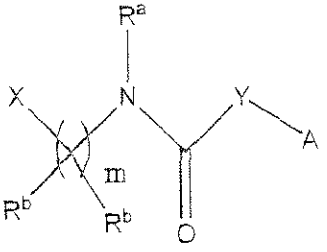
injury, peptic ulcer Helicobacter related diseases, virally-induced angiogenic disorders, preeclampsia, menometrorrhagia, cat scratch fever, rubeosis, neovascular glaucoma, diabetic retinopathy, retinopathy of prematurity, age-related macular degeneration, hyperproliferative disorder, thyroid hyperplasia, Grave's disease, cysts, hypervascularity of ovarian stroma characteristic of polycystic ovarian syndrome, Stein-Leventhal syndrome, or polycystic kidney.

In another embodiment the invention provides a method according to the previous embodiment wherein the disease or condition is osteoarthritis, rheumatoid arthritis, juvenile arthritis, psoriatic arthritis, degenerative joint disease or systemic lupus erythematosus.

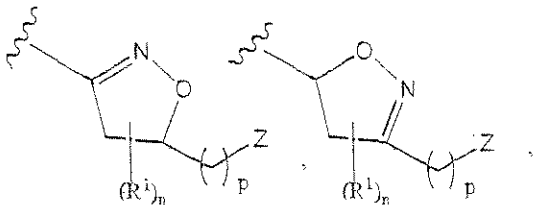
In another embodiment, the invention provides a compound of Formula (I)



Formula (I)

stereoisomers, and pharmaceutically acceptable salts thereof wherein

A is



X is phenyl, benzo[1,4]oxazin-3-onyl, benzoxazolyl, 2,3-dihydrobenzo[1,4]dioxinyl, 2,3-dihydrobenzo furanyl, pyrazinonyl, pyridinyl, pyridonyl, or quinolinyl, each of which is optionally substituted with one or more substituents independently selected from Br, Cl, F, CF_3, CN, COOH, OCF_3, OH, $(C_1$-$C_6)$alkyl, or $(C_1$-$C_6)$alkoxy;

Y is pyridazinonyl, pyridinyl, or pyrimidinyl, each of which is optionally substituted by one or more substituents independently selected from CF_3, halogen, $(C_1$-$C_6)$alkyl, $(C_1$-$C_6)$alkoxy,

Z is $-C(O)NH_2$, CN, -C(O)-morpholinyl, -C(O)-pyrrolidinyl, $-C(O)-N(H)CH_2CH_2OH$, $(C_3$-$C_6)$cycloalkyl, 1,4-dioxanyl, imidazolyl, morpholinyl, phenyl, piperazinyl, piperidinyl, pyrazinyl, pyrazolyl, pyrazolo[3,4-*b*]pyridinyl, pyridazinyl, pyridazinonyl, pyridinyl, pyridonyl, pyrimidinyl, pyrrolidinyl, tetrahydropyranyl, tetrazolyl, or thienyl, each of which is optionally substituted by one or more substituents independently selected from Br, Cl, F, CF_3, CN, -C(O)OH, $-C(O)OCH_3$, $-C(O)CH_3$, $-C(O)CH_2OC(O)CH_3$, $-C(O)C(CH_3)_3$, $-C(O)C(H)(CH_3)_2$, $-C(O)CH_2CN$, $-C(O)CH_2OH$, $-C(O)CH_2OCH_3$, $-C(O)CH_2CH_2N(CH_3)_2$, $-C(O)C(CH_3)_2OH$, $-C(O)OC(CH_3)_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)CH_2N(CH_3)_2$, $-C(O)CH_2N(H)S(O)_2CH_3$, $-C(O)CH_2N(H)C(O)CH_3$, $-C(O)CH_2N(H)C(O)N(H)CH_3$, $-C(O)CH_2OCH_3$, $-C(O)CH_2S(O)_2CH_3$, $-C(O)NH_2$, $-C(O)N(CH_3)_2$, $-C(O)N(H)CH_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)N(H)CH(CH_3)_2$, -

$C(O)N(H)CH_2CH_2OH$, $-C(O)N(H)$-isoxazolyl, $-C(O)N(H)$thiazolyl, $-C(O)N(CH_3)CH_2CH_2OH$, $-C(O)$-cyclopropyl, $-C(O)$-cyclobutyl, $-OCH_2CH_2OH$, $-OC(CH_3)_3$, $-NH_2$, $-N(CH_3)_2$, $-N(H)C(O)CH_3$-, $-N(H)C(O)CH_2OH$, $-N(H)C(O)CH_2CN$, $-N(H)C(O)C(CH_3)_2OH$, $-N(H)C(O)C(H)(OH)CH_3$, $-N(H)C(O)CH_2N(CH_3)_2$, $-N(H)C(O)CH_2N(H)CH_3$, $-N(H)C(O)(OH)CH_3$, $-N(R^c)C(O)N(R^c)(R^e)$, $-N(H)C(O)N(CH_3)_2$, $-N(H)C(O)N(H)CH_3$-$N(H)C(O)N(H)CH_2CH_2OH$, $-N(H)C(O)N(H)C(H)(CH_2OH)_2$, $-N(H)C(O)N(CH_3)CH_2CH_2OH$, $-N(H)C(O)N(CH_3)CH_2C(H)(OH)CH_2OH$, $-N(H)C(O)O(CH_3)_3$, $-N(H)S(O)_2NH_2$, $-N(H)S(O)_2CH_3$, $-N(R^c)S(O)_2N(R^c)(R^e)$, $-NO_2$, 1,2,4-oxadiazolyl, oxo, $-OH$, $-S(O)_2(R^e)$, $-S(O)CH_3$, $-S(O)CH_2CH_3$, $-S(O)CH_2CH_2CH_3$, $-S(O)_2C(H)(CH_3)_2$, $-S(O)_2$cyclopropyl, $-S(O)_2$-imidazolyl, $-S(O)_2$-isoxazolyl, $-CH_2CN$, $-CH_2C(O)N(H)CH_3$, $-CH_2C(O)N(H)$-cyclopropyl, $-CH_2OCH_3$, $-CH_2OH$, $-CH_2CH_2OH$, $-CH_2OS(O)_2$-phenyl, $-C(CH_3)OH$, $-C(H)(CH_2)_2OH$, $-CH_2NH_2$, $-CH_2N(CH_3)_2$, $-CH_2N(H)C(O)CH_3$, $-CH_2N(H)C(O)C(H)(OH)CH_3$, $-CH_2N(H)C(O)CH_2CN$, $-CH_2N(H)C(O)CH_2OH$, $-CH_2N(H)C(O)C(CH_3)_2OH$, $-CH_2N(H)C(O)N(H)CH_3$, $-CH_2N(H)C(O)N(CH_3)_2$, $-CH_2S(O)_2CH_3$, $-CH_2$-morpholinyl, $-CH_2$-thiomorpholinyl, $-CH_2$-thiomorpholinyl 1,1-dioxide, morpholinyl, oxadiazolyl, oxazolyl, piperidinyl, $(C_1$-$C_6)$alkyl, $(C_1$-$C_6)$alkoxy, or $(C_3$-$C_6)$cycloalkyl;

R^c is independently H, $(C_1$-$C_3)$alkyl or $(C_3$-$C_6)$cycloalkyl;

R^e is H, $-C(O)R^c$, $(C_1$-$C_6)$alkyl, $(C_1$-$C_6)$hydroxyalkyl, $(C_3$-$C_6)$cycloalkyl, heteroaryl optionally substituted with $(C_1$-$C_3)$alkyl, or heterocyclyl; or

R^e is independently H, $(C_1$-$C_3)$alkyl or $(C_3$-$C_6)$cycloalkyl,

R^b is independently H, CF_3, $(C_1$-$C_3)$alkyl or $(C_3$-$C_6)$cycloalkyl;

R^1 is independently Br, Cl, F, $(C_1$-$C_3)$alkyl or $(C_3$-$C_6)$cycloalkyl;

m is 0, 1 or 2;

n is 0; and

p is 0 or 1.

In another embodiment, the invention provides a compound selected from the group consisting of:

6-(5-(1-Acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-Carbamoylphenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(morpholine-4-carbonyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(2-(2-hydroxypropan-2-yl)pyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(2*H*-tetrazol-5-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-

methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide hydrochloride;

6-(5-(1-(ethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclopropanecarbonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-acetamidoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-cyanoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylsulfonyl) acetyl) piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxy ethyl carbamoyl) piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(isoxazol-3-ylcarbamoyl) piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*, 5*R*)-5-(hydroxymethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*, 5*R*)-5-(hydroxymethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(methylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*, 5*R*)-5-(cyanomethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5- ((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(2*S*,5*R*)-5-((*S*)-3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxane-2-carboxylic acid;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(morpholinomethyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*S*)-5-Carbamoyl-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1*H*-imidazol-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-(2-hydroxyethoxy)phenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-(4-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-phenyl-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(4-(hydroxymethyl)phenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-6-(5-(4-methoxyphenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-(4-morpholinophenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-(4-(piperidin-1-yl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(4-(dimethylamino)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(3-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-cyclohexyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-cyclohexyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-(hydroxymethyl)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-carbamoylphenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-cyclohexyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-hydroxycyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

tert-butyl 4-(3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)piperidine-1-carboxylate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-3-yl)-4,5-dihydroisoxazol-3-

yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(hydroxymethyl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(4-(hydroxymethyl)piperidin-1-yl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidin-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(4-(morpholinomethyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-cyano-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(4- (thiomorpholinomethyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-cyano-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(3-carbamoylphenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(3-(morpholinomethyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(3-(methylcarbamoyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

methyl 4-(3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(4-(methylcarbamoyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(thiophen-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-{5-[4-(1,1,1,-Dioxo-thiomorpholine-4-yl methyl)phenyl]-4,5-dihydro-isoxazol-3-yl}-2-methyl pyrimidine-4-carboxylate;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(2-cyanoacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(sulfamoylamino)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-3-yl)-4,5-dihydro-1*H*-pyrazol-3-yl)pyrimidine-4-carboxamide;

6-(3-(4-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-5-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(3-phenyl-4,5-dihydroisoxazol-5-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(3-(4-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-5-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(3-phenyl-4,5-dihydroisoxazol-5-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(2-hydroxyethylcarbamoyl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1-acetylpiperidin-4-yl)methyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrimidin-5-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(2-hydroxypropan-2-yl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1-(2-hydroxyacetyl)piperidin-4-yl)methyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1-(methylsulfonyl)piperidin-4-yl)methyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(2-carbamoylpyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1*H*-pyrazol-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((1*r*,4*R*)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*R*)-5-((1*r*,4*R*)-4-(methylsulfonamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((1*r*,4*R*)-4-(2-hydroxyacetamido)cyclohexyl)-

4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrimidin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(tetrahydro-2*H*-pyran-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(methylsulfonamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(2-hydroxyacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrazin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxy-2-methylpropanoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclopropylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(ethylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-isobutyrylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(propylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-methoxybenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-methoxybenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclopropanecarbonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(dimethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(ethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-(dimethylamino)acetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-acetamidoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(isopropylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-cyanoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-methoxyacetyl)piperidin-4-yl)-4,5-

dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclobutanecarbonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylsulfonamido)acetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(5-cyanopyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(5-(hydroxymethyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(methylcarbamoyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(3-methylureido)acetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(5-(1-hydroxyethyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-((2-hydroxyethyl)(methyl)carbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(isopropylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(1-methyl-1*H*-imidazol-2-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(1-methyl-1*H*-imidazol-4-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-(1,2-dimethyl-1*H*-imidazol-4-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(3,5-dimethylisoxazol-4-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(methylsulfonyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(5-acetylpyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(3-methyl-1,2,4-oxadiazol-5-yl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

((2*S*,5*R*)-5-(3-(6-(3-chloro-4-fluorobenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxan-2-yl)methyl 4-methylbenzenesulfonate;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide ;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

((2*S*,5*R*)-5-((*S*)-3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxan-2-yl)methyl 4-methylbenzenesulfonate;

methyl 4-(3-(2-(3-methoxybenzylcarbamoyl)pyridin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoate;

N-(3-methoxybenzyl)-4-(5-(4-methoxyphenyl)-4,5-dihydroisoxazol-3-yl)picolinamide;

4-(5-(4-chlorophenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)picolinamide;

4-(3-(2-(3-methoxybenzylcarbamoyl)pyridin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoic acid

4-(5-(4-((dimethylamino)methyl)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)picolinamide;

4-(3-(2-(4-methoxybenzylcarbamoyl)pyridin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoic acid;

4-(5-(4-chlorophenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)picolinamide;

N-(4-methoxybenzyl)-4-(5-(pyrazin-2-yl)-4,5-dihydroisoxazol-3-yl)picolinamide;

N-(4-methoxybenzyl)-4-(5-(4-methoxyphenyl)-4,5-dihydroisoxazol-3-yl)picolinamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-N-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((R)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*R*)-5-(methylsulfonamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-(methylsulfonylmethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(thiazol-2-ylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(Acetamidomethyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((1r,4*R*)-4-(acetamidomethyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(oxazol-5-yl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(methylsulfonyl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(3-methyl-1,2,4-oxadiazol-5-yl)pyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(6-acetylpyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)pyridazine-3-carboxamide 2,2,2-trifluoroacetate;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((1*r*,4*R*)-4-((2-hydroxyacetamido)methyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-((2-hydroxyacetamido)methyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-3-(3-methyl-1,2,4-oxadiazol-5-yl)-1*H*-pyrazol-5-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(3-carbamoyl-1-methyl-1*H*-pyrazol-5-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-((2-cyanoacetamido)methyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-*N*-methylpyridazine-3-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(3-methyl-1,2,4-oxadiazol-5-yl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5R)-5-((2-hydroxyacetamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(oxazol-5-yl)pyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-(hydroxymethyl)pyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(3-(2-hydroxyethyl)ureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(3-(2-hydroxyethyl)-3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(methylsulfonyl)pyrazin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-

methylpyrimidine-4-carboxamide;

6-(5-(1-(3-(dimethylamino)propanoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

(2*R*,5*S*)-5-((*S*)-3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxane-2-carboxylic acid;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3,3-dimethylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-dichlorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-dichlorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-((2-cyanoacetamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-((2-hydroxy-2-methylpropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*R*)-5-((3-methylureido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-((3,3-dimethylureido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrrolidine-1-carbonyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(2-(dimethylamino)acetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1*H*-pyrazolo[3,4-*b*]pyridin-5-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-1*H*-pyrazolo[3,4- *b*]pyridin-5-

yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(bromomethyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-*N*-(2-hydroxyethyl)-4,5-dihydroisoxazole-5-carboxamide;

3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazole-5-carboxamide;

methyl 3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazole-5-carboxylate;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5S)-5-(2-hydroxypropan-2-yl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-(methylcarbamoyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(methylcarbamoyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-((1r,4r)-4-carbamoylcyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(2-hydroxy-2-methylpropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((3-(hydroxymethyl)pyrrolidin-1-yl)methyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-((4-acetylpiperazin-1-yl)methyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-(6-(dimethylamino)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(pyrrolidin-1-yl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-((1r,4r)-4-(3-(1,3-dihydroxypropan-2-yl)ureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-((2-hydroxyethyl)carbamoyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3-(2,3-dihydroxypropyl)-3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-Acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-

methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(2-(methylamino)acetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(3-methylbenzyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(3-methylbenzyl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(3-chloro-4-fluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(3-chloro-4-fluorobenzyl)-6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzofuran-5-yl)methyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((2-methylbenzo[d]oxazol-5-yl)methyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((2-methylbenzo[*d*]oxazol-5-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(methoxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(methoxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(((*S*)-2-hydroxypropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-(5-((1S,4r)-4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzo[*b*][1,4]dioxin-6-yl)methyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzo[*b*][1,4]dioxin-6-yl)methyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

tert-butyl ((1r,4r)-4-(3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)cyclohexyl)carbamate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-isopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(pyridin-4-ylmethyl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(pyridin-4-ylmethyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(pyridin-4-ylmethyl)pyrimidine-4-carboxamide;

6 6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((1-methyl-2-oxo-1,2-dihydropyridin-4-yl)methyl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-((*R*)-1-(4-fluoro-3-methoxyphenyl)ethyl)-2-methyl-6-(5-phenyl-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-((*S*)-1-(4-fluoro-3-methoxyphenyl)ethyl)-2-methyl-6-(5-phenyl-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((1-methyl-2-oxo-1,2-dihydropyridin-4-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((1-methyl-2-oxo-1,2-dihydropyridin-4-yl)methyl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(quinolin-4-ylmethyl)pyrimidine-4-carboxamide;

(*S*)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((1r,4S)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-

methyl-*N*-(quinolin-4-ylmethyl)pyrimidine-4-carboxamide;

6-((R)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(quinolin-4-ylmethyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-((2-hydroxy-2-methylpropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((S)-5-((1r,4S)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluorobenzyl)-6-(5-((1S,4r)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-((2-oxo-1,2,3,4,4a,8a-hexahydroquinolin-7-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((2-oxo-1,2,3,4-tetrahydroquinolin-7-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-2-oxo-

1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(S)-*N*-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(R)-*N*-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxypyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxypyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

(S)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(R)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(methoxy(methyl)amino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(5-chloro-1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxy-5,6-dihydropyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-(dimethylamino)-2-oxoethyl)-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((1s,4R)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((1r,4R)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((1-methyl-6-oxo-1,6-dihydropyridin-3-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(isopropylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1S,4r)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((3-oxo-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-(5-(3-chloro-5-cyclopropyl-4-oxocyclohexyl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(S)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(R)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(R)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((3-oxo-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-ca)rboxamide;

6-(5-(1-(2-(cyclopropylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-1-methyl-6-oxo-5-(5-phenyl-4,5-dihydroisoxazol-3-yl)-1,6-dihydropyridazine-3-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(1-methoxyethyl)-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide; and

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(isopropylamino)allyl)-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide; or

a pharmaceutically acceptable salt thereof.

In another embodiment, the invention provides a pharmaceutical composition comprising a compound as defined herein, stereoisomer, or pharmaceutically acceptable salt thereof and a pharmaceutically acceptable carrier or excipient.

In another embodiment, the invention provides a compound or a pharmaceutically acceptable salt thereof, wherein the compound is

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide.

DETAILED DESCRIPTION OF THE INVENTION

Many autoimmune diseases and disease associated with chronic inflammation, as well as acute responses, have been linked to excessive or unregulated production or activity of one or more cytokines.

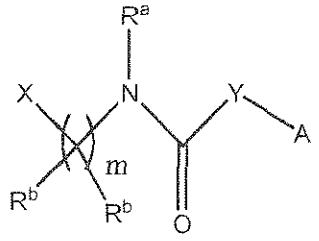
The compounds of the invention are useful in the treatment of diseases mediated by MMP-13 enzyme such as Osteoarthritis as well as other related diseases such as Rheumatoid arthritis, Juvenile arthritis, Psoriatic arthritis, Gouty arthritis, Degenerative joint disease and Systemic lupus erythematosus.

The compounds of the invention are also useful in the treatment of cardiovascular disorders, such as acute myocardial infarction, acute coronary syndrome, chronic heart failure, myocardial infarction, atherosclerosis, viral myocarditis, cardiac allograft rejection, and sepsis-associated cardiac dysfunction. Furthermore, the compounds of the present invention are also useful for the treatment of central nervous system disorders such as meningococcal meningitis, Alzheimer's disease and Parkinson's disease.

The compounds of the invention are also useful in the treatment of an ocular condition, a cancer, rheumatoid arthritis, ankylosing spondilitis, a solid tumor, a sarcoma, fibrosarcoma, osteoma, melanoma, retinoblastoma, a rhabdomyosarcoma, glioblastoma, neuroblastoma, teratocarcinoma, hypersensitivity reactions. hyperkinetic movement disorders, hypersensitivity pneumonitis, hypertension, hypokinetic movement disorders, aordic and peripheral aneuryisms, hypothalamic-pituitary-adrenal axis evaluation, aortic dissection, arterial hypertension, arteriosclerosis, arteriovenous fistula, ataxia, spinocerebellar degenerations, streptococcal myositis, structural lesions of the cerebellum, subacute sclerosing panencephalitis, Syncope, syphilis of the cardiovascular system, systemic anaphalaxis, systemic inflammatory response

What is claimed:

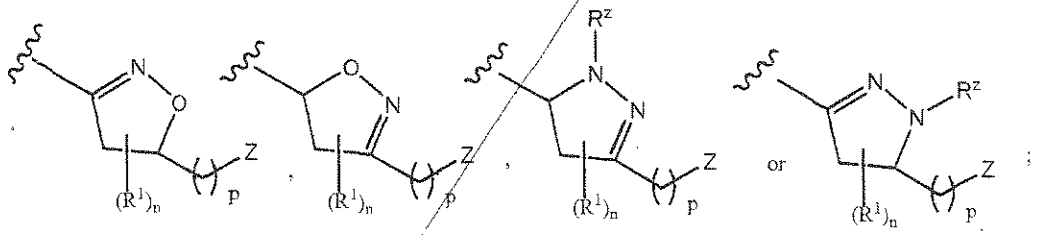
1. A compound of Formula (I)



Formula (I)

biologically active metabolites, pro-drugs, isomers, stereoisomers, solvates, hydrates and pharmaceutically acceptable salts thereof wherein

A is



X is an optionally substituted ring system which is aromatic, partly aromatic or non-aromatic having one or more fused ring systems;

Y is optionally substituted heteroaryl;

Z is $-C(O)NR^cR^d$, $-C(O)OCH_3$, $-NR^cR^d$, optionally substutited (C_1-C_6)alkyl, optionally substituted (C_3-C_6)cycloalkyl, optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl;

R^a is independently H, optionally substituted (C_1-C_3)alkyl or optionally substituted (C_3-C_6)cycloalkyl;

R^b is independently H, CF_3, optionally substituted (C_1-C_3)alkyl or optionally substituted (C_3-C_6)cycloalkyl;

R^c is H and R^d is H or optionally substituted (C_1-C_6)alkyl; or

R^e and R^d, together with the nitrogen atom to which they are attached, form a 5- or 6-membered saturated or unsaturated, aromatic or nonaromatic heterocyclic ring system;

R^1 is independently H, OR^a, Br, Cl, F, optionally substituted $(C_1$-$C_3)$alkyl or optionally substituted $(C_3$-$C_6)$cycloalkyl;

R^z is H, optionally substituted $(C_1$-$C_3)$alkyl or optionally substituted $(C_3$-$C_6)$cycloalkyl;

m is 0, 1 or 2;

n is 0, 1, 2 or 3; and

p is 0, 1, 2 or 3;

provided that when Y is pyridinyl it is not substituted by optionally substituted phenyl.

2. The compound of claim 1 wherein

X is optionally substituted naphthyl, optionally substituted phenyl, optionally substituted azaindolyl, optionally substituted benzo[1,4]oxazin-3-onyl, optionally substituted benzo(b)thienyl, optionally substituted benzimidazolyl, optionally substituted benzo[1,3]dioxolyl, optionally substituted benzofuranyl, optionally substituted benzoxazolyl, optionally substituted benzothiazolyl, optionally substituted benzothiadiazolyl, optionally substituted benzoxadiazolyl, optionally substituted 2,3-dihydrobenzo[1,4]dioxinyl, optionally substituted 2,3-dihydrobenzo furanyl, optionally substituted furanyl, optionally substituted imidazolyl, optionally substituted imidazopyridinyl, optionally substituted indolyl, optionally substituted indazolyl, optionally substituted isoxazolyl, optionally substituted isothiazolyl, optionally substituted oxadiazolyl, optionally substituted oxazolyl, optionally substituted purinyl, optionally substituted pyranyl, optionally substituted pyrazinonyl, optionally substituted pyrazinyl, optionally substituted pyrazolyl, optionally substituted pyridazinyl, optionally substituted pyridazinonyl, optionally substituted pyridinyl, optionally substituted pyridonyl, optionally substituted pyrimidinonyl, optionally substituted pyrimidinyl, optionally substituted pyrrolyl, optionally substituted pyrrolo[2,3-*d*]pyrimidinyl, optionally substituted pyrazolo[3,4-*d*]pyrimidinyl, optionally substituted 1*H*-quinoxalin-2-only, optionally substituted quinolinyl, optionally substituted quinazolinyl, optionally substituted triazolyl, optionally substituted thiazolyl, optionally substituted tetrazolyl, optionally substituted thiadiazolyl, or optionally substituted thienyl.

3. The compound of claim 1 wherein Y is optionally substituted furanyl, optionally substituted imidazolyl, optionally substituted isoxazolyl, optionally substituted isothiazolyl, optionally substituted oxazolyl, optionally substituted pyrazolyl, optionally substituted pyridazinyl, optionally substituted pyridazinonyl, optionally substituted pyridinyl, optionally substituted pyrimidinyl, optionally substituted pyrrolyl, optionally substituted thienyl, optionally substituted triazinyl, or optionally substituted 1,2,4-triazolyl.

4. The compound of claim 1 wherein Z is $-C(O)NH_2$, -C(O)-optionally substituted morpholinyl, -C(O)-optionally substituted pyrrolidinyl, $-C(O)-N(H)CH_2CH_2OH$, optionally substituted (C_3-C_6)cycloalkyl, optionally substituted azaindolyl, optionally substituted azepinyl, optionally substituted azetidinyl, optionally substituted benzo(b)thienyl, optionally substituted benzimidazolyl, optionally substituted benzofuranyl, optionally substituted benzoxazolyl, optionally substituted benzothiazolyl, optionally substituted benzothiadiazolyl, optionally substituted benzoxadiazolyl, optionally substituted 1,4-dioxanyl, optionally substituted furanyl, optionally substituted imidazolyl, optionally substituted imidazopyridinyl, optionally substituted indolinyl, optionally substituted indolyl, optionally substituted indazolyl, optionally substituted isoindolinyl, optionally substituted isoxazolyl, optionally substituted isothiazolyl, optionally substituted morpholinyl, optionally substituted naphthyl, optionally substituted oxadiazolyl, optionally substituted phenyl, optionally substituted oxazolyl, optionally substituted piperazinyl, optionally substituted piperidinyl, optionally substituted purinyl, optionally substituted pyranyl, optionally substituted pyrazinyl, optionally substituted pyrazolyl, optionally substituted pyrazolo[3,4-*b*]pyridinyl, optionally substituted *N*-methylpyrazonyl, optionally substituted pyridazinyl, optionally substituted pyrazinonyl, optionally substituted pyridazinonyl, optionally substituted pyridinyl, optionally substituted pyridonyl, optionally substituted pyrimidinonyl, optionally substituted *N*-methylpyridazonyl, optionally substituted *N*-(C_1-C_6)alkylpyridonyl, optionally substituted pyrimidinyl, optionally substituted *N*-(C_1-C_6)alkylpyrimidonyl, optionally substituted pyrrolidinyl, optionally substituted pyrrolyl, optionally substituted pyrrolo[2,3-*d*]pyrimidinyl, optionally substituted pyrazolo[3,4-*d*]pyrimidinyl, optionally substituted quinolinyl, optionally substituted quinazolinyl, optionally substituted quinucludinyl, optionally substituted tetrahydrofuranyl, optionally substituted tetrahydroindolyl, optionally substituted tetrahydropyranyl, optionally substituted thiomorpholinyl, optionally substituted triazolyl, optionally substituted thiazolyl, optionally substituted tetrazolyl, optionally substituted thiadiazolyl, or optionally substituted thienyl, optionally substituted thiomorpholinyl, or optionally substituted tropanyl.

5. The compound of claim 1 wherein

X is optionally substituted naphthyl, optionally substituted phenyl, optionally substituted azaindolyl, optionally substituted benzo[1,4]oxazin-3-onyl, optionally substituted benzo(b)thienyl, optionally substituted benzimidazolyl, optionally substituted benzo[1,3]dioxolyl, optionally substituted benzofuranyl, optionally substituted benzoxazolyl, optionally substituted benzothiazolyl, optionally substituted benzothiadiazolyl, optionally substituted benzoxadiazolyl, optionally substituted 2,3-dihydrobenzo[1,4]dioxinyl, optionally substituted 2,3-dihydrobenzo furanyl, optionally substituted furanyl, optionally substituted imidazolyl, optionally substituted imidazopyridinyl, optionally substituted indolyl, optionally substituted indazolyl, optionally substituted isoxazolyl, optionally substituted isothiazolyl, optionally substituted oxadiazolyl, optionally substituted oxazolyl, optionally substituted purinyl, optionally substituted pyranyl, optionally substituted pyrazinonyl, optionally substituted pyrazinyl, optionally substituted pyrazolyl, optionally substituted pyridazinyl, optionally substituted pyridazinonyl, optionally substituted pyridinyl, optionally substituted pyridonyl, optionally substituted pyrimidinonyl, optionally substituted pyrimidinyl, optionally substituted pyrrolyl, optionally substituted pyrrolo[2,3-*d*]pyrimidinyl, optionally substituted pyrazolo[3,4-*d*]pyrimidinyl, optionally substituted 1*H*-quinoxalin-2-only, optionally substituted quinolinyl, optionally substituted quinazolinyl, optionally substituted triazolyl, optionally substituted thiazolyl, optionally substituted tetrazolyl, optionally substituted thiadiazolyl, or optionally substituted thienyl;

Y is optionally substituted furanyl, optionally substituted imidazolyl, optionally substituted isoxazolyl, optionally substituted isothiazolyl, optionally substituted oxazolyl, optionally substituted pyrazolyl, optionally substituted pyridazinonyl, optionally substituted pyridazinyl, optionally substituted pyridinyl, optionally substituted pyrimidinyl, optionally substituted pyrrolyl, optionally substituted thienyl, optionally substituted triazinyl, or optionally substituted 1,2,4-triazolyl; and

Z is $-C(O)NH_2$, -C(O)-optionally substituted morpholinyl, -C(O)-optionally substituted pyrrolidinyl, $-C(O)-N(H)CH_2CH_2OH$, optionally substituted (C_3-C_6)cycloalkyl, optionally substituted azaindolyl, optionally substituted azepinyl, optionally substituted azetidinyl, optionally substituted benzo(b)thienyl, optionally substituted benzimidazolyl, optionally substituted benzofuranyl, optionally substituted benzoxazolyl, optionally substituted benzothiazolyl, optionally substituted benzothiadiazolyl, optionally substituted benzoxadiazolyl, optionally substituted 1,4-dioxanyl, optionally substituted furanyl, optionally substituted imidazolyl, optionally substituted imidazopyridinyl, optionally substituted indolinyl, optionally substituted indolyl, optionally substituted indazolyl, optionally substituted isoindolinyl, optionally substituted isoxazolyl, optionally substituted isothiazolyl, optionally substituted morpholinyl, optionally substituted naphthyl, optionally substituted oxadiazolyl, optionally substituted phenyl, optionally substituted oxazolyl, optionally substituted piperazinyl, optionally substituted

piperidinyl, optionally substituted purinyl, optionally substituted pyranyl, optionally substituted pyrazinyl, optionally substituted pyrazolyl, optionally substituted pyrazolo[3,4-*b*]pyridinyl, optionally substituted *N*-methylpyrazonyl, optionally substituted pyridazinyl, optionally substituted pyrazinonyl, optionally substituted pyridazinonyl, optionally substituted pyridinyl, optionally substituted pyridonyl, optionally substituted pyrimidinonyl, optionally substituted *N*-methylpyridazonyl, optionally substituted *N*-(C_1-C_6)alkylpyridonyl, optionally substituted pyrimidinyl, optionally substituted *N*-(C_1-C_6)alkylpyrimidonyl, optionally substituted pyrrolidinyl, optionally substituted pyrrolyl, optionally substituted pyrrolo[2,3-*d*]pyrimidinyl, optionally substituted pyrazolo[3,4-*d*]pyrimidinyl, optionally substituted quinolinyl, optionally substituted quinazolinyl, optionally substituted quinucludinyl, optionally substituted tetrahydrofuranyl, optionally substituted tetrahydroindolyl, optionally substituted tetrahydropyranyl, optionally substituted thiomorpholinyl, optionally substituted triazolyl, optionally substituted thiazolyl, optionally substituted tetrazolyl, optionally substituted thiadiazolyl, or optionally substituted thienyl, optionally substituted thiomorpholinyl, or optionally substituted tropanyl.

6. The compound of claim 5 wherein X is optionally substituted naphthyl, optionally substituted phenyl, optionally substituted benzo[1,4]oxazin-3-onyl, optionally substituted benzoxazolyl, optionally substituted 2,3-dihydrobenzo[1,4]dioxinyl, optionally substituted 2,3-dihydrobenzo furanyl, optionally substituted furanyl, optionally substituted imidazolyl, optionally substituted isoxazolyl, optionally substituted isothiazolyl, optionally substituted oxadiazolyl, optionally substituted oxazolyl, optionally substituted pyrazinyl, optionally substituted pyrazolyl, optionally substituted pyrazinonyl, optionally substituted pyridazinonyl, optionally substituted pyridinyl, optionally substituted pyridonyl, optionally substituted pyrimidinonyl, optionally substituted pyrimidinyl, optionally substituted pyrrolyl, optionally substituted triazolyl, optionally substituted thiazolyl, optionally substituted tetrazolyl, or optionally substituted thienyl.

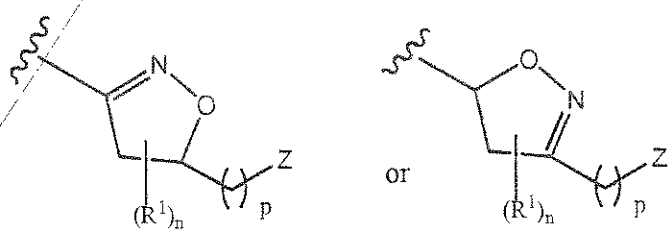
7. The compound of claim 6 wherein Z is optionally substituted (C_3-C_6)cycloalkyl, optionally substituted azetidinyl, optionally substituted 1,4-dioxanyl, optionally substituted imidazolyl, optionally substituted isothiazolyl, optionally substituted morpholinyl, optionally substituted phenyl, optionally substituted piperazinyl, optionally substituted piperidinyl, optionally substituted purinyl, optionally substituted pyranyl, optionally substituted pyrazinonyl, optionally substituted pyrazinyl, optionally substituted pyrazolyl, optionally substituted pyrazolo[3,4-*b*]pyridinyl, optionally substituted pyridazinyl, optionally substituted pyridazinonyl,

optionally substituted pyridinyl, optionally substituted pyridonyl, optionally substituted pyrimidinonyl, optionally substituted pyrimidinyl, optionally substituted pyrrolidinyl, optionally substituted pyrrolyl, optionally substituted tetrahydropyranyl, optionally substituted triazolyl, optionally substituted thiazolyl, optionally substituted tetrazolyl, optionally substituted thiadiazolyl, optionally substituted thienyl, or optionally substituted thiomorpholinyl.

8. The compound of claim 7 wherein X is optionally substituted phenyl, optionally substituted benzo[1,4]oxazin-3-onyl, optionally substituted benzoxazolyl, optionally substituted 2,3-dihydrobenzo[1,4]dioxinyl, optionally substituted 2,3-dihydrobenzo furanyl, optionally substituted pyrazinonyl, optionally substituted pyridazinonyl, optionally substituted pyridinyl, optionally substituted pyridonyl, optionally substituted pyrimidinonyl, optionally substituted pyrimidinyl, optionally substituted pyrrolyl, or optionally substituted thienyl.

9. The compound of claim 8 wherein Z is optionally substituted (C_3-C_6)cycloalkyl, optionally substituted azetidinyl, optionally substituted 1,4-dioxanyl, optionally substituted morpholinyl, optionally substituted phenyl, optionally substituted piperazinyl, optionally substituted piperidinyl, optionally substituted pyrazinyl, optionally substituted pyrazolyl, optionally substituted pyridazinyl, optionally substituted pyridazinonyl, optionally substituted pyridonyl, optionally substituted pyridinyl, optionally substituted pyrimidinyl, optionally substituted pyrimidinonyl, optionally substituted pyrrolidinyl, optionally substituted tetrahydropyranyl, optionally substituted triazolyl, optionally substituted tetrazolyl, optionally substituted thiadiazolyl, optionally substituted thienyl, or optionally substituted thiomorpholinyl.

10. The compound of claim 9 wherein A is



11. The compound of claim 10 wherein m is 1, 2 or 3.

12. The compound of claim 11wherein n is 0 or 1.

13. The compound of claim 12 wherein p is 0.

14. The compound of claim 13 wherein X is optionally substituted phenyl, optionally substituted benzo[1,4]oxazin-3-only, optionally substituted benzoxazolyl, optionally substituted 2,3-dihydrobenzo[1,4]dioxinyl, optionally substituted 2,3-dihydrobenzo furanyl, optionally substituted pyrazinonyl, optionally substituted pyridazinonyl, optionally substituted pyridinyl, optionally substituted pyridonyl, or optionally substituted pyrimidinonyl.

15. The compound of claim 14 wherein X is optionally substituted with one or more substituents independently selected from the group consisting of Br, Cl, F, CF_3, CN, COR^b, COOH, OCF_3, OH, $-N(R^c)(R^d)$, $-NO_2$, optionally substituted (C_1-C_6)alkyl, optionally substituted (C_1-C_6)alkoxy, $-S(O)R^d$, or $-S(O)_2(R^d)$; wherein

R^c is independently H, optionally substituted (C_1-C_3)alkyl or optionally substituted (C_3-C_6)cycloalkyl;

R^d is H, $-C(O)R^b$, optionally substituted (C_1-C_6)alkyl, optionally substituted (C_3-C_6)cycloalkyl, optionally substituted azetidinyl, or optionally substituted pyrrolidinyl.

16. The compound of claim 14 or 15 wherein Y is optionally substituted imidazolyl, optionally substituted pyrazolyl, optionally substituted pyridazinyl, optionally substituted pyridinyl, optionally substituted pyridazinonyl, optionally substituted pyridinyl, optionally substituted pyrimidinyl or optionally substituted triazinyl.

17. The compound of claim 16 wherein Z is optionally substituted (C_3-C_6)cycloalkyl, optionally substituted azetidinyl, optionally substituted 1,4-dioxanyl, optionally substituted morpholinyl, optionally substituted phenyl, optionally substituted piperidinyl, optionally substituted pyrazinonyl, optionally substituted pyrazinyl, optionally substituted pyrazolyl, optionally substituted pyridazinyl, optionally substituted pyridazinonyl, optionally substituted pyridinyl, optionally substituted pyridonyl, optionally substituted pyrimidinonyl, optionally

substituted pyrimidinyl, optionally substituted tetrahydropyranyl, optionally substituted tetrazolyl, optionally substituted thiadiazolyl, optionally substituted thienyl, or optionally substituted thiomorpholinyl.

18. The compound of claim 17 wherein Z is optionally substituted by one or more substituents independently selected from Br, Cl, F, CF_3, CN, $-COR^f$, $-C(O)CH_2N(R^c)(R^e)$, $-CON(R^c)(R^e)$, $COOR^e$, $-N(R^c)(R^e)$, $-N(R^a)C(O)OR^a-$, $-N(R^c)C(O)R^e$, $-N(R^c)C(O)N(R^c)(R^e)$, $-N(R^c)S(O)_2R^e$, $-N(R^c)S(O)_2N(R^c)(R^e)$, $-NO_2$, oxo, -OH, $-S(O)R^e$, $-S(O)_2(R^e)$, optionally substituted (C_1-C_6)alkyl, optionally substituted (C_1-C_6)alkoxy, optionally substituted (C_3-C_6)cycloalkyl, $-(C(R^b)_2)_rR^f$; wherein

R^b is independently H, CF_3, optionally substituted (C_1-C_3)alkyl or optionally substituted (C_3-C_6)cycloalkyl; or

$(R^b)_2$, together with the carbon atom to which it is attached, can form a 3- to 6-membered cycloalkyl

R^e is H, $-C(O)R^c$, optionally substituted (C_1-C_6)alkyl, optionally substituted (C_3-C_6)cycloalkyl, optionally substituted heteroaryl or optionally substituted heterocyclyl; or

R^c and R^e together with the atom to which they are attached can form an optionally substituted heterocycle;

R^f is H, Br, Cl, F, CF_3, CN, OCF_3, $-COR^e$, $-C(O)CH_2N(R^c)(R^e)$, $-CON(R^c)(R^e)$, $COOR^e$, $-N(R^c)(R^e)$, $-N(R^c)C(O)R^e$, $-N(R^c)C(O)N(R^c)(R^e)$, $-N(R^c)S(O)_2R^e$, $-NO_2$, oxo, optionally substituted (C_1-C_6)alkyl, optionally substituted (C_3-C_6)cycloalkyl, OR^c, $-S(R^c)$, $-S(R^c)$, $-S(O)R^c$, $-S(O)R^c$, $-S(O)_2(R^c)$, $-S(O)_2(R^e)$, $-S(O)_2N(R^c)(R^e)$, optionally substituted azetidinyl, optionally substituted 1,4-dioxanyl, optionally substituted imidazolyl, optionally substituted isoxazolyl, optionally substituted morpholinyl, optionally substituted oxazolyl, optionally substituted oxadiazolyl, optionally substituted piperazinyl, optionally substituted piperidinyl, optionally substituted pyrazolyl, optionally substituted pyridinyl, optionally substituted pyrimidinyl, optionally substituted tetrazolyl, optionally substituted thienyl, optionally substituted thiomorpholinyl or optionally substituted triazolyl; and

r is 0, 1 or 2;

provided that r is not 0 when R^f is OCF_3, OR^c, $-S(R^c)$, $-S(R^c)$, $-S(O)R^c$, $-S(O)R^c$, $-S(O)_2(R^c)$, $-S(O)_2(R^e)$, or $-S(O)_2N(R^c)(R^e)$.

19. The compound of claim 18 wherein Z is optionally substituted by one or more substituents independently selected from Br, Cl, F, CF_3, CN, -C(O)OH, $-C(O)OCH_3$, $-C(O)CH_3$, $-C(O)CH_2OC(O)CH_3$, $-C(O)C(CH_3)_3$, $-C(O)C(H)(CH_3)_2$, $-C(O)CH_2CN$, $-C(O)CH_2OH$, $-C(O)CH_2OCH_3$, $-C(O)CH_2CH_2N(CH_3)_2$, $-C(O)C(CH_3)_2OH$, $-C(O)OC(CH_3)_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)CH_2N(CH_3)_2$, $-C(O)CH_2N(H)S(O)_2CH_3$, $-C(O)CH_2N(H)C(O)CH_3$, $-C(O)CH_2N(H)C(O)N(H)CH_3$, $-C(O)CH_2OCH_3$, $-C(O)CH_2S(O)_2CH_3$, $-C(O)NH_2$, $-C(O)N(CH_3)_2$, $-C(O)N(H)CH_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)N(H)CH(CH_3)_2$, $-C(O)N(H)CH_2CH_2OH$, -C(O)N(H)-isoxazolyl, -C(O)N(H)thiazolyl, $-C(O)N(CH_3)CH_2CH_2OH$, -C(O)-cyclopropyl, -C(O)-cyclobutyl, $-OCH_3$, $-OCH_2CH_2OH$, $-OC(CH_3)_3$, $-NH_2$, $-N(H)C(O)CH_3$-, $-N(H)C(O)CH_2OH$, $-N(H)C(O)CH_2CN$, $-N(H)C(O)C(CH_3)_2OH$, $-N(H)C(O)C(H)(OH)CH_3$, $-N(H)C(O)CH_2N(CH_3)_2$, $-N(H)C(O)CH_2N(H)CH_3$, $-N(H)C(O)(OH)CH_3$, $-N(R^e)C(O)N(R^e)(R^e)$, $-N(H)C(O)N(CH_3)_2$, $-N(H)C(O)N(H)CH_3$-$N(H)C(O)N(H)CH_2CH_2OH$, $-N(H)C(O)N(H)C(H)(CH_2OH)_2$, $-N(H)C(O)N(CH_3)CH_2CH_2OH$, $-N(H)C(O)N(CH_3)CH_2C(H)(OH)CH_2OH$, $-N(H)C(O)O(CH_3)_3$, $-N(H)S(O)_2NH_2$, $-N(H)S(O)_2CH_3$, $-N(R^e)S(O)_2N(R^e)(R^e)$, $-NO_2$, oxo, -OH, $-S(O)CH_3$, $-S(O)CH_2CH_3$, $-S(O)CH_2CH_2CH_3$, $-S(O)_2C(H)(CH_3)_2$, $-S(O)_2$cyclopropyl, $-S(O)_2$-optionally substituted imidazolyl, $-S(O)_2$-optionally substituted isoxazolyl, $-CH_2C(O)N(H)CH_3$, $-CH_2OCH_3$, $-CH_2OH$, $-CH_2CH_2OH$, $-CH_2OS(O)_2$-optionally substituted phenyl, $-C(CH_3)OH$, $-C(H)(CH_2)_2OH$, $-CH_2NH_2$, $-CH_2N(CH_3)_2$, $-CH_2N(H)C(O)CH_3$, $-CH_2N(H)C(O)C(H)(OH)CH_3$, $-CH_2N(H)C(O)CH_2CN$, $-CH_2N(H)C(O)CH_2OH$, $-CH_2N(H)C(O)C(CH_3)_2OH$, $-CH_2N(H)C(O)N(H)CH_3$, $-CH_2N(H)C(O)N(CH_3)_2$, $-CH_2S(O)_2CH_3$, optionally substituted (C_1-C_6)alkyl, optionally substituted (C_1-C_6)alkoxy, optionally substituted morpholinyl, optionally substituted piperidinyl, optionally substituted (C_3-C_6)cycloalkyl, and $-(C(R^b)_2)_rR^f$; wherein

R^b is independently H, CF_3, optionally substituted (C_1-C_3)alkyl or optionally substituted (C_3-C_6)cycloalkyl; or

$(R^b)_2$, together with the carbon atom to which it is attached, can form a 3- to 6-membered cycloalkyl

R^e is H, $-C(O)R^c$, optionally substituted (C_1-C_6)alkyl, optionally substituted (C_3-C_6)cycloalkyl, optionally substituted heteroaryl or optionally substituted heterocyclyl; or

R^c and R^e together with the atom to which they are attached can form an optionally substituted heterocycle;

R^f is H, Br, Cl, F, CF_3, CN, OCF_3, $-COR^e$, $-C(O)CH_2N(R^c)(R^e)$, $-CON(R^c)(R^e)$, $COOR^c$, $-N(R^c)(R^e)$, $-N(R^c)C(O)R^e$, $-N(R^c)C(O)N(R^c)(R^e)$, $-N(R^c)S(O)_2R^e$, $-NO_2$, oxo, optionally substituted (C_1-C_6)alkyl, optionally substituted (C_3-C_6)cycloalkyl, OR^c, $-S(R^c)$, $-S(R^c)$, $-S(O)R^c$, $-S(O)R^c$, $-S(O)_2(R^c)$, $-S(O)_2(R^e)$, $-S(O)_2N(R^c)(R^e)$, optionally substituted azetidinyl, optionally

substituted 1,4-dioxanyl, optionally substituted imidazolyl, optionally substituted isoxazolyl, optionally substituted morpholinyl, optionally substituted 1,2,4-oxadiazolyl, optionally substituted oxazolyl, optionally substituted oxadiazolyl, optionally substituted piperazinyl, optionally substituted piperidinyl, optionally substituted pyrazolyl, optionally substituted pyridazinyl, optionally substituted pyridinyl, optionally substituted pyrimidinyl, optionally substituted pyrrolidinyl, optionally substituted tetrazolyl, optionally substituted thienyl, optionally substituted thiomorpholinyl or optionally substituted triazolyl; and

r is 0, 1 or 2;

provided that r is not 0 when R^f is OCF_3, OR^e, $-S(R^e)$, $-S(R^e)$, $-S(O)R^e$, $-S(O)R^e$, $-S(O)_2(R^e)$, $-S(O)_2(R^e)$, or $-S(O)_2N(R^e)(R^e)$.

20. The compound of claim 19 wherein Z is optionally substituted by one or more substituents independently selected from Br, Cl, F, CF_3, CH_3, CN, $-C(O)OH$, $-C(O)OCH_3$, $-C(O)CH_3$, $-C(O)CH_2OC(O)CH_3$, $-C(O)C(CH_3)_3$, $-C(O)C(H)(CH_3)_2$, $-C(O)CH_2CN$, $-C(O)CH_2OH$, $-C(O)CH_2OCH_3$, $-C(O)CH_2CH_2N(CH_3)_2$, $-C(O)C(CH_3)_2OH$, $-C(O)OC(CH_3)_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)CH_2N(CH_3)_2$, $-C(O)CH_2N(H)S(O)_2CH_3$, $-C(O)CH_2N(H)C(O)CH_3$, $-C(O)CH_2N(H)C(O)N(H)CH_3$, $-C(O)CH_2OCH_3$, $-C(O)CH_2S(O)_2CH_3$, $-C(O)NH_2$, $-C(O)N(CH_3)_2$, $-C(O)N(H)CH_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)N(H)CH(CH_3)_2$, $-C(O)N(H)CH_2CH_2OH$, -C(O)N(H)-isoxazolyl, -C(O)N(H)thiazolyl, $-C(O)N(CH_3)CH_2CH_2OH$, -C(O)-cyclopropyl, -C(O)-cyclobutyl, $-OCH_3$, $-OCH_2CH_2OH$, $-OC(CH_3)_3$, $-NH_2$, $-N(H)C(O)CH_3$-, $-N(H)C(O)CH_2OH$, $-N(H)C(O)CH_2CN$, $-N(H)C(O)C(CH_3)_2OH$, $-N(H)C(O)C(H)(OH)CH_3$, $-N(H)C(O)CH_2N(CH_3)_2$, $-N(H)C(O)CH_2N(H)CH_3$, $-N(H)C(O)(OH)CH_3$, $-N(R^e)C(O)N(R^e)(R^e)$, $-N(H)C(O)N(CH_3)_2$, $-N(H)C(O)N(H)CH_3$-$N(H)C(O)N(H)CH_2CH_2OH$, $-N(H)C(O)N(H)C(H)(CH_2OH)_2$, $-N(H)C(O)N(CH_3)CH_2CH_2OH$, $-N(H)C(O)N(CH_3)CH_2C(H)(OH)CH_2OH$, $-N(H)C(O)O(CH_3)_3$, $-N(H)S(O)_2NH_2$, $-N(H)S(O)_2CH_3$, $-N(R^e)S(O)_2N(R^e)(R^e)$, $-NO_2$, 1,2,4-oxadiazolyl, oxo, -OH, $-S(O)CH_3$, $-S(O)CH_2CH_3$, $-S(O)CH_2CH_2CH_3$, $-S(O)_2C(H)(CH_3)_2$, $-S(O)_2$cyclopropyl, $-S(O)_2$-optionally substituted imidazolyl, $-S(O)_2$-optionally substituted isoxazolyl, $-CH_2CN$, $-CH_2C(O)N(H)CH_3$, $-CH_2OCH_3$, $-CH_2OH$, $-CH_2CH_2OH$, $-CH_2OS(O)_2$-optionally substituted phenyl, $-C(CH_3)OH$, $-C(H)(CH_2)_2OH$, $-CH_2NH_2$, $-CH_2N(CH_3)_2$, $-CH_2N(H)C(O)CH_3$, $-CH_2N(H)C(O)C(H)(OH)CH_3$, $-CH_2N(H)C(O)CH_2CN$, $-CH_2N(H)C(O)CH_2OH$, $-CH_2N(H)C(O)C(CH_3)_2OH$, $-CH_2N(H)C(O)N(H)CH_3$, $-CH_2N(H)C(O)N(CH_3)_2$, $-CH_2S(O)_2CH_3$, $-CH_2$-morpholinyl, $-CH_2$-thiomorpholinyl, optionally substituted morpholinyl, optionally substituted oxadiazolyl, optionally substituted oxazolyl, optionally substituted piperidinyl, and optionally substituted (C_3-C_6)cycloalkyl

wherein R^e is H, -C(O)R^c, optionally substituted (C_1-C_6)alkyl, optionally substituted (C_3-C_6)cycloalkyl, optionally substituted azetidinyl, optionally substituted morpholinyl, optionally substituted piperidinyl, or optionally substituted pyrrolidinyl.

21. The compound of claim 20 wherein Y is optionally substituted by one or more substituents independently selected from the group consisting of CF_3, CN, -COH, -COOH, NO_2, -$N(R^a)_2$, -$N(R^a)C(O)N(R^a)_2$, -$C(O)N(R^a)_2$, -$S(O)R^a$, -$S(O)_2R^a$, halogen, optionally substituted (C_1-C_6)alkyl, optionally substituted (C_1-C_6)alkoxy, optionally substituted aryl, optionally substituted heteroaryl, and optionally substituted heterocyclyl.

22. The compound of claim 21 wherein

X is optionally substituted phenyl, optionally substituted benzo[1,4]oxazin-3-onyl, optionally substituted benzoxazolyl, optionally substituted 2,3-dihydrobenzo[1,4]dioxinyl, optionally substituted 2,3-dihydrobenzo furanyl, optionally substituted pyrazinonyl, optionally substituted pyridazinonyl, optionally substituted pyridazinyl, or optionally substituted pyridinyl;

Y is optionally substituted pyrimidinyl; and

Z is optionally substituted cyclohexyl, optionally substituted 1,4-dioxanyl, optionally substituted pyridonyl, or optionally substituted pyridinyl.

23. The compound of claim 1 wherein the compound is

6-(5-(1-Acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-Carbamoylphenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(morpholine-4-carbonyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(2-(2-hydroxypropan-2-yl)pyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(2*H*-tetrazol-5-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide hydrochloride; this is preparation #13 – no A# - should it be included?

6-(5-(1-(ethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclopropanecarbonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-acetamidoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-cyanoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylsulfonyl) acetyl) piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxy ethyl carbamoyl) piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(isoxazol-3-ylcarbamoyl) piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*, 5*R*)-5-(hydroxymethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*, 5*R*)-5-(hydroxymethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(methylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*, 5*R*)-5-(cyanomethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5- ((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(2*S*,5*R*)-5-((*S*)-3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxane-2-carboxylic acid;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(morpholinomethyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*S*)-5-Carbamoyl-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1*H*-imidazol-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-(2-hydroxyethoxy)phenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-(4-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-phenyl-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(4-(hydroxymethyl)phenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-6-(5-(4-methoxyphenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-(4-morpholinophenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-(4-(piperidin-1-yl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(4-(dimethylamino)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(3-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-cyclohexyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-cyclohexyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-(hydroxymethyl)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-carbamoylphenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-cyclohexyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-hydroxycyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

tert-butyl 4-(3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)piperidine-1-carboxylate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(hydroxymethyl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(4-(hydroxymethyl)piperidin-1-yl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidin-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(4-(morpholinomethyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-cyano-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(4- (thiomorpholinomethyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-cyano-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(3-carbamoylphenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(3-(morpholinomethyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(3-(methylcarbamoyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

methyl 4-(3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(4-(methylcarbamoyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(thiophen-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-{5-[4-(1,1,1,-Dioxo-thiomorpholine-4-yl methyl)phenyl]-4,5-dihydro-isoxazol-3-yl}-2-methyl pyrimidine-4-carboxylate;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(2-cyanoacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(sulfamoylamino)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-3-yl)-4,5-dihydro-1H-pyrazol-3-yl)pyrimidine-4-carboxamide;

6-(3-(4-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-5-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(3-phenyl-4,5-dihydroisoxazol-5-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(3-(4-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-5-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(3-phenyl-4,5-dihydroisoxazol-5-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(2-hydroxyethylcarbamoyl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1-acetylpiperidin-4-yl)methyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrimidin-5-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(2-hydroxypropan-2-yl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1-(2-hydroxyacetyl)piperidin-4-yl)methyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1-(methylsulfonyl)piperidin-4-yl)methyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(2-carbamoylpyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1*H*-pyrazol-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((1*r*,4*R*)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*R*)-5-((1*r*,4*R*)-4-(methylsulfonamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((1*r*,4*R*)-4-(2-hydroxyacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrimidin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(tetrahydro-2*H*-pyran-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(methylsulfonamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(2-hydroxyacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrazin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxy-2-methylpropanoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclopropylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(ethylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-isobutyrylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(propylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-methoxybenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-methoxybenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclopropanecarbonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(dimethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(ethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-(dimethylamino)acetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-acetamidoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(isopropylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-cyanoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-methoxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclobutanecarbonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylsulfonamido)acetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(5-cyanopyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(5-(hydroxymethyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(methylcarbamoyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(3-methylureido)acetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(5-(1-hydroxyethyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-((2-hydroxyethyl)(methyl)carbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(isopropylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(1-methyl-1*H*-imidazol-2-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(1-methyl-1*H*-imidazol-4-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-(1,2-dimethyl-1*H*-imidazol-4-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(3,5-dimethylisoxazol-4-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(methylsulfonyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(5-acetylpyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(3-methyl-1,2,4-oxadiazol-5-yl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

((2*S*,5*R*)-5-(3-(6-(3-chloro-4-fluorobenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxan-2-yl)methyl 4-methylbenzenesulfonate;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide ;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

((2*S*,5*R*)-5-((*S*)-3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxan-2-yl)methyl 4-methylbenzenesulfonate;

methyl 4-(3-(2-(3-methoxybenzylcarbamoyl)pyridin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoate;

N-(3-methoxybenzyl)-4-(5-(4-methoxyphenyl)-4,5-dihydroisoxazol-3-yl)picolinamide;

4-(5-(4-chlorophenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)picolinamide;

4-(3-(2-(3-methoxybenzylcarbamoyl)pyridin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoic acid;

4-(5-(4-((dimethylamino)methyl)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)picolinamide;

4-(3-(2-(4-methoxybenzylcarbamoyl)pyridin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoic acid;

4-(5-(4-chlorophenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)picolinamide;

N-(4-methoxybenzyl)-4-(5-(pyrazin-2-yl)-4,5-dihydroisoxazol-3-yl)picolinamide;

N-(4-methoxybenzyl)-4-(5-(4-methoxyphenyl)-4,5-dihydroisoxazol-3-yl)picolinamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-N-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((2S,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((R)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*R*)-5-(methylsulfonamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-(methylsulfonylmethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(thiazol-2-ylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(Acetamidomethyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((1r,4*R*)-4-(acetamidomethyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(oxazol-5-yl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(methylsulfonyl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(3-methyl-1,2,4-oxadiazol-5-yl)pyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(6-acetylpyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)pyridazine-3-carboxamide 2,2,2-trifluoroacetate;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((1*r*,4*R*)-4-((2-hydroxyacetamido)methyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-((2-hydroxyacetamido)methyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-3-(3-methyl-1,2,4-oxadiazol-5-yl)-1*H*-pyrazol-5-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(3-carbamoyl-1-methyl-1*H*-pyrazol-5-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-((2-cyanoacetamido)methyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-*N*-methylpyridazine-3-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(3-methyl-1,2,4-oxadiazol-5-yl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5R)-5-((2-hydroxyacetamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(oxazol-5-yl)pyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-(hydroxymethyl)pyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(3-(2-hydroxyethyl)ureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(3-(2-hydroxyethyl)-3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(methylsulfonyl)pyrazin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(3-(dimethylamino)propanoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

(2*R*,5*S*)-5-((*S*)-3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxane-2-carboxylic acid;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3,3-dimethylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-dichlorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-dichlorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-((2-cyanoacetamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-((2-hydroxy-2-methylpropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*R*)-5-((3-methylureido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-((3,3-dimethylureido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrrolidine-1-carbonyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(2-(dimethylamino)acetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1*H*-pyrazolo[3,4-*b*]pyridin-5-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-1*H*-pyrazolo[3,4- *b*]pyridin-5-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(bromomethyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-*N*-(2-hydroxyethyl)-4,5-dihydroisoxazole-5-carboxamide;

3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazole-5-carboxamide;

methyl 3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazole-5-carboxylate;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5S)-5-(2-hydroxypropan-2-yl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-(methylcarbamoyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(methylcarbamoyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-((1r,4r)-4-carbamoylcyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(2-hydroxy-2-methylpropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((3-(hydroxymethyl)pyrrolidin-1-yl)methyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-((4-acetylpiperazin-1-yl)methyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-(6-(dimethylamino)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(pyrrolidin-1-yl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-((1r,4r)-4-(3-(1,3-dihydroxypropan-2-yl)ureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-((2-hydroxyethyl)carbamoyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3-(2,3-dihydroxypropyl)-3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-Acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(2-(methylamino)acetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(3-methylbenzyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(3-methylbenzyl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(3-chloro-4-fluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(3-chloro-4-fluorobenzyl)-6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzofuran-5-yl)methyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((2-methylbenzo[d]oxazol-5-yl)methyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((2-methylbenzo[*d*]oxazol-5-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(methoxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(methoxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(((*S*)-2-hydroxypropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-(5-((1S,4r)-4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzo[*b*][1,4]dioxin-6-yl)methyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzo[*b*][1,4]dioxin-6-yl)methyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

tert-butyl ((1r,4r)-4-(3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)cyclohexyl)carbamate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-isopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(pyridin-4-ylmethyl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(pyridin-4-ylmethyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(pyridin-4-ylmethyl)pyrimidine-4-carboxamide;

6 6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((1-methyl-2-oxo-1,2-dihydropyridin-4-yl)methyl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-((*R*)-1-(4-fluoro-3-methoxyphenyl)ethyl)-2-methyl-6-(5-phenyl-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-((*S*)-1-(4-fluoro-3-methoxyphenyl)ethyl)-2-methyl-6-(5-phenyl-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((1-methyl-2-oxo-1,2-dihydropyridin-4-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((1-methyl-2-oxo-1,2-dihydropyridin-4-yl)methyl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(quinolin-4-ylmethyl)pyrimidine-4-carboxamide;

(*S*)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((1r,4S)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(quinolin-4-ylmethyl)pyrimidine-4-carboxamide;

6-((R)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(quinolin-4-ylmethyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-((2-hydroxy-2-methylpropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((S)-5-((1r,4S)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluorobenzyl)-6-(5-((1S,4r)-4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-((2-oxo-1,2,3,4,4a,8a-hexahydroquinolin-7-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((2-oxo-1,2,3,4-tetrahydroquinolin-7-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(S)-*N*-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(R)-*N*-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxypyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxypyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

(S)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(R)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(methoxy(methyl)amino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(5-chloro-1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxy-5,6-dihydropyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-(dimethylamino)-2-oxoethyl)-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((1s,4R)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((1r,4R)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((1-methyl-6-oxo-1,6-dihydropyridin-3-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(isopropylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1S,4r)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((3-oxo-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-(5-(3-chloro-5-cyclopropyl-4-oxocyclohexyl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(S)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(R)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(R)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((3-oxo-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-ca)rboxamide;

6-(5-(1-(2-(cyclopropylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-1-methyl-6-oxo-5-(5-phenyl-4,5-dihydroisoxazol-3-yl)-1,6-dihydropyridazine-3-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(1-methoxyethyl)-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide; or

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(isopropylamino)allyl)-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide.

24. The compound according to claim 23 wherein the compound is

N-(4-Fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((1*r*,4*R*)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((1*r*,4*R*)-4-(2-hydroxyacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-(methylsulfonylmethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*, 5*R*)-5-(hydroxymethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*R*)-5-(methylsulfonamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*R*)-5-((1r,4*R*)-4-(acetamidomethyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(methylsulfonyl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(*S*)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

(2*R*,5*S*)-5-((*S*)-3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxane-2-carboxylic acid;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3,3-dimethylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-dichlorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-((2-cyanoacetamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*R*)-5-((3-methylureido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(2*S*,5*R*)-5-((*S*)-3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxane-2-carboxylic acid;

6-((*S*)-5-((2*R*,5*R*)-5-((3,3-dimethylureido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(2-(dimethylamino)acetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5S)-5-(2-hydroxypropan-2-yl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-(methylcarbamoyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(methylcarbamoyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-((1r,4r)-4-carbamoylcyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(2-hydroxy-2-methylpropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3-(1,3-dihydroxypropan-2-yl)ureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(2-cyanoacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*S*)-5-Carbamoyl-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-3-yl)-4,5-dihydro-1H-pyrazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-((2-hydroxyethyl)carbamoyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3-(2,3-dihydroxypropyl)-3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-Acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(2-(methylamino)acetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(3-methylbenzyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(3-methylbenzyl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(3-chloro-4-fluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzofuran-5-yl)methyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((2-methylbenzo[d]oxazol-5-yl)methyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((2-methylbenzo[*d*]oxazol-5-yl)methyl)pyrimidine-4-carboxamide;

6-((S)-5-((1r,4S)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(methoxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(methoxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(((*S*)-2-hydroxypropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-(5-((1S,4r)-4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzo[*b*][1,4]dioxin-6-yl)methyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzo[*b*][1,4]dioxin-6-yl)methyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluorobenzyl)-6-(5-((1S,4r)-4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

(*S*)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((1r,4S)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

(S)-*N*-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(R)-*N*-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxypyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

(S)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(methoxy(methyl)amino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(5-chloro-1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((1s,4R)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((1r,4R)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(isopropylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1S,4r)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-(5-(3-chloro-5-cyclopropyl-4-oxocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(S)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(R)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-(5-(1-(2-(cyclopropylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-1-methyl-6-oxo-5-(5-phenyl-4,5-dihydroisoxazol-3-yl)-1,6-dihydropyridazine-3-carboxamide;

25. A pharmaceutical composition comprising a compound, biologically active metabolite, pro-drug, isomer, stereoisomer, solvate, hydrate or pharmaceutically acceptable salt of Formula (I) and a pharmaceutically acceptable carrier or excipient.

26. A kit comprising a composition of claim 25, packaging and instructions for use.

27. A method of treating a disease or condition comprising administering a therapeutically effective amount of a compound of Formula (I).

28. The method of claim 27 wherein the disease or condition is osteoarthritis, traumatic joint injury (anterior cruciate ligament tear/meniscal tear), rheumatoid arthritis, juvenile arthritis, psoriatic arthritis, gouty arthritis, degenerative joint disease, systemic lupus erythematosus, cardiovascular disorders, such as acute myocardial infarction, acute coronary syndrome, chronic heart failure, myocardial infarction, atherosclerosis, viral myocarditis, cardiac allograft rejection, and sepsis-associated cardiac dysfunction. Furthermore, the compounds of the present invention are also useful for the treatment of central nervous system disorders such as meningococcal meningitis, Alzheimer's disease and Parkinson's disease, the treatment of an ocular condition, a cancer, ankylosing spondylitis, a solid tumor, a sarcoma, fibrosarcoma, osteoma, melanoma, retinoblastoma, a rhabdomyosarcoma, glioblastoma, neuroblastoma, teratocarcinoma, hypersensitivity reactions, hyperkinetic movement disorders, hypersensitivity pneumonitis, hypertension, hypokinetic movement disorders, aordic and peripheral aneuryisms, hypothalamic-pituitary-adrenal axis evaluation, aortic dissection, arterial hypertension, arteriosclerosis,

arteriovenous fistula, ataxia, spinocerebellar degenerations, streptococcal myositis, structural lesions of the cerebellum, subacute sclerosing panencephalitis, Syncope, syphilis of the cardiovascular system, systemic anaphalaxis, systemic inflammatory response syndrome, systemic onset juvenile rheumatoid arthritis, T-cell or FAB ALL, telangiectasia, thromboangitis obliterans, transplants, trauma/hemorrhage, type III hypersensitivity reactions, type IV hypersensitivity, unstable angina, uremia, urosepsis, urticaria, valvular heart diseases, varicose veins, vasculitis, venous diseases, venous thrombosis, ventricular fibrillation, viral and fungal infections, vital encephalitis/aseptic meningitis, vital-associated hemaphagocytic syndrome, Wernicke-Korsakoff syndrome, Wilson's disease, xenograft rejection of any organ or tissue, heart transplant rejection, hemachromatosis, hemodialysis, hemolytic uremic syndrome/thrombolytic thrombocytopenic purpura, hemorrhage, idiopathic pulmonary fibrosis, antibody mediated cytotoxicity, Asthenia, infantile spinal muscular atrophy, inflammation of the aorta, influenza A, ionizing radiation exposure, iridocyclitis/uveitis/optic neuritis, juvenile spinal muscular atrophy, lymphoma, myeloma, leukaemia, malignant ascites, hematopoietic cancers, a diabetic condition such as insulin-dependent diabetes mellitus glaucoma, diabetic retinopathy or microangiopathy, sickle cell anaemia, chronic inflammation, glomerulonephritis, graft rejection, Lyme disease, von Hippel Lindau disease, pemphigoid, Paget's disease, fibrosis, sarcoidosis, cirrhosis, thyroiditis, hyperviscosity syndrome, Osler-Weber-Rendu disease, chronic occlusive pulmonary disease, asthma or edema following burns, trauma, radiation, stroke, hypoxia, ischemia, ovarian hyperstimulation syndrome, post perfusion syndrome, post pump syndrome, post-MI cardiotomy syndrome, preeclampsia, menometrorrhagia, endometriosis, pulmonary hypertension, infantile hemangioma, or infection by Herpes simplex, Herpes Zoster, human immunodeficiency virus, parapoxvirus, protozoa or toxoplasmosis, progressive supranucleo palsy, primary pulmonary hypertension, radiation therapy, Raynaud's phenomenon, Raynaud's disease, Refsum's disease, regular narrow QRS tachycardia, renovascular hypertension, restrictive cardiomyopathy, sarcoma, senile chorea, senile dementia of Lewy body type, shock, skin allograft, skin changes syndrome, ocular or macular edema, ocular neovascular disease, scleritis, radial keratotomy, uveitis, vitritis, myopia, optic pits, chronic retinal detachment, post-laser treatment complications, conjunctivitis, Stargardt's disease, Eales disease, retinopathy, macular degeneration, restenosis, ischemia/reperfusion injury, ischemic stroke, vascular occlusion, carotid obstructive disease, ulcerative colitis, inflammatory bowel disease, diabetes, diabetes mellitus, insulin dependent diabetes mellitus, allergic diseases, dermatitis scleroderma, graft versus host disease, organ transplant rejection, bone marrow transplant rejection, acute or chronic immune disease associated with organ transplantation, sarcoidosis, disseminated intravascular coagulation, Kawasaki's disease, nephrotic syndrome, chronic fatigue syndrome, Wegener's granulomatosis, Henoch-Schoenlein purpurea, microscopic vasculitis of the kidneys, chronic active hepatitis, septic shock, toxic shock syndrome, sepsis syndrome, cachexia, infectious diseases, parasitic diseases, acquired

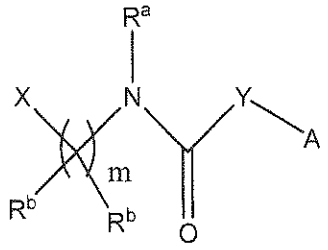
immunodeficiency syndrome, acute transverse myelitis, Huntington's chorea, stroke, primary biliary cirrhosis, hemolytic anemia, malignancies, Addison's disease, idiopathic Addison's disease, sporadic, polyglandular deficiency type I and polyglandular deficiency type II, Schmidt's syndrome, adult respiratory distress syndrome, alopecia, alopecia areata, seronegative arthopathy, arthropathy, Reiter's disease, psoriatic arthropathy, ulcerative colitic arthropathy, enteropathic synovitis, chlamydia, yersinia and salmonella associated arthropathy, atheromatous disease/arteriosclerosis, atopic allergy, autoimmune bullous disease, pemphigus vulgaris, pemphigus foliaceus, pemphigoid, linear IgA disease, autoimmune haemolytic anaemia, Coombs positive haemolytic anaemia, acquired pernicious anaemia, juvenile pernicious anaemia, peripheral vascular disorders, peritonitis, pernicious anemia, myalgic encephalitis/Royal Free Disease, chronic mucocutaneous candidiasis, giant cell arteritis, primary sclerosing hepatitis, cryptogenic autoimmune hepatitis, Acquired Immunodeficiency Disease Syndrome, Acquired Immunodeficiency related diseases, Hepatitis A, Hepatitis B, Hepatitis C, His bundle arrythmias, HIV infection/HIV neuropathy, common varied immunodeficiency, common variable hypogammaglobulinaemia, dilated cardiomyopathy, female infertility, ovarian failure, premature ovarian failure, fibrotic lung disease, chronic wound healing, cryptogenic fibrosing alveolitis, post-inflammatory interstitial lung disease, interstitial pneumonitis, pneumocystis carinii pneumonia, pneumonia, connective tissue disease associated interstitial lung disease, mixed connective tissue disease, associated lung disease, systemic sclerosis associated interstitial lung disease, rheumatoid arthritis associated interstitial lung disease, systemic lupus erythematosus associated lung disease, dermatomyositis/polymyositis associated lung disease, Sjögren's disease associated lung disease, ankylosing spondylitis associated lung disease, vasculitic diffuse lung disease, haemosiderosis associated lung disease, drug-induced interstitial lung disease, radiation fibrosis, bronchiolitis obliterans, chronic eosinophilic pneumonia, lymphocytic infiltrative lung disease, postinfectious interstitial lung disease, gouty arthritis, autoimmune hepatitis, type-1 autoimmune hepatitis (classical autoimmune or lupoid hepatitis), type-2 autoimmune hepatitis (anti-LKM antibody hepatitis), autoimmune mediated hypoglycaemia, type B insulin resistance with acanthosis nigricans, hypoparathyroidism, acute immune disease associated with organ transplantation, chronic immune disease associated with organ transplantation, osteoarthritis, primary sclerosing cholangitis, psoriasis type 1, psoriasis type 2, idiopathic leucopaenia, autoimmune neutropaenia, renal disease NOS, glomerulonephritides, microscopic vasulitis of the kidneys, Lyme disease, discoid lupus erythematosus, male infertility idiopathic or NOS, sperm autoimmunity, multiple sclerosis (all subtypes), sympathetic ophthalmia, pulmonary hypertension secondary to connective tissue disease, acute pain, chronic pain, Goodpasture's syndrome, pulmonary manifestation of polyarteritis nodosa, acute rheumatic fever, rheumatoid spondylitis, Still's disease, systemic sclerosis, Sjögren's syndrome, Takayasu's disease/arteritis, autoimmune thrombocytopaenia, toxicity, transplants, diabetic retinopathy, retinopathy of prematurity,

choroidal neovascularization due to age-related macular degeneration, infantile hemangiomas, ascites, effusions, exudates, macular edema, cerebral edema, acute lung injury, adult respiratory distress syndrome, proliferative disorders, restenosis, fibrotic disorders, hepatic cirrhosis, atherosclerosis, mesangial cell proliferative disorders, diabetic nephropathy, malignant nephrosclerosis, thrombotic microangiopathy syndromes, glomerulopathies, myocardial angiogenesis, coronary and cerebral collaterals, ischemic limb angiogenesis, ischemia/reperfusion injury, peptic ulcer Helicobacter related diseases, virally-induced angiogenic disorders, preeclampsia, menometrorrhagia, cat scratch fever, rubeosis, neovascular glaucoma, diabetic retinopathy, retinopathy of prematurity, age-related macular degeneration, hyperproliferative disorder, thyroid hyperplasia, Grave's disease, cysts, hypervascularity of ovarian stroma characteristic of polycystic ovarian syndrome, Stein-Leventhal syndrome, or polycystic kidney.

29. The method of claim 28 wherein the disease or condition is osteoarthritis, rheumatoid arthritis, juvenile arthritis, psoriatic arthritis, degenerative joint disease or systemic lupus erythematosus.

CLAIMS

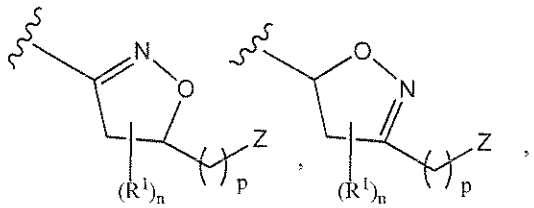
1. A compound of Formula (I)



Formula (I)

stereoisomers, and pharmaceutically acceptable salts thereof wherein

A is



X is phenyl, benzo[1,4]oxazin-3-onyl, benzoxazolyl, 2,3-dihydrobenzo[1,4]dioxinyl, 2,3-dihydrobenzo furanyl, pyrazinonyl, pyridinyl, pyridonyl, or quinolinyl, each of which is optionally substituted with one or more substituents independently selected from Br, Cl, F, CF_3, CN, COOH, OCF_3, OH, $(C_1$-$C_6)$alkyl, or $(C_1$-$C_6)$alkoxy;

Y is pyridazinonyl, pyridinyl, or pyrimidinyl, each of which is optionally substituted by one or more substituents independently selected from CF_3, halogen, $(C_1$-$C_6)$alkyl, $(C_1$-$C_6)$alkoxy,

Z is $-C(O)NH_2$, CN, -C(O)-morpholinyl, -C(O)-pyrrolidinyl, $-C(O)-N(H)CH_2CH_2OH$, $(C_3$-$C_6)$cycloalkyl, 1,4-dioxanyl, imidazolyl, morpholinyl, phenyl, piperazinyl, piperidinyl, pyrazinyl, pyrazolyl, pyrazolo[3,4-*b*]pyridinyl, pyridazinyl, pyridazinonyl, pyridinyl, pyridonyl, pyrimidinyl, pyrrolidinyl, tetrahydropyranyl, tetrazolyl, or thienyl, each of which is optionally substituted by one or more substituents independently selected from Br, Cl, F, CF_3, CN, -C(O)OH, $-C(O)OCH_3$, $-C(O)CH_3$, $-C(O)CH_2OC(O)CH_3$, $-C(O)C(CH_3)_3$, $-C(O)C(H)(CH_3)_2$, $-C(O)CH_2CN$, $-C(O)CH_2OH$, $-C(O)CH_2OCH_3$, $-C(O)CH_2CH_2N(CH_3)_2$, $-C(O)C(CH_3)_2OH$, $-C(O)OC(CH_3)_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)CH_2N(CH_3)_2$, $-C(O)CH_2N(H)S(O)_2CH_3$, $-C(O)CH_2N(H)C(O)CH_3$, $-C(O)CH_2N(H)C(O)N(H)CH_3$, $-C(O)CH_2OCH_3$, $-C(O)CH_2S(O)_2CH_3$, $-C(O)NH_2$, $-C(O)N(CH_3)_2$, $-C(O)N(H)CH_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)N(H)CH(CH_3)_2$, $-C(O)N(H)CH_2CH_2OH$, -C(O)N(H)-isoxazolyl, -C(O)N(H)thiazolyl, $-C(O)N(CH_3)CH_2CH_2OH$, -C(O)-cyclopropyl, -C(O)-cyclobutyl, $-OCH_2CH_2OH$, $-OC(CH_3)_3$, $-NH_2$, $-N(CH_3)_2$, $-N(H)C(O)CH_3$-, $-N(H)C(O)CH_2OH$, $-N(H)C(O)CH_2CN$, $-N(H)C(O)C(CH_3)_2OH$, $-N(H)C(O)C(H)(OH)CH_3$, $-N(H)C(O)CH_2N(CH_3)_2$, $-N(H)C(O)CH_2N(H)CH_3$, -

$N(H)C(O)(OH)CH_3$, $-N(R^c)C(O)N(R^c)(R^e)$, $-N(H)C(O)N(CH_3)_2$, $-N(H)C(O)N(H)CH_3$-$N(H)C(O)N(H)CH_2CH_2OH$, $-N(H)C(O)N(H)C(H)(CH_2OH)_2$, $-N(H)C(O)N(CH_3)CH_2CH_2OH$, -$N(H)C(O)N(CH_3)CH_2C(H)(OH)CH_2OH$, $-N(H)C(O)O(CH_3)_3$, $-N(H)S(O)_2NH_2$, - $N(H)S(O)_2CH_3$, $-N(R^c)S(O)_2N(R^c)(R^e)$, $-NO_2$, 1,2,4-oxadiazolyl, oxo, -OH, $-S(O)_2(R^e)$, $-S(O)CH_3$, -$S(O)CH_2CH_3$, $-S(O)CH_2CH_2CH_3$, $-S(O)_2C(H)(CH_3)_2$, $-S(O)_2$cyclopropyl, $-S(O)_2$-imidazolyl, -$S(O)_2$-isoxazolyl, $-CH_2CN$, $-CH_2C(O)N(H)CH_3$, $-CH_2C(O)N(H)$-cyclopropyl, $-CH_2OCH_3$, -CH_2OH, $-CH_2CH_2OH$, $-CH_2OS(O)_2$-phenyl, $-C(CH_3)OH$, $-C(H)(CH_2)_2OH$, $-CH_2NH_2$, -$CH_2N(CH_3)_2$, $-CH_2N(H)C(O)CH_3$, $-CH_2N(H)C(O)C(H)(OH)CH_3$, $-CH_2N(H)C(O)CH_2CN$, -$CH_2N(H)C(O)CH_2OH$, $-CH_2N(H)C(O)C(CH_3)_2OH$, $-CH_2N(H)C(O)N(H)CH_3$, -$CH_2N(H)C(O)N(CH_3)_2$, $-CH_2S(O)_2CH_3$, $-CH_2$-morpholinyl, $-CH_2$-thiomorpholinyl, $-CH_2$-thiomorpholinyl 1,1-dioxide, morpholinyl, oxadiazolyl, oxazolyl, piperidinyl, $(C_1$-$C_6)$alkyl, $(C_1$-$C_6)$alkoxy, or $(C_3$-$C_6)$cycloalkyl;

R^c is independently H, $(C_1$-$C_3)$alkyl or $(C_3$-$C_6)$cycloalkyl;

R^e is H, $-C(O)R^c$, $(C_1$-$C_6)$alkyl, $(C_1$-$C_6)$hydroxyalkyl, $(C_3$-$C_6)$cycloalkyl, heteroaryl optionally substituted with $(C_1$-$C_3)$alkyl, or heterocyclyl; or

R^a is independently H, $(C_1$-$C_3)$alkyl or $(C_3$-$C_6)$cycloalkyl,

R^b is independently H, CF_3, $(C_1$-$C_3)$alkyl or $(C_3$-$C_6)$cycloalkyl;

R^1 is independently Br, Cl, F, $(C_1$-$C_3)$alkyl or $(C_3$-$C_6)$cycloalkyl;

m is 0, 1 or 2;

n is 0; and

p is 0 or 1.

2. The compound of claim 1, wherein p is 0.

3. The compound of claim 2, wherein X is phenyl, benzo[1,4]oxazin-3-onyl, benzoxazolyl, 2,3-dihydrobenzo[1,4]dioxinyl, 2,3-dihydrobenzo furanyl, pyrazinonyl, pyridinyl, or pyridonyl, each of which is optionally substituted with one or more substituents independently selected from Br, Cl, F, CF_3, OCF_3, $(C_1$-$C_6)$alkyl, $(C_1$-$C_6)$alkoxy.

4. The compound of claim 3, wherein Y is pyrimidinyl optionally substituted by one or more substituents independently selected from CF_3, or $(C_1$-$C_6)$alkyl.

5. The compound of claim 4, wherein Z is $(C_3$-$C_6)$cycloalkyl, 1,4-dioxanyl, morpholinyl, phenyl, piperidinyl, pyrazolyl, pyridazinyl, pyridazinonyl, pyridinyl, pyridonyl, pyrimidinyl, tetrahydropyranyl, tetrazolyl, or thienyl, each of which is optionally substituted by one or more substituents independently selected from Br, Cl, F, CF_3, CH_3, CN, -C(O)OH, -$C(O)OCH_3$, $-C(O)CH_3$, $-C(O)CH_2OC(O)CH_3$, $-C(O)C(CH_3)_3$, $-C(O)C(H)(CH_3)_2$, $-C(O)CH_2CN$, -

$C(O)CH_2OH$, $-C(O)CH_2OCH_3$, $-C(O)CH_2CH_2N(CH_3)_2$, $-C(O)C(CH_3)_2OH$, $-C(O)OC(CH_3)_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)CH_2N(CH_3)_2$, $-C(O)CH_2N(H)S(O)_2CH_3$, $-C(O)CH_2N(H)C(O)CH_3$, $-C(O)CH_2N(H)C(O)N(H)CH_3$, $-C(O)CH_2S(O)_2CH_3$, $-C(O)NH_2$, $-C(O)N(CH_3)_2$, $-C(O)N(H)CH_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)N(H)CH(CH_3)_2$, $-C(O)N(H)CH_2CH_2OH$, -C(O)N(H)-isoxazolyl, -C(O)N(H)thiazolyl, $-C(O)N(CH_3)CH_2CH_2OH$, -C(O)-cyclopropyl, -C(O)-cyclobutyl, $-OCH_3$, $-OCH_2CH_2OH$, $-OC(CH_3)_3$, $-NH_2$, $-N(CH_3)_2$, $-N(H)C(O)CH_3$-, $-N(H)C(O)CH_2OH$, $-N(H)C(O)CH_2CN$, $-N(H)C(O)C(CH_3)_2OH$, $-N(H)C(O)C(H)(OH)CH_3$, $-N(H)C(O)CH_2N(CH_3)_2$, $-N(H)C(O)CH_2N(H)CH_3$, $-N(H)C(O)(OH)CH_3$, $-N(H)C(O)N(CH_3)_2$, $-N(H)C(O)N(H)CH_3$-$N(H)C(O)N(H)CH_2CH_2OH$, $-N(H)C(O)N(H)C(H)(CH_2OH)_2$, $-N(H)C(O)N(CH_3)CH_2CH_2OH$, $-N(H)C(O)N(CH_3)CH_2C(H)(OH)CH_2OH$, $-N(H)C(O)O(CH_3)_3$, $-N(H)S(O)_2NH_2$, $-N(H)S(O)_2CH_3$, $-NO_2$, 1,2,4-oxadiazolyl, oxo, -OH, $-S(O)CH_3$, $-S(O)CH_2CH_3$, $-S(O)CH_2CH_2CH_3$, $-S(O)_2CH_3$, $-S(O)_2C(H)(CH_3)_2$, $-S(O)_2$cyclopropyl, $-S(O)_2$-imidazolyl, $-S(O)_2$-isoxazolyl, $-CH_2CN$, $-CH_2C(O)N(H)CH_3$, $-CH_2C(O)N(H)$-cyclopropyl, $-CH_2OCH_3$, $-CH_2OH$, $-CH_2CH_2OH$, $-CH_2OS(O)_2$-phenyl, $-C(CH_3)_2OH$, $-C(H)(CH_2)_2OH$, $-CH_2NH_2$, $-CH_2N(CH_3)_2$, $-CH_2N(H)C(O)CH_3$, $-CH_2N(H)S(O)_2CH_3$, $-CH_2N(H)C(O)C(H)(OH)CH_3$, $-CH_2N(H)C(O)CH_2CN$, $-CH_2N(H)C(O)CH_2OH$, $-CH_2N(H)C(O)C(CH_3)_2OH$, $-CH_2N(H)C(O)N(H)CH_3$, $-CH_2N(H)C(O)N(CH_3)_2$, $-CH_2S(O)_2CH_3$, $-CH_2$-morpholinyl, $-CH_2$-thiomorpholinyl, $-CH_2$-thiomorpholinyl 1,1-dioxide, morpholinyl, oxadiazolyl, oxazolyl, piperidinyl, or (C_3-C_6)cycloalkyl.

6. The compound of claim 5, wherein

X is phenyl, benzo[1,4]oxazin-3-onyl, benzoxazolyl, 2,3-dihydrobenzo[1,4]dioxinyl, 2,3-dihydrobenzo furanyl, pyrazinonyl, or pyridinyl, each of which is optionally substituted with one or more substituents independently selected from Br, Cl, F, CF_3, OCF_3, (C_1-C_6)alkyl, (C_1-C_6)alkoxy;

Y is 2-methyl pyrimidinyl; and

Z is cyclohexyl, 1,4-dioxanyl, phenyl, pyridonyl, piperidinyl, or pyridinyl, each of which is optionally substituted by one or more substituents independently selected from Br, Cl, F, CF_3, CH_3, CN, -C(O)OH, $-C(O)OCH_3$, $-C(O)CH_3$, $-C(O)CH_2OC(O)CH_3$, $-C(O)C(CH_3)_3$, $-C(O)C(H)(CH_3)_2$, $-C(O)CH_2CN$, $-C(O)CH_2OH$, $-C(O)CH_2OCH_3$, $-C(O)CH_2CH_2N(CH_3)_2$, $-C(O)C(CH_3)_2OH$, $-C(O)OC(CH_3)_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)CH_2N(CH_3)_2$, $-C(O)CH_2N(H)S(O)_2CH_3$, $-C(O)CH_2N(H)C(O)CH_3$, $-C(O)CH_2N(H)C(O)N(H)CH_3$, $-C(O)CH_2S(O)_2CH_3$, $-C(O)NH_2$, $-C(O)N(CH_3)_2$, $-C(O)N(H)CH_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)N(H)CH(CH_3)_2$, $-C(O)N(H)CH_2CH_2OH$, -C(O)N(H)-isoxazolyl, -C(O)N(H)thiazolyl, $-C(O)N(CH_3)CH_2CH_2OH$, -C(O)-cyclopropyl, -C(O)-cyclobutyl, $-OCH_3$, $-OCH_2CH_2OH$, $-OC(CH_3)_3$, $-NH_2$, $-N(CH_3)_2$, $-N(H)C(O)CH_3$-, $-N(H)C(O)CH_2OH$, $-N(H)C(O)CH_2CN$, -

$N(H)C(O)C(CH_3)_2OH$, $-N(H)C(O)C(H)(OH)CH_3$, $-N(H)C(O)CH_2N(CH_3)_2$, $-N(H)C(O)CH_2N(H)CH_3$, $-N(H)C(O)(OH)CH_3$, $-N(H)C(O)N(CH_3)_2$, $-N(H)C(O)N(H)CH_3$-$N(H)C(O)N(H)CH_2CH_2OH$, $-N(H)C(O)N(H)C(H)(CH_2OH)_2$, $-N(H)C(O)N(CH_3)CH_2CH_2OH$, $-N(H)C(O)N(CH_3)CH_2C(H)(OH)CH_2OH$, $-N(H)C(O)O(CH_3)_3$, $-N(H)S(O)_2NH_2$, $-N(H)S(O)_2CH_3$, $-NO_2$, 1,2,4-oxadiazolyl, oxo, $-OH$, $-S(O)CH_3$, $-S(O)CH_2CH_3$, $-S(O)CH_2CH_2CH_3$, $-S(O)_2CH_3$, $-S(O)_2C(H)(CH_3)_2$, $-S(O)_2$cyclopropyl, $-S(O)_2$-imidazolyl, $-S(O)_2$-isoxazolyl, $-CH_2CN$, $-CH_2C(O)N(H)CH_3$, $-CH_2C(O)N(H)$-cyclopropyl, $-CH_2OCH_3$, $-CH_2OH$, $-CH_2CH_2OH$, $-CH_2OS(O)_2$-phenyl, $-C(CH_3)_2OH$, $-C(H)(CH_2)_2OH$, $-CH_2NH_2$, $-CH_2N(CH_3)_2$, $-CH_2N(H)C(O)CH_3$, $-CH_2N(H)S(O)_2CH_3$, $-CH_2N(H)C(O)C(H)(OH)CH_3$, $-CH_2N(H)C(O)CH_2CN$, $-CH_2N(H)C(O)CH_2OH$, $-CH_2N(H)C(O)C(CH_3)_2OH$, $-CH_2N(H)C(O)N(H)CH_3$, $-CH_2N(H)C(O)N(CH_3)_2$, $-CH_2S(O)_2CH_3$, $-CH_2$-morpholinyl, $-CH_2$-thiomorpholinyl, $-CH_2$-thiomorpholinyl 1,1-dioxide, morpholinyl, oxadiazolyl, oxazolyl, piperidinyl, or (C_3-C_6)cycloalkyl.

7. A compound selected from the group consisting of:

6-(5-(1-Acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-Carbamoylphenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(morpholine-4-carbonyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(2-(2-hydroxypropan-2-yl)pyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(2*H*-tetrazol-5-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-

methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide hydrochloride;

6-(5-(1-(ethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclopropanecarbonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-acetamidoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-cyanoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylsulfonyl) acetyl) piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxy ethyl carbamoyl) piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(isoxazol-3-ylcarbamoyl) piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*, 5*R*)-5-(hydroxymethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*, 5*R*)-5-(hydroxymethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(methylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*, 5*R*)-5-(cyanomethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5- ((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(2*S*,5*R*)-5-((*S*)-3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxane-2-carboxylic acid;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(morpholinomethyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*S*)-5-Carbamoyl-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1*H*-imidazol-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-(2-hydroxyethoxy)phenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-(4-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-phenyl-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(4-(hydroxymethyl)phenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-6-(5-(4-methoxyphenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-(4-morpholinophenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-(4-(piperidin-1-yl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(4-(dimethylamino)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(3-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-cyclohexyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-cyclohexyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-(hydroxymethyl)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(4-carbamoylphenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-cyclohexyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-hydroxycyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

tert-butyl 4-(3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)piperidine-1-carboxylate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-3-yl)-4,5-dihydroisoxazol-3-

yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(hydroxymethyl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(4-(hydroxymethyl)piperidin-1-yl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidin-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(4-(morpholinomethyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-cyano-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(4- (thiomorpholinomethyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-cyano-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(3-carbamoylphenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(3-(morpholinomethyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(3-(methylcarbamoyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

methyl 4-(3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(4-(methylcarbamoyl)phenyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(thiophen-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-{5-[4-(1,1,1,-Dioxo-thiomorpholine-4-yl methyl)phenyl]-4,5-dihydro-isoxazol-3-yl}-2-methyl pyrimidine-4-carboxylate;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(2-cyanoacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(sulfamoylamino)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-3-yl)-4,5-dihydro-1H-pyrazol-3-yl)pyrimidine-4-carboxamide;

6-(3-(4-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-5-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(3-phenyl-4,5-dihydroisoxazol-5-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(3-(4-(2-hydroxyethoxy)phenyl)-4,5-dihydroisoxazol-5-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-methoxybenzyl)-2-methyl-6-(3-phenyl-4,5-dihydroisoxazol-5-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(2-hydroxyethylcarbamoyl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1-acetylpiperidin-4-yl)methyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrimidin-5-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(4-(2-hydroxypropan-2-yl)phenyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1-(2-hydroxyacetyl)piperidin-4-yl)methyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1-(methylsulfonyl)piperidin-4-yl)methyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(2-carbamoylpyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1*H*-pyrazol-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((1*r*,4*R*)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*R*)-5-((1*r*,4*R*)-4-(methylsulfonamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((1*r*,4*R*)-4-(2-hydroxyacetamido)cyclohexyl)-

4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrimidin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(tetrahydro-2*H*-pyran-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(methylsulfonamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-Fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(2-hydroxyacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrazin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxy-2-methylpropanoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclopropylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(ethylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-isobutyrylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(propylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-methoxybenzyl)-2-methyl-6-(5-(pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-acetylpiperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-methoxybenzyl)-2-methyl-6-(5-(1-(methylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-(2-hydroxyacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclopropanecarbonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(dimethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(ethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-(dimethylamino)acetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-acetamidoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(isopropylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-cyanoacetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-methoxyacetyl)piperidin-4-yl)-4,5-

dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(cyclobutanecarbonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylsulfonamido)acetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(5-cyanopyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(5-(hydroxymethyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(methylcarbamoyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyethylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(3-methylureido)acetyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(5-(1-hydroxyethyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-((2-hydroxyethyl)(methyl)carbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(isopropylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(1-methyl-1*H*-imidazol-2-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(1-methyl-1*H*-imidazol-4-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-(1,2-dimethyl-1*H*-imidazol-4-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(3,5-dimethylisoxazol-4-ylsulfonyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(methylsulfonyl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(5-acetylpyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(3-methyl-1,2,4-oxadiazol-5-yl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

((2*S*,5*R*)-5-(3-(6-(3-chloro-4-fluorobenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxan-2-yl)methyl 4-methylbenzenesulfonate;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide ;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

((2*S*,5*R*)-5-((*S*)-3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxan-2-yl)methyl 4-methylbenzenesulfonate;

methyl 4-(3-(2-(3-methoxybenzylcarbamoyl)pyridin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoate;

N-(3-methoxybenzyl)-4-(5-(4-methoxyphenyl)-4,5-dihydroisoxazol-3-yl)picolinamide;

4-(5-(4-chlorophenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)picolinamide;

4-(3-(2-(3-methoxybenzylcarbamoyl)pyridin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoic acid;

4-(5-(4-((dimethylamino)methyl)phenyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)picolinamide;

4-(3-(2-(4-methoxybenzylcarbamoyl)pyridin-4-yl)-4,5-dihydroisoxazol-5-yl)benzoic acid;

4-(5-(4-chlorophenyl)-5-methyl-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)picolinamide;

N-(4-methoxybenzyl)-4-(5-(pyrazin-2-yl)-4,5-dihydroisoxazol-3-yl)picolinamide;

N-(4-methoxybenzyl)-4-(5-(4-methoxyphenyl)-4,5-dihydroisoxazol-3-yl)picolinamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-N-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*R*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((2S,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((R)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*R*)-5-(methylsulfonamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-(methylsulfonylmethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(thiazol-2-ylcarbamoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(Acetamidomethyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((1r,4*R*)-4-(acetamidomethyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(oxazol-5-yl)pyridin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(methylsulfonyl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(3-methyl-1,2,4-oxadiazol-5-yl)pyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(6-acetylpyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(3-(6-(4-fluoro-3-methoxybenzylcarbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)pyridazine-3-carboxamide 2,2,2-trifluoroacetate;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((1*r*,4*R*)-4-((2-hydroxyacetamido)methyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-((2-hydroxyacetamido)methyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-3-(3-methyl-1,2,4-oxadiazol-5-yl)-1*H*-pyrazol-5-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(3-carbamoyl-1-methyl-1*H*-pyrazol-5-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-((2-cyanoacetamido)methyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-*N*-methylpyridazine-3-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(3-methyl-1,2,4-oxadiazol-5-yl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5R)-5-((2-hydroxyacetamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(oxazol-5-yl)pyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-(hydroxymethyl)pyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(3-(2-hydroxyethyl)ureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(3-(2-hydroxyethyl)-3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(5-(methylsulfonyl)pyrazin-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-

methylpyrimidine-4-carboxamide;

6-(5-(1-(3-(dimethylamino)propanoyl)piperidin-4-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

(2*R*,5*S*)-5-((*S*)-3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxane-2-carboxylic acid;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3,3-dimethylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-dichlorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-dichlorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-((2-cyanoacetamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-((2-hydroxy-2-methylpropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*R*)-5-((3-methylureido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-((3,3-dimethylureido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyrrolidine-1-carbonyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(2-(dimethylamino)acetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1*H*-pyrazolo[3,4-*b*]pyridin-5-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-1*H*-pyrazolo[3,4- *b*]pyridin-5-

yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(bromomethyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-*N*-(2-hydroxyethyl)-4,5-dihydroisoxazole-5-carboxamide;

*3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-*dihydroisoxazole-5-carboxamide;

methyl 3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazole-5-carboxylate;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*S*)-5-(2-hydroxypropan-2-yl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-(methylcarbamoyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(methylcarbamoyl)cyclohexyl)-4,5-*dihydroisoxazol-3-yl)pyrimidine-4-carboxamide* 2,2,2-*trifluoroacetate;*

6-(5-((1r,4r)-4-carbamoylcyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(2-hydroxy-2-methylpropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((3-(hydroxymethyl)pyrrolidin-1-yl)methyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-((4-acetylpiperazin-1-yl)methyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-(6-(dimethylamino)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(pyrrolidin-1-yl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-((1r,4r)-4-(3-(1,3-dihydroxypropan-2-yl)ureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

*N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-((2-hydroxyethyl)carbamoyl)cyclohexyl)-*4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3-(2,3-dihydroxypropyl)-3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-Acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-

methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(2-(methylamino)acetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(3-methylbenzyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(3-methylbenzyl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(3-chloro-4-fluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(3-chloro-4-fluorobenzyl)-6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzofuran-5-yl)methyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((2-methylbenzo[d]oxazol-5-yl)methyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((2-methylbenzo[*d*]oxazol-5-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(methoxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(methoxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(((*S*)-2-hydroxypropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-(5-((1S,4r)-4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzo[*b*][1,4]dioxin-6-yl)methyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzo[*b*][1,4]dioxin-6-yl)methyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

tert-butyl ((1r,4r)-4-(3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)cyclohexyl)carbamate;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-isopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(pyridin-4-ylmethyl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(pyridin-4-ylmethyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(pyridin-4-ylmethyl)pyrimidine-4-carboxamide;

6 6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((1-methyl-2-oxo-1,2-dihydropyridin-4-yl)methyl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-((*R*)-1-(4-fluoro-3-methoxyphenyl)ethyl)-2-methyl-6-(5-phenyl-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-((*S*)-1-(4-fluoro-3-methoxyphenyl)ethyl)-2-methyl-6-(5-phenyl-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((1-methyl-2-oxo-1,2-dihydropyridin-4-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((1-methyl-2-oxo-1,2-dihydropyridin-4-yl)methyl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(quinolin-4-ylmethyl)pyrimidine-4-carboxamide;

(*S*)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((1r,4S)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-

methyl-*N*-(quinolin-4-ylmethyl)pyrimidine-4-carboxamide;

6-((R)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(quinolin-4-ylmethyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-((2-hydroxy-2-methylpropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((S)-5-((1r,4S)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluorobenzyl)-6-(5-((1S,4r)-4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-((2-oxo-1,2,3,4,4a,8a-hexahydroquinolin-7-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((2-oxo-1,2,3,4-tetrahydroquinolin-7-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-2-oxo-

1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(S)-*N*-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(R)-*N*-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxypyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxypyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

(S)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(R)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(methoxy(methyl)amino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(5-chloro-1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxy-5,6-dihydropyridin-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-(2-(dimethylamino)-2-oxoethyl)-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((1s,4R)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((1r,4R)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((1-methyl-6-oxo-1,6-dihydropyridin-3-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(isopropylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1S,4r)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((3-oxo-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-(5-(3-chloro-5-cyclopropyl-4-oxocyclohexyl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(S)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(R)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(R)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((3-oxo-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-ca)rboxamide;

6-(5-(1-(2-(cyclopropylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-1-methyl-6-oxo-5-(5-phenyl-4,5-dihydroisoxazol-3-yl)-1,6-dihydropyridazine-3-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(1-methoxyethyl)-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide; and

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(isopropylamino)allyl)-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide; or

a pharmaceutically acceptable salt thereof.

8. The compound according to claim 7, wherein the compound is

N-(4-Fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydro-isoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*S*,5*S*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*R*)-5-((1*r*,4*R*)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((1*r*,4*R*)-4-(2-hydroxyacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-(methylsulfonylmethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*, 5*R*)-5-(hydroxymethyl)-1, 4-dioxan-2-yl)-4, 5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*R*)-5-(methylsulfonamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*R*)-5-((1r,4*R*)-4-(acetamidomethyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(6-(methylsulfonyl)pyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(*S*)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

(2*R*,5*S*)-5-((*S*)-3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxane-2-carboxylic acid;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3,3-dimethylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3,4-dichlorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-((2-cyanoacetamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*R*)-5-((3-methylureido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(2*S*,5*R*)-5-((*S*)-3-(6-((4-fluoro-3-methoxybenzyl)carbamoyl)-2-methylpyrimidin-4-yl)-4,5-dihydroisoxazol-5-yl)-1,4-dioxane-2-carboxylic acid;

6-((*S*)-5-((2*R*,5*R*)-5-((3,3-dimethylureido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(2-(dimethylamino)acetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5S)-5-(2-hydroxypropan-2-yl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-(methylcarbamoyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(methylcarbamoyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-(5-((1r,4r)-4-carbamoylcyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-(2-hydroxy-2-methylpropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3-(1,3-dihydroxypropan-2-yl)ureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(2-cyanoacetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*S*)-5-Carbamoyl-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(pyridin-3-yl)-4,5-dihydro-1H-pyrazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-((1r,4r)-4-((2-hydroxyethyl)carbamoyl)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-(3-(2,3-dihydroxypropyl)-3-methylureido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-Acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-((1r,4r)-4-(2-(methylamino)acetamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(acetamidomethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-2-methyl-6-((*S*)-5-((2*R*,5*S*)-5-((methylsulfonyl)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(3-methylbenzyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-(3-methylbenzyl)pyrimidine-4-carboxamide;

6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-hydroxyethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

N-(3-chloro-4-fluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-(5-(6-cyanopyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3-chloro-4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluorobenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzofuran-5-yl)methyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-

dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-fluorobenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-fluorobenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((2-methylbenzo[d]oxazol-5-yl)methyl)pyrimidine-4-carboxamide;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((2-methylbenzo[*d*]oxazol-5-yl)methyl)pyrimidine-4-carboxamide;

6-((S)-5-((1r,4S)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-N-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(methoxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*R*)-5-((2*R*,5*R*)-5-(methoxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2*R*,5*R*)-5-(((*S*)-2-hydroxypropanamido)methyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1r,4r)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(3,4-difluorobenzyl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-(5-((1S,4r)-4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzo[*b*][1,4]dioxin-6-yl)methyl)-6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-((2,3-dihydrobenzo[*b*][1,4]dioxin-6-yl)methyl)-6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-Fluorobenzyl)-6-(5-((1S,4r)-4-((*S*)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methylbenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((S)-5-((1S,4S)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

N-(3-chloro-4-fluorobenzyl)-6-((R)-5-((1S,4R)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-((*S*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

6-((*R*)-5-((2*R*,5*R*)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-N-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide 2,2,2-trifluoroacetate;

(*S*)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-methyl-2-oxo-1,2-dihydropyridin-4-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

6-((*S*)-5-((1r,4S)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methylbenzyl)-2-methylpyrimidine-4-carboxamide;

(S)-*N*-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

(R)-*N*-(3,4-difluorobenzyl)-2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(6-methoxypyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

(S)-*N*-(4-fluoro-3-methoxybenzyl)-2-methyl-6-(5-(1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)pyrimidine-4-carboxamide;

2-methyl-6-(5-(1-methyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(methoxy(methyl)amino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-(5-chloro-1-(2-(methylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-((S)-5-((1s,4R)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-((R)-5-((1r,4R)-4-acetamidocyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[b][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-6-(5-(1-(2-(isopropylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide;

6-(5-((1S,4r)-4-((S)-2-hydroxypropanamido)cyclohexyl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-(5-(3-chloro-5-cyclopropyl-4-oxocyclohexyl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(S)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

(R)-6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridazin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

6-(5-(1-cyclopropyl-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-2-methyl-*N*-((3-oxo-3,4-dihydro-2*H*-benzo[*b*][1,4]oxazin-6-yl)methyl)pyrimidine-4-carboxamide;

6-(5-(1-(2-(cyclopropylamino)-2-oxoethyl)-6-oxo-1,6-dihydropyridin-3-yl)-4,5-dihydroisoxazol-3-yl)-*N*-(4-fluoro-3-methoxybenzyl)-2-methylpyrimidine-4-carboxamide;

N-(4-fluoro-3-methoxybenzyl)-1-methyl-6-oxo-5-(5-phenyl-4,5-dihydroisoxazol-3-yl)-1,6-dihydropyridazine-3-carboxamide; or

N-(4-fluoro-3-methoxybenzyl)-6-((*S*)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide; or

a pharmaceutically acceptable salt thereof.

9. A pharmaceutical composition comprising a compound of claim 1, stereoisomer, or pharmaceutically acceptable salt thereof and a pharmaceutically acceptable carrier or excipient.

10. A kit comprising a pharmaceutical composition of claim 9, packaging and instructions for use.

11. The compound of claim 6, wherein R^a is H; R^b is H; and m is 1.

12. The compound of claim 11, wherein Z is 1,4-dioxanyl, phenyl, pyridonyl, or piperidinyl, each of which is optionally substituted by one or more substituents independently selected from Br, Cl, F, $-CF_3$, $-CH_3$, CN, $-C(O)OH$, $-C(O)OCH_3$, $-C(O)CH_3$, $-C(O)C(H)(CH_3)_2$, -C(O)-cyclopropyl, -C(O)-cyclobutyl, $-C(O)CH_2CN$, $-C(O)CH_2OH$, $-C(O)C(CH_3)_2OH$, $-C(O)CH_2OCH_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)CH_2N(CH_3)_2$, $-C(O)CH_2CH_2N(CH_3)_2$, $-C(O)CH_2N(H)C(O)CH_3$, $-C(O)CH_2N(H)C(O)N(H)CH_3$, -

$C(O)CH_2S(O)_2CH_3$, $-C(O)NH_2$, $-C(O)N(CH_3)_2$, $-C(O)N(H)CH_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)N(H)CH(CH_3)_2$, $-C(O)N(H)CH_2CH_2OH$, $-C(O)N(CH_3)CH_2CH_2OH$, $-OCH_3$, $-OCH_2CH_2OH$, $-NH_2$, $-N(CH_3)_2$, $-N(H)C(O)CH_3$, $-N(H)C(O)C(H)(OH)CH_3$, $-N(H)C(O)CH_2N(H)CH_3$, $-N(H)C(O)N(CH_3)_2$, $-N(H)C(O)N(H)CH_3$, $-N(H)C(O)N(H)CH_2CH_2OH$, $-N(H)C(O)N(H)C(H)(CH_2OH)_2$, $-N(H)C(O)N(CH_3)CH_2CH_2OH$, $-N(H)C(O)N(CH_3)CH_2C(H)(OH)CH_2OH$, $-N(H)S(O)_2CH_3$, -OH, $-S(O)_2CH_3$, $-S(O)_2C(H)(CH_3)_2$, $-CH_2CN$, $-CH_2C(O)N(H)CH_3$, $-CH_2C(O)N(H)$-cyclopropyl, $-CH_2OCH_3$, $-CH_2OH$, $-CH_2CH_2OH$, $-C(H)(CH_3)_2OH$, $-CH_2NH_2$, $-CH_2N(CH_3)_2$, $-CH_2N(H)C(O)CH_3$, $-CH_2N(H)C(O)N(H)CH_3$, $-CH_2N(H)C(O)N(CH_3)_2$, $-CH_2N(H)C(O)CH_2OH$, $-CH_2N(H)S(O)_2CH_3$, $-CH_2N(H)C(O)C(H)(OH)CH_3$, $-CH_2N(H)C(O)C(CH_3)_2OH$, $-CH_2N(H)C(O)CH_2CN$, $-CH_2S(O)_2CH_3$, $-CH_2$-morpholinyl, or cyclopropyl.

13. *The compound of claim* 11, *wherein Z is* 1,4-*dioxanyl or piperidinyl, each of which is* optionally substituted by one or more substituents independently selected from Br, Cl, F, $-CF_3$, $-CH_3$, CN, $-C(O)OH$, $-C(O)OCH_3$, $-C(O)CH_3$, $-C(O)C(H)(CH_3)_2$, $-C(O)$-cyclopropyl, $-C(O)$-cyclobutyl, $-C(O)CH_2CN$, $-C(O)CH_2OH$, $-C(O)C(CH_3)_2OH$, $-C(O)CH_2OCH_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)CH_2N(CH_3)_2$, $-C(O)CH_2CH_2N(CH_3)_2$, $-C(O)CH_2N(H)C(O)CH_3$, $-C(O)CH_2N(H)C(O)N(H)CH_3$, $-C(O)CH_2S(O)_2CH_3$, $-C(O)NH_2$, $-C(O)N(CH_3)_2$, $-C(O)N(H)CH_3$, $-C(O)N(H)CH_2CH_3$, $-C(O)N(H)CH(CH_3)_2$, $-C(O)N(H)CH_2CH_2OH$, $-C(O)N(CH_3)CH_2CH_2OH$, $-S(O)_2CH_3$, $-S(O)_2C(H)(CH_3)_2$, $-CH_2CN$, $-CH_2OCH_3$, $-CH_2OH$, $-CH_2CH_2OH$, $-C(H)(CH_3)_2OH$, $-CH_2NH_2$, $-CH_2N(CH_3)_2$, $-CH_2N(H)C(O)CH_3$, $-CH_2N(H)C(O)N(H)CH_3$, $-CH_2N(H)C(O)N(CH_3)_2$, $-CH_2N(H)S(O)_2CH_3$, $-CH_2N(H)C(O)CH_2OH$, $-CH_2N(H)C(O)C(CH_3)_2OH$, $-CH_2N(H)C(O)C(H)(OH)CH_3$, $-CH_2N(H)C(O)CH_2CN$, or $-CH_2S(O)_2CH_3$.

14. A compound or a pharmaceutically acceptable salt thereof, wherein the compound is N-(4-fluoro-3-methoxybenzyl)-6-((S)-5-((2R,5R)-5-(hydroxymethyl)-1,4-dioxan-2-yl)-4,5-dihydroisoxazol-3-yl)-2-methylpyrimidine-4-carboxamide.