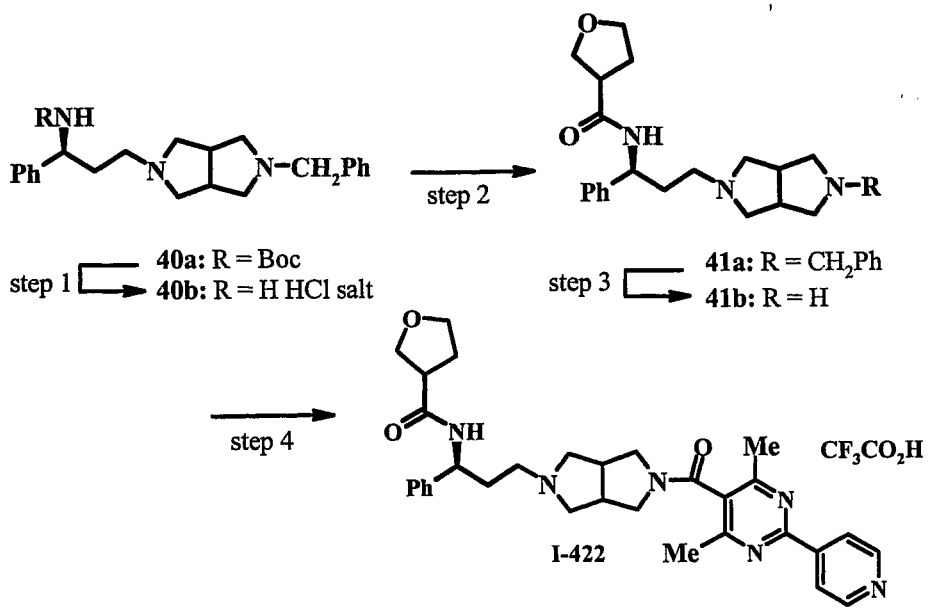
step 1 - Prepared as described in step 1 of Example 5 except 2,4-dimethyl-3-pyridine carboxylic acid N-oxide was replaced with 2,4-dimethyl-3-pyridine carboxylic acid to afford 0.643 g (61%) of **39a**: ms (ES+) m/z 479 $(M + H)^+$.

step 2 - Prepared as described in step 2 of Example 5 except 39a from step 1 was used to afford 0.790 g (100%) of 39b HCl salt: ms (ES+) m/z 379 $(M + H)^+$.

step 3 - Cyclohexylacetic acid (10.6 mg, 75 μmol) was weighed into the reaction vessel and 39b.2HCl (50 μmol) in DCM (500 μl containing with 60 μl DIPEA) was added. The reaction was shaken for 18 h and the solvent was evaporated under reduced pressure. The residue was purified by reverse-phase preparative HPLC to afford I-378 as a TFA salt: ms (ES+) m/z 503 $(M + H)^+$.

Example 7

Tetrahydro-furan-3-carboxylic acid {3-[5-(4,6-dimethyl-2-pyridin-4-yl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide, TFA salt (I-422)



step 1 - To a solution of 40a (5.85 g, 13.4 mmol) in DCM (50 ml) was added a solution of HCl (1 N in ether, 50 ml) and the reaction stirred 18 h. The solution was evaporated under reduced pressure to afford 6.3 g (100%) **40b** HCl as a white solid: ms (ES+) m/z 336 $(M + H)^+$.

step 2 - To a solution of 40b HCl (6.3 g, 13.4 mmol) in DMF (50 ml) at RT was added tetrahydro-3-furoic acid (1.29 ml, 13.4 mmol). To the resulting solution were added sequentially EDCI (2.7 g, 14.1 mmol), HOBt (1.81 g, 13.4 mmol) and $NaHCO_3$ (4.5g, 53.6 mmol). The mixture was stirred overnight at RT. The solvent was evaporated under reduced pressure and the residue was dissolved in DCM (25ml). The organic phase was washed with 2% HCl, water, saturated $NaHCO_3$ solution, dried (Na_2SO_4) and concentrated *in vacuo*. The crude product was purified by flash column chromatography on silica gel (3.5% MeOH/DCM) to afford 5.23 g (90% theory) of 41a: ms (ES+) m/z 434 $(M + H)^+$.

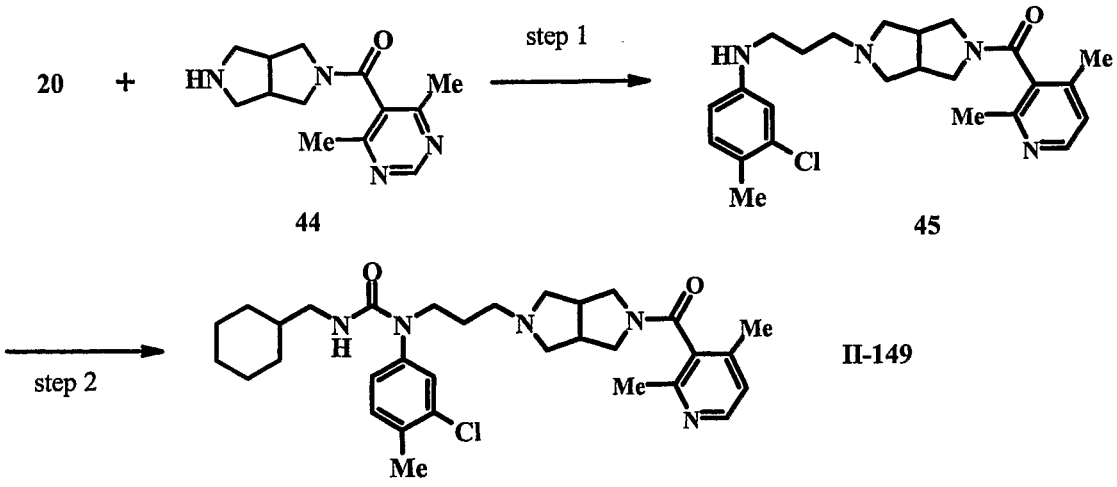
step 3 - Palladium hydroxide (0.2 g) and ammonium formate (3.0 g, 46 mmol) were added to a solution of 41a (2.0 g, 4.6 mmol) in EtOH (100 mL). The solution was heated to reflux for 5 h and filtered through a CELITE® pad. The resulting solution was concentrated in vacuo to afford 1.40 g (89%) of 41b: ms (ES+) m/z 344 $(M + H)^+$.

step 3 - 4,6-Dimethyl-2-pyridin-4-yl-pyrimidine-5-carboxylic acid (18.7 mg, 75 µmol, prepared as described in T. Gebhard *et al. J. Med. Chem.* 2004 47(8):1939-1955 and WO2002081449 A1) was weighed into a reaction vessel and resin-bound carbodimide (120 mg, 2.0 equiv) was added. HOBt (85 µmol, in 10% DMF in DCM, 1.0 ml) was added. The reaction was shaken for 18 h. A solution of 41b (50 µmol) in DCM (500 µl containing 60 µl DIEA) was added. The reaction shakin continued for 18 h. The resin

was filtered and washed with DMF (1.0 ml) and DCM (2 x 1.0 mL). The residue was purified by reverse-phase preparative HPLC to afford I-422: ms (ES+) m/z 555 $(M + H)^+$.

Example 8

1-(3-Chloro-4-methyl-phenyl)-3-cyclohexylmethyl-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea, TFA salt (II-149)

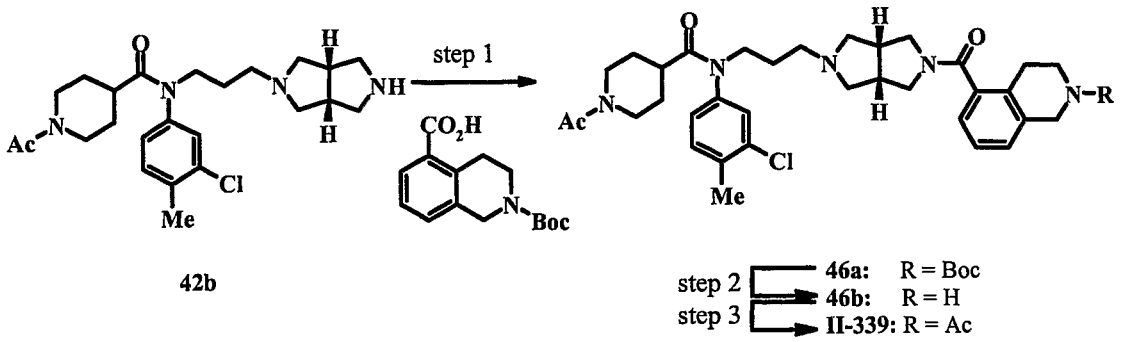


step 1 - To a solution of 44 (0.90 g, 3.7 mmol) in MeCN (80 ml) were added sequentially 20 (0.887 mg, 4.1 mmol), KI (0.681 g, 4.1 mmol) and DIPEA (1.3 mL, 7.4 mmol). The reaction mixture was heated at 80° C for 18h. The reaction was quenched with water and thrice extracted with EtOAc. The organic phase was dried (Na_2SO_4), filtered and evaporated. The crude product was purified by flash column chromatography on silica gel eluting with 3.5% MeOH/DCM (containing 0.4% ammonia) to afford 1.06 g (67%) of 45: ms (ES+) m/z 426 $(M + H)^+$.

step 2 - A flask was charged with cyclohexaneaminemethyl isocyanate (7.5 mg, 52.5 μmol) and 45 (35 μmol) in DCM (500 μL) was added. The reaction was shaken for 18 h. The residue was purified by reverse-phase preparative HPLC to II-149: ms (ES+) m/z 566 $(M + H)^+$.

Example 9

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-acetyl-1,2,3,4-tetrahydro-isoquinoline-5-carbonyl)hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide (II-339)



1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-[3-(hexahydro-pyrrolo[3,4-c]pyrrol-2-yl) -propyl]-amide (**42b**) was prepared by the procedures described in Example 2 except **4a** was first alkylated with **18** as described in step 4 and the benzyl protecting group was removed by catalytic hydrogenolysis as described in step 3.

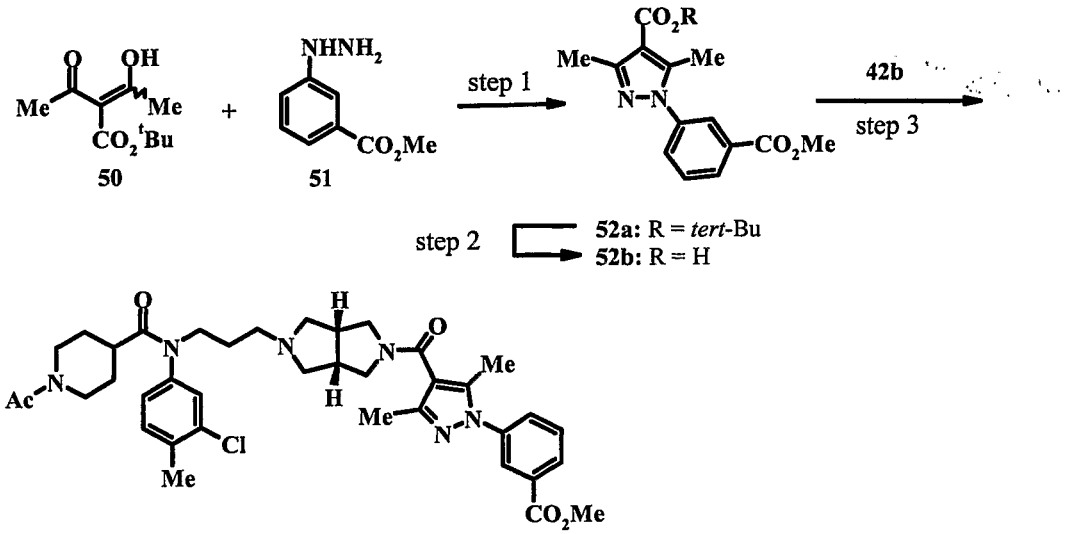
step 1 - To a solution of **42b** (0.1 g, 0.224 mmol), boc-5-hydroxycarbonyl-1,2,3,4-tetrahydroisoquinoline (62 mg, 0.224 mmol, Arch Corp. catalog # AR02230), EDCI (51 mg, 0.268 mmol) and HOBT (41 mg, 0.268 mmol) in DMF (2 mL) at RT was added DIPEA (80 □L, 0.336 mmol). The resulting mixture was stirred at RT for 16 h then partitioned between EtOAc and water. The aqueous layer was back extracted twice with EtOAc. The combined organic layers were dried (Na_2SO_4), filtered and evaporated. The residue was purified by SiO_2 chromatography eluting with a gradient of 100% DCM (2 min isocratic run time) to a 1:1 mixture DCM of a solution of DCM/MeOH/NH_4OH (80/10/1) over 20 minutes at a 25 mL/min flow rate to afford 0.142 g (90%) of **46a** as a light pink foam: ^{13}C NMR ($CDCl_3$) δ 20.19, 21.77, 26.62, 27.20, 28.74, 28.89, 29.56, 39.78, 41.09, 41.28, 42.47, 45.93, 48.47, 50.99, 53.00, 54.34, 60.27, 60.75, 80.31, 124.34, 126.71, 128.88, 131.62, 132.39, 135.64, 136.88, 141.53, 155.20, 168.99, 169.15, 174.30 ; MS (ES+) m/z 706 $(M + H)^+$.

step 2 - To a solution of **46a** (0.11 g, 0.156 mmol) in DCM (2.5 mL)at RT was added TFA (0.5 mL). The resulting solution was stirred at RT for 2 h then evaporated. The residue was purified by SiO_2 chromatography eluting with a gradient of 100% DCM to 100% of a solution of DCM/MeOH/NH_4OH (80/10/1) over 20 minutes then maintained this composition for another 10 min at a flow rate of 25 mL/min which afforded 0.060 g (64%) of **46b**: ^1H NMR (CD_3OD) δ ^{1}1.51-1.78 (m, 6H), 2.30-2.56 (m, 11H), 2.61 (dd, 1H, *J*= 9, 8 Hz), 2.69-2.98 (m, 6H), 3.00-3.14 (m, 3H), 3.42 (dd, 1H, *J*= 12, 6 Hz), 3.58 (dd, 1H, *J*= 15, 3 Hz), 3.63-3.93 (m, 5H), 4.02 (m, 2H), 4.43 (bd, 1H, *J*= 12 Hz), 7.03-7.26 (m, 4H), 7.37-7.47 (m, 2H); ^{13}C NMR (CD_3OD) δ 20.19, 21.54, 26.90, 28.05, 29.78, 30.37, 41.28, 42.34, 42.54, 43.52, 44.29, 47.16, 50.27, 52.30, 54.21, 55.72, 61.91, 125.38, 127.75, 128.40, 128.93, 130.11, 132.08, 133.77, 137.16, 138.32, 138.85, 171.40, 171.83 ; MS (ES+) m/z 606 $(M + H)^+$.

step 3 - A solution of **46b** (30 mg, 49.5 □mol), pyridine (0.5 mL) and Ac_2O (0.5 mL) was stirred at RT for 16 h then diluted with MeOH (2 mL). The resulting mixture was stirred at RT for 1 h then evaporated and co-evaporated with toluene. The residue was purified by SiO_2 chromatography eluting with a gradient of 100% DCM to a 1:1 solution of DCM and DCM/MeOH/NH_4OH (80/10/1) over 20 minutes at a flow rate of 25 mL/min which afforded 22 mg (66%) **II-339**; ^1H NMR (CD_3OD) δ 1.51-1.78 (m, 6H), 2.05 (s, 3H), 2.19 and 2.16 (2s, 3H), 2.28-2.55 (m, 8H), 2.60 (m, 1H), 2.73-2.96 (m, 5H), 3.04 (m, 1H), 3.43 (m, 1H), 3.55-3.90 (m, 7H), 4.43 (bd, 1H, *J*= 12 Hz), 4.71 (d, 2H, *J*= 15 Hz), 7.16 (m, 2H), 7.29 (m, 2H), 7.42 (m, 2H), two proton signals are hidden under the deuterated methanol signals; ^{13}C NMR (CD_3OD) δ 19.77, 21.12, 21.42, 21.66, 26.81, 27.59, 29.37, 29.95, 40.62, 40.85, 41.92, 42.08, 43.13, 44.82, 45.16, 46.74, 52.00, 53.78, 55.42, 61.05, 61.51, 125.37, 125.52, 127.90, 128.00, 128.46, 128.71, 129.71, 133.35, 135.46, 135.81, 136.21, 138.01, 142.38, 170.66, 171.42, 172.21, 176.17 ; MS (ES+) m/z 648 $(M + H)^+$.

Example 10

3-[4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-3,5-dimethyl-pyrazol-1-yl]-benzoic acid methyl ester (II-390)



II-390

step 1 - To a suspension of 3-hydrazinobenzoic acid (1.52 g, 9.99 mmol) (CAS # 38235-71-1) in water (20 mL) and HOAc (20 mL) at RT was added **50** (2 g, 9.99 mmol, prepared according to the procedure described in *Org. Prep. Proc. Int.* 2001 34(4):357-409). The resulting mixture was stirred at RT for 30 min before being poured into 400 mL of a 1:1 mixture of iced cold water and EtOAc. The aqueous layer was extracted once with EtOAc. The combined organic layers were washed with saturated aqueous $NaHCO_3$ until

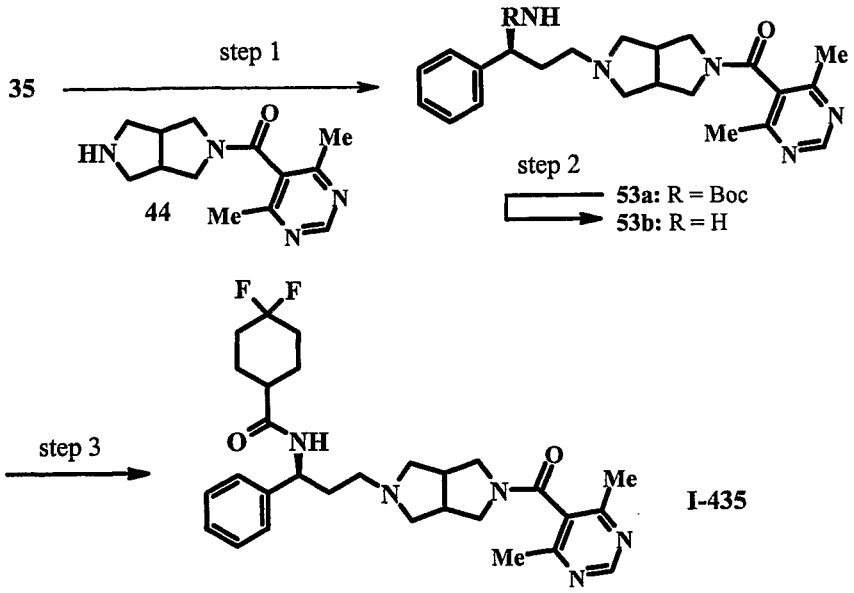
the aqueous extract remained basic. The organic layers were dried (Na_2SO_4), filtered and evaporated. The residue was used directly in without further purification or characterization. To the residue dissolved in DCM (100 mL) was added trimethylsilyldiazomethane (5 mL of 2M in hexane solution, 10 mmol) at RT. The resulting mixture was stirred at RT for 30 min then quenched by adding methanol. The mixture was stirred at RT for 30 min then evaporated. The residue was purified by SiO_2 chromatography eluting with a hexane/EtOAc gradient (100% hexane for 2 minutes then a gradient to 20% EtOAc over 20 minutes which was maintained for an additional 10 minutes) at a flow rate of 55 mL/min to afford 1.09 g (33%) of **52a**: ^{1}H NMR (CDCl3) δ 1.59 (s, 9H), 2.48 (s, 3H), 2.52 (s, 3H), 3.94 (s, 3H), 7.54-7.64 (m, 1H), 8.07 (m, 1H), 8.09 (m, 1H) ; MS (ES+) m/z 331 (M + H).

step 2 - To a solution of **52a** (1.088 g, 3.293 mmol) in DCM (50 mL) and triethylsilane (5 mL) was added TFA (5 mL). The resulting mixture was stirred at RT for 48 h then evaporated to afford 0.9 g (99% theory) of **52b**: ^{1}H NMR (DMSO-d_6) δ 2.27 (s, 3H), 2.39 (s, 3H), 3.78 (s, 3H), 7.58 (m, 1H), 7.71 (m, 1H), 7.87-7.95 (m, 2H) ; MS (ES+) m/z 502 $(M + H)^+$.

step 3 - Diisopropylethylamine was added at RT To a solution of **42b** (75 mg, 0.168 mmol), **52b** (51 mg, 0.185 mmol), EDCI (39 mg, 0.201 mmol), HOBT (31 mg, 0.201 mmol) and DMF (1.5 mL) was added DIPEA(0.15 mL, 0.839 mmol). The resulting mixture was stirred at RT for 16 h then partitioned between EtOAc and water. The aqueous layer was extracted twice with EtOAc. The combined extracts were dried (Na_2SO_4), filtered and evaporated. The residue was purified by SiO_2 chromatography eluting with a gradient of 100% DCM (1 min isocratic) to a 1:1 solution of DCM and DCM/MeOH/NH_4OH (80/10/1) over 20 minutes followed by 2:8 for 10 minat a flow rate of 25 mL/min which afforded 66 mg (%^% theory) of **II-390**: ^{1}H NMR (CD_3OD) δ 1.45-1.78 (m, 6H), 2.04 (s, 3H), 2.27 (s, 3H), 2.30 (s, 3H), 2.31-2.53 (m, 9H), 2.53-2.73 (m, 2H), 2.77-2.97 (m, 3H), 3.58-3.78 (m, 6H), 3.84 (bd, 1H, *J*= 10 Hz), 3.94 (s, 3H), 4.42 (bd, 1H, *J*= 12 Hz), 7.16 (dd, 1H, *J*= 6, 2 Hz), 7.38 (1H, *J*= 2 Hz), 7.41 (d, 1H, *J*= 6 Hz), 7.67 (m, 1H), 7.75 (m, 1H), 8.11 (m, 2H); ^{13}C NMR (CD_3OD) δ 12.26, 12.99, 20.18, 24.54, 28.02, 29.76, 30.35, 41.25, 42.32, 43.54, 47.14, 50.28, 52.22, 53.43, 54.22, 55.24, 61.41, 61.90, 118.72, 127.55, 128.39, 130.09, 130.69, 131.13, 131.30, 133.24, 133.75, 136.62, 138.31, 140.79, 140.93, 142.75, 149.05, 166.81, 167.77, 171.82, 176.56 ; MS (ES+) m/z 703 $(M + H)^+$.

Example 11

4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide (I-435)



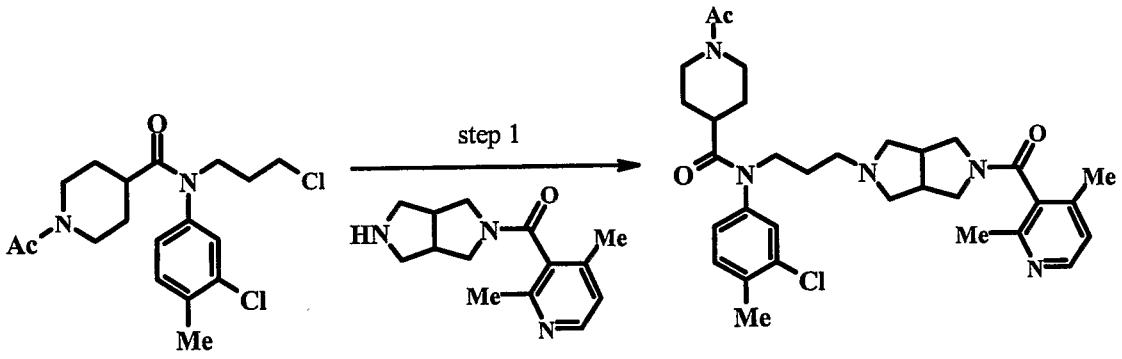
step1 - To the mixture of 35 (0.20 g, 0.82 mmol, CAS 135865-78-0) and 44 (0.18 g, 0.75 mmol) in DCM (10 mL) was added sodium triacetoxyborohydride (0.24g, 1.12 mmol) and the resulting solution was stirred at RT for 3 h. The mixture was diluted with DCM and washed with saturated $NaHCO_3$. The organic layer was dried ($MgSO_4$), filtered and evaporated. The crude product and purified by SiO_2 column chromatography eluting with DCM:MeOH:NH_4OH (140:10:1) to afford 0.29 g (80 %) of 53a: ms (ES+) m/z 480 (M + H).

step 2 - Methanolic HCl (10mL, 1.25 M) was added to 53a (0.29 g, 0.60 mmol) and the mixture was heated at 50° C for 2h. The mixture was concentrated *in vacuo* to afford 53b which used in the next step without additional without purification:

step3 - To a solution of 4,4-difluoro-cyclohexanecarboxylic acid (0.13 g, 0.78 mmol) in toluene (5 mL) was added thionyl chloride (66 µL, 0.90 mmol). It was heated to reflux for 2h. The solvent and the excess of thionyl chloride were evaporated and the residual acid chloride was diluted with DCM (4 mL) and toluene (2 mL). The acid chloride solution was added to a mixture of **53b** in saturated Na_2CO_3 (5 mL) and H_2O (3 mL) and stirred at RT for 4 h. The reaction mixture was diluted with H_2O and extracted with DCM. The organic layer was dried ($MgSO_4$), filtered evaporated. The crude product was purified by SiO_2 column chromatography eluting with DCM:MeOH:NH_4OH (120:10:1) afford 0.16 g (50 %) of I-435: mp 131.8-133.2 °C; ms (ES+) m/z 526 (M + H);

Example 12

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide (II-3)

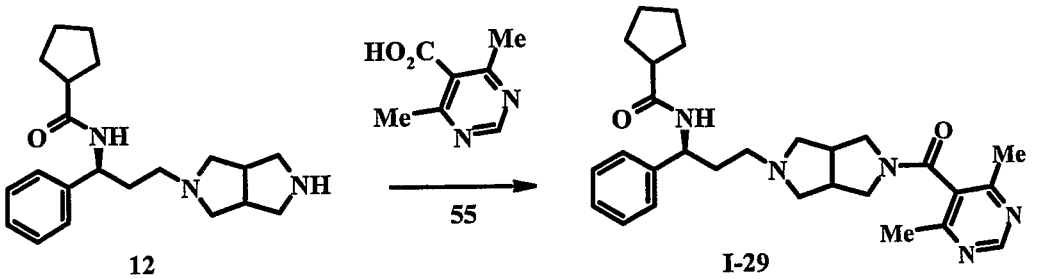


18 54 II-3

To a solution of 18 (0.35 g, 1.15 mmol), 54 (0.18 g, 0.76 mmol) and MeCN (10 mL) were added K_2CO_3 (0.31 g, 2.30 mmol) and KI (0.25 g, 1.53 mmol) and the resulting solution was heated at reflux overnight. The mixture was cooled, diluted with water and extracted with EtOAc. The combined EtOAc extracts were washed with brine and dried ($MgSO_4$) and evaporated. The crude product was purified by SiO_2 column chromatography eluting with DCM:MeOH:NH_4OH (150:10:1) to afford 0.25 g (57 %) of II-3:: mp 81.5-83.6 °C; ^1H NMR (DMSO-d_6) δ 1.3-1.6(m, 8H), 1.95 (s, 3H), 2.15 (s, 3H), 2.3 (s, 3H), 2.35 (s, 3H), 2.65-2.85 (m, 5H), 3.2-3.3 (m, 1H), 3.4-3.8 (m, 6H), 4.2-4.3 (d, 1H), 7.1 (m, 1H), 7.15-7.25 (d, 1H), 7.45 (d, 1H), 8.3 (d, 1H) ; ms (ES+) m/z 580 (M + H); Anal. ($C_{32}H_{42}ClN_5O_3$. 0.3M CH_2Cl_2) C; calcd, 65.78; found, 65.74; H; calcd, 7.38; found, 7.30; N; calcd, 12.07; found, 11.58.

Example 13

Cyclopentanecarboxylic acid {3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide (I-29)



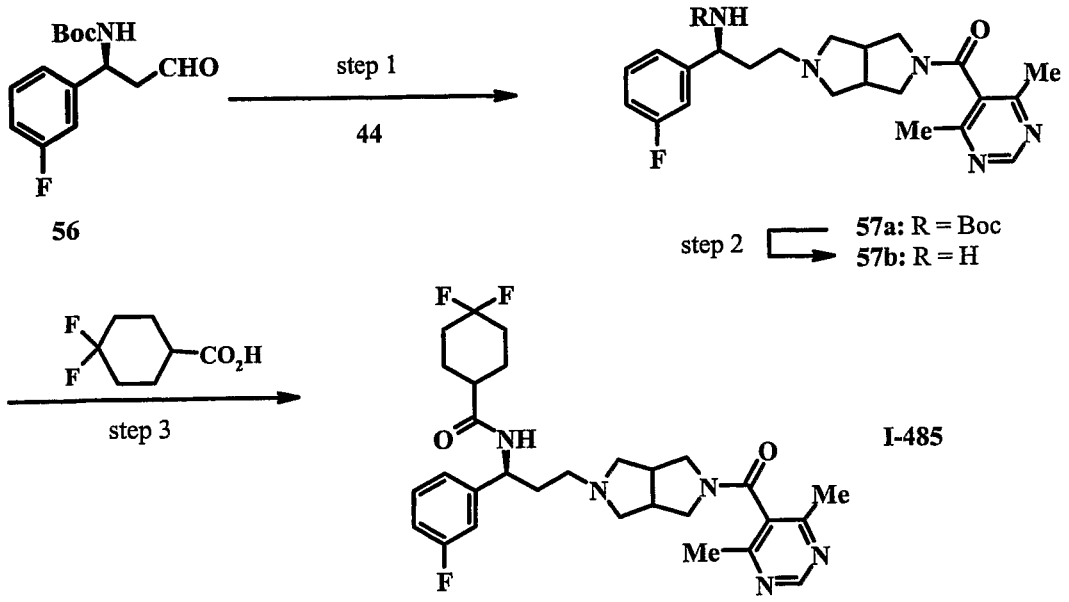
12 I-29

To a solution of 12 (0.24 g, 0.70 mmol) in DCM (10 mL) were added 4,6-dimethyl-pyrimidine-5-carboxylic acid (55, 0.12 g, 0.84 mmol) , EDCI (0.17 g, 0.91 mmol), HOBt (0.12 g, 0.91 mmol) and DIPEA (0.36 mL, 2.10 mmol). The mixture was stirred at RT for

3 h. The reaction mixture was washed with saturated $NaHCO_3$ and the organic layer was dried (Na_2SO_4). The crude product was purified by SiO_2 column chromatography eluting with DCM:MeOH:NH_4OH (150:10:1) to afford 0.27 g (81 %) of I-29:: mp 48.0-49.0 °C; ms (ES+) m/z 476 (M + H); Anal. ($C_{28}H_{37}N_5O_2$. 0.2M CH_2Cl_2) C; calcd, 68.76; found, 68.61; H; calcd, 7.65; found, 7.51; N; calcd, 14.22; found, 14.28

Example 14

(S)-4,4-Difluoro-cyclohexanecarboxylic acid [3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-amide (I-485)



step 1 - To a solution of 56 (562 mg, 2.1 mmol, prepared as described in WO2004/018425) and 44 (518 mg, 2.1 mmol) in DCM (20mL) containing HOAc (0.31 mL) was added $NaBH(OAc)_3$ (579 mg, 2.73 mmol) in 1 portion and the reaction mixture was stirred for 18 hrs at RT. The reaction was quenched by the addition of 10% K_2CO_3 (20 mL) and stirring continued for 30 min. The product was twice extracted with DCM (25 mL). The combined extracts were dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica eluting with DCM/ 5% MeOH (containing 2% NH_4OH) to afford 821 mg (79% theory) of 57a as a white foam: ms (ES+) m/z 498 $(M+H)^+$.

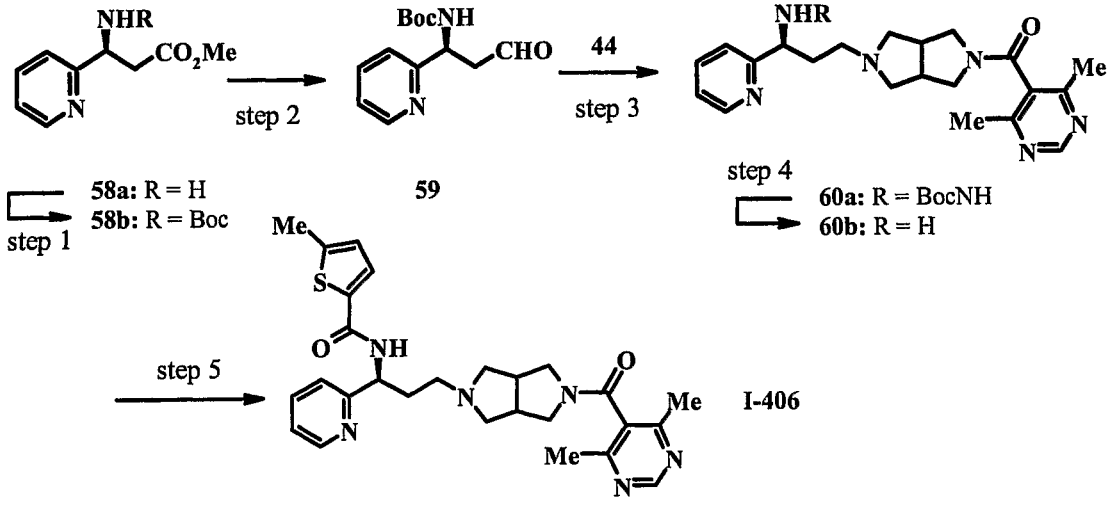
step 2 - A solution of 57a (821 mg, 1.65 mmol) dissolved in 10 M HCl in MeOH (40 mL) was heated at 65° C for 2 h. The MeOH was evaporated under reduced pressure and the residue cautiously partitioned between DCM (35 mL) and 20% K_2CO_3 solution. The aqueous layer was extracted with DCM (2 x 35 mL). The combined organic extracts

were dried (Na_2SO_4) and concentrated *in vacuo* to afford 641 mg (98%) of 57b as a viscous liquid: ms (ES+) m/z 398 $(M+H)^+$.

step 3 - To a solution of 57b (98 mg, 0.25 mmol) in DCM (4 mL) at RT was added 4,4-difluorocyclohexanecarboxylic acid (49 mg, 0.30mmol). To the resulting solution was added sequentially EDCI (61.4 mg, 0.32 mmol), HOBt (43 mg. 0.32 mmol) and DIPEA (0.13 mL, 0.74 mmol). The mixture was stirred for 4 h. The reaction mixture washed with brine and dried (Na_2SO_4), then concentrated *in vacuo*. The crude product was flash chromatographed on silica eluting with DCM/ 7.5% MeOH (containing 2% NH_4OH) to afford 113 mg (84%) of I-485 a white foam: ms (ES+) m/z 544 $(M+H)^+$.

Example 15

5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid (I-406)



step 1 - To a cold (0°) mixture of Na_2CO_3 (418 mg, 3.95 mmol) in THF:H_2O (2:1) was added in one portion (S)-3-amino-3-pyridin-2-yl-propionic acid methyl ester dihydrochloride (58a, 200mg, 0.79 mmol; prepared as described for the isomeric 3-pyridyl analog in *J. Org. Chem.* 2002 67:7819). A solution of $(BOC)_2O$ (189 mg, 0.86 mmol) in THF (2mL) was added in 1 portion and the reaction stirred at 0° C for 1 h, then at RT for 2 hrs. The product was extracted with EtOAc (2 x 25 mL). The combined organic extracts were dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica eluting with DCM/ 5.0% MeOH (containing 2% NH_4OH) to afford 157 mg (71%) of 58b as a viscous liquid: ms (ES+) m/z 281 $(M+H)^+$.

step 2 - To a soln of **58b** (561 mg, 2.0 mmol) in DCM (20 mL) cooled to -78° was added DIBAL-H (4.0 mL of a 1.0 M DCM solution, 4.0 mmol) dropwise at such a rate to maintain the temperature below -70° C. After 1 h the reaction was quenched by the addition of MeOH (5 mL) and H_2O (1 mL) at -78° C, then allowed to warm to RT. The mixture was filtered through a CELITE® pad. The filtrate was dried (Na_2SO_4) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica eluting with DCM/6.0% MeOH (containing 2% NH_4OH) to afford 416 mg (83%) of **59** as a viscous liquid; ms (ES+) m/z 251 $(M+H)^+$.

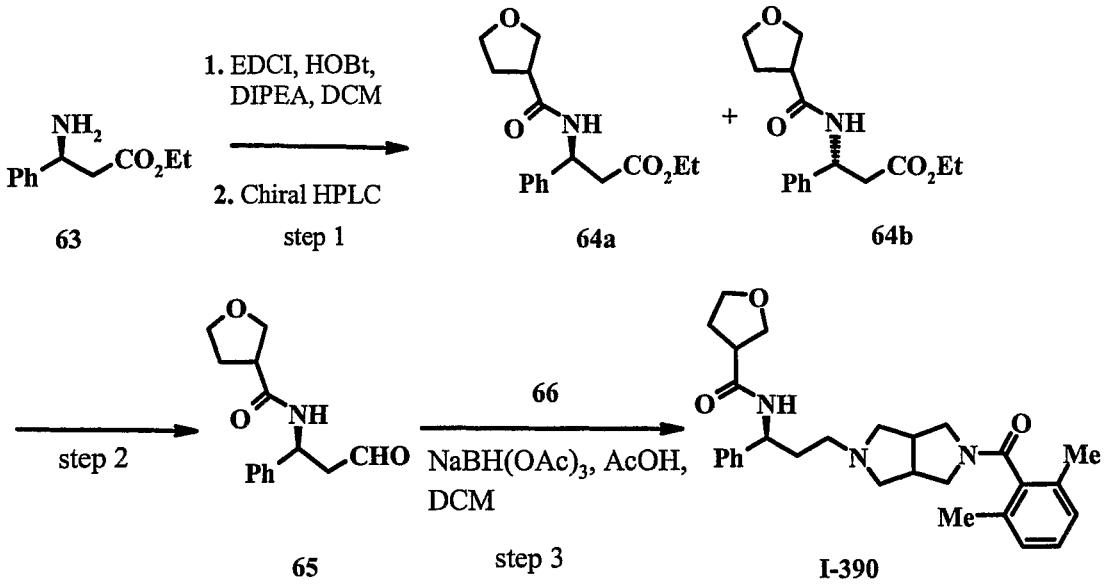
step 3 - To a solution of **59** (220 mg, 0.88 mmol) and **44** (215 mg, 0.88 mmol) in DCM (10 mL) containing HOAc (0.13 mL) was added $NaBH(OAc)_3$ (224 mg, 1.05 mmol) in 1 portion and the reaction was stirred for 5 h. The reaction was quenched by adding10% K_2CO_3 soln (10 mL) and stirred for an additional 30 min. The product was extracted with DCM (2 x 20 mL). The combined extracts were dried (Na_2SO_4) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica eluting with DCM/7.5% MeOH (containing 2% NH_4OH) to afford 304 mg (72%) of **60a** as a viscous liquid: ms (ES+) m/z 479 $(M+H)^+$.

step 4 - A solution of 60a (304 mg, 0.64 mmol) in 10 M HCl and MeOH (10 mL) heated at 65° C for 2 h. The MeOH was evaporated under reduced pressure and the residue cautiously partitioned between DCM (35 mL) and 20% K_2CO_3 solution (20 mL) . The aqueous layer was reextracted with DCM (2 x 35 mL). The combined organic extracts were dried (Na_2SO_4) and concentrated *in vacuo* to afford **60b** as a viscous liquid. (254 mg) which was used in the next step without further purification: ms (ES+) m/z 379 $(M+H)^+$

step 5 - To 5-methyl-2-thiophencarboxylic acid (10.7 mg, 0.075mmol) and HOBt (1.0 mL 0.06 M soln in DCM:DMF, 9:1) was added resin bound-carbodiimide (81 mg) and the mixture was stirred for 18 h. A solution of 60b (500 µL of 0.1 M soln in DCM) was added and stirring was continued for 18 h. The reaction mixture was concentrated *in vacuo* and purified by reverse phase HPLC to afford **I-406**: ms (ES+) m/z 503 $(M + H)^+$.

Example 16

(3S, 3'S)-Tetrahydro-furan-3-carboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide (**I-390**)



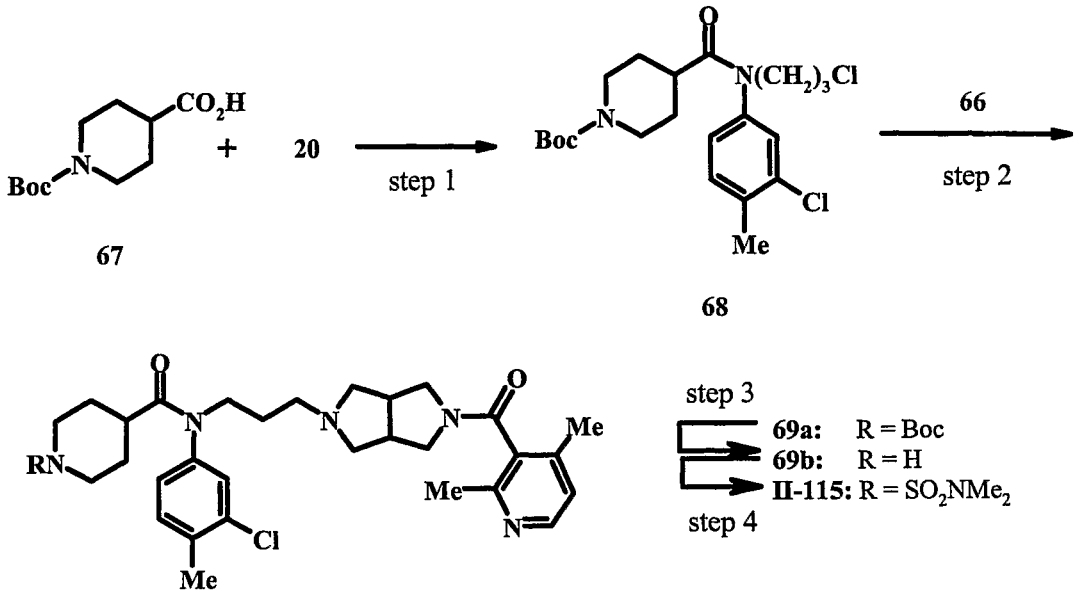
step 1 -To a mixture of tetrahydro-3-furoic acid (1.55 g, 13.34 mmol) and (S)-3-amino-3-phenylpropanoic acid ethyl ester hydrochloride (63, 3.06 g, 13.34 mmol) in 50 mL of DCM at RT were added sequentially EDCI (3.42 g, 16.01 mmol), HOBt (2.16 g, 16.01 mmol) and DIPEA (11.62 mL, 66.74 mmol). The mixture was stirred overnight at RT. The reaction mixture was washed with 5% $NaHCO_3$ solution, dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by preparative HPLC using a Chiralcel OD-H column, eluting with 20% 2-propanol/hexanes to afford 1.5 g of (3S, 3'S)-3-phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propionic acid ethyl ester(64a): retention time: 14:71 min, mp 84.6-85.9 °C, ms (ES+) m/z = 291 M^+.

step 2 – A DIBAL-H (1M solution in DCM, 7.5 mL, 7.5 mmol) was cooled to -78 °C and added drop-wise to a solution of 64a (1.1 g, 3.7 mmol) and DCM (20 mL) cooled to -78 °C with stirring. Stirring was continued for 2 h at -78° C, then 2N hydrochloric acid (1 mL)was added drop-wise at -78° C and mixture allowed to warm to RT. Additional 2N HCl (20 mL)was added, the layers were separated and aqueous phase thrice extracted with DCM (3x). Combined organic layers were dried ($MgSO_4$) and concentrated. The crude product was purified by SiO_2 chromatography eluting with EtOAc/DCM (0 to 100% EtOAc) to afford 0.33 g of 65: MS (ES+) m/z = 248 $(M + H)^+$.

Step 3 was carried out as described in step 3 of Example 14 to afford I-390: MS (ES+) m/z = 476 $(M + H)^+$.

Example 17

1-Dimethylsulfamoyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide (I-115)



step 1 - To a stirred mixture of Boc-isonipecotic acid (67, 2.12 g, 9.22 mmol), pyridine (1.9 mL, 23.6 mmol), and DCM (13 mL) under a nitrogen atmosphere and at RT was added $SOCl_2$ (0.8 mL, 11 mmol) was added. After 25 min, a solution of 20 (2.21 g, 10.16 mmol), TEA (4.5 mL, 32.33 mmol), and DMAP (0.12 g, 0.92 mmol) in DCM (16 mL) was added dropwise and stirring was continued for 72 h. 5% HCl (20 mL) was added dropwise, layers separated and aqueous layer thrice extracted with DCM. The combined organic layers were dried ($MgSO_4$) and concentrated. The crude product was purified by SiO_2 chromatography eluting with a EtOAc/hexanes gradient (0 to 30%) to afford 2.2 g of 68: ms (ES+) m/z = 451 $(M + Na)^+$.

step 2 - 4-To a solution of 68 (0.52 g, 1.2 mmol) and 66 (0.3 g, 1.2 mmol) dissolved in MeCN (30 mL) were added K_2CO_3 (0.33 g, 2.4 mmol) and potassium iodide (0.22 g, 1.32 mmol) and the resulting mixture was heated at reflux overnight. Additional 68 was added (0.2 g, 0.46 mmol) and heating continued for 6 h. A third aliquot of 68 was added (0.2 g, 0.46 mmol) and reaction continued for an additional 18 h. The mixture was cooled to RT, diluted with water and extracted with EtOAc. The combined EtOAc extracts were washed with brine, dried ($MgSO_4$) and evaporated. The crude product was purified by flash chromatography on silica gel eluting with MeOH containing 10% NH_4OH)/DCM (0 to 4%) to afford 0.244 g of **69a**: ms(ES+) m/z 637 $(M + H)^+$.

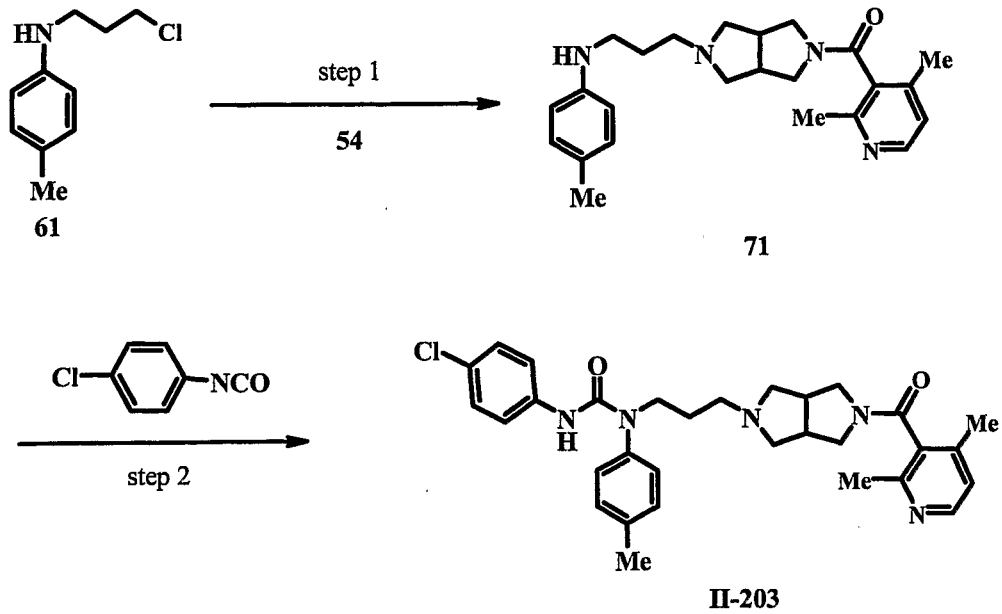
step 3 - To **69a** (0.24 g, 0.37 mmol) dissolved in DCM (9 mL) was added TFA (1 mL) and the mixture was stirred at RT overnight. The solvents were evaporated, and the residue suspended in toluene and re-evaporated (2x). The residue was purified by flash

chromatography on silica gel eluting with MeOH containing 10% NH_4OH)/DCM (0 to 7%) to afford 0.194 g of **69b**: MS(ES+) m/z 537 $(M + H)^+$.

step 4 - To a RT solution of **69b** (0.019 g, 0.035 mmol) and DIPEA (0.02 mL, 0.105 mmol) in 1 mL of DCM was added dimethylsulfamoyl chloride (0.01 g, 0.07 mmol) and the mixture stirred overnight at rt. The reaction mixture was concentrated *in vacuo* and purified by reverse phase HPLC to afford **II-115**: ms (ES+) m/z 644 $(M + H)^+$.

Example 18

3-(4-Chloro-phenyl)-1-{3-[(3aS,6aR)-5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-1-*p*-tolyl-urea (**II-203**



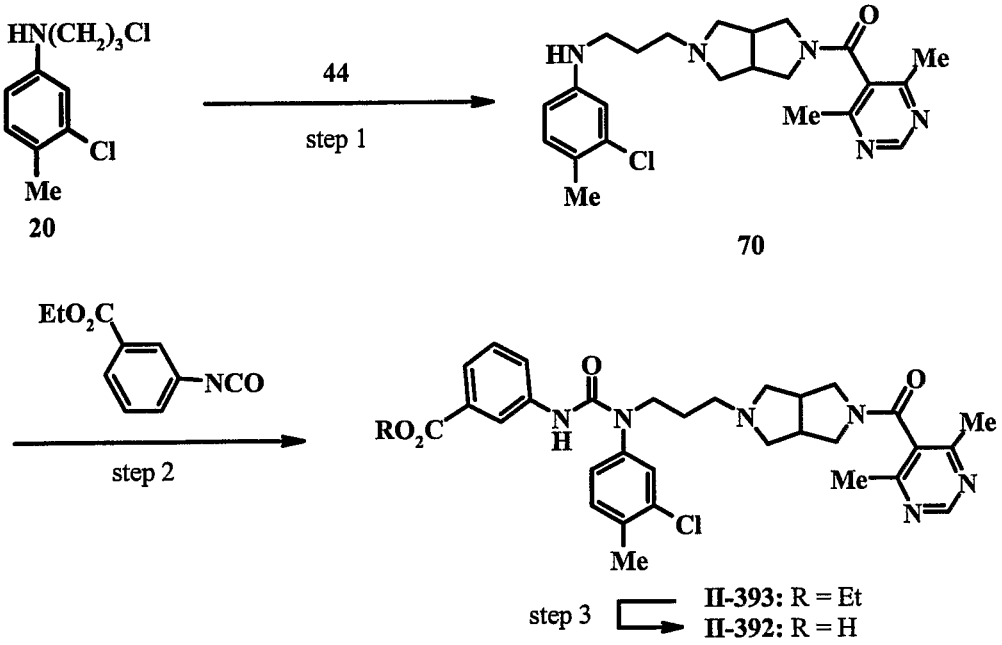
step 1 – To a olution of **61** (0.20 g, 1.13 mmol) and **54** (0.18 g, 0.75 mmol) dissolved in MeCN (10 mL) were K_2CO_3 (0.20 g, 1.51 mmol) and potassium iodide (0.18 g, 1.13 mmol) and the resulting solution was heated at reflux overnight. The mixture was cooled, diluted with water and extracted with EtOAc. The combined EtOAc extracts were washed with brine, dried ($MgSO_4$) and evaporated. The crude product was purified by flash column chromatography on silica gel (DCM:MeOH:NH_4OH / 130:10:1) to afford 0.16 g (54%) of **71**: ms (ES+) m/z 393 (M + H);

step 2 - To a solution of **71** (0.16 g, 0.41 mmol) in DCM (5 mL) were added 1-chloro-4-isocyanato-benzene (0.09 g, 0.62 mmol) and TEA (0.13 mL, 0.91 mmol). The mixture was allowed to stir at RT for 2h then was diluted with DCM and washed with water. The organic layer was dried (Na_2SO_4), filtered and concentrated. The crude product was purified by flash column chromatography on silica gel

(CH_2Cl_2:MeOH:NH_4OH / 200:10:1) to afford 0.19 g (yield 83 %) of II-203: mp 60.5-62.5 °C, ms (ES+) m/z 546 (M + H),

Example 19

3-(3-(3-Chloro-4-methyl-phenyl)-3-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-ureido)-benzoic acid ethyl ester; compound with trifluoro-acetic acid (II-393) and 3-(3-(3-chloro-4-methyl-phenyl)-3-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-ureido)-benzoic acid (II-392)

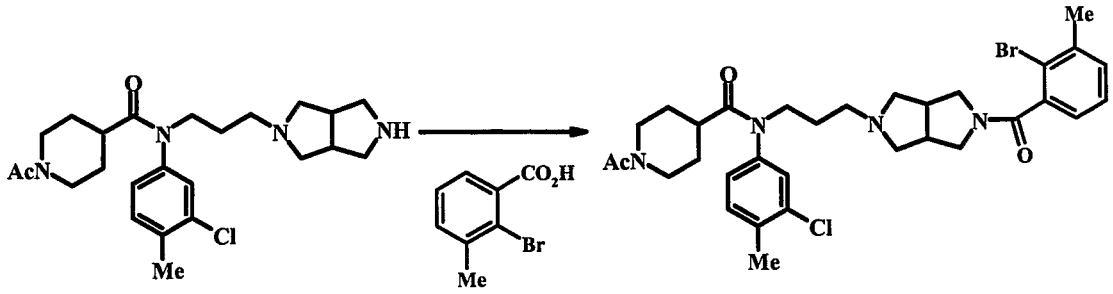


step 1 - To a solution of 20 (1.77 g, 8.12 mmol) and 44 (2.0 g, 8.12 mmol) dissolved in MeCN (10 mL) was added $NaHCO_3$ (1.36 g, 16.24 mmol) and KI (1.34 g, 8.12 mmol). The resulting mixture was heated at reflux overnight. The mixture was cooled, diluted with water and extracted with EtOAc. The combined EtOAc extracts were washed with brine, dried ($MgSO_4$) and evaporated. The crude product was purified by flash chromatography on silica gel eluting with MeOH (containing 10% NH_4OH)/DCM (0 to 4%) to afford 2.7 g of 70: ms (ES+) m/z 428 $(M + H)^+$.

step 2 and 3 - A solution of ethyl 3-isocyanatobenzoate and 70 and THF is shaken overnight at RT. The solvent is evaporated which afforded II-393 which could be purified by SiO_2 chromatography. The crude product was dissolved in MeOH (1 mL) and treated with a 10% solution of NaOH (excess). The reaction mixture was shaken overnight at RT. The reaction mixture is acidified with TFA, solvents evaporated, the crude was dissolved in 5% MeOH/DCM, filtered and concentrated to afford the carboxylic acid II-392: ms (ES+) m/z 591 $(M + H)^+$.

Example 20

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide (II-244)

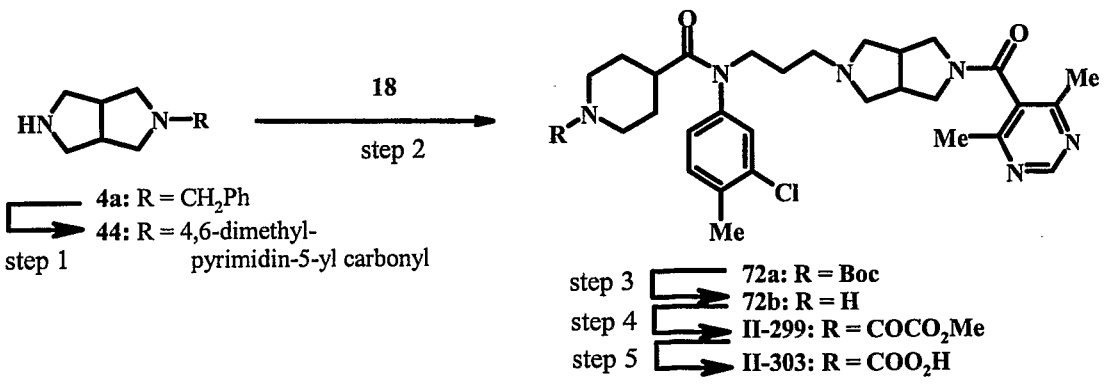


42b 62 II-244

A mixture of 2-bromo-3-methyl-benzoic acid (62, 0.016 g, 0.075 mmol), resin-bound carbodiimide (0.078 g, 0.15 mmol) and HOBT (0.012 g, 0.085 mmol) in DCM:DMF (1 mL, 10:1) was shaken for 18 h. A solution of 42b (0.022 g, 0.05 mmol) and DIPEA (0.03 mL, 0.17 mmol) and DCM (1 mL) was added. Shaking was continued for 24 h, the resin filtered and washed with DMF (1 mL) and twice with DCM (1 mL). The reaction mixture was concentrated *in vacuo* and purified by reverse phase HPLC to afford II-244: ms (ES+) m/z 644 $(M + H)^+$.

Example 21

[4-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-piperidin-1-yl]-oxo-acetic acid, hydrochloride salt (II-303)



step 1 - To a mixture of 4,6-dimethyl-pyrimidine-5-carboxylic acid (0.85 g, 5.58 mmol, T. J. Kress *et. al. Heterocycles* 1994 38:1375) and 4a (1.13 g, 5.58 mmol, C. J.

Ohnmacht *et al. J. Heterocycl. Chem.* 1983 20:321) in DCM (25 mL) at RT was added sequentially EDCI (1.43 g, 6.7 mmol), HOBt (0.9 g, 6.7 mmol) and DIPEA (3.9 mL, 22.34 mmol) and the mixture was stirred overnight at RT. The reaction mixture was washed with 5% $NaHCO_3$ solution, dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica gel eluting with MeOH (containing 10% NH_4OH)/DCM (0 to 4%) to afford 1.5 g of (5-benzyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl)-(4,6-dimethyl-pyrimidin-5-yl)-methanone: ms (ES+) m/z 337 $(M + H)^+$. To a solution of amide from the previous step (1.5 g, 4.45 mmol) in MeOH (50 mL) was added ammonium formate (2.81 g, 44.58 mmol). Palladium on charcoal previously wetted with MeOH was slowly added and the mixture heated to reflux for 8h. The catalyst was filtered and solvent evaporated. The residue was purified by flash chromatography on silica gel eluting with MeOH (containing 10% NH_4OH)/DCM (0 to 4%) to afford 0.941 g of **44b**: ms (ES+) m/z 247 $(M + H)^+$.

step 2 - To a solution of **18** (0.6 g, 1.39 mmol) and **44** (0.31 g, 1.27 mmol) and MeCN (20 mL) were added DIPEA (0.44 mL, 2.54 mmol) and KI (0.23 g, 1.39 mmol) and the resulting mixture was heated at reflux overnight. The mixture was cooled, diluted with water and extracted with EtOAc. The combined EtOAc extracts were washed with brine, dried ($MgSO_4$) and evaporated. The crude product was purified by flash chromatography on silica gel with eluting with MeOH (containing 10% NH_4OH)/DCM (0 to 4%) to afford 0.347 g of **72a**: ms (ES+) m/z 639 $(M + H)^+$.

step 3 - To a solution of **72a** dissolved in DCM (9 mL) was added TFA (1 mL) and the mixture was stirred at T overnight. The solvents were evaporated and the residue suspended in toluene and evaporated again (2x). The residue purified by flash chromatography on silica gel eluting with gradient DCM/MeOH (containing 10% NH_4OH) (0 to 4% MeOH) to afford 0.347 g of **72b**: ms(ES+) m/z 539 $(M + H)^+$.

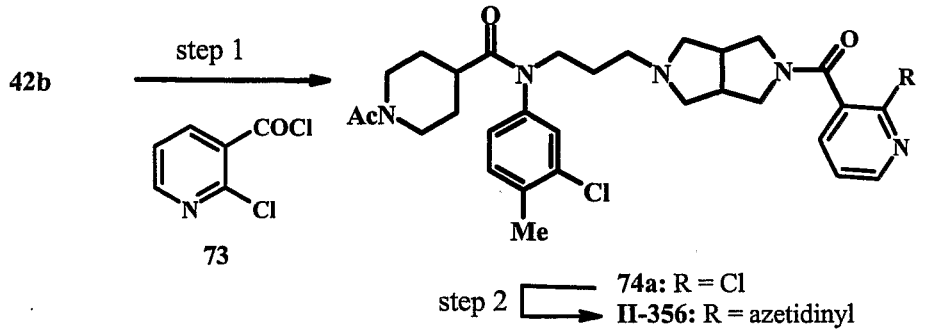
step 4 - To a mixture of **72b** (0.3 g, 0.55 mmol) and methyl oxalyl chloride (0.057 mL, 0.61 mmol) in DCM (5 mL) at RT was added DIPEA (0.145 mL, 0.83 mmol). The reaction mixture was stirred overnight at RT then washed with 5% $NaHCO_3$ solution, back extracted with DCM (5x), dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica gel eluting with MeOH (containing 10% NH_4OH)/DCM (0 to 4%) to afford 0.304 g of **II-299**: mp, 60.3-62 °C; Anal, calcd. for $C_{32}H_{41}ClN_6O_5$ (containing 0.15 mol of CH_2Cl_2): C, 60.53; H, 6.53; N, 13.17. Found: C, 60.59; H, 6.49; N, 12.99; MS(ES+) m/z 625 $(M + H)^+$.

step 5 - To a solution of **II-299** (0.194 g, 0.31 mmol) in MeOH (3 mL) was added a solution of NaOH (0.019 g, 0.47 mmol) in water (1 mL). The mixture was stirred overnight at RT. The solvents were evaporated with a nitrogen stream, redissolved in

MeOH, acidified with HCl (1M solution in ethyl ether), stirred for 15 min at RT then concentrated *in vacuo*. The residue dissolved in DCM, stirred for 15 min and filtered to remove NaCl. Cyclohexane was added and an oily phase separated. The solvents were decanted and product dried *in vacuo* to afford 0.12 g of II-303: mp 172.4-175.2° C; Anal. calcd. for $C_{31}H_{39}ClN_6O_5$ (containing 1 mol of HCl and 0.55 mol of CH_2Cl_2): C, 54.58; H, 5.97; N, 12.10. Found: C, 54.66; H, 5.88; N, 12.03; MS(ES+) m/z 611 $(M + H)^+$.

Example 22

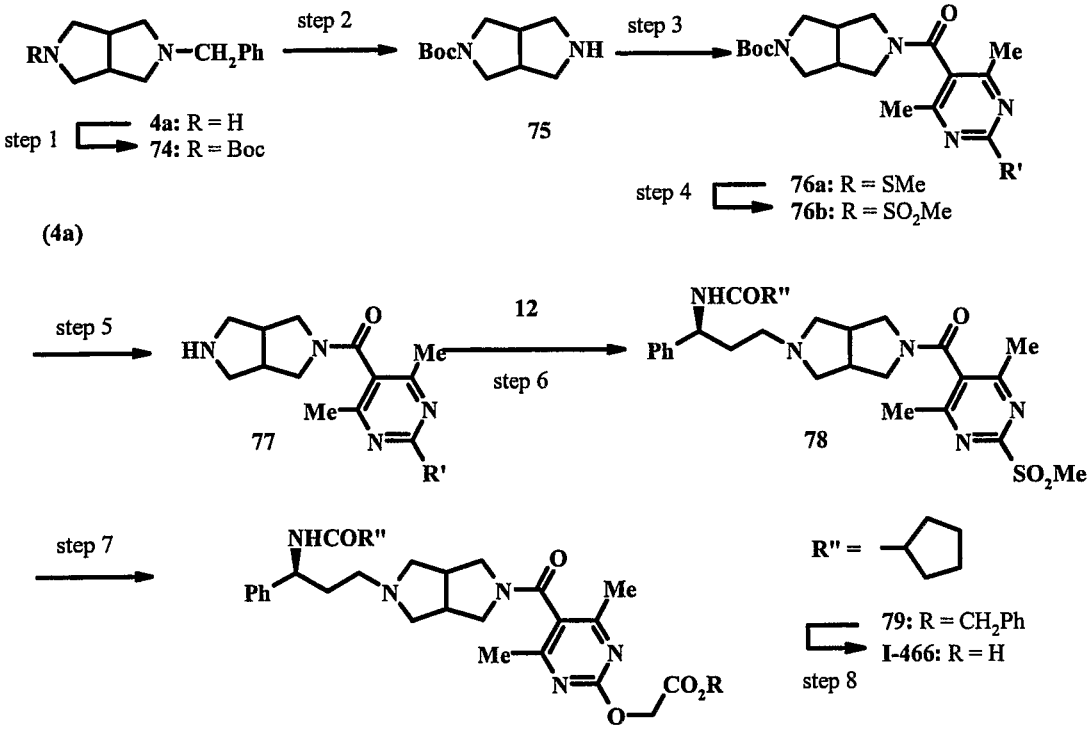
1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-azetidin-1-yl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide (II-356)



To a mixture of 42b (0.063 g, 0.138 mmol) and 2-chloronicotinoyl chloride (73, 0.027 g, 0.15 mmol) in DCM (3 mL) at RT was added DIPEA (0.05 mL, 0.27 mmol). The mixture was stirred overnight at RT and the solvent was evaporated. The residue as obtained contained 74a and was treated with a solution of azetidine (0.1 mL, excess) in THF (2 mL). The reaction mixture was heated at 80° C with stirring for 4 h. The reaction mixture was cooled to RT and the solvents evaporated *in vacuo*. The crude product was purified by flash chromatography on silica gel eluting with MeOH (containing 10% NH_4OH)/DCM (0 to 4%). The recovered material was dissolved in DCM and cyclohexane was added which resulted in the separation of an oil. The solvents were decanted and product dried *in vacuo* to afford 0.045 g of II-356: mp 72.9-74° C; Anal. calcd. for $C_{33}H_{43}ClN_6O_3$ (containing 0.5 mol of C_6H_{12} and 0.5 mol of H_2O): C, 65.69; H, 7.66; N, 12.77. Found: C, 65.99; H, 7.47; N, 12.61; ms (ES+) m/z 607 $(M + H)^+$.

Example 23

(5-{(5-[(S)-3-(Cyclopentanecarbonyl-amino)-3-phenyl-propyl]-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl}-4,6-dimethyl-pyrimidin-2-yloxy)-acetic acid; compound with trifluoro-acetic acid, (I-466)



step 1 - A solution of $(BOC)_2O$ (15.7 g, 72 mmol) in THF (20 mL) was added to a solution of 4a (12g, 59.3 mmol) in THF (100 mL) at RT, and resulting solution was stirred for 18 h, concentrated in *vacuo*. The residue was dissolved in EtOAc (100 mL), and the organic phase was washed with saturated $NaHCO_3$ (25 mL), 1M citric acid (25 mL) and brine (1x25 mL), dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by flash chromatography on SiO_2 eluting with 10%EtOAc/DCM to afford 11.7g (65%) of 74:^{1}H NMR ($CDCl_3$) δ 1.46 (s, 9H), 2.37-2.40 (m, 2H), 2.62-2.68 (m, 2H), 2.76-2.79 (m, 2H), 3.25 (m, 2H), 3.51-3.57 (m, 2H), 3.58 (s, 2H), 7.23-7.31 (m, 5H); MS (ES+) m/z 303 $(M + H)^+$.

step 2 - Palladium hydroxide (0.1 g) was added to a solution of 74 (1.0 g, 3.3 mmol) in EtOH (50 mL). The solution was hydrogenated under atmospheric pressure overnight and filtered through a CELITE® pad. The resulting solution was concentrated *in vacuo* to afford 0.6 g (85%) of 75: MS (ES+) m/z 213 $(M + H)^+$.

step 3 - A solution of 75 (0.6 g, 2.82 mmol), dimethylsulfamoyl chloride (0.3 mL, 2.82 mmol), 4,6-dimethyl-2-methylsulfanyl-pyrimidine-5- carboxylic acid (0.56 g, 2.82 mmol), N,N-dimethylbutyl amine (1.16 mL, 8.4 mmol), and DMAP (34 mg, 0.000282 mol) in DMF (5 mL) was stirred at 65° C overnight. DMF was removed *in vacuo*. The residue was dissolved in EtOAc (30 mL), washed with saturated $NaHCO_3$ (25 mL), 1M citric acid (25 mL), brine (25 mL), dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by flash chromatography on SiO_2 eluting with 20%MeOH/DCM to afford 0.6g (54%) of 76a: MS (ES+) m/z 393 $(M + H)^+$.

step 4 - A solution of **76a** (1 g, 2.54 mmol) in DCM (15 mL) was cooled to 0° C and MCPBA (1.3 g, 7.6 mmol) was added in portions. Stirring was continued for 2 h. The reaction mixture was diluted with DCM (15 mL), washed with saturated $NaHCO_3$ (25 mL), brine (25 mL), dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica gel eluting with 10%MeOH/CH_2Cl_2 to afford 0.34g (31%) of **76b**: MS (ES+) m/z 425 $(M + H)^+$.

step 5 - The carbamate **76b** (0.12 g, 0.00028 mole) was dissolved in 50% TFA/ DCM (5 mL) and stirred at RT for 1 h. Solvents were removed *in vacuo*, and residue was dried *in vacuo* for 5 h at 40°C to afford 0.12 g of **77**: ms (ES+) m/z 325 $(M + H)^+$.

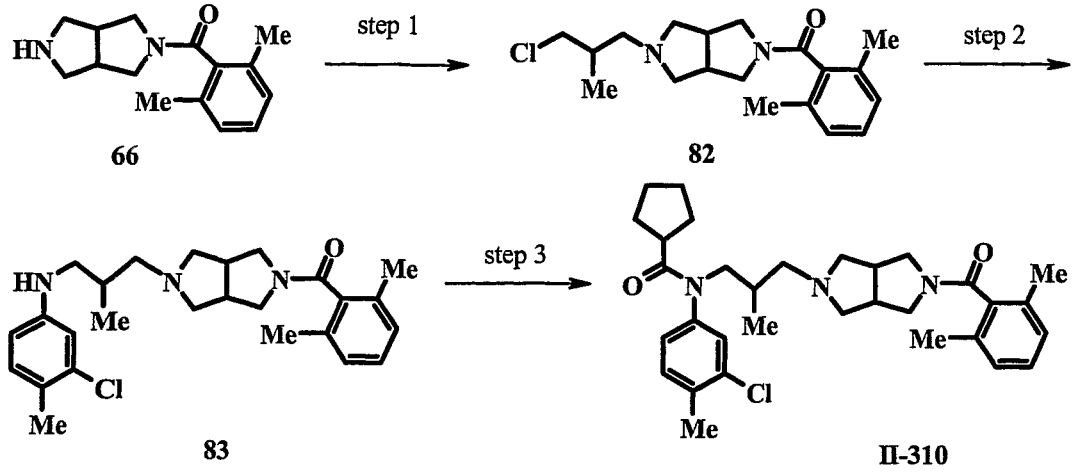
step 6 - Cyclopentanecarboxylic acid (3-oxo-1-phenyl-propyl)-amide (**12**, 0.41g, 1.68 mmol) was added to a solution of **77** TFA salt (0.51 g, 1.18 mmol) in DCM (5 mL), and $NaBH(OAc)_3$ (0.35 g, 1.65 mmol) was added. The reaction was stirred at RT for 18 h, diluted with EtOAc (25 mL), washed with saturated $NaHCO_3$ (25 mL), brine (25 mL), dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica gel eluting with 10% MeOH/DCM to afford 0.4g (61%) of **78**: MS (ES+) m/z 554 $(M + H)^+$.

step 7 - To a mixture of **78** (0.14 g, 2.51 mmol) and Cs_2CO_3 (0.16 g, 5.0 mmol) in DMF (2 mL) was added benzyl glycolate (0.05g, 3 mmol), and resulting mixture was stirred at 70° C for 8 h. DMF was removed *in vacuo* and residue was purified by flash chromatography on silica gel eluting with 10%MeOH/DCM to afford 0.1g (62%) **79**: MS (ES+) m/z 640 $(M + H)^+$.

step 8 - Palladium on carbon (20 mg) was added to a solution of **79** (0.1 g, 0.15 mmol) on EtOH (20 mL). The solution was hydrogenated under atmospheric pressure overnight and filtered through a CELITE® pad. The resulting solution was concentrated *in vacuo* to afford 78 mg (91%) of **I-466**: mp 117.3-119.0° C, ms (ES+) m/z 550 $(M + H)^+$.

Example 24

Cyclopentanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-methyl-propyl}-amide (**II-310**)



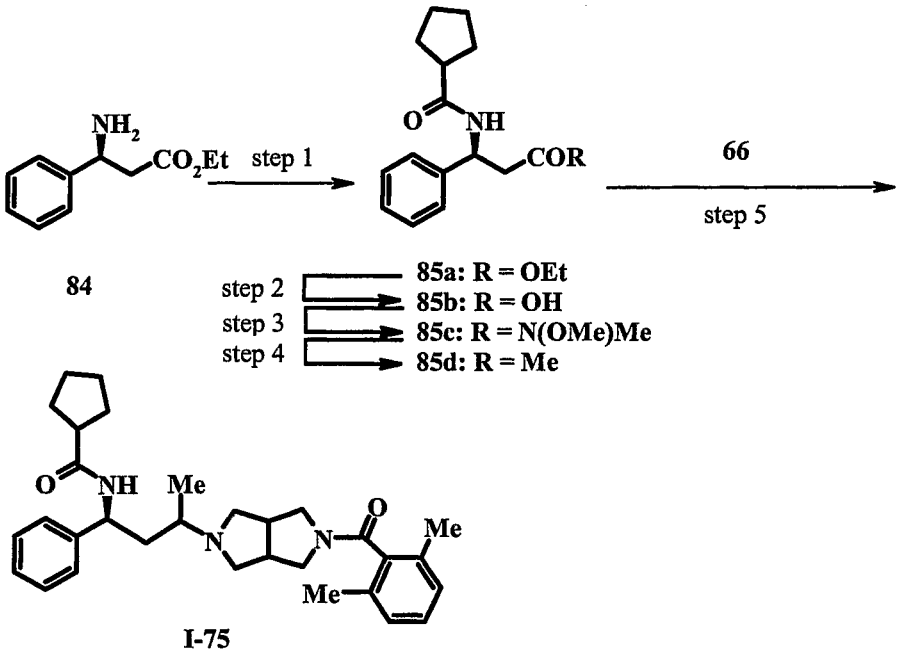
step 1 - 1-Bromo-3-chloro-2-methyl-propane (0.17 mL, 1.45 mmol) and TEA (0.24 mL, 1.71 mmol) were added to a solution of (2,6-dimethyl-phenyl)-(hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-methanone (**66**, 358 mg, 1.47 mmol) in DMF (6 mL). The mixture was stirred overnight at RT. Water was added and the mixture was extracted with DCM. The organic layer was dried (Na_2SO_4) and concentrated. The crude product was purified by SiO_2 column chromatography on silica gel eluting with a DCM: MeOH: NH_4OH gradient (98:1.4:0.14 to 96:3.5:0.35 over 40 min.) to afford **82** (224 mg, 46%): ms (LCMS) m/z 335 (M + H).

step 2 - To a solution of **82** (224 mg, 0.67 mmol) dissolved in DMF (3 mL) was added KI (120 mg, 0.72 mmol), 3-chloro-4-methyl-phenylamine (105 mg, 0.74 mmol) and DIPEA (0.12 mL, 0.69 mmol) and the reaction mixture was stirred at 80° C for 4 h. The mixture was allowed to cool to RT, diluted with water and extracted with DCM. The combined DCM extracts were dried (Na_2SO_4) and concentrated. The crude product was purified by SiO_2 column chromatography eluting with a CH_2Cl_2: MeOH: NH_4OH (99:0.7:0.07) to afford **83** (146 mg, 49%) as an off-white solid: ms (LCMS) m/z 440 (M + H).

step 3 - Cyclopentanecarbonyl chloride (0.07 mL, 0.06 mmol) and DIPEA (0.02 mL, 0.11 mmol) were added to a solution of **83** (28.5 mg, 0.06 mmol) in toluene (1 mL). The resulting solution was stirred at 50° C for 1 hour. The mixture was allowed to cool to room temperature, diluted with water and extracted with DCM. The combined DCM extracts were dried (Na_2SO_4) and concentrated. The crude product was purified by SiO_2 column chromatography eluting with a DCM: MeOH: NH_4OH gradient (99:0.7:0.07 to 96:3.5:0.35 over 50 min.) to afford **II-310** (14 mg, 40%): ^1H NMR ($CDCl_3$) δ 0.9 (d, 3H), 1.35-1.45 (m, 2H), 1.55 (s, 3H), 1.6-1.85 (m, 5), 2.1-2.5 (m, 15H), 2.65-2.9 (m, 3H), 3.22-3.3 (m, 1H), 3.4-3.9 (m, 4H), 6.95-7.4 (m, 3H), 7.1-7.3 (m, 3H); ms (ES+) m/z 536 (M + H).

Example 25

Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-butyl}-amide (I-75)



I-75

steps 1 and 2- To a solution of (S)-3-amino-3-phenylpropanoic acid ethyl ester hydrochloride (84, 5.0 g, 21.76 mmol) in H_2O (50 mL), saturated Na_2CO_3 (50 mL), DCM (50 mL) and toluene (20 mL) was added cyclopentylcarboxylic acid chloride (2.9 mL, 23.93 mmol). The reaction was stirred overnight at RT. The mixture was extracted with DCM and the organic layer was dried ($MgSO_4$), filtered and concentrated *in vacuo*. The residue containing **85a** was redissolved in H_2O (50 mL) and THF (50 mL) $LiOH.H_20$ (2.73 g, 65.06 mmol) was added. After 3h, the mixture was washed with ether. The aqueous layer was acidified with 2N HCl and extracted with EtOAc. The organic layer was dried ($MgSO_4$), filtered and evaporated to afford 5.56 g (97%) of **85b**; ^1H NMR ($CDCl_3$) δ 1.5-1.65(m, 2H), 1.65-1.95 (m, 6H), 2.55 (t, 1H), 2.8-3.0 (qd, 2H), 5.4-5.5 (m, 1H), 6.4-6.45 (d, 1H), 7.2-7.45 (m, 5H).

step 3 - N,O-dimethylhydroxylamine hydrochloride (0.92 g, 9.50 mmol), O-(7-azabenzotriazol-1-yl)N,N,N',N'-tetramethyluronium hexafluorophosphate (3.61 g, 9.50 mmol) and DIPEA (5.5 mL, 31.68 mmol) were added to a solution of **85b** (2.07 g, 7.92 mmol) in DCM (80 mL). The reaction mixture was stirred overnight at RT, poured into 2N NaOH solution and stirred for 10 min. The organic layer was washed with H_2O, 2N HCl and brine, dried ($MgSO_4$) and concentrated *in vacuo*. The crude product was purified by SiO_2 column chromatography eluting with n-hexane:EtOAc (1:2) to afford 1.24 g of **85c** (yield 51 %): ^1H NMR ($CDCl_3$) δ 1.55-1.65(m, 2H), 1.65-1.95 (m, 6H), 2.65

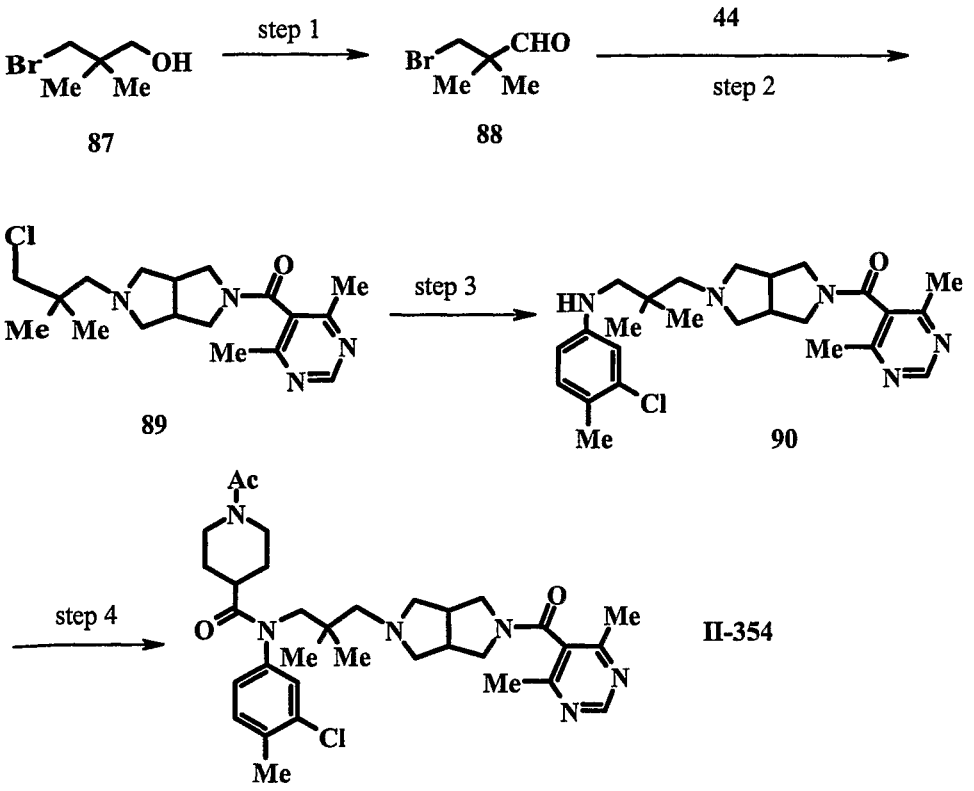
(q, 1H), 2.8 (dd, 1H), 3.1 (s, 3H) 3.1-3.2 (dd, 1H), 3.45 (s, 3H), 5.43 (m, 1H), 7.15-7.45 (m, 5H), 7.4 (d 1H);

step 4 - To a solution of **85c** (1.06g, 3.50 mmol) in THF (15 mL) was added dropwise at -78 °C MeMgCl (3.7 mL, 3M in THF). The mixture was warmed to RT over 3h and it was stirred at RT one additional hour. The reaction was quenched with1N K_2HPO_4 extracted with Et_2O. The organic layer was washed with saturated $NaHCO_3$ and brine, dried ($MgSO_4$), filtered and concentrated *in vacuo*. The crude product was purified by SiO_2 column chromatography on eluting with n-hexane:EtOAc (1:1) to afford 0.89 g (98%)of **85d**.

step 5 - To a solution of (2,6-dimethyl-phenyl)-(hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-methanone (**66**, 0.12 g, 0.52 mmol) in DCM (7 mL) and THF (7 mL) was added cyclopentanecarboxylic acid (3-oxo-1-phenyl-butyl)-amide (**85d**, 0.15 g, 0.57 mmol). Titanium tetraisopropoxide (0.34 mL, 1.15 mmol) was added to the mixture. After 30 min $NaBH(OAc)_3$ (0.16 g, 0.78 mmol) was added and the mixture stirred at RT for 4 h. Saturated $NaHCO_3$ was added to the mixture and it was stirred for 10 min. The mixture was extracted with DCMand the organic layer was dried ($MgSO_4$), filtered and evaporated. The crude product was purified by SiO_2 chromatography on silica gel eluting with DCM:MeOH:NH_4OH (150:10:1) to afford I-75: ms (ES+) m/z 488 (M + H); Anal. ($C_{31}H_{41}N_3O_2$. 0.3M H_2O) C; calcd, 75.51; found, 75.53; H; calcd, 8.50; found, 8.29; N; calcd, 8.52; found, 8.53.

Example 26

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2,2-dimethyl-propyl}-amide (II-354)



step 1 - The Dess-Martin reagent (1.12 g, 2.64 mmol) was added to a solution of 3-bromo-2,2-dimethyl-propan-1-ol (87, 0.37 mL, 3.00 mmol) in DCM (20 mL) maintained 0° C. The reaction mixture was stirred at 0° C for 1 h. The ice bath was removed and the reaction mixture was stirred at RT for additional 30 min. Et_2O (40 mL) was added, and the mixture was poured onto 20 mL of an 9:1 mixture of aqueous saturated $NaHCO_3$ and aqueous saturated $Na_2S_2O_3$. The mixture was stirred until a clear solution was obtained (10 minutes). The layers were separated, and the Et_2O layer was washed with saturated aqueous $NaHCO_3$ (2x) and water (2x). The organic layer was dried (Na_2SO_4) and concentrated to afford **88** (393 mg). The crude product was used for the next step without further purification.

step 2 - A solution of 3-bromo-2,2-dimethyl-propionaldehyde (**88**, 393 mg, 23.8 mmol) and 44 (620 mg, 25.2 mmol) in DCM (6 mL) was treated with HOAc (0.2 mL, 34.9 mmol) and $NaBH(OAc)_3$ (530 mg, 25.0 mmol). The reaction mixture was stirred for 2 h at RT and then quenched by the addition of aqueous NaOH (2 M, 2.4 mL, 0.48 mmol). The mixture was extracted with DCM. The combined DCM extracts were dried (Na_2SO_4) and concentrated. The crude product was purified by SiO_2 column chromatography eluting with a CH_2Cl_2: MeOH: NH_4OH gradient (99:0.7:0.07 to 96:3.5:0.35 over 50 min.) to afford **89** (95 mg, 10%): ms (LCMS) m/z 397 (M + H).

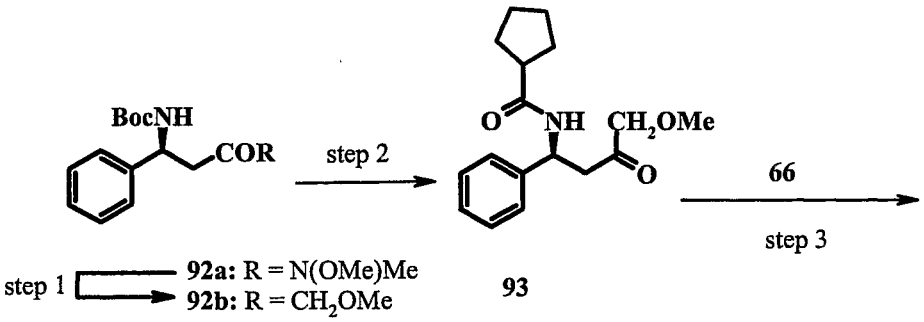
step 3 - A solution of **89** (95 mg, 0.24 mmol) in MeCN (1 mL) was treated with 3-chloro-4-methyl-phenylamine (76 mg, 0.53 mmol) and TEA (0.035 mL, 0.25mmol). The

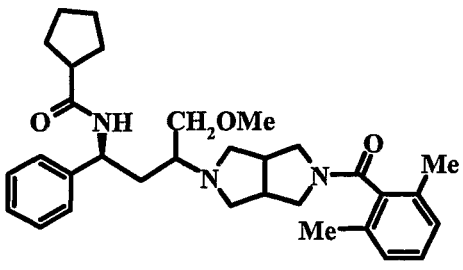
reaction mixture was stirred at 50° C overnight. The mixture was allowed to cool to RT, diluted with water and extracted with DCM. The combined CH_2Cl_2 extracts were dried (Na_2SO_4) and concentrated. The crude product was purified by SiO_2 column chromatography eluting with a CH_2Cl_2: MeOH: NH_4OH gradient (99:0.7:0.07 to 96:3.5:0.35 over 50 min.) to afford 90 (25 mg, 23%): ms (LCMS) m/z 456 (M + H).

step 4 - 1-Acetyl-piperidine-4-carbonyl chloride (24 mg, 0.13 mmol) and TEA (0.02 mL, 0.14 mmol) were added to a solution of 90 (25 mg, 0.054 mmol) in DCM (2 mL). The resulting solution was stirred overnight. The mixture was diluted with water and extracted with DCM. The combined DCM extracts were dried (Na_2SO_4) and concentrated. The crude product was purified by SiO_2 column chromatography eluting with a CH_2Cl_2: MeOH: NH_4OH gradient (98:1.4:0.14 to 93:6:0.6 over 50 min.) to afford II-354 (9 mg, 27%): ^1H NMR (DMSO-d_6) δ 0.75 (s, 6H), 1.2-1.6 (m, 4H), 1.95 (s, 3H), 2.1 (s, 2H), 2.2-2.5 (m, 16H), 2.65-2.85 (m, 4H), 3.25-3.35 (m, 1H), 3.6-3.8 (m, 4H), 4.2-4.3 (m, 1H), 7.25-7.5 (m, 3H), 8.95 (s, 1H); ms (ES+) m/z 609 (M + H).

Example 27

Cyclopentanecarboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-4-methoxy-1-phenyl-butyl}-amide. (I-354)





I-354

step 1 - To a solution of tributyl-methoxymethyl-stannane (0.71 g, 2.14 mmol, T. S. Kaufman *Syn. Lett.* 1997 12:1377-1378) in THF (5 mL) was added n-BuLi (1.4 mL, 1.5M in hexane) dropwise at -78 °C and the reaction was stirred for 10 min at -78 °C. A solution of 92a (0.22 g, 0.71 mmol) in THF (3 mL) was added to the mixture slowly and

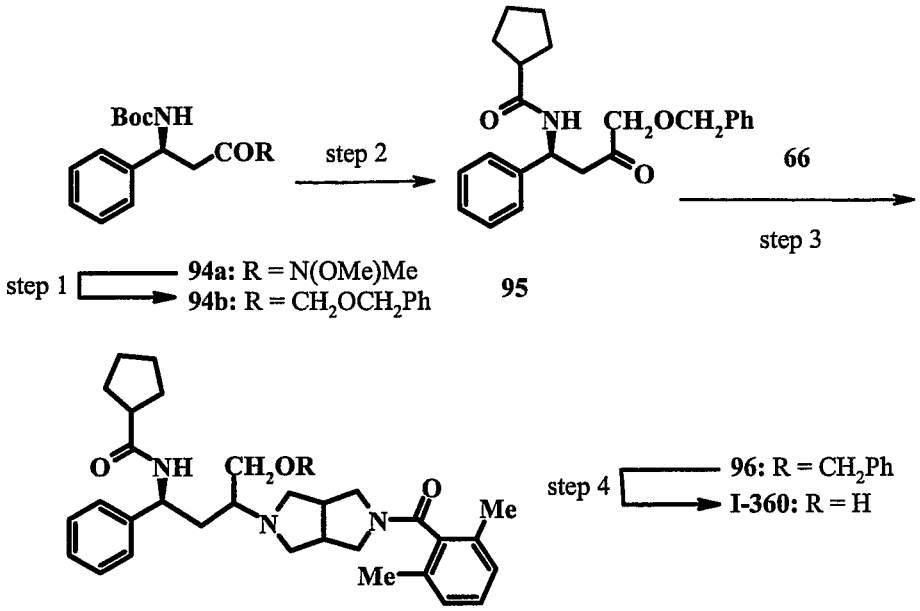
after the addition the reaction mixture was stirred for 30 min at -78 °C. Sat. NH_4Cl was added to the mixture and it was warmed up to RT. The mixture was extracted with EtOAc and the organic layer was washed with brine, dried ($MgSO_4$), filtered and concentrated *in vacuo*. The crude product was purified by SiO_2 column chromatography eluting with n-hexane:EtOAc (2:1) to afford 0.13 g (62 %) of **92b**: ^1H NMR ($CDCl_3$) δ 1.42 (s, 9H), 2.85-3.1 (dd, 2H), 3.35 (s, 3H), 3.85 (s, 2H), 5.1 (br, 1H), 5.38 (br, 1H), 7.2-7.4 (m, 5H).

step 2 - Methanolic HCl (17 mL, 1.25 M) was added to **92b** (0.31 g, 1.05 mmol) and the mixture was heated at 50 °C for 2h. The mixture was concentrated *in vacuo* and dried *in vacuo*. The crude product was dissolved in a mixture of saturated Na_2CO_3 (4 mL), H_2O (2 mL), DCM (2 mL) and toluene (1 mL) and the resulting mixture was treated with cyclopentylcarboxylic acid chloride (0.18 mL, 1.48 mmol). After stirring overnight the solution was extracted with DCM and the organic layer was washed with brine, dried (Na_2SO_4) and concentrated *in vacuo*. The crude product was purified by SiO_2 column chromatography eluting with n-hexane:EtOAc (1:1) to afford 0.23 g (76 %)of **93**. mp 89.5-92.2 °C; ms (ES+) m/z 290 (M + H).

step 3 - To a solution of **93** (0.15 g, 0.63 mmol) in DCE (7 mL) and THF (7 mL) was added **66** (0.18 g, 0.63 mmol). Titanium tetraisopropoxide (0.41 mL, 1.39 mmol) was added to the mixture and after 30 min $NaBH(OAc)_3$ (0.20 g, 0.95 mmol) was added and the reaction stirred at rt overnight. Saturated $NaHCO_3$ was added to the mixture and stirring continued for 10 min. The mixture was filtered through a CELITE® pad and extracted with DCM. The organic layer was dried ($MgSO_4$), filtered and concentrated *in vacuo*. The crude product was purified by SiO_2 column chromatography eluting with 5% MeOH/EtOAc to afford 0.32g (50 %) of **II-354**: ms (ES+) m/z 517 (M + H); Anal. ($C_{32}H_{43}N_3O_3 \cdot 0.15M$ DCM) C; calcd, 72.80; found, 72.55; H; calcd, 8.23; found, 8.22; N; calcd, 7.92; found, 7.93.

Example 28

Cyclopentanecarboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-4-hydroxy-1-phenyl-butyl}-amide (**I-360**)

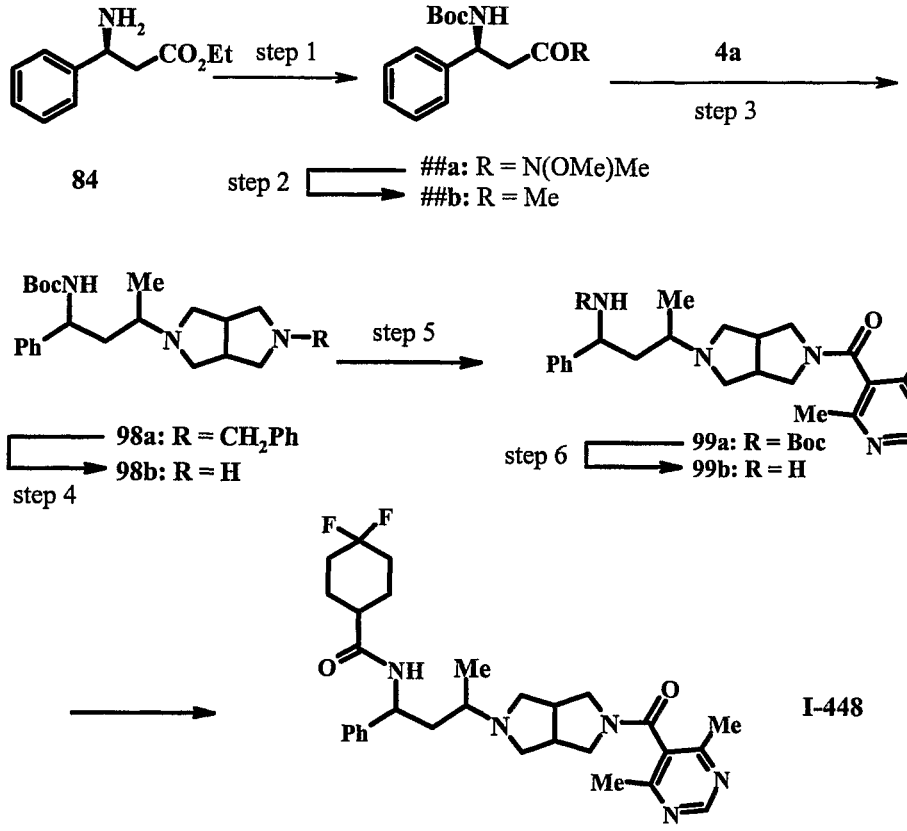


Steps 1 to 3 were carried out as described in Example 27 except benzyloxymethyl-tributyl-stannane was used in place of methoxymethyl-tributyl-stannane in step 1.

step 4 - To a solution of 96 (0.21g, 0.35 mmol) and EtOH (6 mL) was added 2N HCl (2 mL) and Pd/C (0.02 g). The mixture was stirred at RT overnight under H_2 atmosphere. The catalyst was filtered through a CELITE® pad and the filtrate was concentrated *in vacuo*. The crude product was purified by SiO_2 column chromatography eluting with CH_2Cl_2:MeOH:NH_4OH (120:10:1) to afford 0.10 g (56 %) of I-360: mp 78.9-84.9 °C; ms (ES+) m/z 504 (M + H).

Example 29

4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-butyl}-amide (I-448)



Formation of the N-Boc derivative of **84** was carried out using stand protocols. Steps 2 and 3 were carried out as described for steps 4 and 5 of example 25 except the N-protecting groups were Boc and benzyl in the present example. Silica chromatography of **98a** eluting with DCM:MeOH:NH_4OH (150:10:1) afforded 0.72 g (57%): ^1H NMR (DMSO-d_6) δ 0.9-0.95 (m, 3H), 1.35 (s, 9H), 1.45-1.7 (m 1H), 1.8-2.2 (m, 1H), 2.25-2.38 (m, 4H), 2.45-2.7 (m, 6H), 3.55 (q, 2H), 4.6-4.75 (1H), 7.25-7.32 (m, 10H).

step 4 - Palladium on carbon (0.07 g) and ammonium formate (1.01 g, 16.01 mmol) were added to a solution of **98a** (0.72 g, 1.60 mmol) in MeOH (30 mL). The solution was heated at reflux for 2 h and filtered through a CELITE® pad. The resulting solution was concentrated *in vacuo* and crude product purified by SiO_2 column chromatography eluting with DCM: MeOH: NH_4OH(90:10:1) to afford **98b**.

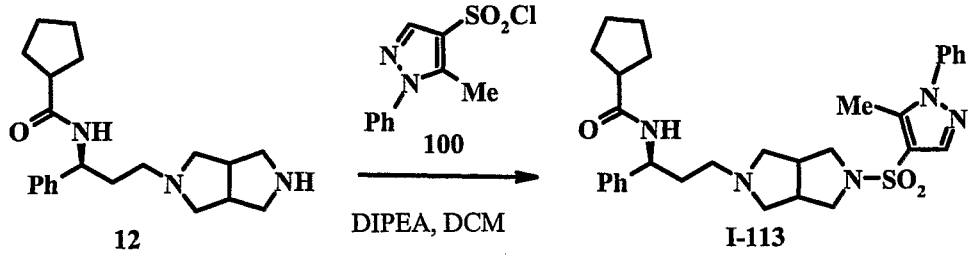
step 5 - To a solution of [3-(hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-butyl]-carbamic acid tert-butyl ester **98b** (0.22 g, 0.63 mmol) in DCM (10 mL) were added 4,6-dimethyl-pyrimidine-5-carboxylic acid (0.11 g, 0.76 mmol) , EDCI (0.16 g, 0.82 mmol), HOBt (0.11 g, 0.82 mmol) and DIPEA (0.33 mL, 1.90 mmol). The mixture was stirred at RT overnight. The mixture was washed with saturated. $NaHCO_3$ and the organic layer was dried (Na_2SO_4). The crude product was purified by SiO_2 column chromatography eluting wit CH_2Cl_2:MeOH:NH_4OH (160:10:1) to afford 0.30 g (95 %) of **99a**:

step 6 - Methanolic HCl (5 mL, 1.25 M) was added to **99a** (0.30 g, 0.61 mmol) and the mixture was heated at 50 °C for 2 h. The mixture was concentrated in vacuo and purified by SiO_2 column chromatography eluting with CH_2Cl_2:MeOH:NH_4OH (120:10:1) to afford 0.17 g (70 %) of **99b**:

step 7 - To a solution of **99b** (0.17 g, 0.43 mmol) in DCM (10 mL) were added 4,4-difluoro-cyclohexanecarboxylic acid (0.08 g, 0.51 mmol) , EDCI (0.10 g, 0.56 mmol), HOBt (0.07 g, 0.56 mmol) and DIPEA (0.22 mL, 1.29 mmol). The mixture was stirred at RT overnight. The mixture was washed with saturated. $NaHCO_3$ and the organic layer was dried (Na_2SO_4). The crude product was purified by SiO_2 column chromatography eluting with DCM:MeOH:NH_4OH (150:10:1) to afford 0.22 g (95 %) of **I-448**: mp 86.4-87.0 °C; ms (ES+) m/z 540 (M + H).

Example 30

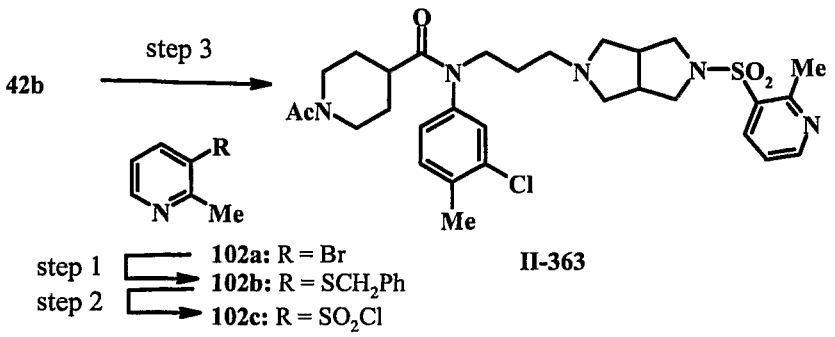
Cyclopentanecarboxylic acid {3-[5-(5-methyl-1-phenyl-1H-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide (I-113)



To a solution of 12 (, 0.012 g, 0.035 mmol), DIPEA (0.02 mL, 0.105 mmol) and DCM (1 mL) at RT was added 5-methyl-1-phenyl-1*H*-pyrazole-4-sulfonyl chloride (**100** 0.018 g, 0.07 mmol) and the mixture stirred overnight at rt. The reaction mixture was concentrated *in vacuo* and purified by reverse phase HPLC: MS (ES+) m/z 562 $(M + H)^+$ to afford **I-113**.

Example 31

4-Acetyl-piperidine-1-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-pyridine-3-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide (II-363)



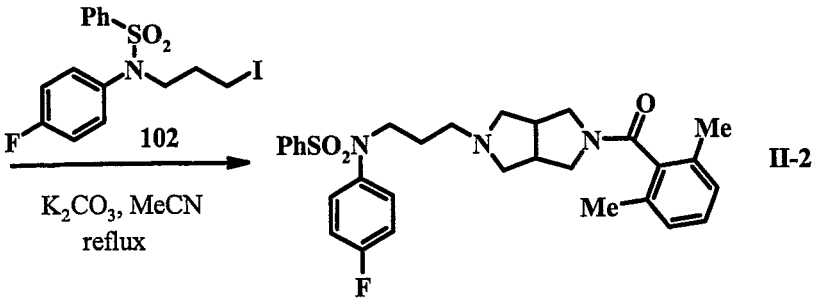
step 1 - Benzylmercaptan (664 mg, 0.63 mL, 5.34 mmol)was dissolved in 8.5 mL DMF under an atmosphere of N_2. NaH(214 mg 60% as an oil dipersion, 5.34 mmol) was added portionwise with stirring, then stirred for an additional 15 mins. To the solution was added 3-bromo-2-methyl-pyridine(102a, 707 mg, 4.11 mmol) in one portion and the mixture heated in an oil bath at 130° C for 4 h. The reaction mixture was cooled to RT and partitioned between H_2O (50 mL) and hexane (50 mL). The hexane layer was separated, washed with H_2O (50 mL), dried over $MgSO_4$, then concentrated *in vacuo*. The crude product was purified by flash chromatography on silica gel eluting with EtOAc:hexane (1:2) to afford 522 mg (44%) of **102b** as a viscous liquid: ms (ES+) m/z 216 $(M+H)^+$.

step 2 - A solution of **102b** (414 mg, 1.92 mmol) in glacial HOAc (8.3 mL) and H_2O (0.83 mL) was cooled to 0° C. Chlorine gas was bubbled into the mixture for 15 min, then stirred for an additional 20 min. The mixture was diluted with DCM (50 mL), washed with brine, saturated $NaHCO_3$ solution, brine, dried ($MgSO_4$), and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica eluting with EtOAc:hexane (1:2) to afford 285 mg (76%) of **102c** as a pale yellow liquid: ms (ES+) m/z 191 (M^+).

step 3 - A solution of **102c** (47 mg, 0.25 mmol), DIPEA (35 mg,0.05 mL, 0.27 mmol) and **42b** (100 mg,0.22 mmol) in DCM (3.5 mL) was stirred for 2 h then concentrated *in vacuo*. The crude product was purified by flash chromatography on silica eluting with DCM/ 8% MeOH (containing 2% NH_4OH) to afford 133 mg (99%) of II-363as a viscous liquid.

Example 32

N-{3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-N-(4-fluoro-phenyl)-benzenesulfonamide (II-2)

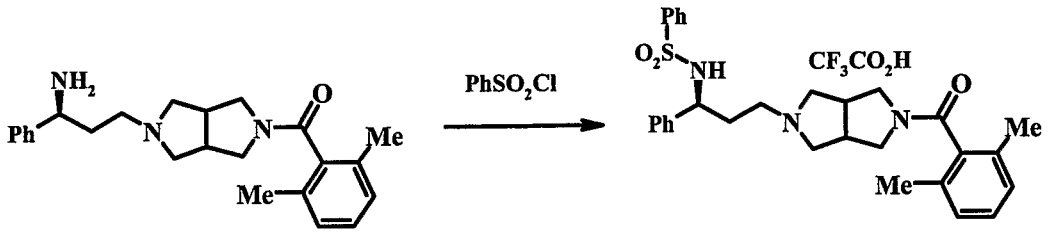


66 II-2

To a solution of **66** (0.012 g, 0.05 mmol) in MeCN (1 mL) at RT were added *N*-(4-fluoro-phenyl)-*N*-(3-iodo-propyl)-benzenesulfonamide (**102**, 0.025 g, 0.06 mmol) and K_2CO_3 (0.01 g, 0.075 mmol). The reaction mixture was shaken at 80 °C for 48 h. After cooling to RT, the mixture was diluted with DCM (5 mL), filtered and evaporated *in vacuo*. The crude product was purified by reverse phase HPLC to afford **II-2**: MS (ES+) m/z = 536 $(M + H)^+$.

Example 33

N-{3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-benzenesulfonamide (**I-32**)

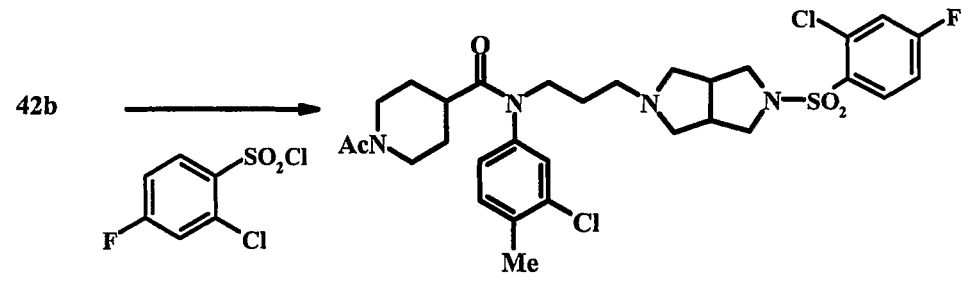


103 **II-32**

A solution of **103** (35 µmol) in DCM was added to benzenesulfonyl chloride (9.3 mg 53 µmol) and DIPEA (30 µl) was added. The reaction was shaken for 18 h. The solution was evaporated under reduced pressure and purified by preperative HPLC to afford **II-32**: ms (ES+) m/z 518 $(M + H)^+$.

Example 34

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-4-fluoro-benzenesulfonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide (**II-296**)

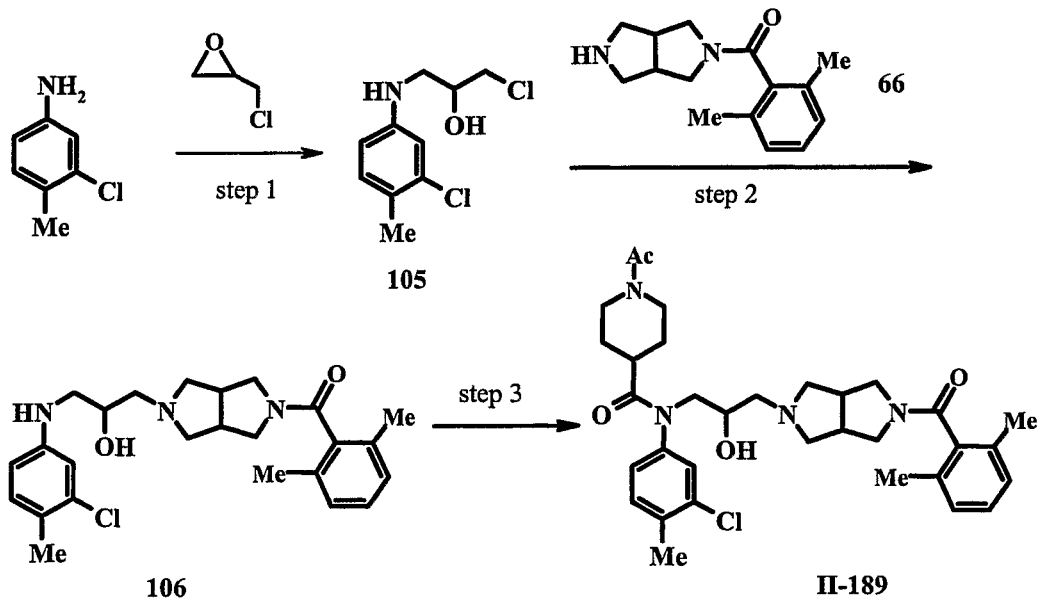


104 II-296

To a solution of 42b (0.016 g, 0.035 mmol) and DIPEA(0.02 mL, 0.105 mmol) in DCM (1 mL) at RT was added 2-chloro-4-fluorobenzenesulfonyl chloride (104, 0.016 g, 0.07 mmol) and the mixture shaken overnight at rt. The reaction mixture was concentrated *in vacuo* and purified by reverse phase HPLC to afford **II-296:** ms (ES+) m/z 639 $(M + H)^+$.

Example 35

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-hydroxy-propyl}-amide(II-189)



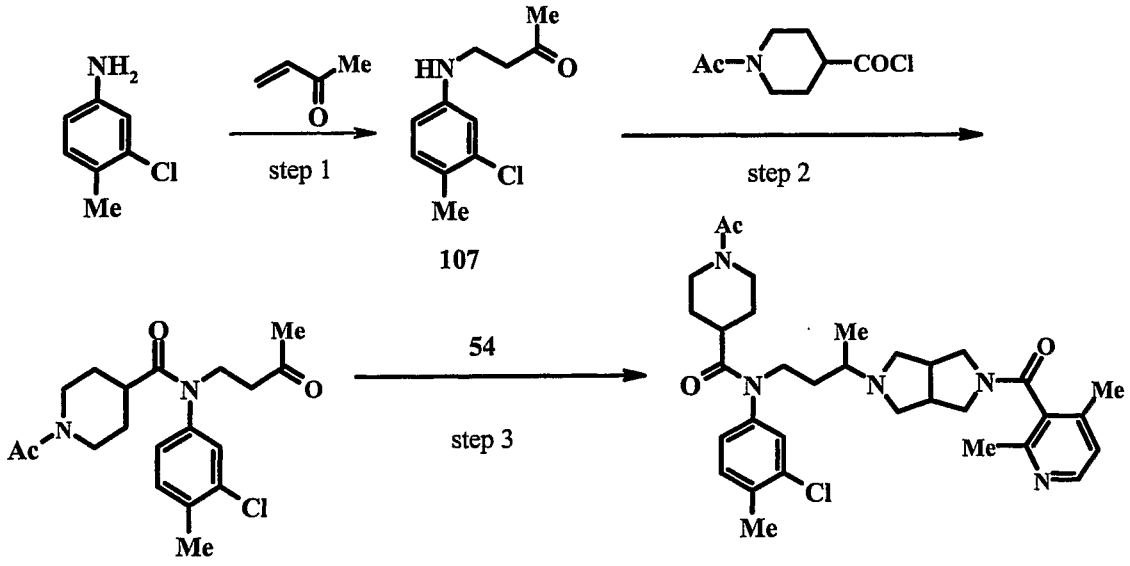
step 1 – A solution of 3-chloro-4-methyl-phenylamine (3.0 g, 21.18 mmol) and 2-chloromethyl-oxirane (0.83 mL, 10.59 mmol) and EtOH (30 mL) were heated to reflux overnight. The mixture was concentrated *in vacuo* and purified by SiO_2 column chromatography eluting with n-hexane:EtOAc (5:1) to afford 2.23 g (90 %) of **105:** ^1H NMR ($CDCl_3$) δ 2.25 (s, 3H), 3.18 (dd, 1H), 3.35 (dd, 1H), 3.6-3.75 (m, 2H), 4.0-4.15 (m, 1H) 6.45 (dd, 1H), 6.65 (dd, 1H), 7.0 (d, 1H)

step2 – To a solution of 105 (0.12 g, 0.52 mmol) and 66 (0.11 g, 0.47 mmol) and MeCN (10 mL) was added K_2CO_3 (0.13 g, 0.94 mmol) and KI (0.09 g, 0.56 mmol) and the resulting solution was heated at reflux overnight. The mixture was cooled, diluted with water and extracted with EtOAc. The combined EtOAc extracts were washed with brine, dried ($MgSO_4$), filtered and evaporated. The crude product was purified by SiO_2 column chromatography eluting with DCM:MeOH:NH_4OH (120:10:1) to afford 0.13 g (62%) of 106.

step 3 - To a solution of 106 (0.13 g, 0.29 mmol) in DCE (5 mL) were added 1-acetyl-piperidine-4-carbonyl chloride (0.16 g, 0.82 mmol) and pyridine (0.94 mL, 1.17 mL). The solution was heated at 50 °C overnight. The mixture was cooled, diluted with DCM and washed with saturated $NaHCO_3$. The organic layer was dried (Na_2SO_4), filtered and evaporated. The crude product was purified by SiO_2 column chromatography eluting with CH_2Cl_2:MeOH:NH_4OH (150:10:1) to afford 0.12 g (68 %) of II-189: mp 107.9-109.8 °C; ms (ES+) m/z 595 (M + H).

Example 36

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-butyl}-amide (II-36)



step 1 - To a mixture of 3-chloro-4-methyl-phenylamine (2.5 g, 17.65 mmol) and trifluoromethanesulfonimide (0.33 g, 1.20 mmol) in MeCN (20 mL) was added methyl vinyl ketone (1 mL, 12.05 mmol) at RT. After 1 h silica gel and Na_2CO_3 (200 mg) were added to the mixture and it was concentrated *in vacuo*. The crude product was purified

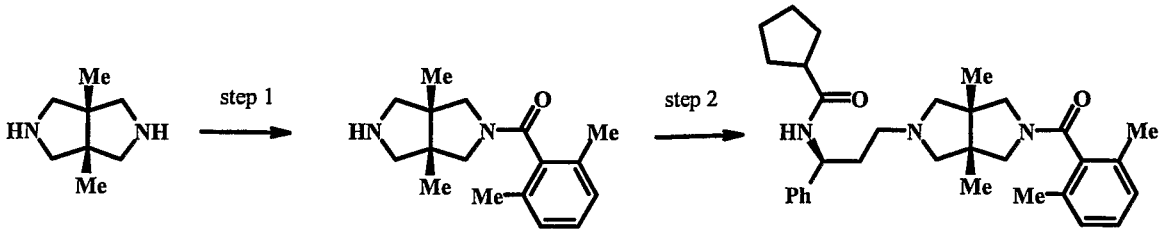
by SiO_2 column chromatography eluting with n-hexane:EtOAc (4:1) to afford 1.3 g (51 %) of **107**: NMR ($CDCl_3$) δ 2.15 (s, 3H), 2.25 (s, 3H), 2.73 (t, 2H), 3.35 (t, 2H), 3.93 (br, 1H), 6.4 (dd, 1H), 6.6 (d, 1H), 6.98 (d, 1H);

step 2 - To a solution of **107** (1.3 g, 6.14 mmol) in DCM (30 mL) were added 1-acetyl-piperidine-4-carbonyl chloride (3.49 g, 18.42 mmol) and TEA (3 mL, 22.09 mmol) at 0° C. After 20 min the solution was heated at 40 °C overnight. The mixture was diluted with DCM and washed sequentially with H_2O, 2N HCl, saturated $NaHCO_3$ and brine. The organic layer was dried (Na_2SO_4), filtered and evaporated. The crude product was purified by SiO_2 column chromatography eluting with 5% MeOH/EtOAc to afford 1.28 g 957 %) of **108**: ^1H NMR ($CDCl_3$) δ 1.6-1.85 (m, 4H), 2.05 (s,3H), 2.45 (s, 3H), 2.68 (t, 2H), 2.85 (t, 1H), 3.28 (d, 1H), 3.85-3.95 (m. 2H), 4.5 (d, 1H), 6.98 (dd, 1H), 7.2 (d, 1H), 7.33 (d, 1H).

step 3 - To a solution of **108** (0.17 g, 0.48 mmol) in THF (7 mL) was added a solution of **54** (0.10 g, 0.40 mmol) in DCM (7 mL). Titanium tetra-isopropoxide (0.26 mL, 0.89 mmol) was added to the mixture. After stirring for 40 min, $NaBH(OAc)_3$ (0.13 g, 0.61 mmol) was added to the mixture and stirring was continued at RT overnight. Saturated $NaHCO_3$ was added to the mixture and it was stirred for 10 min. The mixture was filtered through a CELITE pad and the filtrate was extracted with DCM. The organic layer was dried over ($MgSO_4$) and purified by SiO_2 column chromatography eluting with DCM:MeOH:NH_4OH (150:10:1) afford 0.15 g (64 %) of **II-36**: mp 60.9-62.4 °C; ms (ES+) m/z 594 (M + H).

Example 37

Cyclopentanecarboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-3a,6a-dimethyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide (**I-446**)



110 **111** **I-446**

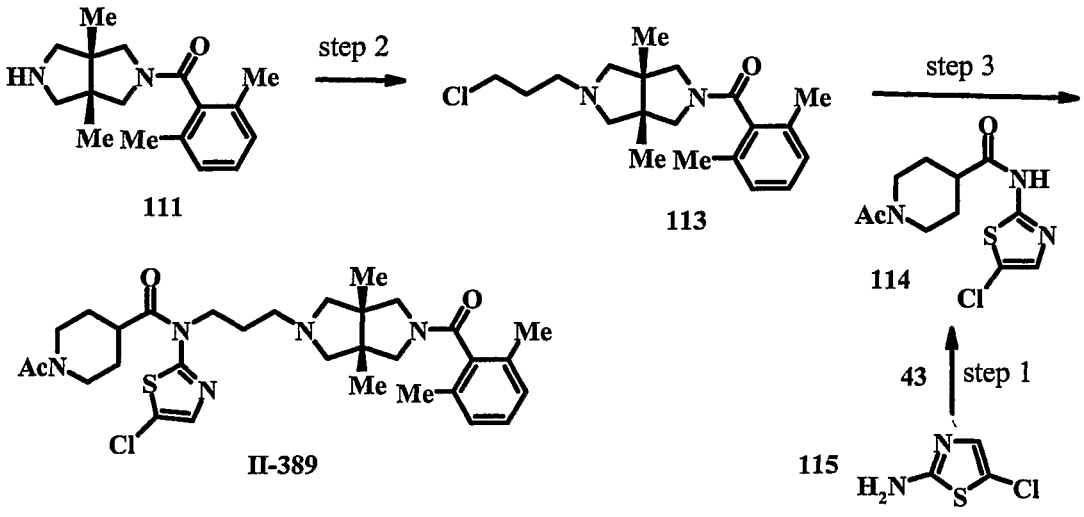
step 1 - Butyllithium 2.5M in hexanes (2 mL, 5 mmol) was added dropwise at 0° C to a suspension of **110** (0.35 g, 3 mmol, prepared as described in *J. Org. Chem.* 1996, 61:8897-8903) () in THF (8 mL). The resulting mixture was stirred at 0° C for 15 min then 2,6-dimethyl-benzoyl chloride (0.421 g, 3 mmol) in THF (2mL) was added. The reaction

mixture was stirred at 0° C for 15 min then quenched by addition of MeOH (7 mL) and evaporated. The residue was partitioned between EtOAc and water. The organic layer was dried ($MgSO_4$), filtered and evaporated. The residue was purified by flash chromatography on silica gel eluting with a gradient from 100% DCM to a 1:1 solution of DCM and DCM/MeOH/NH_4OH 80/10/1 to afford 52 mg (8%) of 111: ^1H NMR ($CDCl_3$) δ 1.01 (s, 3H), 1.11 (s, 3H), 2.26 (s, 6H), 2.83 (d, 1H, *J*= 12 Hz), 2.90 (d, 1H, *J*= 12 Hz), 2.95 (d, 1H, *J*= 12 Hz), 2.96 (d, 1H, *J*= 12 Hz), 3.09 (d, 1H, *J*= 12 Hz), 3.15 (d, 1H, *J*= 12 Hz), 3.54 (d, 1H, *J*= 12 Hz), 3.84 (d, 1H, *J*= 12 Hz), 7.03 (bd, 2H, *J*= 9 Hz), 7.15 (dd, 1H, *J*= 9; 8 Hz) ; MS (ES+) m/z 273 $(M + H)^+$.

step 2 - A mixture of 11 (52 mg, 0.191 mmol), aldehyde 10 (56 mg, 0.228 mmol), sodium triacetoxyborohydride (81 mg, 0.382 mmol) and HOAc (30 □l, 0.524 mmol) in DCM (5 mL) was stirred at RT for 48 h before being partitioned between DCM and a solution of 5% aqueous $NaHCO_3$. The aqueous layer was back extracted three times with DCM. The combined organic layers were dried over magnesium sulfate, filtered and evaporated. The residue was purified by SiO_2 flash chromatography on silica gel eluting with gradient of DCM (100%) to a solution 6:4 solution of DCM and DCM/MeOH/NH_4OH (80/10/1) to afford (**I-446**) (50 mg, 54% theoretical). ^1H NMR (DMSO-d_6) δ TO BE ADDED; ^{13}C NMR (DMSO-d_6) δ 18.86, 18.99, 25.96, 26.01, 30.13, 30.43, 35.74, 41.69, 44.57, 50.50, 51.06, 52.37, 53.12, 60.37, 126.64, 126.68, 126.89, 127.54, 127.61, 128.32, 128.53, 133.06, 138.25, 144.40, 167.72, 174.81 ; MS (ES+) m/z 502 $(M + H)^+$.

Example 38

1-Acetyl-piperidine-4-carboxylic acid (5-chloro-thiazol-2-yl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide (II-389)



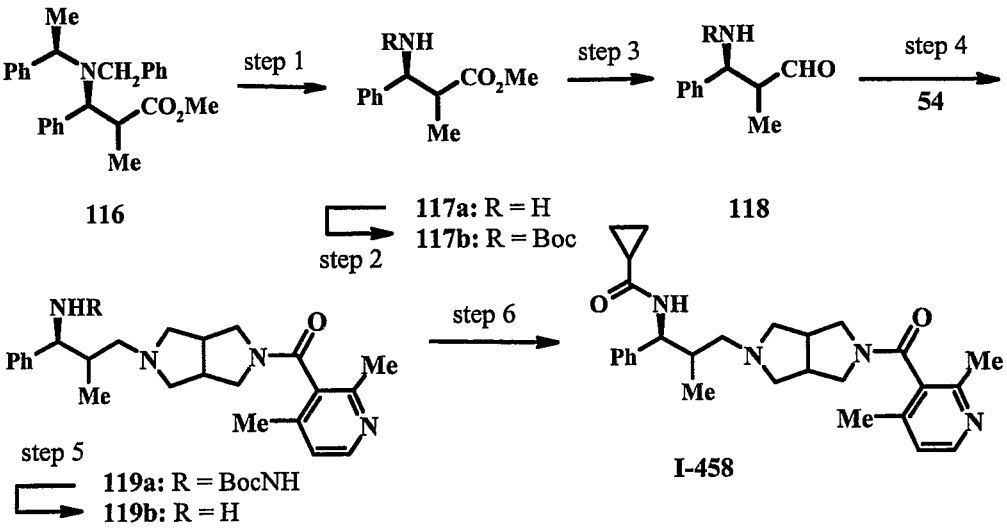
step 1 - To a solution of 5-chloro-thiazol-2-ylamine hydrochloride (115, 216 mg, 1.26 mmol) in DCM (3 mL) was added 1-acetyl-piperidine-4-carbonyl chloride (43, 282 mg, 1.49 mmol) and TEA (0.41 mL, 2.92 mmol). The mixture was stirred for 12 h at RT. The reaction was quenched with water and the mixture was extracted with DCM. The organic layer was dried (Na_2SO_4), concentrated and purified by SiO_2 chromatography on (100% EtOAc) to afford 93 mg (26%) of 114: ^1H NMR ($CDCl_3$, 1 H not observed) δ 1.66-2.04 (m, 4H), 2.12 (s, 3H), 2.54-2.71 (m, 1H), 2.72-2.84 (m, 1H), 3.10-3.25 (m, 1H), 3.85-3.99 (m, 1H), 4.54-4.65 (m, 1H), 7.23 (s, 1H).

step 2 - To a solution of (2,6-dimethyl-phenyl)-(hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-methanone (111, 964 mg, 3.95 mmol) in DMF (7 mL) was added 1-chloro-3-iodo-propane (0.6 mL, 5.58 mmol) and Cs_2CO_3 (1.7 g, 5.21 mol) The mixture was stirred overnight at RT. Water was added to the mixture and the resulting solution was extracted with EtOAc. The organic layer was dried (Na_2SO_4) and concentrated. The crude product was purified by flash column chromatography on silica gel eluting with a DCM: MeOH: NH_4OH gradient (99:0.7:0.07 to 96:3.5:0.35 over 50 min.) to afford 0.828 g (65%) of 113: ms (LCMS) m/z 321 (M + H).

step 3 - 1-Acetyl-piperidine-4-carboxylic acid (5-chloro-thiazol-2-yl)-amide (114, 79 mg, 0.28 mmol) was dissolved in DMF (3 mL) and the resulting solution was cooled to 0° C. NaH (29 mg, 60% dispersion in mineral oil) was added and the mixture was heated to 50° C for 2 h. A solution of 113 (88 mg, 0.28 mmol) and DMF (0.6 mL) was added dropwise and the reaction mixture was stirred at 90° C for 9 h. The mixture was cooled, diluted with water and extracted with DCM. The combined DCM extracts were dried ($MgSO_4$) and concentrated. The crude product was purified by preparative HPLC to afford 20.5 mg (13%) of II-389 as a TFA salt: ^1H NMR (DMSO-d_6) δ 1.4-1.6 (m, 2H), 1.85-1.95 (m, 2H), 2.0 (s, 3H), 2.05-2.22 (m, 9H), 2.55-2.65 (m, 1H), 2.9-3.0 (m, 1H), 3.2-3.3 (m, 7H), 3.5-3.9 (m, 7H), 4.15-4.2 (m, 2H), 7.05-7.2 (m, 3H), 7.6 (s, 1H); ms (ES+) m/z 572 (M + H).

Example 39

(S)-Cyclopropanecarboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-amide (I-458)



step 1 - A solution of (2R, 3S, αR)3-[benzyl-(1-phenyl-ethyl)-amino]-2-methyl-3-phenyl-propionic acid methyl ester (116, 1.00g, 2.58 mmol, prepared as described in *J. Chem. Soc. Perkin Trans. 1* 1994 1129) and MeOH:EtOAc:10% HCl solution (25 mL) containing $Pd(OH)_2$-C (0.50 g) and hydrogenated (1atm) for 24 h. The reaction mixture was filtered through a CELITE® pad to remove the catalyst. The filtrate was concentrated *in vacuo* and the residue partitioned between Et_2O (40 mL) and saturated $NaHCO_3$ solution (25 mL). The organic layer was dried ($MgSO_4$) and concentrated *in vacuo* to afford 408 mg (80%) of 117a as a pale yellow liquid: ms (ES+) m/z 194 $(M+H)^+$.

step 2 - A solution of (2R, 3S)-3-amino-2-methyl-3-phenyl-propionic acid methyl ester (117a, 400 mg, 2.06 mmol) in THF (5 mL) was cooled to 0° C. A cold solution of NaOH (166 mg, 4.14 mmol) in H_2O (3.75 mL) was added to the above solution followed by a solution of $(BOC)_2O$ in THF (2.5 mL) and the mixture stirred at RT for 5 h. The reaction mixture was extracted with EtOAc (2 x 50 mL) and the combined organic extracts were dried ($MgSO_4$) and concentrated *in vacuo* to afford 117b as a waxy solid: ms (ES+) m/z 237 $(M-C_4H_8)^+$.

step 3 - To a solution of 117b (355 mg, 1.21 mmol) in DCM (20 mL) cooled to -78° C was added DIBAL-H (2.42 mL of 1 M DCM solutionn, 2.42 mmol) dropwise at such a rate to maintain the temperature below -70° C. After 2 h the reaction was quenched by the slow addition of MeOH (2 mL) and then allowed to warm to RT. The reaction mixture was filtered through a CELITE® pad. The filtrate was dried (Na_2SO_4) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica eluting with EtOAc:hexane (1:3) to afford 118 as a white solid: ^{1}H-NMR showed this material to be a 1:1.38 ratio of diastereomers.

step 4 - To a solution of 118 (197 mg, 0.75 mmol) and 54 (184 mg, 0.75 mmol) in DCM (16 mL) containing HOAc (0.11 mL) was added $NaBH(OAc)_3$ (191 mg, 0.90

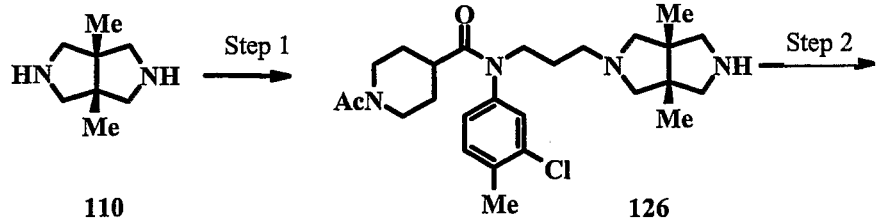
mmol) in 1 portion and the reaction was stirred for 18 h at RT. The reaction was quenched by the addition of 10% K_2CO_3 solution (10 mL) and stirred for 20 min. The product was extracted with DCM (2 x 20 mL) and the combined extracts were dried (Na_2SO_4) and concentrated *in vacuo*. The crude product was purified by flash chromatography on silica eluting with DCM/ 7.5% MeOH (containing 2% NH_4OH) to afford 290 mg (79%) of **119aas** an off-white foam: ms (ES+) m/z 493 $(M+H)^+$.

step 5 - A solution of **119a** (258 mg, 0.52 mmol) dissolved in 10 M HCl in MeOH (8 mL) was heated at 65° C for 2 h. The MeOH was evaporated under reduced pressure and the residue cautiously partitioned between DCM (25 mL) and 20% K_2CO_3 soln (15 mL). The aqueous layer was reextracted with DCM (2 x 20 mL). The combined extracts were dried (Na_2SO_4) and concentrated *in vacuo* to afford 194 mg (95%) of **119b** as a viscous liquid: MS (ES+) m/z 393 $(M+H)^+$.

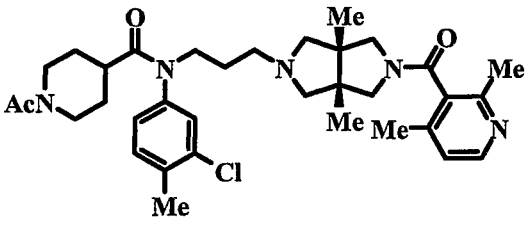
step 6 - To a solution of **119b** (500µL of 0.1 M DCM soln, 0.050 mmol) and DIPEA (0.03mL) was added cyclopropanecarbonyl chloride (6.8 µL, 7.8 mg, 0.075 mmol) and the resulting mixture stirred at room temp for 18 hrs. The reaction mixture was concentrated in a stream of N_2 and purified by reverse phase HPLC to afford **I-458:** ms (ES+) m/z 461 $(M+H)^+$.

Example 40

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-3a,6a-dimethyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide (II-174)



110 126



II-174

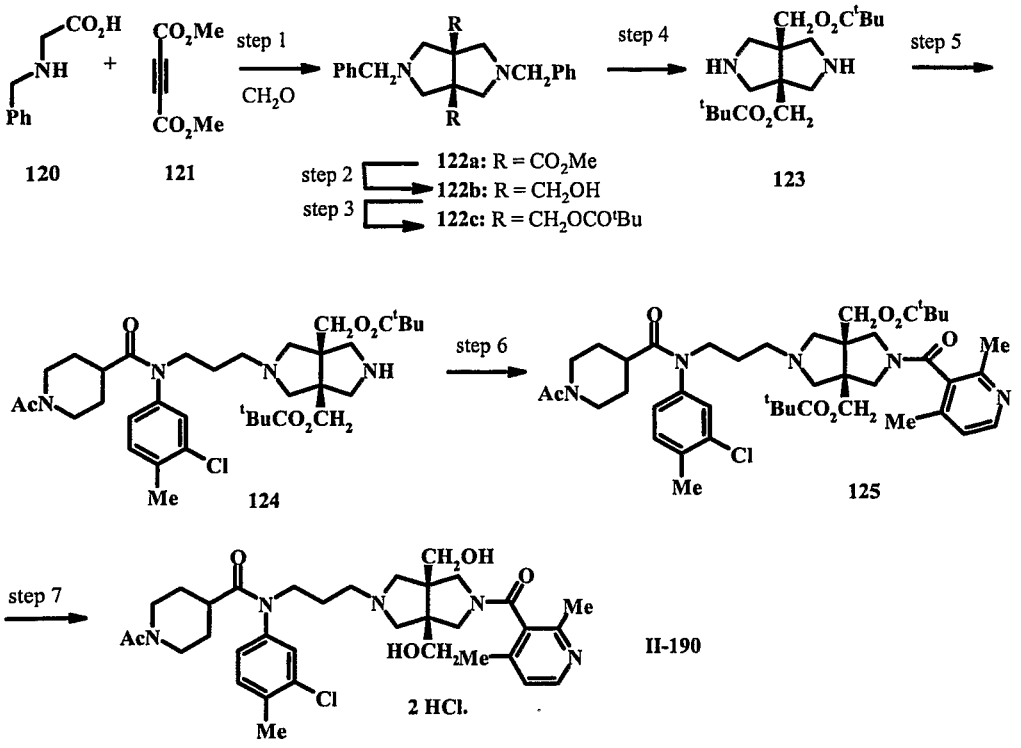
step 1 - A mixture of **110** (1.2 g, 8 mmol), **18** (0.75 g, 2 mmol), KI (0.55 g, 3 mmol) and K_2CO_3 (0.825 g, 6 mmol) in DMF (30 mL) was stirred at 80° C for 16 h before being partitioned between water and EtOAc. The aqueous layer was extracted twice with

EtOAc. The combined organic layers were dried over ($MgSO_4$), filtered and evaporated. The residue was purified by flash chromatography on silica gel eluting with a gradient from 100% DCM to a 1:9 solution of DCM and DCM/MeOH/NH_4OH 60/10/1 to afford 0.100 g (3%) of **126**: ^1H NMR ($CDCl_3$) δ 1.53-2.46 (m, 15H), 2.56 (d, 2H, *J*= 9 Hz), 2.67 (d, 2H, *J*= 9 Hz), 2.78-2.95 (m, 3H), 3.48 (s, 6H), 4.52 (db, 1H, *J*= 15 Hz), 6.97 (dd, 1H, *J*= 9; 3 Hz), 7.18 (d, 1H, *J*= 3 Hz), 7.31 (d, 1H, *J*= 3 Hz); MS (ES+) m/z 608 $(M + H)^+$.

step 2 - DIPEA (73 □L, 0.42 mmol) was added at RT to a solution of **126** (100 mg, 0.21 mmol), 2,4-dimethyl-3-pyridyl carboxylic acid (32 mg, 0.21 mmol), EDCI (48 mg, 0.25 mmol) and HOBT (34 mg, 0.25 mmol) in DMF (1 mL). The resulting mixture was stirred at RT for 16 h then partitioned between EtOAc and water. The aqueous layer was extracted twice with EtOAc. The combined organic layers were dried (Na_2SO_4), filtered and evaporated. The residue was purified by SiO_2 flash chromatography eluting with a gradient of 100% DCM to a 1:1 mixture DCM of a solution of DCM/MeOH/NH_4OH (80/10/1) over 20 minutes, followed by 8/2 DCM/MeOH solution for 10 min, 15 mL/min) to afford 87 mg (64%) of **II-174**:^1H NMR (CD_3OD) δ δ 0.90-3.34 (m, 35H), 3.39-3.84 (m, 4H), 3.92 (m, 1H), 4.52 (bd, 1H, *J*= 12 Hz), 6.97 (m, 2H), 7.17 (bs, 1H), 7.31 (d, 1H, *J*= 6 Hz), 8.36 (d, 1H, *J*= 3 Hz); ^{13}C NMR ($CDCl_3$) δ 19.10, 19.18, 20.19, 20.82, 21.14, 21.77, 22.52, 22.62, 27.05, 28.71, 29.22, 39.76, 41.07, 45.91, 48.41, 49.35, 49.38, 50.12, 50.69, 53.36, 58.30, 60.95, 67.41, 67.72, 123.02, 123.10, 126.69, 128.83, 132.40, 133.10, 135.64, 136.91, 141.50, 143.49, 143.58, 149.17, 149.22, 153.88, 154.06, 167.45, 169.19, 174

Example 41

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[(5-(2,4-dimethyl-pyridine-3-carbonyl)-3a,6a-bis-hydroxymethyl-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide (**II-190**)



step - 1 - Formaldehyde (37 wt.% in water) was added dropwise over 3 h to a refluxing suspension of N-benzylglycine (10 g, 60.37 mmol) and dimethylacetylenedicarboxylate (3.7 mL, 30.19 mmol) in 200 mL of toluene. The resulting mixture stirred and heated to reflux for 14 h while water was continuously removed using a Dean Stark trap. The reraction misxture was cooled to RT and evaporated. The residue was purified by SiO_2 chromatography eluting with a hexane/EtOAc gradient (100 % EtOAc to 75/25) over 40 minutes with and 80 mL/min flow rate to afford 3.4 g (28%) of 122a: ^{1}H NMR ($CDCl_3$) δ 2.71 (d, 2H, *J*= 9 Hz), 3.08 (d, 2H, *J*= 9 Hz), 3.66 (s, 5H), 7.17-7.42 (m, 5H); MS (ES+) m/z 409 $(M + H)^+$.

step 2 - An LAH solution (33 mL, 33 mmol, 1M solution in THF) was added to a solution of 122a (3.4 g, 8.323 mmol) and THF (100 mL) cooled to 0° C and maintained under nitrogen. The resulting reaction mixture was stirred at 0° C for 2 h before being quenched by addition of saturated $NaHCO_3$. The aqueous layer was extracted twice with EtOAc and combined organic layers were dried (Na_2SO_4), filtered and evaporated to afford 2.9 g (99%) of 122b: ^{1}H NMR ($CDCl_3$) δ 2.56 (s, 4H), 3.58 (s, 2H), 3.63 (s, 2H), 7.18-7.38 (m, 5H); MS (ES+) m/z 353 $(M + H)^+$.

step 3 - Trimethyl acetyl chloride (1.7 mL, 13.8 mmol) was added to a solution of 122b (1.9 g, 5.39 mmol) in DCM (25 mL) and pyridine (2mL). The resulting mixture was stirred at RT for 24 h before adding 5 mL of MeOH. After stirring at RT for 1 h, the reaction mixture was diluted with DCM and washed with 1M aqueous HCl, aqueous saturated $NaHCO_3$ and water. The organic layer was dried (Na_2SO_4), filtered and

evaporated. The residue was purified by SiO_2 chromatography eluting with a hexane/EtOAc gradient (100% hexane to 10% EtOAc) to afford 1.14 g (41% theory) of 122c: ^{1}H NMR ($CDCl_3$) δ 1.15 (s, 9H), 2.53 (d, 2H, *J*= 9 Hz), 2.57 (d, 2H, *J*= 9 Hz), 3.58 (s, 2H), 4.12 (s, 2H), 7.18-7.34 (m, 5H); MS (ES+) m/z 521 $(M + H)^+$.

step 4- A mixture of 122c (1.14 g, 2.189 mmol) and $Pd(OH)_2/C$ (0.5 g; 20 wt. % on carbon) in EtOH (40 mL) and 10 mL of cyclohexane was heated to reflux with stirring for 24 h. The reaction mixture was cooled to RT, filtered and evaporated. The residue was dissolved in DCM and 5 mL of 1M HCl in Et_2O were added. The precipitate was filtered off and rinsed with DCM to afford 0.365 g (40%) of 123 as a light brown solid:^{1}H NMR (DMSO-d_6) δ 1.18 (s, 9H), 3.37 (d, 2H, *J*= 9 Hz), 3.60 (d, 2H, *J*= 9 Hz), 4.26 (s, 2H), 10.01 (bs, 1H) ; MS (ES+) m/z 341 $(M + H)^+$.

step 5 - A mixture of 123 (90 mg, 0.242 mmol) and KI (44 mg, 0.266 mmol) in DMF (1 mL) was stirred at 80° C for 8 h then added to a mixture of chloride 18 (0.1 g, 0.242 mmol), DIPEA (110 □L, 0.605 mmol) in DMF (1 mL). The resulting cloudy mixture was stirred at 40° C for 15 h and then at 80° C degrees for 2 h before being cooled to RT and diluted with EtOAc. The organic layer was washed with water and the aqueous layer was back extracted once with EtOAc. Combined organic layers were dried over (Na_2SO_4), filtered and evaporated. The residue was purified by SiO_2 chromatography gel eluting with a gradient from 100% DCM to a 1:1 solution of DCM and DCM/MeOH/NH_4OH 60/10/1 over 20 minutes at a 25 mL/min flow rate to afford 0.064 g (39%) of 124:MS (ES+) m/z 341 $(M + H)^+$.

step 6 - 2,4-Dimethyl-nicotinoyl chloride hydrochloride (39 mg / 188.7 □mol) prepared from 2,4-dimethyl-nicotinic acid was added to a mixture of 124 (64 mg / 94.33 □mol) and DIPEA (30 □L / 188.7 □mol) in 0.5 mL of DCM. The resulting reaction mixture was stirred at RT for 5 h before being diluted with DCM, and washed with water. The aqueous layer was extracted once with DCM. Combined organic layers were dried over sodium sulfate, filtered and evaporated. The residue was purified by SiO_2 chromatography eluting with a gradient from 100% DCM to a 1:1 solution of DCM and DCM/MeOH/NH_4OH 60/10/1 over 20 minutes at a 25 mL/min flow rate to afford 0.062 g (81%) of 125: ^{1}H NMR ($CDCl_3$) δ, mixture of rotamers, selected data: 4.51 (bd, 1H, *J*= 12 Hz), 6.93-7.04 (m, 2H), 7.16 (m, 1H), 7.32 (d, 1H, *J*= 9 Hz), 8.37 (dd, 1H, *J*= 3, 1 Hz) ; MS (ES+) m/z 808 $(M + H)^+$.

step 7 - Sodium methoxide (0.5 mL, 25 wt.% in MeOH) was added at RT to a solution of 125 (0.060 g, 74.21 □mol) in MeOH (4 mL). The resulting mixture was stirred at RT overnight then neutralized by addition of Dowex 50WX8-200. The resin was filtered and rinsed with MeOH. The filtrate was evaporated and the residue was purified

by SiO_2 chromatography (13 g RediSep column, DCM/[DCM/MeOH/NH_4OH 60/10/1] 10/0 to 1/1 over 20 minutes, 25mL/min) to afford 23 mg (48%) of II-190 as a white foam: ^1H NMR ($CDCl_3$) δ 1.51-1.85 (m, 6H), 2.05 (s, 3H), 2.22-2.56 (m, 16H), 2.61 (m, 1H), 2.84 (m, 1H), 3.07 (m, 1H), 3.39 (m, 1H), 3.51 (m, 1H), 3.38-3.57 (m, 8H), 4.51 (bd, 1H, *J*= 12 Hz), 6.96 (dd, 1H, *J*= 9, 1 Hz), 6.99 (dd, 1H, *J*= 3, 1 Hz), 7.17 (m, 1H), 7.32 (d, 1H, *J*= 9 Hz), 8.34 (t, 1H, *J*= 3 Hz); ^{13}C NMR ($CDCl_3$) δ 19.09, 19.31, 20.20, 21.76, 22.41, 22.68, 26.87, 28.72, 29.22, 30.08, 39.73, 41.07, 45.91, 48.16, 52.60, 53.82, 55.07, 55.13, 55.97, 56.05, 56.48, 63.91, 64.16, 123.16, 123.24, 126.61, 128.73, 132.48, 135.72, 137.07, 141.35, 149.09, 149.24, 153.73, 167.70, 167.92, 169.24, 174.37 ; MS (ES+) m/z 640 $(M + H)^+$.

Example 42

Human CCR5 receptor-ligand binding assay protocol

Human CCR5 receptor (Genebank ID: 29169292) was cloned into mammalian expression vector, pTarget (Promega). The construct was transfected into CHO-$G_{\alpha 16}$ cells by using Fugene Reagent (Roche). Clones were selected under antibiotic pressure (G418 and Hygromycin) and sorted 4 times with a fluorescence activates cell sorter and a monoclonal antibody specific for CCR5 receptor (BD Biosciences Pharmigen , Mab 2D7, Cat. No. 555993). The clone with highest expression (100,000 copies per cell) was chosen for the binding assays.

Adherent cells in 225 mL tissue culture flask (~90% confluent) were harvested using 1 mM EDTA in PBS (phosphate-buffered saline) without Ca^{2+} and Mg^{2+} . Cells were washed twice with PBS containing no Ca^{2+} and Mg^{2+}. CHO-$G_{\alpha 16}$-hCCR5 cells were then resuspended (1×10^6/ml) in ice cold binding buffer (50 mM HEPES, 1 mM $CaCl_2$, 5 mM $MgCl_2$, 0.5% BSA, 0.05% NaN_3,

pH 7.24), pH 7.4), supplemented with freshly made 0.5% BSA and 0.05% NaN_3.

Eighty µl CHO-$G_{\alpha 16}$ -hCCR5 (1×10^6/ml) cells were added to 96 well plates. All dilutions were made in binding buffer (50 mM HEPES, 1 mM $CaCl_2$, 5 mM $MgCl_2$, 0.5% BSA, 0.05% NaN_3,

pH 7.24).

The plates were incubated on a cell shaker at RT for 2 h with a final concentration of 0.1 nM ^{125}I RANTES or ^{125}I MIP-1α or ^{125}I MIP-1β. The compound dilutions were made in PBS, 1% BSA. Total reaction volume was 100 µl per well. The test compounds were added to the cells prior to the addition of radioligand.

After incubation, the cells were harvested onto GF/C filter plates using Packard cell harvester. Filters were pretreated with 0.3% PEI /0.2% BSA for 30 min. The filter plate was washed rapidly 5 times with 25 mM HEPES, 500 mM NaCl, 1 mM $CaCl_2$ and 5mM $MgCl_2$ adjusted to pH 7.1. Plates were dried in oven (70° C) for 20 min, added with 40 µl scintillation fluid and sealed with Packard TopSeal-A. Packard Top Count was used to measure of the radioactivity for 1 min per well.

Total binding was determined with control wells added with radioisotope and buffer and the non-specific binding was determined using an excess cold RANTES to some of the control wells. Specific binding was determined by subtracting the non-specific form total binding. Results are expressed as the percentage of specific ^{125}I RANTES binding. IC_{50} values were determined using varying concentrations of the test ligand in triplicates and the data was analyzed using GraphPad Prism (GraphPad, San Diego, CA).

Compound No.	Binding IC_{50} (□M) RANTES	Compound No.	Binding IC_{50} (□M) RANTES
II-303	0.1019	II-279	0.0015
I-466	0.011	I-39	0.0032
I-449	0.001	II-152	0.0022
II-3	0.0184	II-355	0.0027
I-78	0.037	II-182	0.0028
II-1	0.0042	I-52	0.0028
I-29	0.016	II-175	0.0029
I-4	0.0191	I-47	0.0037
I-14	0.0019		

Example 43

Pharmaceutical compositions of the subject Compounds for administration via several routes were prepared as described in this Example.

Composition for Oral Administration (A)

Ingredient	% wt./wt.
Active ingredient	20.0%
Lactose	79.5%
Magnesium stearate	0.5%

The ingredients are mixed and dispensed into capsules containing about 100 mg each; one capsule would approximate a total daily dosage.

Composition for Oral Administration (B)

Ingredient	% wt./wt.
Active ingredient	20.0%
Magnesium stearate	0.5%
Crosscarmellose sodium	2.0%
Lactose	76.5%
PVP (polyvinylpyrrolidine)	1.0%

The ingredients are combined and granulated using a solvent such as methanol. The formulation is then dried and formed into tablets (containing about 20 mg of active compound) with an appropriate tablet machine.

Composition for Oral Administration (C)

Ingredient	% wt./wt.
Active compound	1.0 g
Fumaric acid	0.5 g
Sodium chloride	2.0 g
Methyl paraben	0.15 g
Propyl paraben	0.05 g
Granulated sugar	25.5 g
Sorbitol (70% solution)	12.85 g
Veegum K (Vanderbilt Co.)	1.0 g
Flavoring	0.035 ml
Colorings	0.5 mg
Distilled water	q.s. to 100 ml

The ingredients are mixed to form a suspension for oral administration.

Parenteral Formulation (D)

Ingredient	% wt./wt.
Active ingredient	0.25 g
Sodium Chloride	qs to make isotonic
Water for injection to	100 ml

The active ingredient is dissolved in a portion of the water for injection. A sufficient quantity of sodium chloride is then added with stirring to make the solution

isotonic. The solution is made up to weight with the remainder of the water for injection, filtered through a 0.2 micron membrane filter and packaged under sterile conditions.

Suppository Formulation (E)

Ingredient	% wt./wt.
Active ingredient	1.0%
Polyethylene glycol 1000	74.5%
Polyethylene glycol 4000	24.5%

The ingredients are melted together and mixed on a steam bath, and poured into molds containing 2.5 g total weight.

Topical Formulation (F)

Ingredients	grams
Active compound	0.2-2
Span 60	2
Tween 60	2
Mineral oil	5
Petrolatum	10
Methyl paraben	0.15
Propyl paraben	0.05
BHA (butylated hydroxy anisole)	0.01
Water	q.s. 100

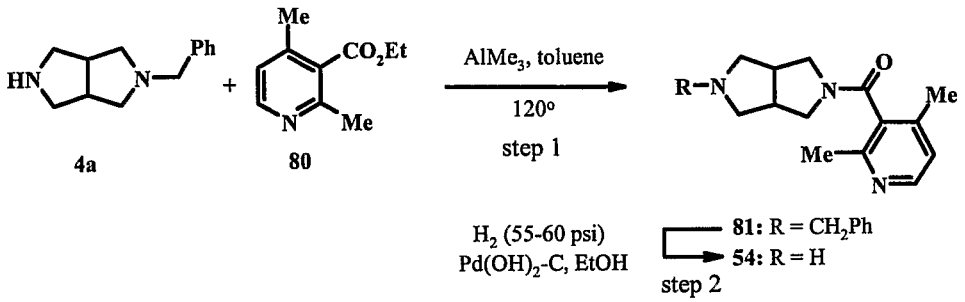
All of the ingredients, except water, are combined and heated to about 60°C with stirring. A sufficient quantity of water at about 60°C is then added with vigorous stirring to emulsify the ingredients, and water then added q.s. about 100 g.

Nasal Spray Formulations (G)

Several aqueous suspensions containing from about 0.025-0.5 percent active compound are prepared as nasal spray formulations. The formulations optionally contain inactive ingredients such as, for example, microcrystalline cellulose, sodium carboxymethylcellulose, dextrose, and the like. Hydrochloric acid may be added to adjust pH. The nasal spray formulations may be delivered via a nasal spray metered pump typically delivering about 50-100 microliters of formulation per actuation. A typical dosing schedule is 2-4 sprays every 4-12 hours.

Referential Example

(2,4-Dimethyl-pyridin-3-yl)-(hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-methanone (54)



step 1 - A 25 mL 3-neck flask equipped with stir bar, addition funnel, N_2 inlet tube and reflux condenser was charged with **4a** (1.0g, 4.94 mmol) and (dry) toluene (4 mL) and cooled to 0° C. Trimethylaluminum (2.5 mL of a 2.0 M toluene solutionn, 5.0 mmol) was added dropwise, then allowed to warm to RT. 2,4-Dimethyl-nicotinic acid ethyl ester (**80**, 0.93 g, 5.2 mmol) in toluene (1 mL) was added dropwise then heated at 120° C for 18 h. The reaction mixture was cooled in an ice bath and MeOH (2 mL) was added. The mixture was brought to RT, heated at reflux for 10 min then recooled to RT. The mixture was filtered through a CELITE® pad. The filtrate was washed with brine, dried ($MgSO_4$) then concentrated *in vacuo*. The crude product was purified by flash chromatography on silica eluting with 7.5% MeOH (containing 2% NH_4OH) / DCM to afford 1.17 g, (71%) of **81** as a pale yellow liquid: ms (ES+) m/z 336 $(M+H)^+$.

step 2 - A solution of **81** (5.55g , 16.5 mmol) in EtOH (250 mL) containing 20% $Pd(OH)_2$ (2.75 g) was hydrogenated at 55 psi to 60 psi for 18 hrs. The mixture was filtered through a CELITE® pad to remove the catalyst and the filtrate concentrated *in vacuo* to afford 3.88 g (95%) **54** as a dark viscous liquid: ms (ES+) m/z 246 $(M+H)^+$.

The features disclosed in the foregoing description, or the following claims, expressed in their specific forms or in terms of a means for performing the disclosed function, or a method or process for attaining the disclosed result, as appropriate, may, separately, or in any combination of such features, be utilized for realizing the invention in diverse forms thereof.

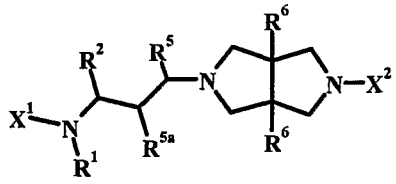
The foregoing invention has been described in some detail by way of illustration and example, for purposes of clarity and understanding. It will be obvious to one of skill in the art that changes and modifications may be practiced within the scope of the appended claims. Therefore, it is to be understood that the above description is intended to be illustrative and not restrictive. The scope of the invention should, therefore, be determined not with reference to the above description, but should instead be

determined with reference to the following appended claims, along with the full scope of equivalents to which such claims are entitled.

All patents, patent applications and publications cited in this application are hereby incorporated by reference in their entirety for all purposes to the same extent as if each individual patent, patent application or publication were so individually denoted.

Claims

1. A compound according to formula I wherein:



(I)

X^1 is selected from the group consisting $-C(=O)R^3$, $-S(=O)_2R^3$ and $-C(=O)OR^3$;

X^2 is selected from the group consisting $-C(=O)R^4$, $-S(=O)_2R^4$, benzyl or $-CH_2$(pyridin-3-yl) said benzyl optionally substituted with halogen or C_{1-3} alkyl;

one of R^1 and R^2 is:

(i) phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of halogen, C_{1-6} alkyl and C_{1-6} alkoxy;

(ii)heteroaryl selected from the group consisting of pyridinyl, pyridinyl-N-oxide, pyrimidinyl and thiazolyl said heteroaryl optionally substituted with one to three substituents independently selected from the group consisting of halogen, C_{1-6} alkyl, C_{1-6} haloalkyl and C_{1-6} alkoxy; and,

the other of R^1 and R^2 is hydrogen;

R^3 is selected from the group consisting of:

(i) phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of:

(a) C_{1-10} alkyl,

(b) C_{1-10} heteroalkyl,

(c) C_{1-6} haloalkyl,

(d) C_{1-6} alkoxy,

(e) C_{1-6} thioalkyl,

(f) amino,

(g) C_{1-6} alkyl amino,

(h) C_{1-6} dialkylamino,

(g) C_{1-6} acylamino,

(i) carbamoyl, N-C_{1-6} alkylcarbamoyl or N,N-C_{1-6} dialkylcarbamoyl,

(j) ureido,

(k) nitro,

(l) cyano,

(m) halogen,

(n) C_{1-6} alkylsulfonyl,

(o) sulfamoyl, N-C_{1-6} alkylsulfamoyl or N, N-C_{1-6} dialkylsulfamoyl,

(p) C_{1-6} alkylsulfonamido or optionally substituted phenylsulfonamido,

(q) optionally substituted phenoxy,

(r) optionally substituted heteroaryloxy, and,

(s) $-Y(CH_2)_nR^{11}$ wherein R^{11} is selected from the group consisting of cyano, $-CO_2R^{12}$, $-CONR^{12}R^{13}$, $-SO_2N^{12}R^{13}$, $-NHSO_2R^{12}$ and $-NHSO_2NR^{12}R^{13}$,

(t) CO_2R^{12},

(u) C_{1-6} acyloxy,

(v) C_{1-6} alkylcarbonyl, and,

(w) C_{1-6} haloalkoxy;

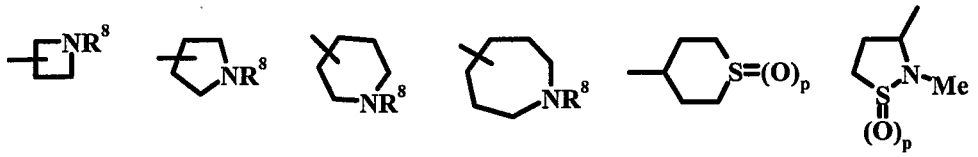
(ii) phenyl C_{1-6} alkyl wherein phenyl as described in (i) above;

(iii)heteroaryl selected from the group consisting of pyridinyl, pyridinyl-N-oxide, pyrimidinyl, pyrazinyl, pyridazinyl, isoxazolyl, isothiazolyl, pyrrolyl, furyl, thiazolyl, oxazolyl, pyrazolyl, oxadiazolyl, thiadiazolyl, imidazolyl, triazolyl, tetrazolyl, indolyl, quinolinyl, isoquinolinyl, 2,4-dimethyl-6-oxo-6H-pyranyl and thienyl said heteroaryl optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, phenyl, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, C_{1-6} alkoxycarbonyl, carbamoyl, acetyl,

nitro, cyano, amino, C_{1-3} alkylamino, C_{1-3}dialkylamino, N-morpholino, sulfamoyl and halogen,

(iv) heteroaryl C_{1-6} alkyl wherein heteroaryl is as described above,

(v) heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is selected from the group consisting of:

(i) C_{1-6} alkyl optionally substituted with one or two groups independently selected from the group consisting of C_{1-3} alkoxy, C_{1-3} acyloxy, hydroxyl, phenyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, aminocarbonylpyridyl wherein said aminocarbonylpyridyl can optionally be the N-oxide and aminobenzoyl wherein said phenyl or said benzoyl radical is optionally substituted with one to three groups independently selected from the group consisting of amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, hydroxyl, halogen and C_{1-3} alkoxy,

(ii) phenyl optionally substituted with one or two groups independently selected from the group consisting of sulfamoyl, acetylamino, halogen, C_{1-6} alkyl, C_{1-6} haloalkyl, and C_{1-6}alkoxy,

(iii) NH_2, C_{1-6} alkylamino, C_{1-6} dialkylamino,

(iv) $C(=O)NH_2$, $C(=O)OC_{1-6}$ alkyl, or C(=O)OH,

(v) 1H indole-3-carbonyl,

(vi) furyl, pyridinyl, or pyridinyl N-oxide,

(vii) N-acetylpiperidin-4-yl,

(viii) furfuryl,

(ix) C_{3-8} cycloalkyl,

(x) C_{1-6} haloalkyl,

(xi) phenyl C_{1-3} alkyl, and,

(xii) C_{1-6} alkoxy;

R^{10} is selected from the group consisting of:

(i) C_{1-6} alkyl,

(ii) C_{1-3} haloalkyl,

(iii) C_{3-8} cycloalkyl, and,

(iv) NH_2, C_{1-3} alkylamino, C_{1-3} dialkylamino;

(vi) heterocycle C_{1-6} alkyl wherein heterocycle is as defined above;

(vii) hydrogen;

(viii) C_{1-10} alkyl optionally independently substituted in each occurrence with one to three substituents independently selected at each occurrence from the group consisting of hydroxy, halogen, C_{1-6} alkoxy, C_{1-6} thioalkyl, C_{1-6} acyloxy, C_{1-6} alkoxycarbonyl, C_{1-6} acylamino, $NR^{14a}R^{14b}$, cyano, C_{1-6} alkylsulfonyl, phenyl, N-methyl-methyl-sulfonamido, N-piperidinyl, N-pyrrolidinyl, imidazolyl, 2-aza-bicyclo[2.2.1]hept-2-yl, carbamoyl, C_{1-6} alkoxycarbonyl, and optionally substituted phenyl as defined in R^3 (*i*) above;

(ix) C_{3-7} cycloalkyl or [3.1.0]bicyclohexyl, 4-oxo-cyclohexyl or 3-oxo-cyclobutyl said cycloalkyl optionally substituted with 1 to 4 fluorine, cyano, hydroxyl, C_{1-3} alkyl or phenyl;

(x) C_{1-3} alkyl-C_{3-7} cycloalkyl said cycloalkyl optionally substituted with 1 to 4 fluorine, C_{1-3} alkyl or phenyl;

(xi) $NR^{14a}R^{14b}$; and,

(xii) $C(=O)OR^{14c}$;

R^4 is selected from the group consisting of:

(i) aryl selected from the group consisting of:

(a) phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of a phenyl substituent (i)(a) to (i)(w) as described in R^3, C_{1-6} haloalkoxy, pyridinyl, 2-oxo-pyrrolidin-1-yl, 2,5-dimethyl-pyrrolidin-1-yl, 3-methyl-5-oxo-4,5-dihydro-pyrazol-1-yl and $NR^{14a}R^{14b}$ wherein R^{14a} and R^{14b} taken together along with the nitrogen to which they are attached are $(CH_2)_r$ or $[(CH_2)_2X^3(CH_2)_2]$, $-NMe(CH_2)_2OH$;

(b) 2,3-dihydrobenzofuran-4-yl,

(c) benzofuran-4-yl,

(d) 2,3-dihydrobenzofuran-7-yl,

(e) quinolin-8-yl,

(f) 1,2,3,4-tetrahydro-isoquinolin-5-yl,

(g) N-acetyl-1,2,3,4-tetrahydro-isoquinolin-5-yl,

(h) N-Boc-1,2,3,4-tetrahydro-isoquinolin-5-yl,

(i) 1,2,3,4-tetrahydro-isoquinolin-8-yl,

(j) N-Boc-1,2,3,4-tetrahydro-isoquinolin-8-yl,

(k) isoquinolin-7-yl,

(l) 1H-indol-5-yl,

(m) 2-acetyl-1,2,3,4-tetrahydro-isoquinolin-5-yl, and

(n) 6-fluoro-benzo[1,3]dioxin-8-yl, and,

(o) quinolin-6-yl;

(ii) phenyl C_{1-6} alkyl wherein phenyl as described above;

(iii)heteroaryl selected from the group consisting of pyridinyl, 2-azetidin-1-yl, pyridinyl N-oxide, pyrimidinyl, pyrazinyl, pyridazinyl, isoxazolyl, isothiazolyl,

pyrrolyl, furyl, thiazolyl, oxazolyl, pyrazolyl, oxadiazolyl, thiadiazolyl, imidazolyl, triazolyl, tetrazolyl, indolyl, quinolyl, isoquinolyl, quinoxalin-2-yl, indazole, benzofuranyl, 4,5,6,7-tetrahydro-benzofuranyl, 2,4-dimethyl-6-oxo-6H-pyranyl, benzo[b]thiophenyl, 4,5,6,7-tetrahydroindazole, 1,4,5,6-tetrahydrocyclopentylpyrazolyl, imidazo[2,1-b]thiazolyl, 6-fluoro-4H-benzo[1,3]dioxin-8-yl, cinnolin-4-yl, and thienyl said heteroaryl optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of substituents (i)(a) to (i)(w) as described for R^3 above, benzyl, pyridinyl, 2-methyl-thiazolyl, acetyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, aminobenzyl, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, C_{1-6} alkylsulfonyl, C_{1-6} alkoxycarbonyl, carboxyl, carbamoyl, nitro, cyano, sulfamoyl, $-OCH_2CO_2R^{14c}$, $-SCH_2CO_2R^{14c}$ and halogen;

(iv) heteroaryl C_{1-6} alkyl wherein heteroaryl is as described above;

(v) heterocycle selected from the group consisting of **IIa-f** as described in (v) of R^3 above, furyl, oxetanyl, tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, piperazinyl, imidazolinyl, N-benzyl-morpholine, N-methyl-1,2,3,4-tetrahydroquinolin-2-yl, 4,5-dihydro-pyrazolyl, and 1,2,3,4-tetrahydroisoquinolinyl said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, acetyl, hydroxy, C_{1-6} alkylcarbonyl, C_{1-6} alkoxycarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo);

(vi) heterocycle C_{1-6} alkyl wherein heterocycle is as defined above;

(vii) hydrogen;

(viii) C_{1-10} alkyl substituted with one to three substituents independently selected at each occurrence from the group consisting of hydroxy, halogen, C_{1-6} alkoxy, C_{1-6} thioalkyl, acyloxy, C_{1-6} acylamino, $NR^{14a}R^{14b}$, cyano, C_{1-6} alkylsulfonyl, N-methyl-methyl-sulfonamido, phenyl, N-piperidinyl, N-pyrrolidinyl, imidazolyl, carbamoyl and C_{1-6} alkoxycarbonyl;

(ix) C_{3-7} cycloalkyl optionally substituted with 1 to 4 fluorine, C_{1-3} alkyl or phenyl;

(x) C_{1-3} alkyl-C_{3-7} cycloalkyl optionally substituted with 1 to 4 fluorine, C_{1-3} alkyl or phenyl; and,

(xi) $NR^{14a}R^{14b}$;

R^5 is hydrogen, C_{1-6} alkyl, C_{1-6} haloalkyl, C_{1-6} hydroxyalkyl, methoxymethyl, $-CO_2R^{12}$ or $CONR^{12}R^{13}$;

R^{5a} is hydrogen or C_{1-6} alkyl;

R^6 is independently selected from the group consisting of hydrogen, C_{1-10} alkyl, hydroxy, C_{1-6} alkoxy, C_{1-3} hydroxyalkyl and C_{1-3} alkoxy-C_{1-3} alkyl;

R^7 is hydrogen or Boc;

R^{12} and R^{13} (i) are independently hydrogen, C_{1-6} alkyl or C_{1-6} heteroalkyl, or (*ii*) R^{12} and R^{13}, when both attached to a nitrogen atom, together can be C_{2-5} alkylene;

R^{14a} and R^{14b} (i) taken independently are R^3, or (ii) taken together along with the nitrogen to which they are attached are $(CH_2)_r$ or $[(CH_2)_2X^3(CH_2)_2]$;

R^{14c} is hydrogen or C_{1-6} alkyl;

R^{15} is hydrogen, methyl, *iso*-propyl, *iso*-butyl, *sec*-butyl, $-CH_2OH$, $-CH(OH)CH_3$, $-CH_2SH$, $-CH_2CH_2SMe$, $-(CH_2)_sCOR^{16}$ wherein R^{16} is -OH or $-NH_2$ and s is 1 or 2, $-(CH_2)_t-NH_2$ where t is 3 or 4, $-(CH_2)_3-NHC(=NH)NH_2$, $-CH_2C_6H_5$, $-CH_2$-*p*-C_6H_4-OH, (3-indolinyl)methylene or (4-imidazolyl)methylene;

X^3 is $-O-$, $-S(O)_p-$, NR^{14c};

Y is a direct bond, -O-, -S- or $-NR^{12}-$;

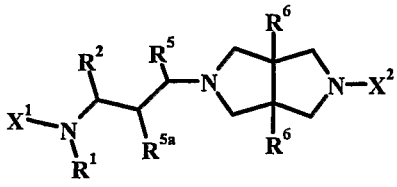
n is an integer from 1 to 6;

p is an integer from 0 to 2;

r is an integer from 3 to 6; and,

pharmaceutically acceptable salts, hydrates, solvates and stereoisomers thereof.

2. A compound according to formula I



(I)

wherein:

one of R^1 and R^2 is:

phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of halogen and C_{1-6} alkyl;

heteroaryl selected from the group consisting of pyridinyl, pyrimidinyl and thiazolyl said heteroaryl optionally substituted with one to three substituents independently selected from the group comprising halogen, C_{1-6} alkyl and C_{1-6} haloalkyl; and,

the other of R^1 and R^2 is hydrogen;

R^5 is hydrogen, C_{1-6} alkyl methoxymethyl or C_{1-3} hydroxyalkyl;

R^{5a} is hydrogen, hydroxyl or C_{1-6} alkyl;

R^6 is hydrogen, C_{1-6} alkyl or C_{1-3} hydroxyalkyl;

X^1 is selected from the group consisting of $-C(=O)R^3$, $-S(=O)_2R^3$ and $-C(=O)OR^3$;

X^2 is selected from the group consisting of $-C(=O)R^4$, $-S(=O)_2R^4$ or benzyl, said benzyl optionally substituted with halogen or C_{1-3} alkyl;

R^3 is selected from the group consisting of:

hydrogen, C_{1-6} alkyl, $-CH_2NH2$, $-CH_2CH_2NH_2$, $-C(CH_3)NH_2$, $-CH_2CH_2N(ethyl)_2$,, $-N(CH_3)_2$, $-N(CH_2CH_3)_2$, -N(iso-propyl)$_2$, $-NCH_3$-benzyl, $-NHCH_2COOCH_2CH_3$, $-CH_2NHCOCH_3$, $CH_2CH_2CONH_2$, $-C(NH_2)CH_2C(CH_3)_2$, $-CH(NH_2)CH(CH_3)_2$,-CH_2CN,

$-C_{1-6}$ alkyl-OH, $-CH_2CH_2COOCH_3$, $-C(CH_3)_2OCOCH_3$, $-CH_2OCOCH_3$, -$CH_2SO_2CH_3$,

$-CH_2(CH_3)SO_2CH_3$,

heteroalkyl, said heteroalkyl is optionally substituted with $-NHCOCH_3$,

$-(CH_2)_t$-C_{3-8} cycloalkyl, wherein t is 0 to 2, and

said $-C_{3-8}$ cycloalkyl is optionally substituted with one to three substituents selected independently from the group comprising of C_{1-6} alkyl, halogen, hydroxyl, cyano, oxo, phenyl,

$-(CHR^m)_u$-phenyl, wherein R^m is H, hydroxyl or C_{1-6} alkyl, u is 0 to 3, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of:

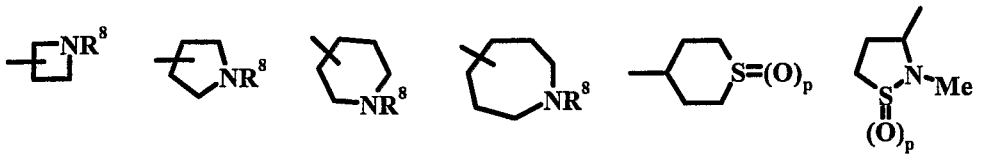
C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, -SO_2NH_2,
-$NHCOCH_3$ or -$COOCH_3$,

-$(CH_2)_v$-heteroaryl selected from the group consisting of pyridinyl, pyridinyl-N-oxide, pyrimidinyl, pyrazinyl, pyridazinyl, isoxazolyl, isothiazolyl, pyrrolyl, furyl, thiazolyl, oxazolyl, pyrazolyl, oxadiazolyl, thiadiazolyl, imidazolyl, triazolyl, tetrazolyl, indolyl, quinolinyl, isoquinolinyl, 2,4-dimethyl-6-oxo-6H-pyranyl and thienyl,

wherein v 0 to 3, and

said heteroaryl is optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, phenyl, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, C_{1-6} alkoxycarbonyl, carbamoyl, acetyl, nitro, cyano, amino, C_{1-3} alkylamino, C_{1-3} dialkylamino, N-morpholino, sulfamoyl and halogen,

-$(CHR^n)_w$-heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl,

wherein R^n is hydrogen or C_{1-6} alkyl, w is 0 to 3, and

said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is C_{1-6} alkyl, C_{1-6} haloalkyl, furanyl, C_{1-6} alkoxy, C_{3-5} cycloalkyl, C_{1-6} alkylamino, -COO- C_{1-6} alkyl, -COOH, $-CH_2OCH_2OOH$, $-CH_2CH_2COOH$;

R^{10} is C_{1-6} alkyl, C_{1-6} dialkylamino, C_{3-8} cycloalkyl;

R^7 is hydrogen or Boc;

R^{15} is hydrogen or C_{1-6} alkyl;

$-NH-(CH_2)_x$-phenyl, wherein x is 0 or 1, and

said phenyl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, $-OCF_3$, heteroalkyl, haloalkyl, acetyl, dialkylamin, cyano, -COOH, $COOCH_2CH_3$,

$-NH(CH_2)_y$-heteroaryl, wherein y is 0 or 1, and

said heteroaryl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, haloalkyl,

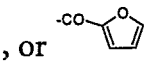
-NH-heterocycle, wherein said heteroaryl is optionally substituted with 1 to 2 substituents independently selected in each occurrence from the group consisting of $-COCF_3$, $COOCH_2$-phenyl,

$-NH(CH_2)_z-C_{3-8}$ cycloalkyl, wherein z is 0 or 1;

R^4 is selected from the group consisting of:

C_{1-6} alkyl, $-CH_2CH_2CONH_2$, $-CH_2NCH_3SO_2CH_3$, $-CH(CH(CH_3)_2)NHCOCH_3$,

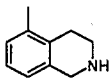
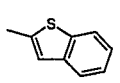
$-(CH_2)_eC_{3-8}$ cycloalkyl, wherein e is 0 to 2, and said cycloalkyl is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl,

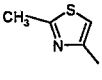
C_{3-8} heterocycle, wherein said heterocyle is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxyl, -CO- C_{1-6} alkyl, oxo, benzyl, , or ,

$-(CHR^f)_f$-phenyl, wherein R^f is hydrogen or –NHCOCH3, f is 0 or 1, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10}

heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, NO_2, -CO-C_{1-6} alkyl, $-COOCH_3$, $COOC(CH_3)_3$, $-NHSO_2CH_3$, NH_2, $-N(CH_3)_2$, $-N(CH_3)CH_2CH_2OH$, $-NHCOCH_3$,$-OCF_3$ $-OCH_2COOH$, pyridine-4-yl,  ;

$(CH_2)_g$-heteroaryl, wherein g is 0 or 1, sais heteroary selected from the group consisting of furanyl, thiophenyl, isoxazolyl, thiazolyl, oxazolyl, pyrroly, pyrazolyl, imidazolyl, indolyl, benzofuranyl, pyranyl, pyridinyl, pyrimidinyl, chinoxalinyl, chinolinyl, , , , , and ,

said heteroaryl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-6} alkyl, oxo, oxido, heteroalkyl, halogen, haloalkyl, methoxy-phenyl, haloalkyl, $-NH_2$, $-N(CH_3)_2$, -NHCH2-phenyl, -NHCOCH3, cyano, $-SO_2NH_2$, $-SO_2CH_3$, $-OCH_2COOCH_3$, $-OCH_2COOH$, $-SCH_2COOH$, $-COCH_3$, $-COOC(CH_3)_3$, $-COOC(CH_3)_3$, $-COOCH_2CH_3$,- COOH, pyridine-4-yl, , , benzyl, phenyl, said phenyl is optionally substituted with one to three substitutuents consisting of C_{1-6} alkyl , halogen, C_{1-6} alkoxyl, $-COOCH_3$;

$-(NR^h)-(CH_2)_h$-phenyl, wherein R^h is hydrogen or C_{1-6} alkyl, h is 0 or 1, and said phenyl is optionally substituted with one to three substituents selected from C_{1-6} alkyl, and

pharmaceutically acceptable salts, hydrates, solvates and stereoisomers thereof.

3. A compound according to claim 2, wherein:

R^2 is phenyl optionally substituted with one to four halogen, or pyridinyl;

R^1 is hydrogen;

R^5 is hydrogen, C_{1-6} alkyl or methoxymethyl, C_{1-3} hydroxyalkyl;

R^{5a} is hydrogen hydroxyl or C_{1-6} alkyl;

R^6 is hydrogen, C_{1-6} alkyl or C_{1-3} hydroxyalkyl;

X^1 is selected from the group consisting of $-C(=O)R^3$, $-S(=O)_2R^3$ and $-C(=O)OR^3$;

X^2 is selected from the group consisting of $-C(=O)R^4$, $-S(=O)_2R^4$ or benzyl, said benzyl optionally substituted with halogen or C_{1-3} alkyl;

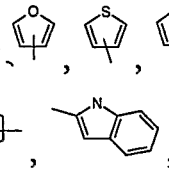
R^3 is selected from the group consisting of:

hydrogen, C_{1-6} alkyl, $-CH_2NH2$, $-CH_2CH_2NH_2$, $-C(CH_3)NH_2$, $-CH_2CH_2N(ethyl)_2$, $-N(CH_3)_2$, $-N(CH_2CH_3)_2$, -N(iso-propyl)$_2$, $-NCH_3$-benzyl, $-NHCH_2COOCH_2CH_3$, -NH-phenyl, $-CH_2NHCOCH_3$, $CH_2CH_2CONH_2$, $-C(NH_2)CH_2C(CH_3)_2$, $-CH(NH_2)CH(CH_3)_2$,-CH2CN, $-C_{1-6}$ alkyl-OH, $-CH_2CH_2COOCH_3$, $-C(CH_3)_2OCOCH_3$,
$-CH_2OCOCH_3$, -CH2SO2CH3, -CH2(CH3)SO2CH3, heteroalkyl,

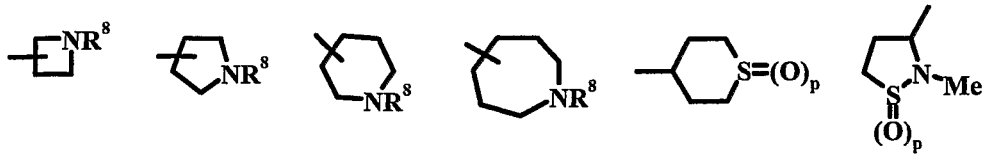
$-(CH_2)_t$-C_{3-8} cycloalkyl selected from cyclopropanyl, cyclobutanyl, cyclopentanyl, cyclohexanyl, cycloheptanyl and , wherein t is 0 to 2, and
said $-C_{3-8}$ cycloalkyl is optionally substituted with one to three substituents selected independently from the group comprising of C_{1-6} alkyl, halogen, hydroxyl, cyano, oxo, phenyl,

$-(CHR^m)_u$-phenyl, wherein R^m is H, hydroxyl or C_{1-6} alkyl, u is 0 to 2, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, $-SO_2NH_2$, $-NHCOCH_3$ or $-COOCH_3$,

$-(CH_2)_v$-heteroaryl selected from the group consisting of , wherein v is 0 to 3, and said heteroaryl is optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, halogen, phenyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, $-NH_2$, SO2NH2, -COCH3, oxo, oxido, ,

$-(CHR^n)_w$-heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl,

wherein R^n is hydrogen or C_{1-6} alkyl, w is 0 to 3, and

said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is C_{1-6} alkyl, C_{1-6} haloalkyl, furanyl, C_{1-6} alkoxy, C_{3-5} cycloalkyl, C_{1-6} alkylamino, -COO- C_{1-6} alkyl, -COOH, $-CH_2OCH_2OOH$, $-CH_2CH_2COOH$;

R^{10} is C_{1-6} alkyl, C_{1-6} dialkylamino, C_{3-8} cycloalkyl;

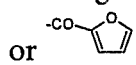
R^7 is hydrogen or Boc;

R^{15} is hydrogen or C_{1-6} alkyl;

R^4 is selected from the group consisting of:

C_{1-6} alkyl, $-CH_2CH_2CONH_2$, $-CH_2NCH_3SO_2CH_3$, $-CH(CH(CH_3)_2)NHCOCH_3$,

$-(CH_2)_eC_{3-8}$ cycloalkyl selected from cyclopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, wherein e is 0 to 2, and said cycloalkyl is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl,

C_{3-8} heterocycle, wherein said heterocyle is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxyl, -CO- C_{1-6} alkyl, oxo, benzyl, , or ,

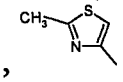
$-(CHR^f)_f$-phenyl, wherein R^f is hydrogen or –NHCOCH3, f is 0 or 1, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, NO_2, -CO-C_{1-6} alkyl, $-COOCH_3$, $COOC(CH_3)_3$,

$-NHSO_2CH_3$, NH_2, $-N(CH_3)_2$, $-N(CH_3)CH_2CH_2OH$, $-NHCOCH_3$,$-OCF_3$

$-OCH_2COOH$, pyridine-4-yl,  ;

$(CH_2)_g$-heteroaryl, wherein g is 0 or 1, sais heteroary selected from the group consisting of furanyl, thiophenyl, isoxazolyl, thiazolyl, oxazolyl, pyrroly, pyrazolyl, imidazolyl, indolyl, benzofuranyl, pyranyl, pyridinyl, pyrimidinyl, chinoxalinyl, chinolinyl, , , and ,

said heteroaryl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-6} alkyl, oxo, oxido, heteroalkyl, halogen, haloalkyl, methoxy-phenyl, haloalkyl, $-NH_2$, $-N(CH_3)_2$, -NHCH2-phenyl, -NHCOCH3, cyano, $-SO_2NH_2$, $-SO_2CH_3$, $-OCH_2COOCH_3$, $-OCH_2COOH$, $-SCH_2COOH$, $-COCH_3$, $-COOC(CH_3)_3$, $-COOC(CH_3)_3$, $-COOCH_2CH_3$,- COOH, pyridine-4-yl, , , benzyl, phenyl, said phenyl is optionally substituted with one to three substitutuents consisting of C_{1-6} alkyl , halogen, C_{1-6} alkoxyl, $-COOCH_3$;

-NH-phenyl, wherein said phenyl is optionally substituted with one to three substituents selected from C_{1-6} alkyl, and

pharmaceutically acceptable salts, hydrates, solvates and stereoisomers thereof.

4. A compound according to claim 3, wherein

X^1 is $C(=O)R^3$, wherein

R^3 is selected from the group consisting of:

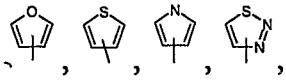
hydrogen, C_{1-6} alkyl, $-CH_2NH2$, $-CH_2CH_2NH_2$, $-C(CH_3)NH_2$, $-CH_2CH_2N(ethyl)_2$,, $-N(CH_3)_2$, $-N(CH_2CH_3)_2$, $-N(iso\text{-}propyl)_2$, $-NCH_3$-benzyl, $-NHCH_2COOCH_2CH_3$, -NH-phenyl, $-CH_2NHCOCH_3$, $CH_2CH_2CONH_2$, $-C(NH_2)CH_2C(CH_3)_2$, $-CH(NH_2)CH(CH_3)_2$,-CH2CN, $-C_{1-6}$ alkyl-OH, $-CH_2CH_2COOCH_3$, $-C(CH_3)_2OCOCH_3$, $-CH_2OCOCH_3$, -CH2SO2CH3, -CH2(CH3)SO2CH3, heteroalkyl,

$-(CH_2)_t$-C_{3-8} cycloalkyl selected from cyclopropanyl, cyclobutanyl, cyclopentanyl, cyclohexanyl, cycloheptanyl or , wherein t is 0 to 2, and

said -C_{3-8} cycloalkyl is optionally substituted with one to three substituents selected independently from the group comprising: C_{1-6} alkyl, halogen, hydroxyl, cyano, oxo, phenyl,

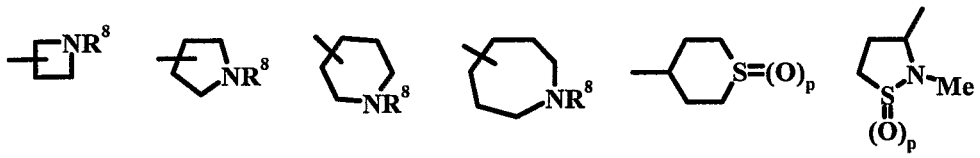
-$(CHR^m)_u$-phenyl, wherein R^m is H, hydroxyl or C_{1-6} alkyl, u is 0 to 2, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, -SO_2NH_2, -$NHCOCH_3$ or -$COOCH_3$,

-$(CH_2)_v$-heteroaryl selected from the group consisting of , wherein v 0 to 3, and

said heteroaryl is optionally substituted with one to three substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, halogen, phenyl, C_{1-6} alkoxy, C_{1-6} thioalkyl, -NH_2, SO2NH2, -COCH3, oxo, oxido, ,

-$(CHR^n)_w$-heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl,

wherein R^n is hydrogen or C_{1-6} alkyl, w is 0 to 3, and

said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is C_{1-6} alkyl, C_{1-6} haloalkyl, furanyl, C_{1-6} alkoxy, C_{3-5} cycloalkyl, C_{1-6} alkylamino, -COO- C_{1-6} alkyl, -COOH, $-CH_2OCH_2OOH$, $-CH_2CH_2COOH$;

R^{10} is C_{1-6} alkyl, C_{1-6} dialkylamino, C_{3-8} cycloalkyl;

R^7 is hydrogen or Boc;

R^{15} is hydrogen or C_{1-6} alkyl.

5. A compound according to claim 3, wherein

X^1 is $S(=O)_2R^3$, wherein $R^3$is phenyl optionally substituted with one to three halogen.

6. A compound according to claim 3, wherein

X^1 is $C(=O)OR^3$, wherein R^3 is C_{1-6} alkyl.

7. A compound according to claim 3, wherein

X^2 is $C(=O)R^4$, wherein

R^4 is selected from the group consisting of:

C_{1-6} alkyl, $-CH_2CH_2CONH_2$, $-CH_2NCH_3SO_2CH_3$, $-CH(CH(CH_3)_2)NHCOCH_3$,

$-(CH_2)_eC_{3-8}$ cycloalkyl, wherein e is 0 to 2, and said cycloalkyl is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl,

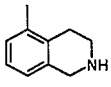
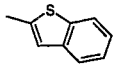
C_{3-8} heterocycle, wherein said heterocyle is optionally substituted with 1 to 4 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxyl, -CO- C_{1-6} alkyl, oxo, benzyl, , or ,

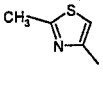
$-(CHR^f)_f$-phenyl, wherein R^f is hydrogen or –NHCOCH3, f is 0 or 1, and

said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, NO_2, -CO-C_{1-6} alkyl, $-COOCH_3$, $COOC(CH_3)_3$,

$-NHSO_2CH_3$, NH_2, $-N(CH_3)_2$, $-N(CH_3)CH_2CH_2OH$, $-NHCOCH_3$,$-OCF_3$

$-OCH_2COOH$, pyridine-4-yl, ;

$(CH_2)_g$-heteroaryl, wherein g is 0 or 1, sais heteroary selected from the group consisting of furanyl, thiophenyl, isoxazolyl, thiazolyl, oxazolyl, pyrroly, pyrazolyl, imidazolyl, indolyl, benzofuranyl, pyranyl, pyridinyl, pyrimidinyl, chinoxalinyl, chinolinyl, , , , , and ,

said heteroaryl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-6} alkyl, oxo, oxido, heteroalkyl, halogen, haloalkyl, methoxy-phenyl, haloalkyl, $-NH_2$, $-N(CH_3)_2$, -NHCH2-phenyl, -NHCOCH3, cyano, $-SO_2NH_2$, $-SO_2CH_3$, $-OCH_2COOCH_3$, $-OCH_2COOH$, $-SCH_2COOH$, $-COCH_3$, $-COOC(CH_3)_3$, $-COOC(CH_3)_3$, $-COOCH_2CH_3$,- COOH, pyridine-4-yl, , , benzyl, phenyl, said phenyl is optionally substituted with one to three substitutuents consisting of C_{1-6} alkyl , halogen, C_{1-6} alkoxyl, $-COOCH_3$;

-NH-phenyl, wherein said phenyl is optionally substituted with one to three substituents selected from C_{1-6} alkyl.

8. A compound according to claim 3, wherein

X^2 is $S(=O)_2R^4$, wherein

R^4 is phenyl optionally substituted with one to three C_{1-6} alkyl, or

heteroaryl selected from , , , ,and , and said heteroaryl is optionally substituted with one to four substitutuents consisting C_{1-6} alkyl, halogen, $-COOCH_3$ or phenyl.

9. A compound according to any one of claims 3 to 8, which compound is

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoxaline-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(3,4-dichloro-benzyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid [(S)-1-phenyl-3-(5-phenylacetyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-propyl]-amide; compound with trifluoro-acetic acid,
5-[(S)-3-(Cyclopentanecarbonyl-amino)-3-phenyl-propyl]-hexahydro-pyrrolo[3,4-c]pyrrole-2-carboxylic acid benzyl ester,
Cyclopentanecarboxylic acid {(R)-2-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-ethyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,5-dimethyl-2H-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,
Cyclopentanecarboxylic acid {(S)-3-[5-(1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,
Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(1H-[1,2,4]triazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,
Cyclopentanecarboxylic acid {(S)-3-[5-(4-methoxy-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dichloro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2-chloro-6-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dichloro-4-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(4-butoxy-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(4-ethoxy-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with hydrochloric acid,
Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2-bromo-6-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-difluoro-4-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(4-chloro-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,3-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2-methoxy-4-methylsulfanyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(1H-pyrrole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(3,5-dimethyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,
Cyclopentanecarboxylic acid [(S)-3-(5-acetyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,2-dimethyl-propionyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-benzenesulfonamide; compound with trifluoro-acetic acid,
Cyclopropanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Furan-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-isobutyramide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-methyl-butyramide; compound with trifluoro-acetic acid,
Thiophene-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid

N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2,2-dimethyl-propionamide; compound with trifluoro-acetic acid,
Cyclohexanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Morpholine-4-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-hydroxy-2-methyl-propionamide; compound with trifluoro-acetic acid,
Cyclobutanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Pyrrolidine-1-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-methoxy-acetamide; compound with trifluoro-acetic acid
N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-hydroxy-2-phenyl-acetamide; compound with trifluoro-acetic acid,
2-Cyclopentyl-N-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-methanesulfonyl-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid,
Furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
1H-Pyrrole-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
1-Methyl-1H-pyrrole-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
2-Acetylamino-N-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-sulfamoyl-benzamide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2-Oxo-thiazolidine-4-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Pyrazine-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-hydroxy-acetamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-methyl-butyramide; compound with trifluoro-acetic acid,

Tetrahydro-furan-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2-Dimethylamino-N-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,

1-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-phenyl-urea; compound with trifluoro-acetic acid,

1-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-isopropyl-urea; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid [(S)-3-(5-isobutyryl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(3-methyl-butyryl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid [(S)-3-(5-butyryl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid [(S)-3-(5-cyclopropanecarbonyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid [(S)-3-(5-cyclobutanecarbonyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2-methyl-butyryl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2-ethyl-butyryl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-pyridin-3-yl-propyl}-carbamic acid tert-butyl ester,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-butyl}-amide,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-pyridin-3-yl-propyl}-amide,
Cyclopentanecarboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-butyl}-amide,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,
Cyclopentanecarboxylic acid {(S)-3-[5-(1-acetyl-piperidine-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid [(S)-3-(5-benzyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trifluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(1-benzyl-5-oxo-pyrrolidine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
(1S,5R)-Bicyclo[3.1.0]hexane-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
4,4-Difluoro-cyclohexanecarboxylic acid [(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-amide; compound with trifluoro-acetic acid,
N-[(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-acetamide; compound with trifluoro-acetic acid,
(2S,3S)-1-Methyl-5-oxo-2-pyridin-3-yl-pyrrolidine-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
4-Dimethylamino-N-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-butyramide; compound with trifluoro-acetic acid,

1-Methyl-pyrrolidine-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-morpholin-4-yl-propionamide; compound with trifluoro-acetic acid,

2-(2-Aza-bicyclo[2.2.1]hept-2-yl)-N-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,

1-(Furan-2-carbonyl)-4-hydroxy-pyrrolidine-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-(5-morpholin-4-yl-tetrazol-2-yl)-acetamide; compound with trifluoro-acetic acid,

1-Methyl-piperidine-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

1-Methyl-azepane-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2-Methyl-1,2,3,4-tetrahydro-isoquinoline-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,,

3-(2-Aza-bicyclo[2.2.1]hept-2-yl)-N-{(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-propionamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-piperidin-1-yl-acetamide; compound with trifluoro-acetic acid,

1-Isopropyl-5-oxo-pyrrolidine-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

1-Ethyl-piperidine-4-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

1-Isobutyl-5-oxo-pyrrolidine-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-pyrrolidin-1-yl-acetamide; compound with trifluoro-acetic acid,

1-Methyl-5-sulfamoyl-1H-pyrrole-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

(S)-1-Acetyl-pyrrolidine-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2-methyl-1,2,3,4-tetrahydro-isoquinoline-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(1-pyrimidin-2-yl-piperidine-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(1-isobutyl-5-oxo-pyrrolidine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(1-methyl-5-sulfamoyl-1H-pyrrole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid ((S)-3-{5-[1-(furan-2-carbonyl)-4-hydroxy-pyrrolidine-2-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(4-benzyl-morpholine-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(6-chloro-imidazo[2,1-b]thiazole-5-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(5-methyl-1-phenyl-1H-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(5-methyl-2-phenyl-2H-[1,2,3]triazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid ((S)-3-{5-[3-(2-chloro-phenyl)-5-methyl-isoxazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(4,5,6,7-tetrahydro-2H-indazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(1,4,5,6-tetrahydro-cyclopentapyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(5-ethyl-2-methyl-2H-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

[(S)-3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-carbamic acid tert-butyl ester,

Cyclopentanecarboxylic acid {(S)-3-[5-(4-fluoro-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

2-Chloro-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-fluoro-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-trifluoromethyl-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-methoxy-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-methoxy-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3,3-dimethyl-butyramide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-succinamic acid methyl ester; compound with trifluoro-acetic acid,

2-Chloro-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-cyano-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2,4-difluoro-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2,6-difluoro-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3,4-difluoro-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-fluoro-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-fluoro-benzamide; compound with trifluoro-acetic acid,

Furan-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-isobutyramide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-methyl-butyramide; compound with trifluoro-acetic acid,
3-Chloro-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-fluoro-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-thiophen-2-yl-acetamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-trifluoromethyl-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-cyano-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-ethoxy-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3,5-difluoro-benzamide; compound with trifluoro-acetic acid,
Thiophene-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
4-Chloro-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-methoxy-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-fluoro-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3,4-dimethoxy-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2,2-dimethyl-propionamide; compound with trifluoro-acetic acid,
Cyclohexanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Morpholine-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-oxalamic acid ethyl ester; compound with trifluoro-acetic acid,

Acetic acid 1-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propylcarbamoyl}-1-methyl-ethyl ester; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-phenyl-acetamide; compound with trifluoro-acetic acid,

(1R,2R)-2-Phenyl-cyclopropanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclobutanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Isoxazole-5-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

5-Chloro-4-methoxy-thiophene-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

4-Methyl-[1,2,3]thiadiazole-5-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

3-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-1,1-diethyl-urea; compound with trifluoro-acetic acid,

3-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-1,1-diisopropyl-urea; compound with trifluoro-acetic acid,

3-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-1,1-dimethyl-urea; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-methoxy-acetamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-cyclopentyl-propionamide; compound with trifluoro-acetic acid,

Acetic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propylcarbamoyl}-methyl ester; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-methoxy-phenyl)-acetamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-cyclopentyl-acetamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-dimethylamino-benzamide; compound with trifluoro-acetic acid,
3,5-Dimethyl-isoxazole-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
3-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-1-methyl-1-phenyl-urea; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-methylsulfanyl-acetamide; compound with trifluoro-acetic acid,
3-Chloro-thiophene-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-carbamic acid methyl ester; compound with trifluoro-acetic acid,
1-Benzyl-3-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-1-methyl-urea; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dichloro-4-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
3-Acetylamino-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid,
4-Acetylamino-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-benzamide; compound with trifluoro-acetic acid,
1-Acetyl-piperidine-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
3-Amino-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-propionamide; compound with trifluoro-acetic acid,
2-Amino-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,
(S)-Pyrrolidine-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-cyano-acetamide; compound with trifluoro-acetic acid,
Cycloheptanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-cyclohexyl-acetamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-cyclopropyl-acetamide; compound with trifluoro-acetic acid,
Furan-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-isonicotinamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-terephthalamic acid methyl ester; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-piperidin-1-yl-propionamide; compound with trifluoro-acetic acid,
1H-Pyrrole-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-succinamide; compound with trifluoro-acetic acid,
1H-Indole-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
1-Methyl-1H-pyrrole-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
2-Acetylamino-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,
Pyridine-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-methylamino-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-methylamino-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-dimethylamino-benzamide; compound with trifluoro-acetic acid,

5-Amino-1-phenyl-1H-pyrazole-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2,2,3,3-Tetramethyl-cyclopropanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2-Acetylamino-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-methylsulfanyl-butyramide; compound with trifluoro-acetic acid,

2-Ethyl-5-methyl-2H-pyrazole-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

5-Amino-2H-[1,2,4]triazole-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

5-Chloro-thiophene-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

(S)-2-Acetylamino-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-methylsulfanyl-butyramide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-diethylamino-propionamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-methyl-nicotinamide; compound with trifluoro-acetic acid,

3-Methyl-furan-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-nicotinamide; compound with trifluoro-acetic acid,

Pyrazine-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-(1-methyl-1H-imidazol-4-yl)-acetamide; compound with trifluoro-acetic acid,

3H-Imidazole-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-methanesulfonyl-acetamide; compound with trifluoro-acetic acid,

2-Methyl-thiazole-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-imidazol-1-yl-acetamide; compound with trifluoro-acetic acid,

1-Methyl-1H-pyrazole-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

1-Methyl-1H-imidazole-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dichloro-4-methylsulfanyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

5-{(S)-3-Phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carboxylic acid phenylamide; compound with trifluoro-acetic acid,

5-{(S)-3-Phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carboxylic acid (2,6-dimethyl-phenyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-benzenesulfonyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(thiophene-2-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3,5-dimethyl-isoxazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-chloro-1,3-dimethyl-1H-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2,5-Dimethyl-4-(5-{(S)-3-phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-sulfonyl)-furan-3-carboxylic acid methyl ester; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(1,3,5-trimethyl-1H-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Methyl-5-(5-{(S)-3-phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-sulfonyl)-1H-pyrrole-2-carboxylic acid methyl ester; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-cyano-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(1H-pyrrole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methoxy-1H-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(pyridine-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(pyrazine-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(tetrahydro-furan-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-acetylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(isoquinoline-7-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(3,5-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(1H-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(benzofuran-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(benzo[b]thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(1-methyl-1H-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-chloro-6-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methoxy-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-methylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[4-(3-methyl-5-oxo-4,5-dihydro-pyrazol-1-yl)-benzoyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(tetrahydro-furan-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2-thiophen-2-yl-acetyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-chloro-thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-5-phenyl-furan-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-fluoro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[3-(2-oxo-pyrrolidin-1-yl)-benzoyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(benzo[b]thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

4-(5-{(S)-3-Phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-benzoic acid tert-butyl ester; compound with trifluoro-acetic acid,

3-(5-{(S)-3-Phenyl-3-[(tetrahydro-furan-3-carbonyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-benzoic acid tert-butyl ester; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3,5-dimethyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(1H-indole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(benzofuran-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,3-dihydro-benzofuran-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3,3-dimethyl-butyryl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-cyclobutanecarbonyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-cyclohexyl-acetyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-cyclopentanecarbonyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-cyclopentyl-acetyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dichloro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-isobutyryl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-methyl-butyryl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid [(S)-1-phenyl-3-(5-propionyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-propyl]-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-cyano-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-chloro-2-ethyl-2H-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-chloro-2-methyl-2H-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,5-trimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-fluoro-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-benzoyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,

1-Methyl-cyclohexanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-morpholin-4-yl-propionamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-(4-methyl-piperazin-1-yl)-propionamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-(1H-[1,2,4]triazol-3-yl)-propionamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-(1H-tetrazol-5-yl)-acetamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-furan-2-yl-propionamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-thiophen-2-yl-propionamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-pyridin-3-yl-propionamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-(3H-imidazol-4-yl)-propionamide; compound with trifluoro-acetic acid,

Tetrahydro-furan-2-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-(methanesulfonyl-methyl-amino)-acetamide; compound with trifluoro-acetic acid,

4-Methoxy-thiophene-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3,3,3-trifluoro-propionamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-dimethylamino-acetamide; compound with trifluoro-acetic acid,

2-Methyl-cyclopropanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

1-Cyano-cyclopropanecarboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

5-Acetyl-2,4-dimethyl-1H-pyrrole-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2,4-Dimethyl-6-oxo-6H-pyran-3-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

4,6-Dimethyl-pyrimidine-5-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2,4-dimethyl-nicotinamide; compound with trifluoro-acetic acid,

(S)-2-Amino-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-propionamide; compound with trifluoro-acetic acid,

(S)-2-Amino-4-methyl-pentanoic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid ,

(S)-2-Amino-N-{(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-methyl-butyramide; compound with trifluoro-acetic acid ,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-butyramide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2,3-dimethyl-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2,6-dimethyl-benzamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-fluoro-phenyl)-acetamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-phenyl-propionamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-methanesulfonyl-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-methyl-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-methyl-benzamide; compound with trifluoro-acetic acid,
N-{(S)-3-[5-(2-Chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-4-methyl-benzamide; compound with trifluoro-acetic acid,
Cyclopentanecarboxylic acid {(S)-3-[5-(3,4-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-amino-2,6-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-6-oxo-6H-pyran-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid
Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(1-acetyl-piperidine-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-cyclohexanecarbonyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,
Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(furan-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(furan-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,
Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-ethyl-5-methyl-2H-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-methyl-furan-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-fluoro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-chloro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-thiazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(1-methyl-1H-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-methoxy-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-acetylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-acetylamino-2-phenyl-acetyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-((S)-pyrrolidine-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-chloro-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(isoquinoline-1-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-6-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-8-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-carbamoyl-propionyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid [(S)-3-(5-cyclopropanecarbonyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-1-phenyl-propyl]-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-butyryl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[2-(methanesulfonyl-methyl-amino)-acetyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-difluoro-4-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trifluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-4-methoxy-1-phenyl-butyl}-amide,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-((R)-2-acetylamino-3-methyl-butyryl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(1-methyl-5-oxo-pyrrolidine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid;,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-2-methylsulfanyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-4-hydroxy-1-phenyl-butyl}-amide,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,3,6-trichloro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methyl-2-phenyl-2H-[1,2,3]triazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2-Oxo-imidazolidine-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid,

2-Cyclohexyl-N-{(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamidetrifluoro-acetic acid,

N-{(S)-3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-fluoro-phenyl)-acetamidetrifluoro-acetic acid,

N-{(S)-3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-methoxy-phenyl)-acetamidetrifluoro-acetic acid,

1-{(S)-3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-isopropyl-ureatrifluoro-acetic acid,

2-Oxo-thiazolidine-4-carboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid,

Furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid,

N-{(S)-3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3,4-difluoro-benzamidetrifluoro-acetic acid,

1-{(S)-3-[5-(2,4-Dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-phenyl-ureatrifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid,

Tetrahydro-pyran-4-carboxylic acid {(S)-3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2-Cyclohexyl-N-{(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-fluoro-phenyl)-acetamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-2-(4-methoxy-phenyl)-acetamide; compound with trifluoro-acetic acid,

Cyclohexanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3,4-difluoro-benzamide; compound with trifluoro-acetic acid,

1-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3-phenyl-urea; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-pyran-4-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-pyran-4-carboxylic acid {(S)-3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

(R)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

(R)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

(R)-Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,5-trimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

(S)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

(S)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

(S)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

(S)-Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2,4,5-trimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methyl-3-phenyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-6-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(quinoline-8-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

(R)-Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-4-pyridin-4-yl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(4-cyano-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopropanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid,

5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-pyridin-2-yl-propyl}-amide; compound with trifluoro-acetic acid,

1,1-Dioxo-hexahydro-1λ6-thiopyran-4-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[5-amino-1-(4-methoxy-phenyl)-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(1-phenyl-5-trifluoromethyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[1-(4-methoxy-phenyl)-5-trifluoromethyl-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[1-(2-methoxy-phenyl)-5-trifluoromethyl-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[1-(4-chloro-phenyl)-5-trifluoromethyl-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(1-p-tolyl-5-trifluoromethyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[1-(4-fluoro-phenyl)-5-trifluoromethyl-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methyl-1-phenyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-methyl-1-p-tolyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[2-(4-methoxy-phenyl)-5-methyl-2H-pyrazole-3-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-2-phenyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[4,6-dimethyl-2-(2-methyl-thiazol-4-yl)-pyrimidine-5-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-2-pyridin-4-yl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-bromo-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-fluoro-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-chloro-pyridine-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methoxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-1-phenyl-3-[5-(2-trifluoromethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(5-amino-1-phenyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[5-amino-1-(4-fluoro-phenyl)-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[5-amino-1-(2-methoxy-phenyl)-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid ((S)-3-{5-[1-(4-chloro-phenyl)-5-methyl-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-1-phenyl-propyl)-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-bromo-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-butyl}-amide,

4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

Pentanoic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclobutanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2-Cyclopentyl-N-{(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-methyl-5-trifluoromethyl-oxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2-amino-6-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-4-nitro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amidetrifluoro-acetic acid;,

N-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-propionamide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-butyramide; compound with trifluoro-acetic acid,

Cyclopropanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

{(S)-3-[5-(2,4-Dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-carbamic acid tert-butyl ester,

Cyclopentanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-3a,6a-dimethyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

5-Oxo-1-(2,2,2-trifluoro-ethyl)-pyrrolidine-3-carboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-butyl}-amide,

4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-butyl}-amide,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(6-chloro-2-dimethylamino-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-cyano-2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-iodo-2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(2,5-di-tert-butyl-4-cyano-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

4-Oxo-cyclohexanecarboxylic acid {(S)-3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

Cyclopentanecarboxylic acid {(S)-3-[5-(2-ethoxy-4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

Cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopropanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-methyl-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(4-bromo-2,5-di-tert-butyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3,5-diethyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2-methanesulfonyl-4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

(5-{5-[(S)-3-(Cyclopentanecarbonyl-amino)-3-phenyl-propyl]-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl}-4,6-dimethyl-pyrimidin-2-yloxy)-acetic acid methyl ester,

Cyclopentanecarboxylic acid {(S)-3-[5-(2-benzylamino-4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

(5-{5-[(S)-3-(Cyclopentanecarbonyl-amino)-3-phenyl-propyl]-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl}-4,6-dimethyl-pyrimidin-2-yloxy)-acetic acid; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-6-oxo-6H-pyran-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

Cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(2,4,6-trimethyl-benzenesulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

Cyclopentanecarboxylic acid {(R)-3-[5-(2-dimethylamino-4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(4,6-Dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3,4-difluoro-benzamide; compound with trifluoro-acetic acid,

Cyclobutanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2-Cyclopentyl-N-{(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,

5-Methyl-thiophene-2-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2-Oxo-thiazolidine-4-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

2-Cyclohexyl-N-{(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-acetamide; compound with trifluoro-acetic acid,

3,3-Difluoro-cyclobutanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

3-Oxo-cyclobutanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

N-{(S)-3-[5-(4,6-Dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-3,4-difluoro-benzenesulfonamide; compound with trifluoro-acetic acid,

Cyclopropanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Thiophene-2-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(3-methyl-benzofuran-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-furan-3-carboxylic acid {(S)-3-[5-(7-methoxy-3-methyl-1H-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Tetrahydro-pyran-3-carboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid [(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-amide,

4,4-Difluoro-cyclohexanecarboxylic acid [(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-(3-fluoro-phenyl)-propyl]-amide,

4-Hydroxy-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide,

4-Hydroxy-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide, or

3,3-Difluoro-cyclobutanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-1-phenyl-propyl}-amide.

10. A compound according to claim 2, wherein:

R^1 is phenyl optionally substituted with one to four substituents selected independently in each incidence from the group consisting of halogen and C_{1-6} alkyl, or

heteroaryl selected from the group consisting of pyridinyl, pyrimidinyl and thiazolyl said heteroaryl optionally substituted with one to three substituents independently selected from the group consisting of halogen, C_{1-6} alkyl and C_{1-6} haloalkyl;

R^2 is hydrogen;

R^5 is hydrogen or C_{1-6} alkyl;

R^{5a} is hydrogen, hydroxyl or C_{1-6} alkyl;

R^6 is hydrogen or C_{1-3} hydroxyalkyl;

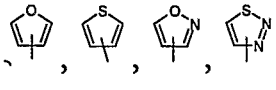
X^1 is selected from the group consisting $-C(=O)R^3$, $-S(=O)_2R^3$;

X^2 is selected from the group consisting $-C(=O)R^4$, $-S(=O)_2R^4$ or benzyl;

R^3 is selected from the group consisting of:

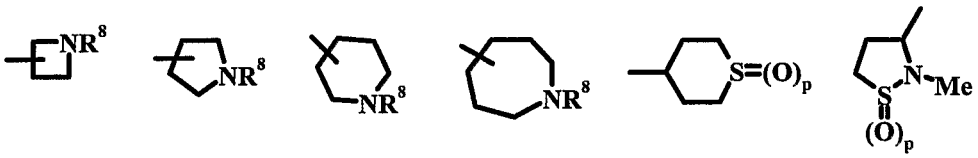
C_{1-6} alkyl, heteroalkyl, -phenyl-SO_2CH_3,

$-(CH_2)_t-C_{3-8}$ cycloalkyl selected from the group consisting of cycopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, wherein t is 0 or 2, and said $-C_{3-8}$ cycloalkyl is optionally substituted with one to three substituents selected independently from the group consisting of halogen, hydroxyl and oxo,

heteroaryl selected from the group consisting of , 

or , wherein v 0 to 3, and said heteroaryl is optionally substituted with one to three C_{1-10} alkyl,

heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl, and

said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is C_{1-6} alkyl, C_{1-6} haloalkyl, furanyl, C_{1-6} alkoxy, C_{3-5} cycloalkyl, C_{1-6} alkylamino, -COO- C_{1-6} alkyl, -COOH, $-CH_2OCH_2OOH$, $-CH_2CH_2COOH$;

R^{10} is C_{1-6} alkyl, C_{1-6} dialkylamino, C_{3-8} cycloalkyl;

R^7 is hydrogen or Boc;

R^{15} is hydro[illegible] or C_{1-6} alkyl;

$-NH-(CH_2)_x$-phenyl, wherein x is 0 or 1, and

said phenyl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, $-OCF_3$, heteroalkyl, haloalkyl, acetyl, dialkylamin, cyano, -COOH, $COOCH_2CH_3$,

$-NH(CH_2)_y$-heteroaryl, wherein y is 0 or 1, and

said heteroaryl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, haloalkyl,

-NH-heterocycle, wherein said heteroaryl is optionally substituted with 1 to 2 substituents independently selected in each occurrence from the group consisting of $-COCF_3$, $COOCH_2$-phenyl,

$-NH(CH_2)_z-C_{3-8}$ cycloalkyl selected from cycopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, wherein z is 0 or 1;

R^4 is selected from the group consisting of:

C_{1-6} alkyl,

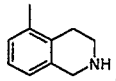
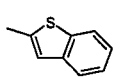
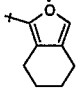
C_{3-8} cycloalkyl selected from selected from cyclopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, and said cycloalkyl is optionally substituted with 1 to 4 substituents consisting of C_{1-6} alkyl,

C_{3-8} heterocycle selected from tetrahydrofuranyl and ,wherein said heterocyle is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl and -CO- C_{1-6} alkyl,
Phenyl, and said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, NO_2, -CO-C_{1-6}

alkyl,

$-COOCH_3$, $COOC(CH_3)_3$,$-NHSO_2CH_3$, NH_2, $-N(CH_3)_2$, $-N(CH_3)CH_2CH_2OH$,

$-NHCOCH_3$,$-OCF_3$ $-OCH_2COOH$, pyridine-4-yl, , , ,

heteroaryl selected from the group consisting of furanyl, thiophenyl, isoxazolyl, thiazolyl, oxazolyl, pyrroly, pyrazolyl, imidazolyl, indolyl, benzofuranyl, pyranyl, pyridinyl, pyrimidinyl, chinoxalinyl, chinolinyl, , , , , and ,

said heteroaryl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-6} alkyl, oxo, oxido, heteroalkyl, halogen, haloalkyl, methoxy-phenyl, haloalkyl, $-NH_2$, $-N(CH_3)_2$, -NHCH2-phenyl, -NHCOCH3, cyano, $-SO_2NH_2$, $-SO_2CH_3$, $-OCH_2COOCH_3$, $-OCH_2COOH$, $-SCH_2COOH$, $-COCH_3$, $-COOC(CH_3)_3$, $-COOC(CH_3)_3$, $-COOCH_2CH_3$,- COOH, pyridine-4-yl, , benzyl, phenyl, said phenyl is optionally substituted with one to three substituents consisting of C_{1-6} alkyl , halogen, C_{1-6} alkoxyl, $-COOCH_3$;

$-(NR^h)-(CH2)_h$-phenyl, wherein R^h is hydrogen or C_{1-6} alkyl, h is 0 or 1, and said phenyl is optionally substituted with one to three substituents selected from C_{1-6} alkyl, and

pharmaceutically acceptable salts, hydrates, solvates and stereoisomers thereof.

11. A compound according to claim 10, wherein

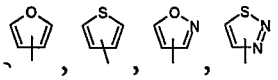
X^1 is $C(=O)R^3$, wherein

R^3 is selected from the group consisting of:

is selected from the group consisting of:

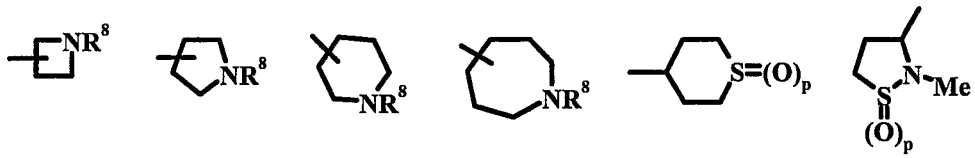
C_{1-6} alkyl, heteroalkyl, $-phenyl-SO_2CH_3$,

$-(CH_2)_t-C_{3-8}$ cycloalkyl selected from the group consisting of cycopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, wherein t is 0 or 2, and said $-C_{3-8}$ cycloalkyl is optionally substituted with one to three substituents selected independently from the group consisting of halogen, hydroxyl and oxo,

heteroaryl selected from the group consisting of 

or , wherein v 0 to 3, and said heteroaryl is optionally substituted with one to three C_{1-10} alkyl,

heterocycle selected from the group consisting of IIa-f, oxetanyl,



IIa IIb IIc IId IIe IIf

tetrahydrofuranyl, tetrahydropyranyl, thiazolidinyl, oxazolidinyl, isothiazolidinyl, piperazinyl, imidazolinyl, 1,2,3,4-tetrahydroquinolinyl and N-methyl-1,2,3,4-tetrahydroisoquinolin-3-yl, and

said heterocycle optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, hydroxy, C_{1-6} alkylcarbonyl, carbamoyl, amino, C_{1-6} alkylamino, C_{1-6} dialkylamino, pyridinyl and phenylamino, and two hydrogens on a carbon bonded to a nitrogen can be replaced by oxygen (oxo), wherein:

R^8 is hydrogen, C_{1-6}-alkyl, C_{1-6} haloalkyl, C_{1-6} heteroalkyl, phenyl-C_{1-6} alkyl, C_{1-3} alkoxy-C_{1-6} alkyl, pyrimidin-2-yl, COR^9, $COCHR^{15}NHR^7$ or SO_2R^{10};

R^9 is C_{1-6} alkyl, C_{1-6} haloalkyl, furanyl, C_{1-6} alkoxy, C_{3-5} cycloalkyl, C_{1-6} alkylamino, -COO- C_{1-6} alkyl, -COOH, $-CH_2OCH_2OOH$, $-CH_2CH_2COOH$;

R^{10} is C_{1-6} alkyl, C_{1-6} dialkylamino, C_{3-8} cycloalkyl;

R^7 is hydrogen or Boc;

R^{15} is hydrogen or C_{1-6} alkyl;

$-NH-(CH_2)_x$-phenyl, wherein x is 0 or 1, and

said phenyl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, $-OCF_3$, heteroalkyl, haloalkyl, acetyl, dialkylamin, cyano, -COOH, $COOCH_2CH_3$,

$-NH(CH_2)_y$-heteroaryl, wherein y is 0 or 1, and

said heteroaryl is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl, halogen, haloalkyl,

-NH-heterocycle, wherein said heteroaryl is optionally substituted with 1 to 2 substituents independently selected in each occurrence from the group consisting of $-COCF_3$, $COOCH_2$-phenyl,

$-NH(CH_2)_z-C_{3-8}$ cycloalkyl selected from cycopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, wherein z is 0 or 1.

12. A compound according to claim 10, wherein

X^1 is $-S(=O)_2R^3$, wherein

R^3 phenyl.

13. A compound according to claim 10, wherein

X^2 is $-C(=O)R^4$,

R^4 is selected from the group consisting of:

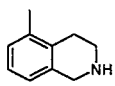
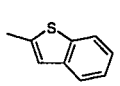
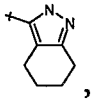
C_{1-6} alkyl,

C_{3-8} cycloalkyl selected from selected from cyclopropanyl, cyclobutanyl, cyclopentanyl and cyclohexanyl, and said cycloalkyl is optionally substituted with 1 to 4 substituents consisting of C_{1-6} alkyl,

C_{3-8} heterocycle selected from tetrahydrofuranyl and ,wherein said heterocyle is optionally substituted with 1 to 3 substituents independently selected in each occurrence from the group consisting of C_{1-6} alkyl and -CO- C_{1-6} alkyl,
Phenyl, and said phenyl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-10} alkyl, C_{1-10} heteroalkyl, C_{1-6} haloalkyl, cyano, halogen, C_{1-6} alkylsulfonyl, NO_2, -CO-C_{1-6} alkyl,
$-COOCH_3$, $COOC(CH_3)_3$,$-NHSO_2CH_3$, NH_2, $-N(CH_3)_2$, $-N(CH_3)CH_2CH_2OH$,

$-NHCOCH_3$,$-OCF_3$ $-OCH_2COOH$, pyridine-4-yl, ,

heteroaryl selected from the group consisting of furanyl, thiophenyl, isoxazolyl, thiazolyl, oxazolyl, pyrroly, pyrazolyl, imidazolyl, indolyl, benzofuranyl, pyranyl,

pyridinyl, pyrimidinyl, chinoxalinyl, chinolinyl, , , , , and ,

said heteroaryl is optionally substituted with one to four substituents selected independently in each incidence from the group consisting of C_{1-6} alkyl, oxo, oxido, heteroalkyl, halogen, haloalkyl, methoxy-phenyl, haloalkyl, $-NH_2$, $-N(CH_3)_2$, -NHCH2-phenyl, -NHCOCH3, cyano, $-SO_2NH_2$, $-SO_2CH_3$, $-OCH_2COOCH_3$, $-OCH_2COOH$, $-SCH_2COOH$, $-COCH_3$, $-COOC(CH_3)_3$, $-COOC(CH_3)_3$, $-COOCH_2CH_3$,- COOH, pyridine-4-yl, , benzyl, phenyl, said phenyl is optionally substituted with one to three substituents consisting of C_{1-6} alkyl , halogen, C_{1-6} alkoxyl, $-COOCH_3$;

$-(NR^h)-(CH2)_h$-phenyl, wherein R^h is hydrogen or C_{1-6} alkyl, h is 0 or 1, and said phenyl is optionally substituted with one to three substituents selected from C_{1-6} alkyl.

14. A compound according to claim 10, wherein

X^2 is $-S(=O)_2R^4$,

R^4 is phenyl optionally substituted with one to four substituents consiting of C_{1-6} alkyl, halogen, cyano, $-COOCH_3$ and COOH, or

heteroaryl selected from isoxazolyl, pyrazolyl, furanyl, pyrrolyl and pyridinyl, and said heteroaryl is optionally substituted with one to four substituents consisting of C_{1-6} alkyl, halogen, phenyl, $-N(CH_3)_2$, -COOH, $-COOCH_3$ and $-COOCH_2CH_3$.

15. A compound according to any one of claim 10 to 14, which is

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,
N-{3-[5-(2,6-Dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-N-(4-fluoro-phenyl)-benzenesulfonamide; compound with trifluoro-acetic acid,
1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,
1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid [3-(5-benzyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-propyl]-(3-chloro-4-methyl-phenyl)-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,5-trimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-difluoro-4-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-6-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-6-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dichloro-4-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,6-trimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-chloro-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-acetyl-2,4-dimethyl-1H-pyrrole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with hydrochloric acid,

Furan-2-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

N-(3-Chloro-4-methyl-phenyl)-N-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-isobutyramide; compound with trifluoro-acetic acid,

Thiophene-2-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Morpholine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Isoxazole-5-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4-Methyl-[1,2,3]thiadiazole-5-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Pyrrolidine-1-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3,3-dimethyl-urea; compound with trifluoro-acetic acid,

N-(3-Chloro-4-methyl-phenyl)-N-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-acetamide; compound with trifluoro-acetic acid,

N-(3-Chloro-4-methyl-phenyl)-2-cyclopentyl-N-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-acetamide; compound with trifluoro-acetic acid,

3,5-Dimethyl-isoxazole-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

N-(3-Chloro-4-methyl-phenyl)-N-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-4-methanesulfonyl-benzamide; compound with trifluoro-acetic acid,

Piperidine-1-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Methyl-1H-pyrrole-2-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-(4-chloro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-butyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-p-tolyl-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-{5-[5-chloro-2-(2,5-dimethyl-pyrrol-1-yl)-benzoyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-propyl)-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-quinoline-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-chloro-4-methyl-6-pyrrolidin-1-yl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[2-(1,1-dioxo-1λ6-thiomorpholin-4-yl)-benzoyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-bromo-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-2-pyridin-4-yl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carboxylic acid (2-isopropyl-phenyl)-amide; compound with trifluoro-acetic acid,

5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carboxylic acid (2,6-dimethyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,6-trimethyl-benzenesulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(thiophene-2-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-isoxazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-chloro-1,3-dimethyl-1H-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-sulfonyl)-2,5-dimethyl-furan-3-carboxylic acid methyl ester; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1,3,5-trimethyl-1H-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(1-acetyl-piperidine-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-cyano-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(furan-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(furan-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(pyridine-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

3-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-benzoic acid methyl ester; compound with trifluoro-acetic acid,

4-((3-Chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-piperidine-1-carboxylic acid tert-butyl ester,

Piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-phenyl-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-benzoic acid methyl ester; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1H-pyrrole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-methyl-1H-pyrrole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(pyridine-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methoxy-2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-ethyl-5-methyl-2H-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-methyl-furan-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-fluoro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3H-imidazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-thiazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-methyl-1H-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(tetrahydro-furan-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methoxy-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(isoquinoline-1-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-cyclohexanecarbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(quinoline-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1H-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(benzofuran-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(benzo[b]thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-methyl-1H-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methoxy-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1H-indole-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-chloro-thiophene-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-5-phenyl-furan-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-fluoro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(1-benzyl-3,5-dimethyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,6-trimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-chloro-2,6-dimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-1-phenyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-difluoro-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-fluoro-6-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(6-chloro-2-fluoro-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-6-fluoro-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,6-trifluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-chloro-2-fluoro-6-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Methanesulfonyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Cyclopropanecarbonyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Piperidine-1,4-dicarboxylic acid 4-((3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide) 1-dimethylamide; compound with trifluoro-acetic acid,

Piperidine-1,4-dicarboxylic acid 4-((3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide) 1-ethylamide; compound with trifluoro-acetic acid,

1-Dimethylsulfamoyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(2,2,2-Trifluoro-acetyl)-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-pyridin-2-yl-amide,

1-Acetyl-piperidine-4-carboxylic acid (3,4-dichloro-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-amino-1-phenyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-{5-[5-amino-1-(4-methoxy-phenyl)-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-propyl)-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-phenyl-5-trifluoromethyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[1-(4-methoxy-phenyl)-5-trifluoromethyl-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[1-(2-methoxy-phenyl)-5-trifluoromethyl-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[1-(4-chloro-phenyl)-5-trifluoromethyl-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-p-tolyl-5-trifluoromethyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[1-(4-fluoro-phenyl)-5-trifluoromethyl-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-amino-1-p-tolyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-{5-[5-amino-1-(4-fluoro-phenyl)-1H-pyrazole-4-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-propyl)-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-1-phenyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-1-p-tolyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-2-p-tolyl-2H-pyrazole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-(3-{5-[2-(4-methoxy-phenyl)-5-methyl-2H-pyrazole-3-carbonyl]-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl}-propyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-phenyl-amide,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-acetylamino-4-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-fluoro-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-chloro-pyridine-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methoxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methanesulfonylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-(2,6-dichloro-pyridin-4-yl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-(3,5-dimethyl-isoxazol-4-yl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-cyclopentyl-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-thiophen-3-yl-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-furan-2-ylmethyl-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-[1-(2,2,2-trifluoro-acetyl)-piperidin-3-yl]-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-[1-(2,2,2-trifluoro-acetyl)-piperidin-4-yl]-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-cyclohexylmethyl-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-(5-methyl-2-trifluoromethyl-furan-3-yl)-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-phenyl-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-(3-chloro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-(4-fluoro-phenyl)-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-p-tolyl-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-(4-trifluoromethoxy-phenyl)-urea; compound with trifluoro-acetic acid,

(3-(3-Chloro-4-methyl-phenyl)-3-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-ureido)-acetic acid ethyl ester; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-(4-fluoro-benzyl)-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-(4-methoxy-benzyl)-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-(4-trifluoromethyl-phenyl)-urea; compound with trifluoro-acetic acid,

3-(4-Acetyl-phenyl)-1-(3-chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-(3,4-difluoro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-6-oxo-6H-pyran-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-(4-dimethylamino-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

4-(3-(3-Chloro-4-methyl-phenyl)-3-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-ureido)-piperidine-1-carboxylic acid benzyl ester; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-(4-methoxy-phenyl)-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-cyclohexyl-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-(4-methylsulfanyl-phenyl)-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-(2,4-dichloro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-(4-chloro-2-trifluoromethyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-(4-methyl-benzyl)-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-(3,4-dichloro-benzyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-3-(4-cyano-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-urea; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3,4-dimethyl-phenyl)-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-3a,6a-dimethyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-6-oxo-6H-pyran-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(5-acetyl-2,4-dimethyl-1H-pyrrole-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,5-trimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-1-phenyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dichloro-4-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-2-pyridin-4-yl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-2-phenyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-3-phenyl-isoxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclobutanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-[1-(2,2,2-trifluoro-acetyl)-piperidin-3-yl]-urea; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-hydroxy-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-bis-hydroxymethyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-p-tolyl-amide,

4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-phenyl-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-5-trifluoromethyl-oxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-phenyl-5-trifluoromethyl-oxazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-amino-6-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-4-nitro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amidetrifluoro-acetic acid;,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1,3,5-trimethyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,5-dimethyl-1H-pyrazole-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

N-(3-Chloro-4-methyl-phenyl)-N-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-propionamide; compound with trifluoro-acetic acid,

N-(3-Chloro-4-methyl-phenyl)-N-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-butyramide; compound with trifluoro-acetic acid,

Pentanoic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopropanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

3-(4-Chloro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-1-p-tolyl-urea,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-chloro-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid [3-(5-benzoyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-propyl]-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,4-bis-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3-dihydro-benzofuran-7-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3-dimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-chloro-5-methylsulfanyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-fluoro-3-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-chloro-2-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-ethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-fluoro-5-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-4,5-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4,5-trifluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dichloro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-dichloro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3-dichloro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-difluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-4-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-chloro-2-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-trifluoromethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Acetic acid 2-(5-{3-[(1-acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-phenyl ester; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-bromo-2-chloro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-acetyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-dimethoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

2-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-benzoic acid methyl ester; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,5-bis-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,3,4-trifluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-fluoro-3-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dichloro-5-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-fluoro-2-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-fluoro-5-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-acetyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-bromo-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-5-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-bromo-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3,5-bis-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-fluoro-4-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-bromo-5-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-3-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-dimethylamino-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-iodo-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-ethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-[3-(5-{4-[(2-hydroxy-ethyl)-methyl-amino]-benzoyl}-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl)-propyl]-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(6-fluoro-4H-benzo[1,3]dioxine-8-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-dibromo-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methoxy-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-fluoro-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-methoxy-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-fluoro-2-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(cinnoline-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethoxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-trifluoromethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-methyl-pyridine-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-pyrimidin-2-yl-amide; compound with trifluoro-acetic acid,

5-Oxo-1-(2,2,2-trifluoro-ethyl)-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(6-chloro-2-dimethylamino-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-cyano-2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-iodo-2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-dimethyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3,6-dimethoxy-pyridazine-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-bromo-2,5-di-tert-butyl-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,5-di-tert-butyl-4-cyano-thiophene-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

3-(4-Chloro-phenyl)-1-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-1-phenyl-urea; compound with trifluoro-acetic acid,

1,1-Dioxo-hexahydro-1λ6-thiopyran-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-hydroxy-propyl}-amide,

Cyclopentanecarboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-cyclohexyl}-phenyl-amide,

Cyclopentanecarboxylic acid {3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-phenyl-amide; compound with trifluoroacetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-chloro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-chloro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-chloro-3-trifluoromethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(3-bromo-2-methoxy-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-chloro-2-methyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-chloro-pyridine-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-methyl-2-methylsulfanyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-4-methanesulfonyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-3-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4-bromo-2-chloro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-cyano-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,2,3,3-tetramethyl-cyclopropanecarbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-ethyl-butyryl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-chloro-4-fluoro-benzenesulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-cyano-benzenesulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

{1-[4-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-piperidine-1-carbonyl]-2-methyl-propyl}-carbamic acid tert-butyl ester; compound with trifluoro-acetic acid,

[4-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-piperidin-1-yl]-oxo-acetic acid methyl ester; compound with trifluoro-acetic acid,

4-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-piperidine-1-carboxylic acid methyl ester; compound with trifluoro-acetic acid,

1-(2-Amino-3-methyl-butyryl)-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(2-Methoxy-ethyl)-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

[4-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-piperidin-1-yl]-oxo-acetic acid; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(6-chloro-2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-6-methylsulfanyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(4-trifluoromethyl-pyrimidin-2-yl)-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-cyano-2-fluoro-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-dimethylamino-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-dimethylamino-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-methyl-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-3a,6a-dimethyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-3a,6a-dimethyl-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-dimethylamino-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-dimethylamino-2-methylsulfanyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4-dimethylamino-pyridine-3-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

[4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-3-chloro-phenoxy]-acetic acid; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(5-bromo-2-methylsulfanyl-pyrimidine-4-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide; compound with trifluoro-acetic acid,

1-Dimethylsulfamoyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(2,2,2-Trifluoro-acetyl)-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Dimethylsulfamoyl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(2,2,2-Trifluoro-acetyl)-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Methanesulfonyl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2-hydroxy-propyl}-amide; compound with trifluoro-acetic acid,

4-Oxo-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

[5-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-4,6-dimethyl-pyridin-2-ylsulfanyl]-acetic acid; compound with trifluoro-acetic acid,

Cyclopentanecarboxylic acid {3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-fluoro-phenyl)-amide; compound with trifluoro-acetic acid,

4,4-Difluoro-cyclohexanecarboxylic acid {3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-fluoro-phenyl)-amide,

3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-pyrrolidine-1-carboxylic acid tert-butyl ester,

1-Acetyl-piperidine-4-carboxylic acid {2-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-ethyl}-(3-fluoro-phenyl)-amide,

2-Cyclopentyl-N-{2-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-ethyl}-N-(3-fluoro-phenyl)-acetamide,

N-{2-[5-(4,6-Dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-ethyl}-N-(3-fluoro-phenyl)-propionamide,

N-{2-[5-(4,6-Dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-ethyl}-N-(3-fluoro-phenyl)-3-methyl-butyramide,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-p-tolyl-amide,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,4-dimethyl-1-oxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-p-tolyl-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(6-dimethylamino-2-ethoxy-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1,1-Dioxo-hexahydro-1λ6-thiopyran-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

5-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-3,4-dihydro-1H-isoquinoline-2-carboxylic acid tert-butyl ester,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1,2,3,4-tetrahydro-isoquinoline-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-acetyl-1,2,3,4-tetrahydro-isoquinoline-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide,

4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-3,5-dimethyl-1H-pyrrole-2-carboxylic acid ethyl ester,

[3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-pyrrolidin-1-yl]-oxo-acetic acid methyl ester,

5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carboxylic acid methyl-phenyl-amide; compound with trifluoro-acetic acid,

5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carboxylic acid benzylamide; compound with trifluoro-acetic acid,

1-(3-Chloro-4-methyl-phenyl)-1-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-3-methyl-3-phenyl-urea; compound with trifluoro-acetic acid,

1-Acetyl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Propionyl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

{2-[3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-azetidin-1-yl]-2-oxo-ethoxy}-acetic acid; compound with trifluoro-acetic acid,

4-[3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-azetidin-1-yl]-4-oxo-butyric acid; compound with trifluoro-acetic acid,

3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-azetidine-1-carboxylic acid methyl ester; compound with trifluoro-acetic acid,

[3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-azetidin-1-yl]-oxo-acetic acid methyl ester; compound with trifluoro-acetic acid,

1-Cyclopropanecarbonyl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Butyryl-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(2-Methoxy-acetyl)-azetidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-2,2-dimethyl-propyl}-amide,

4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-sulfonyl)-3,5-dimethyl-1H-pyrrole-2-carboxylic acid ethyl ester,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2-azetidin-1-yl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide,

4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-1,3,5-trimethyl-1H-pyrrole-2-carboxylic acid ethyl ester,

5-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-1,2,4-trimethyl-1H-pyrrole-3-carboxylic acid ethyl ester,

4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-3,5-dimethyl-1H-pyrrole-2-carboxylic acid,

1-Acetyl-piperidine-4-carboxylic acid {3-[5-(2,6-bis-dimethylamino-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-(3-chloro-4-methyl-phenyl)-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(5-methyl-1-phenyl-1H-pyrazole-4-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,
4-Oxo-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,
1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2-methyl-pyridine-3-sulfonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,
1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(3-methyl-benzofuran-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(7-methoxy-3-methyl-1H-indole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
Tetrahydro-pyran-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
N-(3-Chloro-4-methyl-phenyl)-N-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-acetamide; compound with trifluoro-acetic acid,
Cyclopropanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
1-Acetyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
1-(2,2,2-Trifluoro-acetyl)-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
1-Methanesulfonyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
1-Cyclopropanecarbonyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,
3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-pyrrolidine-1-carboxylic acid methyl ester; compound with trifluoro-acetic acid,
1-Dimethylsulfamoyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

[3-((3-Chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-carbamoyl)-pyrrolidin-1-yl]-oxo-acetic acid; compound with trifluoro-acetic acid,

2-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-sulfonyl)-benzoic acid methyl ester; compound with trifluoro-acetic acid,

2-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-sulfonyl)-benzoic acid; compound with trifluoro-acetic acid,

1-Methanesulfonyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

Pyrrolidine-1,3-dicarboxylic acid 3-((3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide) 1-dimethylamide; compound with trifluoro-acetic acid,

1-Cyclopropanesulfonyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Isobutyryl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-Propionyl-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

1-(2,2,2-Trifluoro-ethyl)-pyrrolidine-3-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-sulfonyl)-3,5-dimethyl-1H-pyrrole-2-carboxylic acid,

8-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-3,4-dihydro-1H-isoquinoline-2-carboxylic acid tert-butyl ester,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1-methyl-1H-imidazole-2-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

4-Hydroxy-cyclohexanecarboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(1,2,3,4-tetrahydro-isoquinoline-8-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide,

1-Acetyl-piperidine-4-carboxylic acid (5-chloro-thiazol-2-yl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

3-[4-(5-{3-[(1-Acetyl-piperidine-4-carbonyl)-(3-chloro-4-methyl-phenyl)-amino]-propyl}-hexahydro-pyrrolo[3,4-c]pyrrole-2-carbonyl)-3,5-dimethyl-pyrazol-1-yl]-benzoic acid methyl ester,

1-Acetyl-piperidine-4-carboxylic acid (5-chloro-thiazol-2-yl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

3-(3-(3-Chloro-4-methyl-phenyl)-3-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-ureido)-benzoic acid; compound with trifluoro-acetic acid, or

3-(3-(3-Chloro-4-methyl-phenyl)-3-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-ureido)-benzoic acid ethyl ester; compound with trifluoro-acetic acid.

16. A compound according to claim 1, which is:

1-acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,6-dimethyl-benzoyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide,

1-acetyl-piperidine-4-carboxylic acid (3-chloro-4-methyl-phenyl)-{3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide,

[4-((3-chloro-4-methyl-phenyl)-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-carbamoyl)-piperidin-1-yl]-oxo-acetic acid; compound with trifluoro-acetic acid,

cyclopentanecarboxylic acid {(S)-1-phenyl-3-[5-(pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-propyl}-amide; compound with trifluoro-acetic acid,

cyclopentanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide,

cyclopentanecarboxylic acid {(S)-3-[5-(2,4-dimethyl-pyridine-3-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-propyl}-amide,

3-(3-(3-chloro-4-methyl-phenyl)-3-{3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-c]pyrrol-2-yl]-propyl}-ureido)-benzoic acid; compound with trifluoro-acetic acid,

4,4-difluoro-cyclohexanecarboxylic acid {(S)-3-[5-(4,6-dimethyl-pyrimidine-5-carbonyl)-hexahydro-pyrrolo[3,4-*c*]pyrrol-2-yl]-1-phenyl-butyl}-amide, or

(5-{5-[(S)-3-(cyclopentanecarbonyl-amino)-3-phenyl-propyl]-hexahydro-pyrrolo[3,4-*c*]pyrrole-2-carbonyl}-4,6-dimethyl-pyrimidin-2-yloxy)-acetic acid; compound with trifluoro-acetic acid.

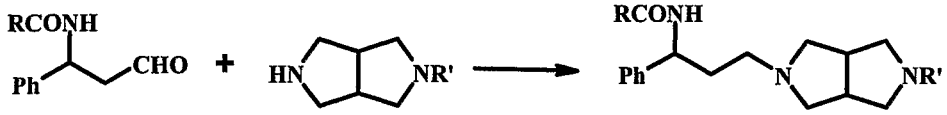
17. A compound of formula I according to any one of claim 1 to 16 for use as medicament.

18. Use of the compound of formula I according to any one of claims 1 to 16 for the manufacture of a medicament for the treatment of diseases mediated by the human immunodeficiency virus (HIV).

19. A pharmaceutical composition comprising a pharmaceutically effective amount of compound or a pharmaceutically acceptable salt thereof as defined in any one of claim 1 to 16 and a pharmaceutical inert carrier.

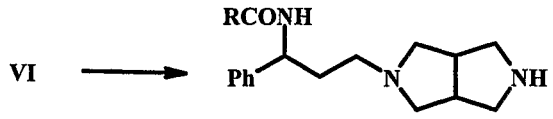
20. A pharmaceutical composition according to claim 14 for the use in the treatment of diseases mediated by the human immunodeficiency virus (HIV).

21. A process for the preparation of a compound according to formula I wherein R^1 is hydrogen and R^2 is optionally substituted phenyl or optionally substituted heteroaryl comprising the steps of:



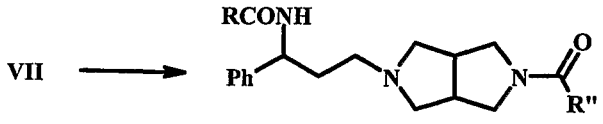
IV V VI

(a) contacting V wherein R' is a benzyl, optionally substituted benzoyl or *tert*-butoxycarbonyl protecting group with an β-acylaminoaldehyde of formula IV, wherein R is a group R^3 according to claim 1, and sodium triacetoxyborohydride under acidic conditions sufficient to promote formation of an imine and reduction of the corresponding iminium salt to afford VI;



VII

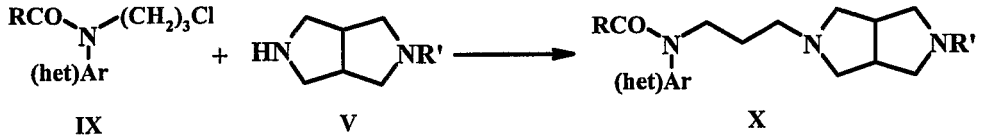
(b) optionally exposing VI to a conditions sufficient to effect deprotection when R' is a protecting group to afford VII;



VIII

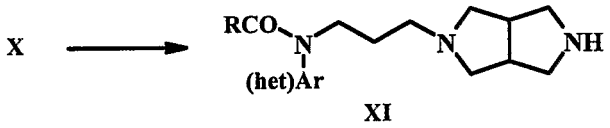
(c) contacting VII with a carboxylic acid, R"CO_2H, or an activated carboxyl acid derivative, under conditions sufficient to promote acylation to afford VIII wherein R" is a group R^4 according to claim 1.

22. A process for the preparation of a compound according to formula I wherein R^1 is optionally substituted phenyl or optionally substituted heteroaryl and R^2 is hydrogen comprising the steps of:



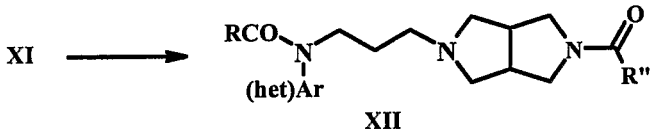
IX V X

(a) contacting V wherein R' is a benzyl, optionally substituted benzoyl or *tert*-butoxycarbonyl protecting group with an β-acylaminoaldehyde of formula IX, where R is a group R^3 according to claim 1 and (het)Ar denotes an optionally substituted phenyl or heterocyclic group according to claim 1, under conditions sufficient to afford X;



XI

(b) optionally exposing X to a reagent sufficient to effect deprotection when R' is a protecting group to afford XI;



XII

(c) contacting XI with a carboxylic acid, R"CO_2H, or an activated carboxyl acid derivative under conditions wherein acylation occurs to afford XII wherein R" is a group R^4 according to claim 1.

23. The invention as hereinbefore described.