PYRIMIDINE DERIVATIVES FOR THE TREATMENT OF VIRAL INFECTIONS.

This invention relates to pyrimidine derivatives, processes for their preparation, pharmaceutical compositions, and their use in treating viral infections, like HBV or HCV.

The present invention relates to the use of pyrimidine derivatives in the treatment of viral infections, immune or inflammatory disorders, whereby the modulation, or agonism, of toll-like-receptors (TLRs) is involved. Toll-Like Receptors are primary transmembrane proteins characterized by an extracellular leucine rich domain and a cytoplasmic extension that contains a conserved region. The innate immune system can recognize pathogen-associated molecular patterns via these TLRs expressed on the cell surface of certain types of immune cells. Recognition of foreign pathogens activates the production of cytokines and upregulation of co-stimulatory molecules on phagocytes. This leads to the modulation of T cell behaviour.

It has been estimated that most mammalian species have between ten and fifteen types of Toll-like receptors. Thirteen TLRs (named TLR1 to TLR13) have been identified in humans and mice together, and equivalent forms of many of these have been found in other mammalian species. However, equivalents of certain TLR found in humans are not present in all mammals. For example, a gene coding for a protein analogous to TLR10 in humans is present in mice, but appears to have been damaged at some point in the past by a retrovirus. On the other hand, mice express TLRs 11, 12, and 13, none of which are represented in humans. Other mammals may express TLRs which are not found in humans. Other non-mammalian species may have TLRs distinct from mammals, as demonstrated by TLR14, which is found in the Takifugu pufferfish. This may complicate the process of using experimental animals as models of human innate immunity.

For detailed reviews on toll-like receptors see the following journal articles. Hoffmann, J.A., Nature, **426**, p33-38, 2003; Akira, S., Takeda, K., and Kaisho, T., Annual Rev. Immunology, **21**, p335-376, 2003; Ulevitch, R. J., Nature Reviews: Immunology, **4**, p512-520, 2004.

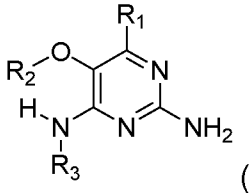
Compounds indicating activity on Toll-Like receptors have been previously described such as purine derivatives in WO 2006/117670, adenine derivatives in WO 98/01448 and WO 99/28321, and pyrimidines in WO 2009/067081.

However, there exists a strong need for novel Toll-Like receptor modulators having preferred selectivity, higher potency, higher metabolic stability, and an improved safety profile compared to the compounds of the prior art.

In the treatment of certain viral infections, regular injections of interferon (IFNα) can be administered, as is the case for hepatitis C virus (HCV), (Fried et. al. Peginterferon-alfa plus ribavirin for chronic hepatitis C virus infection, *N Engl J Med* 2002; 347: 975-82). Orally available small molecule IFN inducers offer the potential advantages of reduced immunogenicity and convenience of administration. Thus, novel IFN inducers are potentially effective new class of drugs for treating virus infections. For an example in the literature of a small molecule IFN inducer having antiviral effect see De Clercq, E.; Descamps, J.; De Somer, P. *Science* **1978**, *200*, 563-565.

IFNα is also given in combination with other drugs in the treatment of certain types of cancer (Eur. J. Cancer 46, 2849–57, and Cancer Res. **1992**, *52*, 1056). TLR 7/8 agonists are also of interest as vaccine adjuvants because of their ability to induce pronounced Th1 response (Hum. Vaccines 2010, 6, 1−14; Hum. Vaccines 2009, 5, 381−394).

In accordance with the present invention a compound of formula (I) is provided



(I)

or a pharmaceutically acceptable salt, tautomer(s), solvate or polymorph thereof, wherein

R_1 is hydrogen, methyl. C_{1-2}alkyl, cyclopropyl, methoxy, halogen, hydroxyl, trifluoromethyl, or difluoromethyl,

R_2 is C_{1-8}alkyl, (C_{1-4})alkoxy-(C_{1-4})alkyl, C_{3-7}cycloalkyl, C_{4-7}heterocycle, aromatic, bicyclic heterocycle, arylalkyl, heteroaryl, heteroarylalkyl each of which is optionally substituted by one or more substituents independently selected from halogen, hydroxyl, amino, C_{1-6}alkyl, di-(C_{1-6})alkylamino, C_{1-6}alkylamino, C_{1-6}alkyl, C_{1-6} alkoxy, C_{3-6} cycloalkyl, carboxylic acid, carboxylic ester,

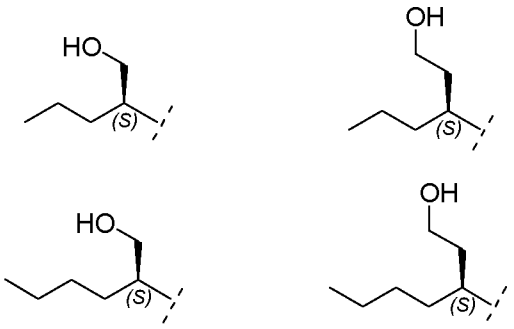
carboxylic amide, heterocycle, aryl, alkenyl, alkynyl, arylalkyl, heteroaryl, heteroarylalkyl, nitrile, and

R_3 is C_{4-8}alkyl, C_{4-8}alkoxy, C_{2-6} alkenyl or C_{2-6}alkynyl, each of which is optionally substituted by one or more substituents independently selected from halogen, hydroxyl, amino, C_{1-3}alkyl, C_{1-3}alkoxy or C_{3-6} cycloalkyl, nitrile.

In a first embodiment the present invention provides compounds of formula (I) wherein R_3 is butyl or pentyl and wherein R_2 and R_1 are as specified above.

In a further embodiment the invention concerns compounds of formula (I) wherein R_3 is C_{4-8}alkyl substituted with hydroxyl, and wherein R_2 and R_1 are as specified above.

Another embodiment relates to compounds of formula (I) wherein R_3, when being C_{4-8}alkyl substituted with hydroxyl, is one of the following

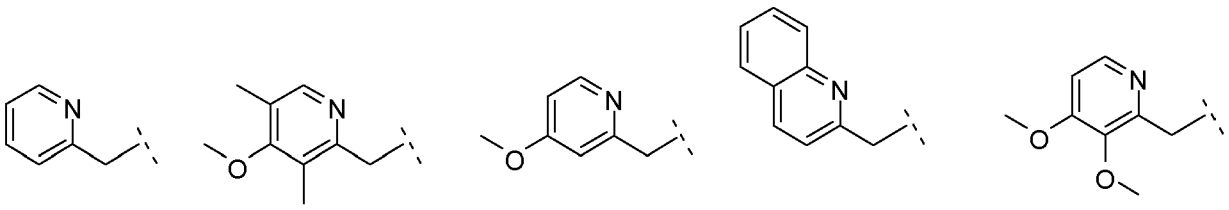


Furthermore the present invention also provides compounds of formula (I) wherein R_1 is hydrogen or $-CH_3$ and wherein R_2 and R_3 are as specified above.

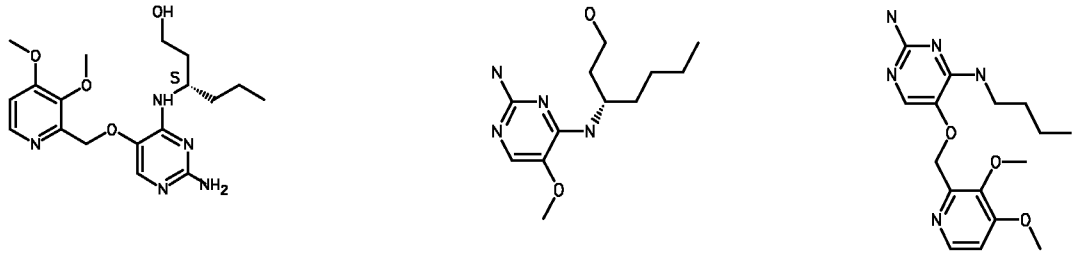
In another embodiment the present invention provides compounds of formula (I) wherein wherein R_2 is arylalkyl or heteroarylalkyl, substituted with C_{1-3}alkyl, hydroxyl, alkoxy, nitrile, heterocycle or ester and wherein R_1 and R_3 are as specified above.

In a further embodiment the current invention concerns compounds of formula (I) wherein R_2 is C_{1-3}alkyl substituted by aryl, heterocycle, or heteroaryl which is further substituted by C_{1-3}alkyl, alkoxy, carboxylic ester or carboxylic amide and wherein R_1 and R_3 are as specified above

Furthermore the invention relates to compounds of formula (I) wherein R_2 is one of the following examples that can be further substituted with C_{1-3}alkyl, hydroxyl, alkoxy, nitrile, heterocycle or ester.



The preferred compounds according to the invention are:



The compounds of formula (I) and their pharmaceutically acceptable salt, tautomer(s), solvate or polymorph thereof have activity as pharmaceuticals, in particular as modulators of Toll-Like Receptors (especially TLR7 and/or TLR8) activity.

In a further aspect the present invention provides a pharmaceutical composition comprising a compound of formula (I) or a pharmaceutically acceptable salt, solvate or polymorph thereof together with one or more pharmaceutically acceptable excipients, diluents or carriers.

Furthermore a compound of formula (I) or a pharmaceutically acceptable salt, solvate or polymorph thereof according to the current invention, or a pharmaceutical composition comprising said compound of formula (I) or a pharmaceutically acceptable salt, solvate or polymorph thereof can be used as a medicament.

Another aspect of the invention is that a compound of formula (I) or a pharmaceutically acceptable salt, solvate or polymorph thereof, or said pharmaceutical composition comprising said compound of formula (I) or a pharmaceutically acceptable salt, solvate or polymorph thereof can be used accordingly in the treatment of a disorder or disease in which the modulation of TLR7 and /or TLR8 is involved.

The term “alkyl” refers to a straight-chain or branched-chain saturated aliphatic hydrocarbon containing the specified number of carbon atoms.

The term “halogen” refers to fluorine, chlorine, bromine or iodine.

The term “alkenyl” refers to an alkyl as defined above consisting of at least two carbon atoms and at least one carbon-carbon double bond.

The term “ alkynyl” refers to an alkyl as defined above consisting of at least two carbon atoms and at least one carbon-carbon triple bond.

The term “cycloalkyl” refers to a carbocyclic ring containing the specified number of carbon atoms.

The term “heteroaryl” means an aromatic ring structure as defined for the term “aryl” comprising at least 1 heteroatom selected from N, O and S, in particular from N and O.

The term “aryl” means an aromatic ring structure optionally comprising one or two heteroatoms selected from N, O and S, in particular from N and O. Said aromatic ring structure may have 4, 5, 6 or 7 ring atoms. In particular, said aromatic ring structure may have 5 or 6 ring atoms.

The term “bicyclic heterocycle” means an aromatic ring structure, as defined for the term “aryl” comprised of two fused aromatic rings. Each ring is optionally comprised of heteroatoms selected from N, O and S, in particular from N and O.

The term arylalkyl” means an aromatic ring structure as defined for the term “aryl” optionally substituted with an alkyl group.

The term “heteroarylalkyl” means an aromatic ring structure as defined for the term “heteroaryl” optionally substituted by an alkyl group.

The term “alkoxy” refers to an alkyl (carbon and hydrogen chain) group singular bonded to oxygen like for instance a methoxy group or ethoxy group.

Heterocycle refers to molecules that are saturated or partially saturated and include ethyloxide, tetrahydrofuran, dioxane or other cyclic ethers. Heterocycles containing nitrogen include, for example azetidine, morpholine, piperidine, piperazine, pyrrolidine, and the like. Other heterocycles include, for example, thiomorpholine, dioxolinyl, and cyclic sulfones.

Heteroaryl groups are heterocyclic groups which are aromatic in nature. These are monocyclic, bicyclic, or polycyclic containing one or more heteroatoms selected from N, O or S. Heteroaryl groups can be, for example, imidazolyl, isoxazolyl, furyl, oxazolyl, pyrrolyl, pyridonyl, pyridyl, pyridazinyl, or pyrazinyl.

Pharmaceutically acceptable salts of the compounds of formula (I) include the acid addition and base salts thereof. Suitable acid addition salts are formed from acids which form non-toxic salts. Suitable base salts are formed from bases which form non-toxic salts.

The compounds of the invention may also exist in unsolvated and solvated forms. The term "solvate" is used herein to describe a molecular complex comprising the compound of the invention and one or more pharmaceutically acceptable solvent molecules, for example, ethanol.

The term "polymorph" refers to the ability of the compound of the invention to exist in more than one form or crystal structure.

The compounds of the present invention may be administered as crystalline or amorphous products. They may be obtained for example as solid plugs, powders, or films by methods such as precipitation, crystallization, freeze drying, spray drying, or evaporative drying. They may be administered alone or in combination with one or more other compounds of the invention or in combination with one or more other drugs. Generally, they will be administered as a formulation in association with one or more pharmaceutically acceptable excipients. The term "excipient" is used herein to describe any ingredient other than the compound(s) of the invention. The choice of excipient depends largely on factors such as the particular mode of administration, the effect of the excipient on solubility and stability, and the nature of the dosage form.

The compounds of the present invention or any subgroup thereof may be formulated into various pharmaceutical forms for administration purposes. As appropriate compositions there may be cited all compositions usually employed for systemically administering drugs. To prepare the pharmaceutical compositions of this invention, an effective amount of the particular compound, optionally in addition salt form, as the active ingredient is combined in intimate admixture with a pharmaceutically acceptable carrier, which carrier may take a wide variety of forms depending on the form of preparation desired for administration. These pharmaceutical compositions are desirably in unitary dosage form suitable, for example, for oral, rectal, or percutaneous administration. For example, in preparing the compositions in oral dosage form, any of the usual pharmaceutical media may be employed such as, for example, water, glycols, oils, alcohols and the like in the case of oral liquid preparations such as suspensions, syrups, elixirs, emulsions, and solutions; or solid carriers

such as starches, sugars, kaolin, diluents, lubricants, binders, disintegrating agents and the like in the case of powders, pills, capsules, and tablets. Because of their ease in administration, tablets and capsules represent the most advantageous oral dosage unit forms, in which case solid pharmaceutical carriers are obviously employed. Also included are solid form preparations that can be converted, shortly before use, to liquid forms. In the compositions suitable for percutaneous administration, the carrier optionally comprises a penetration enhancing agent and/or a suitable wetting agent, optionally combined with suitable additives of any nature in minor proportions, which additives do not introduce a significant deleterious effect on the skin. Said additives may facilitate the administration to the skin and/or may be helpful for preparing the desired compositions. These compositions may be administered in various ways, e.g., as a transdermal patch, as a spot-on, as an ointment. The compounds of the present invention may also be administered via inhalation or insufflation by means of methods and formulations employed in the art for administration via this way. Thus, in general the compounds of the present invention may be administered to the lungs in the form of a solution, a suspension or a dry powder.

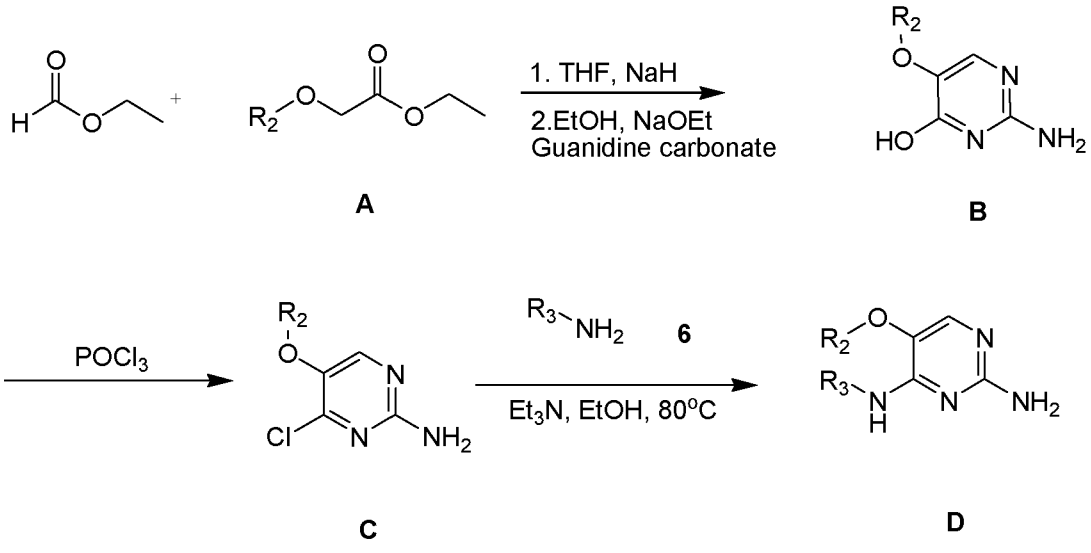
It is especially advantageous to formulate the aforementioned pharmaceutical compositions in unit dosage form for ease of administration and uniformity of dosage. Unit dosage form as used herein refers to physically discrete units suitable as unitary dosages, each unit containing a predetermined quantity of active ingredient calculated to produce the desired therapeutic effect in association with the required pharmaceutical carrier. Examples of such unit dosage forms are tablets (including scored or coated tablets), capsules, pills, powder packets, wafers, suppositories, injectable solutions or suspensions and the like, and segregated multiples thereof.

Those of skill in the treatment of infectious diseases will be able to determine the effective amount from the test results presented hereinafter. In general it is contemplated that an effective daily amount would be from 0.01 mg/kg to 50 mg/kg body weight, more preferably from 0.1 mg/kg to 10 mg/kg body weight. It may be appropriate to administer the required dose as two, three, four or more sub-doses at appropriate intervals throughout the day. Said sub-doses may be formulated as unit dosage forms, for example, containing 1 to 1000 mg, and in particular 5 to 200 mg of active ingredient per unit dosage form.

The exact dosage and frequency of administration depends on the particular compound of formula (I) used, the particular condition being treated, the severity of the condition being treated, the age, weight and general physical condition of the particular patient as well as other medication the individual may be taking, as is well known to those skilled in the art. Furthermore, it is evident that the effective amount may be lowered or increased depending on the response of the treated subject and/or depending on the evaluation of the physician prescribing the compounds of the instant invention. The effective amount ranges mentioned above are therefore only guidelines and are not intended to limit the scope or use of the invention to any extent.

Preparation of compounds.

Compounds of formula (I), where R_1 is hydrogen atom are prepared according to **scheme 1.**



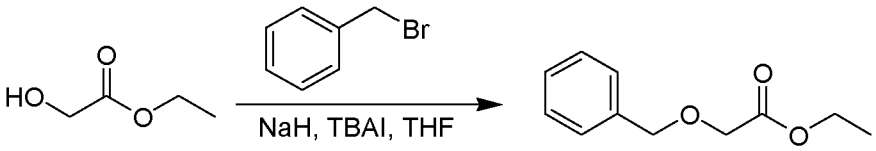
Scheme 1

Compounds of type A, in scheme 1 are made by either

(i) Reaction of a heterocyclic alcohol with a halogenated ester and an suitable base, for example potassium carbonate, cesium carbonate, or sodium hydride. Example shown in **scheme 2a**.

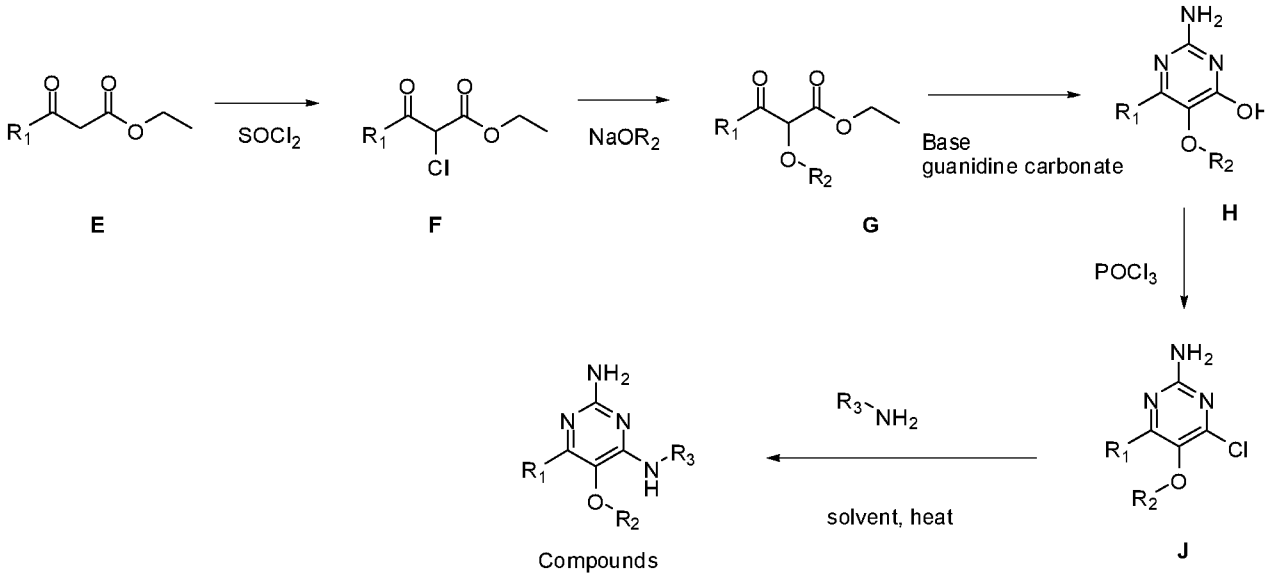
(ii) Reaction of an alcohol, or hydroxy ester, for example 2-hydroxy ethyl acetate, with an alkyl halide using an appropriate base, for example sodium hydride. Example shown in **scheme 2b**.

Scheme 2a

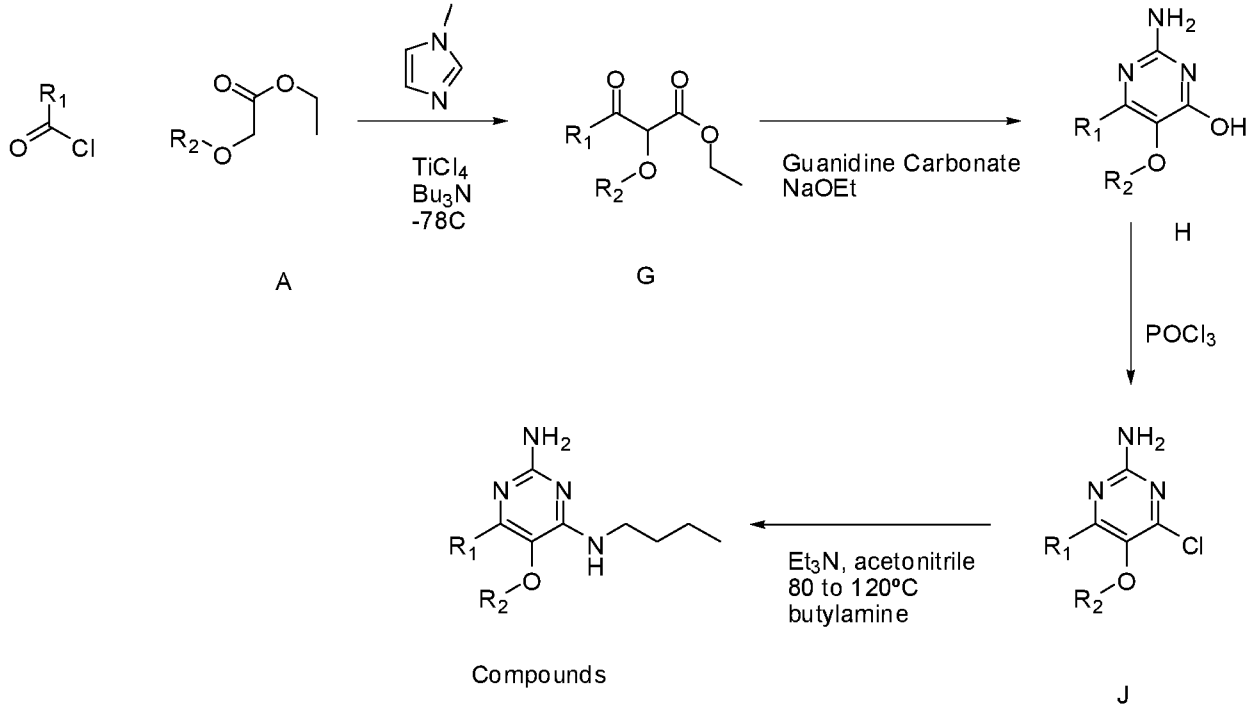


Scheme 2b

Compounds of formula (I), when R_1 is alkyl, cycloalkyl, trifluoromethyl, or alkoxy and where R_2 is aryl or heteroaryl, are prepared as in scheme 3 below. The betaketoester (E) can be chlorinated using, for example, thionyl chloride to provide the 2-chloro-beta-ketoester intermediate (F). The phenol or hetero-aromatic alcohol (R2OH) is combined with an equimolar ratio of aqueous sodium hydroxide. The solvents are then removed under reduced pressure to afford the phenol or heteroaromatic alcohol salt of R2. This salt is combined with the 2-chloro-β-ketoester intermediate (F) to afford intermediate G according to literature procedure. Intermediate G is then combined, with or without base, with guanidine carbonate in an appropriate solvent, for example, ethanol. Intermediate H is then reacted with phosphorous oxychloride to form the chloropyrimidine intermediate (J). The products are then formed as a result of heating (J) in the presence of excess amine and optionally excess organic base, for example triethylamine, at elevated temperature. This is a general scheme using methods known to a skilled person, see for instance Organic Syntheses volume 33, p.43 (1953).



Scheme 3



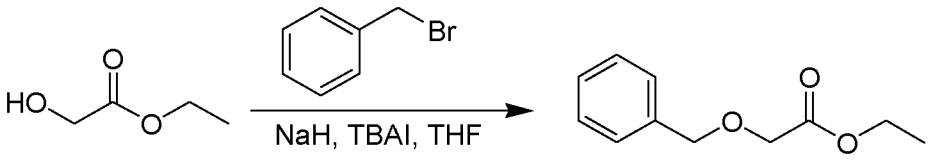
Scheme 4

Compounds of formula (I), when R_1 is alkyl, cycloalkyl, trifluoromethyl, or alkoxy and where R_2 is aromatic or aliphatic, can be prepared according scheme 4. This reaction scheme begins with a crossed-Claisen reaction where an acyl chloride reacts with ester intermediate A (shown in scheme 1) to form intermediates (G) as in scheme 3. From intermediate G, the reaction scheme follows the same pathway to the products as in scheme 3. This is a general

scheme using methods known to a skilled person, see for instance The Journal of American Chemical Society volume 127, page 2854 (2005).

Experimental Section.

Synthesis of Intermediate A-1.

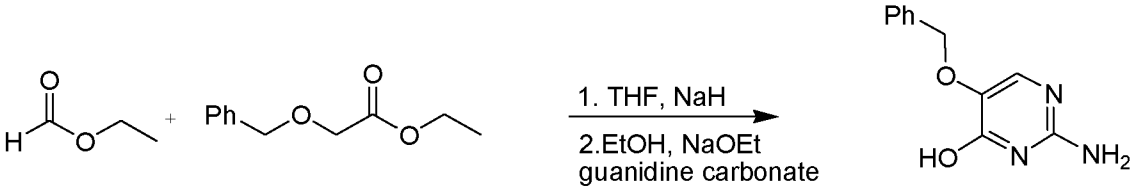


A-1

To a mixture of ethyl glycolate [623-50-7] (250.00 g, 2.40 mol), NaH (105.65 g, 2.64 mol), tetrabutylammonium iodide (TBAI) (88.70 g, 240.14 mmol) in anhydrous THF (2 L) was added benzyl bromide (451.80 g, 2.64 mol) dropwise at 0°C. The resulting mixture was stirred at 25°C for 16 hours. The reaction mixture was quenched with saturated, aqueous ammonium chloride (1L), and the aqueous layer was extracted with ethyl acetate (3 x 1L). The combined organic layers were washed with brine (1 L), dried over magnesium sulfate, the solids were removed via filtration, and the solvents of the filtrate were concentrated under reduced pressure. The residue was purified by silica gel column chromatography (petroleum ether: ethyl acetate = 6:1) to obtain intermediate **A-1** (200 g).

^{1}H NMR ($CDCl_3$ 400MHz) δ ppm 7.37-7.27 (m, 5H); 4.62 (s, 2H), 4.24-4.19 (q, J = 6.8 Hz, 2H); 4.07 (s, 2H); 1.29-1.25 (t, J = 6.8 Hz, 3H).

Procedure for preparation of Intermediate B-1.



A-1 **B-1**

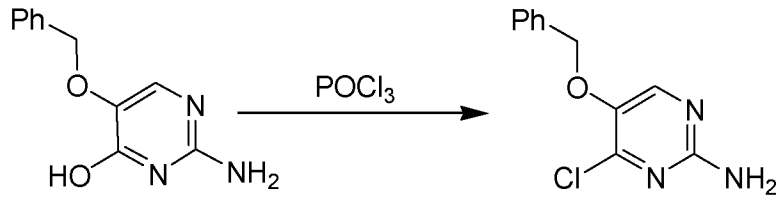
To a stirred suspension of NaH (45.30 g, 1.13 mol) in anhydrous THF (1.2 L) was added ethyl formate (114.42 g, 1.54 mol). The suspension was cooled in an ice bath, and then compound **A-1** (200 g, 1.03 mol) in anhydrous THF (300 mL) was added dropwise via an addition funnel. The white mixture was stirred at 0°C to room temperature for 5 hours. During this time, the reaction was exothermic and turned yellow. In a separate flask, guanidine carbonate [593-85-1] (111.31 g, 0.618 mol) was treated with a sodium ethoxide solution,

freshly prepared by the careful addition of Na (28.41 g, 1.24 mol) to anhydrous ethanol (750 mL) at room temperature. The off-white slurry obtained after stirring for 1 hour, was then added to the yellow solution prepared above. The resulting pale yellow reaction mixture was heated to reflux for 15 hours. The solvent was removed, and then the crude residue was dissolved in water (1.5 L). The mixture was adjusted to pH=5 with acetic acid. The solid was collected, washed extensively with water and ethanol to give intermediate **B-1** (160 g).

^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 4.90 (s, 2 H), 6.33 (br. s., 2 H), 7.25 (s, 1 H), 7.29 - 7.42 (m, 5 H), 11.21 (br. s., 1 H)

Procedure for preparation of intermediate C-1.

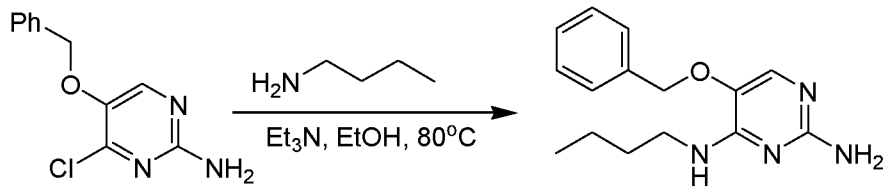
Reaction Scheme:



B-1 **C-1**

A suspension of intermediate **B-1** (160 g, 0.74 mol) in $POCl_3$ (900 mL) was heated to 100°C under N_2 with stirring for 5 hours. The reaction mixture was cooled to room temperature. The excess $POCl_3$ was removed under reduced pressure, the oil residue was poured into cold, sat. aq. $NaHCO_3$ (2 L) that was stirred for 30 minutes. The mixture was extracted with ethyl acetate (3 x 1.5 L). The combined organic layers were separated and washed with brine (1 L), dried over sodium sulfate, the solids were removed via filtration, and the solvents of the filtrate were concentrated to afford intermediate **C-1** (70 g) as a yellow solid. The product was used in the next step without further purification.

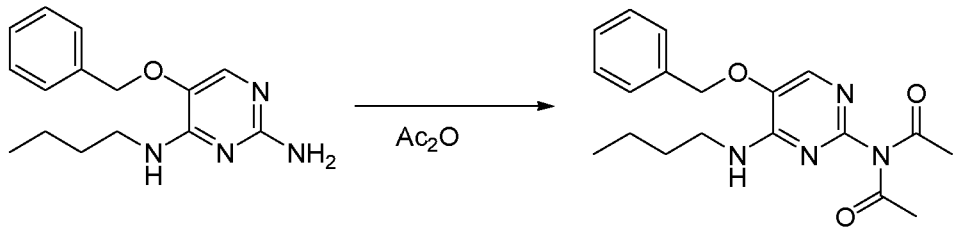
Procedure for preparation of **compound 1.**



C-1 **1**

To a suspension of **C-1** (70.00 g, 297.03 mmol) in ethanol (1.4 L) was added *n*-butylamine (217.24 g, 2.97 mol) and triethylamine (60.11 g, 594.05 mmol). The reaction mixture was heated to reflux for 16 hours. The reaction mixture was cooled to room temperature and the solvents were removed under reduced pressure. The residue was purified by silica gel flash chromatography using a petroleum ether to ethyl acetate gradient to obtain **1** (26 g) as a pale yellow solid.

^{1}H NMR (400 MHz, METHANOL-d_4) δ ppm 0.96 (t, *J*=7.3 Hz, 3 H), 1.32 - 1.43 (m, 2 H), 1.52 - 1.61 (m, 2 H), 3.38 (t, *J*=7.2 Hz, 2 H), 5.01 (s, 2 H), 7.28 (s, 1 H), 7.31 - 7.46 (m, 5 H)



1 **D-1**

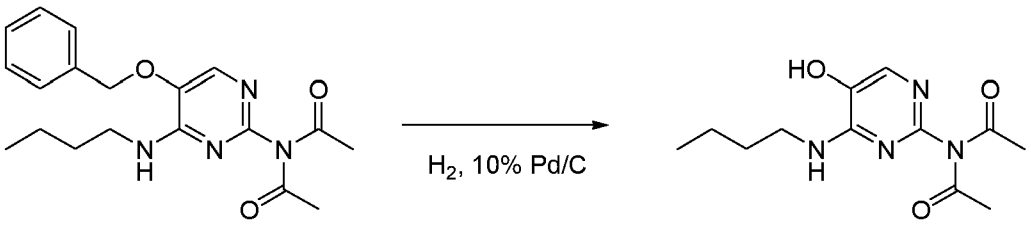
Preparation of intermediate D-1.

Into a 100 mL round bottom flask equipped with a magnetic stir bar was placed **1** (1g, 3.67 mmol) in acetic anhydride (40 mL). The yellow solution was allowed to stir at reflux for 15 hours. The solvents were removed under reduced pressure. The crude was purified via silica gel chromatography using a heptane to ethyl acetate gradient. The best fractions were collected and the solvents were removed under reduced pressure to afford a white solid, **D-1**.

LC-MS: Anal. Calcd. For $C_{19}H_{24}N_4O_3$: 356.19; found 357$[M+H]^+$

^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.94 (t, *J*=7.4 Hz, 3 H), 1.31 - 1.45 (m, 2 H), 1.50 - 1.67 (m, 2 H), 2.31 (s, 6 H), 3.44 (m, *J*=6.0 Hz, 2 H), 5.12 (s, 2 H), 5.41 - 5.52 (m, 1 H), 7.43 (m, *J*=1.5 Hz, 5 H), 7.79 (s, 1 H)

Preparation of intermediate D-2.



D-1 D-2

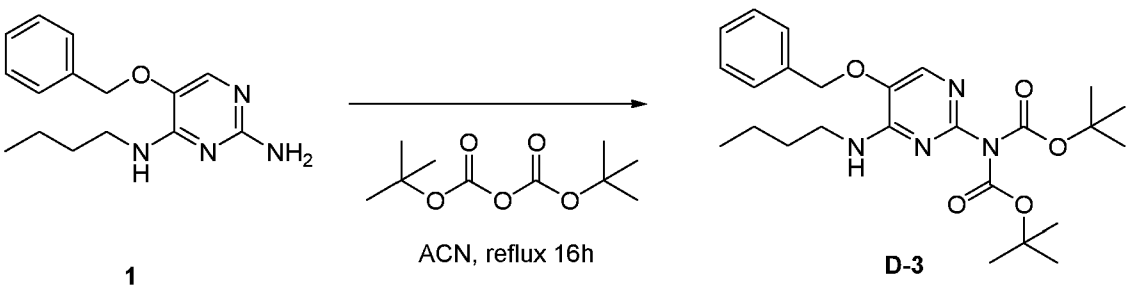
Method A. Into a 250 mL erlenmeyer flask equipped with a magnetic stir bar was placed intermediate D-1 (1g), and ethanol (100 mL). The flask is sparged with nitrogen, followed by the addition of 10% Pd on carbon (100 mg). The flask was sealed and the atmosphere removed and replaced with hydrogen. The reaction was allowed to stir at room temperature for 15 hours. The heterogeneous mixture was filtered through packed celite and the solvents of the filtrate were removed under reduced pressure to afford D-2 in quantitative yield.

Method B. A 0.1 M solution of starting material in methanol was run through the H-cube, equipped with a 10% Pd/C cartridge, at 0.5 mL/min and 30 bar pressure of hydrogen. LC-MS shows complete conversion. The solvents were removed under reduced pressure. The crude was purified via silica gel chromatography using a dichloromethane to 10% methanol in dichloromethane gradient. The best fractions were pooled; the solvents were removed under reduced pressure to afford a white solid, **D-2**.

LC-MS: Anal. Calcd. For $C_{12}H_{18}N_4O_3$: 266.14; found 267$[M+H]^+$

^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.4 Hz, 3 H), 1.28 (dd, J=14.9, 7.4 Hz, 2 H), 1.49 (t, J=7.2 Hz, 2 H), 2.15 (s, 6 H), 3.20 - 3.37 (m, 2 H), 7.02 - 7.12 (m, 1 H), 7.58 (s, 1 H), 10.27 (br. s, 1 H)

Preparation of intermediate D-3.



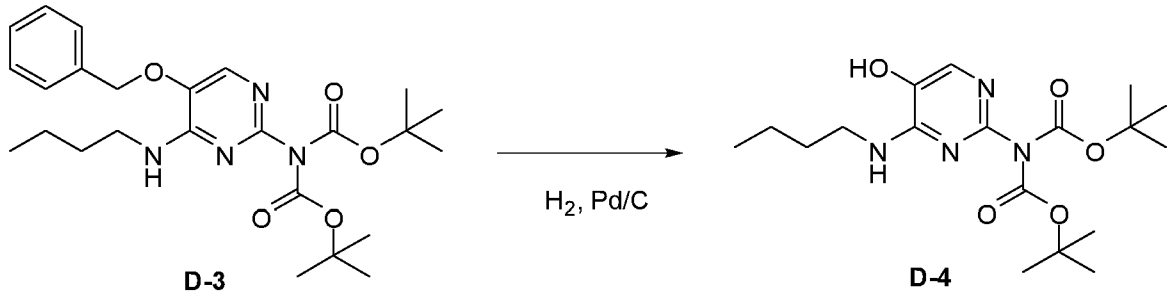
1 D-3

Into a 100 mL round bottom flask was placed **1** (1 g, 3.67 mmol), di-*tert*-butyl dicarbonate (7.5 g), and acetonitrile (50 mL). The yellow solution was stirred at reflux for 16 hours. The solvents were removed under reduced pressure. The residue was purified via silica chromatography using a prepacked 80g silica column and a heptane to ethyl acetate gradient autocollecting at 254nm. The best fractions were pooled to afford a yellow oil, **D-3**.

LC-MS: Anal. Calcd. For $C_{25}H_{36}N_4O_5$: 472.269; found 473$[M+H]^+$

^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.94 (t, *J*=7.4 Hz, 3 H), 1.33 - 1.42 (m, 2 H), 1.46 (s, 18 H), 1.50 - 1.65 (m, 2 H), 3.35 - 3.51 (m, 2 H), 5.09 (s, 2 H), 5.31 - 5.38 (m, 1 H), 7.36 - 7.48 (m, 5 H), 7.75 (s, 1 H)

Preparation of intermediate D-4.

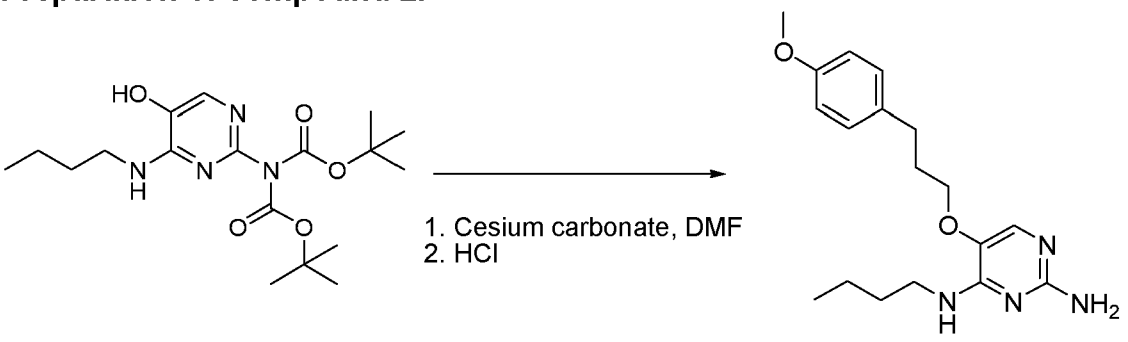


D-3 **D-4**

Intermediate D-4 is prepared according to the procedure to prepare intermediate D-2, employing either method A or B.

LC-MS: Anal. Calcd. For $C_{18}H_{30}N_4O_5$: 382.222; found 383$[M+H]^+$
^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.95 (t, *J*=7.3 Hz, 3 H), 1.39 (s, 18 H), 1.40 - 1.45 (m, 2 H), 1.53 - 1.64 (m, 2 H), 3.42 - 3.51 (m, 2 H), 5.66 (s, 1 H), 7.43 (s, 1 H)

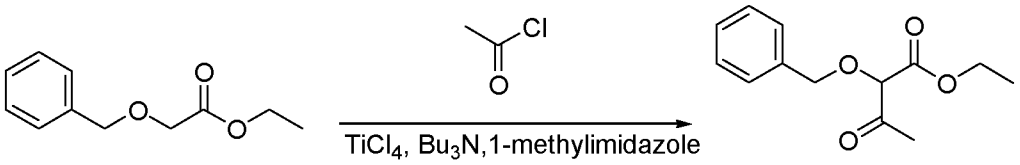
Preparation of compound 2.



D-4 2

Into a 30 mL vial was placed intermediate D-4 (200 mg, 0.52 mmol), DMF (5 mL), 1-(3-bromopropyl)-4-methoxybenzene (130 mg, 0.57 mmol), and cesium carbonate (508 mg, 1.56 mmol). The reaction was allowed to stir for 15 hours at room temperature. The solids were removed via filtration. The solvents of the filtrate were removed under reduced pressure and the crude was reconstituted in methanol and to it was added HCl (6M in isopropanol) and the reaction was allowed to stir 15 hours at room temperature. The solvents were removed under reduced pressure and the crude was purified via reverse phase separation to afford **2** as the free base.

Preparation of intermediate G-1.

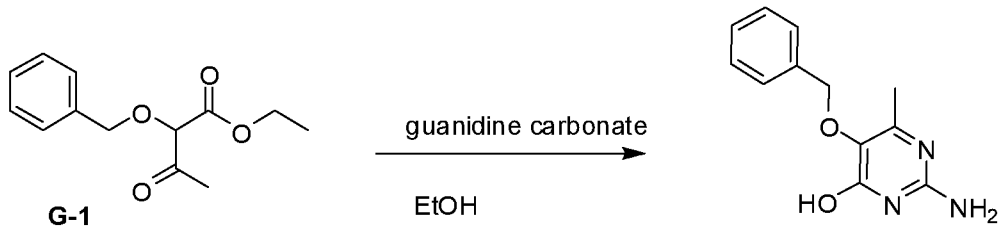


A-1 G-1

To a stirred solution of **A-1** (60 g, 309 mmol, 1eq) and 1-methylimidazole (30.4 g, 370 mmol, 1.2 eq) in CH_2Cl_2 (1 L) was added acetyl chloride (24.3 g, 309 mmol, 1eq) at -45 ^oC under N_2. After stirring for 20 min, $TiCl_4$ (210 g, 1.08 mol, 3.5 eq) and tributylamine (230 g, 1.24 mol, 4 eq) were added to the mixture at -45^oC under N_2, and continues to stir for 50 minutes at -45^oC under N_2. After completion, water and ethyl acetate were added. The organic layer was separated and the aqueous layer was extracted with ethyl acetate twice. The organic layer was washed with brine and dried over sodium sulfate. The solids were removed by filtration and the solvents of the filtrate were removed under reduced pressure. The crude was purified via silica column chromatography using a heptane to ethyl acetate gradient to afford a pale yellow oil, **G-1**.

^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 1.30 (t, J=7.2 Hz, 3 H), 2.28 (s, 3 H), 4.27 (q, J=7.2 Hz, 2 H), 4.41 (s, 1 H), 4.58 (d, J=11.8 Hz, 1 H), 4.75 (d, J=11.8 Hz, 1 H), 7.32 - 7.43 (m, 5 H)

Preparation of intermediate H-1.

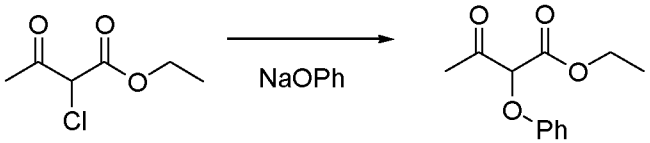


G-1 H-1

Into a 20 mL microwave vial was placed intermediate G-1 (500 mg, 2.12 mmol), ethanol (5mL), and guanidine carbonate (200 mg, 2.22 mmol). The vial was sealed and allowed to react at 120ºC with stirring for 4 hours. The solvents were removed under reduced pressure. Water (25 mL) was added. The mixture was brought to pH=5 via careful addition of acetic acid. The precipitate was isolated via filtration to afford a white solid, **H-1**.

^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 1.88 (s, 3 H), 4.85 (s, 2 H), 6.38 (br. s., 2 H), 7.24 - 7.49 (m, 5 H), 11.16 (s, 1 H)

Preparation of intermediate G-2.



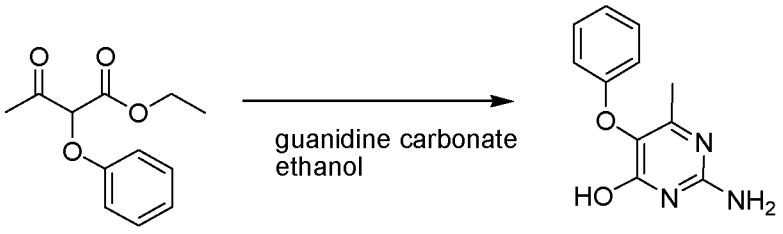
F-1 G-2

Step 1. Sodium phenolate was prepared by evaporating equimolar portions of phenol and sodium hydroxide in a 1L round bottom flask on the rotary evaporator. Toluene is used in the azeotropic removal of water.

Step 2. Sodium phenolate (116 g, 1 mol) prepared in step 1 and toluene (1 L) were placed in a 2L three-necked flask fitted with mechanical stirrer, addition funnel, and reflux condenser with drying tube. The suspension was heated to reflux, then ethyl α-chloroacetoacetate (165 g, 1 mol) was added with stirring through the addition funnel where the reaction continues heating at reflux for 4 hours. The light brown suspension is cooled to room temperature, extracted

with water (2 x 500 mL), and dried (anhydrous magnesium sulfate). The solids were removed via filtration and the solvents of the filtrate were removed under reduced pressure. The crude is used in the next step without purification.

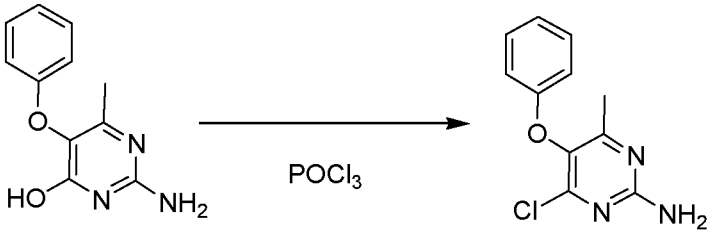
Preparation of intermediate H-2.



G-2 H-2

Into a 100 mL round bottom flask equipped with a magnetic stir bar and reflux condenser was added intermediate G-2 (1g, 4.5 mmol), ethanol (50 mL), and guanidine carbonate [593-85-1](203 mg, 2.25 mmol). The reaction mixture is brought to reflux for 15 hours. The solvent was removed under reduced pressure. Water (25 mL) was added. The mixture was brought to pH=5 via careful addition of acetic acid. The precipitate was isolated via filtration to afford a white solid, **H-2**. This is used without further purification in the next step.

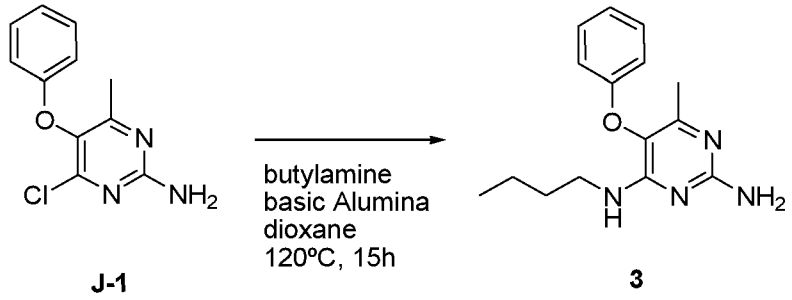
Preparation of intermediate J-1.



H-2 J-1

Into a 50 mL round bottom flask equipped with a magnetic stir bar and reflux condenser was added intermediate H-2 (500 mg, 2.3 mmol) and $POCl_3$ (20 mL). The suspension was heated to reflux with stirring for 6 hours. The solvents were removed under reduced pressure to afford a crude brown oil, **J-1**. No further purification was done. The compound was used as such in the subsequent step.

Preparation of 3.



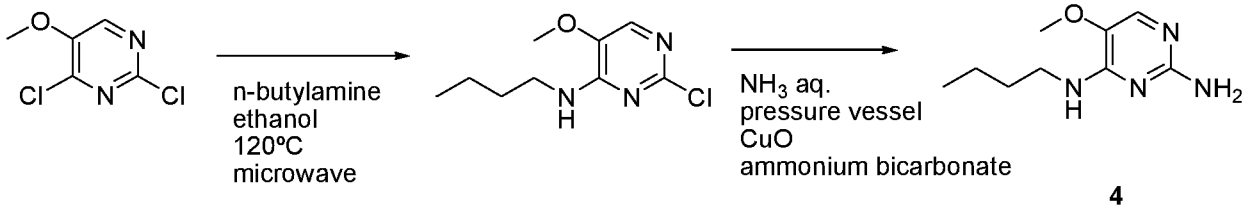
J-1 3

Into a 50 mL sealed tube equipped with a magnetic stir bar was placed intermediate J-1 (150 mg, 0.64 mmol), n-butylamine (70 mg, 0.96 mmol), basic alumina (100 mg), and dioxane (10 mL). The tube was sealed, placed in an oil bath at 120ºC, and the reaction was heated with stirring for 15 hours. The vessel was cooled to room temperature and the cap was carefully removed. The contents were poured into a round bottom flask where the solvents were removed under reduced pressure. The crude was purified via silica gel column chromatography using a dichloromethane to 5% methanol in dichloromethane gradient. The best fractions were pooled, and the solvents were removed under reduced pressure to afford **3**.

LC-MS: Anal. Calcd. For $C_{15}H_{20}N_4O$: 272.16; found 273 $[M+H]^+$

^1H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.80 (t, J=7.3 Hz, 3 H), 1.20 (dq, J=15.0, 7.3 Hz, 2 H), 1.33 - 1.47 (m, 2 H), 1.98 (s, 3 H), 3.20 - 3.34 (m, 2 H), 4.74 (br. s., 2 H), 4.79 (br. s., 1 H), 6.78 - 6.84 (m, 2 H), 6.91 - 7.01 (m, 1 H), 7.18 - 7.28 (m, 2 H)

Preparation of 4



4

Step 1.

Into a 20 mL microwave vial was added commercially available 2,4-dichloro-5-methoxy pyrimidine (300 mg, 1.68 mmol), ethanol (5 mL), and n-butylamine (0.166 mL, 1.68 mmol). The vial is sealed then heated in the microwave for

10 minutes at 120°C. LC-MS shows complete conversion. The solvents were removed under reduced pressure. The crude is used as such in step 2.

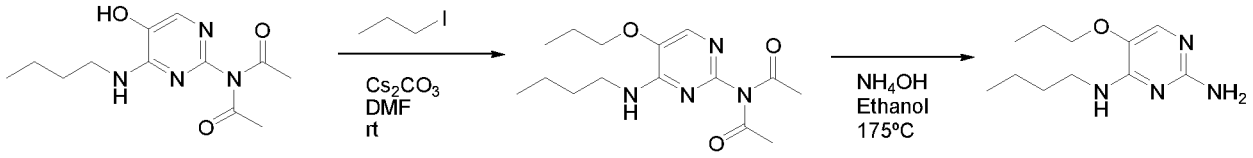
Step 2.

Compound from step 1 was placed into a 20 mL pressure vessel with aqueous ammonia (10 mL) and to this was added ammonium bicarbonate (200 mg, 2.6 mmol), and CuO (24 mg, 0.17 mmol, 0.1eq). The vessel was sealed and the mixture was heated to 120°C with stirring for 24 hours. The reaction mixture was extracted 3 times with 5mL dichloromethane:methanol 9:1 and the volatiles were removed under reduced pressure. The compound was filtered through silica eluting with dichloromethane:methanol 9:1 and the volatiles were removed under reduced pressure. The residue was purified by reversed phase chromatography.

LC/MS: Anal. Calcd. For $C_9H_{16}N_4O$: 196.13; found 197$[M+H]^+$

^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.97 (t, *J*=7.3 Hz, 3 H), 1.35 - 1.48 (m, 2 H), 1.56 - 1.68 (m, 2 H), 3.44 - 3.52 (m, 2 H), 3.80 (s, 3 H), 5.86 (s, 1 H), 5.97 (s, 2 H), 7.07 - 7.14 (m, 1 H)

Preparation of 5.



D-2 5

Step 1.

Into a 16 x 100 test tube was placed intermediate D-2 (180 mg, 0.66 mmol), DMF (5 mL), propyl iodide (111 mg, 0.656 mmol), and cesium carbonate (320 mg, 0.98 mmol). The reaction was allowed to stir at room temperature for 15 hours. The solids were removed by filtration, and the solvents of the filtrate were removed under reduced pressure. The crude was purified via silica gel chromatography using a dichloromethane to 10% methanol in dichloromethane gradient. The best fractions were pooled, the solvents were removed under reduced pressure to afford a white solid.

Step 2.

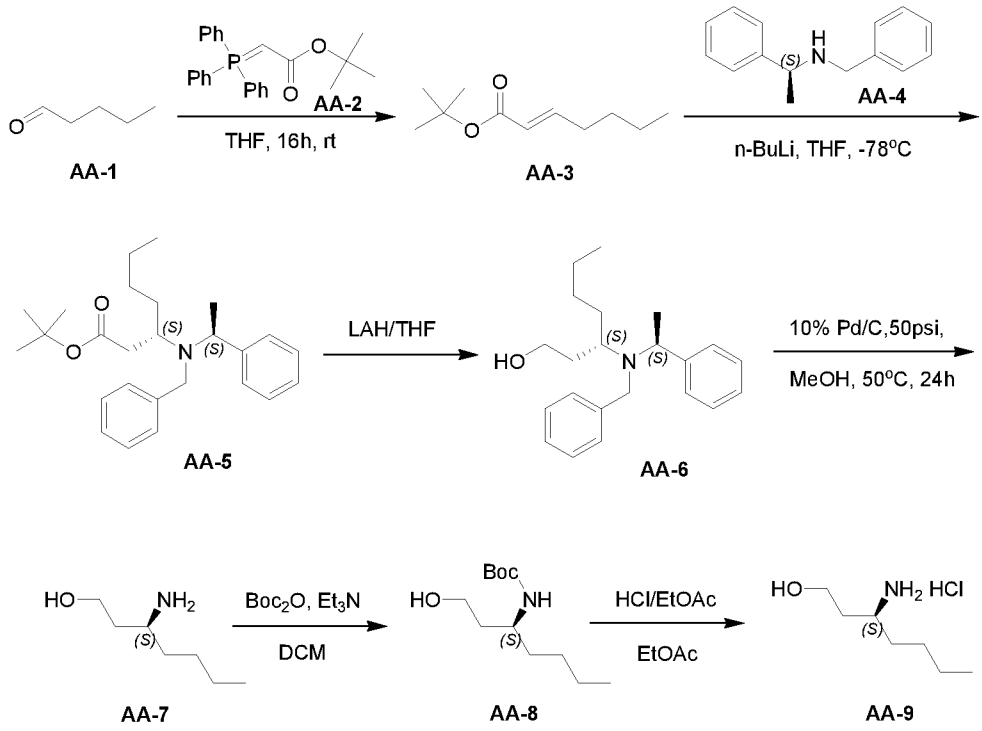
Into a 10 mL microwave vial was placed the above white solid (100 mg), ammonium hydroxide (1 mL) and ethanol (1 mL). The vial was sealed and

heated with stirring to 175ºC for 10 minutes. LC-MS shows complete conversion to product. The solvents were removed under reduced pressure. The crude was purified via silica gel chromatography using a dichloromethane to 10% methanol in dichloromethane gradient. The best fractions were pooled, the solvents were removed under reduced pressure to afford a colorless oil. Addition of one equivalent of HCl (using 5 to 6N HCl in isopropanol) affords a white solid, **5**.

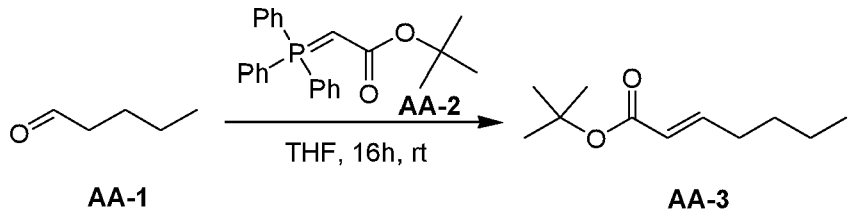
LC/MS: Anal. Calcd. For $C_{11}H_{20}N_4O$: 224.16; found 225$[M+H]^+$

^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 0.98 (t, J=7.4 Hz, 3 H), 1.20 - 1.35 (m, 2 H), 1.54 (t, J=7.2 Hz, 2 H), 1.69 - 1.75 (m, 2 H), 3.40 (d, J=7.0 Hz, 2 H), 3.87 (t, J=6.5 Hz, 2 H), 7.39 (d, J=5.5 Hz, 1 H), 7.46 (br. s., 2 H), 8.28 - 8.37 (m, 1 H)

Synthetic Scheme for the preparation of AA-9



Synthesis of intermediate AA-3

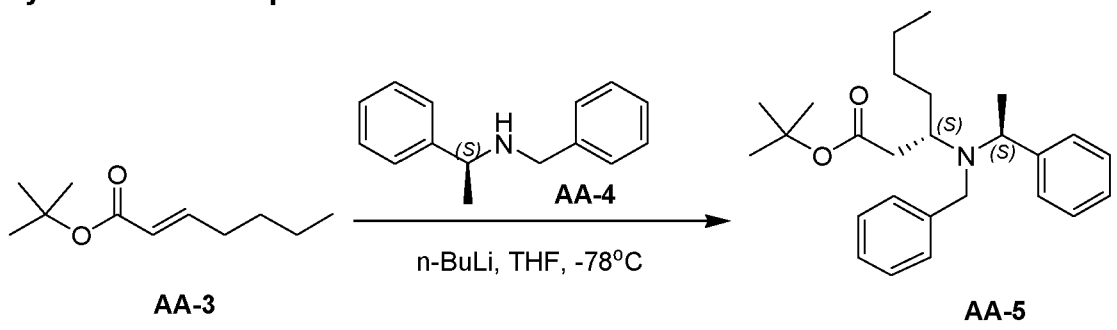


AA-1 **AA-3**

To a solution of valeraldehyde (43 g, 500 mmol) in THF (1 L) was added **AA-2** (200 g, 532 mmol) and the reaction mixture was stirred for 16 hours at room temperature. The solvents were evaporated and the residue was diluted in petroleum ether and filtered. The solvents of the filtrate were removed under reduced pressure and the residue was purified by silica chromatography using a petroleum ether to 3% ethyl acetate in petroleum ether gradient to give **AA-3** (90 g) as a colorless oil.

^{1}H NMR (400 MHz, $CDCl_3$): δ ppm 6.81-6.77 (m, 1H), 5.68-5.64 (td, J=1.2Hz, 15.6 Hz, 1H), 2.11-2.09 (m, 2H), 1.406 (s, 9H), 1.38-1.26(m, 4H), 0.85-0.81(t, J=7.2Hz, 3H).

Synthesis of compound AA-5

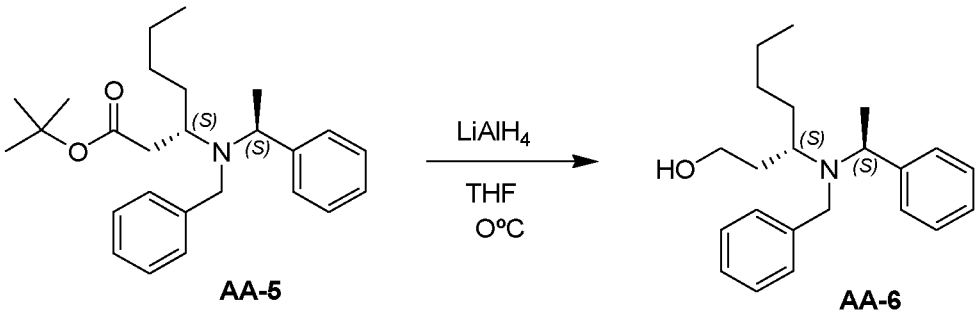


AA-3 **AA-5**

n-butyl lithium (290mL, 725mmol, 1.5eq.) was added to a stirred solution of AA-4 (165 g, 781 mmol) in THF (800 mL) at -78°C. The reaction mixture was stirred for 30 minutes then AA-3 (90 g, 488.4 mmol) in THF (400 mL) was added and the reaction was stirred for 2 hours at -78°C. The mixture was quenched with sat., aq. NH_4Cl solution and warmed to room temperature. The product was partitioned between ethyl acetate and water. The organic phase was washed with brine, dried and evaporated. The residue was purified by column chromatography eluting with 5% ethyl acetate in petroleum ether to afford a colorless oil, **AA-5** (132 g).

^{1}H NMR (400 MHz, $CDCl_3$): δ ppm 7.36-7.16 (m, 10H), 3.75-3.70 (m, 2H), 3.43-3.39 (d, J=15.2Hz, 1H), 3.33-3.15 (m, 1H), 1.86-1.80 (m, 2H), 1.47-1.37 (m, 2H), 1.32 (s, 9H), 1.26-1.17 (m, 7H), 0.83-0.79 (t, J=7.2Hz, 3H).

Synthesis of AA-6

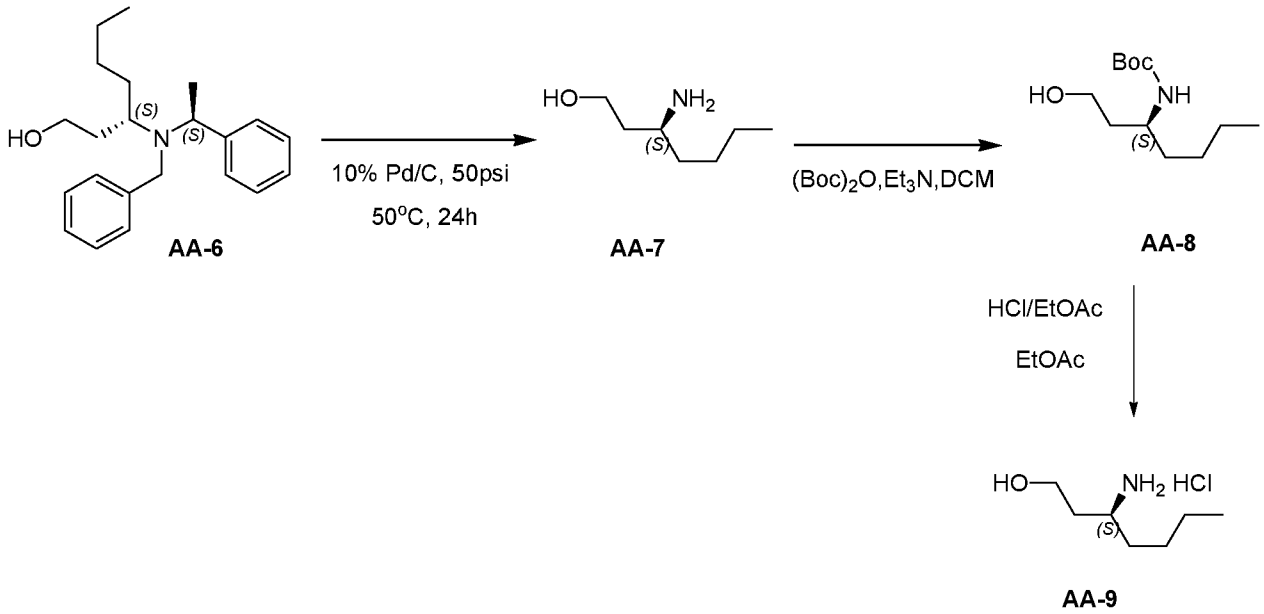


AA-5 **AA-6**

AA-5 (130 g, 328 mmol) was dissolved in THF (1.5 L) and LAH (20 g, 526 mmol) was added at 0°C in small portions. The resulting mixture was stirred at the same temperature for 2 hours and then allowed to warm to room temperature. The mixture was quenched with a sat. aq. NH_4Cl solution. The product was partitioned between ethyl acetate and water. The organic phase was washed with brine, dried and evaporated. The combined organic layers were dried over sodium sulfate, the solids were removed via filtration and concentrated to afford crude **AA-6** (100 g), which was used in the next step without further purification.

^{1}H NMR (400 MHz, $CDCl_3$): δ ppm 7.33-7.14 (m, 10H), 3.91-3.86 (m, 1H), 3.80-3.77 (d, J=13.6Hz, 1H), 3.63-3.60 (d, J=13.6Hz, 1H), 3.43-3.42 (m, 1 H), 3.15-3.10 (m, 1H), 2.70-2.63 (m, 2H), 1.65-1.28 (m, 10H), 0.89-0.81 (m, 3H).

Synthesis of AA-9



A solution of AA-6 (38 g, 116.75 mmol) and 10% Pd/C in methanol (200 mL) was hydrogenated under 50 PSI hydrogen at 50°C for 24 hours. The reaction mixture was filtered and the solvent was evaporated to give crude product **AA-7** (17 g).

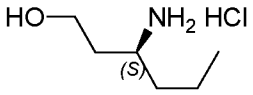
The crude product was dissolved in dichloromethane (200 mL), triethylamine (26.17 g, 259.1 mmol) and di-*tert*-butyl dicarbonate (84.7 g, 194.4 mmol) was added at 0°C. The resulting mixture was stirred at room temperature for 16 hours. The mixture was partitioned between dichloromethane and water. The organic phase was washed with brine, dried and evaporated. The residue was purified by silica gel chromatography eluting with 20% ethyl acetate in petroleum ether to give **AA-8** (13 g) as colorless oil.

^{1}H NMR (400 MHz, $CDCl_3$): δ ppm 4.08-4.03 (br, 1H), 3.68 (m, 1H), 3.58-3.55 (m, 2H), 3.20-2.90(br, 1H), 1.80-1.73 (m, 1H), 1.42-1.17 (m, 15 H), 0.85-0.82(t, J=6.8Hz, 3H).

AA-8 (42 g, 0.182 mol) was dissolved in dioxane (200 mL) and dioxane/HCl (4M, 200 mL) was added at 0°C. The resulting mixture was stirred at room temperature for 2h. The solvent was evaporated to afford the crude product. A dichloromethane/petroleum ether mixture (50 mL, 1:1, v/v) was added to the crude product, and the supernatant was decanted. This procedure was repeated two times to obtain an oil, **AA-9** (26.6 g).

^{1}H NMR (400 MHz, DMSO-d_6): δ ppm 8.04 (s, 3H), 3.60-3.49 (m, 2H), 3.16-3.15 (m, 1H), 1.71-1.67 (m, 2H), 1.60-1.55(m, 2H), 1.33-1.26 (m, 4H), 0.90-0.87 (t, J=6.8Hz, 3H).

Preparation of AA-10

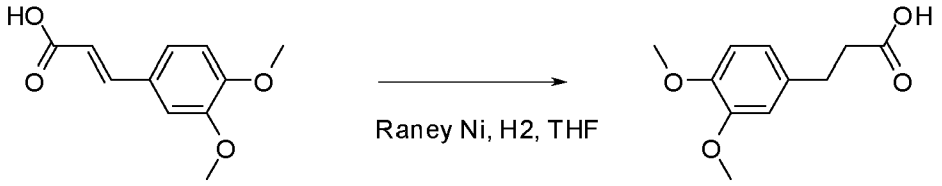


AA-10

AA-10 was prepared according to the preparation of AA-9, using butyraldehyde instead of valeraldehyde.

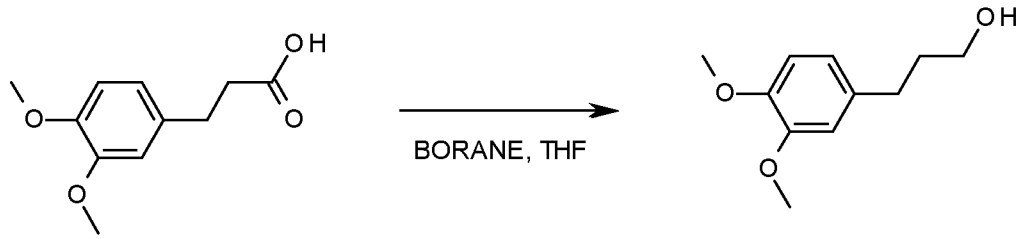
^{1}H NMR (400 MHz, DMSO-d_6):δ ppm 8.07 (s, 3H), 4.85 (br, 1H), 3.57-3.45 (m, 2H), 3.14-3.12 (m, 1H), 1.70-1.64 (m, 2H), 1.56-1.49 (m, 2H), 1.38-1.30 (m, 2H), 0.90-0.80 (t, J=6.8Hz, 3H).

Preparation of 74

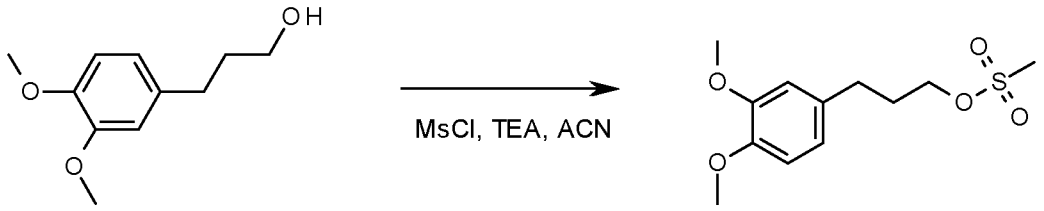


Step 1. 3,4-dimethoxycinnamic acid (5 g, 24 mmol) was dissolved in THF (100 mL). Raney Nickel was added to this solution under a N_2 atmosphere. The reaction mixture was exposed to a hydrogen atmosphere and stirred 15 hours at room temperature. The reaction mixture was filtered over a cartridge packed with diatomateous earth and the solvent of the filtrate was removed under reduced pressure. The residue was used as such in the next step.

LC-MS: Anal. Calcd. For $C_{11}H_{14}O_4$: 210.09; found 209[M-H]

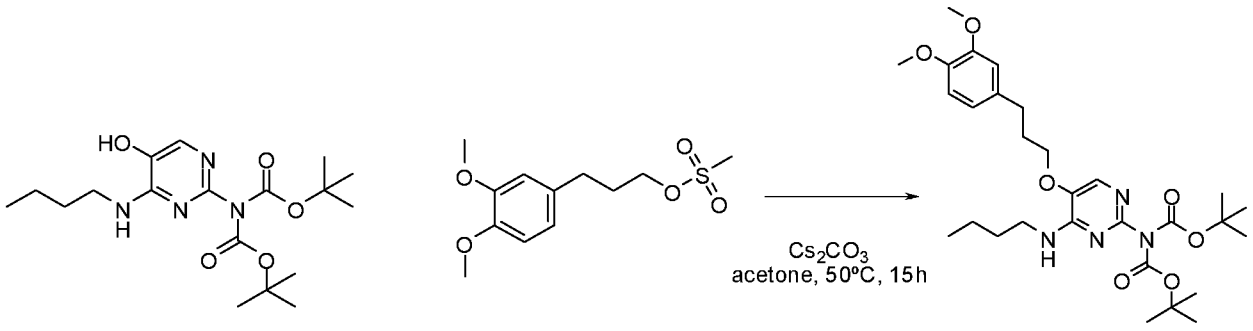


Step 2. 3-(3,4-dimethoxyphenyl)propanoic acid was dissolved in THF (100 mL). Borane-dimethyl sulfide complex (2M in diethyl ether, 20 mL, 40 mmol) was added. The reaction mixture was stirred overnight at room temperature. Methanol was added slowly to quench the reaction mixture, then silica was added and the volatiles were removed under reduced pressure. The residue was purified on silica using a heptane to ethyl acetate gradient yielding the product as an oil. This was used as such in the next step.
LC-MS: Anal. Calcd. For $C_{11}H_{16}O_3$: 196.11; found 195[M-H]



Step 3. 3-(3,4-dimethoxyphenyl)propan-1-ol (3.8 g, 19.5 mmol) and triethylamine (3.8 mL, 27.3 mmol) were dissolved in acetonitrile (15 mL) and then methanesulfonyl chloride (1.5 mL, 19.5 mmol) was added. The reaction mixture was shaken overnight at room temperature. The volatiles were removed under reduced pressure and the residue was purified via silica gel chromatography using a heptane to ethyl acetate gradient yielding the product as a clear oil.

^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 1.91 - 2.01 (m, 2 H), 2.58 - 2.64 (m, 2 H), 3.17 (s, 3 H), 3.72 (s, 3 H), 3.75 (s, 3 H), 4.19 (t, J=6.4 Hz, 2 H), 6.71 - 6.76 (m, 1 H), 6.81 - 6.89 (m, 2 H)

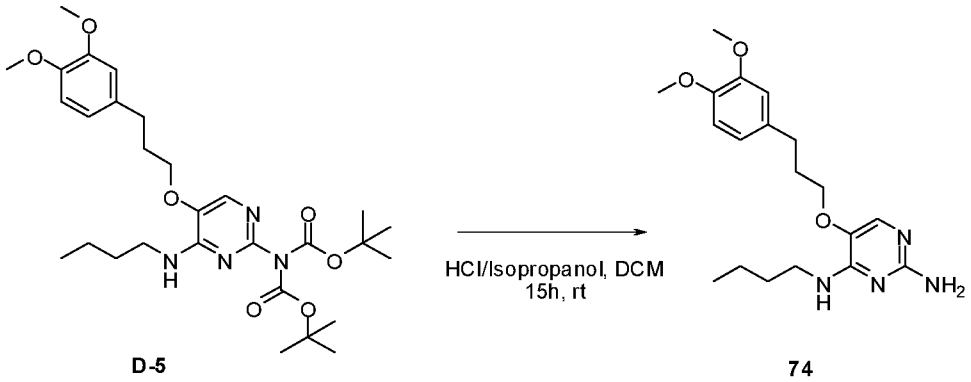


D-4 D-5

Step 4. A solution of **D-4** (400 mg, 1 mmol), cesium carbonate (511 mg, 1.6 mmol) and 3-(3,4-dimethoxyphenyl)propyl methanesulfonate (430 mg,

1.6 mmol) in acetone (50 mL) was heated to 50ºC for 15 hours. The reaction mixture was placed in the centrifuge and the supernatant was decanted then evaporated to dryness. The residue was purified via silica column chromatography using a gradient from heptane to ethyl acetate. The fractions containing the product were pooled and the solvents were removed under reduced pressure to afford **D-5**.

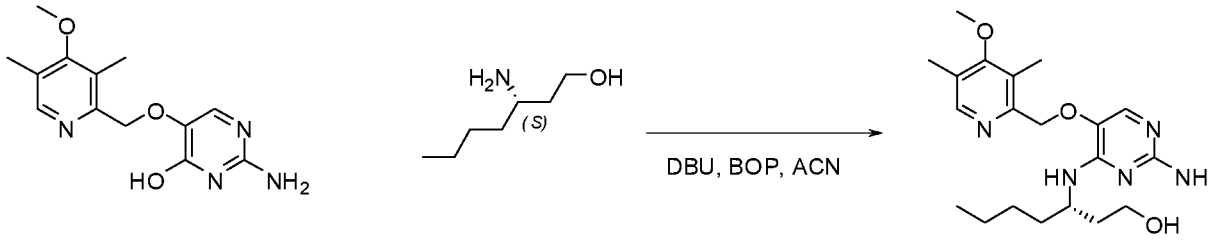
LC-MS: Anal. Calcd. For $C_{29}H_{44}N_4O_7$: 560.32; found 561 $[M+H]^+$



D-5 74

Step 5. The boc-protected compound was dissolved in dichloromethane (5 mL) and 6M HCL in isopropanol (3 mL) was added. The reaction mixture was stirred 15 hours at room temperature. The volatiles were removed under reduced pressure. Ether (5 mL) was added and a precipitate formed, **74** was isolated by filtration then dried in the vacuum oven for 15 hours.

Preparation of **75**



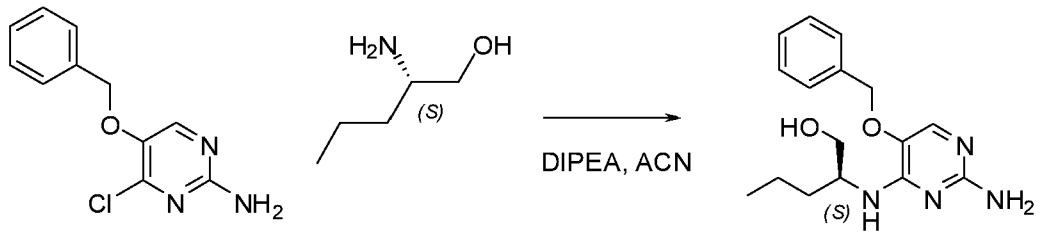
B-2 AA-9 75

Step 1. Intermediate **B-2** was prepared according to the method described for the preparation of intermediate **B-1**.

Step 2. To a solution of **B-2** (1 g, 3.62 mmol) and DBU (5.4 mL, 36 mmol) in acetonitrile (20 mL) was added BOP (2.08 g, 4.71 mmol) and the reaction mixture became transparent and was stirred for 15 minutes at room

temperature. **AA-9** (910 mg, 5.43 mmol) was added and the reaction mixture was stirred for 2 days at 50ºC. The volatiles were removed under reduced pressure and the residue was purified on silica using a dichloromethane to 10% methanol in dichloromethane gradient. The best fractions were pooled and the solvents were removed under reduced pressure. The crude was reconstituted in dichloromethane (2 mL) then HCl in diethylether was added to form the HCl salt. The precipitate was isolated by filtration and dried in the vacuum oven to afford compound **75**.

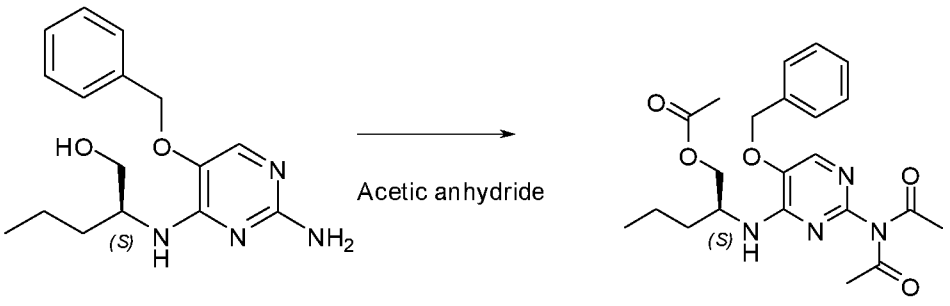
Preparation of **76**



C-1 **D-6**

Step 1. **C-1** (2 g, 8.49 mmol), L-norvalinol (1.75 g, 17 mmol) and diisopropylethylamine (5.85 mL, 34 mmol) were dissolved in acetonitrile (200 mL) in a 500 mL teflon coated pressure vessel and heated to 130ºC for 15 hours. The mixture was allowed to cool to room temperature, the volatiles were removed under reduced pressure and the crude was purified via silica gel column chromatography using a gradient from dichloromethane to 10% methanol in dichloromethane. The best fractions were pooled and the solvents were removed under reduced pressure to afford intermediate **D-6**.

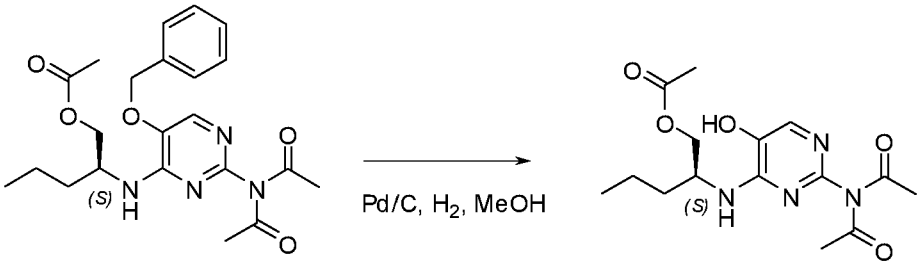
LC-MS: Anal. Calcd. For $C_{16}H_{22}N_4O_2$: 302.17; found 303 $[M+H]^+$



D-6 **D-7**

Step 2. **D-6** (2 g, 6.61 mmol) was heated to reflux in acetic anhydride (100 mL) in a 250 mL round bottom flask for 4 hours. The volatiles were removed under reduced pressure and the residue was purified via silica gel column chromatography using a heptane to ethyl acetate gradient yielding a yellow oil, **D-7**.

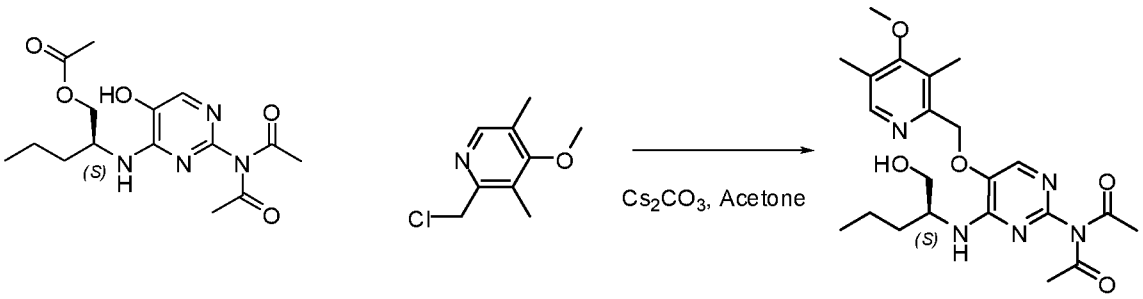
LC-MS: Anal. Calcd. For $C_{22}H_{28}N_4O_5$: 428.21; found 429 $[M+H]^+$



D-7 **D-8**

Step 3. **D-8** was prepared according to the method to prepare intermediate D-2.

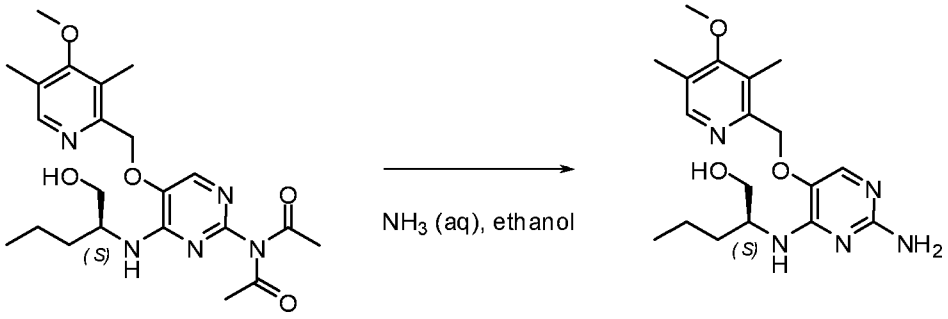
LC-MS: Anal. Calcd. For $C_{15}H_{22}N_4O_5$: 338.16; found 339 $[M+H]^+$



D-8 **D-9**

Step 4. Intermediate **D-9** was prepared according to the method described in example **75** from intermediate **D-4**.

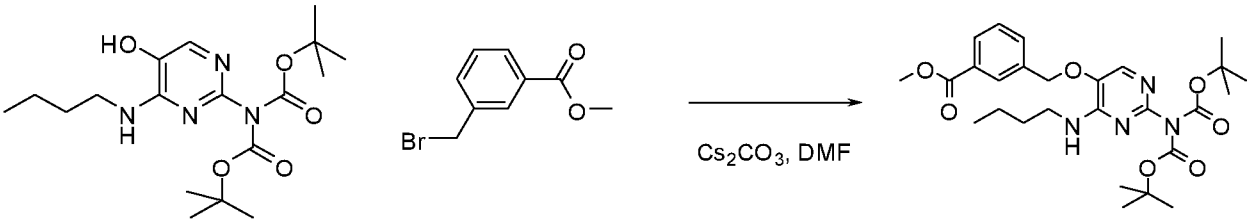
LC-MS: Anal. Calcd. For $C_{15}H_{22}N_4O_5$: 338.16; found 339 $[M+H]^+$



D-9 76

Step 5. Deprotection of D-9 was performed according to the method described in step 2 of compound 5 to afford **76**.

Preparation of compound **77**

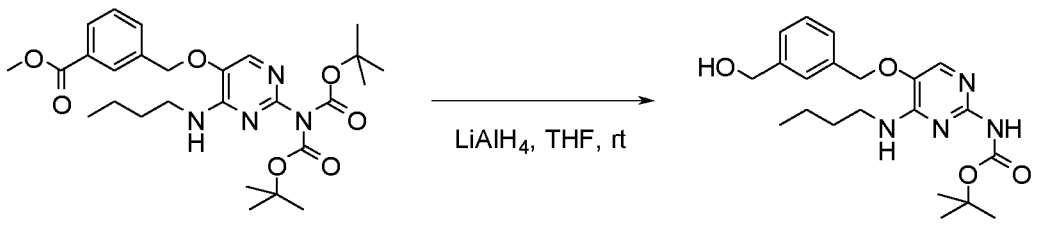


D-4 D-10

Step 1. **D-10** was prepared from D-4 according to the method to prepare example **5**, purification via silica column with heptane to ethyl acetate gradient.

LC-MS: Anal. Calcd. For $C_{27}H_{38}N_4O_7$: 530.27; found 531 $[M+H]^+$

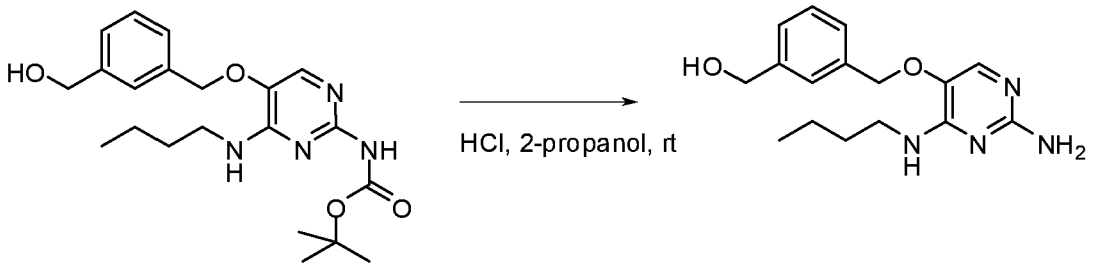
^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.93 (t, J=7.3 Hz, 3 H), 1.37 (dd, J=14.9, 7.4 Hz, 2 H), 1.53 - 1.62 (m, 2 H), 3.40 - 3.50 (m, 2 H), 3.92 - 3.95 (m, 3 H), 5.13 (s, 2 H), 5.33 (s, 1 H), 7.46 - 7.52 (m, 1 H), 7.56 - 7.62 (m, 1 H), 7.73 (s, 1 H), 8.05 (dt, J=7.7, 1.4 Hz, 1 H), 8.09 (d, J=1.5 Hz, 1 H)



D-10 D-11

Step 2. D-10 (2.14 g, 3.91 mmol) was dissolved in anhydrous THF (250 mL). Lithium aluminum hydride (1M in THF, 5.87 mL, 5.87 mmol) was added

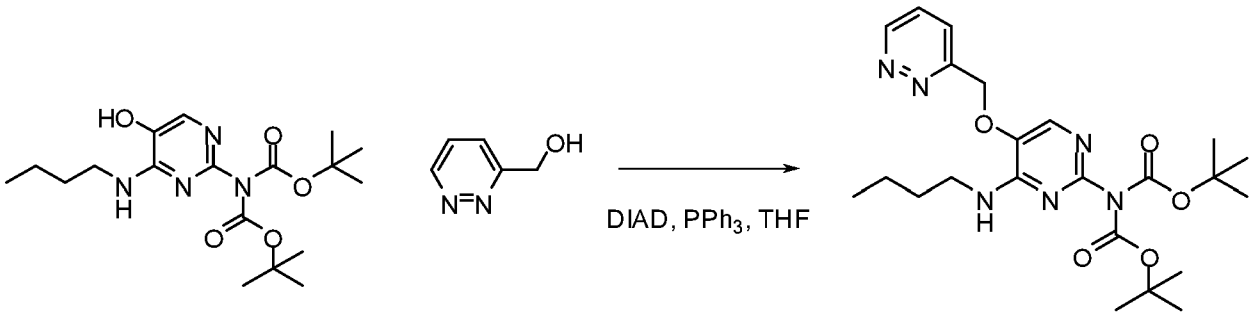
dropwise and the reaction mixture was stirred for 3 hours at room temperature. NH_4Cl (sat., aq.) was added drop wise to the reaction mixture and the precipitated salts were removed by filtration and washed with THF. The filtrate was evaporated to dryness and crude **D-11** was used as such in the next step. LC-MS: Anal. Calcd. For $C_{21}H_{30}N_4O_4$: 402.23; found 403 $[M+H]^+$



D-11 **77**

Step 3. D-11 (1.57 g, 3.91 mmol) was dissolved in dichloromethane (20 mL) and to it was added HCl (6 M in isopropanol, 50 mL). The reaction mixture stirred for 16 hours at room temperature. The volatiles were removed under reduced pressure and the crude was purified via silica column using a dichloromethane to 10% dichloromethane in methanol gradient yielding **77** as an oil which solidified on standing.

Preparation of **78**

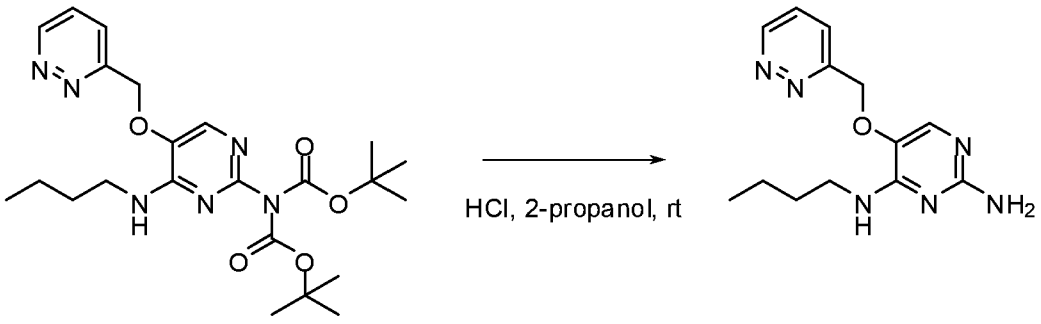


D-4 **D-12**

Step 1. A solution of D-4 (0.5 g, 1.31 mmol), 3-pyridazinylmethanol (158 mg, 1.44 mmol) and triphenylphosphine (377 mg, 1.44 mmol) in anhydrous THF (4 mL) was cooled to 0°C and a solution of DIAD (0.28 mL, 1.44 mmol) was added dropwise at 0°C. After addition, the reaction mixture was stirred for 3 hours at ambient temperature. The solvent was quenched with water (10 mL), stirred for 10 minutes and the volatiles were removed under reduced pressure. The water layer was extracted with dichloromethane, the organic layers were combined, and the solvent was removed under reduced pressure. The crude

was purified via silica gel column chromatography using a heptane to ethyl acetate gradient. The best fractions were combined, the solvents were removed under reduced pressure to afford **D-12**.

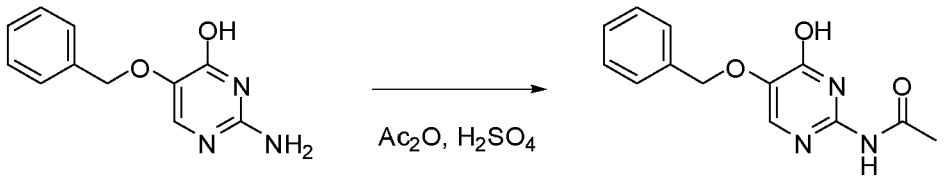
LC-MS: Anal. Calcd. For $C_{23}H_{34}N_6O_5$: 474.26; found 475 $[M+H]^+$



D-12 **78**

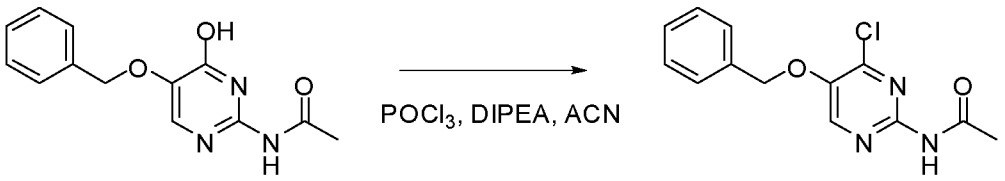
Step 2. D-11 (620 mg, 1.31 mmol) was dissolved in dichloromethane (10 mL) and to it was added HCl (6 M in isopropanol, 10 mL). The reaction mixture stirred for 15 hours at room temperature. The volatiles were removed under reduced pressure and the residue was purified by reversed phase chromatography to afford **78**.

Preparation of **79**



B-1 B-5

Step 1. In a 500 mL flask a mixture of B-1 (30 g, 138 mmol) and sulfuric acid (3 mL) in acetic anhydride (300 mL) was heated to 90°C for 3 hours. The reaction cooled to room temperature and the precipitate was isolated by filtration, washed with diisopropylether and dried in vacuo at 50°C to obtain a white solid, **B-5**.

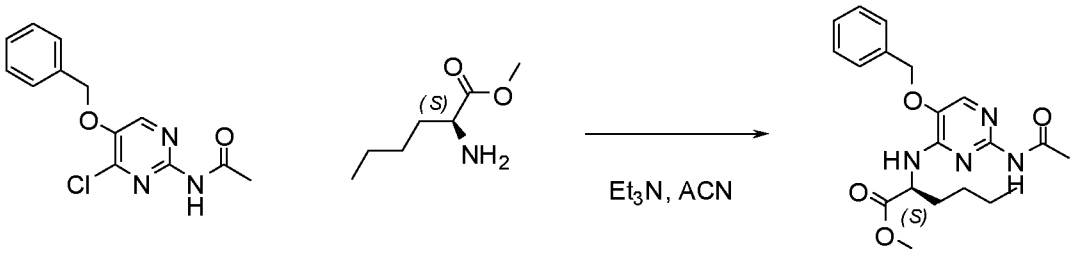


B-5 C-2

Step 2. In a 400 mL multimax reactor a mixture of B-5 (21.8 g, 84 mmol) in acetonitrile (244 mL) was stirred at 30°C under a gentle stream of nitrogen. Phosphoryl chloride (18.14 mL, 195 mmol) was added dropwise over a period of 5 minutes. After addition, the reaction mixture was heated to 45°C and the mixture was stirred for 15 minutes, then DIPEA (33 mL, 195 mmol) was added slowly over a period of 1.5 hours. The reaction was stirred at 45°C until completion (monitored by LC-MS). A solution of sodium ethanoate (65 g) in water (732 mL) was heated in a 2 L flask to 35°C and the reaction mixture was portioned into this solution over a period of 5 minutes. The temperature is kept between 35-40°C via an external cooling bath. The mixture was allowed to reach ambient temperature and stirring was continued for 1 hour. The precipitate was isolated by filtration, washed with water and dried in vacuo at 50°C to obtain **C-2** as a solid.

LC-MS: Anal. Calcd. For $C_{13}H_{12}ClN_3O_2$: 277.06; found 278 $[M+H]^+$

^1H NMR (400 MHz, DMSO-d_6) δ ppm 2.11 (s, 3 H), 5.31 (s, 2 H), 7.33 - 7.39 (m, 1 H), 7.43 (t, *J*=7.2 Hz, 2 H), 7.46 - 7.51 (m, 2 H), 8.59 (s, 1 H), 10.65 (s, 1 H)

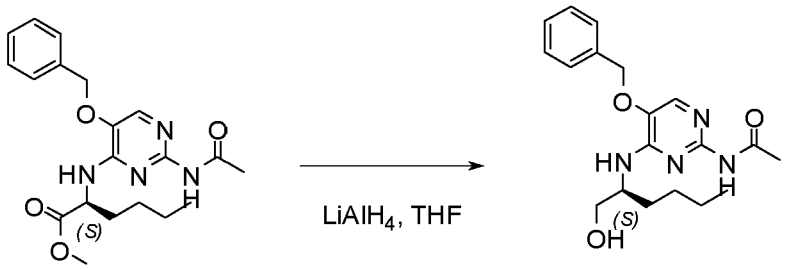


C-2 **D-13**

Step 3. A solution of intermediate C-2 (5.9 g, 21.2 mmol), methyl (2S)-2-aminohexanoate (5.79 g, 31.9 mmol) and triethylamine (14.8 mL, 106 mmol) in acetonitrile (100 mL) was heated to reflux for 4 days. The reaction mixture was cooled to room temperature and the solvent was removed under reduced pressure. The residue was dissolved in dichloromethane and washed with

brine. The organic layer was dried (magnesium sulfate) then purified directly via silica column using a gradient of dichloromethane to 10% methanol in dichloromethane. The best fractions were pooled and the solvents were removed under reduced pressure to afford **D-13**.

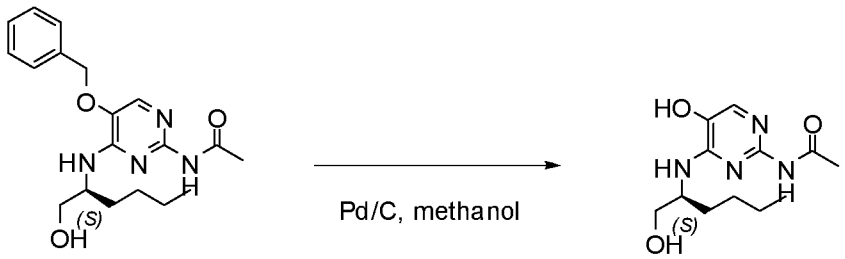
LC-MS: Anal. Calcd. For $C_{20}H_{26}N_4O_4$: 386.20; found 387 $[M+H]^+$



D-13 **D-14**

Step 2. D-13 (3.7 g, 9.57 mmol) was dissolved in anhydrous THF (100 mL). Lithium aluminum hydride (1M in THF, 9.6 mL, 9.6 mmol) was added dropwise and the reaction mixture stirred for 3 hours at room temperature. NH_4Cl (sat., aq.) was added drop wise to the reaction mixture and the precipitated salts were removed via filtration and washed with THF. The filtrate was evaporated to dryness and the residue was purified via silica gel column chromatography using a dichloromethane to 10% methanol in dichloromethane gradient. The best fractions were combined and the solvents were removed under reduced pressure to afford **D-14**.

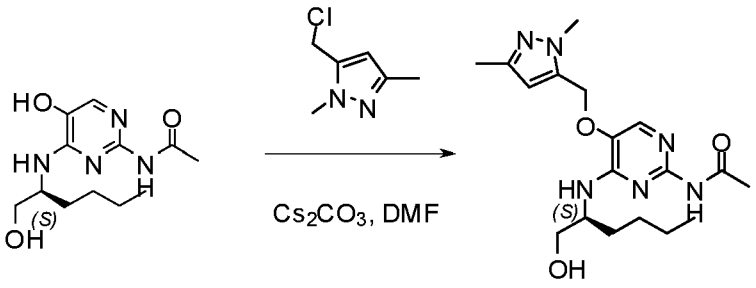
LC-MS: Anal. Calcd. For $C_{19}H_{26}N_4O_3$: 358.20; found 359 $[M+H]^+$



D-14 **D-15**

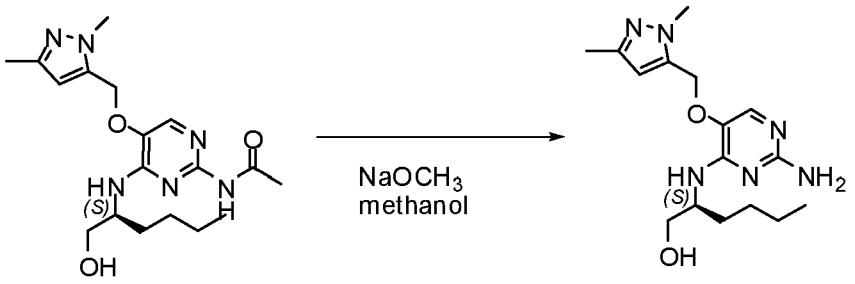
Step 3. **D-15** was prepared according to the method described for intermediate D-2. Used without purification in the next step.

LC-MS: Anal. Calcd. For $C_{12}H_{20}N_4O_3$: 268.15; found 269 $[M+H]^+$



D-15 **D-16**

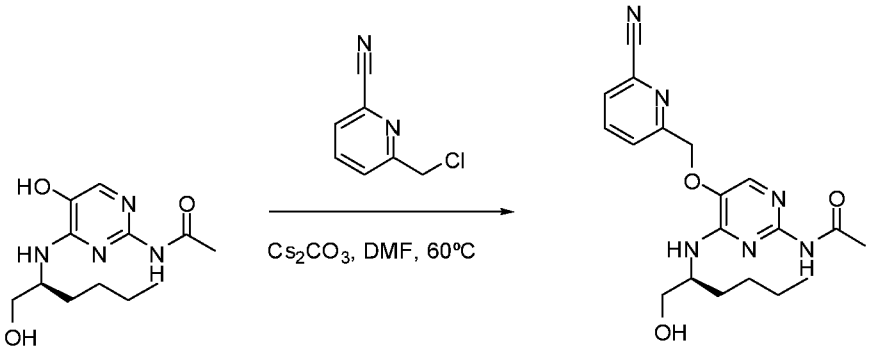
Step 4. A mixture of D-15 (210 mg, 0.78 mmol) and cesium carbonate (765 mg, 2.35 mmol) in DMF (25 mL) was heated to 60°C with stirring then a solution of 5-(chloromethyl)-1,3-dimethyl-1H-pyrazole (113 mg, 0.78 mmol) in DMF(10 mL) was added drop wise. The reaction mixture was stirred for 1 hour at 60°C. The solids were removed by filtration and the solvent was removed under reduced pressure. Crude **D-16** was used as such in the next step.
LC-MS: Anal. Calcd. For $C_{18}H_{28}N_6O_3$: 376.22; found 377 $[M+H]^+$



D-16 **79**

Step 5. Into a 30 mL glass tube was placed D-16 (295 mg, 0.78 mmol) and $NaOCH_3$ (30% in methanol, 2 mL) and methanol (20 mL) and the mixture was stirred at 60°C overnight. The reaction mixture was purified via reverse phase liquid chromatography (Sunfire Prep C18 OBD 10mm, 30 x 150 mm. Mobile phase 0.25% NH_4OAc solution in water, methanol) to afford **79** as the free base.

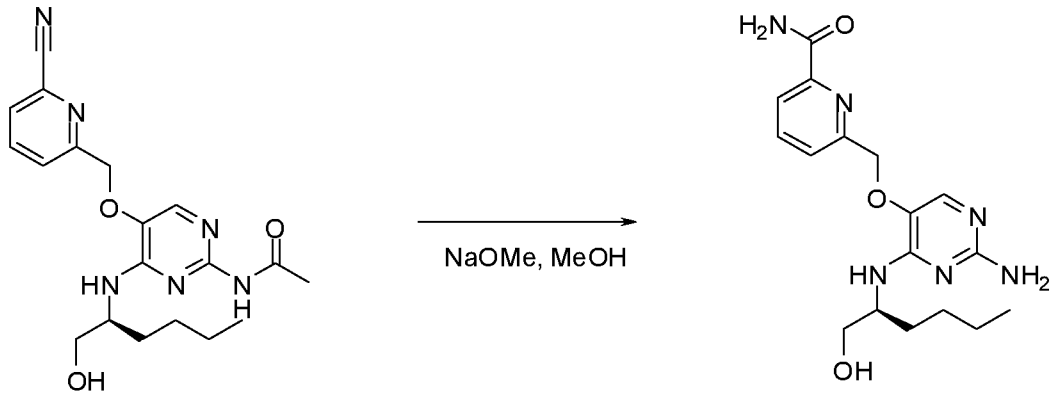
Preparation of **80**



D-15 **D-17**

Step 1. Intermediate **D-17** was prepared according to the method used for D-16 via alkylation of D-15.

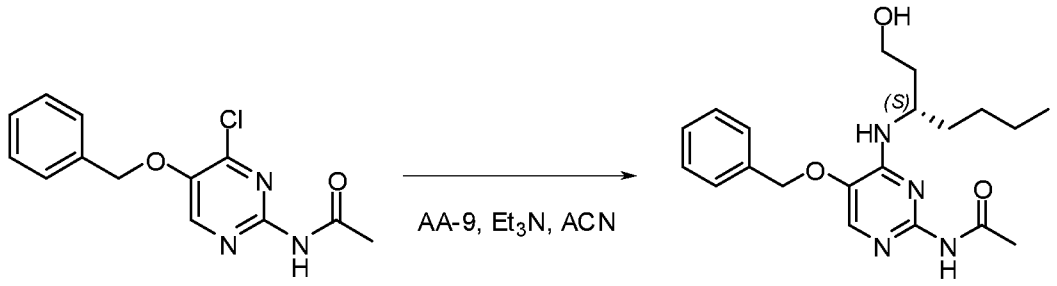
LC-MS: Anal. Calcd. For $C_{19}H_{24}N_6O_3$: 384.19; found 385 $[M+H]^+$



D-17 **80**

Step 2. In a 30 mL glass tube D-17 (301 mg, 0.78 mmol) and $NaOCH_3$ (30% in methanol, 2 mL) were dissolved in methanol (20 mL) and stirred at 60°C overnight. 10 mL of water was added to the reaction mixture and it was stirred for 2 hours at 60°C. The reaction mixture was purified via reverse phase liquid chromatography (Sunfire Prep C18 OBD 10mm, 30 x 150 mm. Mobile phase 0.25% NH_4OAc solution in water, methanol) yielding **80** as a powder.

Preparation of **81**

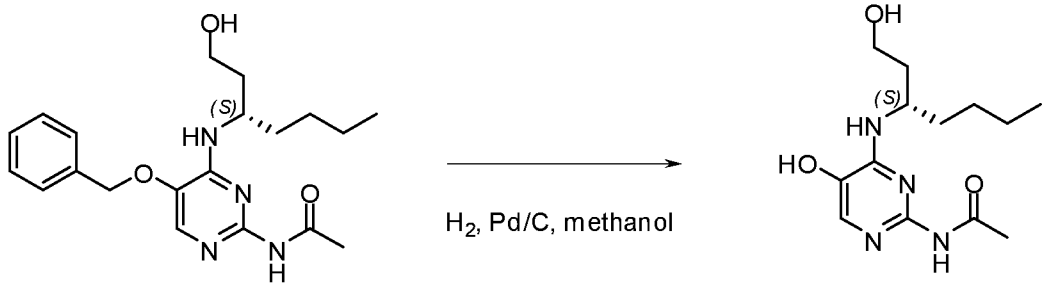


C-2 **D-18**

A solution of intermediate C-2 (2 g, 7.2 mmol), AA-9 (3.02 g, 18 mmol) and triethylamine (5 mL, 36 mmol) in acetonitrile (75 mL) was heated to reflux for 6 hours. The reaction mixture was cooled down and the solvent was removed under reduced pressure. The residue was dissolved in dichloromethane and washed with brine. The organic layer was loaded on a silica cartridge and a gradient of dichloromethane to 10% methanol in dichloromethane was applied. The fractions containing the product were evaporated to dryness yielding a white powder, **D-18**.

LC-MS: Anal. Calcd. For $C_{20}H_{28}N_4O_3$: 372.22; found 373 $[M+H]^+$

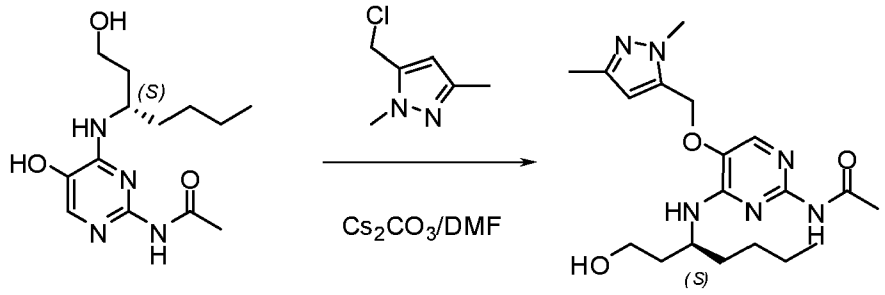
^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.77 - 0.92 (m, 3 H) 1.15 - 1.36 (m, 4 H) 1.42 - 1.72 (m, 4 H) 2.12 (s, 3 H) 3.35 - 3.42 (m, 2 H) 4.11 - 4.24 (m, 1 H) 4.35 - 4.52 (m, 1 H) 6.42 (d, *J*=8.80 Hz, 1 H) 7.42 (s, 1 H) 9.63 (br. s., 1 H)



D-18 **D-19**

D-19 was prepared from D-18 according to the method employed for intermediate D-2.

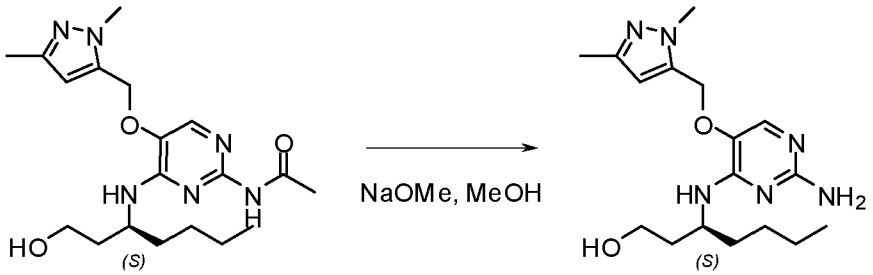
LC-MS: Anal. Calcd. For $C_{13}H_{22}N_4O_3$: 282.1; found 283 $[M+H]^+$



D-19 **D-20**

D-20 was prepared from D-19 according to the method to prepare D-17.

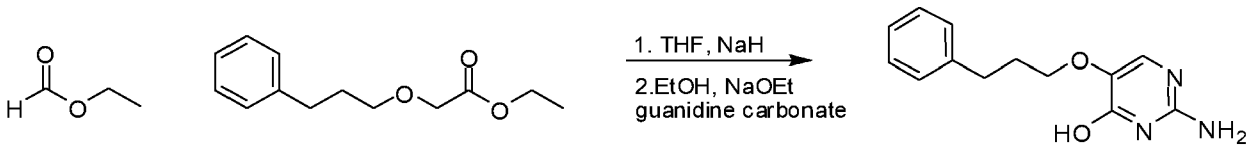
LC-MS: Anal. Calcd. For $C_{19}H_{30}N_6O_3$: 390.24; found 391 $[M+H]^+$



D-20 **81**

81 was prepared from D-20 according to the method to prepare compound 79.

Preparation of **82**

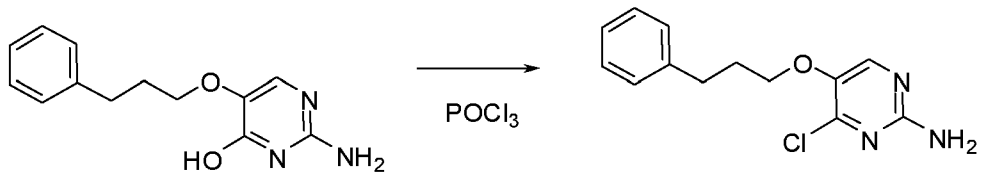


B-3

Step 1. Intermediate **B-3** was prepared according to the method described for B-1.

LC-MS: Anal. Calcd. For $C_{13}H_{15}N_3O_2$: 245.12; found 246 $[M+H]^+$

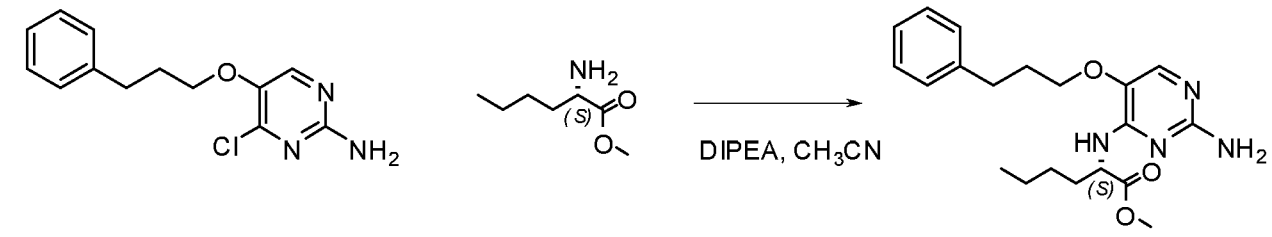
^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 1.79 - 1.93 (m, 2 H), 2.66 (t, J=7.8 Hz, 2 H), 3.76 (t, J=6.4 Hz, 2 H), 6.54 (br. s., 2 H), 7.11 - 7.21 (m, 3 H), 7.22 - 7.29 (m, 3 H), 11.46 (br. s, 1 H)



B-3 **C-3**

Step 2. In a 250 mL round bottom flask a mixture of B-3 (15 g, 61.15 mmol) in $POCl_3$ (150 mL) was heated to reflux and stirred for 2 hours. The reaction was allowed to cool and the solvent was removed under reduced pressure. The residual fraction was triturated with diisopropylether. The formed precipitate isolated by filtration, washed with diisopropylether and dried under vacuo at 50°C to obtain a solid, **C-3**, used as such in the next step.

LC-MS: Anal. Calcd. For $C_{13}H_{14}ClN_3O$: 263.08; found 264 $[M+H]^+$

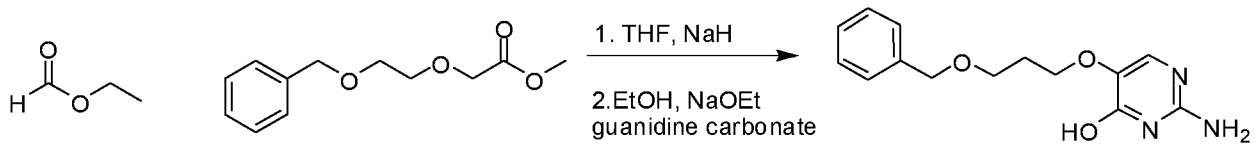


C-3 **82**

Step 3. Into a 20 mL tube was placed C-3 (0.45 g, 1.05 mmol), L-2-aminohexanoic acid-methyl ester HCl (0.48 g, 2.62 mmol), DIPEA (1.18 mL, 6.82 mmol), and acetonitrile (5 mL). The tube was sealed and heated in the microwave for 1.5 hours at 120°C. The reaction was allowed to cool and the solvent was removed under reduced pressure.

The crude mixture was purified by Prep HPLC on (RP Vydac Denali C18 - 10 µm, 250 g, 5 cm). Mobile phase (0.25% NH_4OAc solution in water, methanol), the desired fractions were collected and evaporated to dryness. The residual fraction was dissolved in a mixture of dichloromethane/methanol and poured over a acid modified solid phase extraction cartridge (SCX). The product was released using with NH_3 7N in methanol. The collected solution was concentrated under reduced pressure to obtain the desired solid, **82**.

Preparation of **83**

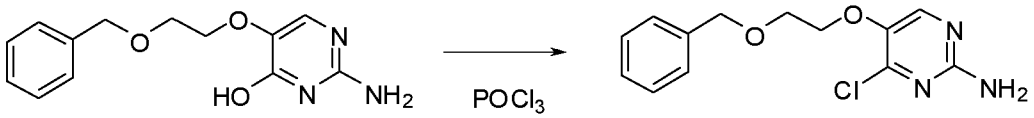


B-4

Step 1. Intermediate **B-4** was prepared according to the method to prepare B-1.

LC-MS: Anal. Calcd. For $C_{14}H_{17}N_3O_3$: 275.13; found 276 $[M+H]^+$

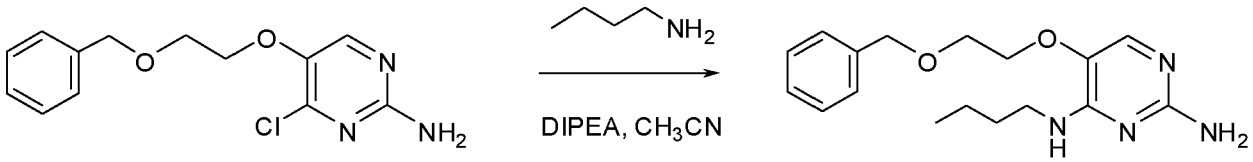
^1H NMR (400 MHz, DMSO-d_6) δ ppm 3.63 (dd, J=5.4, 3.9 Hz, 2 H), 3.95 (dd, J=5.4, 3.6 Hz, 2 H), 4.50 (s, 2 H), 6.33 (br. s., 2 H), 7.22 - 7.29 (m, 2 H), 7.30 - 7.36 (m, 4 H), 10.71 - 11.58 (m, 1 H)



B-4 **C-4**

Step 2. Into a 250 mL round bottom flask was placed B-4 (10 g, 38.27 mmol) and $POCl_3$ (75 mL). The mixture was heated to reflux and stirred for 5 hours. The reaction mixture was allowed to reach room temperature and stirred for 15 hours. The solvent was removed under reduced pressure. Crude **C-4** was used as such in the next step.

LC-MS: Anal. Calcd. For $C_{12}H_{12}ClN_3O_2$: 265.06; found 266 $[M+H]^+$

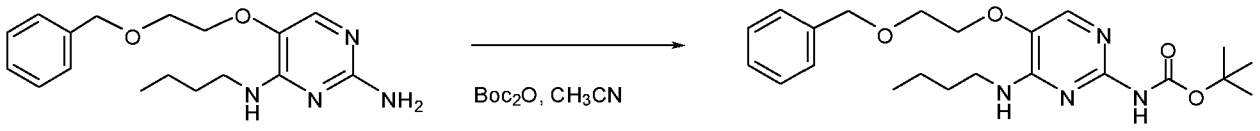


C-4 **83**

Step 3. Into a 50 mL tubes was placed C-4 (10 g, 35.75 mmol), *n*-butylamine (10.6 mL, 107.25 mmol) and DIPEA (30.8 mL, 178.75 mmol) in acetonitrile (40 mL). The mixture was heated to 120°C under microwave irradiation for 3 hours. The combined reaction mixtures were concentrated under reduced

pressure and the residual oil was dissolved in dichloromethane and washed with 1N HCl and water. The organic layer was dried (magnesium sulfate), the solids were removed by filtration and the solvent of the filtrate were removed under reduced pressure to obtain an red-brown foam, **83**.

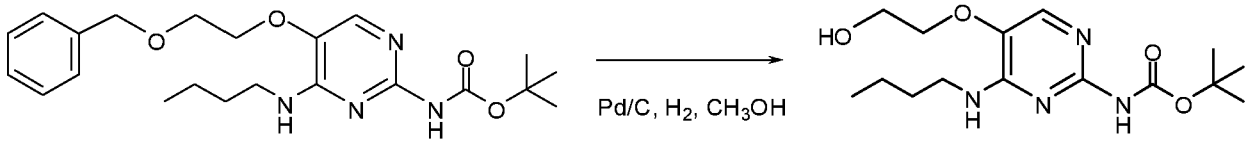
Preparation of **84**



83 D-20

Step 1. Into a 500 mL round bottom flask was placed 83 (13.5 g, 25.6 mmol), Boc-anhydride (27.94 g, 128 mmol) and acetonitrile (150 mL). The yellow solution was stirred at reflux for 16 hours. The solvent was removed under reduced pressure. The residual fraction was dissolved in dichloromethane and washed with a saturated aqueous $NaHCO_3$ solution and water. The organic layer was dried (magnesium sulfate), the solids were removed via filtration, and the solvents of the filtrate were removed under reduced pressure to obtain an oil, **D-20**.

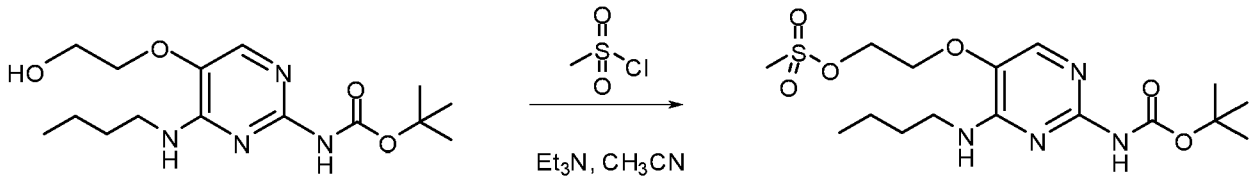
LC-MS: Anal. Calcd. For $C_{22}H_{32}N_4O_4$: 416.24; found 417 $[M+H]^+$



D-20 D-21

Step 2. Into a 1L erlenmeyer was suspended 10% Pd/C (4 g) in methanol (350 mL) under N_2 gas flow, then D-20 (14.3 g, 34.33 mmol) was added. The mixture was stirred at 50°C under a hydrogen atmosphere until 1 equivalent of hydrogen was absorbed. The catalyst was removed by filtration over packed decalite. The solvent of the filtrate was removed under reduced pressure to obtain an oil, **D-21**. The residue was used as such in the next step.

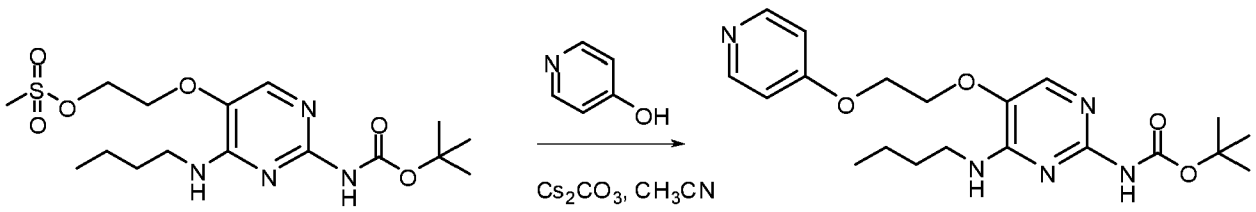
LC-MS: Anal. Calcd. For $C_{15}H_{26}N_4O_4$: 326.20; found 327 $[M+H]^+$



D-21 D-22

Step 3. Into a 1L round bottom flask a solution of D-21 (8.7 g, 26.66 mmol) and triethylamine (7.41 mL, 53.31 mmol) in acetonitrile (300 mL) was stirred at ambient temperature and methanesulfonyl chloride (3.1 mL, 40 mmol) was added. After addition, the reaction mixture was stirred for 1.5 hours at room temperature. The solvent was removed under reduced pressure. The crude was dissolved in ethyl acetate and washed with saturated aqueous $NaHCO_3$. The organic layers were combined, dried (magnesium sulfate), the solids were removed by filtration and the solvent of the filtrate were evaporated to dryness to obtain **D-22** as an oil.

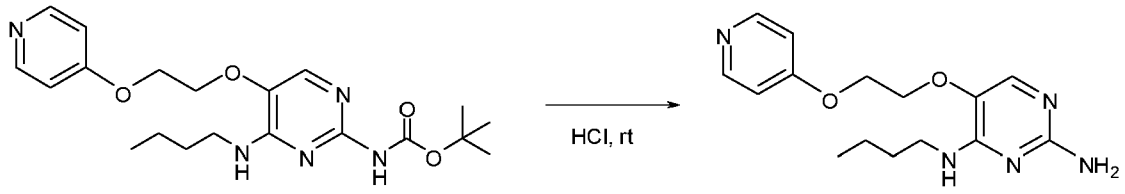
LC-MS: Anal. Calcd. For $C_{16}H_{28}N_4O_6S$: 404.17; found 405 $[M+H]^+$



D-22 D-23

Step 4. Into a 30 mL glass tube was placed a mixture of 4-hydroxypiridine (94 mg, 0.99 mmol) and Cs_2CO_3 (0.8 g, 2.47 mmol) in acetonitrile (10 mL). The vial was sealed and shaken at ambient temperature for 1 hour. D-22 (400 mg, 0.99 mmol) as a solution in acetonitrile (10 mL) was added to the reaction mixture and shaken for an additional 18 hours at room temperature. Cesium carbonate (320 mg, 1 mmol) was added and the mixture was shaken for 1 day at room temperature. The solvent was removed under reduced pressure and the crude was treated with a mixture of dichloromethane/methanol, 95/5 and shaken for 1h, then filtered over 2 g of packed silica. The filtrate was concentrated under reduced pressure and **D-23** was used as such in the next step.

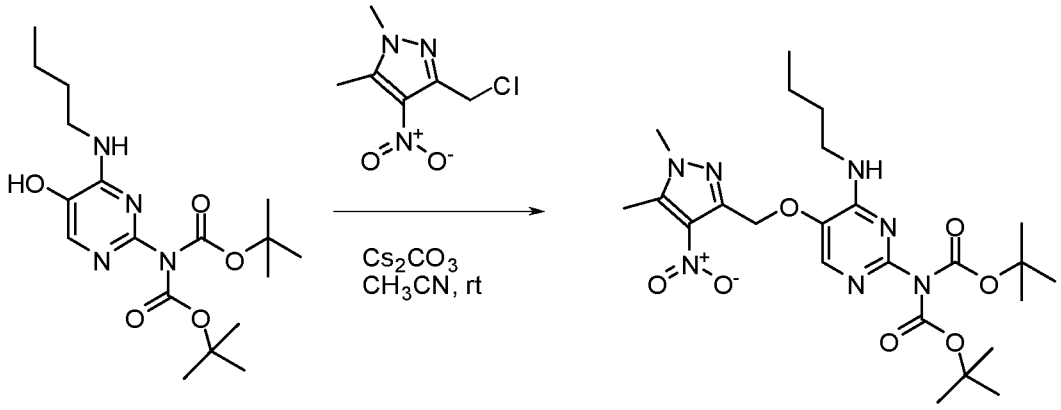
LC-MS: Anal. Calcd. For $C_{20}H_{29}N_5O_4$: 403.22; found 404 $[M+H]^+$



D-23 84

Step 5. D-23 was deprotected to afford **84** using the method applied to deprotect 78.

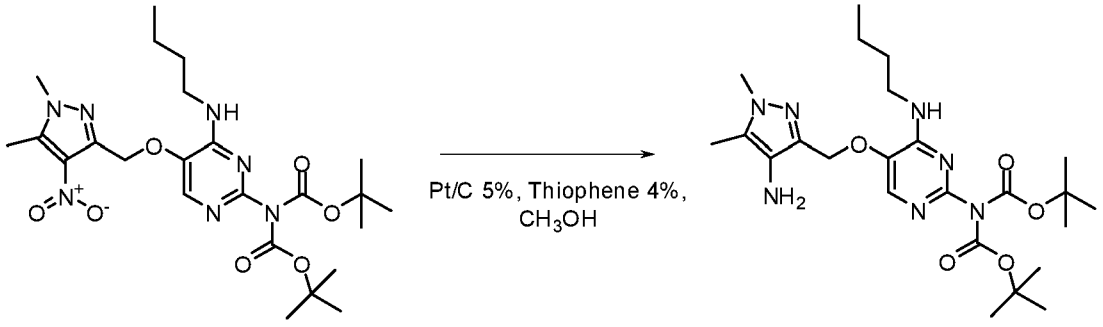
Preparation of **85**



D-4 D-24

Step 1. Into a 250 mL round bottem flask equipped with a magnetic stir bar was placed D-4 (0.35 g, 5.23 mmol) and cesium carbonate (0.89 g, 2.75 mmol) in acetonitrile (20 mL). The mixture was stirred at ambient temperature for 30 minutes. A solution of the alkyl halide (0.19 g, 1 mmol) in acetonitrile (5mL) was added and the reaction mixture was stirred for 1 day at room temperature. The reaction was completed and the salts were removed by filtration. The filtrate was concentrated under reduced pressure and the crude was purified by silica column chromatography using a heptane to ethyl acetate gradient to afford intermediate **D-24**.

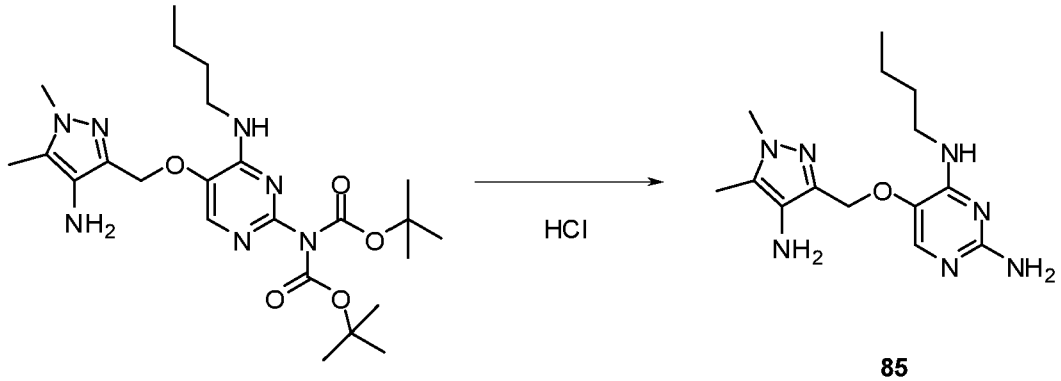
LC-MS: Anal. Calcd. For $C_{24}H_{37}N_7O_7$: 535.28; found 536 $[M+H]^+$



D-24 D-25

Step 2. Into a 100 mL erlenmeyer flask was suspended Pt/C, 5% (100 mg) in thiophene (0.25 mL) and methanol (20 mL) under a blanket of nitrogen gas, then D-24 (130 mg, 0.24 mmol) was added. The reaction mixture was stirred at 50°C under a hydrogen atmosphere. The catalyst was removed by filtration over packed decalite. The solvents of the filtrate were removed under reduce pressure to obtain **D-25** as an oil, that was used as such in the next step.

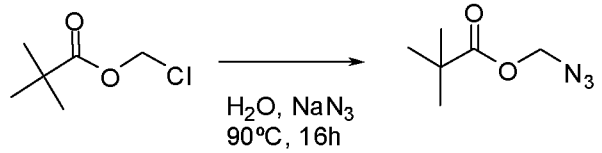
LC-MS: Anal. Calcd. For $C_{24}H_{39}N_7O_5$: 505.30; found 506 $[M+H]^+$



D-25 85

Step 3. Intermediate D-25 is deprotected to afford **85** according to the method used to prepare 78.

Preparation of **86**



A-2

Step 1. Into a 100 mL round bottom flask was placed sodium azide (6.85 g, 103.76 mmol) in water (12.5 mL) then chloromethyl pivalate (10.6 g,

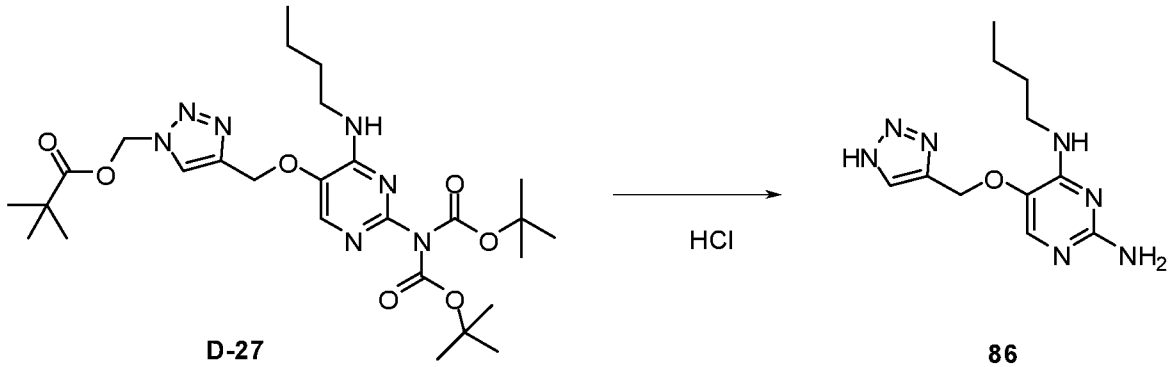
70.38 mmol) and stirred vigorously at 90°C for 16 hours. The reaction mixture was allowed to cool to room temperature and dichloromethane (20 mL) was added. The organic layer was separated, dried over anhydrous sodium sulfate, the solids were removed by filtration and the solvent of the filtrate was removed under reduced pressure to obtain **A-2** as an oil.

LC-MS: Anal. Calcd. For $C_6H_{11}N_3O_2$: 157.09; found 158 $[M+H]^+$

D-26 D-27

Step 2. Into a 25 mL tube was placed of D-26 (100 mg, 0.238 mmol), A-2 (37.9 mg, 0.238 mmol), *t*-butanol (2.5 mL) and water (2.5 mL). The tube was sealed and the mixture was stirred at ambient temperature. Copper(II) sulfate pentahydrate (3 mg, 0.012 mmol) and L-ascorbic acid sodium salt (15.5 mg, 0.079 mmol) were added. The reaction mixture was stirred for 18 hours at room temperature, then water (2.5 mL) was added. The precipitate was isolated by filtration, washed with water and dried in vacuo at 60°C to obtain a white powder, **D-27**.

LC-MS: Anal. Calcd. For $C_{27}H_{43}N_7O_7$: 577.32; found 578 $[M+H]^+$



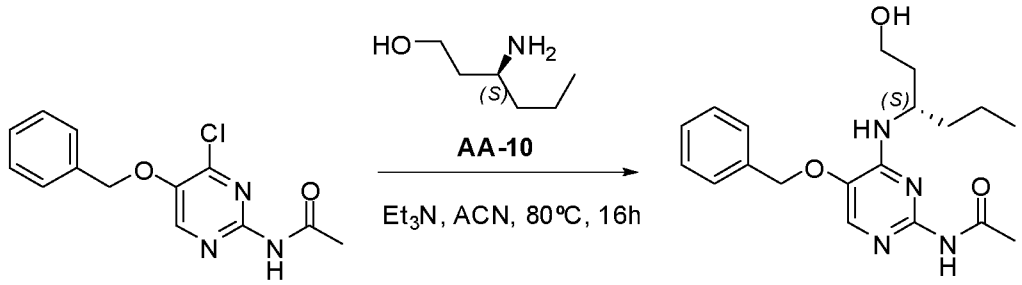
D-27 86

Step 3. In a 100 mL round bottom flask a mixture of D-27 (0.1 g, 0.17 mmol) in HCl (5 mL 6M in isopropanol) and dichloromethane (5 mL) was stirred at

ambient temperature for 16 hours. The reaction was heated to 65°C and stirred for an additional 16 hours. The solvent was removed under reduced pressure.

The crude product was purified by reverse phase liquid chromatography (RP Vydac Denali C18 - 10µm, 250 g, 5 cm). Mobile phase (0.25% NH_4HCO_3 solution in water, methanol), the desired fractions were collected, evaporated, dissolved in methanol and treated with 2M HCl in ether. The solid was isolated by filtration to afford **86** as the HCl salt.

Preparation of **87**

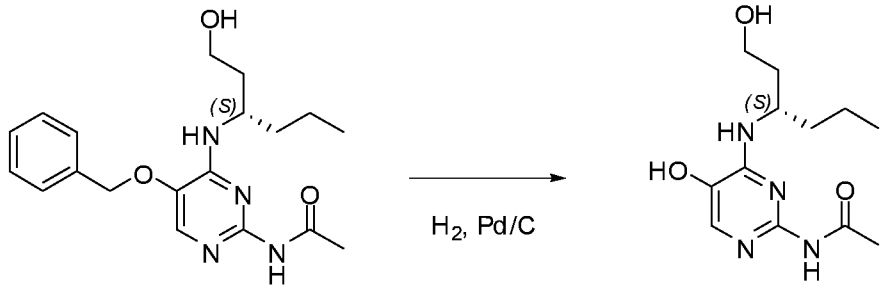


C-2 **D-28**

Step 1. Into a 100 mL round bottom flask was placed a solution of C-2 (500 mg, 1.8 mmol), AA-10 (692 mg, 4.5 mmol) and triethylamine (0.75 mL, 5.4 mmol) in acetonitrile (30 mL). The mixture was heated to 80°C for 16 hours with stirring. The reaction was allowed to cool and the solvent was removed under reduced pressure. The crude was dissolved in dichloromethane and washed with brine. The organic layer was dried (magnesium sulfate), the solids were removed by filtration and the solvent of the filtrate was removed to obtain an oil, **D-28**.

LC-MS: Anal. Calcd. For $C_{19}H_{26}N_4O_3$: 358.20; found 359 $[M+H]^+$

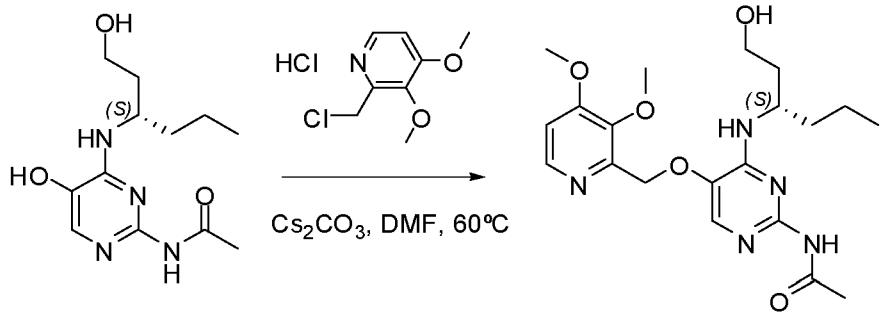
^1H NMR (360 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.32 Hz, 3 H) 1.19 - 1.37 (m, 2 H) 1.38 - 1.53 (m, 1 H) 1.53 - 1.75 (m, 3 H) 2.13 (s, 3 H) 3.38 - 3.48 (m, 2 H) 4.19 - 4.31 (m, 1 H) 5.16 (s, 2 H) 6.69 (d, J=9.15 Hz, 1 H) 7.29 - 7.41 (m, 3 H) 7.45 - 7.53 (m, 2 H) 7.66 (s, 1 H) 9.77 (s, 1 H)



D-28 **D-29**

Step2. **D-29** was prepared according to the method used to prepare D-21. THF was added to increase the solubility of **D-29**.

LC-MS: Anal. Calcd. For $C_{12}H_{20}N_4O_3$: 268.15; found 269 $[M+H]^+$



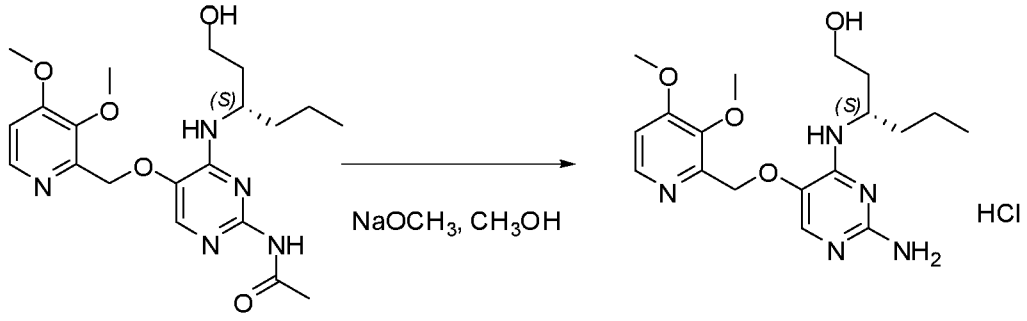
D-29 **D-30**

Step 3. In a 250 mL round bottom flask a mixture of D-29 (5 g, 18.6 mmol) and cesium carbonate (18.2 g, 55.9 mmol) in DMF (80 mL) was stirred at ambient temperature for 30 minutes. The mixture was heated to 60°C and a solution of 2-chloromethyl-3,4-dimethoxy pyridine hydrochloride (3.97 g, 17.7 mmol) in DMF (60 mL) was added dropwise. The reaction mixture was stirred for 2 hours at 60°C. The reaction was allowed to cool and the salts were removed by filtration. The reaction mixture was concentrated under reduced pressure and **D-30** was used as such in the next step.

LC-MS: Anal. Calcd. For $C_{20}H_{29}N_5O_5$: 419.22; found 420 $[M+H]^+$

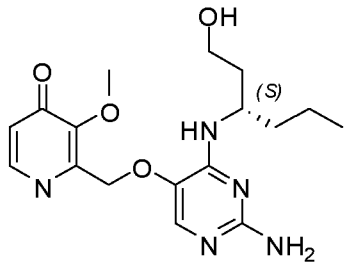
^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.83 (t, J=7.4 Hz, 3 H), 1.18 - 1.32 (m, 2 H), 1.41 - 1.71 (m, 4 H), 2.14 (s, 3 H), 3.34 - 3.40 (m, 2 H), 3.78 (s, 3 H), 3.91 (s, 3 H), 4.17 - 4.29 (m, 1 H), 4.41 (t, J=5.3 Hz, 1 H), 5.09 (s, 2 H), 6.79 (d,

J=8.8 Hz, 1 H), 7.15 (d, *J*=5.7 Hz, 1 H), 7.75 (s, 1 H), 8.24 (d, *J*=5.5 Hz, 1 H), 9.75 (s, 1 H)



D-30 **87**

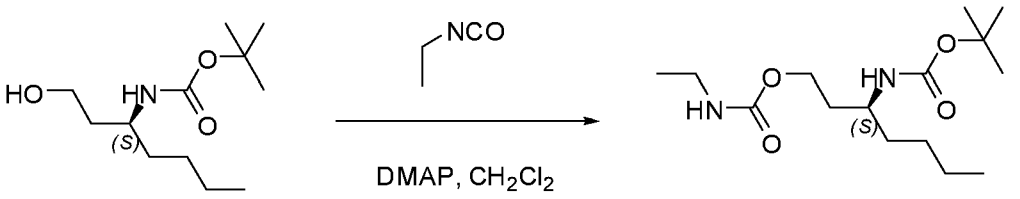
Step 4. **87** was prepared according to the same method used to prepare 79 from intermediate D-16. 87 was purified by reverse phase chromatography (Hyperprep C18 HS BDS. Mobile phase (Gradient from 90% ammonium bicarbonate in water 0.25%, 10% acetonitrile to 0% ammonium bicarbonate in water 0.25%, 100% acetonitrile). The best fractions were pooled, the solvents were removed under reduced pressure, reconstituted in methanol and treated with 2M HCl in ether and then concentrated under reduced pressure to obtain a white solid, the HCl salt of **87**.



88

Isolation of the HCl salt of 87 via reverse phase liquid chromatography led to the concomitant isolation of **88** in low yield. The best fractions were pooled, and the solvents were removed under reduced pressure to afford a white solid, **88**.

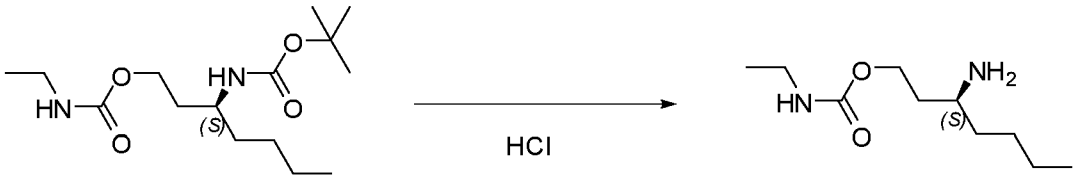
Preparation of **89**



AA-8 **AA-12**

Step 1. Into a 100 mL round bottom flask was placed AA-8 (2 g, 8.65 mmol), dichloromethane (6 mL), ethyl isocyanate (1.6 mL, 10.38 mmol), and DMAP (21 mg, 0.173 mmol). The reaction mixture was allowed to stir for 16 hours at room temperature. The solvent was removed under reduced pressure and **AA-12** was used in the subsequent step without further purification.

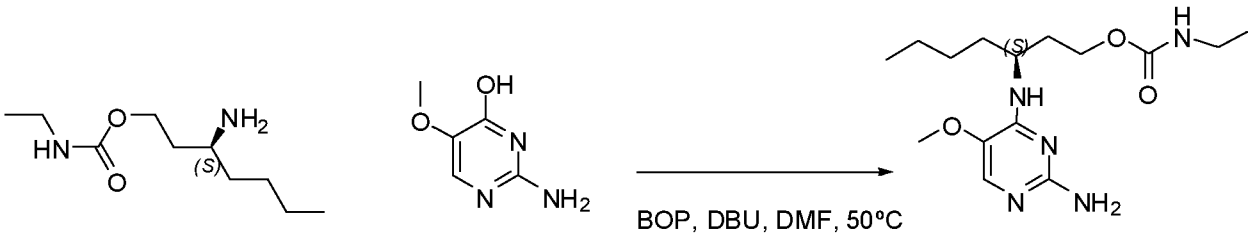
LC-MS: Anal. Calcd. For $C_{15}H_{30}N_2O_4$: 302.22; found 303 $[M+H]^+$



AA-12 **AA-13**

Step 2. Into a 100 mL round bottom flask was placed crude AA-12 (2.61 g, 8.65 mmol), and dichloromethane (30 mL). To this solution was added HCl (20 mL, 4M in dioxane). The reation was allowed to stir 3 hours at room temperature.

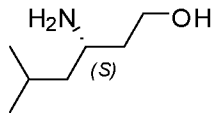
LC-MS: Anal. Calcd. For $C_{10}H_{22}N_2O_2$: 202.17; found 203 $[M+H]^+$



AA-13 4763-35-3 **89**

Step 3. Into a 100 mL round bottom flask equipped with a magnetic stir bar was placed 2-Amino-4-hydroxy-5-methoxy-pyrimidine (500 mg, 3.54 mmol), anhydrous DMF (30 mL), AA-13 (1.27 g, 5.31 mmol), DBU (2.12 mL, 14.17 mmol), and BOP (1.96 g, 4.43 mmol). The reaction mixture was allowed to stir at room temperature for 30 minutes then at 50ºC for 16 hours. The solvent was removed under reduced pressure and the residue was partitioned between brine and ethyl acetate. The organic layers were combined, dried (magnesium sulfate), the solids were removed via filtration, and the solvents of the filtrate were removed under reduced pressure. The crude was purified via reverse phase liquid chromatography (RP Vydac Denali C18 - 10µm, 250 g, 5 cm. Mobile phase 0.25% NH_4HCO_3 solution in water, methanol), the best fractions were pooled, the solvents were removed under reduced pressure to afford **89**.

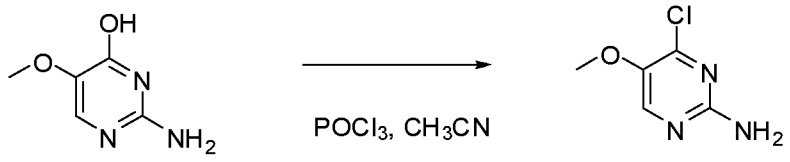
Preparation of **264**



AA-14

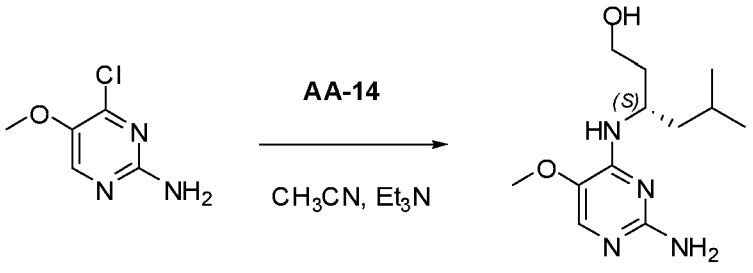
Step 1. **AA-14** was prepared according to the procedure to prepare AA-10, employing the appropriate starting aldehyde.
LC-MS: Anal. Calcd. For $C_7H_{17}NO$: 131.13; found 132 $[M+H]^+$

^1H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.81 - 0.89 (m, 6 H), 1.15 - 1.25 (m, 2 H), 1.33 - 1.47 (m, 1 H), 1.54 - 1.69 (m, 2 H), 2.71 (br. s., 3 H), 2.88 - 2.98 (m, 1 H), 3.69 - 3.80 (m, 2 H)



4763-35-3 C-5

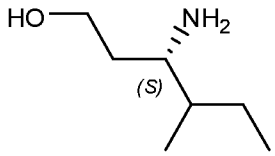
Step 2. **C-5** was prepared according to the method used to prepare C-2 from the available starting material. The crude was used without further purification.
LC-MS: Anal. Calcd. For $C_5H_6ClN_3O$: 159.02; found 160 $[M+H]^+$



C-5 **264**

Step 3. C-5 was combined with AA-14 according to the method used to prepare compound 1, except that acetonitrile was used as a solvent, to afford **264**.

Preparation of **278**

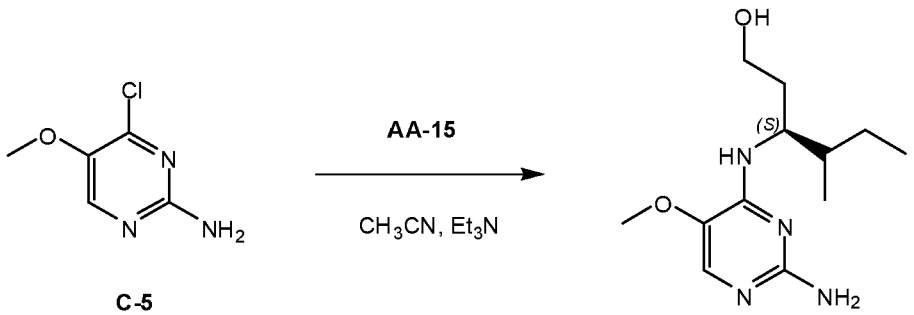


AA-15

Step 1. **AA-15** was prepared according to the procedure to prepare AA-10, employing the appropriate starting aldehyde.

LC-MS: Anal. Calcd. For $C_7H_{17}NO$: 131.13; found 132 $[M+H]^+$

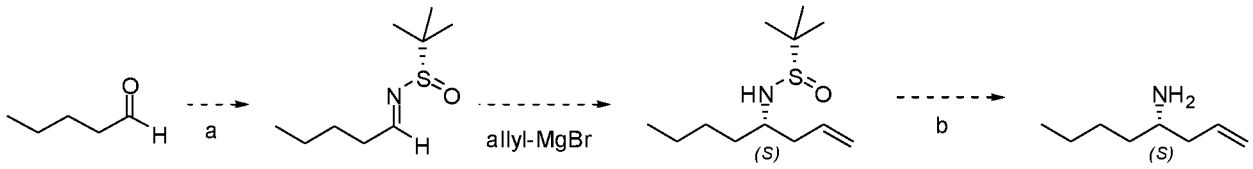
^1H NMR (400 MHz, DMSO-d_6) δ ppm 0.81 - 0.89 (m, 6 H), 1.05 - 1.20 (m, 1 H), 1.27 - 1.40 (m, 1 H), 1.43 - 1.77 (m, 3 H), 3.05 - 3.19 (m, 1 H), 3.44 - 3.57 (m, 2 H), 4.82 (br. s., 1 H), 7.94 (d, J=18.6 Hz, 2 H)



C-5 **278**

Step 2. C-5 was combined with AA-15 according to the method used to prepare compound 1, except that acetonitrile was used as a solvent, to afford **278.**

Preparation of **295**

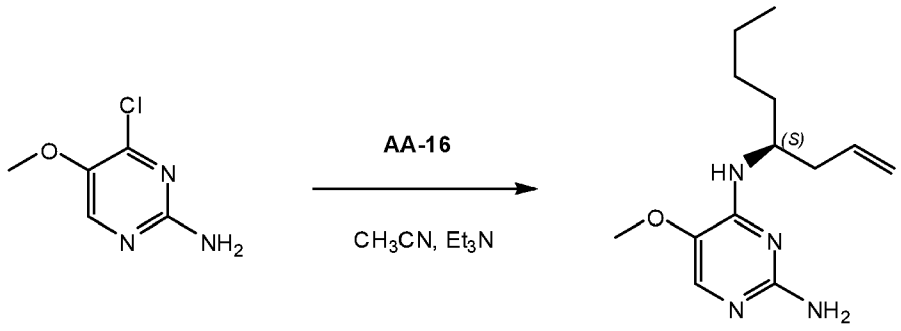


valeraldehyde
110-62-3

AA-16

a) MgSO4 (5 equiv), PPTS (0.05 equiv), CH_2Cl_2, rt b) HCl in ether, filter off byproduct

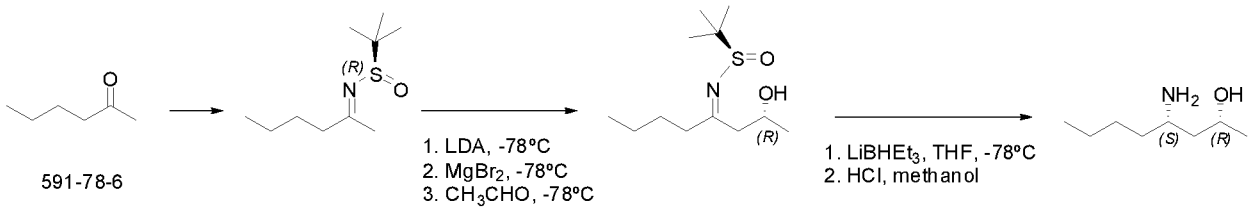
Step 1. **AA-16** was prepared according to the procedures outlined in Chem. Rev., 2010, Vol. 110, No. 6, 3600-3740.
LC-MS: Anal. Calcd. For $C_8H_{17}N$: 127.14; found 128 $[M+H]^+$



C-5 **278**

Step 2. C-5 was combined with AA-16 according to the method used to prepare compound 1, except that acetonitrile was used as a solvent, to afford **295.**

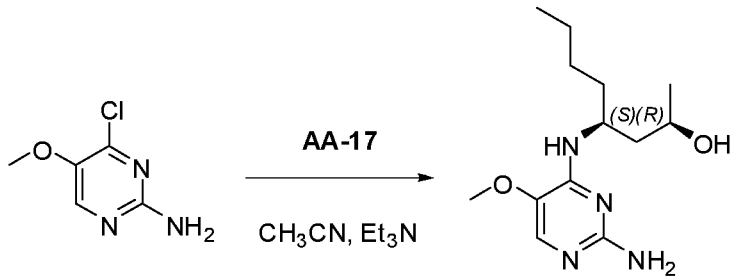
Preparation of **304**



591-78-6

AA-17

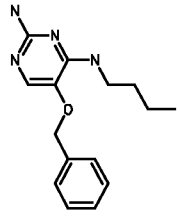
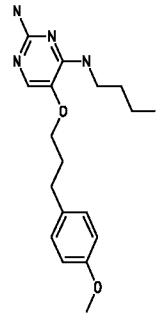
Step 1. **AA-17** was prepared according to the procedures outlined in Chem. Rev., 2010, Vol. 110, No. 6, 3600-3740.
LC-MS: Anal. Calcd. For $C_8H_{19}NO$: 145.15; found 146 $[M+H]^+$

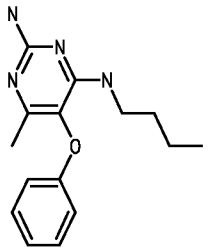
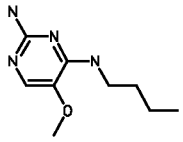
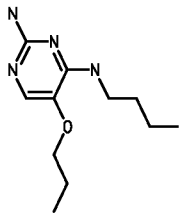
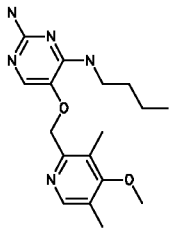
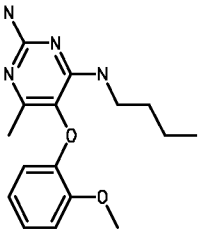


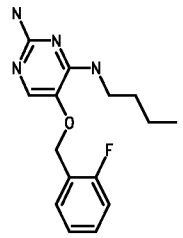
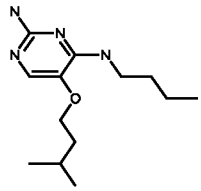
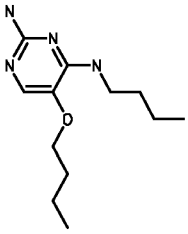
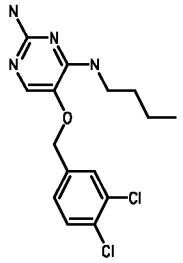
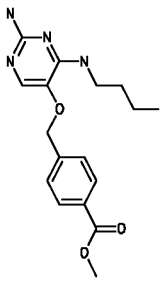
C-5 **304**

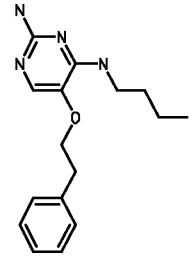
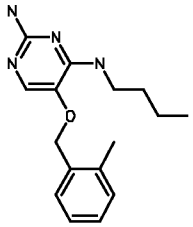
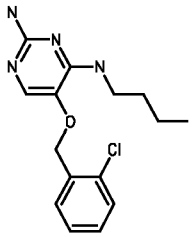
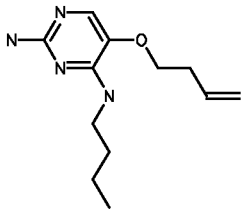
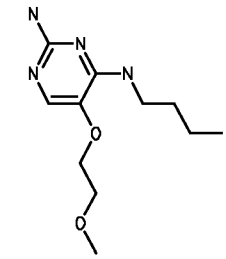
Step 2. C-5 was combined with AA-17 according to the method used to prepare compound 1, except that acetonitrile was used as a solvent, to afford **304.**

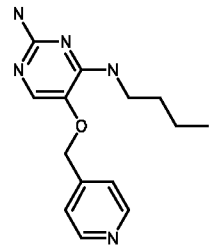
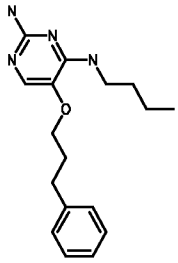
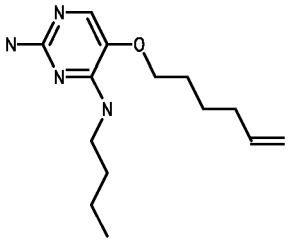
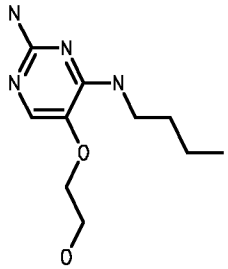
Table I: **Compounds of formula (I)**.

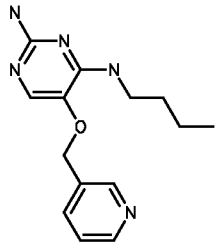
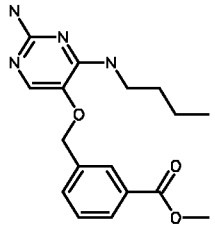
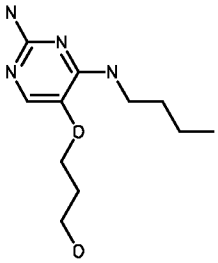
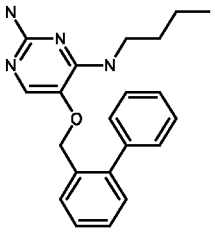
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
1		272.16	273	4.51,B	^{1}H NMR (400 MHz, METHANOL-d_4) δ ppm 0.96 (t, J=7.3 Hz, 3 H), 1.32 - 1.43 (m, 2 H), 1.52 - 1.61 (m, 2 H), 3.38 (t, J=7.2 Hz, 2 H), 5.01 (s, 2 H), 7.28 (s, 1 H), 7.31 - 7.46 (m, 5 H)
2		330.21	331	2.46, E	^{1}H NMR (300 MHz, CHLOROFORM-d) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.24 - 1.40 (m, 2 H), 1.43 - 1.59 (m, 2 H), 1.88 - 2.07 (m, 2 H), 2.65 (t, J=7.4 Hz, 2 H), 3.24 - 3.37 (m, 2 H), 3.72 (s, 3 H), 3.82 (t, J=6.3 Hz, 2 H), 4.54 (br. s., 2 H), 4.99 - 5.14 (m, 1 H), 6.72 - 6.82 (m, 2 H), 7.04 (d, J=8.5 Hz, 2 H), 7.19 (s, 1 H)

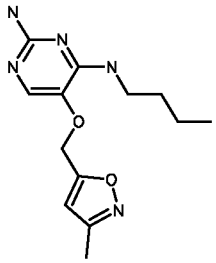
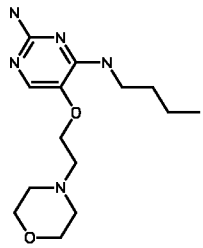
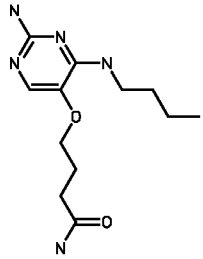
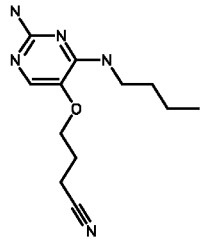
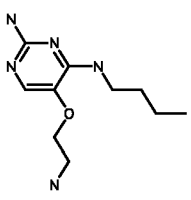
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
3		272.16	273	1.54, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.80 (t, J=7.3 Hz, 3 H), 1.20 (dq, J=15.0, 7.3 Hz, 2 H), 1.33 - 1.47 (m, 2 H), 1.98 (s, 3 H), 3.20 - 3.34 (m, 2 H), 4.74 (br. s., 2 H), 4.79 (br. s., 1 H), 6.78 - 6.84 (m, 2 H), 6.91 - 7.01 (m, 1 H), 7.18 - 7.28 (m, 2 H)
4		196.13	197	0.49, A	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.97 (t, J=7.3 Hz, 3 H), 1.35 - 1.48 (m, 2 H), 1.56 - 1.68 (m, 2 H), 3.44 - 3.52 (m, 2 H), 3.80 (s, 3 H), 5.86 (s, 1 H), 5.97 (s, 2 H), 7.07 - 7.14 (m, 1 H)
5		224.16	225	0.83, A	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.92 (t, J=7.3 Hz, 3 H), 1.03 (t, J=7.4 Hz, 3 H), 1.30 - 1.40 (m, 2 H), 1.50 - 1.62 (m, 2 H), 1.83 (m, J=7.5 Hz, 2 H), 2.27 (s, 6 H), 3.34 - 3.48 (m, 2 H), 3.99 (t, J=6.4 Hz, 2 H), 5.39 - 5.52 (m, 1 H), 7.63 (s, 1 H)
6		331.20	332	0.88, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.3 Hz, 3 H), 1.21 - 1.34 (m, 2 H), 1.48 (t, J=7.3 Hz, 2 H), 2.22 (s, 3 H), 2.24 (s, 3 H), 3.26 (q, J=7.0 Hz, 2 H), 3.74 (s, 3 H), 4.96 (s, 2 H), 5.54 (s, 2 H), 6.62 (s, 1 H), 7.39 (s, 1 H), 8.21 (s, 1 H)
7		302.17	303	1.55, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.80 (t, J=7.2 Hz, 3 H), 1.12 - 1.29 (m, 2 H), 1.34 - 1.47 (m, 2 H), 2.03 (s, 3 H), 3.21 - 3.31 (m, 2 H), 3.89 (s, 3 H), 4.67 (br. s., 2 H), 4.93 - 5.04 (m, 1 H), 6.55 - 6.62 (m, 1 H), 6.76 (td, J=7.4, 2.3 Hz, 1 H), 6.90 - 6.96 (m, 2 H)

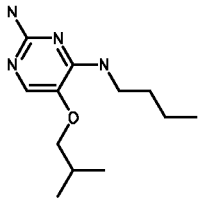
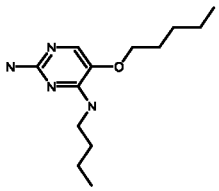
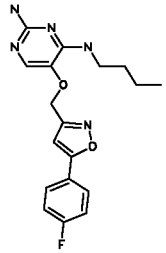
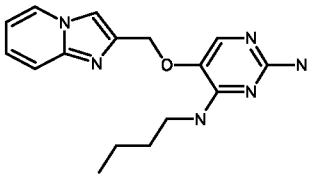
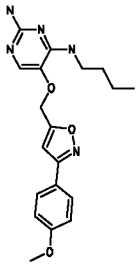
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
8		290.15	291	1.64, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.82 - 0.94 (m, 3 H), 1.22 - 1.39 (m, 2 H), 1.41 - 1.56 (m, 2 H), 3.24 - 3.38 (m, 2 H), 4.51 (br. s., 2 H), 4.92 (s, 2 H), 5.16 (br. s., 1 H), 6.97 - 7.15 (m, 2 H), 7.23 - 7.37 (m, 2 H), 7.40 (s, 1 H)
9		252.20	253	2.33, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.84 - 0.93 (m, 9 H), 1.24 - 1.39 (m, 2 H), 1.45 - 1.55 (m, 2 H), 1.53 - 1.62 (m, 2 H), 1.70 (dd, J=13.5, 6.7 Hz, 1 H), 3.28 - 3.38 (m, 2 H), 3.84 (t, J=6.6 Hz, 2 H), 4.47 (br. s., 2 H), 5.04 - 5.16 (m, 1 H), 7.20 (s, 1 H)
10		238.18	239	2.15, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.85 - 0.90 (m, 3 H), 0.89 - 0.95 (m, 3 H), 1.25 - 1.44 (m, 4 H), 1.45 - 1.58 (m, 2 H), 1.61 - 1.73 (m, 2 H), 3.27 - 3.39 (m, 2 H), 3.82 (t, J=6.5 Hz, 2 H), 4.57 (br. s., 2 H), 5.05 - 5.21 (m, 1 H), 7.25 (s, 1 H)
11		340.09	341	1.98, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.85 (t, J=7.3 Hz, 3 H), 1.20 - 1.36 (m, 2 H), 1.40 - 1.54 (m, 2 H), 3.24 - 3.36 (m, 2 H), 4.55 (br. s., 2 H), 4.80 (s, 2 H), 5.00 - 5.11 (m, 1 H), 7.11 (dd, J=8.2, 1.9 Hz, 1 H), 7.35 (s, 1 H), 7.38 (d, J=2.5 Hz, 2 H)
12		330.17	331	1.66, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.85 (t, J=7.3 Hz, 3 H), 1.28 (dd, J=15.2, 7.2 Hz, 2 H), 1.39 - 1.54 (m, 2 H), 3.25 - 3.35 (m, 2 H), 3.84 (s, 3 H), 4.61 (br. s., 2 H), 4.91 (s, 2 H), 5.07 - 5.17 (m, 1 H), 7.17 (s, 1 H), 7.35 (d, J=8.1 Hz, 2 H), 7.97 (d, J=8.2 Hz, 2 H)

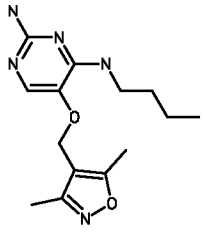
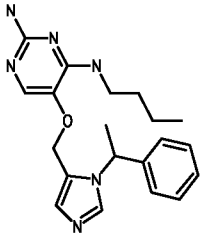
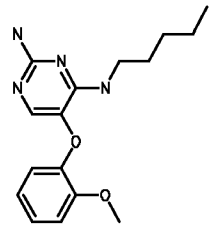
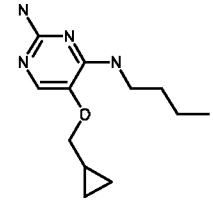
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
13		286.18	287	2.29, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.86 (t, J=1.0 Hz, 3 H), 1.17 - 1.31 (m, 2 H), 1.33 - 1.46 (m, 2 H), 2.98 (t, J=6.5 Hz, 2 H), 3.17 - 3.27 (m, 2 H), 4.03 (t, J=6.6 Hz, 2 H), 4.61 (br. s., 2 H), 4.83 - 4.97 (m, 1 H), 7.15 - 7.22 (m, 3 H), 7.23 - 7.31 (m, 3 H)
14		286.18	287	1.75, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.28 (dd, J=15.3, 7.3 Hz, 2 H), 1.41 - 1.54 (m, 2 H), 2.29 (s, 3 H), 3.26 - 3.37 (m, 2 H), 4.79 - 4.84 (m, 1 H), 4.87 (s, 2 H), 7.11 - 7.27 (m, 4 H), 7.31 (s, 1 H)
15		306.12	307	1.79, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.87 (t, J=7.3 Hz, 3 H), 1.24 - 1.37 (m, 2 H), 1.42 - 1.57 (m, 2 H), 3.24 - 3.38 (m, 2 H), 4.54 (br. s., 2 H), 4.97 (s, 2 H), 5.14 - 5.24 (m, 1 H), 7.17 - 7.27 (m, 2 H), 7.31 - 7.39 (m, 3 H)
16		236.16	237	1.98, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.88 (t, J=7.3 Hz, 3 H), 1.25 - 1.39 (m, 2 H), 1.44 - 1.57 (m, 2 H), 2.43 (q, J=6.6 Hz, 2 H), 3.27 - 3.37 (m, 2 H), 3.87 (t, J=6.5 Hz, 2 H), 4.51 (br. s., 2 H), 5.02 - 5.09 (m, 2 H), 5.10 - 5.18 (m, 1 H), 5.79 (ddt, J=17.1, 10.3, 6.7, 6.7 Hz, 1 H), 7.20 (s, 1 H)
17		240.16	241	1.52, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.88 (t, J=7.3 Hz, 3 H), 1.33 (dq, J=15.0, 7.2 Hz, 2 H), 1.44 - 1.57 (m, 2 H), 3.32 (m, J=7.1, 7.1, 5.7 Hz, 2 H), 3.36 (s, 3 H), 3.54 - 3.62 (m, 2 H), 3.90 - 3.96 (m, 2 H), 4.57 (br. s., 2 H), 5.55 - 5.69 (m, 1 H), 7.38 (s, 1 H)

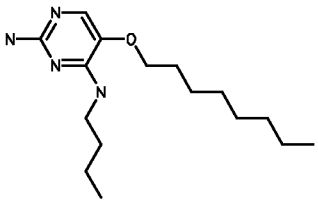
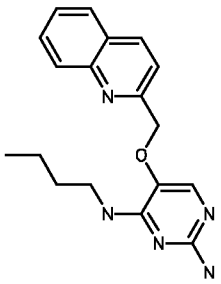
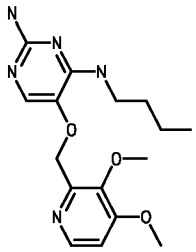
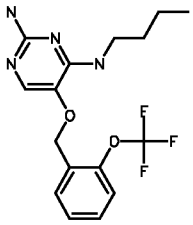
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
18		273.16	274	0.58, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.32 (dq, J=15.0, 7.3 Hz, 2 H), 1.45 - 1.59 (m, 2 H), 3.35 (td, J=7.0, 6.0 Hz, 2 H), 4.59 (br. s., 2 H), 4.92 (s, 2 H), 5.11 - 5.19 (m, 1 H), 7.20 (s, 1 H), 7.23 (s, 2 H), 8.54 - 8.59 (m, 2 H)
19		300.20	301	2.46, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.90 (t, J=7.1 Hz, 3 H), 1.25 - 1.39 (m, 2 H), 1.50 (m, J=6.7 Hz, 2 H), 2.04 - 2.19 (m, 2 H), 2.66 - 2.79 (m, 2 H), 3.37 (d, J=4.5 Hz, 2 H), 3.79 - 3.94 (m, 2 H), 5.68 - 5.88 (m, 1 H), 7.05 - 7.37 (m, 6 H)
20		264.20	265	2.38, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.91 (t, J=7.3 Hz, 3 H), 1.27 - 1.42 (m, 2 H), 1.44 - 1.61 (m, 4 H), 1.65 - 1.80 (m, 2 H), 2.07 (q, J=7.2 Hz, 2 H), 3.36 (td, J=7.0, 5.9 Hz, 2 H), 3.84 (t, J=6.5 Hz, 2 H), 4.60 (br. s., 2 H), 4.90 - 4.98 (m, 1 H), 5.02 (q, J=1.6 Hz, 1 H), 5.09 - 5.21 (m, 1 H), 5.77 (ddt, J=17.0, 10.3, 6.6, 6.6 Hz, 1 H), 7.27 (s, 1 H)
21		226.14	227	0.82, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.93 (t, J=7.3 Hz, 3 H), 1.30 - 1.46 (m, 2 H), 1.49 - 1.62 (m, 2 H), 3.20 (br. s., 1 H), 3.32 - 3.43 (m, 2 H), 3.88 - 3.94 (m, 2 H), 3.95 - 4.00 (m, 2 H), 4.62 (br. s., 2 H), 5.68 (t, J=5.2 Hz, 1 H), 7.39 (s, 1 H)

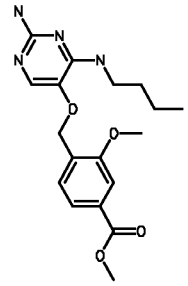
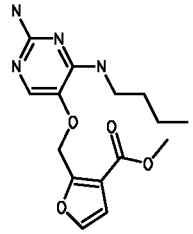
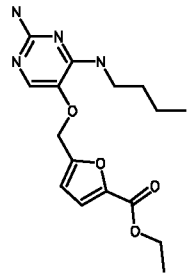
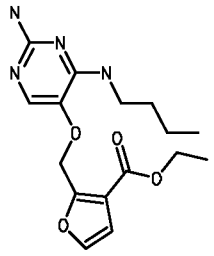
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
22		273.16	274	0.807, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.94 (t, J=7.3 Hz, 3 H), 1.29 - 1.45 (m, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 3.39 (q, J=6.8 Hz, 2 H), 4.57 (br. s., 2 H), 4.97 (s, 2 H), 5.08 - 5.19 (m, 1 H), 7.34 (dd, J=7.8, 4.9 Hz, 1 H), 7.44 (s, 1 H), 7.71 (m, J=7.8 Hz, 1 H), 8.62 (dd, J=4.7, 1.3 Hz, 1 H), 8.67 (d, J=1.5 Hz, 1 H)
23		330.17	331	1.65, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.94 (t, J=7.3 Hz, 3 H), 1.30 - 1.47 (m, 2 H), 1.56 (quin, J=7.3 Hz, 2 H), 3.35 - 3.45 (m, 2 H), 3.94 (s, 3 H), 4.62 (br. s., 2 H), 5.00 (s, 2 H), 5.15 - 5.25 (m, 1 H), 7.40 (s, 1 H), 7.49 (d, J=7.6 Hz, 1 H), 7.55 - 7.63 (m, 1 H), 7.99 - 8.13 (m, 2 H)
24		240.16	241	0.97, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.95 (t, J=7.2 Hz, 3 H), 1.31 - 1.46 (m, 2 H), 1.51 - 1.65 (m, 2 H), 2.01 (quin, J=6.0 Hz, 2 H), 2.61 (br. s., 1 H), 3.30 - 3.45 (m, 2 H), 3.84 (t, J=5.9 Hz, 2 H), 4.01 (t, J=6.0 Hz, 2 H), 4.55 (br. s., 2 H), 5.31 - 5.42 (m, 1 H), 7.35 (s, 1 H)
25		348.20	349	2.02, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.95 (t, J=7.3 Hz, 3 H), 1.28 - 1.45 (m, 2 H), 1.47 - 1.60 (m, 2 H), 3.30 - 3.40 (m, 2 H), 4.60 (br. s., 2 H), 4.87 (s, 2 H), 5.10 (m, J=5.2 Hz, 1 H), 7.20 (s, 1 H), 7.31 - 7.47 (m, 8 H), 7.49 - 7.56 (m, 1 H)

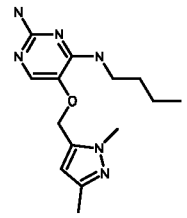
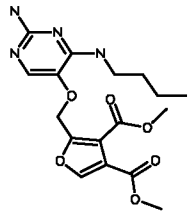
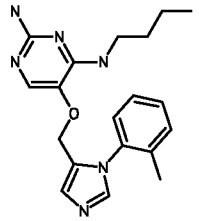
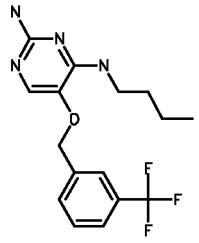
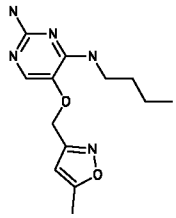
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
26		277.15	278	1.69, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.96 (t, J=7.3 Hz, 3 H), 1.28 - 1.46 (m, 2 H), 1.50 - 1.63 (m, 2 H), 2.32 (s, 3 H), 3.39 (td, J=7.1, 5.9 Hz, 2 H), 4.70 (br. s., 2 H), 5.00 (s, 2 H), 5.18 - 5.27 (m, 1 H), 6.15 (s, 1 H), 7.45 (s, 1 H)
27		295.20	296	0.67, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.96 (t, J=7.3 Hz, 3 H), 1.34 - 1.47 (m, 2 H), 1.52 - 1.67 (m, 2 H), 2.51 - 2.60 (m, 4 H), 2.69 (t, J=5.4 Hz, 2 H), 3.41 (td, J=7.1, 5.9 Hz, 2 H), 3.71 - 3.81 (m, 4 H), 3.98 (t, J=5.4 Hz, 2 H), 4.60 (br. s., 2 H), 5.85 - 5.98 (m, 1 H), 7.44 (s, 1 H)
28		267.17	268	0.94, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.98 (t, J=7.3 Hz, 3 H), 1.34 - 1.50 (m, 2 H), 1.55 - 1.70 (m, 2 H), 2.06 (d, J=3.4 Hz, 2 H), 2.15 (dt, J=13.0, 6.4 Hz, 2 H), 2.37 - 2.47 (m, 2 H), 3.42 (td, J=7.1, 5.8 Hz, 2 H), 3.96 (t, J=6.0 Hz, 2 H), 4.70 (br. s., 2 H), 5.34 - 5.44 (m, 1 H), 7.32 (s, 1 H)
29		249.16	250	1.18, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.98 (t, J=7.3 Hz, 3 H), 1.34 - 1.50 (m, 2 H), 1.56 - 1.69 (m, 2 H), 2.15 (dt, J=13.0, 6.4 Hz, 2 H), 2.39 - 2.47 (m, 2 H), 3.42 (td, J=7.1, 5.8 Hz, 2 H), 3.96 (t, J=6.0 Hz, 2 H), 4.70 (br. s., 2 H), 5.45 - 5.59 (m, 1 H), 7.32 (s, 1 H)
30		225.16	226	0.20, E	^{1}H NMR (300 MHz, METHANOL-d_4) δ ppm 0.86 (t, J=7.4 Hz, 3 H), 1.22 - 1.37 (m, 2 H), 1.49 (t, J=7.5 Hz, 2 H), 2.89 (t, J=5.0 Hz, 2 H), 3.29 (t, J=7.2 Hz, 2 H), 3.81 (t, J=5.1 Hz, 2 H), 7.16 (s, 1 H)

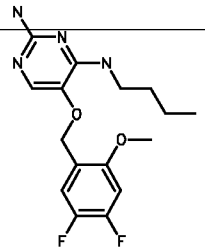
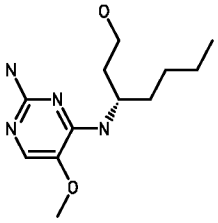
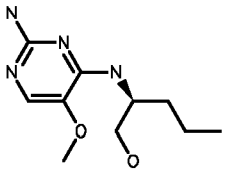
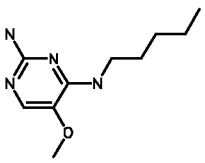
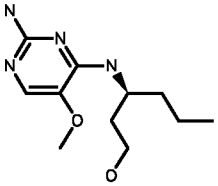
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
31		238.18	239	2.16, E	^{1}H NMR (300 MHz, METHANOL-d_4) δ ppm 0.97 (t, J=1.0 Hz, 3 H), 1.05 (d, J=6.7 Hz, 6 H), 1.27 - 1.48 (m, 2 H), 1.54 - 1.73 (m, 2 H), 1.99 - 2.22 (m, 1 H), 3.45 - 3.60 (m, 2 H), 3.68 - 3.79 (m, 2 H), 7.15 - 7.22 (m, 1 H)
32		252.20	253	2.36, E	^{1}H NMR (300 MHz, METHANOL-d_4) δ ppm 1.00 - 1.13 (m, 6 H), 1.38 - 1.60 (m, 6 H), 1.65 - 1.78 (m, 2 H), 1.87 - 1.97 (m, 2 H), 3.56 - 3.64 (m, 2 H), 3.66 - 3.78 (m, 1 H), 4.00 - 4.09 (m, 2 H)
33		357.16	358	1.01, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.3 Hz, 3 H), 1.19 - 1.33 (m, 2 H), 1.41 - 1.53 (m, 2 H), 3.28 (q, J=6.6 Hz, 2 H), 5.04 (s, 2 H), 5.63 (s, 2 H), 6.52 (t, J=5.9 Hz, 1 H), 7.23 (s, 1 H), 7.37 - 7.45 (m, 2 H), 7.50 (s, 1 H), 7.91 - 7.98 (m, 2 H)
34		312.17	313	0.71, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.3 Hz, 3 H), 1.19 - 1.35 (m, 2 H), 1.40 - 1.53 (m, 2 H), 3.26 (q, J=7.0 Hz, 2 H), 5.00 (s, 2 H), 5.58 (s, 2 H), 6.62 (t, J=5.7 Hz, 1 H), 6.90 (t, J=6.6 Hz, 1 H), 7.21 - 7.30 (m, 1 H), 7.46 - 7.57 (m, 2 H), 8.00 (s, 1 H), 8.53 (d, J=7.0 Hz, 1 H)
35		369.18	370	0.98, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.5 Hz, 3 H), 1.23 - 1.32 (m, 2 H), 1.42 - 1.53 (m, 2 H), 3.23 - 3.31 (m, 2 H), 3.82 (s, 3 H), 5.09 (s, 2 H), 5.63 (s, 2 H), 6.48 - 6.56 (m, 1 H), 7.07 (d, J=8.4 Hz, 2 H), 7.15 (s, 1 H), 7.46 (s, 1 H), 7.81 (d, J=8.4 Hz, 2 H)

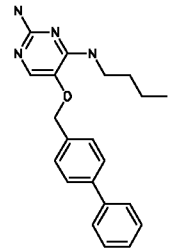
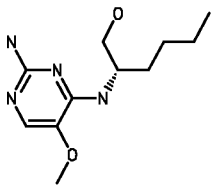
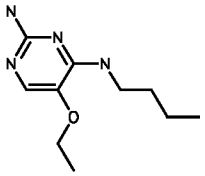
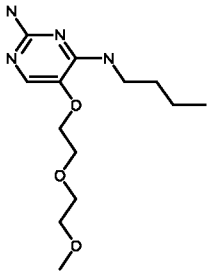
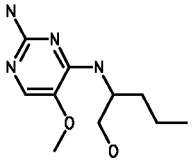
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
36		291.17	292	0.78, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.3 Hz, 3 H), 1.18 - 1.31 (m, 2 H), 1.38 - 1.51 (m, 2 H), 2.20 (s, 3 H), 2.33 (s, 3 H), 3.18 - 3.29 (m, 2 H), 4.72 (s, 2 H), 5.57 (s, 2 H), 6.40 (t, J=5.9 Hz, 1 H), 7.38 (s, 1 H)
37		366.22	367	0.84, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.3 Hz, 3 H), 1.20 - 1.32 (m, 2 H), 1.38 - 1.49 (m, 2 H), 1.81 (d, J=7.0 Hz, 3 H), 3.21 (dt, J=13.4, 6.9 Hz, 2 H), 4.62 (d, J=12.8 Hz, 1 H), 4.87 (d, J=12.4 Hz, 1 H), 5.52 - 5.61 (m, 3 H), 6.12 (t, J=5.9 Hz, 1 H) , 7.00 (s, 1 H), 7.15 (d, J=7.0 Hz, 2 H), 7.25 - 7.37 (m, 4 H), 7.99 (s, 1 H)
38		302.17	303	0.99, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.86 - 0.94 (m, 3 H), 1.26 (s, 1 H), 1.29 - 1.39 (m, 4 H), 1.60 (t, J=7.2 Hz, 2 H), 3.40 - 3.49 (m, 2 H), 3.87 (s, 3 H), 5.50 - 5.64 (m, 1 H), 5.74 - 5.84 (m, 1 H), 6.92 (dd, J=7.3, 1.3 Hz, 1 H), 6.95 - 7.01 (m, 2 H), 7.11 - 7.17 (m, 1 H), 7.26 (s, 1 H)
39		236.16	237	1.91, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.31 - 0.43 (m, 2 H), 0.63 - 0.78 (m, 2 H), 0.99 (t, J=7.3 Hz, 3 H), 1.16 - 1.31 (m, 1 H), 1.35 - 1.49 (m, 2 H), 1.65 (quin, J=7.4 Hz, 2 H), 3.43 - 3.59 (m, 2 H), 3.72 (d, J=7.0 Hz, 2 H), 6.02 - 6.18 (m, 1 H), 7.01 (s, 1 H)

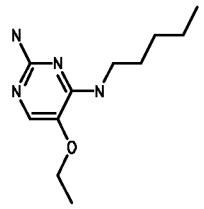
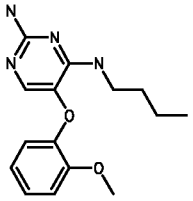
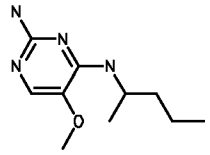
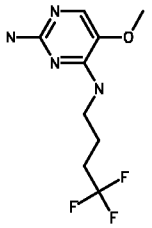
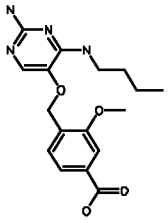
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
40		294.24	295	2.83, E	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.77 - 0.85 (m, 3 H), 0.88 (t, J=7.3 Hz, 3 H), 1.15 - 1.40 (m, 8 H), 1.45 - 1.58 (m, 2 H), 1.62 - 1.73 (m, 2 H), 1.77 (m, J=13.3 Hz, 2 H), 3.33 (td, J=7.0, 5.9 Hz, 2 H), 3.53 - 3.62 (m, 1 H), 3.66 - 3.74 (m, 1 H), 3.81 (t, J=6.6 Hz, 2 H), 4.41 (br. s., 2 H), 5.03 - 5.14 (m, 1 H), 7.27 (s, 1 H)
41		323.17	324	0.90, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.23 - 1.36 (m, 2 H), 1.52 (t, J=7.1 Hz, 2 H), 3.27 - 3.33 (m, 2 H), 5.20 (s, 2 H), 5.57 (s, 2 H), 6.78 (s, 1 H), 7.43 (s, 1 H), 7.59 - 7.66 (m, 1 H), 7.74 - 7.82 (m, 2 H), 8.01 (d, J=8.4 Hz, 2 H), 8.43 (d, J=8.4 Hz, 1 H)
42		333.18	334	0.76, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.5 Hz, 3 H), 1.21 - 1.36 (m, 2 H), 1.42 - 1.54 (m, 2 H), 3.23 - 3.30 (m, 2 H), 3.75 (s, 3 H), 3.90 (s, 3 H), 4.90 (s, 2 H), 5.59 (s, 2 H), 6.72 (t, J=5.5 Hz, 1 H), 7.14 (d, J=5.9 Hz, 1 H), 7.44 (s, 1 H), 8.23 (d , J=5.5 Hz, 1 H)
43		356.15	357	1.07, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.28 (dq, J=14.9, 7.3 Hz, 2 H), 1.49 (quin, J=7.2 Hz, 2 H), 3.28 (q, J=6.6 Hz, 2 H), 4.98 (s, 2 H), 5.60 (s, 2 H), 6.40 (t, J=5.9 Hz, 1 H), 7.35 (s, 1 H), 7.37 - 7.54 (m, 3 H), 7.70 (dd, J=7.3, 1.5 Hz, 1 H)

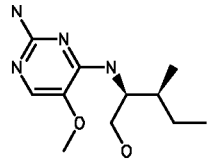
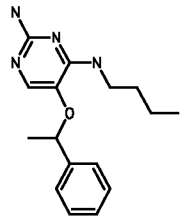
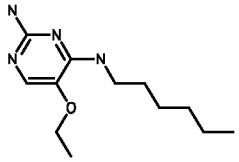
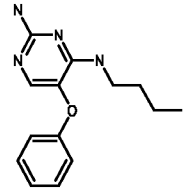
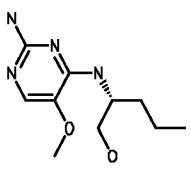
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
44		360.18	361	0.95, D	^{1}H NMR (400 MHz, CHLOROFORM- *d*) δ ppm 0.95 (t, J=7.3 Hz, 3 H), 1.38 (dq, J=15.1, 7.4 Hz, 2 H), 1.57 (quin, J=7.3 Hz, 2 H), 3.36 - 3.44 (m, 2 H), 3.92 (s, 3 H), 3.93 (s, 3 H), 4.63 - 4.72 (m, 2 H), 5.00 (s, 2 H), 5.32 (br. s., 1 H), 7.40 (d, J=7.8 Hz, 1 H), 7.43 (s, 1 H), 7.57 (d, J=1.0 Hz, 1 H), 7.66 (dd, J=7.8, 1.5 Hz, 1 H)
45		320.15	321	0.83, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.22 - 1.36 (m, 2 H), 1.49 (s, 2 H), 3.22 - 3.31 (m, 2 H), 3.82 (s, 3 H), 5.09 (s, 2 H), 5.57 (s, 2 H), 6.52 (t, J=5.9 Hz, 1 H), 6.94 (d, J=1.5 Hz, 1 H), 7.36 (s, 1 H), 7.95 (d, J=1.8 Hz, 1 H)
46		334.16	335	0.89, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.5 Hz, 3 H), 1.21 - 1.29 (m, 2 H), 1.29 (t, J=7.0 Hz, 3 H), 1.47 (quin, J=7.4 Hz, 2 H), 3.25 (q, J=6.8 Hz, 2 H), 4.29 (q, J=7.1 Hz, 2 H), 4.95 (s, 2 H), 5.60 (s, 2 H), 6.41 (t, J=5.9 Hz, 1 H), 6.76 (d, J=3.7 Hz, 1 H), 7.28 (d, J=3.3 Hz, 1 H), 7.39 (s, 1 H)
47		334.16	335	0.93, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.24 (t, J=7.0 Hz, 3 H), 1.26 - 1.34 (m, 2 H), 1.43 - 1.54 (m, 2 H), 3.21 - 3.30 (m, 2 H), 4.21 (q, J=7.0 Hz, 2 H), 5.11 (s, 2 H), 5.62 (s, 2 H), 6.42 (t, J=5.9 Hz, 1 H), 6.78 (d, J=1.5 Hz, 1 H), 7.29 (s, 1 H), 7.78 - 7.86 (m, 1 H)

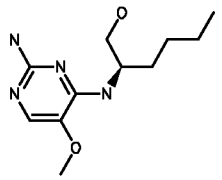
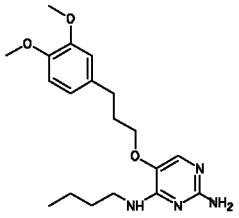
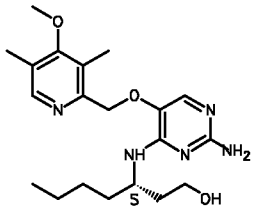
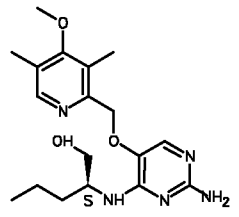
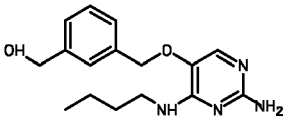
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
48		290.19	291	0.73, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.3 Hz, 3 H), 1.26 (dd, J=15.2, 7.5 Hz, 2 H), 1.41 - 1.53 (m, 2 H), 2.10 (s, 3 H), 3.21 - 3.29 (m, 2 H), 3.73 (s, 3 H), 4.91 (s, 2 H), 5.55 (s, 2 H), 6.11 (s, 1 H), 6.44 (t, J=5.9 Hz, 1 H), 7.39 (s, 1 H)
49		378.15	379	0.83, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.3 Hz, 3 H), 1.27 (dd, J=15.2, 7.5 Hz, 2 H), 1.46 (t, J=7.1 Hz, 2 H), 3.20 - 3.29 (m, 2 H), 3.74 (s, 3 H), 3.77 (s, 3 H), 5.00 (s, 2 H), 5.68 (s, 2 H), 6.38 - 6.48 (m, 1 H), 7.29 (s, 1 H), 8.46 (s, 1 H)
50		352.20	353	0.82, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.18 - 1.31 (m, 2 H), 1.37 - 1.49 (m, 2 H), 2.00 (s, 3 H), 3.19 (q, J=6.8 Hz, 2 H), 4.61 (br. s., 2 H), 5.53 (s, 2 H), 5.93 (t, J=5.9 Hz, 1 H), 7.01 (s, 1 H), 7.21 (s, 1 H), 7.32 (dd, J=8.6, 3.5 Hz, 1 H), 7.40 - 7.45 (m, 3 H), 7.82 (s, 1 H)
51		340.15	341	1.06, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.23 - 1.34 (m, 2 H), 1.49 (t, J=7.3 Hz, 2 H), 3.23 - 3.31 (m, 2 H), 5.05 (s, 2 H), 5.54 (s, 2 H), 6.57 (s, 1 H), 7.42 (s, 1 H), 7.63 (m, J=7.7 Hz, 1 H), 7.66 - 7.71 (m, 1 H), 7.75 - 7.84 (m, 2 H)
52		277.15	278	0.78, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.21 - 1.35 (m, 2 H), 1.42 - 1.54 (m, 2 H), 2.41 (s, 3 H), 3.27 (q, J=6.7 Hz, 2 H), 4.95 (s, 2 H), 5.61 (s, 2 H), 6.41 (s, 1 H), 6.50 (t, J=5.7 Hz, 1 H), 7.42 (s, 1 H)

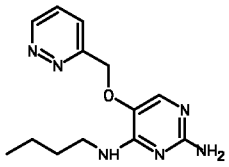
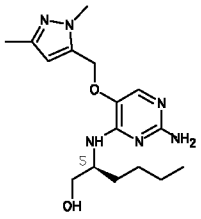
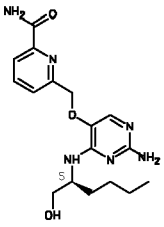
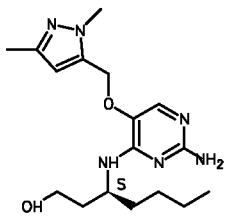
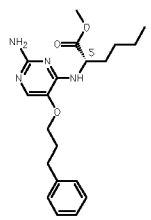
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
53		338.16	339	1.00, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.21 - 1.35 (m, 2 H), 1.49 (quin, J=7.3 Hz, 2 H), 3.29 (q, J=6.6 Hz, 2 H), 3.81 (s, 3 H), 4.83 (s, 2 H), 5.56 (s, 2 H), 6.45 (t, J=5.9 Hz, 1 H), 7.20 (dd, J=12.8, 6.6 Hz, 1 H), 7.38 (s, 1 H), 7.60 (dd, J=11.0, 9.5 Hz, 1 H)
54		254.17	255	0.68, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=6.9 Hz, 3 H), 1.14 - 1.33 (m, 4 H), 1.44 - 1.54 (m, 2 H), 1.56 - 1.72 (m, 2 H), 3.40 (t, J=6.4 Hz, 2 H), 3.67 (s, 3 H), 4.05 - 4.18 (m, 1 H), 4.39 (br. s., 1 H), 5.45 (s, 2 H), 6.13 (d, J=9.0 Hz, 1 H), 7.34 (s, 1 H)
55		226.14	227	0.52, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.4 Hz, 3 H), 1.19 - 1.35 (m, 2 H), 1.45 (dt, J=13.5, 4.4 Hz, 1 H), 1.50 - 1.62 (m, 1 H), 3.30 - 3.49 (m, 2 H), 3.67 (s, 3 H), 4.05 (td, J=8.8, 5.0 Hz, 1 H), 4.36 - 4.96 (m, 1 H), 5.46 (s, 2 H), 5.89 (d, J=9.0 Hz, 1 H), 7.35 (s, 1 H)
56		210.15	211	0.75, D	^{1}H NMR (400 MHz, CHLOROFORM-d) δ ppm 0.86 - 0.98 (m, 3 H), 1.29 - 1.43 (m, 4 H), 1.55 - 1.65 (m, 2 H), 3.39 (td, J=7.2, 5.8 Hz, 2 H), 3.78 (s, 3 H), 4.42 (br. s., 2 H), 5.14 (br. s., 1 H), 7.37 (s, 1 H)
57		240.16	241	0.58, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.4 Hz, 3 H), 1.15 - 1.34 (m, 2 H), 1.37 - 1.54 (m, 2 H), 1.56 - 1.73 (m, 2 H), 3.40 (t, J=6.4 Hz, 2 H), 3.67 (s, 3 H), 4.04 - 4.22 (m, 1 H), 4.40 (br. s., 1 H), 5.46 (s, 2 H), 6.13 (d, J=8.8 Hz, 1 H), 7.35 (br. s., 1 H)

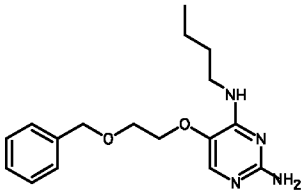
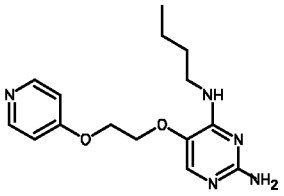
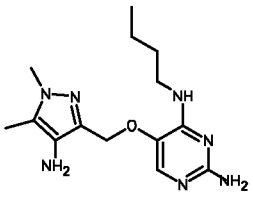
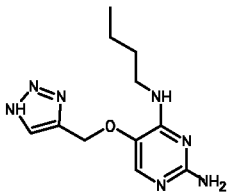
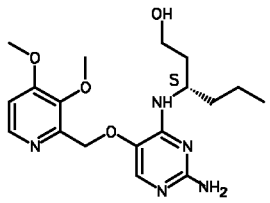
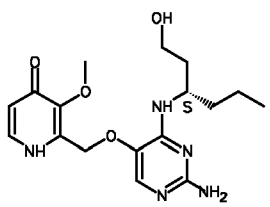
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
58		348.20	349	1.16, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 - 0.93 (m, 3 H), 1.20 - 1.36 (m, 2 H), 1.53 (t, J=7.4 Hz, 2 H), 3.33 - 3.45 (m, 2 H), 5.11 (s, 2 H), 7.33 - 7.40 (m, 1 H), 7.43 - 7.50 (m, 2 H), 7.51 - 7.60 (m, 4 H), 7.64 - 7.73 (m, 3 H), 8.42 - 8.50 (m, 1 H), 12.15 (d, J=4.8 Hz, 0 H)
59		240.16	241	0.62, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.79 - 0.90 (m, 3 H), 1.16 - 1.34 (m, 4 H), 1.37 - 1.49 (m, 1 H), 1.53 - 1.67 (m, 1 H), 3.17 - 3.51 (m, 2 H), 3.68 (s, 3 H), 3.95 - 4.11 (m, 1 H), 4.67 (br. s., 1 H), 5.45 (s, 2 H), 5.89 (d, J=9.0 Hz, 1 H), 7.36 (s, 1 H)
60		210.15	211	3.93, B	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.83 - 0.92 (m, 3 H), 1.22 - 1.29 (m, 2 H), 1.32 (t, J=7.0 Hz, 3 H), 1.52 (quin, J=7.3 Hz, 2 H), 3.36 - 3.42 (m, 2 H), 3.96 (q, J=6.9 Hz, 2 H), 7.41 (s, 1 H), 7.48 (br. s., 2 H), 8.36 (t, J=5.9 Hz, 1 H)
61		284.18	285	0.68, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.23 - 1.37 (m, 2 H), 1.44 - 1.55 (m, 2 H), 3.26 (s, 3 H), 3.26 - 3.31 (m, 2 H), 3.47 (dd, J=5.5, 3.7 Hz, 2 H), 3.56 - 3.60 (m, 2 H), 3.65 (dd, J=5.5, 3.7 Hz, 2 H), 3.90 (dd, J=5.3, 3.8 Hz, 2 H), 5.60 (s, 2 H), 6.28 (t, J=5.9 Hz, 1 H), 7.41 (s, 1 H)
62		226.14	227	0.52, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.96 (t, J=7.3 Hz, 3 H), 1.33 - 1.48 (m, 2 H), 1.50 - 1.67 (m, 2 H), 3.61 (dd, J=10.9, 6.9 Hz, 1 H), 3.76 (d, J=3.0 Hz, 1 H), 3.79 (s, 3 H), 3.87 - 4.00 (m, 1 H), 4.01 - 4.13 (m, 1 H), 4.45 (br. s., 2 H), 5.22 (d, J=6.8 Hz, 1 H), 7.39 (s, 1 H)

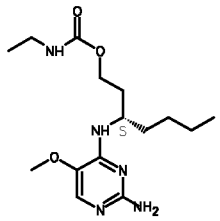
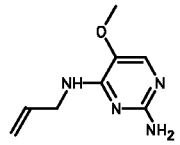
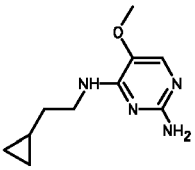
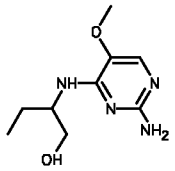
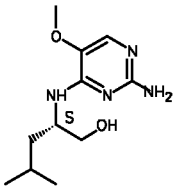
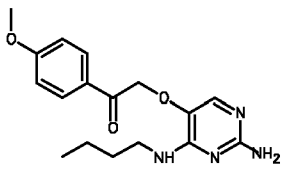
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
63		224.16	225	3.23, C	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.0 Hz, 3 H), 1.29 (dd, J=15.3, 7.5 Hz, 2 H), 1.24 (m, J=3.0 Hz, 2 H), 1.32 - 1.38 (m, 3 H), 1.51 - 1.62 (m, 2 H), 3.40 - 3.44 (m, 2 H), 3.98 (q, J=6.9 Hz, 2 H), 7.42 (s, 1 H), 7.49 (br. s., 2 H), 8.39 (t, J=5.8 Hz, 1 H)
64		288.16	289	0.91, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.94 (t, J=7.3 Hz, 3 H), 1.26 (s, 1 H), 1.37 (dd, J=15.1, 7.5 Hz, 2 H), 1.52 - 1.63 (m, 2 H), 3.39 - 3.50 (m, 2 H), 3.88 (s, 3 H), 5.31 - 5.44 (m, 1 H), 5.60 - 5.71 (m, 1 H), 6.87 - 7.01 (m, 3 H), 7.08 - 7.15 (m, 1 H), 7.33 (s, 1 H).
65		210.15	211	0.73, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.86 - 0.99 (m, 3 H), 1.18 (d, J=6.5 Hz, 3 H), 1.28 - 1.39 (m, 2 H), 1.44 - 1.55 (m, 2 H), 3.76 (s, 3 H), 4.08 - 4.22 (m, 1 H), 4.40 (br. s., 2 H), 4.94 (d, J=7.8 Hz, 1 H), 7.34 (s, 1 H)
66		250.10	251	0.66, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 1.82 - 1.93 (m, 2 H), 2.07 - 2.25 (m, 2 H), 3.50 (q, J=6.6 Hz, 2 H), 3.77 (s, 3 H), 4.54 (br. s., 2 H), 5.21 - 5.31 (m, 1 H), 7.39 (s, 1 H)
67		346.16	347	0.56, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.24 - 1.35 (m, 2 H), 1.50 (t, J=7.2 Hz, 2 H), 3.25 - 3.33 (m, 2 H), 3.83 (s, 3 H), 4.88 (s, 2 H), 5.57 (s, 1 H), 6.32 (s, 1 H), 7.33 (d, J=7.8 Hz, 1 H), 7.36 (s, 1 H), 7.48 (dd, J=7.7, 1.1 Hz, 1 H), 7.54 (d, J=1.0 Hz, 1 H)

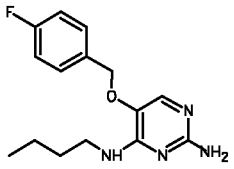
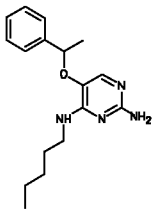
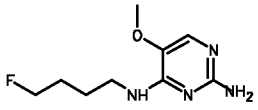
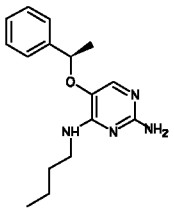
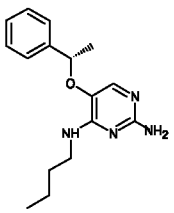
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
68		240.16	241	0.61, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.91 (t, J=7.4 Hz, 3 H), 0.96 (d, J=7.0 Hz, 3 H), 1.19 (ddd, J=13.6, 8.8, 7.3 Hz, 1 H), 1.53 (ddd, J=13.5, 7.5, 4.1 Hz, 1 H), 1.75 (ddd, J=6.6, 4.2, 2.3 Hz, 1 H), 3.65 - 3.71 (m, 1 H), 3.75 (s, 3 H), 3.77 (d, J=3.0 Hz, 1 H), 3.80 (d, J=3.3 Hz, 1 H), 3.90 - 4.00 (m, 1 H), 4.64 (br. s., 2 H), 5.39 (d, J=7.8 Hz, 1 H), 7.32 (s, 1 H)
69		286.18	287	1.00, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.91 (t, J=7.4 Hz, 3 H), 1.23 - 1.36 (m, 2 H), 1.49 - 1.57 (m, 2 H), 1.58 (d, J=6.5 Hz, 3 H), 3.37 - 3.47 (m, 2 H), 5.39 (d, J=6.5 Hz, 1 H), 7.21 (s, 1 H), 7.27 - 7.33 (m, 2 H), 7.34 - 7.40 (m, 2 H), 7.41 - 7.46 (m, 1 H), 8.43 (s, 1 H), 11.05 - 11.32 (m, 1 H)
70		238.18	239	3.56, C	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.83 - 0.92 (m, 3 H), 1.22 - 1.31 (m, 7 H), 1.35 (t, J=6.9 Hz, 3 H), 1.49 - 1.63 (m, 2 H), 3.40 - 3.44 (m, 2 H), 3.99 (q, J=6.9 Hz, 2 H), 7.47 (br. s., 2 H), 8.39 (t, J=5.8 Hz, 1 H)
71		258.15	259	0.94,A	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.86 - 0.91 (m, 3 H), 1.24 - 1.30 (m, 2 H), 1.44 - 1.54 (m, 2 H), 3.37 (td, J=7.1, 5.9 Hz, 2 H), 4.97 (br. s., 3 H), 6.92 - 6.97 (m, 2 H), 7.01 - 7.06 (m, 1 H), 7.25 - 7.31 (m, 2 H), 7.58 (s, 1 H)
72		226.14	227	0.52, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.18 - 1.36 (m, 2 H), 1.45 (dd, J=8.9, 4.9 Hz, 1 H), 1.51 - 1.62 (m, 1 H), 3.40 (d, J=16.6 Hz, 2 H), 3.67 (s, 3 H), 3.95 - 4.13 (m, 1 H), 4.65 (br. s., 1 H), 5.44 (s, 2 H), 5.88 (d, J=9.0 Hz, 1 H), 7.35 (s, 1 H)

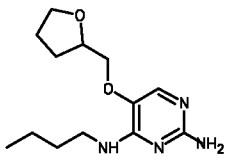
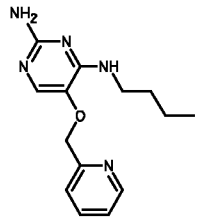
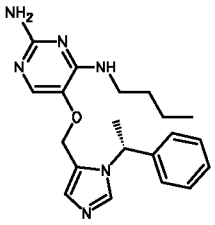
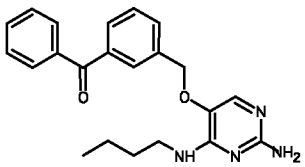
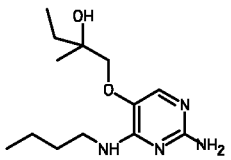
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
73		240.16	241	0.63, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.81 - 0.90 (m, 3 H), 1.17 - 1.37 (m, 3 H), 1.39 - 1.51 (m, 1 H), 1.54 - 1.66 (m, 1 H), 2.51 (dt, J=3.7, 1.8 Hz, 1 H), 3.34 - 3.41 (m, 1 H), 3.41 - 3.48 (m, 1 H), 3.68 (s, 3 H), 4.04 (td, J=8.7, 5.0 Hz, 1 H), 4.43 - 4.91 (m, 1 H), 5.47 (s, 2 H), 5.90 (d, J=9.0 Hz, 1 H), 7.36 (s, 1 H)
74		360.22	361	0.94, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.91 (t, J=7.4 Hz, 3 H), 1.30 (dq, J=14.9, 7.4 Hz, 2 H), 1.49 - 1.61 (m, 2 H), 1.95 - 2.09 (m, 2 H), 2.70 (t, J=7.7 Hz, 2 6 H), 3.42 (q, J=6.8 Hz, 2 H), 3.71 (s, 3 H), 3.72 (s, 3 H), 3.89 (t, J=6.3 Hz, 2 H), 6.72 (dd, J=8.2, 1.9 Hz, 1 H), 6.81 (d, J=1.8 Hz, 1 H), 6.86 (d, J=8.3 Hz, 1 H), 7.36 (d, J=5.8 Hz, 1 H), 7.43 (br. s., 2 H), 8.32 (t, J=6.0 Hz, 1 H), 11.77 (d, J=5.3 Hz, 1 H)
75		389.24	390	0.88, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=6.9 Hz, 3 H), 1.15 - 1.38 (m, 4 H), 1.58 (m, J=13.3, 13.3, 7.0 Hz, 1 H), 1.67 - 1.83 (m, 2 H), 1.84 - 1.99 (m, 6 1 H), 2.27 (s, 3 H), 2.38 (s, 3 H), 3.41 (t, J=6.4 Hz, 2 H), 3.97 (s, 3 H), 4.38 (dt, J=9.0, 4.7 Hz, 1 H), 5.35 (s, 2 H), 7.51 (br. s, 2 H), 7.77 (s, 1 H), 8.53 (s, 1 H), 8.96 (br. s., 1 H), 12.20 (br. s., 1 H)
76		361.21	362	0.75, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.18 - 1.36 (m, 2 H), 1.36 - 1.50 (m, 1 H), 1.50 - 1.63 (m, 1 H), 2.22 (s, 3 H), 2.24 (s, 3 H), 6 3.29 - 3.48 (m, 2 H), 3.74 (s, 3 H), 4.03 (td, J=8.7, 4.6 Hz, 1 H), 4.68 (br. s., 1 H), 4.91 - 5.05 (m, 2 H), 5.53 (s, 2 H), 6.19 (d, J=8.8 Hz, 1 H), 7.44 (s, 1 H), 8.21 (s, 1 H)
77		302.17	303	0.75, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.22 - 1.35 (m, 2 H), 1.50 (quin, J=7.3 Hz, 2 H), 3.24 - 3.30 (m, 2 H), 4.51 (d, J=5.3 Hz, 2 H), 6 4.95 (s, 2 H), 5.19 (t, J=5.6 Hz, 1 H), 5.52 (s, 2 H), 6.42 (t, J=5.8 Hz, 1 H),

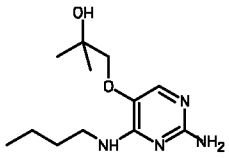
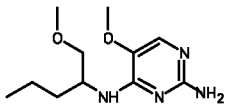
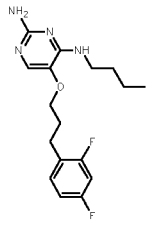
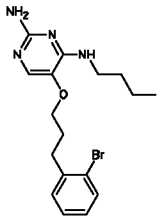
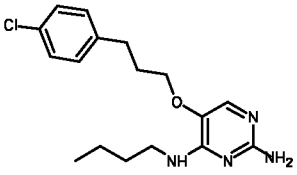
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					7.24 - 7.29 (m, 1 H), 7.29 - 7.34 (m, 2 H), 7.35 - 7.40 (m, 2 H)
78		274.15	275	0.59, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 1.30 (dq, J=14.9, 7.3 Hz, 2 H), 1.51 (quin, J=7.3 Hz, 2 H), 3.26 - 3.32 (m, 2 H), 5.24 (s, 2 H), 6 5.68 (s, 2 H), 6.78 (t, J=5.8 Hz, 1 H), 7.46 (s, 1 H), 7.76 (dd, J=8.4, 4.9 Hz, 1 H), 7.93 (dd, J=8.5, 1.5 Hz, 1 H), 9.21 (dd, J=5.0, 1.5 Hz, 1 H)
79		334.21	335	0.7, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.0 Hz, 3 H), 1.15 - 1.34 (m, 4 H), 1.36 - 1.50 (m, 1 H), 1.51 - 1.64 (m, 1 H), 2.11 (s, 3 H), 3.39 - 3.46 (m, 2 6 H), 3.73 (s, 3 H), 4.02 (td, J=8.8, 4.8 Hz, 1 H), 4.66 (br. s., 1 H), 4.94 (s, 2 H), 5.56 (s, 2 H), 5.85 (d, J=8.8 Hz, 1 H), 6.09 (s, 1 H), 7.43 (s, 1 H)
80		360.19	361	0.63, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=6.9 Hz, 3 H), 1.14 - 1.34 (m, 4 H), 1.37 - 1.51 (m, 1 H), 1.52 - 1.67 (m, 1 H), 3.36 - 3.48 (m, 2 H), 3.99 - 6 4.11 (m, 1 H), 4.69 (br. s., 1 H), 5.10 (s, 2 H), 5.54 (s, 2 H), 6.00 (d, J=8.8 Hz, 1 H), 7.46 (s, 1 H), 7.68 (br. s., 1 H), 7.72 (dd, J=7.3, 1.3 Hz, 1 H), 7.93 - 8.02 (m, 2 H), 8.03 (s, 1 H)
81		348.23	349	0.73, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=7.2 Hz, 3 H), 1.11 - 1.37 (m, 4 H), 1.47 - 1.63 (m, 2 H), 1.63 - 1.79 (m, 2 H), 2.13 (s, 3 H), 3.40 (t, J=6.3 Hz, 6 2 H), 3.75 (s, 3 H), 4.30 (m, J=8.0 Hz, 1 H), 5.08 (s, 2 H), 6.21 (s, 1 H), 7.48 (br. s., 2 H), 7.55 (d, J=5.3 Hz, 1 H), 8.11 (d, J=8.8 Hz, 1 H), 11.96 (d, J=5.3 Hz, 1 H)
82		372.22	373	1.15, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.77 - 0.91 (m, 3 H) 1.17 - 1.37 (m, 4 H) 1.75 - 1.91 (m, 2 H) 1.93 - 2.09 (m, 2 H) 2.74 (t, J=7.65 Hz, 2 H) 3.63 (s, 3 H) 3.87 (q, J=6.02 Hz, 2 H) 4.58 (q, J=7.28 Hz, 1 H) 5.71 (br. s., 2 H) 6.53 (d, J=8.28 Hz, 1 H) 7.15 - 7.35 (m, 5 H) 7.43 (br. s., 1 H)

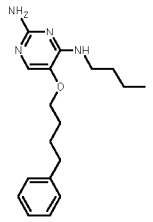
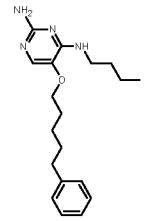
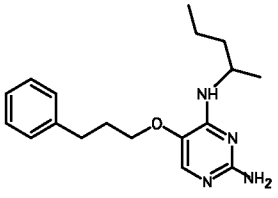
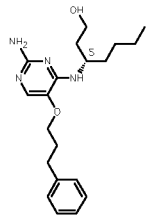
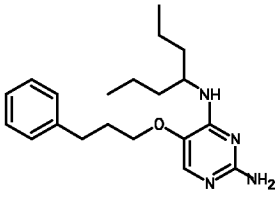
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
83		316.19	317	0.95, D	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.3 Hz, 3 H), 1.17 - 1.31 (m, 2 H), 1.33 - 1.46 (m, 2 H), 3.16 - 3.25 (m, 2 H), 3.71 (dd, J=5.5, 3.7 Hz, 2 H), 3.96 (dd, J=5.5, 3.7 Hz, 2 H), 4.55 (s, 2 H), 5.60 (s, 2 H), 6.26 (t, J=5.5 Hz, 1 H), 7.26 - 7.37 (m, 5 H), 7.41 (s, 1 H)
84		303.17	304	0.65, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.28 Hz, 3 H) 1.20 - 1.31 (m, 2 H) 1.36 - 1.49 (m, 2 H) 3.17 - 3.27 (m, 2 H) 4.16 (dd, J=5.27, 3.26 Hz, 2 H) 4.26 - 4.44 (m, 2 H) 5.57 (s, 2 H) 6.25 (s, 1 H) 6.93 - 7.09 (m, 2 H) 7.44 (s, 1 H) 8.30 - 8.52 (m, 2 H)
85		305.20	306	0.45, D	not available
86		263.15	264	0.58, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 - 0.92 (m, 2 H) 1.18 - 1.36 (m, 2 H) 1.42 - 1.57 (m, 2 H) 3.23 - 3.52 (m, 2 H) 5.16 (s, 2 H) 7.53 (br. s., 2 H) 7.61 (d, J=5.02 Hz, 1 H) 8.09 (s, 1 H) 8.38 (br.s, 1 H) 12.08 (s, 1 H)
87		377.21	378	0.7, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.40 Hz, 3 H) 1.19 - 1.32 (m, 2 H) 1.49 - 1.59 (m, 1 H) 1.66 - 1.79 (m, 2 H) 1.83 - 1.93 (m, 1 H) 3.35 - 3.48 (m, 2 H) 3.90 (s, 3 H) 4.09 (s, 3 H) 4.35 - 4.51 (m, 1 H) 5.32 (s, 2 H) 7.52 (br. s., 2 H) 7.56 (br. s., 1 H) 7.71 (d, J=5.27 Hz, 1 H) 8.53 (d, J=6.27 Hz, 1 H) 8.82 (br. s., 1 H) 12.01 (d, J=4.27 Hz, 1 H)
88		395.16	396	0.48, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 - 0.91 (m, 3 H), 1.21 - 1.36 (m, 2 H), 1.47 - 1.55 (m, 2 H), 1.57 - 1.78 (m, 2 H), 3.45 (dd, J=6.9, 6.1 Hz, 4 H), 3.76 (s, 3 H), 4.06 - 4.22 (m, 1 H), 4.89 (s, 2 H), 5.31 (s, 2 H), 6.07 (br. s., 1 H), 6.40 (d, J=6.1 Hz, 1 H), 7.47 (s, 1 H), 7.67 (d, J=6.5 Hz, 1 H)

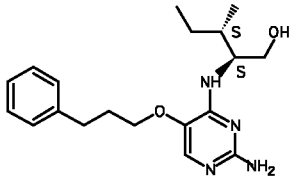
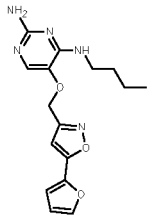
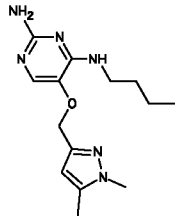
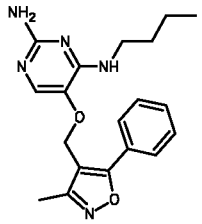
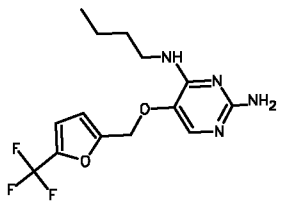
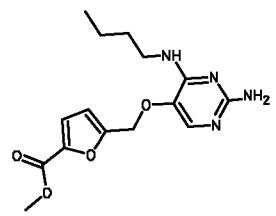
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
89		325.21	326	0.87, H	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=6.78 Hz, 3 H) 0.99 (t, J=7.15 Hz, 3 H) 1.11 - 1.36 (m, 4 H) 1.49 (m, J=5.00 Hz, 2 H) 1.77 (q, J=6.78 Hz, 2 H) 2.97 (quin, J=6.78 Hz, 2 H) 3.67 (s, 3 H) 3.90 (m, J=4.00 Hz, 2 H) 4.05 - 4.25 (m, 1 H) 5.40 (br. s., 2 H) 6.17 (d, J=9.03 Hz, 1 H) 6.99 (br. t, J=1.00, 1.00 Hz, 1 H) 7.35 (s, 1 H)
90		180.10	181	0.47, D	^{1}H NMR (400 MHz, CHLOROFORM- *d*) δ ppm 3.78 (s, 3 H), 4.06 (tt, J=5.7, 1.5 Hz, 2 H), 4.44 (br. s., 2 H), 5.15 (dq, J=10.3, 1.4 Hz, 1 H), 5.23 (br. s, 1 H), 5.23 (dq, J=17.1, 1.7 Hz, 1 H), 5.94 (ddt, J=17.2, 10.3, 5.6, 5.6 Hz, 1 H), 7.39 (s, 1 H)
91		208.13	209	0.65, D	^{1}H NMR (400 MHz, CHLOROFORM- *d*) δ ppm 0.05 - 0.13 (m, 2 H), 0.42 - 0.52 (m, 2 H), 0.65 - 0.80 (m, 1 H), 1.50 (q, J=7.0 Hz, 2 H), 3.49 (td, J=7.0, 5.9 Hz, 2 H), 3.73 - 3.80 (m, 3 H), 4.42 (br. s., 2 H), 5.27 (br. s., 1 H), 7.36 (s, 1 H)
92		212.13	213	0.42, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=7.5 Hz, 3 H), 1.37 - 1.54 (m, 1 H), 1.54 - 1.71 (m, 1 H), 3.38 (dt, J=10.7, 5.3 Hz, 1 H), 3.45 (dt, J=10.4, 5.1Hz, 1 H), 3.68 (s, 3 H), 3.85 - 4.02 (m, 1 H), 4.66 (t, J=5.4 Hz, 1 H), 5.45 (br. s, 1 H), 5.88 (d, J=8.8 Hz, 1 H), 7.36 (s, 1 H)
93		240.16	241	0.61, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (dd, J=6.7, 4.9 Hz, 6 H), 1.31 - 1.49 (m, 2 H), 1.50 - 1.64 (m, 1 H), 3.37 - 3.44 (m, 2 H), 3.67 (s, 3 H), 4.15 (tq, 6 J=9.7, 4.8 Hz, 1 H), 4.65 (br. s., 1 H), 5.42 (s, 2 H), 5.88 (d, J=9.3 Hz, 1 H), 7.35 (s, 1 H)
94		330.17	331	1.65, E	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.98 (t, J=7.3 Hz, 3 H), 1.35 - 1.49 (m, 2 H), 1.54 - 1.74 (m, 2 H), 3.36 - 3.47 (m, 2 H), 3.88 - 3.96 (m, 3 H), 4.84 (br. s., 2 H), 5.19 (s, 2 H), 6.15 (br. s., 1 H), 6.94 - 7.05 (m, 2

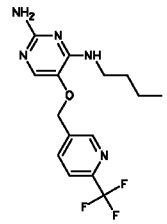
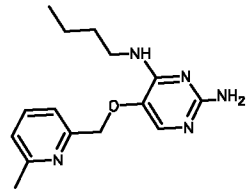
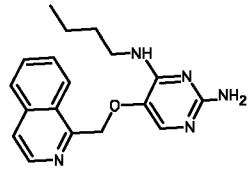
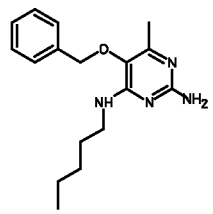
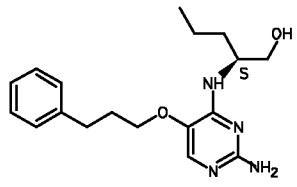
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					H), 7.45 (s, 1 H), 7.86 - 7.98 (m, 2 H)
95		290.15	291	1.67, E	^{1}H NMR (300 MHz, METHANOL-d_4) δ ppm 0.84 (t, J=7.3 Hz, 3 H), 1.17 - 1.33 (m, 2 H), 1.36 - 1.52 (m, 2 H), 3.26 (t, J=7.1 Hz, 2 H), 4.86 (s, 2 H), 6.94 - 7.05 (m, 2 H), 7.17 (s, 1 H), 7.29 - 7.40 (m, 2 H), 3 labile protons not seen.
96		300.20	301	1.09, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.0 Hz, 3 H), 1.21 - 1.36 (m, 4 H), 1.46 - 1.51 (m, 2 H), 1.52 (d, J=6.5 Hz, 3 H), 3.22 - 3.29 (m, 2 H), 5.17 (q, J=6.3 Hz, 1 H), 5.41 (s, 2 H), 6.34 (t, J=5.9 Hz, 1 H), 7.20 (s, 1 H), 7.23 - 7.29 (m, 1 H), 7.29 - 7.36 (m, 2 H), 7.38 - 7.44 (m, 2 H)
97		214.12	215	0.53, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 1.68 - 1.87 (m, 4 H), 3.46 (q, J=6.5 Hz, 2 H), 3.77 (s, 2 H), 4.43 (br. s, 2 H), 4.38 - 4.48 (m, 1 H), 4.55 (t, J=5.9 Hz, 1 H), 5.19 (br. s., 1 H), 7.37 (s, 1 H)
98		286.18	287	0.96, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.29 (dq, J=14.9, 7.3 Hz, 2 H), 1.45 - 1.50 (m, 2 H), 1.52 (d, J=6.5 Hz, 3 H), 3.23 - 3.30 (m, 2 H), 5.16 (q, J=6.4 Hz, 1 H), 5.41 (s, 2 H), 6.33 (t, J=5.9 Hz, 1 H), 7.20 (s, 1 H), 7.23 - 7.29 (m, 1 H), 7.29 - 7.36 (m, 2 H), 7.37 - 7.44 (m, 2 H)
99		286.18	287	0.97, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.29 (dq, J=15.0, 7.3 Hz, 2 H), 1.44 - 1.50 (m, 2 H), 1.52 (d, J=6.3 Hz, 3 H), 3.23 - 3.29 (m, 2 H), 5.17 (q, J=6.3 Hz, 1 H), 5.42 (s, 2 H), 6.35 (t, J=5.9 Hz, 1 H), 7.20 (s, 1 H), 7.22 - 7.29 (m, 1 H), 7.29 - 7.36 (m, 2 H), 7.38 - 7.44 (m, 2 H)

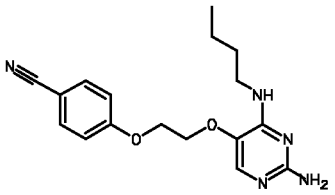
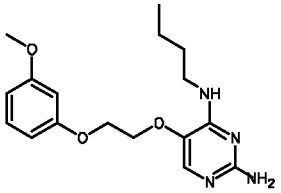
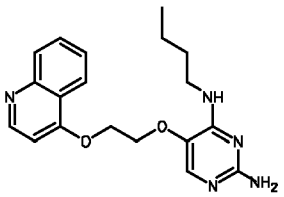
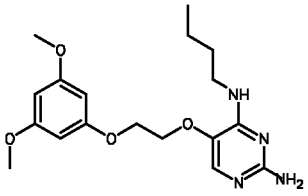
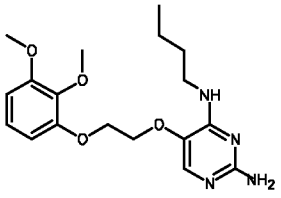
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
100		266.17	267	1.4, E	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.95 (t, J=7.3 Hz, 3 H), 1.32 - 1.48 (m, 2 H), 1.51 - 1.61 (m, 2 H), 1.62 - 1.73 (m, 1 H), 1.88 - 1.98 (m, 2 H), 1.98 - 2.10 (m, 1 H), 3.38 (td, J=7.0, 5.8 Hz, 2 H), 3.73 - 3.81 (m, 1 H), 3.82 - 3.95 (m, 3 H), 4.13 - 4.27 (m, 1 H), 4.73 (br. s., 2 H), 5.84 (br. s., 1 H), 7.42 (s, 1 H)
101		273.16	274	1.28, F	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.89 (t, J=7.4 Hz, 3 H), 1.26 - 1.43 (m, 2 H), 1.47 - 1.61 (m, 2 H), 3.35 (td, J=7.0, 5.8 Hz, 2 H), 4.53 (br. s., 2 H), 4.97 (s, 2 H), 5.91 (br. s., 1 H), 7.16 - 7.24 (m, 1 H), 7.30 (d, J=7.8 Hz, 1 H), 7.38 (s, 1 H), 7.66 (td, J=7.7, 1.6 Hz, 1 H), 8.55 (d, J=4.7 Hz, 1 H)
102		366.22	367	0.83, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.88 (t, J =7.3 Hz, 3 H), 1.26 (dq, J =14.8, 7.3 Hz, 2 H), 1.38 - 1.50 (m, 2 H), 1.82 (d, J =7.3 Hz, 3 H), 3.12 - 3.29 (m, 2 H), 4.63 (d, J =12.5 Hz, 1 H), 4.87 (d, J =12.9 Hz, 1 H), 5.51 (s, 2 H), 5.58 (q, J =6.9 Hz, 1 H), 6.08 (t, J =5.9 Hz, 1 H), 7.01 (s, 1 H), 7.12 - 7.18 (m, 2 H), 7.25 - 7.30 (m, 1 H), 7.27 (s, 1 H), 7.30 - 7.37 (m, 2 H), 7.97 (s, 1 H)
103		376.19	377	2.52, F	^{1}H NMR (300 MHz, CHLOROFORM-d) δ ppm 0.82 (t, J=7.3 Hz, 3 H), 1.18 - 1.33 (m, 2 H), 1.38 - 1.51 (m, 2 H), 3.22 - 3.34 (m, 2 H), 4.50 (br. s., 2 H), 4.92 (s, 2 H), 5.05 - 5.15 (m, 1 H), 7.13 (s, 1 H), 7.32 - 7.41 (m, 4 H), 7.43 - 7.52 (m, 1 H), 7.63 - 7.74 (m, 4 H)
104		268.19	269	1.74, F	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.89 (td, J=7.4, 5.0 Hz, 6 H), 1.20 (s, 3 H), 1.25 - 1.39 (m, 2 H), 1.44 - 1.62 (m, 4 H), 1.81 - 2.20 (m, 1 H), 3.33 (td, J=7.0, 5.8 Hz, 2 H), 3.60 - 3.69 (m, 2 H), 4.55 (br. s., 2 H), 5.40 (br. s., 1 H), 7.19 (s, 1 H)

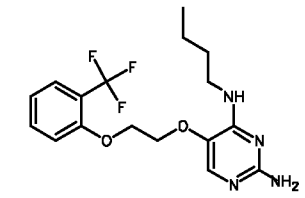
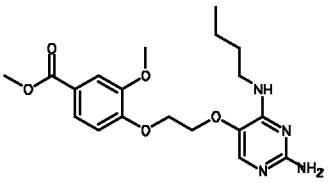
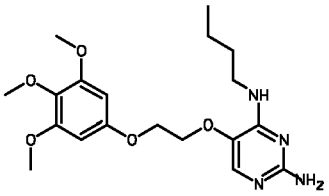
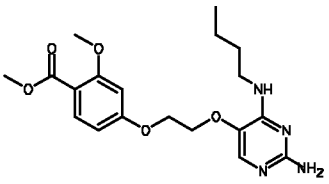
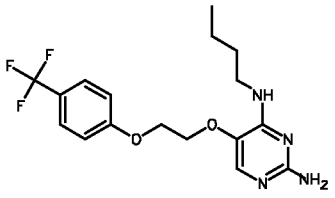
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
105		254.17	255	1.56, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.84 - 0.93 (m, 3 H), 1.27 (s, 6 H), 1.28 - 1.39 (m, 2 H), 1.45 - 1.58 (m, 2 H), 3.34 (td, J=7.0, 5.8 Hz, 2 H), 3.62 - 3.65 (m, 3 H), 4.62 (br. s., 2 H), 5.37 - 5.55 (m, 1 H), 7.32 (s, 1 H)
106		240.16	241	0.64, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.15 - 1.35 (m, 2 H), 1.44 - 1.60 (m, 2 H), 3.23 (s, 2 H), 3.35 - 3.38 (m, 1 H), 3.40 - 3.47 (m, 1H), 3.77 (s, 3 H), 4.36 - 4.49 (m, 1 H), 7.39 (s, 1 H), 7.44 (br. s., 2 H), 8.16 (d, J=8.8 Hz, 1 H), 11.88 (br. s., 1 H)
107		336.18	337	2.57, F	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.97 (t, J=7.2 Hz, 3 H), 1.32 - 1.48 (m, 2 H), 1.52 - 1.66 (m, 2 H), 2.01 - 2.14 (m, 2 H), 2.78 (t, J=7.5 Hz, 2 H), 3.33 - 3.47 (m, 2 H), 3.91 (t, J=6.1 Hz, 2 H), 4.55 (br. s., 2 H), 5.12 (br. s., 1 H), 6.74 - 6.88 (m, 2 H), 7.07 - 7.22 (m, 1 H), 7.31 (s, 1 H)
108		378.11	379	2.62, F	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.24 - 1.40 (m, 2 H), 1.44 - 1.58 (m, 2 H), 1.96 - 2.09 (m, 2 H), 2.73 - 2.90 (m, 2 H), 3.26 - 3.43 (m, 2 H), 3.87 (t, J=6.1 Hz, 2 H), 4.43 (br. s., 2 H), 5.09 (br. s., 1 H), 6.93 - 7.06 (m, 1 H), 7.11 - 7.22 (m, 3 H), 7.48 (d, J=8.2 Hz, 1 H)
109		334.16	335	2.68, F	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.83 - 0.97 (m, 3 H), 1.25 - 1.41 (m, 2 H), 1.50 (dt, J=14.6, 7.3 Hz, 2 H), 1.93 - 2.05 (m, 2 H), 2.68 (t, J=7.5 Hz, 2 H), 3.24 - 3.40 (m, 2 H), 3.82 (t, J=6.2 Hz, 2 H), 4.42 (br. s., 2 H), 4.95 (br. s., 1 H), 7.01 - 7.12 (m, 2 H), 7.16 - 7.22 (m, 3 H)

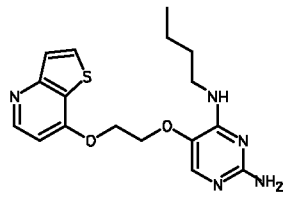
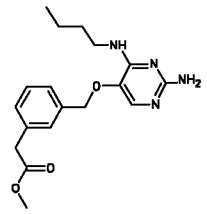
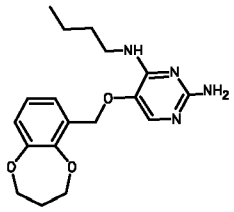
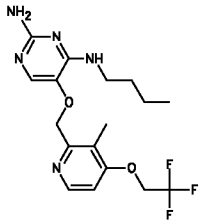
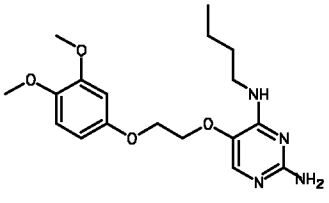
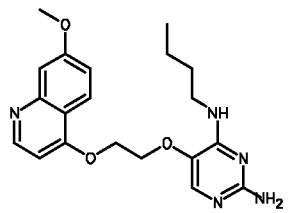
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
110		314.21	315	2.64, F	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.88 (t, J=7.3 Hz, 3 H), 1.31 (dq, J=15.0, 7.3 Hz, 2 H), 1.43 - 1.57 (m, 2 H), 1.65 - 1.78 (m, 4 H), 2.61 (t, J=6.9 Hz, 2 H), 3.27 - 3.38 (m, 2 H), 3.77 - 3.89 (m, 2 H), 4.48 (br. s., 2 H), 5.09 (br. s., 1 H), 7.09 - 7.16 (m, 3 H), 7.18 - 7.23 (m, 3 H)
111		328.23	329	2.75, F	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.88 (t, J=7.3 Hz, 3 H), 1.24 - 1.43 (m, 4 H), 1.44 - 1.55 (m, 2 H), 1.55 - 1.65 (m, 2 H), 1.66 - 1.77 (m, 2 H), 2.57 (t, J=7.6 Hz, 2 H), 3.25 - 3.37 (m, 2 H), 3.80 (t, J=6.5 Hz, 2 H), 4.45 (br. s., 2 H), 5.07 (br. s., 1 H), 7.07 - 7.15 (m, 3 H), 7.17 - 7.24 (m, 3 H)
112		314.21	315	1.12, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.28 Hz, 3 H) 1.11 (d, J=6.53 Hz, 3 H) 1.20 - 1.35 (m, 2 H) 1.36 - 1.59 (m, 2 H) 1.94 - 2.05 (m, 2 H) 2.65 - 2.78 (m, 2 H) 3.83 (t, J=6.40 Hz, 2 H) 4.07 - 4.18 (m, 1 H) 5.60 (s, 2 H) 5.99 (d, J=8.53 Hz, 1 H) 7.14 - 7.32 (m, 5 H) 7.33 (s, 1 H)
113		358.24	359	1.04, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.79 - 0.87 (m, 3 H) 1.18 - 1.31 (m, 4 H) 1.42 - 1.59 (m, 2 H) 1.59 - 1.75 (m, 2 H) 1.94 - 2.02 (m, 2 H) 2.66 - 2.75 (m, 2 H) 3.41 - 3.50 (m, 2 H) 3.79 - 3.87 (m, 2 H) 4.10 - 4.18 (m, 1 H) 4.44 - 4.49 (m, 1 H) 5.71 (br. s., 2 H) 6.25 (br. s., 1 H) 7.12 - 7.31 (m, 5 H) 7.33 (s, 1 H)
114		342.24	343	1.25, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.28 Hz, 6 H) 1.13 - 1.38 (m, 4 H) 1.38 - 1.54 (m, 4 H) 1.95 - 2.09 (m, 2 H) 2.72 (t, J=7.15 Hz, 2 H) 3.71 - 3.85 (m, 2 H) 4.01 - 4.21 (m, 1 H) 5.59 (br. s., 2 H) 5.92 (d, J=9.03 Hz, 1 H) 7.29 (s, 1 H) 7.15 - 7.43 (m, 5 H)

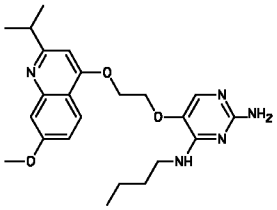
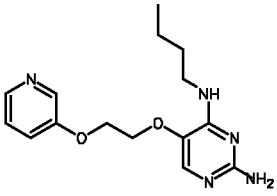
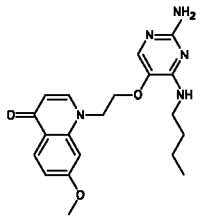
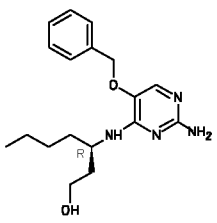
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
115		344.22	345	0.98, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.82 - 0.88 (m, 3 H) 0.89 (d, J=6.86 Hz, 3 H) 1.06 - 1.17 (m, 1 H) 1.44 - 1.53 (m, 1 H) 1.71 - 1.78 (m, 1 H) 1.95 - 2.04 (m, 2 H) 2.72 (t, J=7.67 Hz, 2 H) 3.48 - 3.60 (m, 2 H) 3.84 - 3.90 (m, 2 H) 3.90 - 3.96 (m, 1 H) 4.38 (t, J=5.25 Hz, 1 H) 5.21 (br. s., 2 H) 5.55 (d, J=8.88 Hz, 1 H) 7.14 - 7.31 (m, 5 H) 7.37 (s, 1 H)
116		329.15	330	0.9, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.40 Hz, 3 H) 1.21 - 1.33 (m, 2 H) 1.42 - 1.53 (m, 2 H) 3.24 - 3.31 (m, 2 H) 5.04 (s, 2 H) 5.58 (s, 2 H) 6.52 (t, J=5.90 Hz, 1 H) 6.73 (dd, J=3.51, 1.76 Hz, 1 H) 6.99 (s, 1 H) 7.14 (d, J=3.26 Hz, 1 H) 7.49 (s, 1 H) 7.96 (dd, J=1.76, 0.50 Hz, 1 H)
117		290.19	291	0.75, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.40 Hz, 3 H) 1.28 (quin, J=1.00 Hz, 2 H) 1.43 - 1.52 (m, 2 H) 2.22 (s, 3 H) 3.21 - 3.27 (m, 2 H) 3.68 (s, 3 H) 4.76 (s, 2 H) 5.48 (s, 2 H) 6.10 (s, 1 H) 6.26 (t, J=5.65 Hz, 1 H) 7.40 (s, 1 H)
118		353.19	354	0.97, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.40 Hz, 3 H) 1.20 - 1.30 (m, 2 H) 1.40 - 1.49 (m, 2 H) 2.27 (s, 3 H) 3.21 - 3.29 (m, 2 H) 4.87 (s, 2 H) 5.56 (s, 2 H) 6.40 (t, J=5.77 Hz, 1 H) 7.37 (s, 1 H) 7.53 - 7.60 (m, 3 H) 7.71 - 7.77 (m, 2 H)
119		330.13	331	0.99, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.28 Hz, 3 H) 1.21 - 1.33 (m, 2 H) 1.41 - 1.52 (m, 2 H) 3.16 - 3.29 (m, 2 H) 4.95 (s, 2 H) 5.58 (s, 2 H) 6.39 (t, J=5.77 Hz, 1 H) 6.78 (d, J=3.01 Hz, 1 H) 7.21 (dd, J=3.51, 1.25 Hz, 1 H) 7.38 (s, 1 H)
120		320.15	321	0.79, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.40 Hz, 3 H) 1.22 - 1.32 (m, 2 H) 1.42 - 1.52 (m, 2 H) 3.21 - 3.28 (m, 2 H) 3.81 (s, 3 H) 4.94 (s, 2 H) 5.57 (s, 2 H) 6.38 (t, J=5.65 Hz, 1 H) 6.75 (d, J=3.51 Hz, 1 H) 7.29 (d, J=3.51 Hz, 1 H)

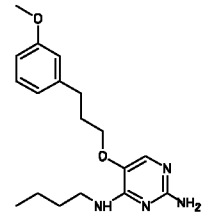
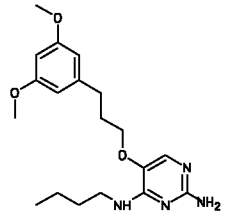
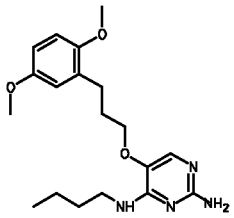
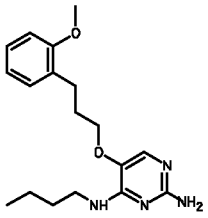
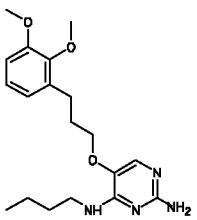
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					7.39 (s, 1 H)
121		341.15	342	0.89, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.28 Hz, 3 H) 1.20 - 1.35 (m, 2 H) 1.39 - 1.55 (m, 2 H) 3.21 - 3.30 (m, 2 H) 5.11 (s, 2 H) 5.54 (s, 2 H) 6.58 (s, 1 H) 7.47 (s, 1 H) 7.93 (d, J=8.03 Hz, 1 H) 8.14 - 8.22 (m, 1 H) 8.85 - 8.93 (m, 1 H)
122		287.17	288	0.79, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.28 Hz, 3 H) 1.23 - 1.37 (m, 2 H) 1.45 - 1.58 (m, 2 H) 2.48 (s, 3 H) 3.29 - 3.33 (m, 2 H) 4.93 (s, 2 H) 5.54 (s, 2 H) 6.75 (s, 1 H) 7.20 (d, J=7.78 Hz, 1 H) 7.37 (d, J=7.53 Hz, 1 H) 7.40 (s, 1 H) 7.71 (t, J=7.65 Hz, 1 H)
123		323.17	324	0.87, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.82 - 0.91 (m, 3 H) 1.18 - 1.28 (m, 2 H) 1.38 - 1.47 (m, 2 H) 3.19 - 3.27 (m, 2 H) 5.50 (s, 2 H) 5.52 (s, 2 H) 6.49 (s, 1 H) 7.44 (s, 1 H) 7.71 (ddd, J=8.41, 7.03, 1.13 Hz, 1 H) 7.81 (ddd, J=8.09, 6.96, 1.25 Hz, 1 H) 7.85 (d, J=5.52 Hz, 1 H) 8.02 (d, J=8.03 Hz, 1 H) 8.38 - 8.42 (m, 1 H) 8.48 (d, J=5.77 Hz, 1 H)
124		300.20	301	1.08, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.2 Hz, 3 H), 1.16 - 1.37 (m, 4 H), 1.53 (quin, J=7.3 Hz, 2 H), 2.03 (s, 3 H), 3.37 (q, J=6.6 Hz, 2 H), 4.36 (br. s., 2 H), 4.83 (s, 2 H), 7.29 - 7.58 (m, 5 H), 8.30 (t, J=5.9 Hz, 1 H), 12.68 (br. s., 1 H)
125		330.21	331	0.9, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.28 Hz, 3 H) 1.19 - 1.38 (m, 2 H) 1.40 - 1.51 (m, 1 H) 1.51 - 1.62 (m, 1 H) 1.94 - 2.02 (m, 2 H) 2.66 - 2.76 (m, 2 H) 3.38 - 3.48 (m, 2 H) 3.83 (td, J=6.34, 2.64 Hz, 2 H) 4.00 - 4.10 (m, 1 H) 4.69 (br. s., 1 H) 5.48 (s, 2 H) 5.72 - 5.79 (m, 1 H) 7.05 - 7.33 (m, 5 H) 7.35 (s, 1 H)

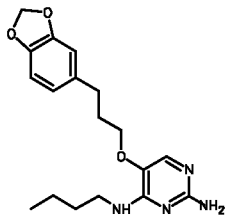
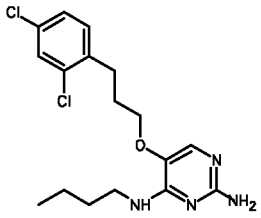
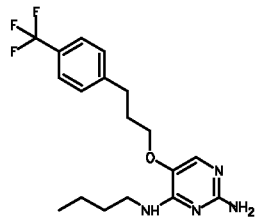
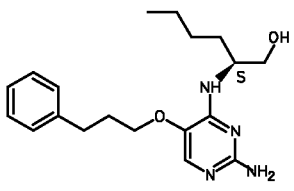
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
126		327.17	328	0.84, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.40 Hz, 3 H) 1.19 - 1.31 (m, 2 H) 1.37 - 1.48 (m, 2 H) 3.20 - 3.27 (m, 2 H) 4.13 - 4.23 (m, 2 H) 4.30 - 4.42 (m, 2 H) 5.57 (s, 2 H) 6.22 (s, 1 H) 7.12 - 7.20 (m, 2 H) 7.45 (s, 1 H) 7.75 - 7.83 (m, 2 H)
127		332.18	333	0.94, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.80 - 0.91 (m, 3 H) 1.20 - 1.32 (m, 2 H) 1.37 - 1.50 (m, 2 H) 3.17 - 3.28 (m, 2 H) 3.73 (s, 3 H) 4.13 (dd, J=5.52, 3.26 Hz, 2 H) 4.23 (dd, J=5.52, 3.26 Hz, 2 H) 5.56 (s, 2 H) 6.20 (s, 1 H) 6.49 - 6.59 (m, 3 H) 7.16 - 7.22 (m, 1 H) 7.45 (s, 1 H)
128		353.19	354	0.79, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.76 (t, J=7.28 Hz, 3 H) 1.11 - 1.21 (m, 2 H) 1.32 (t, J=7.15 Hz, 2 H) 3.15 - 3.22 (m, 2 H) 4.32 - 4.36 (m, 2 H) 4.52 - 4.56 (m, 2 H) 5.57 (s, 2 H) 6.24 (s, 1 H) 7.07 (d, J=5.27 Hz, 1 H) 7.55 (s, 1 H) 7.52 - 7.58 (m, 1 H) 7.74 (ddd, J=8.41, 6.90, 1.25 Hz, 1 H) 7.95 (d, J=8.03 Hz, 1 H) 8.12 (dd, J=8.28, 1.00 Hz, 1 H) 8.73 (d, J=5.27 Hz, 1 H)
129		362.20	363	0.92, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.78 - 0.93 (m, 3 H) 1.12 - 1.35 (m, 2 H) 1.39 - 1.54 (m, 2 H) 3.18 - 3.28 (m, 2 H) 4.07 - 4.17 (m, 2 H) 4.21 (dd, J=5.52, 3.01 Hz, 2 H) 5.58 (br. s., 2 H) 6.09 - 6.12 (m, 1 H) 6.14 (d, J=2.26 Hz, 2 H) 6.21 (s, 1 H) 7.45 (s, 1 H)
130		362.20	363	0.87, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.28 Hz, 3 H) 1.20 - 1.31 (m, 2 H) 1.39 - 1.48 (m, 2 H) 3.21 - 3.28 (m, 2 H) 3.67 (s, 3 H) 3.77 (s, 3 H) 4.11 - 4.18 (m, 2 H) 4.22 - 4.29 (m, 2 H) 5.56 (s, 2 H) 6.18 (t, J=5.90 Hz, 1 H) 6.66 - 6.74 (m, 2 H) 6.96 - 7.01 (m, 1 H) 7.47 (s, 1 H)

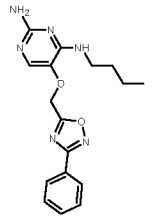
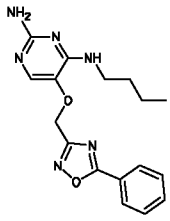
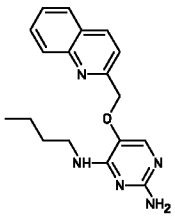
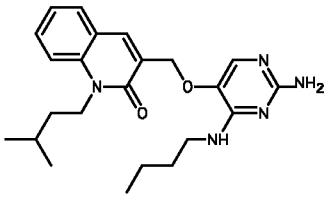
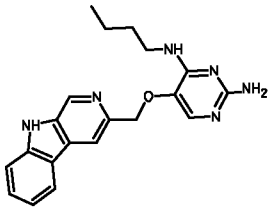
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
131		370.16	371	1.05, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.28 Hz, 3 H) 1.18 - 1.30 (m, 2 H) 1.36 - 1.47 (m, 2 H) 3.17 - 3.28 (m, 2 H) 4.12 - 4.22 (m, 2 H) 4.33 - 4.43 (m, 2 H) 5.61 (s, 2 H) 5.98 (s, 1 H) 7.09 - 7.15 (m, 1 H) 7.33 (d, J=8.53 Hz, 1 H) 7.47 (s, 1 H) 7.63 (d, J=7.78 Hz, 2 H)
132		390.19	391	0.86, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.28 Hz, 3 H) 1.19 - 1.32 (m, 2 H) 1.37 - 1.49 (m, 2 H) 3.17 - 3.28 (m, 2 H) 3.83 (s, 3 H) 3.82 (s, 3 H) 4.16 (dd, J=5.27, 3.26 Hz, 2 H) 4.32 (dd, J=5.27, 3.26 Hz, 2 H) 5.59 (s, 2 H) 6.18 (s, 1 H) 7.14 (d, J=8.53 Hz, 1 H) 7.45 - 7.53 (m, 2 H) 7.59 (dd, J=8.53, 2.01 Hz, 1 H)
133		392.21	393	0.84, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.40 Hz, 3 H) 1.20 - 1.34 (m, 2 H) 1.40 - 1.52 (m, 2 H) 3.14 - 3.28 (m, 2 H) 3.58 (s, 3 H) 3.75 (s, 6 H) 4.13 (dd, J=5.52, 3.26 Hz, 2 H) 4.23 (dd, J=5.52, 3.01 Hz, 2 H) 5.58 (s, 2 H) 6.22 (s, 1 H) 6.28 (s, 2 H) 7.46 (s, 1 H)
134		390.19	391	0.83, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.40 Hz, 3 H) 1.20 - 1.30 (m, 2 H) 1.39 - 1.48 (m, 2 H) 3.20 - 3.28 (m, 2 H) 3.73 (s, 3 H) 3.81 (s, 3 H) 4.13 - 4.19 (m, 2 H) 4.34 (dd, J=5.27, 3.26 Hz, 2 H) 5.56 (s, 2 H) 6.20 (s, 1 H) 6.63 (dd, J=8.66, 2.38 Hz, 1 H) 6.68 (d, J=2.26 Hz, 1 H) 7.46 (s, 1 H) 7.71 (d, J=8.53 Hz, 1 H)
135		370.16	371	1.06, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.40 Hz, 3 H) 1.18 - 1.34 (m, 2 H) 1.36 - 1.47 (m, 2 H) 3.17 - 3.27 (m, 2 H) 4.13 - 4.23 (m, 2 H) 4.29 - 4.41 (m, 2 H) 5.57 (s, 2 H) 6.21 (s, 1 H) 7.17 (m, J=8.53 Hz, 2 H) 7.46 (s, 1 H) 7.67 (m, J=8.53 Hz, 2 H)

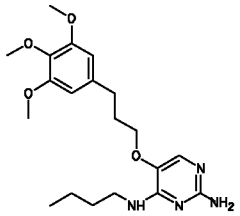
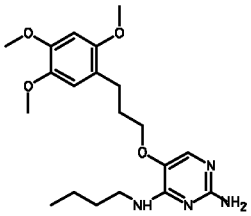
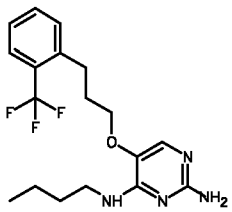
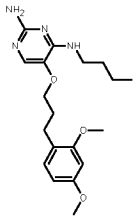
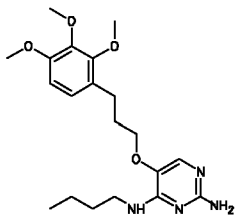
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
136		359.14	360	0.77, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.81 (t, J=7.40 Hz, 3 H) 1.16 - 1.25 (m, 2 H) 1.34 - 1.42 (m, 2 H) 3.19 - 3.25 (m, 2 H) 4.24 - 4.28 (m, 2 H) 4.55 - 4.60 (m, 2 H) 5.57 (s, 2 H) 6.16 (s, 1 H) 7.07 (d, J=5.27 Hz, 1 H) 7.50 (s, 1 H) 7.53 (d, J=5.52 Hz, 1 H) 8.07 (d, J=5.52 Hz, 1 H) 8.55 (d, J=5.52 Hz, 1 H)
137		344.18	345	0.88, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.40 Hz, 3 H) 1.22 - 1.34 (m, 2 H) 1.44 - 1.54 (m, 2 H) 3.25 - 3.30 (m, 2 H) 3.61 (s, 3 H) 3.69 (s, 2 H) 4.93 (s, 2 H) 5.50 (s, 2 H) 6.39 (s, 1 H) 7.22 (d, J=6.00 Hz, 1 H) 7.33 (s, 1 H) 7.28 - 7.37 (m, 2 H) 7.38 (s, 1 H)
138		344.18	345	0.94, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.40 Hz, 3 H) 1.20 - 1.37 (m, 2 H) 1.42 - 1.57 (m, 2 H) 2.02 - 2.19 (m, 2 H) 3.26 - 3.32 (m, 2 H) 4.07 - 4.18 (m, 4 H) 4.89 (s, 2 H) 5.52 (s, 2 H) 6.31 (s, 1 H) 6.88 - 7.04 (m, 2 H) 7.12 (d, J=6.70 Hz, 1 H) 7.37 (s, 1 H)
139		385.17	386	0.93, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.28 Hz, 3 H) 1.22 - 1.36 (m, 2 H) 1.42 - 1.57 (m, 2 H) 2.20 (s, 3 H) 3.22 - 3.29 (m, 2 H) 4.84 - 4.98 (m, 2 H) 5.01 (s, 2 H) 5.50 (s, 2 H) 6.59 (s, 1 H) 7.13 (d, J=5.77 Hz, 1 H) 7.40 (s, 1 H) 8.34 (d, J=5.52 Hz, 1 H)
140		362.20	363	0.71, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.28 Hz, 3 H) 1.18 - 1.34 (m, 2 H) 1.37 - 1.52 (m, 2 H) 3.23 - 3.28 (m, 2 H) 3.69 (s, 3 H) 3.74 (s, 3 H) 4.07 - 4.15 (m, 2 H) 4.15 - 4.26 (m, 2 H) 5.56 (s, 2 H) 6.20 (s, 1 H) 6.47 (dd, J=8.66, 2.89 Hz, 1 H) 6.60 (d, J=3.01 Hz, 1 H) 6.85 (d, J=8.78 Hz, 1 H) 7.45 (s, 1 H)
141		383.20	384	0.82, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.77 (t, J=7.28 Hz, 3 H) 1.12 - 1.26 (m, 2 H) 1.28 - 1.37 (m, 2 H) 3.15 - 3.25 (m, 2 H) 3.90 (s, 3 H) 4.29 - 4.34 (m, 2 H) 4.51 (dd, J=5.14, 3.14 Hz, 2 H) 5.58 (s, 2 H) 6.24 (s, 1 H) 6.93 (d, J=5.27 Hz, 1

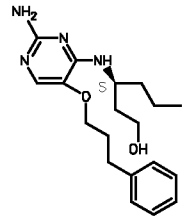
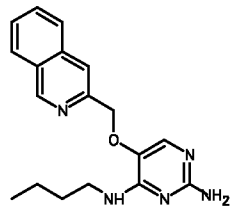
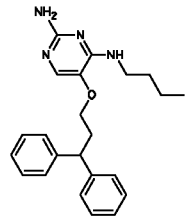
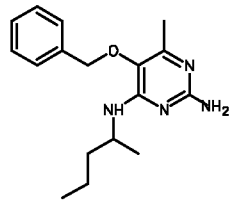
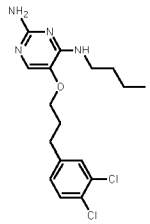
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					H) 7.17 (dd, J=9.16, 2.64 Hz, 1 H) 7.32 (d, J=2.51 Hz, 1 H) 7.52 (s, 1 H) 8.00 (d, J=9.29 Hz, 1 H) 8.65 (d, J=5.27 Hz, 1 H)
142		425.24	426	1, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.77 (t, J=7.40 Hz, 3 H) 1.13 - 1.19 (m, 2 H) 1.28 - 1.35 (m, 6 H) 1.28 - 1.35 (m, 2 H) 3.05 - 3.15 (m, 1 H) 3.16 - 3.21 (m, 2 H) 3.89 (s, 3 H) 4.29 - 4.32 (m, 2 H) 4.50 - 4.52 (m, 2 H) 5.57 (s, 2 H) 6.22 (s, 1 H) 6.83 (s, 1 H) 7.08 (dd, J=9.29, 2.51 Hz, 1 H) 7.25 (d, J=2.51 Hz, 1 H) 7.52 (s, 1 H) 7.93 (d, J=9.03 Hz, 1 H)
143		303.17	304	0.68, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.40 Hz, 3 H) 1.20 - 1.35 (m, 2 H) 1.38 - 1.51 (m, 2 H) 3.22 - 3.28 (m, 2 H) 4.09 - 4.25 (m, 2 H) 4.27 - 4.40 (m, 2 H) 5.60 (s, 2 H) 6.27 (s, 1 H) 7.31 - 7.37 (m, 1 H) 7.41 - 7.45 (m, 1 H) 7.45 (s, 1 H) 8.19 (dd, J=4.52, 1.25 Hz, 1 H) 8.33 (d, J=2.76 Hz, 1 H)
144		383.20	384	0.65, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.28 Hz, 3 H) 1.19 - 1.25 (m, 2 H) 1.39 (t, J=7.40 Hz, 2 H) 3.17 - 3.22 (m, 2 H) 3.90 (s, 3 H) 4.11 - 4.22 (m, 2 H) 4.59 (m, J=4.90, 4.90 Hz, 2 H) 5.53 (s, 2 H) 5.86 (s, 1 H) 5.97 (d, J=7.53 Hz, 1 H) 6.99 (d, J=8.80 Hz, 1 H) 7.11 (d, J=2.26 Hz, 1 H) 7.32 (s, 1 H) 8.04 (d, J=7.78 Hz, 1 H) 8.09 (d, J=9.03 Hz, 1 H)
145		330.21	331	2.18, F	^{1}H NMR (300 MHz, CHLOROFORM-d) δ ppm 0.76 - 0.87 (m, 3 H), 1.14 - 1.31 (m, 5 H), 1.33 - 1.57 (m, 3 H), 1.80 (m, J=11.4, 5.1, 2.7 Hz, 1 H), 3.31 - 3.43 (m, 1 H), 3.45 - 3.56 (m, 1 H), 4.03 (d, J=3.3 Hz, 1 H), 4.44 (s, 2 H), 4.81 - 4.89 (m, 1 H), 4.91 (s, 2 H), 7.27 - 7.35 (m, 5 H), 7.39 (s, 1 H)

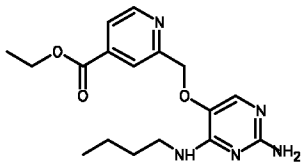
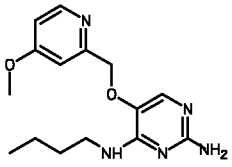
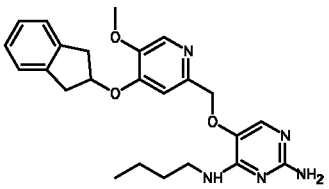
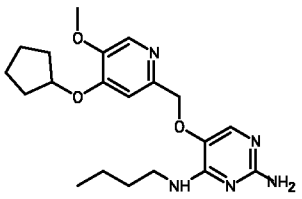
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
146		330.21	331	1.03, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.91 (t, J=7.3 Hz, 3 H), 1.30 (dq, J=14.9, 7.4 Hz, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 1.97 - 2.08 (m, 2 H), 2.69 - 2.78 (m, 6 2 H), 3.42 (q, J=6.8 Hz, 2 H), 3.73 (s, 3 H), 3.90 (t, J=6.3 Hz, 2 H), 6.73 - 6.78 (m, 1 H), 6.78 - 6.83 (m, 2 H), 7.17 - 7.25 (m, 1 H), 7.37 (s, 1 H), 7.43 (br. s., 2 H), 8.32 (t, J=6.0 Hz, 1 H), 11.83 (br. s., 1 H)
147		360.22	361	1.02, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.24 - 1.37 (m, 2 H), 1.55 (t, J=7.3 Hz, 2 H), 1.96 - 2.07 (m, 2 H), 2.65 - 2.74 (m, 2 H), 3.42 (q, J=6.9 Hz, 2 H), 3.71 (s, 6 H), 3.89 (t, J=6.1 Hz, 2 H), 6.31 - 6.35 (m, 1 H), 6.38 (d, J=2.3 Hz, 2 H), 7.34 (s, 1 H), 7.39 (br. s., 2 H), 8.31 (s, 1 H)
148		360.22	361	1.03, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.23 - 1.36 (m, 2 H), 1.49 - 1.60 (m, 2 H), 1.92 - 2.04 (m, 2 H), 2.68 (t, J=7.5 Hz, 2 H), 3.41 (q, 6 J=6.8 Hz, 2 H), 3.67 (s, 3 H), 3.71 (s, 3 H), 3.89 (t, J=6.3 Hz, 2 H), 6.69 - 6.77 (m, 2 H), 6.84 - 6.91 (m, 1 H), 7.34 (s, 1 H), 7.41 (br. s., 2 H), 8.31 (t, J=5.9 Hz, 1 H), 11.70 (s, 1 H)
149		330.21	331	1.06, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.24 - 1.37 (m, 2 H), 1.49 - 1.61 (m, 2 H), 1.92 - 2.05 (m, 2 H), 2.67 - 2.76 (m, 2 H), 3.41 (q, 6 J=6.9 Hz, 2 H), 3.76 (s, 3 H), 3.90 (t, J=6.3 Hz, 2 H), 6.87 (td, J=7.4, 1.0 Hz, 1 H), 6.96 (d, J=7.5 Hz, 1 H), 7.11 - 7.23 (m, 2 H), 7.33 (s, 1 H), 7.40 (br. s., 2 H), 8.31 (t, J=5.9 Hz, 1 H), 11.67 (br. s., 1 H)
150		360.22	361	1.02, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.24 - 1.36 (m, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 1.93 - 2.04 (m, 2 H), 2.69 - 2.76 (m, 2 H), 6 3.41 (q, J=6.8 Hz, 2 H), 3.70 (s, 3 H), 3.78 (s, 3 H), 3.91 (t, J=6.4 Hz, 2 H),

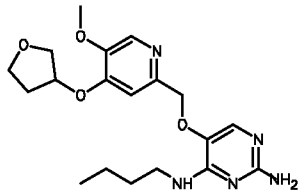
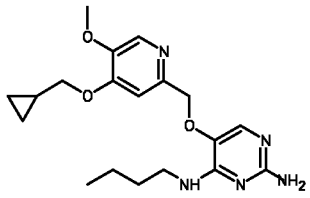
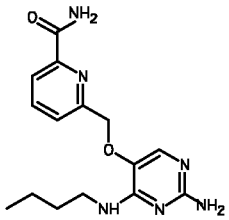
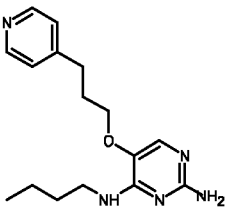
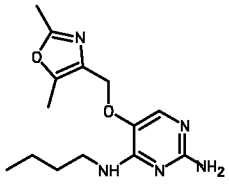
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					6.79 (dd, J=7.5, 1.5 Hz, 1 H), 6.87 - 6.92 (m, 1 H), 6.99 (t, J=7.9 Hz, 1 H), 7.36 (s, 1 H), 7.44 (br. s., 2 H), 8.31 (t, J=6.0 Hz, 1 H), 11.81 (s, 1 H)
151		344.18	345	1, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.91 (t, J=7.3 Hz, 3 H), 1.30 (dq, J=14.9, 7.4 Hz, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 1.93 - 2.04 (m, 2 H), 2.68 (t, J=7.5 6 Hz, 2 H), 3.42 (q, J=6.8 Hz, 2 H), 3.88 (t, J=6.1 Hz, 2 H), 5.94 - 5.99 (m, 2 H), 6.67 (dd, J=7.9, 1.6 Hz, 1 H), 6.82 (d, J=6.0 Hz, 1 H), 6.83 (s, 1 H), 7.36 (s, 1 H), 7.42 (br. s., 2 H), 8.31 (t, J=5.9 Hz, 1 H), 11.77 (br. s., 1 H)
152		368.12	369	1.13, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 - 0.95 (m, 3 H), 1.24 - 1.36 (m, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 1.97 - 2.07 (m, 2 H), 2.82 - 2.90 (m, 2 H), 3.42 6 (q, J=6.8 Hz, 2 H), 3.92 (t, J=6.1 Hz, 2 H), 7.37 (s, 1 H), 7.38 - 7.40 (m, 2 H), 7.43 (br. s., 2 H), 7.55 - 7.61 (m, 1 H), 8.32 (t, J=5.9 Hz, 1 H), 11.80 (br. s., 1 H)
153		368.18	369	1.15, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.91 (t, J=7.3 Hz, 3 H), 1.31 (dq, J=14.9, 7.4 Hz, 2 H), 1.56 (quin, J=7.3 Hz, 2 H), 1.99 - 2.11 (m, 2 H), 2.87 (t, J=7.8 6 Hz, 2 H), 3.38 - 3.47 (m, 2 H), 3.92 (t, J=6.1 Hz, 2 H), 7.38 (s, 1 H), 7.43 (br. s., 1 H), 7.48 (d, J=8.0 Hz, 2 H), 7.66 (d, J=8.0 Hz, 2 H), 8.33 (t, J=6.0 Hz, 1 H), 11.83 (br. s., 1 H)
154		344.22	345	0.98, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=6.90 Hz, 3 H) 1.22 - 1.36 (m, 4 H) 1.44 - 1.67 (m, 2 H) 1.95 - 2.08 (m, 2 H) 2.73 (t, J=7.65 Hz, 2 H) 3.41 - 3.64 (m, 2 H) 3.81 - 3.96 (m, 2 H) 4.05 - 4.20 (m, 1 H) 4.80 (br. s., 1 H) 6.69 (br. s., 2 H) 6.99 (d, J=8.53 Hz, 1 H) 7.14 - 7.34 (m, 5 H) 7.39 (s, 1 H) 7.90 (br. s., 1 H)

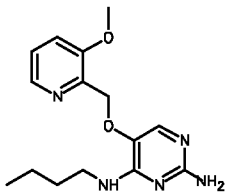
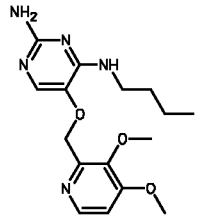
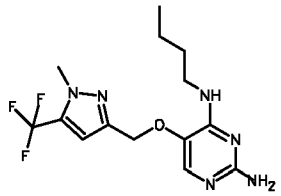
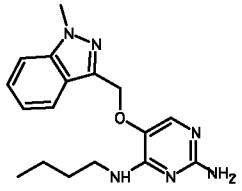
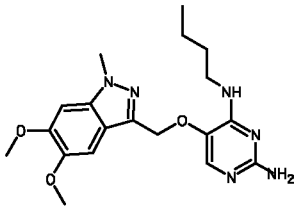
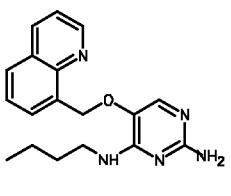
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
155		340.16	341	0.99, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=1.00 Hz, 3 H) 1.20 - 1.37 (m, 2 H) 1.47 - 1.60 (m, 2 H) 3.39 - 3.47 (m, 2 H) 5.53 (s, 2 H) 7.55 - 7.67 (m, 5 H) 7.71 (s, 1 H) 7.97 - 8.08 (m, 2 H) 8.59 (s, 1 H) 12.05 (br. s., 1 H)
156		340.16	341	0.97, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.40 Hz, 3 H) 1.21 - 1.33 (m, 2 H) 1.44 - 1.57 (m, 2 H) 3.36 - 3.46 (m, 2 H) 5.34 (s, 2 H) 7.58 (br. s., 2 H) 7.67 (s, 1 H) 7.63 - 7.70 (m, 2 H) 7.72 - 7.78 (m, 1 H) 8.10 - 8.18 (m, 2 H) 8.50 (s, 1 H) 11.98 (br. s., 1 H)
157		323.17	324	0.33, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.40 Hz, 3 H) 1.25 - 1.37 (m, 2 H) 1.50 - 1.61 (m, 2 H) 3.39 - 3.50 (m, 2 H) 5.39 (s, 2 H) 7.54 (br. s., 2 H) 7.59 (d, J=4.77 Hz, 1 H) 7.69 (t, J=7.40 Hz, 1 H) 7.81 - 7.91 (m, 2 H) 8.08 (d, J=8.28 Hz, 1 H) 8.12 (d, J=8.03 Hz, 1 H) 8.57 (d, J=8.78 Hz, 1 H) 8.68 (br. s., 1 H) 11.94 (br. s., 1 H)
158		409.25	410	1.21, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.91 (t, J=7.28 Hz, 3 H) 1.00 (d, J=6.78 Hz, 6 H) 1.31 - 1.41 (m, 2 H) 1.48 - 1.63 (m, 4 H) 1.70 - 1.80 (m, 1 H) 3.33 - 3.42 (m, 2 H) 4.28 - 4.37 (m, 2 H) 4.75 (s, 2 H) 5.62 (s, 2 H) 7.08 (t, J=1.00 Hz, 1 H) 7.31 (t, J=7.28 Hz, 1 H) 7.49 (s, 1 H) 7.57 (d, J=8.53 Hz, 1 H) 7.66 (dd, J=7.15, 1.38 Hz, 1 H) 7.79 (dd, J=7.78, 1.25 Hz, 1 H) 8.12 (s, 1 H)
159		362.19	363	0.89, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.28 Hz, 3 H) 1.27 - 1.37 (m, 2 H) 1.49 - 1.57 (m, 2 H) 3.32 - 3.39 (m, 2 H) 5.10 (s, 2 H) 5.53 (s, 2 H) 6.83 (s, 1 H) 7.23 - 7.28 (m, 1 H) 7.48 (s, 1 H) 7.56 (dd, J=6.90, 1.13 Hz, 1 H) 7.59 - 7.62 (m, 1 H) 8.25 (d, J=8.03 Hz, 1 H) 8.28 (s, 1 H) 8.88 (d, J=1.00 Hz, 1 H) 11.64 (s, 1 H)

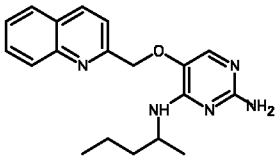
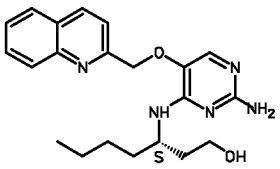
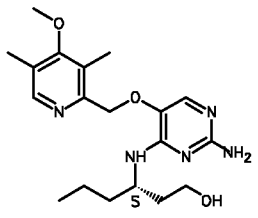
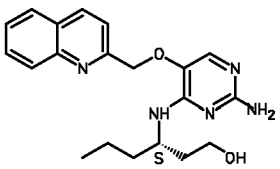
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
160		390.23	391	0.95, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 1.30 (dq, J=14.9, 7.3 Hz, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 1.94 - 2.12 (m, 2 H), 2.70 (t, J=7.7 6 Hz, 2 H), 3.37 - 3.44 (m, 2 H), 3.62 (s, 3 H), 3.70 - 3.79 (m, 6 H), 3.89 (t, J=6.3 Hz, 2 H), 6.51 (s, 2 H), 7.27 (br. s., 2 H), 7.39 (s, 1 H), 8.15 (t, J=5.6 Hz, 1 H)
161		390.23	391	0.97, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 1.30 (dq, J=14.9, 7.4 Hz, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 1.89 - 2.03 (m, 2 H), 2.64 (t, J=7.3 6 Hz, 2 H), 3.38 - 3.46 (m, 2 H), 3.65 (s, 3 H), 3.74 (s, 3 H), 3.76 (s, 3 H), 3.87 (t, J=6.4 Hz, 2 H), 6.66 (s, 1 H), 6.75 (s, 1 H), 7.39 (s, 1 H), 7.48 (br. s., 2 H), 8.30 (t, J=5.9 Hz, 1 H)
162		368.18	369	1.15, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.31 (dq, J=14.9, 7.4 Hz, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 1.98 - 2.09 (m, 2 H), 2.87 - 2.96 (m, 6 2 H), 3.36 - 3.44 (m, 2 H), 3.97 (t, J=6.3 Hz, 2 H), 7.17 (br. s., 2 H), 7.42 (s, 1 H), 7.43 (t, J=7.5 Hz, 1 H), 7.55 (d, J=7.8 Hz, 1 H), 7.63 (t, J=7.5 Hz, 1 H), 7.69 (d, J=7.8 Hz, 1 H), 7.97 (t, J=5.6 Hz, 1 H)
163		360.22	361	1.05, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.30 (dq, J=14.9, 7.4 Hz, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 1.88 - 2.01 (m, 2 H), 2.63 (t, J=7.4 6 Hz, 2 H), 3.37 - 3.44 (m, 2 H), 3.75 (s, 3 H), 3.73 (s, 3 H), 3.86 (t, J=6.4 Hz, 2 H), 6.44 (dd, J=8.3, 2.5 Hz, 1 H), 6.52 (d, J=2.3 Hz, 1 H), 7.02 (d, J=8.0 Hz, 1 H), 7.29 (br. s., 2 H), 7.35 (s, 1 H), 8.14 (t, J=5.9 Hz, 1 H)
164		390.23	391	1.02, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 1.30 (dq, J=14.9, 7.3 Hz, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 1.90 - 2.03 (m, 2 H), 2.66 (t, J=7.5 6 Hz, 2 H), 3.37 - 3.46 (m, 2 H), 3.73 (s, 3 H), 3.76 (s, 6 H), 3.86 - 3.95 (m, 2 H), 6.72 (d, J=8.5 Hz, 1 H), 6.86 (d,

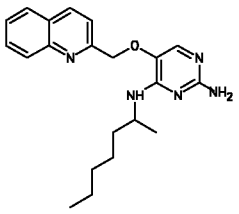
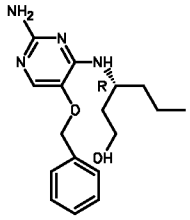
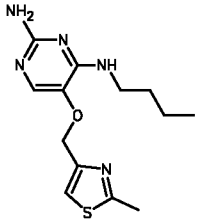
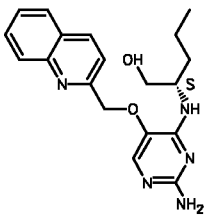
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					J=8.5 Hz, 1 H), 7.38 (s, 1 H), 7.46 (br. s., 2 H), 8.30 (t, J=5.9 Hz, 1 H), 11.94 (br. s., 1 H)
165		344.22	345	0.97, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.40 Hz, 3 H) 1.21 - 1.34 (m, 2 H) 1.41 - 1.57 (m, 2 H) 1.57 - 1.70 (m, 2 H) 1.94 - 2.01 (m, 2 H) 2.69 - 2.75 (m, 2 H) 3.38 - 3.46 (m, 2 H) 3.82 (td, J=6.34, 1.88 Hz, 2 H) 4.11 - 4.18 (m, 1 H) 4.45 (t, J=5.02 Hz, 1 H) 5.48 (s, 2 H) 6.00 (d, J=8.78 Hz, 1 H) 7.11 - 7.31 (m, 5 H) 7.33 (s, 1 H)
166		323.17	324	5.32, G	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.40 Hz, 3 H) 1.02 - 1.14 (m, 2 H) 1.23 - 1.38 (m, 2 H) 1.46 - 1.59 (m, 2 H) 3.36 - 3.46 (m, 2 H) 5.13 (s, 2 H) 5.55 (s, 2 H) 6.79 (br. s., 1 H) 7.48 (s, 1 H) 7.70 (ddd, J=8.16, 6.90, 1.00 Hz, 1 H) 7.78 - 7.85 (m, 1 H) 8.00 (d, J=1.00 Hz, 1 H) 7.99 (s, 1 H) 8.16 (d, J=7.53 Hz, 1 H) 9.34 (s, 1 H)
167		376.23	377	1.18, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.92 (t, J=7.3 Hz, 3 H), 1.27 - 1.38 (m, 2 H), 1.51 - 1.63 (m, 2 H), 3.40 - 3.48 (m, 2 H), 3.78 (t, J=6.1 Hz, 2 H), 4.32 (t, J=8.0 Hz, 1 H), 7.13 - 7.22 (m, 2 H), 7.25 - 7.36 (m, 10 H), 7.49 (br. s., 2 H), 8.33 (t, J=6.0 Hz, 1 H), 12.01 (s, 1 H). 214
168		300.20	301	1.06, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.3 Hz, 3 H), 1.12 (d, J=6.8 Hz, 3 H), 1.19 - 1.31 (m, 2 H), 1.39 - 1.50 (m, 1 H), 1.52 - 1.64 (m, 1 H), 2.03 (s, 3 H), 4.07 (br. s., 2 H), 4.15 - 4.27 (m, 1 H), 4.78 - 4.91 (m, 2 H), 7.35 - 7.43 (m, 3 H), 7.44 - 7.48 (m, 2 H), 7.92 (d, J=8.8 Hz, 1 H)
169		368.12	369	1.19, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.91 (t, J=7.3 Hz, 3 H), 1.31 (dq, J=14.9, 7.4 Hz, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 1.96 - 2.09 (m, 2 H), 2.74 - 2.83 (m, 6 2 H), 3.42 (q, J=6.8 Hz, 2 H), 3.90 (t, J=6.1 Hz, 2 H), 7.25 (dd, J=8.3, 2.0 Hz, 1 H), 7.38 (s, 1 H), 7.44 (br. s., 1 H), 7.53 (d, J=2.0 Hz, 1 H), 7.55

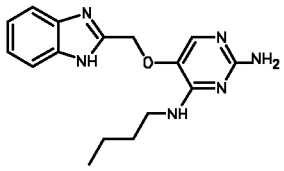
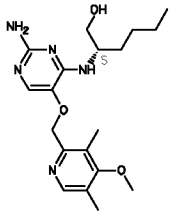
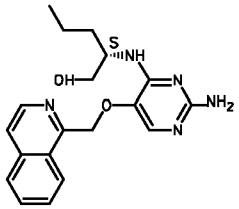
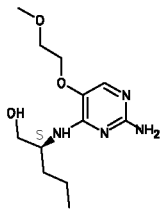
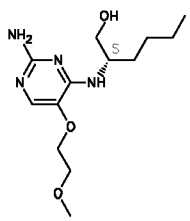
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					(d, J=8.0 Hz, 1 H), 8.31 (t, J=5.9 Hz, 1 H), 11.84 (br. s., 1 H)
170		345.18	346	0.36, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 1.24 - 1.33 (m, 2 H), 1.35 (t, J=7.2 Hz, 3 H), 1.55 (m, J=7.3, 7.3, 7.3, 7.3 Hz, 2 H), 3.42 (q, 6 J=6.9 Hz, 2 H), 4.38 (q, J=7.1 Hz, 1 H), 5.26 (s, 2 H), 7.56 (br. s, 2 H), 7.57 (s, 1 H), 7.85 (dd, J=5.0, 1.5 Hz, 1 H), 8.04 (s, 1 H), 8.61 (t, J=5.8 Hz, 1 H), 8.82 (dd, J=5.0, 0.8 Hz, 1 H), 12.05 (br. s., 1 H)
171		303.17	304	0.75, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.31 (dq, J=14.9, 7.4 Hz, 2 H), 1.59 (quin, J=7.3 Hz, 2 H), 3.44 (q, J=6.9 Hz, 2 H), 4.07 (s, 3 6 H), 5.36 (s, 2 H), 7.41 - 7.51 (m, 1 H), 7.52 - 7.69 (m, 4 H), 8.71 (d, J=6.8 Hz, 1 H), 9.06 (br. s., 1 H), 12.08 (br. s., 1 H)
172		435.23	436	1.02, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.4 Hz, 3 H), 1.22 - 1.36 (m, 2 H), 1.52 (quin, J=7.3 Hz, 2 H), 3.03 (dd, J=17.1, 2.0 Hz, 2 H), 3.32 (q, J=6.9 6 Hz, 2 H), 3.40 (dd, J=17.2, 6.1 Hz, 2 H), 3.77 (s, 3 H), 4.95 (s, 2 H), 5.29 - 5.37 (m, 1 H), 5.94 (br. s., 2 H), 7.07 (t, J=5.6 Hz, 1 H), 7.14 - 7.22 (m, 2 H), 7.22 - 7.30 (m, 3 H), 7.45 (s, 1 H), 8.12 (s, 1 H)
173		387.23	388	0.96, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 1.31 (dq, J=15.0, 7.4 Hz, 2 H), 1.52 - 1.62 (m, 2 H), 1.62 - 1.86 (m, 6 H), 1.97 - 2.13 (m, 2 H), 6 3.43 (q, J=6.9 Hz, 2 H), 3.96 (s, 3 H), 5.11 - 5.20 (m, 1 H), 5.35 (s, 2 H), 7.62 (br. s, 2 H), 7.65 (d, J=3.5 Hz, 1 H), 7.68 (s, 1 H), 8.30 (s, 1 H), 9.06 (t, J=5.4 Hz, 1 H), 12.21 (br. s., 1 H)

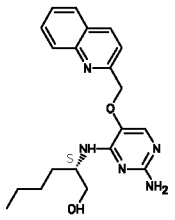
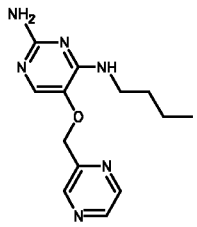
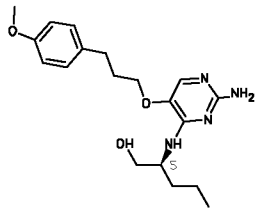
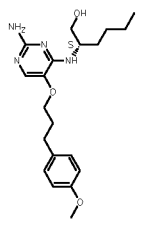
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
174		389.21	390	0.73, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 1.30 (dq, J=15.0, 7.4 Hz, 2 H), 1.57 (quin, J=7.3 Hz, 2 H), 1.98 - 2.10 (m, 1 H), 2.28 - 2.43 (m, 6 1 H), 3.42 (q, J=6.9 Hz, 2 H), 3.79 (td, J=8.4, 4.8 Hz, 1 H), 3.83 - 3.94 (m, 3 H), 3.95 (s, 3 H), 5.27 (s, 2 H), 5.30 - 5.37 (m, 1 H), 7.49 - 7.69 (m, 4 H), 8.31 (s, 1 H), 8.92 (br. s., 1 H), 11.99 - 12.13 (m, 1 H)
175		373.21	374	0.52, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.35 - 0.44 (m, 2 H), 0.61 - 0.69 (m, 2 H), 0.90 (t, J=7.4 Hz, 3 H), 1.23 - 1.39 (m, 1 H), 1.23 - 1.39 (m, 2 H), 1.58 6 (quin, J=7.3 Hz, 2 H), 3.43 (q, J=6.9 Hz, 2 H), 4.00 (s, 3 H), 4.18 (d, J=7.3 Hz, 2 H), 5.33 (s, 2 H), 7.62 (br. s, 2 H), 7.64 (d, J=5.0 Hz, 1 H), 7.69 (s, 1 H), 8.34 (s, 1 H), 9.04 (t, J=5.6 Hz, 1 H), 12.16 (d, J=4.8 Hz, 1 H)
176		316.16	317	0.65, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.4 Hz, 3 H), 1.28 (dq, J=15.0, 7.4 Hz, 2 H), 1.44 - 1.56 (m, 2 H), 3.29 (q, J=6.9 Hz, 2 H), 5.09 (s, 2 H), 5.52 6 (br. s, 2 H), 6.59 (t, J=5.9 Hz, 1 H), 7.43 (s, 1 H), 7.68 (br. s., 1 H), 7.79 (dd, J=7.5, 1.3 Hz, 1 H), 7.96 (dd, J=7.5, 1.3 Hz, 1 H), 7.98 - 8.02 (m, 1 H), 8.04 (br. s, 1 H)
177		301.19	302	0.74, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.20 - 1.38 (m, 2 H), 1.44 - 1.56 (m, 2 H), 1.95 - 2.06 (m, 2 H), 2.73 - 2.80 (m, 2 H), 3.23 - 6 3.32 (m, 2 H), 3.82 (t, J=6.3 Hz, 2 H), 5.49 (s, 2 H), 6.32 (t, J=5.9 Hz, 1 H), 7.23 - 7.29 (m, 2 H), 7.34 (s, 1 H), 8.42 - 8.51 (m, 2 H)
178		291.17	292	0.77, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.20 - 1.34 (m, 2 H), 1.52 (quin, J=7.3 Hz, 2 H), 2.31 (s, 3 H), 2.36 (s, 3 H), 3.37 (q, J=6.8 Hz, 6 2 H), 4.85 (s, 2 H), 7.57 (br. s., 3 H), 8.32 (t, J=5.9 Hz, 1 H), 12.26 (br. s., 1 H)

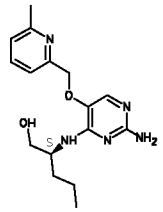
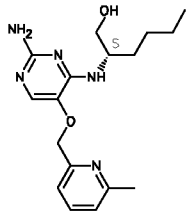
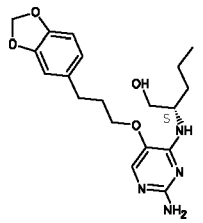
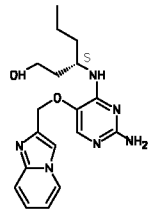
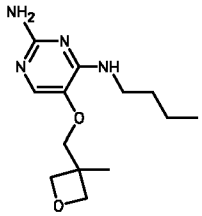
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
179		303.17	304	0.77, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.4 Hz, 3 H), 1.24 - 1.36 (m, 2 H), 1.51 - 1.59 (m, 2 H), 3.37 - 3.45 (m, 2 H), 3.92 (s, 3 H), 5.21 (s, 2 H), 7.50 (br. s., 2 H), 7.58 - 7.66 (m, 2 H), 7.79 (d, J=7.8 Hz, 1 H), 8.28 (d, J=4.3 Hz, 1 H), 8.73 - 8.91 (m, 1 H), 11.86 (d, J=5.5 Hz, 1 H)
180		333.18	334	0.75, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.40 Hz, 3 H) 1.18 - 1.37 (m, 2 H) 1.51 - 1.66 (m, 2 H) 3.33 - 3.53 (m, 2 H) 3.93 (s, 3 H) 4.17 (s, 3 H) 5.44 (s, 2 H) 7.62 (br. s., 2 H) 7.73 (d, J=7.03 Hz, 1 H) 7.83 (br. s., 1 H) 8.63 (d, J=6.78 Hz, 1 H) 9.58 (t, J=5.90 Hz, 1 H) 12.45 (br. s., 1 H)
181		344.16	345	0.93, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.28 Hz, 3 H) 1.17 - 1.32 (m, 2 H) 1.47 - 1.56 (m, 2 H) 3.35 - 3.43 (m, 2 H) 3.97 (s, 3 H) 5.02 (s, 2 H) 7.11 (s, 1 H) 7.48 (br. s., 2 H) 7.56 (s, 1 H) 8.35 (t, J=6.02 Hz, 1 H) 11.85 (br. s., 1 H)
182		326.19	327	0.75, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.28 Hz, 3 H) 1.15 - 1.31 (m, 2 H) 1.32 - 1.48 (m, 2 H) 3.15 - 3.26 (m, 3 H) 4.03 (s, 3 H) 5.26 (s, 2 H) 5.51 (s, 2 H) 6.28 (s, 1 H) 7.15 (td, J=7.53, 0.75 Hz, 1 H) 7.43 (s, 1 H) 7.36 - 7.49 (m, 1 H) 7.62 (d, J=8.53 Hz, 1 H) 7.79 - 7.89 (m, 1 H)
183		386.21	387	0.79, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=1.00 Hz, 3 H) 1.12 - 1.27 (m, 2 H) 1.31 - 1.46 (m, 2 H) 3.13 - 3.27 (m, 2 H) 3.77 (s, 3 H) 3.85 (s, 3 H) 3.96 (s, 3 H) 5.18 (s, 2 H) 5.48 (s, 2 H) 6.26 (s, 1 H) 7.12 (d, J=9.29 Hz, 2 H) 7.42 (s, 1 H)
184		323.17	324	0.93, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.48 - 1.60 (m, 2 H), 1.71 - 1.80 (m, 2 H), 3.43 - 3.49 (m, 2 H), 5.65 (s, 2 H), 7.56 (br. s., 2 H), 7.67 (d, J=5.0 Hz, 1 H), 7.71 - 7.77 (m, 2 H), 8.05 - 8.14 (m, 2 H), 8.60 (dd, J=8.3, 1.3 Hz, 1 H), 8.67 (t, J=5.9

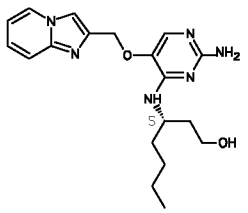
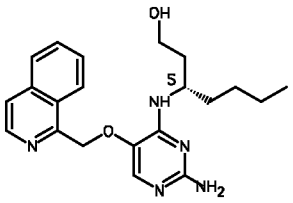
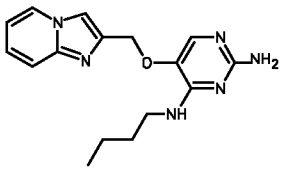
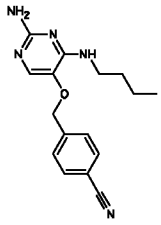
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					Hz, 1 H), 9.04 (dd, J=4.3, 1.8 Hz, 1 H), 12.01 (d, J=4.8 Hz, 1 H)
185		337.19	338	0.95, D	^{1}H NMR (400 MHz, CHLOROFORM- *d*) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.21 (d, J=6.5 Hz, 3 H), 1.28 - 1.40 (m, 2 H), 1.43 - 1.62 (m, 2 H), 3.45 (s, 2 H), 4.23 (dd, J=7.9, 7.2 Hz, 1 H), 5.29 (s, 2 H), 6.70 (d, J=8.5 Hz, 1 H), 7.40 (s, 1 H), 7.54 (d, J=8.5 Hz, 1 H), 7.56 - 7.60 (m, 1 H), 7.74 (ddd, J=8.5, 7.0, 1.4 Hz, 1 H), 7.85 (dd, J=8.0, 1.0 Hz, 1 H), 8.08 (d, J=8.5 Hz, 1 H), 8.22 (d, J=8.3 Hz, 1 H)
186		381.22	382	0.9, D	^{1}H NMR (400 MHz, CHLOROFORM- *d*) δ ppm 0.80 - 0.89 (m, 3 H), 1.20 - 1.35 (m, 5 H), 1.44 (d, J=3.5 Hz, 1 H), 1.59 (dd, J=8.3, 5.8 Hz, 2 H), 1.86 - 1.98 (m, 1 H), 3.11 - 3.40 (m, 2 H), 3.55 (dd, J=10.8, 3.0 Hz, 1 H), 3.59 (dd, J=5.0, 3.3 Hz, 1 H), 4.14 - 4.27 (m, 1 H), 5.25 (s, 2 H), 6.32 (d, J=8.8 Hz, 1 H), 7.46 (s, 1 H), 7.48 (d, J=8.5 Hz, 1 H), 7.57 (ddd, J=8.1, 7.0, 1.3 Hz, 1 H), 7.75 (ddd, J=8.5, 7.0, 1.4 Hz, 1 H), 7.84 (d, J=8.3 Hz, 1 H), 8.07 (d, J=8.5 Hz, 1 H), 8.21 (d, J=8.5 Hz, 1 H) taken on the free base
187		375.23	376	0.81, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.3 Hz, 3 H), 1.21 - 1.36 (m, 2 H), 1.47 - 1.63 (m, 1 H), 1.69 - 1.88 (m, 2 H), 1.89 - 2.04 (m, 1 H), 2.29 (s, 3 6 H), 2.43 (s, 3 H), 3.41 (t, J=6.5 Hz, 2 H), 4.03 (s, 3 H), 4.36 - 4.50 (m, 1 H), 5.41 (s, 2 H), 7.53 (br. s., 2 H), 7.86 (d, J=5.5 Hz, 1 H), 8.62 (s, 1 H), 9.19 (d, J=8.8 Hz, 1 H), 12.35 (d, J=5.3 Hz, 1 H)
188		367.20	368	0.78, H	^{1}H NMR (400 MHz, CHLOROFORM- *d*) δ ppm 0.85 (t, J=7.40 Hz, 3 H) 1.25 - 1.43 (m, 3 H) 1.50 - 1.59 (m, 2 H) 1.82 - 1.94 (m, 1 H) 2.92 - 3.32 (m, 1 H) 3.42 - 3.51 (m, 1 H) 3.53 - 3.60 (m, 1 H) 4.11 - 4.23 (m, 1 H) 4.83 (s, 2 H) 5.22 (s, 2 H) 5.73 (d, J=8.78 Hz,

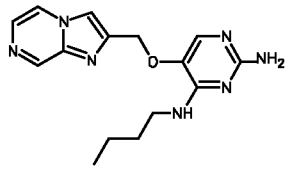
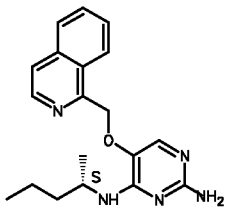
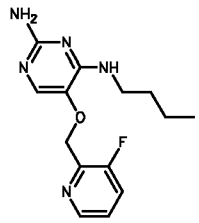
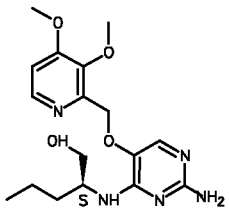
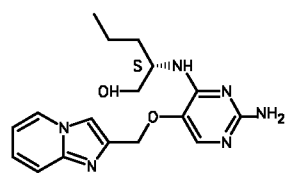
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					1 H) 7.46 (d, J=8.53 Hz, 1 H) 7.53 (s, 1 H) 7.55 - 7.59 (m, 1 H) 7.73 (ddd, J=8.47, 6.96, 1.38 Hz, 1 H) 7.82 (d, J=8.03 Hz, 1 H) 8.08 (d, J=8.28 Hz, 1 H) 8.18 (d, J=8.53 Hz, 1 H)
189		365.22	366	1.13, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.79 - 0.87 (m, 3 H), 1.15 - 1.21 (m, 4 H), 1.22 - 1.28 (m, 6 H), 4.16 - 4.40 (m, 1 H), 5.35 - 5.40 (m, 2 H), 7.40 - 7.48 (m, 2 H), 7.50 - 7.54 (m, 1 H), 7.62 - 7.68 (m, 1 H), 7.73 - 7.77 (m, 1 H), 7.78 - 7.85 (m, 1 H), 7.99 - 8.07 (m, 2 H), 8.31 - 8.38 (m, 1 H), 8.45 - 8.51 (m, 1 H), 11.47 - 11.58 (m, 1 H)
190		316.19	317	0.84, D	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.84 (t, J=7.4 Hz, 3 H), 1.14 - 1.55 (m, 6 H), 1.75 - 1.90 (m, 1 H), 3.30 - 3.43 (m, 1 H), 3.45 - 3.57 (m, 1 H), 4.06 (ddd, J=11.3, 5.2, 3.3 Hz, 1 H), 4.42 (s, 2 H), 4.80 - 4.86 (m, 1 H), 4.90 (s, 2 H), 7.27 - 7.34 (m, 5 H), 7.40 (s, 1 H)
191		293.13	294	0.71, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.28 Hz, 3 H) 1.12 - 1.40 (m, 2 H) 1.43 - 1.60 (m, 2 H) 2.68 (s, 3 H) 3.32 - 3.48 (m, 2 H) 5.07 (s, 2 H) 7.57 (br. s., 2 H) 7.61 (br. s., 1 H) 7.78 (s, 1 H) 8.45 (t, J=5.90 Hz, 1 H) 12.21 (br. s., 1 H)
192		353.19	354	0.78, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.4 Hz, 3 H), 1.16 - 1.35 (m, 2 H), 1.51 - 1.63 (m, 2 H), 3.48 - 3.55 (m, 2 H), 4.28 (d, J=6.0 Hz, 1 H), 5.41 (s, 2 H), 7.51 (br. s., 2 H), 7.58 (d, J=5.5 Hz, 1 H), 7.68 (td, J=7.5, 1.0 Hz, 1 H), 7.79 (d, J=8.5 Hz, 1 H), 7.85 (ddd, J=8.5, 7.0, 1.4 Hz, 1 H), 8.07 (d, J=7.3 Hz, 1 H), 8.11 (d, J=8.5 Hz, 1 H), 8.18 (d, J=9.0 Hz, 1 H), 8.55 (d, J=8.5 Hz, 1 H), 11.83 (d, J=5.5 Hz, 1 H)

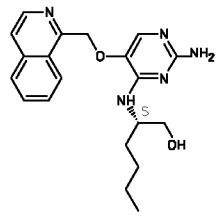
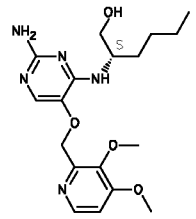
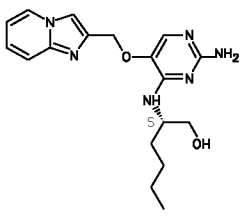
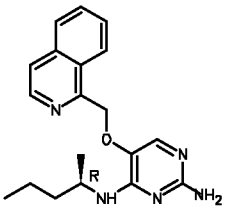
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
193		312.17	313	0.76, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.40 Hz, 3 H) 1.25 - 1.36 (m, 2 H) 1.47 - 1.55 (m, 2 H) 3.25 - 3.29 (m, 2 H) 5.13 (s, 2 H) 5.58 (s, 2 H) 6.71 (t, J=5.77 Hz, 1 H) 7.19 (br. s., 2 H) 7.51 (s, 1 H) 7.53 (br. s., 1 H) 7.59 (br. s., 1 H) 12.60 (br. s., 1 H)
194		375.23	376	0.82, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=6.9 Hz, 3 H), 1.14 - 1.35 (m, 4 H), 1.59 - 1.80 (m, 2 H), 2.29 (s, 3 H), 2.42 (s, 3 H), 3.51 - 3.61 (m, 2 H), 6 4.02 (s, 3 H), 4.26 - 4.39 (m, 1 H), 5.41 (s, 2 H), 7.54 (br. s., 2 H), 7.86 (d, J=3.3 Hz, 1 H), 8.61 (s, 1 H), 9.00 (d, J=8.0 Hz, 1 H), 12.41 (d, J=3.3 Hz, 1 H)
195		353.19	354	0.76, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.6 Hz, 3 H), 1.18 - 1.39 (m, 2 H), 1.56 - 1.69 (m, 1 H), 1.69 - 1.84 (m, 1 H), 3.53 - 3.68 (m, 2 H), 4.33 - 6 4.45 (m, 1 H), 6.08 (s, 2 H), 7.55 (br. s., 2 H), 7.96 - 8.08 (m, 2 H), 8.17 (t, J=7.5 Hz, 1 H), 8.33 (d, J=8.3 Hz, 1 H), 8.41 (d, J=6.3 Hz, 1 H), 8.65 (d, J=6.5 Hz, 1 H), 8.70 (d, J=8.5 Hz, 1 H), 9.10 - 9.28 (m, 1 H), 12.58 (br. s., 1 H)
196		270.17	271	1.37, F	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.95 (t, J=7.1 Hz, 3 H), 1.30 - 1.48 (m, 3 H), 1.49 - 1.67 (m, 2 H), 3.44 (s, 3 H), 3.55 - 3.64 (m, 1 H), 3.67 (t, J=4.4 Hz, 2 H), 3.73 - 3.80 (m, 1 H), 3.97 - 4.04 (m, 2 H), 4.09 (d, J=2.6 Hz, 1 H), 4.80 (br. s., 2 H), 5.91 (d, J=7.0 Hz, 1 H), 7.47 (s, 1 H)
197		284.18	285	1.62, F	^{1}H NMR (300 MHz, CHLOROFORM- *d*) δ ppm 0.86 - 0.97 (m, 3 H), 1.24 - 1.43 (m, 4 H), 1.46 - 1.72 (m, 2 H), 3.40 - 3.45 (m, 3 H), 3.48 (br. s, 1 H), 3.60 (dd, J=11.1, 6.7 Hz, 1 H), 3.67 (t, J=4.3 Hz, 2 H), 3.72 - 3.81 (m, 1 H), 4.00 (q, J=3.9 Hz, 2 H), 4.04 - 4.14 (m, 1 H), 4.92 (br. s., 2 H), 5.96 (d, J=7.4 Hz, 1 H), 7.45 (s,

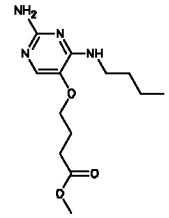
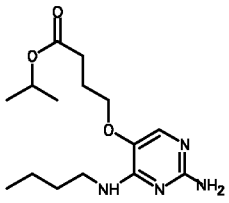
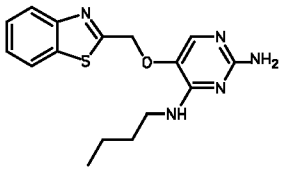
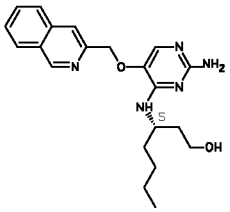
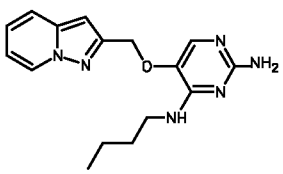
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					1 H)
198		367.20	368	0.85, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.77 - 0.84 (m, 3 H), 1.14 - 1.34 (m, 5 H), 1.48 (d, J=5.8 Hz, 2 H), 1.56 - 1.67 (m, 1 H), 3.39 - 3.51 (m, 2 H), 4.07 (d, J=5.0 Hz, 1 H), 4.72 (br. s., 1 H), 5.63 (s, 2 H), 6.35 (d, J=9.0 Hz, 1 H), 7.47 (s, 1 H), 7.62 (ddd, J=8.1, 6.8, 1.1 Hz, 1 H), 7.69 (d, J=8.5 Hz, 1 H), 7.79 (ddd, J=8.4, 6.9, 1.5 Hz, 1 H), 7.98 - 8.05 (m, 2 H), 8.41 (d, J=8.5 Hz, 1 H)
199		274.15	275	0.65, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.88 - 0.94 (m, 3 H), 1.20 - 1.37 (m, 2 H), 1.55 (quin, J=7.3 Hz, 2 H), 3.42 (q, J=6.8 Hz, 2 H), 5.22 (s, 2 H), 7.59 (br. s., 2 H), 7.66 (br. s., 1 H), 8.51 (t, J=5.9 Hz, 1 H), 8.68 (s, 2 H), 9.02 (s, 1 H), 12.24 (br. s., 1 H)
200		360.22	361	2.21, F	^{1}H NMR (300 MHz, CHLOROFORM-d) δ ppm 0.96 (t, J=7.2 Hz, 3 H), 1.36 - 1.50 (m, 3 H), 1.50 - 1.69 (m, 3 H), 2.00 - 2.14 (m, 2 H), 2.72 (t, J=7.4 Hz, 2 H), 3.58 - 3.66 (m, 1 H), 3.80 (s, 3 H), 3.91 (t, J=6.3 Hz, 2 H), 4.05 (d, J=5.9 Hz, 1 H), 4.59 (br. s., 2 H), 5.25 (d, J=6.9 Hz, 1 H), 6.80 - 6.88 (m, 2 H), 7.11 (d, J=8.5 Hz, 2 H), 7.34 (s, 1 H)
201		374.23	375	2.43, F	^{1}H NMR (300 MHz, CHLOROFORM-d) δ ppm 0.85 - 0.96 (m, 3 H), 1.23 - 1.43 (m, 5 H), 1.46 - 1.71 (m, 2 H), 1.99 - 2.13 (m, 2 H), 2.71 (t, J=7.5 Hz, 2 H), 3.57 - 3.66 (m, 1 H), 3.74 (d, J=3.2 Hz, 1 H), 3.78 (s, 3 H), 3.90 (t, J=6.3 Hz, 2 H), 4.03 (t, J=5.5 Hz, 1 H), 4.63 (br. s., 2 H), 5.26 (d, J=7.1 Hz, 1 H), 6.80 - 6.89 (m, 2 H), 7.10 (d, J=8.5 Hz, 2 H), 7.32 (s, 1 H)

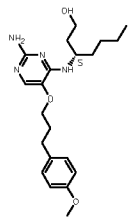
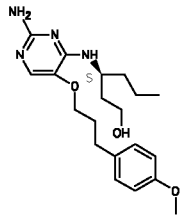
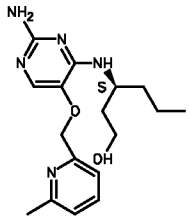
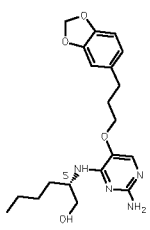
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
202		317.19	318	1.35, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.94 (t, J=7.3 Hz, 3 H), 1.32 - 1.52 (m, 3 H), 1.53 - 1.68 (m, 2 H), 2.59 (s, 3 H), 3.58 - 3.68 (m, 1 H), 3.74 - 3.84 (m, 1 H), 4.12 (td, J=6.9, 3.0 Hz, 1 H), 4.61 (br. s., 2 H), 4.99 (s, 2 H), 5.94 (d, J=7.1 Hz, 1 H), 7.15 (dd, J=11.7, 7.7 Hz, 2 H), 7.49 (s, 1 H), 7.62 (t, J=7.7 Hz, 1 H)
203		331.20	332	1.63, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.85 - 0.96 (m, 3 H), 1.18 - 1.46 (m, 5 H), 1.50 - 1.72 (m, 2 H), 2.59 (s, 3 H), 3.58 - 3.69 (m, 1 H), 3.75 - 3.84 (m, 1 H), 4.09 (td, J=6.9, 2.6 Hz, 1 H), 4.62 (br. s., 2 H), 5.00 (s, 2 H), 5.95 (d, J=7.0 Hz, 1 H), 7.15 (dd, J=12.3, 7.8 Hz, 2 H), 7.49 (s, 1 H), 7.62 (t, J=7.7 Hz, 1 H)
204		374.20	375	2.26, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.97 (t, J=7.2 Hz, 3 H), 1.32 - 1.49 (m, 3 H), 1.51 - 1.70 (m, 3 H), 1.98 - 2.14 (m, 2 H), 2.70 (t, J=7.5 Hz, 1 H), 3.59 - 3.71 (m, 1 H), 3.74 - 3.83 (m, 1 H), 3.91 (t, J=6.4 Hz, 1 H), 3.99 - 4.15 (m, 1 H), 4.68 (br. s., 2 H), 5.26 - 5.33 (m, 2 H), 5.92 - 5.95 (m, 2 H), 6.59 - 6.66 (m, 1 H), 6.69 (d, J=1.4 Hz, 1 H), 6.72 - 6.78 (m, 1 H), 7.33 (s, 1 H)
205		356.20	357	0.66, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=7.28 Hz, 3 H) 1.15 - 1.30 (m, 2 H) 1.43 - 1.57 (m, 1 H) 1.57 - 1.69 (m, 1 H) 1.69 - 1.87 (m, 2 H) 3.37 - 3.45 (m, 2 H) 4.24 - 4.43 (m, 1 H) 5.30 (s, 2 H) 7.28 (t, J=6.53 Hz, 1 H) 7.55 (br. s., 2 H) 7.70 (s, 1 H) 7.62 - 7.77 (m, 1 H) 7.81 (d, J=8.78 Hz, 1 H) 8.31 (s, 1 H) 8.27 - 8.35 (m, 1 H) 8.81 (d, J=6.78 Hz, 1 H) 12.15 (br. s., 1 H)
206		266.17	267	0.72, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.40 Hz, 3 H) 1.15 - 1.33 (m, 2 H) 1.35 (s, 3 H) 1.43 - 1.56 (m, 2 H) 3.12 - 3.30 (m, 2 H) 3.91 (s, 2 H) 4.28 (d, J=5.77 Hz, 2 H) 4.46 (d, J=5.77 Hz, 2 H) 5.50 (s, 2 H) 6.20 (t, J=5.90 Hz, 1 H) 7.41

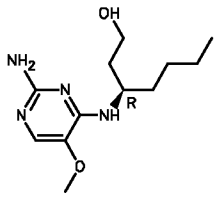
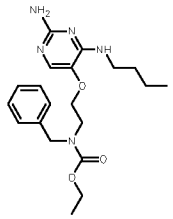
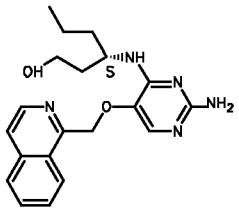
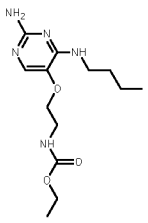
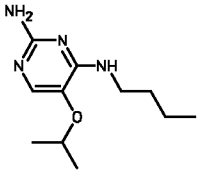
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					(s, 1 H)
207		370.21	371	0.71, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.83 (t, J=1.00 Hz, 3 H) 1.13 - 1.33 (m, 4 H) 1.47 - 1.61 (m, 1 H) 1.61 - 1.78 (m, 2 H) 1.79 - 1.89 (m, 1 H) 3.29 - 3.47 (m, 2 H) 4.27 - 4.38 (m, 1 H) 5.37 (s, 2 H) 7.47 (br. s., 1 H) 7.57 (br. s., 2 H) 7.73 (br. s., 1 H) 7.86 - 8.01 (m, 2 H) 8.35 (d, J=9.03 Hz, 1 H) 8.42 (s, 1 H) 8.94 (d, J=6.27 Hz, 1 H) 12.19 (br. s., 1 H)
208		381.22	382	0.86, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=7.03 Hz, 3 H) 1.13 - 1.36 (m, 4 H) 1.52 - 1.67 (m, 1 H) 1.71 - 1.84 (m, 2 H) 1.88 - 2.00 (m, 1 H) 3.33 - 3.48 (m, 2 H) 4.42 (m, J=8.80, 4.60, 4.60 Hz, 1 H) 6.02 (s, 2 H) 7.51 (br. s., 2 H) 7.96 (t, J=1.00 Hz, 1 H) 7.96 (t, J=1.00 Hz, 1 H) 8.13 (t, J=7.65 Hz, 1 H) 8.21 - 8.47 (m, 1 H) 8.32 (d, J=1.00 Hz, 1 H) 8.65 (s, 1 H) 8.64 (d, J=1.00 Hz, 1 H) 9.17 (br. s., 1 H) 12.34 (br. s., 1 H)"
209		312.17	313	0.69, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.4 Hz, 3 H), 1.29 (dq, J=14.9, 7.3 Hz, 2 H), 1.57 (quin, J=7.3 Hz, 2 H), 3.43 (dd, J=13.6, 6.8 Hz, 2 H), 5.38 6 (s, 2 H), 7.48 (td, J=6.7, 1.3 Hz, 1 H), 7.62 (br. s., 2 H), 7.72 (s, 1 H), 7.87 - 8.02 (m, 2 H), 8.46 (s, 1 H), 8.82 (t, J=5.9 Hz, 1 H), 8.94 (d, J=6.8 Hz, 1 H), 12.29 (br. s., 1 H)
210		297.16	298	0.85, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.22 - 1.35 (m, 2 H), 1.49 - 1.60 (m, 2 H), 3.37 - 3.47 (m, 2 H), 5.18 (s, 2 H), 7.49 - 7.62 (m, 3 6 H), 7.71 (m, J=8.5 Hz, 2 H), 7.86 - 7.93 (m, 2 H), 8.51 (t, J=5.9 Hz, 1 H), 12.17 - 12.31 (m, 1 H)

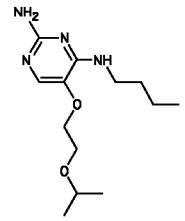
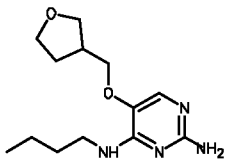
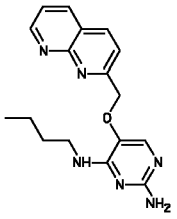
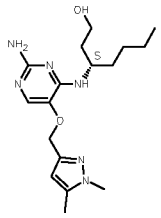
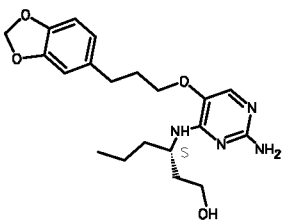
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
211		313.17	314	0.59, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.4 Hz, 3 H), 1.21 - 1.37 (m, 2 H), 1.48 - 1.62 (m, 2 H), 3.41 (q, J=6.8 Hz, 2 H), 5.33 (s, 2 H), 7.60 (br. s., 2 6 H), 7.69 (br. s., 1 H), 8.09 (d, J=4.5 Hz, 1 H), 8.50 - 8.67 (m, 2 H), 8.85 (d, J=4.3 Hz, 1 H), 9.32 (s, 1 H), 12.29 (br. s., 1 H)
212		337.19	338	0.94, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.3 Hz, 3 H), 1.17 (d, J=6.5 Hz, 3 H), 1.19 - 1.29 (m, 2 H), 1.40 - 1.55 (m, 1 H), 1.57 - 1.72 (m, 1 H), 4.21 - 6 4.35 (m, 1 H), 5.81 (s, 2 H), 7.47 (br. s., 2 H), 7.66 (br. s., 1 H), 7.78 - 7.86 (m, 1 H), 7.95 (t, J=7.3 Hz, 1 H), 8.08 (br. s., 1 H), 8.15 (d, J=8.0 Hz, 1 H), 8.48 (d, J=8.3 Hz, 1 H), 8.56 (d, J=5.8 Hz, 1 H), 11.73 (br. s., 1 H)
213		291.15	292	0.75, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.4 Hz, 3 H), 1.27 (dq, J=14.9, 7.4 Hz, 2 H), 1.51 (quin, J=7.3 Hz, 2 H), 3.38 (q, J=6.9 Hz, 2 H), 5.20 (d, 6 J=1.8 Hz, 2 H), 7.51 (br. s., 2 H), 7.54 - 7.62 (m, 2 H), 7.84 (ddd, J=9.9, 8.6, 1.1 Hz, 1 H), 8.39 - 8.53 (m, 2 H), 11.85 (d, J=5.5 Hz, 1 H)
214		363.19	364	0.65, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.18 - 1.35 (m, 2 H), 1.36 - 1.48 (m, 1 H), 1.51 - 1.64 (m, 1 H), 3.31 - 3.49 (m, 2 H), 3.78 (s, 3 6 H), 3.90 (s, 3 H), 3.99 - 4.09 (m, 1 H), 4.68 (br. s., 1 H), 4.86 - 4.97 (m, 2 H), 5.59 (s, 2 H), 6.38 (d, J=8.8 Hz, 1 H), 7.14 (d, J=5.5 Hz, 1 H), 7.49 (s, 1 H), 8.23 (d, J=5.5 Hz, 1 H)
215		342.18	343	0.6, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.3 Hz, 3 H), 1.14 - 1.33 (m, 2 H), 1.49 - 1.72 (m, 2 H), 3.47 - 3.61 (m, 2 H), 4.21 - 4.33 (m, 1 H), 5.41 (s, 2 6 H), 7.50 (td, J=6.5, 1.5 Hz, 1 H), 7.61 (br. s., 2 H), 7.78 (s, 1 H), 7.91 - 8.03 (m, 2 H), 8.22 (d, J=9.0 Hz, 1 H), 8.48 (s, 1 H), 8.97 (d, J=6.8 Hz, 1 H), 12.42 (br. s., 1 H)

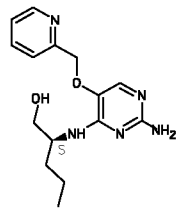
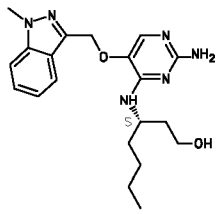
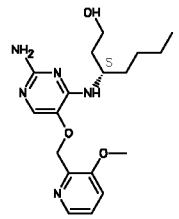
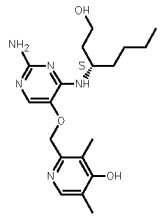
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
216		367.20	368	0.82, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.80 (t, J=7.2 Hz, 3 H), 1.08 - 1.18 (m, 2 H), 1.18 - 1.27 (m, 2 H), 1.27 - 1.37 (m, 1 H), 1.49 - 1.61 (m, 1 H), 3.27 - 6 3.33 (m, 2 H), 3.92 - 4.04 (m, 1 H), 4.65 (br. s., 1 H), 5.47 - 5.63 (m, 4 H), 6.08 (d, J=9.0 Hz, 1 H), 7.51 (s, 1 H), 7.66 - 7.74 (m, 1 H), 7.78 - 7.83 (m, 1 H), 7.85 (d, J=5.5 Hz, 1 H), 8.02 (d, J=8.0 Hz, 1 H), 8.40 (d, J=8.5 Hz, 1 H), 8.48 (d, J=5.8 Hz, 1 H)
217		377.21	378	0.73, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=6.9 Hz, 3 H), 1.15 - 1.37 (m, 4 H), 1.58 - 1.79 (m, 2 H), 3.50 - 3.64 (m, 2 H), 3.93 (s, 3 H), 4.16 (s, 3 H), 6 4.25 - 4.37 (m, 1 H), 5.37 - 5.47 (m, 2 H), 7.58 (br. s., 2 H), 7.71 (d, J=6.8 Hz, 1 H), 7.81 (d, J=4.3 Hz, 1 H), 8.62 (d, J=6.8 Hz, 1 H), 8.89 (d, J=8.8 Hz, 1 H), 12.30 - 12.47 (m, 1 H)
218		356.20	357	0.68, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.82 (t, J=7.0 Hz, 3 H), 1.09 - 1.36 (m, 4 H), 1.61 (q, J=7.2 Hz, 2 H), 3.45 - 3.59 (m, 2 H), 4.18 - 4.31 (m, 1 H), 5.33 - 6 5.45 (m, 2 H), 7.47 (t, J=6.7 Hz, 1 H), 7.59 (br. s., 2 H), 7.76 (s, 1 H), 7.86 - 8.02 (m, 2 H), 8.20 (d, J=9.0 Hz, 1 H), 8.45 (s, 1 H), 8.94 (d, J=6.8 Hz, 1 H), 12.33 (br. s., 1 H)
219		337.19	338	0.94, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.3 Hz, 3 H), 1.18 (d, J=6.5 Hz, 3 H), 1.20 - 1.29 (m, 2 H), 1.41 - 1.56 (m, 1 H), 1.67 (dd, J=13.4, 6.7 Hz, 1H), 4.24 - 4.36 (m, 1 H), 5.84 (br. s., 2 H), 7.47 (br. s., 2 H), 7.70 (br. s., 1 H), 7.80 - 7.89 (m, 1 H), 7.98 (t, J=7.2 Hz, 1 H), 8.11 (br. s., 1 H), 8.17 (d, J=8.3 Hz, 1 H), 8.50 (d, J=8.3 Hz, 1 H), 8.57 (d, J=6.0 Hz, 1 H), 8.71 (br. s., 1 H), 11.79 (br. s., 1 H)

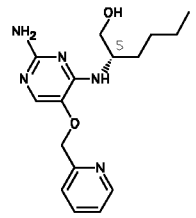
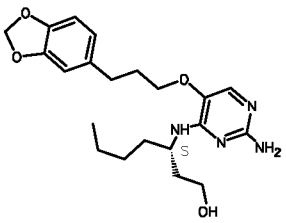
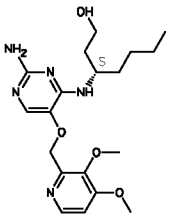
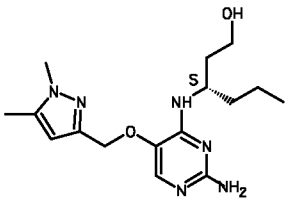
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
220		282.17	283	0.76, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.30 (dq, J=14.9, 7.3 Hz, 2 H), 1.54 (dt, J=14.5, 7.4 Hz, 2 H), 1.97 (quin, J=6.7 Hz, 2 H), 2.55 6 (t, J=7.4 Hz, 2 H), 3.37 - 3.45 (m, 2 H), 3.61 (s, 3 H), 3.93 (t, J=6.1 Hz, 2 H), 7.39 (s, 1 H), 7.47 (br. s., 2 H), 8.34 (t, J=5.8 Hz, 1 H), 11.96 (br. s., 1 H)
221		310.20	311	0.91, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 1.19 (d, J=6.3 Hz, 6 H), 1.30 (dq, J=14.9, 7.4 Hz, 2 H), 1.54 (dt, J=14.5, 7.4 Hz, 2 H), 1.89 - 6 2.02 (m, 2 H), 3.36 - 3.44 (m, 2 H), 3.92 (t, J=6.1 Hz, 2 H), 4.90 (quin, J=6.3 Hz, 1 H), 7.36 (s, 1 H), 7.41 (br. s., 2 H), 8.35 (t, J=6.0 Hz, 1 H), 11.73 (br. s., 1 H)
222		329.13	330	0.27, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.28 Hz, 2 H) 1.22 - 1.40 (m, 2 H) 1.42 - 1.58 (m, 2 H) 3.25 - 3.38 (m, 2 H) 5.39 (s, 2 H) 5.63 (s, 1 H) 6.56 (t, J=5.77 Hz, 1 H) 7.43 - 7.61 (m, 2 H) 8.01 (d, J=7.53 Hz, 1 H) 8.13 (dd, J=7.91, 0.63 Hz, 1 H)
223		381.22	382	0.86, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.83 (t, J=7.00 Hz, 3 H) 1.17 - 1.34 (m, 4 H) 1.53 - 1.67 (m, 2 H) 1.71 - 1.83 (m, 2 H) 3.46 (t, J=6.30 Hz, 2 H) 4.34 (m, J=7.80 Hz, 1 H) 5.33 (s, 2 H) 7.49 (br. s., 2 H) 7.64 (d, J=5.52 Hz, 1 H) 7.79 (t, J=7.50 Hz, 1 H) 7.91 (t, J=7.53 Hz, 1 H) 8.10 (s, 1 H) 8.06 (d, J=8.30 Hz, 1 H) 8.26 (d, J=8.28 Hz, 1 H) 8.46 (d, J=8.78 Hz, 1 H) 9.48 (s, 1 H)
224		312.17	313	0.26, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.88 (t, J=7.40 Hz, 3 H) 1.19 - 1.31 (m, 2 H) 1.51 (quin, J=7.28 Hz, 2 H) 3.39 (m, J=6.80, 6.80, 6.80 Hz, 2 H) 5.24 (s, 2 H) 6.78 (s, 1 H) 6.92 (t, J=6.90 Hz, 1 H) 7.25 (dd, J=8.28, 7.28 Hz, 1 H) 7.47 (br. s., 2 H) 7.55 (d, J=5.77 Hz, 1 H) 7.70 (d, J=9.03 Hz, 1 H) 8.42 (t, J=5.77 Hz, 1 H) 8.65 (d, J=7.03 Hz, 1 H) 11.74 (d, J=5.77 Hz, 1 H)

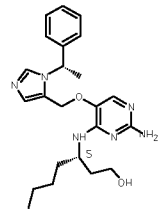
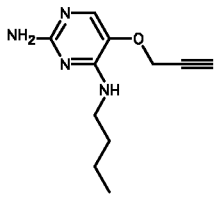
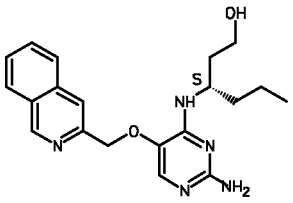
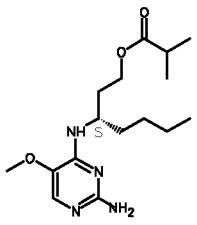
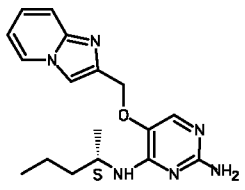
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
225		388.25	389	2.51, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.74 - 0.88 (m, 3 H), 1.14 - 1.36 (m, 4 H), 1.40 - 1.65 (m, 2 H), 1.77 - 1.93 (m, 2 H), 2.00 (quin, J=6.9 Hz, 2 H), 2.64 (td, J=7.4, 2.4 Hz, 2 H), 3.38 - 3.42 (m, 1 H), 3.46 (dd, J=11.4, 2.6 Hz, 1 H), 3.52 (dd, J=5.1, 2.2 Hz, 1 H), 3.72 (s, 3 H), 3.84 (td, J=6.3, 1.8 Hz, 2 H), 4.06 (d, J=2.7 Hz, 1 H), 4.48 (br. s., 2 H), 4.89 (d, J=8.7 Hz, 1 H), 6.72 - 6.80 (m, 2 H), 7.02 (d, J=8.7 Hz, 2 H), 7.25 (s, 1 H)
226		374.23	375	2.36, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.85 (t, J=7.2 Hz, 3 H), 1.24 - 1.39 (m, 3 H), 1.41 - 1.54 (m, 2 H), 1.85 (d, J=5.4 Hz, 1 H), 2.00 (t, J=6.9 Hz, 2 H), 2.64 (td, J=7.4, 2.1 Hz, 2 H), 3.42 (s, 1 H), 3.46 (dd, J=11.4, 2.6 Hz, 1 H), 3.52 (d, J=2.6 Hz, 1 H), 3.72 (s, 3 H), 3.84 (td, J=6.4, 1.4 Hz, 2 H), 4.01 - 4.17 (m, 1 H), 4.46 (br. s., 2 H), 4.85 (br. s., 1 H), 6.71 - 6.82 (m, 2 H), 6.97 - 7.08 (m, 2 H), 7.26 (s, 1 H)
227		331.20	332	1.5, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 1.22 - 1.50 (m, 4 H), 1.52 - 1.67 (m, 2 H), 1.83 - 2.05 (m, 1 H), 2.58 (s, 3 H), 3.43 - 3.55 (m, 1 H), 3.56 - 3.65 (m, 1 H), 4.20 (br. s., 1 H), 4.60 - 4.76 (m, 2 H), 4.99 (s, 2 H), 5.82 (d, J=8.7 Hz, 1 H), 7.14 (t, J=6.8 Hz, 2 H), 7.50 (s, 1 H), 7.61 (t, J=7.7 Hz, 1 H)
228		388.21	389	2.4, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.92 (t, J=1.0 Hz, 3 H), 1.22 - 1.43 (m, 5 H), 1.47 - 1.71 (m, 2 H), 1.97 - 2.12 (m, 2 H), 2.64 - 2.75 (m, 2 H), 3.63 (dd, J=10.9, 6.8 Hz, 1 H), 3.74 - 3.83 (m, 1 H), 3.91 (t, J=6.3 Hz, 2 H), 3.97 - 4.10 (m, 1 H), 4.57 (br. s., 2 H), 5.26 (d, J=6.7 Hz, 1 H), 5.94 (s, 2 H), 6.59 - 6.66 (m, 1 H), 6.69 (d, J=1.5 Hz, 1 H), 6.72 - 6.78 (m, 1 H), 7.35 (s, 1 H)

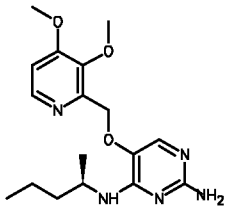
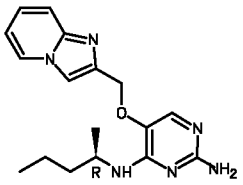
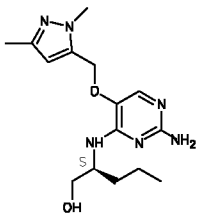
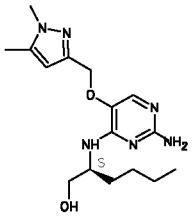
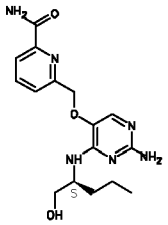
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
229		254.17	255	1.59, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.90 (t, J=1.0 Hz, 3 H), 1.30 - 1.46 (m, 5 H), 1.48 - 1.73 (m, 3 H), 1.95 (tdd, J=11.2, 11.2, 5.5, 2.7 Hz, 1 H), 3.54 (dd, J=11.3, 2.7 Hz, 1 H), 3.58 - 3.67 (m, 1 H), 3.79 (s, 3 H), 4.16 (dd, J=5.7, 3.0 Hz, 1 H), 4.99 (br. s., 2 H), 5.10 (d, J=8.5 Hz, 1 H), 7.32 (s, 1 H)
230		387.23	388	5.75, G	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.32 Hz, 3 H) 1.03 - 1.20 (m, 3 H) 1.29 (m, J=7.70 Hz, 2 H) 1.52 (d, J=6.95 Hz, 2 H) 3.38 (m, J=7.00 Hz, 2 H) 3.48 - 3.63 (m, 2 H) 4.07 (q, J=7.20 Hz, 4 H) 4.53 (s, 2 H) 7.19 - 7.29 (m, 3 H) 7.30 - 7.38 (m, 2 H) 7.42 (s, 1 H) 7.45 - 7.56 (m, 2 H) 8.09 - 8.32 (m, 1 H) 11.84 - 12.01 (m, 1 H)
231		367.20	368	0.8, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.40 Hz, 3 H) 1.19 - 1.30 (m, 2 H) 1.48 - 1.58 (m, 1 H) 1.65 - 1.78 (m, 2 H) 1.82 - 1.92 (m, 1 H) 3.35 - 3.45 (m, 2 H) 4.37 - 4.45 (m, 1 H) 5.93 (s, 2 H) 7.49 (br. s., 2 H) 7.80 (br. s., 1 H) 7.90 (t, J=7.40 Hz, 1 H) 8.04 (t, J=6.90 Hz, 1 H) 8.22 (d, J=8.03 Hz, 2 H) 8.54 - 8.63 (m, 2 H) 8.88 (br. s., 1 H) 12.04 (br. s., 1 H)
232		297.18	298	4.18, G	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.91 (t, J=7.32 Hz, 3 H) 1.17 (t, J=7.14 Hz, 3 H) 1.32 (m, J=7.40, 7.40, 7.40, 7.40, 7.40 Hz, 2 H) 1.56 (m, J=7.30, 7.30, 7.30, 7.30 Hz, 2 H) 3.38 - 3.48 (m, 2 H) 3.88 (t, J=5.12 Hz, 2 H) 4.01 (q, J=7.20 Hz, 2 H) 7.32 - 7.40 (m, 2 H) 7.44 (br. s., 2 H) 8.32 (t, J=5.67 Hz, 1 H) 11.71 (br. s., 1 H)
233		224.16	225	4.53, G	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.32 Hz, 3 H) 1.19 - 1.34 (m, 8 H) 1.45 - 1.58 (m, 2 H) 3.35 - 3.43 (m, 2 H) 4.41 (m, J=6.00, 6.00, 6.00, 6.00 Hz, 1 H) 7.35 - 7.54 (m, 3 H) 8.26 (t, J=6.04 Hz, 1 H) 11.89 (br. s., 1 H)

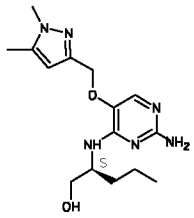
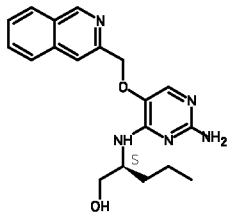
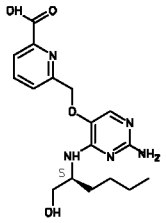
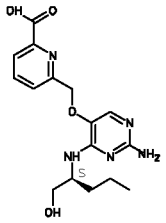
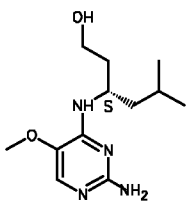
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
234		268.19	269	0.84, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.4 Hz, 3 H), 1.10 (d, J=6.0 Hz, 6 H), 1.24 - 1.35 (m, 2 H), 1.54 (quin, J=7.3 Hz, 2 H), 3.40 (q, J=6.9 Hz, 2 6 H), 3.62 (dt, J=12.2, 6.1 Hz, 1 H), 3.68 (dd, J=5.3, 4.0 Hz, 2 H), 4.01 - 4.07 (m, 2 H), 7.36 - 7.52 (m, 3 H), 8.27 (t, J=5.9 Hz, 1 H), 11.77 (br. s., 1 H)
235		266.17	267	0.71, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.3 Hz, 3 H), 1.30 (dq, J=14.9, 7.3 Hz, 2 H), 1.49 - 1.57 (m, 2 H), 1.57 - 1.67 (m, 1 H), 1.97 - 2.09 (m, 1 H), 6 2.59 - 2.71 (m, 1 H), 3.40 (q, J=6.8 Hz, 2 H), 3.52 (dd, J=8.7, 5.4 Hz, 1 H), 3.65 (q, J=7.7 Hz, 1 H), 3.72 - 3.85 (m, 3 H), 3.86 - 3.93 (m, 1 H), 7.32 - 7.48 (m, 3 H), 8.30 (t, J=5.9 Hz, 1 H), 11.88 (br. s., 1 H)
236		324.17	325	0.71, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.91 (t, J=7.4 Hz, 3 H), 1.32 (sxt, J=7.4 Hz, 2 H), 1.52 - 1.65 (m, 2 H), 3.46 (q, J=6.8 Hz, 2 H), 5.43 (s, 2 H), 7.59 (br. 6 s., 2 H), 7.65 (d, J=4.5 Hz, 1 H), 7.81 (dd, J=8.2, 4.4 Hz, 1 H), 8.05 (d, J=8.5 Hz, 1 H), 8.64 - 8.77 (m, 3 H), 9.20 (dd, J=4.4, 1.9 Hz, 1 H), 12.13 (br. s., 1 H)
237		348.23	349	0.73, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (m, J=7.20, 7.20 Hz, 3 H) 1.14 - 1.34 (m, 4 H) 1.55 (m, J=16.10, 8.00, 8.00 Hz, 2 H) 1.62 - 1.78 (m, 2 H) 2.23 (s, 2 H) 3.39 (m, J=6.40, 6.40 Hz, 2 H) 3.69 (s, 3 H) 4.23 - 4.33 (m, 1 H) 4.93 (s, 2 H) 6.15 (s, 1 H) 7.46 (br. s., 1 H) 7.52 (s, 1 H) 8.04 (d, J=9.03 Hz, 1 H) 11.92 (d, J=5.27 Hz, 1 H)
238		388.21	389	2.34, F	^{1}H NMR (300 MHz, CHLOROFORM-d) δ ppm 0.93 (t, J=7.2 Hz, 3 H), 1.29 - 1.47 (m, 3 H), 1.49 - 1.64 (m, 3 H), 1.87 - 2.00 (m, 1 H), 2.07 (quin, J=6.9 Hz, 2 H), 2.66 - 2.73 (m, 2 H), 3.46 - 3.57 (m, 1 H), 3.58 - 3.68 (m, 1 H), 3.91 (td, J=6.4, 1.4 Hz, 2 H), 4.16 (ddd, J=11.2, 5.4, 3.0 Hz, 1 H), 4.52 (s, 2 H), 4.93 (d, J=8.7

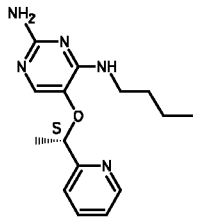
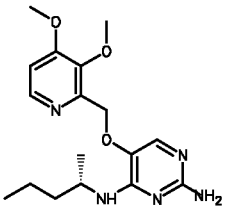
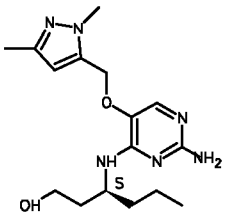
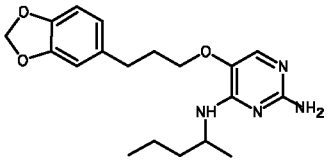
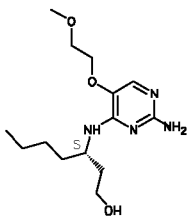
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					Hz, 1 H), 5.94 (s, 2 H), 6.60 - 6.65 (m, 1 H), 6.69 (d, J=1.5 Hz, 1 H), 6.72 - 6.77 (m, 1 H), 7.34 (s, 1 H)
239		303.17	304	1.42, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.88 - 1.01 (m, 3 H), 1.22 - 1.51 (m, 3 H), 1.54 - 1.71 (m, 2 H), 3.62 (dd, J=11.0, 6.7 Hz, 1 H), 3.78 (dd, J=11.0, 3.2 Hz, 1 H), 4.11 (td, J=6.8, 3.0 Hz, 1 H), 4.56 (br. s., 2 H), 4.92 - 5.13 (m, 2 H), 6.21 (d, J=7.0 Hz, 1 H), 7.30 (m, J=5.4 Hz, 1 H), 7.36 (d, J=7.7 Hz, 1 H), 7.52 (s, 1 H), 7.74 (td, J=7.7, 1.6 Hz, 1 H), 8.61 (d, J=4.7 Hz, 1 H)
240		384.23	385	0.88, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.73 - 0.87 (m, 3 H) 1.08 - 1.19 (m, 2 H) 1.19 - 1.31 (m, 2 H) 1.43 - 1.59 (m, 2 H) 1.59 - 1.75 (m, 2 H) 3.35 - 3.42 (m, 2 H) 4.03 (s, 3 H) 4.20 - 4.33 (m, 1 H) 5.44 (s, 2 H) 7.16 (t, J=7.40 Hz, 1 H) 7.43 (br. s, 1 H) 7.43 (t, J=7.70 Hz, 1 H) 7.51 (s, 1 H) 7.65 (d, J=8.53 Hz, 1 H) 7.88 (d, J=8.03 Hz, 1 H) 8.08 (d, J=8.78 Hz, 1 H) 11.70 (s, 1 H)
241		361.21	362	0.88, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (m, J=7.00, 7.00 Hz, 3 H) 1.14 - 1.35 (m, 4 H) 1.53 - 1.66 (m, 2 H) 1.68 - 1.83 (m, 2 H) 3.40 (m, J=6.70, 6.70 Hz, 2 H) 3.91 (s, 3 H) 4.28 - 4.41 (m, 1 H) 5.22 (s, 2 H) 7.49 (br. s., 2 H) 7.61 (d, J=1.00 Hz, 1 H) 7.61 (s, 1 H) 7.77 (d, J=7.78 Hz, 1 H) 8.26 (d, J=4.52 Hz, 1 H) 8.53 (d, J=8.03 Hz, 1 H) 11.84 (d, J=5.50 Hz, 1 H)
242		375.23	376	, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.79 - 0.87 (m, 3 H) 1.16 - 1.34 (m, 4 H) 1.54 - 1.63 (m, 1 H) 1.68 - 1.79 (m, 2 H) 1.85 - 1.95 (m, 1 H) 2.17 (s, 3 H) 2.24 (s, 3 H) 3.38 - 3.46 (m, 2 H) 4.33 - 4.43 (m, 1 H) 5.30 (s, 2 H) 7.48 (br. s., 2 H) 7.74 (d, J=4.77 Hz, 1 H) 8.29 (s, 1 H) 8.87 (d, J=8.53 Hz, 1 H) 11.99 (br. s., 1 H)"

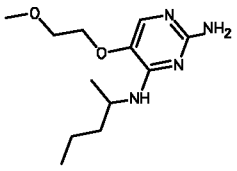
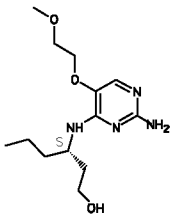
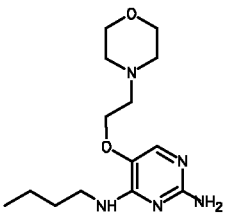
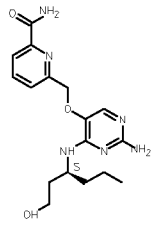
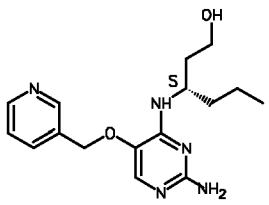
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
243		317.19	318	1.66, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.90 (t, J=1.0 Hz, 3 H), 1.30 - 1.46 (m, 5 H), 1.51 - 1.75 (m, 2 H), 3.57 - 3.68 (m, 1 H), 3.75 - 3.84 (m, 1 H), 4.09 (td, J=6.9, 2.9 Hz, 1 H), 4.63 (br. s., 2 H), 4.94 - 5.12 (m, 2 H), 6.25 (d, J=7.0 Hz, 1 H), 7.28 - 7.32 (m, 1 H), 7.37 (d, J=7.7 Hz, 1 H), 7.52 (s, 1 H), 7.74 (td, J=7.7, 1.8 Hz, 1 H), 8.62 (d, J=4.1 Hz, 1 H)
244		402.23	403	2.46, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.85 - 0.94 (m, 3 H), 1.23 - 1.44 (m, 5 H), 1.46 - 1.71 (m, 2 H), 1.94 (m, J=14.0, 11.3, 5.3, 3.0 Hz, 2 H), 2.06 (quin, J=6.9 Hz, 2 H), 2.70 (td, J=7.4, 1.6 Hz, 2 H), 3.45 - 3.57 (m, 1 H), 3.58 - 3.68 (m, 1 H), 3.85 - 3.98 (m, 2 H), 4.13 (ddd, J=11.2, 5.4, 3.0 Hz, 1 H), 4.53 (s, 2 H), 4.94 (d, J=8.7 Hz, 1 H), 5.93 (s, 2 H), 6.60 - 6.65 (m, 1 H), 6.68 (d, J=1.5 Hz, 1 H), 6.71 - 6.77 (m, 1 H), 7.35 (s, 1 H)
245		391.22	392	0.77, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.78 - 0.87 (m, 3 H) 1.16 - 1.33 (m, 4 H) 1.52 - 1.62 (m, 1 H) 1.63 - 1.78 (m, 2 H) 1.81 - 1.91 (m, 1 H) 3.35 - 3.42 (m, 2 H) 3.89 (s, 3 H) 4.08 (s, 3 H) 4.32 - 4.41 (m, 1 H) 5.29 (s, 2 H) 7.52 (s, 1 H) 7.51 (s, 2 H) 7.68 (d, J=5.52 Hz, 1 H) 8.51 (d, J=6.02 Hz, 1 H) 8.74 (br. s., 1 H) 11.90 (s, 1 H)"
246		334.21	335	0.66, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.78 - 0.90 (m, 3 H) 1.15 - 1.29 (m, 2 H) 1.40 - 1.62 (m, 2 H) 1.63 - 1.78 (m, 2 H) 2.23 (s, 3 H) 3.32 - 3.43 (m, 2 H) 3.70 (s, 3 H) 4.25 - 4.33 (m, 2 H) 4.93 (s, 2 H) 6.15 (s, 1 H) 7.47 (br. s., 2 H) 7.52 (s, 1 H) 8.04 (d, J=8.78 Hz, 1 H) 11.93 (s, 1 H)

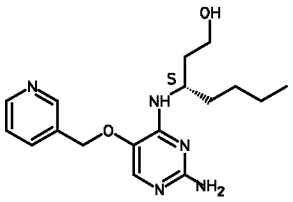
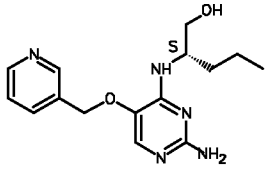
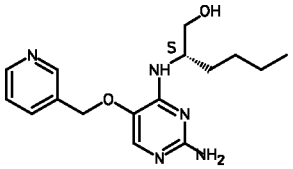
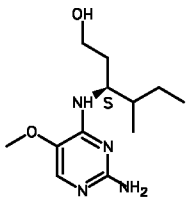
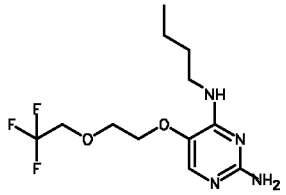
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
247		424.26	425	0.27, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm -0.05 - 0.01 (m, 2 H) 0.77 - 0.87 (m, 3 H) 1.12 - 1.35 (m, 4 H) 1.48 - 1.59 (m, 2 H) 1.66 - 1.79 (m, 2 H) 1.90 (d, J=7.03 Hz, 3 H) 3.41 - 3.47 (m, 2 H) 4.25 - 4.36 (m, 1 H) 4.85 (d, J=13.30 Hz, 1 H) 5.12 (d, J=13.05 Hz, 1 H) 5.81 (d, J=7.03 Hz, 1 H) 7.27 - 7.43 (m, 5 H) 7.45 - 7.61 (m, 2 H) 7.54 (br. s, 1 H) 7.95 - 8.05 (m, 1 H) 9.47 (s, 1 H) 12.16 (br. s., 1 H)
248		220.13	221	0.75, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 - 0.93 (m, 3 H) 1.22 - 1.35 (m, 2 H) 1.54 (m, J=1.00, 1.00, 1.00 Hz, 2 H) 3.33 - 3.43 (m, 2 H) 4.79 (d, J=2.51 Hz, 2 H) 7.50 (d, J=4.02 Hz, 1 H) 7.56 (br. s., 2 H) 8.51 (t, J=5.77 Hz, 1 H) 12.02 (br. s., 1 H)
249		367.20	368	0.84, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.40 Hz, 3 H) 1.21 - 1.31 (m, 2 H) 1.49 - 1.58 (m, 1 H) 1.58 - 1.69 (m, 1 H) 1.70 - 1.85 (m, 2 H) 3.38 - 3.50 (m, 2 H) 4.30 - 4.42 (m, 1 H) 5.35 (s, 2 H) 7.51 (br. s., 2 H) 7.65 (d, J=5.52 Hz, 1 H) 7.81 (t, J=7.53 Hz, 1 H) 7.93 (t, J=7.40 Hz, 1 H) 8.08 (d, J=8.03 Hz, 1 H) 8.13 (s, 1 H) 8.29 (d, J=8.28 Hz, 1 H) 8.46 (d, J=8.78 Hz, 1 H) 9.52 (s, 1 H) 11.84 (d, J=5.27 Hz, 1 H)
250		324.22	325	1.02, D	^{1}H NMR (600 MHz, DMSO-d_6) δ ppm 0.84 (t, J=7.0 Hz, 3 H), 1.05 (d, J=7.0 Hz, 3 H), 1.06 (d, J=7.0 Hz, 3 H), 1.16 - 1.32 (m, 4 H), 1.45 - 1.55 (m, 2 H), 6 1.80 (q, J=6.9 Hz, 2 H), 2.48 (spt, J=6.9 Hz, 1 H), 3.67 (s, 3 H), 3.95 - 4.03 (m, 2 H), 4.13 - 4.21 (m, 1 H), 5.37 (s, 2 H), 6.20 (d, J=9.1 Hz, 1 H), 7.35 (s, 1 H)
251		326.19	327	0.79, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.4 Hz, 3 H), 1.20 (d, J=6.8 Hz, 3 H), 1.21 - 1.28 (m, 2 H), 1.44 - 1.56 (m, 1 H), 1.71 (dd, J=13.4, 7.4 Hz, 1H), 4.21 - 4.36 (m, 1 H), 5.37 (d, J=1.8 Hz, 2 H), 7.47 (t, J=6.7 Hz, 1 H), 7.59 (br. s., 2 H), 7.73 (s, 1 H), 7.86 -

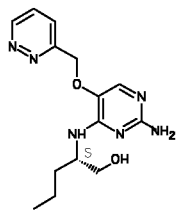
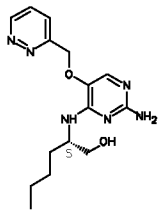
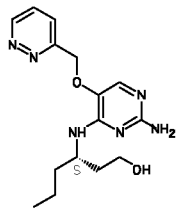
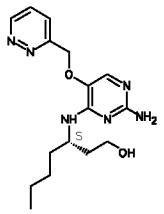
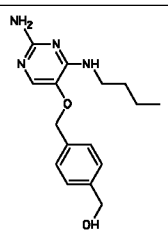
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					8.00 (m, 2 H), 8.36 - 8.46 (m, 2 H), 8.93 (d, J=6.5 Hz, 1 H), 12.24 (br. s., 1 H)
252		347.20	348	0.86, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.4 Hz, 3 H), 1.24 (d, J=6.5 Hz, 3 H), 1.25 - 1.33 (m, 2 H), 1.47 - 1.60 (m, 1 H), 1.71 - 1.86 (m, 1 H), 3.91 6 (s, 3 H), 4.12 (s, 3 H), 4.28 - 4.42 (m, 1 H), 5.36 (s, 2 H), 7.57 (br. s, 2 H), 7.63 (d, J=6.5 Hz, 1 H), 7.72 - 7.80 (m, 1 H), 8.56 (d, J=6.5 Hz, 1 H), 8.99 - 9.10 (m, 1 H), 12.27 (br. s., 1 H)
253		326.19	327	0.79, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.4 Hz, 3 H), 1.21 (d, J=6.5 Hz, 3 H), 1.23 - 1.29 (m, 2 H), 1.45 - 1.58 (m, 1 H), 1.66 - 1.80 (m, 1 H), 4.21 - 6 4.38 (m, 1 H), 5.33 - 5.45 (m, 2 H), 7.44 - 7.55 (m, 1 H), 7.62 (br. s., 2 H), 7.76 (s, 1 H), 7.89 - 8.02 (m, 2 H), 8.40 - 8.52 (m, 2 H), 8.96 (d, J=6.8 Hz, 1 H), 12.39 (br. s., 1 H)
254		320.20	321	0.62, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.17 - 1.35 (m, 2 H), 1.36 - 1.47 (m, 1 H), 1.47 - 1.60 (m, 1 H), 2.11 (s, 3 H), 3.36 - 3.47 (m, 2 6 H), 3.73 (s, 3 H), 4.05 (td, J=8.8, 4.9 Hz, 1 H), 4.66 (br. s., 1 H), 4.94 (s, 2 H), 5.58 (s, 2 H), 5.86 (d, J=9.0 Hz, 1 H), 6.10 (s, 1 H), 7.43 (s, 1 H)
255		334.21	335	0.72, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=7.0 Hz, 3 H), 1.11 - 1.38 (m, 4 H), 1.39 - 1.67 (m, 2 H), 2.23 (s, 3 H), 3.38 - 3.52 (m, 2 H), 3.70 (s, 3 H), 6 4.13 - 4.24 (m, 1 H), 4.93 (s, 2 H), 6.16 (s, 1 H), 7.47 (br. s., 2 H), 7.53 (d, J=5.3 Hz, 1 H), 7.79 (d, J=9.0 Hz, 1 H), 11.96 (d, J=5.3 Hz, 1 H)
256		346.18	347	0.55, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.17 - 1.33 (m, 2 H), 1.39 - 1.50 (m, 1 H), 1.50 - 1.62 (m, 1 H), 3.37 - 3.48 (m, 2 H), 4.01 - 6 4.14 (m, 1 H), 4.69 (br. s., 1 H), 5.10 (s, 2 H), 5.54 (s, 2 H), 6.00 (d, J=9.0 Hz, 1 H), 7.46 (s, 1 H), 7.68 (br. s., 1 H), 7.72 (dd, J=7.5, 1.3 Hz, 1 H), 7.94

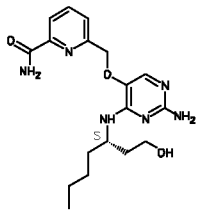
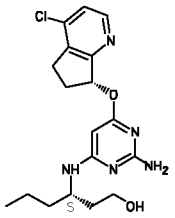
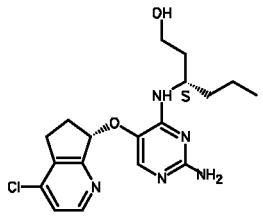
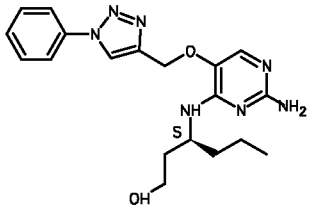
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					- 8.03 (m, 2 H), 8.03 (s, 1 H)
257		320.20	321	0.63, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.13 - 1.34 (m, 2 H), 1.46 - 1.60 (m, 2 H), 2.24 (s, 3 H), 3.38 - 3.53 (m, 2 H), 3.70 (s, 3 H), 4.18 - 4.28 (m, 1 H), 4.93 (s, 2 H), 6.16 (s, 1 H), 7.48 (br. s., 2 H), 7.54 (d, J=5.5 Hz, 1 H), 7.78 (d, J=8.8 Hz, 1 H), 11.97 (d, J=5.5 Hz, 1 H)
258		353.19	354	0.79, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.4 Hz, 3 H), 1.18 - 1.37 (m, 2 H), 1.58 (q, J=7.7 Hz, 2 H), 3.45 - 3.58 (m, 2 H), 4.21 - 4.32 (m, 1 H), 5.37 6 (s, 2 H), 7.54 (br. s., 2 H), 7.69 (d, J=5.0 Hz, 1 H), 7.84 (t, J=7.5 Hz, 1 H), 7.97 (t, J=7.5 Hz, 1 H), 8.11 (d, J=8.3 Hz, 1 H), 8.21 (s, 1 H), 8.32 (t, J=8.5 Hz, 2 H), 9.58 (s, 1 H), 11.98 (d, J=5.0 Hz, 1 H)
259		361.18	362	0.5, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=6.8 Hz, 3 H), 1.15 - 1.35 (m, 4 H), 1.38 - 1.57 (m, 1 H), 1.57 - 1.68 (m, 1 H), 3.38 - 3.50 (m, 2 H), 4.04 - 6 4.17 (m, 1 H), 5.12 (s, 2 H), 6.51 (br. s., 2 H), 6.71 (d, J=7.8 Hz, 1 H), 7.62 - 7.74 (m, 2 H), 7.90 - 7.98 (m, 2 H)
260		347.16	348	0.44, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.26 (dq, J=14.9, 7.3 Hz, 2 H), 1.43 - 1.63 (m, 2 H), 3.38 - 3.50 (m, 2 H), 4.13 (td, J=8.7, 5.1Hz, 1 H), 5.12 (s, 2 H), 6.50 (br. s., 2 H), 6.69 (d, J=8.5 Hz, 1 H), 7.63 - 7.70 (m, 2 H), 7.93 - 7.97 (m, 2 H)
261		254.17	255	3.71, G	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (dd, J=6.02, 4.52 Hz, 6 H) 1.14 - 1.29 (m, 1 H) 1.46 - 1.67 (m, 4 H) 3.34 - 3.43 (m, 2 H) 3.66 (s, 3 H) 4.22 (m, J=8.70, 8.70, 4.40 Hz, 1 H) 4.37 (t, J=5.40 Hz, 1 H) 5.43 (s, 2 H) 6.12 (d, J=9.03 Hz, 1 H) 7.34 (s, 1 H)

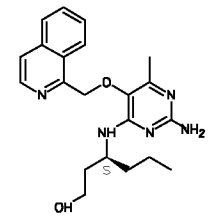
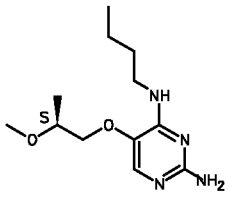
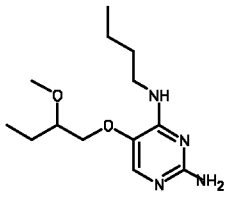
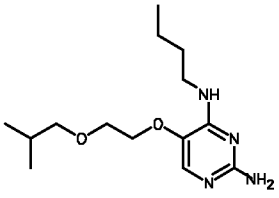
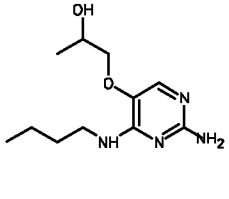
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
262		287.17	288	4.59, G	^{1}H NMR (360 MHz, DMSO-d_6) δ ppm 0.91 (t, J=7.50 Hz, 3 H) 1.31 (dq, J=14.96, 7.46 Hz, 2 H) 1.58 (quin, J=7.41 Hz, 2 H) 1.66 (d, J=6.22 Hz, 3 H) 3.43 (q, J=6.59 Hz, 2 H) 5.66 (q, J=6.10 Hz, 1 H) 7.57 (m, J=4.80 Hz, 3 H) 7.71 (t, J=5.90 Hz, 1 H) 7.87 (d, J=8.05 Hz, 1 H) 8.25 (t, J=7.50 Hz, 1 H) 8.75 (d, J=4.76 Hz, 1 H) 8.84 (t, J=5.85 Hz, 1 H) 12.10 (d, J=4.39 Hz, 1 H)
263		347.20	348	0.86, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.90 (t, J=7.2 Hz, 3 H), 1.17 (d, J=6.5 Hz, 3 H), 1.29 - 1.40 (m, 2 H), 1.40 - 1.59 (m, 2 H), 3.86 (s, 3 H), 3.93 (s, 3 H), 4.07 - 4.20 (m, 1 H), 4.71 (br. s., 2 H), 5.02 (s, 2 H), 6.28 (d, J=8.3 Hz, 1 H), 6.85 (d, J=5.5 Hz, 1 H), 7.59 (s, 1 H), 8.26 (d, J=5.5 Hz, 1 H)
264		334.21	335	0.65, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.16 - 1.30 (m, 2 H), 1.43 - 1.63 (m, 2 H), 1.63 - 1.80 (m, 2 H), 2.14 (s, 3 H), 3.40 (t, J=6.4 Hz, 6 2 H), 3.75 (s, 3 H), 4.26 - 4.39 (m, 1 H), 5.08 (s, 2 H), 6.23 (s, 1 H), 7.53 (br. s., 2 H), 7.59 (d, J=4.8 Hz, 1 H), 8.10 (d, J=8.8 Hz, 1 H), 12.22 (d, J=5.0 Hz, 1 H)
265		358.20	359	2.62, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.94 (t, J=1.0 Hz, 3 H), 1.19 (d, J=6.5 Hz, 3 H), 1.31 - 1.44 (m, 2 H), 1.45 - 1.58 (m, 2 H), 1.98 - 2.11 (m, 2 H), 2.70 (t, J=7.5 Hz, 2 H), 3.89 (t, J=6.3 Hz, 2 H), 4.15 (m, J=8.4, 6.6, 6.6, 6.6 Hz, 1 H), 4.44 (s, 2 H), 4.90 (d, J=8.4 Hz, 1 H), 5.94 (s, 2 H), 6.60 - 6.66 (m, 1 H), 6.69 (d, J=1.5 Hz, 1 H), 6.72 - 6.77 (m, 1 H), 7.32 (s, 1 H)
266		298.20	299	1.7, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.86 - 0.94 (m, 3 H), 1.25 - 1.45 (m, 5 H), 1.46 - 1.69 (m, 2 H), 1.86 - 2.00 (m, 1 H), 2.05 (s, 1 H), 3.43 (s, 3 H), 3.46 - 3.56 (m, 1 H), 3.57 - 3.63 (m, 1 H), 3.64 - 3.69 (m, 2

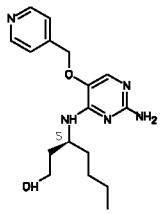
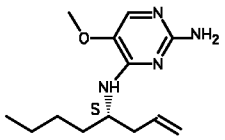
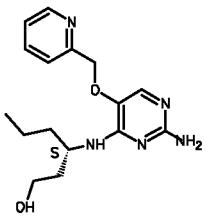
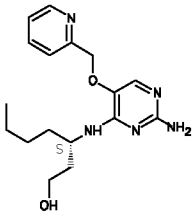
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					H), 3.96 - 4.04 (m, 2 H), 4.06 - 4.24 (m, 1 H), 5.18 (br. s., 2 H), 5.72 (d, J=8.8 Hz, 1 H), 7.45 (s, 1 H)
267		254.17	255	1.75, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.93 (t, J=1.0 Hz, 3 H), 1.18 (d, J=6.5 Hz, 3 H), 1.29 - 1.60 (m, 4 H), 3.44 (s, 3 H), 3.60 - 3.70 (m, 2 H), 3.95 - 4.02 (m, 2 H), 4.05 - 4.21 (m, 1 H), 4.53 (br. s., 2 H), 5.51 (d, J=7.8 Hz, 1 H), 7.46 (s, 1 H)
268		284.18	285	1.46, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.92 (t, J=7.2 Hz, 3 H), 1.30 - 1.49 (m, 4 H), 1.51 - 1.65 (m, 2 H), 1.85 - 1.98 (m, 1 H), 3.43 (s, 3 H), 3.52 (dd, J=11.4, 2.6 Hz, 1 H), 3.60 (td, J=5.9, 2.5 Hz, 1 H), 3.63 - 3.69 (m, 2 H), 3.95 - 4.03 (m, 2 H), 4.14 (ddd, J=8.3, 5.5, 2.7 Hz, 1 H), 4.85 (br. s., 2 H), 5.65 (d, J=8.7 Hz, 1 H), 7.48 (s, 1 H)
269		295.20	296	0.63, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.74 - 0.93 (m, 3 H) 1.30 (m, J=1.00, 1.00, 1.00 Hz, 2 H) 1.43 - 1.65 (m, 2 H) 3.09 - 3.18 (m, 2 H) 3.40 - 3.45 (m, 2 H) 3.49 - 3.60 (m, 2 H) 3.72 - 3.88 (m, 1 H) 3.88 - 4.13 (m, 5 H) 4.25 (t, J=4.77 Hz, 2 H) 7.45 (s, 1 H) 7.51 (br. s., 2 H) 9.31 (t, J=5.77 Hz, 1 H) 11.69 (br. s., 1 H) 12.01 (br. s., 1 H)
270		360.19	361	0.61, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.4 Hz, 3 H), 1.24 (dq, J=14.7, 7.4 Hz, 2 H), 1.39 - 1.56 (m, 2 H), 1.56 - 1.73 (m, 2 H), 3.41 (br. s., 2 H), 6 4.09 - 4.22 (m, 1 H), 4.44 (br. s., 1 H), 5.10 (s, 2 H), 5.54 (s, 2 H), 6.26 (d, J=9.0 Hz, 1 H), 7.45 (s, 1 H), 7.68 (br. s., 1 H), 7.74 (d, J=7.5 Hz, 1 H), 7.93 - 8.03 (m, 2 H), 8.06 (br. s., 1 H)
271		317.19	318	0.64, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.80 - 0.92 (m, 3 H) 1.17 - 1.36 (m, 4 H) 1.47 - 1.65 (m, 2 H) 1.67 - 1.81 (m, 2 H) 4.29 - 4.37 (m, 1 H) 5.26 (s, 2 H) 7.52 (br. s., 2 H) 7.62 (d, J=5.02 Hz, 1 H) 7.99 (dd, J=8.03, 5.52 Hz, 1 H) 8.19 (d,

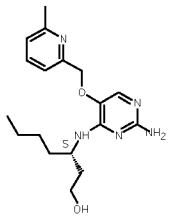
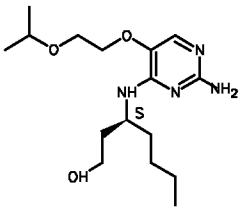
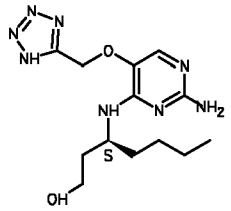
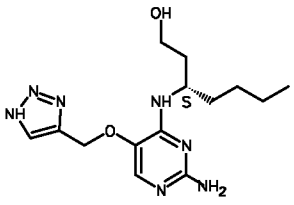
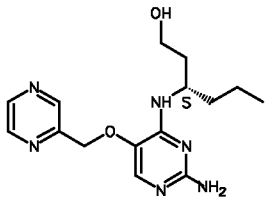
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					J=8.78 Hz, 1 H) 8.51 (d, J=8.03 Hz, 1 H) 8.87 (d, J=5.02 Hz, 1 H) 9.02 (s, 1 H) 11.98 (s, 1 H)
272		331.20	332	0.72, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.15 Hz, 3 H) 1.10 - 1.38 (m, 4 H) 1.56 (dd, J=14.56, 7.53 Hz, 2 H) 1.74 (dd, J=13.68, 5.90 Hz, 2 H) 4.25 - 4.39 (m, 2 H) 4.25 - 4.39 (m, 1 H) 5.19 (s, 2 H) 7.52 (br. s., 2 H) 7.61 (s, 1 H) 7.66 (dd, J=7.78, 5.02 Hz, 1 H) 8.16 (t, J=8.41 Hz, 2 H) 8.69 (d, J=4.27 Hz, 1 H) 8.83 (s, 1 H) 12.08 (br. s., 1 H)
273		303.17	304	0.59, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.81 - 0.91 (m, 3 H) 1.17 - 1.32 (m, 2 H) 1.47 - 1.60 (m, 2 H) 3.41 - 3.54 (m, 2 H) 4.20 - 4.34 (m, 1 H) 5.21 (s, 2 H) 7.50 (br. s., 2 H) 7.59 (d, J=4.77 Hz, 1 H) 7.78 (dd, J=7.65, 5.40 Hz, 1 H) 8.00 (d, J=9.03 Hz, 1 H) 8.29 (d, J=7.53 Hz, 1 H) 8.75 (d, J=4.27 Hz, 1 H) 8.92 (s, 1 H) 11.95 (br. s., 1 H)
274		317.19	318	0.57, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.03 Hz, 3 H) 1.13 - 1.36 (m, 4 H) 1.47 - 1.66 (m, 2 H) 3.40 - 3.52 (m, 2 H) 4.16 - 4.30 (m, 1 H) 5.24 (s, 2 H) 7.53 (br. s., 2 H) 7.62 (d, J=4.02 Hz, 1 H) 7.86 (dd, J=7.91, 5.40 Hz, 1 H) 8.02 (d, J=8.78 Hz, 1 H) 8.39 (d, J=8.03 Hz, 1 H) 8.80 (d, J=4.27 Hz, 1 H) 8.98 (s, 1 H) 12.08 (br. s., 1 H)
275		254.17	255	0.67, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.73 - 0.91 (m, 6 H) 0.94 - 1.16 (m, 1 H) 1.33 - 1.47 (m, 1 H) 1.49 - 1.75 (m, 3 H) 3.38 (m, J=9.00 Hz, 2 H) 3.67 (s, 3 H) 3.93 - 4.18 (m, 1 H) 4.34 (t, J=1.00 Hz, 1 H) 5.44 (br. s., 2 H) 5.94 (d, J=1.00 Hz, 1 H) 7.35 (s, 1 H)
276		308.15	309	4.89, G	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.28 Hz, 3 H) 1.30 (sxt, J=7.43 Hz, 2 H) 1.47 - 1.60 (m, 2 H) 3.41 (q, J=6.78 Hz, 2 H) 3.90 - 3.97 (m, 2 H) 4.09 - 4.13 (m, 2 H) 4.18 (q, J=1.00 Hz, 2 H) 7.46 (s, 1 H) 7.49 (br. s., 1 H) 8.32 (t, J=5.90 Hz, 1 H)

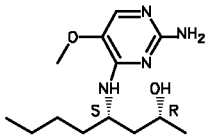
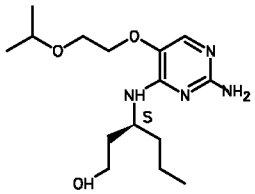
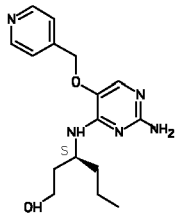
	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
277		304.16	305	0.5, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.4 Hz, 3 H), 1.17 - 1.35 (m, 2 H), 1.47 - 1.62 (m, 2 H), 3.43 - 3.54 (m, 2 H), 4.19 - 4.31 (m, 1 H), 5.39 (s, 2 6 H), 7.55 (br. s., 2 H), 7.65 (d, J=4.0 Hz, 1 H), 7.85 (dd, J=8.5, 5.0 Hz, 1 H), 8.00 (dd, J=8.4, 1.6 Hz, 1 H), 8.07 (d, J=8.8 Hz, 1 H), 9.27 (dd, J=4.9, 1.6 Hz, 1 H), 12.03 - 12.17 (m, 1 H)
278		318.18	319	0.58, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.0 Hz, 3 H), 1.14 - 1.38 (m, 4 H), 1.45 - 1.71 (m, 2 H), 3.42 - 3.53 (m, 2 H), 4.23 (td, J=9.0, 5.4 Hz, 1 H), 6 5.39 (s, 2 H), 7.55 (br. s., 2 H), 7.65 (d, J=3.3 Hz, 1 H), 7.84 (dd, J=8.4, 4.9 Hz, 1 H), 8.00 (dd, J=8.5, 1.5 Hz, 1 H), 8.07 (d, J=8.8 Hz, 1 H), 9.27 (dd, J=5.0, 1.8 Hz, 1 H), 12.10 (br. s., 1 H)
279		318.18	319	0.54, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.3 Hz, 3 H), 1.25 (dq, J=14.9, 7.4 Hz, 2 H), 1.45 - 1.66 (m, 2 H), 1.66 - 1.83 (m, 2 H), 3.43 (t, J=6.4 Hz, 2 6 H), 4.28 - 4.40 (m, 1 H), 5.39 (s, 2 H), 7.56 (br. s., 2 H), 7.66 (d, J=4.0 Hz, 1 H), 7.85 (dd, J=8.5, 5.0 Hz, 1 H), 7.98 (dd, J=8.5, 1.5 Hz, 1 H), 8.26 (d, J=9.0 Hz, 1 H), 9.27 (dd, J=4.9, 1.6 Hz, 1 H), 12.13 (br. s., 1 H)
280		332.20	333	0.62, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.0 Hz, 3 H), 1.15 - 1.35 (m, 4 H), 1.49 - 1.66 (m, 2 H), 1.68 - 1.80 (m, 2 H), 3.43 (t, J=6.4 Hz, 2 H), 4.26 - 6 4.39 (m, 1 H), 5.39 (s, 2 H), 7.55 (br. s., 2 H), 7.66 (d, J=4.3 Hz, 1 H), 7.85 (dd, J=8.5, 5.0 Hz, 1 H), 7.97 (dd, J=8.5, 1.5 Hz, 1 H), 8.26 (d, J=8.8 Hz, 1 H), 9.27 (dd, J=5.0, 1.5 Hz, 1 H), 12.05 - 12.16 (m, 1 H)
281		302.17	303	0.71, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.89 (t, J=7.3 Hz, 3 H), 1.28 (dq, J=14.9, 7.3 Hz, 2 H), 1.49 (quin, J=7.3 Hz, 2 H), 3.23 - 3.31 (m, 2 H), 4.49 (br. s., 2 6 H), 4.93 (s, 2 H), 5.17 (br. s., 1 H), 5.47 (s, 2 H), 6.37 (t, J=5.8 Hz, 1 H), 7.26 -

	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
					7.33 (m, 2 H), 7.33 - 7.42 (m, 3 H)
282		374.21	375	0.66, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=7.0 Hz, 3 H), 1.13 - 1.33 (m, 4 H), 1.42 - 1.56 (m, 2 H), 1.56 - 1.73 (m, 2 H), 3.40 (br. s., 2 H), 4.06 - 4.20 6 (m, 1 H), 4.44 (br. s., 1 H), 5.10 (s, 2 H), 5.55 (s, 2 H), 6.28 (d, J=8.8 Hz, 1 H), 7.45 (s, 1 H), 7.67 (br. s., 1 H), 7.71 - 7.76 (m, 1 H), 7.93 - 8.03 (m, 2 H), 8.06 (br. s., 1 H)
283		377.16	378	0.91, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.82 (t, J=7.40 Hz, 3 H) 1.11 - 1.22 (m, 2 H) 1.43 - 1.55 (m, 2 H) 1.66 - 1.76 (m, 2 H) 2.25 - 2.34 (m, 1 H) 2.52 - 2.65 (m, 1 H) 2.88 - 2.97 (m, 1 H) 3.10 - 3.22 (m, 1 H) 3.43 (t, J=6.40 Hz, 2 H) 4.24 - 4.34 (m, 1 H) 5.61 (dd, J=7.40, 4.14 Hz, 1 H) 7.51 (d, J=7.60 Hz, 1 H) 7.53 (br. s, 2 H) 7.84 (s, 1 H) 8.17 (d, J=8.78 Hz, 1 H) 8.44 (d, J=5.52 Hz, 1 H) 11.77 (br. s., 1 H)
284		377.16	378	0.92, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.82 (t, J=7.40 Hz, 3 H) 1.11 - 1.22 (m, 2 H) 1.43 - 1.55 (m, 2 H) 1.66 - 1.76 (m, 2 H) 2.25 - 2.34 (m, 1 H) 2.52 - 2.65 (m, 1 H) 2.88 - 2.97 (m, 1 H) 3.10 - 3.22 (m, 1 H) 3.43 (t, J=6.40 Hz, 2 H) 4.24 - 4.34 (m, 1 H) 5.61 (dd, J=7.40, 4.14 Hz, 1 H) 7.51 (d, J=7.60 Hz, 1 H) 7.53 (br. s, 2 H) 7.84 (s, 1 H) 8.17 (d, J=8.78 Hz, 1 H) 8.44 (d, J=5.52 Hz, 1 H) 11.77 (br. s., 1 H)
285		383.21	384	0.8, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.62 - 0.92 (m, 3 H) 1.14 - 1.31 (m, 2 H) 1.42 - 1.63 (m, 2 H) 1.63 - 1.82 (m, 2 H) 3.40 (t, J=6.40 Hz, 2 H) 4.25 - 4.36 (m, 1 H) 5.22 (s, 2 H) 7.47 - 7.59 (m, 1 H) 7.47 - 7.59 (m, 2 H) 7.59 - 7.67 (m, 2 H) 7.72 (br. s., 1 H) 7.85 - 7.98 (m, 2 H) 8.08 (d, J=8.78 Hz, 1 H) 9.07 (s, 1 H) 12.16 (br. s., 1 H)

	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
286		381.22	382	0.89, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=7.28 Hz, 3 H) 1.18 - 1.30 (m, 2 H) 1.50 - 1.64 (m, 2 H) 1.75 (dt, J=12.80, 6.40 Hz, 2 H) 2.14 (s, 3 H) 3.40 - 3.44 (m, 2 H) 4.31 (m, J=7.50 Hz, 1 H) 5.64 (s, 2 H) 7.46 (br. s., 2 H) 7.78 - 7.85 (m, 1 H) 7.91 (t, J=7.65 Hz, 1 H) 8.00 (d, J=6.02 Hz, 1 H) 8.11 (d, J=8.28 Hz, 1 H) 8.37 (d, J=8.28 Hz, 1 H) 8.56 (d, J=5.77 Hz, 1 H) 9.30 (br. s., 1 H) 12.20 (s, 1 H)
287		254.17	255	4.21, G	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (t, J=7.40 Hz, 3 H) 1.17 (d, J=6.52 Hz, 3 H) 1.23 - 1.38 (m, 2 H) 1.54 (quin, J=7.34 Hz, 2 H) 3.30 (s, 3 H) 3.41 (q, J=6.69 Hz, 2 H) 3.60 - 3.75 (m, 1 H) 3.78 - 3.98 (m, 2 H) 7.32 - 7.58 (m, 3 H) 8.24 (t, J=5.77 Hz, 1 H)
288		268.19	269	0.86, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.90 (tt, J=7.40, 3.50 Hz, 6 H) 1.23 - 1.36 (m, 2 H) 1.47 - 1.69 (m, 4 H) 3.33 (s, 3 H) 3.36 - 3.52 (m, 3 H) 3.92 (d, J=4.77 Hz, 2 H) 7.19 - 7.68 (m, 3 H) 8.21 (t, J=6.02 Hz, 1 H)
289		282.21	283	5.45, G	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (d, J=6.78 Hz, 6 H) 0.90 (t, J=7.40 Hz, 3 H) 1.22 - 1.37 (m, 2 H) 1.54 (quin, J=7.28 Hz, 2 H) 1.78 (m, J=13.40, 6.70, 6.70 Hz, 1 H) 3.21 (d, J=6.52 Hz, 2 H) 3.40 (q, J=6.69 Hz, 2 H) 3.56 - 3.75 (m, 2 H) 3.99 - 4.14 (m, 2 H) 7.32 - 7.60 (m, 3 H) 8.29 (t, J=5.65 Hz, 1 H)
290		240.16	241	3.38, G	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.91 (t, J=7.28 Hz, 3 H) 1.12 (d, J=6.52 Hz, 3 H) 1.31 (sxt, J=7.43 Hz, 2 H) 1.56 (quin, J=7.34 Hz, 2 H) 3.34 - 3.48 (m, 2 H) 3.61 (dd, J=9.41, 7.40 Hz, 1 H) 3.83 (dd, J=9.54, 3.51 Hz, 1 H) 3.90 - 4.02 (m, 1 H) 6.10 (br. s., 1 H) 7.40 (d, J=5.27 Hz, 1 H) 7.48 (br. s., 2 H) 8.54 (t, J=5.65 Hz, 1 H) 12.02 (br. s., 1 H)

	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
291		331.20	332	0.7, D	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.90 (t, J=6.9 Hz, 3 H), 1.27 - 1.45 (m, 5 H), 1.47 - 1.69 (m, 2 H), 1.87 - 1.99 (m, 1 H), 3.49 - 3.58 (m, 1 H), 3.60 - 3.66 (m, 1 H), 4.17 (ddd, J=10.8, 5.5, 3.0 Hz, 1 H), 5.00 (s, 2 H), 5.15 (d, J=8.5 Hz, 1 H), 7.25 - 7.32 (m, 2 H), 7.39 (s, 1 H), 8.57 - 8.67 (m, 2 H) supports structure but don't see exchangables.
292		250.18	251	4.51, B	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.82 - 0.93 (m, 3 H), 1.22 - 1.37 (m, 4 H), 1.40 - 1.51 (m, 1 H), 1.52 - 1.63 (m, 1 H), 2.20 - 2.39 (m, 2 H), 3.75 (s, 3 H), 4.09 - 4.23 (m, 1 H), 4.72 (br. s., 2 H), 5.04 (s, 1 H), 5.08 (d, J=4.8 Hz, 2 H), 5.70 - 5.87 (m, 1 H), 7.30 (s, 1 H)
293		317.19	318	1.55, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.92 (t, J=7.3 Hz, 3 H), 1.31 - 1.50 (m, 3 H), 1.55 - 1.67 (m, 2 H), 1.94 (m, J=11.2, 11.2, 5.5, 2.6 Hz, 2 H), 3.42 - 3.54 (m, 1 H), 3.56 - 3.69 (m, 1 H), 4.17 (d, J=7.3 Hz, 1 H), 4.53 (br. s., 2 H), 5.04 (s, 2 H), 6.05 (d, J=8.5 Hz, 1 H), 7.29 - 7.38 (m, 2 H), 7.54 (s, 1 H), 7.74 (td, J=7.7, 1.6 Hz, 1 H), 8.63 (d, J=4.7 Hz, 1 H)
294		331.20	332	1.76, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.80 (t, J=6.9 Hz, 3 H), 1.13 - 1.39 (m, 5 H), 1.44 - 1.60 (m, 3 H), 1.80 - 1.95 (m, 1 H), 3.35 - 3.47 (m, 1 H), 3.48 - 3.59 (m, 1 H), 4.08 (ddd, J=11.0, 5.5, 2.7 Hz, 1 H), 4.49 (s, 2 H), 4.97 (s, 2 H), 6.04 (d, J=8.2 Hz, 1 H), 7.20 - 7.24 (m, 1 H), 7.27 (d, J=7.7 Hz, 1 H), 7.45 (s, 1 H), 7.66 (td, J=7.7, 2.2 Hz, 1 H), 8.51 - 8.60 (m, 1 H)

	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
295		345.22	346	1.7, F	^{1}H NMR (300 MHz, CHLOROFORM-*d*) δ ppm 0.88 (t, J=7.0 Hz, 3 H), 1.17 - 1.45 (m, 4 H), 1.50 - 1.81 (m, 4 H), 1.95 (tdd, J=11.2, 11.2, 5.5, 2.6 Hz, 1 H), 2.59 (s, 3 H), 3.42 - 3.54 (m, 1 H), 3.56 - 3.66 (m, 1 H), 4.17 (m, J=11.1, 5.6, 2.8 Hz, 1 H), 4.51 (br. s., 2 H), 5.00 (s, 2 H), 5.77 (d, J=8.7 Hz, 1 H), 7.14 (t, J=6.7 Hz, 2 H), 7.53 (s, 1 H), 7.62 (t, J=7.6 Hz, 1 H)
296		326.23	327	0.84, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=6.8 Hz, 3 H), 1.12 (d, J=6.3 Hz, 6 H), 1.18 - 1.36 (m, 4 H), 1.41 - 1.73 (m, 4 H), 3.41 (t, J=6.4 Hz, 2 H), 3.55 - 3.67 (m, 3 H), 3.82 - 3.90 (m, 2 H), 4.04 - 4.18 (m, 1 H), 4.40 (br. s., 1 H), 5.58 (s, 2 H), 5.86 (d, J=9.0 Hz, 1 H), 7.43 (s, 1 H)
297		322.19	323	0.48, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.85 (t, J=7.00 Hz, 3 H) 1.13 - 1.39 (m, 4 H) 1.51 - 1.65 (m, 2 H) 1.66 - 1.81 (m, 2 H) 3.36 - 3.45 (m, 2 H) 4.28 - 4.39 (m, 1 H) 5.46 (s, 2 H) 7.51 (br. s., 2 H) 7.62 (s, 1 H) 8.23 (d, J=9.03 Hz, 1 H) 11.85 (br. s., 1 H)
298		321.19	322	0.58, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.84 (t, J=7.15 Hz, 3 H) 1.09 - 1.34 (m, 4 H) 1.46 - 1.61 (m, 2 H) 1.61 - 1.77 (m, 2 H) 4.24 - 4.34 (m, 1 H) 5.17 (s, 2 H) 7.47 (br. s., 2 H) 7.59 (d, J=5.52 Hz, 1 H) 8.05 (s, 1 H) 7.99 - 8.11 (m, 1 H) 11.89 (d, J=5.52 Hz, 1 H)
299		318.18	319	0.58, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.40 Hz, 3 H) 1.16 - 1.32 (m, 2 H) 1.47 - 1.65 (m, 2 H) 1.67 - 1.80 (m, 2 H) 4.29 - 4.40 (m, 1 H) 5.03 - 5.20 (m, 2 H) 5.23 (s, 2 H) 7.53 (br. s., 2 H) 7.63 (d, J=5.27 Hz, 1 H) 8.18 (d, J=8.78 Hz, 1 H) 8.61 - 8.71 (m, 2 H) 8.94 (s, 1 H) 12.05 (d, J=5.02 Hz, 1 H)

	STRUCTURE	Exact Mass	Mass Found [M+H]	LCMS Ret Time, Method	1H NMR
300		268.19	269	4.13, B	^{1}H NMR (400 MHz, CHLOROFORM-*d*) δ ppm 0.89 (t, J=7.0 Hz, 3 H), 1.16 (d, J=6.3 Hz, 3 H), 1.26 - 1.44 (m, 6 H), 1.51 (dd, J=8.7, 4.6 Hz, 1 H), 1.57 - 1.67 (m, 2 H), 3.63 - 3.75 (m, 1 H), 3.78 (s, 3 H), 4.07 - 4.24 (m, 1 H), 4.62 (br. s., 2 H), 5.07 (d, J=8.5 Hz, 1 H), 7.34 (s, 1 H)
301		312.22	313	0.75, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.86 (t, J=7.3 Hz, 3 H), 1.12 (d, J=6.0 Hz, 6 H), 1.28 (dt, J=14.7, 7.5 Hz, 2 H), 1.48 (q, J=7.4 Hz, 2 H), 1.54 - 1.62 (m, 1 H), 1.63 - 1.74 (m, 1 H), 3.38 - 3.46 (m, 2 H), 3.54 - 3.68 (m, 3 H), 3.86 (dd, J=5.5, 4.0 Hz, 2 H), 4.14 (d, J=4.8 Hz, 1 H), 4.34 - 4.48 (m, 1 H), 5.58 (s, 2 H), 5.86 (d, J=9.0 Hz, 1 H), 7.43 (s, 1 H)
302		317.19	318	0.61, D	^{1}H NMR (400 MHz, DMSO-d_6) δ ppm 0.87 (t, J=7.4 Hz, 3 H), 1.19 - 1.34 (m, 2 H), 1.41 - 1.60 (m, 2 H), 1.66 (s, 2 H), 3.44 (d, J=6.5 Hz, 2 H), 4.09 - 4.26 (m, 1 H), 4.41 - 4.50 (m, 1 H), 5.04 (s, 2 H), 5.61 (br. s., 2 H), 6.36 (d, J=8.5 Hz, 1 H), 7.39 (s, 1 H), 7.42 - 7.46 (m, 2 H), 8.52 - 8.61 (m, 2 H)

Analytical Methods.

All compounds were characterized by LC-MS. The following LC-MS methods were used:

Method A. Waters Aquity UPLC equipped with a PDA detector (210-400 nm) and a Waters SQD with a dual mode ion source ES+/-. The column used was a Halo C18, 2.7 µm, 2.1 x 50 mm, heated to 50ºC. A gradient of 95% aqueous formic acid (0.1%)/5% acetonitrile to 100% acetonitrile was ramped over 1.5 minutes, held for 0.6 minutes, then returns to 100% aqueous formic acid (0.1%) for 0.5 minutes. The flow rate was 0.6 mL/min.

Method B.

Column	YMC-PACK ODS-AQ, 50×2.0mm 5μm		
Mobile Phase	A :H_2O (0.1%TFA)		
	B:acetonitrile (0.05%TFA)		
Gradient	StopTime:10min		
	PostTime :OFF		
	TIME(min)	A%	B%
	0	100	0
	1	100	0
	5	40	60
	7.5	40	60
	8	100	0
Flow Rate	0.8ml/min		
Wavelength	UV 220nm		
Column Temperture	50°C		
MS polarity	positive		
LCMS	Agilent 1100		

Method C.

Column	YMC-PACK ODS-AQ, 50×2.0mm 5μm		
Mobile Phase	A :H_2O (0.1%TFA)		
	B:acetonitrile (0.05%TFA)		
Gradient	StopTime:10min		
	Post Time :OFF		
	TIME(min)	A%	B%
	0	90	10
	0.8	90	10
	4.5	20	80
	7.5	20	80
	8	90	10
Flow Rate	0.8ml/min		
Wavelength	UV 220nm		
Oven Tem.	50°C		
MS polarity	positive		
LCMS	Agilent 1100		

Method D. Reversed phase UPLC (Ultra Performance Liquid Chromatography) was carried out on a bridged ethylsiloxane/silica hybrid (BEH) C18 column (1.7 µm, 2.1 x 50 mm; Waters Acquity) with a flow rate of 0.8 ml/min. Two mobile phases (10 mM ammonium acetate in H_2O/acetonitrile 95/5; mobile phase B: acetonitrile) were used to run a gradient condition from 95 % A and 5 % B to 5 % A and 95 % B in 1.3 minutes and hold for 0.7 minutes. An injection volume of 0.75 µl was used. Cone voltage was 30 V for positive ionization mode and 30 V for negative ionization mode.

Method E. Using a Phenomenex Kinetex column(XB-C18 50 x 4.6 mm I.D. 2.6u) held at 35ºC. MS detection: API-ES Positive ionization mode, Mass range 100-1200. PDA detection (λ=190-400nm). The following gradient was used with a 2µL injection:

Solvent A	H_2O + 0.1% Formic Acid		
Solvent B	Acetonitrile		
Time (min)	%A	%B	Flow (ml/min)
0.0	95	5	3.0
4.2	5	95	3.0
4.9	5	95	3.0
5.0	95	5	3.0

Method F. Using a YMC ODS-AQ C-18;50 x 4.6 mm, ID = 3µm held at 35ºC. MS detection: API-ES Positive ionization mode, Mass range 100-1400. PDA detection (λ=190-400nm). The following gradient was used with a 2µL injection:

Solvent A	H_2O + 0.1% Formic Acid		
Solvent B	Acetonitrile		
Time (min)	%A	%B	Flow (ml/min)
0.0	95	5	2.6
4.8	5	95	2.6
5.8	5	95	2.6
6.0	95	5	2.6

Method G. Alliance HT 2790 (Waters) system comprising a quaternary pump with degasser, an autosampler, a column oven (set at 40 °C). Flow from the column was split to a MS spectrometer. The MS detector was configured with an electrospray ionization source. The capillary needle voltage was 3 kV and the source temperature was maintained at 140 °C. Nitrogen was used as the nebulizer gas. Xterra MS C18 column (3.5 µm, 4.6 x 100 mm) with a flow rate of 1.6 mL/min. Three mobile phases (mobile phase A: 95% 25 mM ammoniumacetate + 5 % acetonitrile; mobile phase B: acetonitrile; mobile phase C: methanol) were employed to run a gradient condition from 100 % A to 50 % B and 50 % C in 6.5 minutes, to 100 % B in 0.5 minute, 100 % B for 1 minute and re-equilibrate with 100 % A for 1.5 minutes. An injection volume of 10 µl was used.

Method H. Reversed phase UPLC (Ultra Performance Liquid Chromatography) was carried out on a bridged ethylsiloxane/silica hybrid (BEH) C18 column (1.7 µm, 2.1 x 50 mm; Waters Acquity) with a flow rate of 0.8 mL/min. Two mobile phases (mobile phase A: 10mM ammonium acetate in H_2O/acetonitrile 95/5; mobile phase B: acetonitrile) were used to run a gradient condition from 95 % A and 5 % B to 5 % A and 95 % B in 1.3 minutes and hold for 0.2 minutes. An injection volume of 0.5 µl was used. Cone voltage was 10 V for positive ionization mode and 20 V for negative ionization mode.

Biological Activity of compounds of formula (I)

Description of Biological Assays

Assessment of TLR7 and TLR8 activity

The ability of compounds to activate human TLR7 and/or TLR8 was assessed in a cellular reporter assay using HEK293 cells transiently transfected with a TLR7 or TLR8 expression vector and NFκB-luc reporter construct. In one instance the TLR expression construct expresses the respective wild type sequence or a mutant sequence comprising a deletion in the second leucine-rich repeat of the TLR. Such mutant TLR proteins have previously been shown to be more susceptible to agonist activation (US 7498409).

Briefly, HEK293 cells were grown in culture medium (DMEM supplemented with 10% FCS and 2 mM Glutamine). For transfection of cells in 10 cm dishes, cells were detached with Trypsin-EDTA, transfected with a mix of CMV-TLR7

or TLR8 plasmid (750 ng), NFκB-luc plasmid (375 ng) and a transfection reagent and incubated for 48 hours at 37°C in a humidified 5% CO_2 atmosphere. Transfected cells were then detached with Trypsin-EDTA, washed in PBS and resuspended in medium to a density of 1.67 x 10^5 cells/mL. Thirty microliters of cells were then dispensed into each well in 384-well plates, where 10 µL of compound in 4% DMSO was already present. Following 6 hours incubation at 37°C, 5% CO_2, the luciferase activity was determined by adding 15 µl of Steady Lite Plus substrate (Perkin Elmer) to each well and readout performed on a ViewLux ultraHTS microplate imager (Perkin Elmer). Dose response curves were generated from measurements performed in quadruplicates. Lowest effective concentrations (LEC) values, defined as the concentration that induces an effect which is at least two fold above the standard deviation of the assay, were determined for each compound.

Compound toxicity was determined in parallel using a similar dilution series of compound with 30 µL per well of cells transfected with the CMV-TLR7 construct alone (1.67 x 10^5 cells/mL), in 384-well plates. Cell viability was measured after 6 hours incubation at 37°C, 5% CO_2 by adding 15 µL of ATP lite (Perkin Elmer) per well and reading on a ViewLux ultraHTS microplate imager (Perkin Elmer). Data was reported as CC_{50}.

Suppression of HCV replicon replication

Activation of human TLR7 results in robust production of interferon by plasmacytoid dendritic cells present in human blood. The potential of compounds to induce interferon was evaluated by looking at the antiviral activity in the HCV replicon system upon incubation with conditioned media from peripheral blood mononuclear cells (PBMC). The HCV replicon assay is based on a bicistronic expression construct, as described by Lohmann *et al.* (Science (1999) 285: 110-113; Journal of Virology (2003) 77: 3007-15 3019) with modifications described by Krieger *et al.* (Journal of Virology (2001) 75: 4614-4624). The assay utilized the stably transfected cell line Huh-7 luc/neo harboring an RNA encoding a bicistronic expression construct comprising the wild type NS3-NS5B regions of HCV type 1b translated from an Internal Ribosome Entry Site (IRES) from encephalomyocarditis virus (EMCV), preceded by a reporter gene (Firefly-luciferase) and a selectable marker gene (neoR, neomycine phosphotransferase). The construct is flanked by 5' and 3' NTRs (non-translated regions) from HCV type 1b. Continued culture of the replicon cells in the presence of G418 (neoR) is dependent on the replication of

the HCV RNA. The stably transfected replicon cells that replicate HCV RNA autonomously and to high levels, encoding *inter alia* luciferase, were used for profiling of the conditioned cell culture media.

Briefly, PBMCs were prepared from buffy coats of at least two donors using a standard Ficoll centrifugation protocol. Isolated PBMCs were resuspended in RPMI medium supplemented with 10% human AB serum and 2 x 10^5 cells/well were dispensed into 384-well plates containing compounds (70 µL total volume). After overnight incubation, 10 µL of supernatant was transferred to 384-well plates containing 2.2 x 10^3 replicon cells/well in 30 µL (plated the day before). Following 24 hours of incubation, replication was measured by assaying luciferase activity using 40 µL/well Steady Lite Plus substrate (Perkin Elmer) and measured with ViewLux ultraHTS microplate imager (Perkin Elmer). The inhibitory activity of each compound on the Huh7-luc/neo cells were reported as EC_{50} values, defined as the compound concentration applied to the PBMCs resulting in a 50% reduction of luciferase activity which in turn indicates the degree of replication of the replicon RNA on transfer of a defined amount of PBMC culture medium. Recombinant interferon α-2a (Roferon-A) was used as a standard control compound.

Biological activity of compounds of formula (I) . All compounds showed CC50 of >24uM in the HEK 293 TOX assay described above.

Activation of ISRE promoter elements

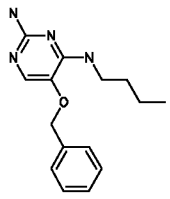
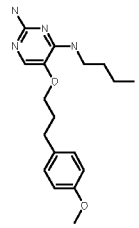
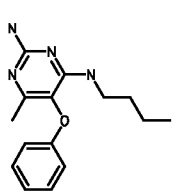
The potential of compounds to induce IFN-I was also evaluated by measuring the activation of interferon-stimulated responsive elements (ISRE) by conditioned media from PBMC. The ISRE element of sequence GAAACTGAAACT is highly responsive to the STAT1-STAT2-IRF9 transcription factor, activated upon binding of IFN-I to their receptor IFNAR (Clontech, PT3372-5W). The plasmid pISRE-Luc from Clontech (ref. 631913) contains 5 copies of this ISRE element, followed by the firefly luciferase ORF. A HEK293 cell line stably transfected with pISRE-Luc (HEK-ISREluc) was established to profile of the conditioned PBMC cell culture media.

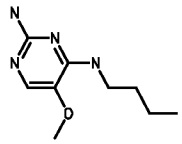
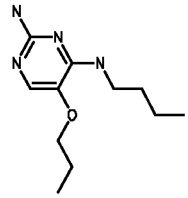
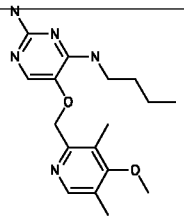
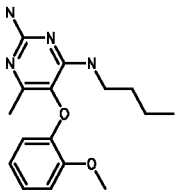
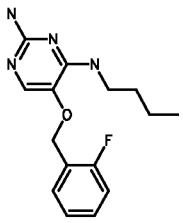
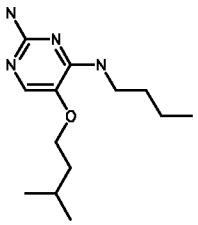
Briefly, PBMCs were prepared from buffy coats of at least two donors using a standard Ficoll centrifugation protocol. Isolated PBMCs were resuspended in RPMI medium supplemented with 10% human AB serum and 2 x 10^5 cells/well were dispensed into 384-well plates containing compounds (70 µL total volume). After overnight incubation, 10 µL of supernatant was transferred to

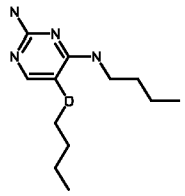
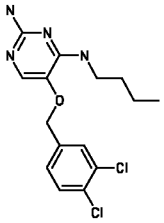
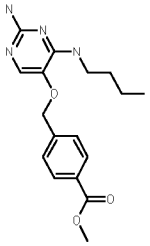
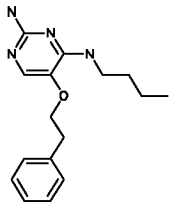
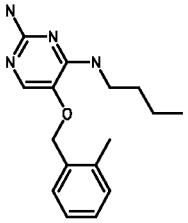
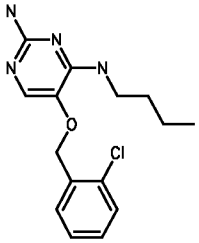
384-well plates containing 5 x 10^3 HEK-ISREluc cells/well in 30 µL (plated the day before). Following 24 hours of incubation, activation of the ISRE elements was measured by assaying luciferase activity using 40 µL/well Steady Lite Plus substrate (Perkin Elmer) and measured with ViewLux ultraHTS microplate imager (Perkin Elmer). The stimulating activity of each compound on the HEK-ISREluc cells was reported as LEC value, defined as the compound concentration applied to the PBMCs resulting in a luciferase activity at least two fold above the standard deviation of the assay. The LEC in turn indicates the degree of ISRE activation on transfer of a defined amount of PBMC culture medium. Recombinant interferon α-2a (Roferon-A) was used as a standard control compound.

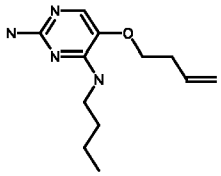
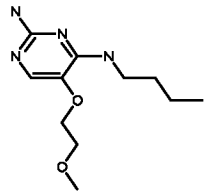
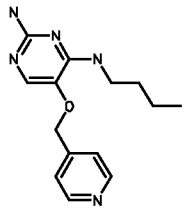
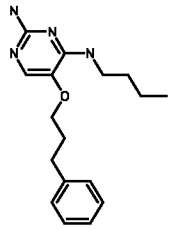
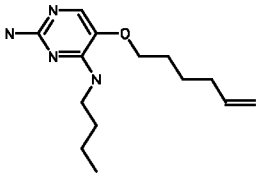
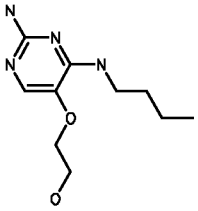
For a given compound, the LEC value obtained from this assay were in the same range as the EC_{50} values obtained from the "suppression of HCV replication assay." Thus, it is possible to compare the potential of compounds to induce IFN-I by PBMC, measured by either of the 2 assays.

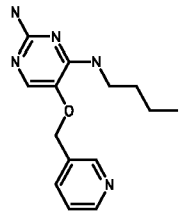
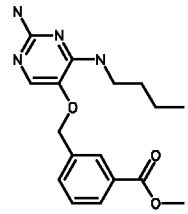
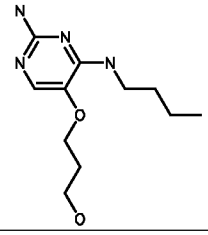
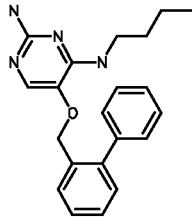
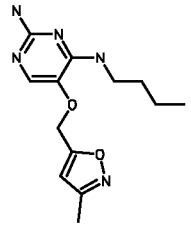
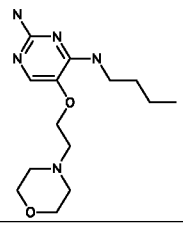
TABLE II BIOLOGICAL ACTIVITY OF THE COMPOUNDS.

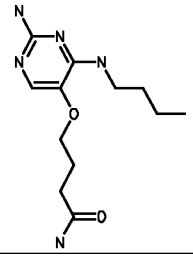
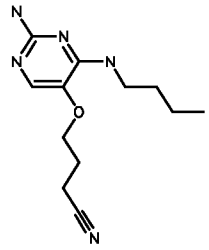
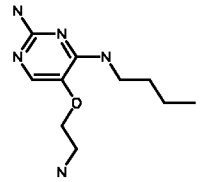
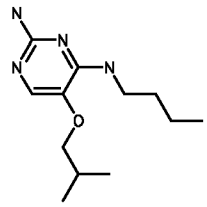
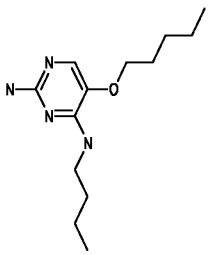
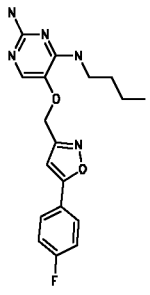
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
1		0.90	0.55	2.42	1.30	0.70
2			0.02	1.34	0.31	0.04
3			21.69	4.91	1.66	10.80

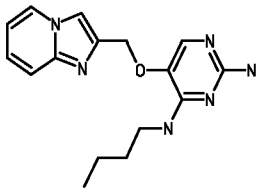
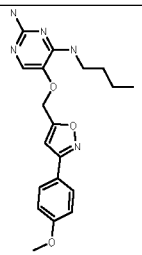
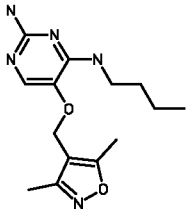
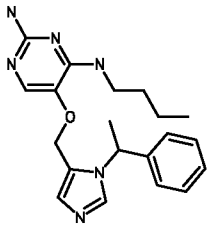
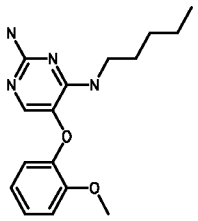
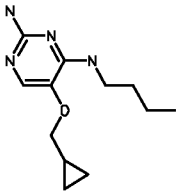
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
4		6.71	1.17	2.56	1.10	1.14
5		6.18	1.69	4.53	2.30	2.65
6			0.01	0.16	0.10	0.02
7			1.11	5.84	3.03	3.11
8			0.38	1.88	0.81	0.37
9			1.55	19.00	9.70	9.72

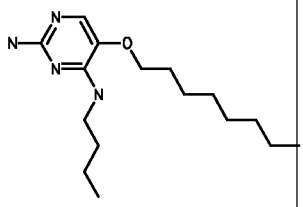
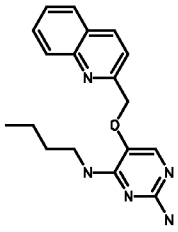
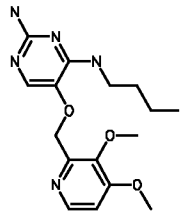
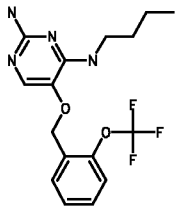
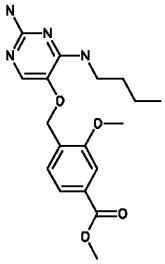
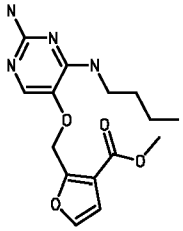
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
10			1.28	8.14	2.82	1.58
11			0.60	4.52	3.50	0.78
12			0.26	1.49	1.12	2.76
13			2.26	13.18	5.23	2.53
14			0.91	4.92	0.91	1.81
15			0.17	2.31	1.06	0.17

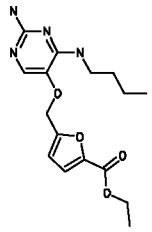
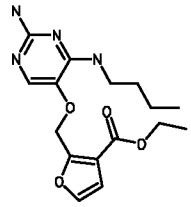
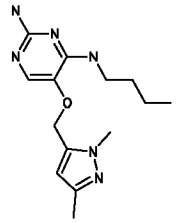
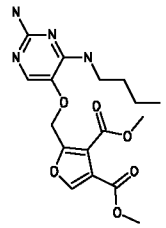
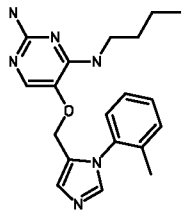
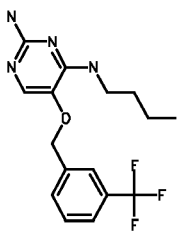
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
16			0.49	2.68	0.59	0.79
17			0.34	2.03	0.67	0.71
18			0.83	1.87	0.85	0.63
19		1.53	0.16	7.94	2.36	0.43
20			0.79	10.21	2.87	1.33
21			1.61	2.69	0.64	3.08

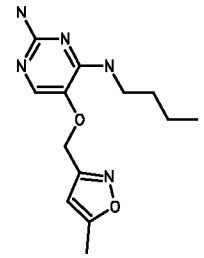
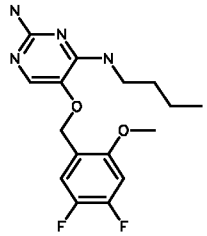
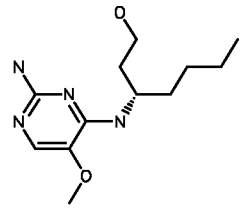
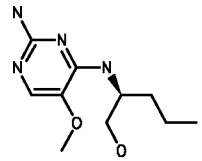
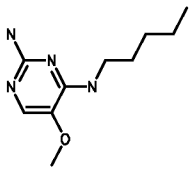
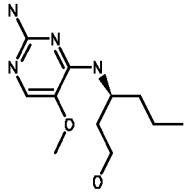
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
22			0.31	2.35	0.94	0.25
23			0.26	2.55	1.45	10.84
24			1.99	2.42	1.50	2.75
25			0.64	>25	>25	1.55
26			0.49	3.90	1.52	0.58
27			0.78	5.36	0.64	1.03

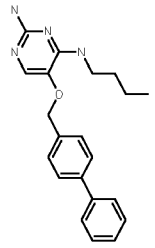
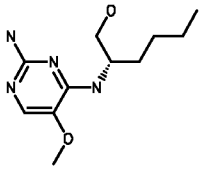
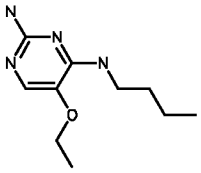
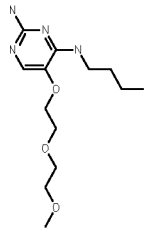
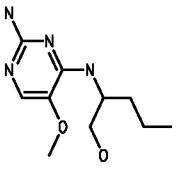
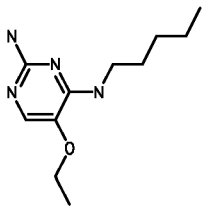
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
28			2.47	9.18	6.99	1.75
29			1.32	2.86	1.19	2.97
30			>25	6.44	1.16	9.07
31		>24.59	5.27	17.53	6.46	10.36
32		10.60	1.35	9.97	4.43	1.06
33			0.36	1.78	1.17	1.48

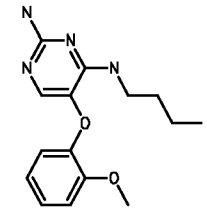
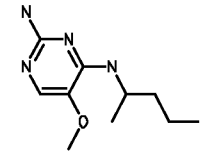
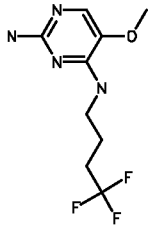
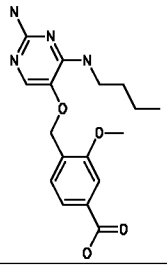
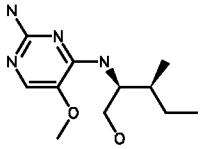
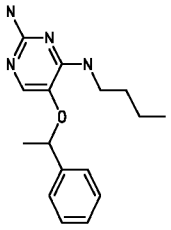
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
34			0.06	0.83	0.61	0.05
35			0.39	1.67	1.66	1.50
36			0.58	1.68	0.82	0.70
37			0.04	9.22	5.69	0.12
38		21.97	2.46	>50	22.88	11.28
39			3.01	14.41	7.10	

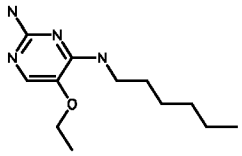
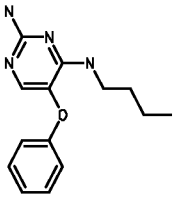
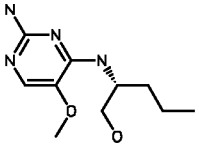
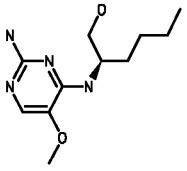
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
40			2.69	>25	>25	
41			0.03	0.83	0.51	0.10
42			0.04	1.15	0.41	0.04
43			0.08	8.22	1.66	0.79
44			0.16	3.11	1.96	0.59
45			0.17	0.58	0.40	0.17

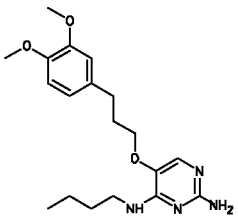
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
46			0.19	3.85	1.96	2.51
47			0.20	1.87	0.66	0.33
48			0.28	1.75	0.60	0.64
49			0.31	3.72	2.07	0.55
50			0.51	>25	>25	0.78
51			0.58	3.92	2.09	0.50

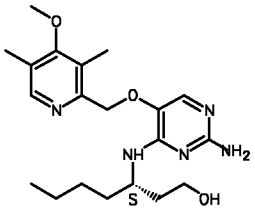
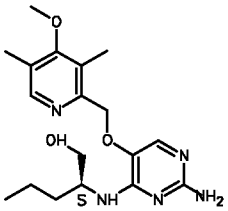
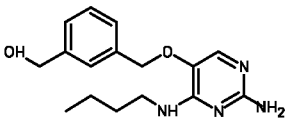
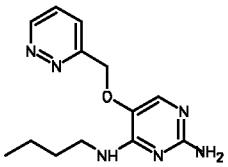
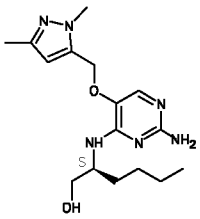
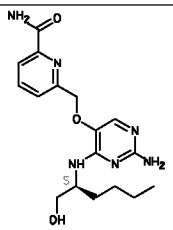
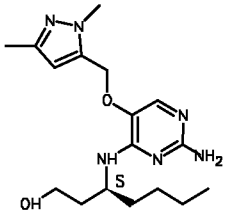
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
52			0.63	3.61	1.65	0.26
53			0.64	3.06	2.15	0.60
54			0.68	1.40	0.69	0.75
55			0.72	0.16	0.12	0.41
56		12.02	0.84	5.55	1.47	0.80
57			0.88	1.80	0.74	0.80

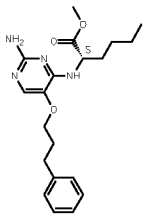
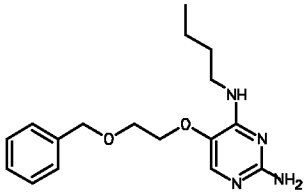
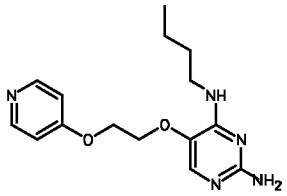
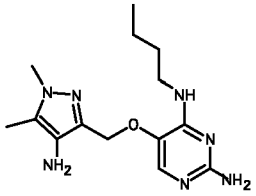
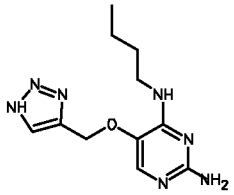
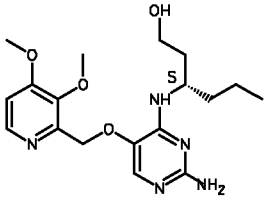
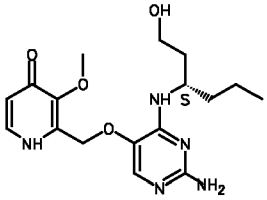
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
58		6.48	0.99	3.84	2.17	2.99
59			1.20	0.36	0.13	0.40
60		5.58	1.38	2.08	0.65	1.91
61			1.38	3.59	1.56	1.91
62		21.26	1.76	0.55	0.15	0.74
63		2.78	1.79	6.35	1.94	2.69

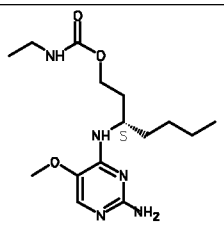
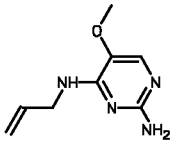
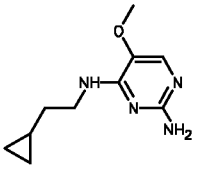
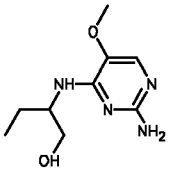
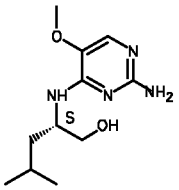
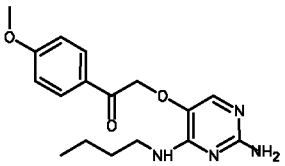
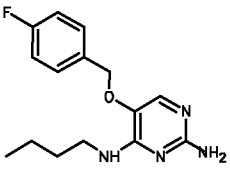
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
64		8.47	2.03	18.43	7.65	4.29
65		21.59	2.04	3.68	1.13	2.30
66			2.29	9.03	1.89	2.27
67			2.31	>24.59	>24.59	2.43
68			2.54	0.56	0.43	1.17
69			3.75	6.43	2.22	6.16

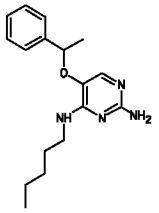
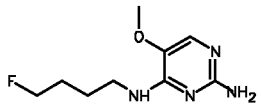
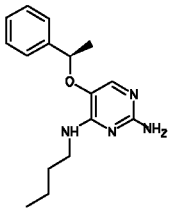
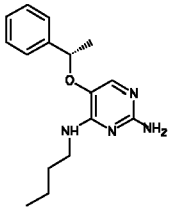
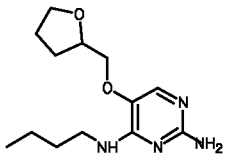
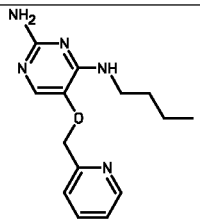
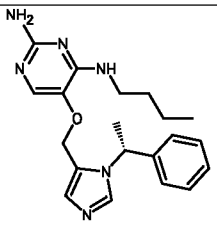
	STRUCTURE	TLR7-wt_LEC	TLR7-dIRR2_LEC	TLR8-wt_LEC	TLR8-dIRR2_LEC	PBMC-HUH7_EC_{50}
70		15.84	4.96	>24.59	>24.59	>23.81
71			>24.59	>24.59	>24.59	4.96
72			>25	6.57	6.24	17.50
73			>25	0.80	0.47	1.39

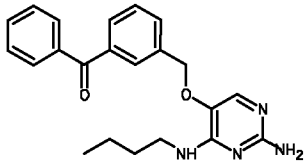
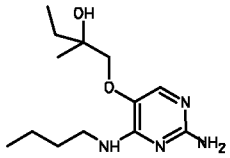
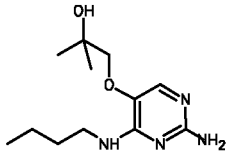
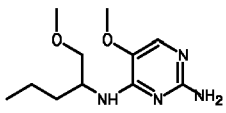
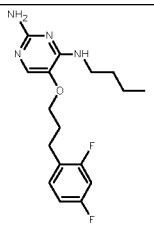
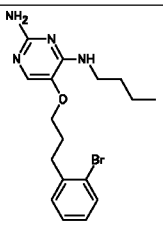
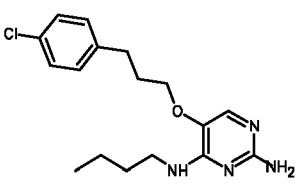
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
74		0.713	1.720	0.157	ND

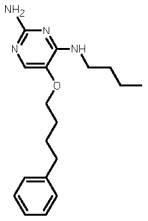
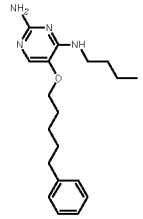
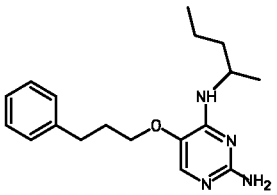
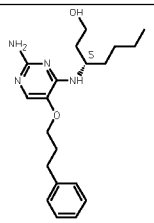
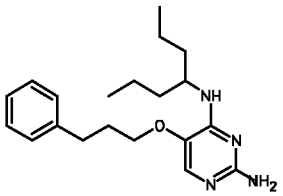
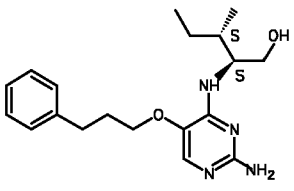
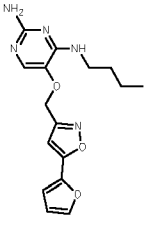
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
75		0.023	0.218	0.007	ND
76		0.021	0.055	ND	0.008
77		0.449	0.623	ND	0.137
78		0.519	0.827	ND	0.123
79		1.620	0.329	ND	0.235
80		0.560	0.041	ND	0.027
81		0.101	0.429	ND	0.086

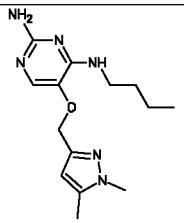
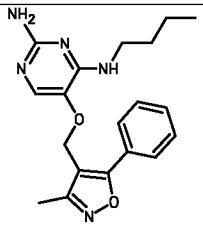
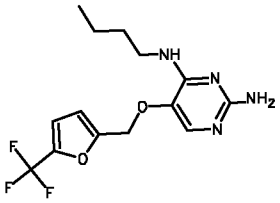
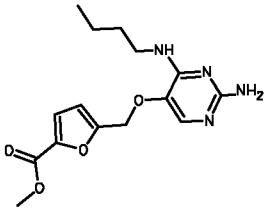
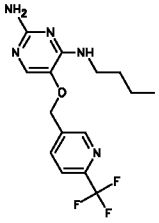
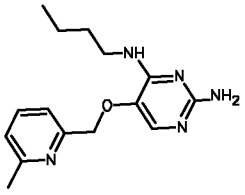
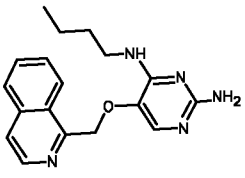
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
82		4.420	13.590	14.020	ND
83		0.997	1.610	0.204	ND
84		0.860	0.250	0.076	ND
85		0.509	2.960	0.209	ND
86		0.646	3.750	ND	0.131
87		0.013	0.567	0.012	ND
88		3.090	6.960	ND	0.050

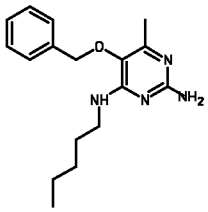
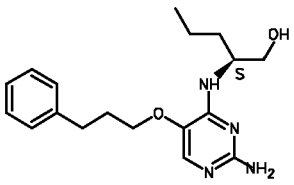
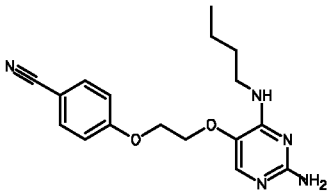
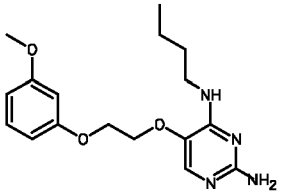
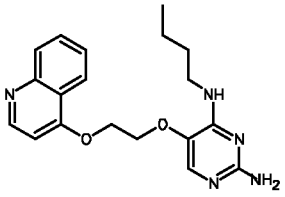
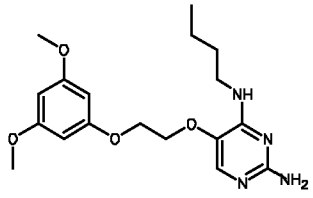
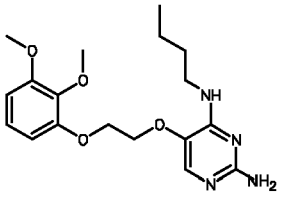
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
89		1.670	6.670	ND	0.526
90		>25	8.460	6.950	ND
91		>25	20.850	7.650	ND
92		>25	14.570	20.160	ND
93		>25	15.880	9.050	ND
94		1.590	3.170	0.696	ND
95		2.730	2.010	0.726	ND

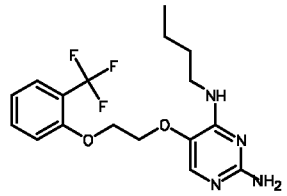
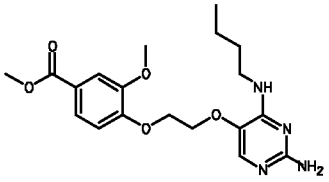
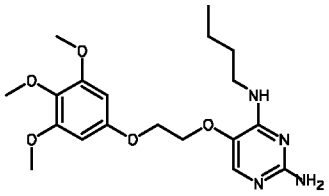
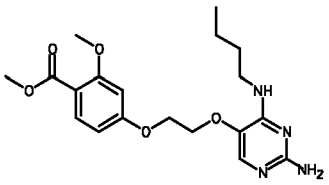
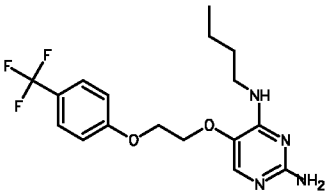
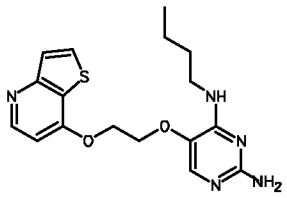
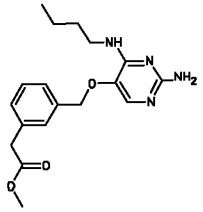
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
96		>25	6.340	4.310	ND
97		21.810	5.070	2.640	ND
98		>25	10.100	21.960	ND
99		8.980	1.820	1.280	ND
100		18.950	6.160	5.120	ND
101		0.277	0.597	0.055	ND
102		0.141	5.690	0.012	ND

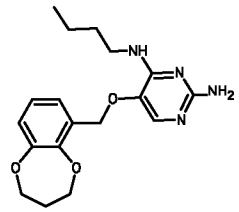
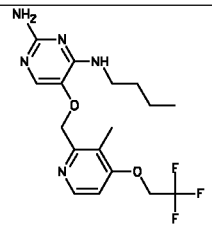
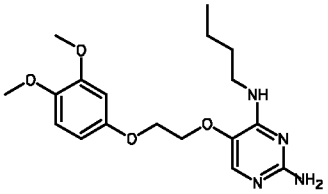
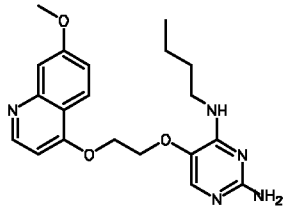
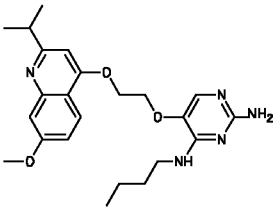
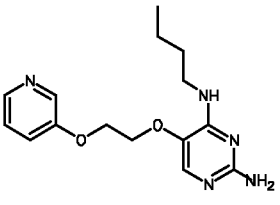
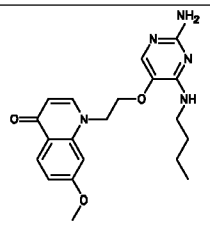
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
103		1.190	1.270	0.725	ND
104		>25	12.390	>23.81	ND
105		>25	22.020	19.050	ND
106		16.100	5.940	3.150	ND
107		2.460	3.940	1.590	ND
108		6.580	>25	6.770	ND
109		0.790	2.230	0.393	ND

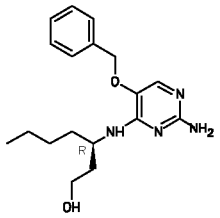
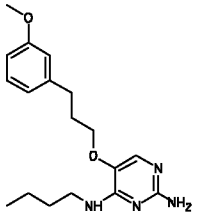
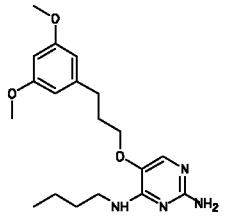
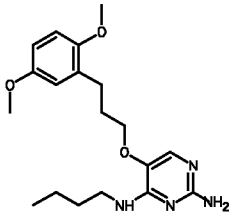
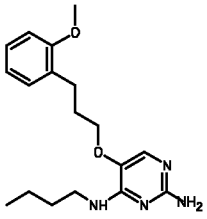
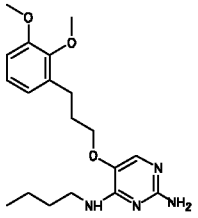
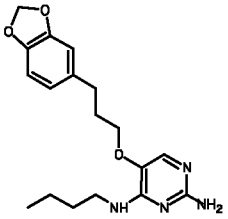
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
110		2.380	3.780	0.740	ND
111		0.257	ND	0.096	ND
112		3.960	5.560	3.350	ND
113		0.433	2.240	0.251	ND
114		2.020	>25	2.000	ND
115		6.180	6.510	3.730	ND
116		0.652	1.610	0.066	ND

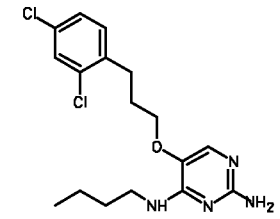
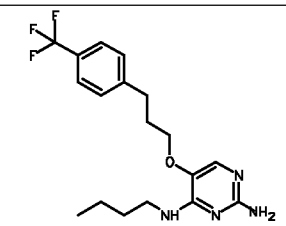
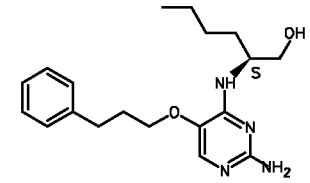
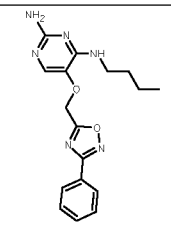
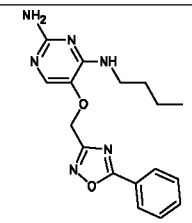
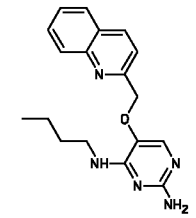
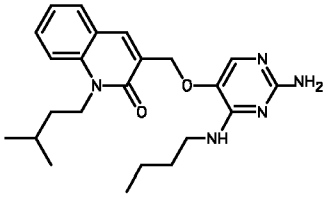
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
117		0.335	1.120	0.088	ND
118		1.670	3.710	0.976	ND
119		1.720	6.120	0.193	ND
120		0.649	3.910	0.273	ND
121		0.797	3.020	0.272	ND
122		0.118	0.628	0.025	ND
123		0.008	0.143	0.002	ND

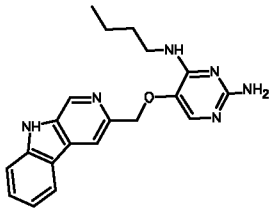
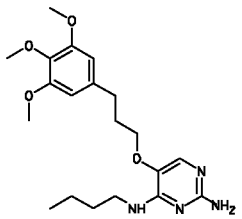
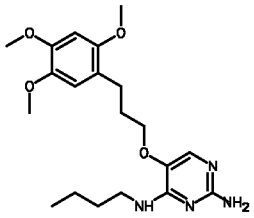
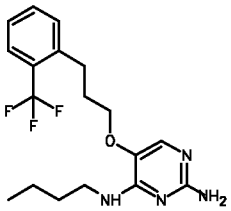
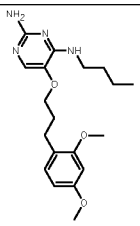
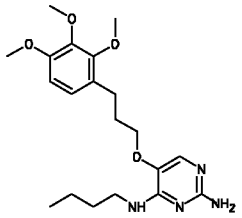
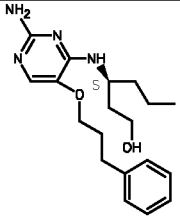
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
124		15.610	13.650	>23.81	ND
125		1.630	0.598	0.336	ND
126		1.000	1.020	0.264	ND
127		1.030	2.050	0.256	ND
128		2.430	3.740	0.284	ND
129		2.090	3.250	0.432	ND
130		0.676	6.560	0.103	ND

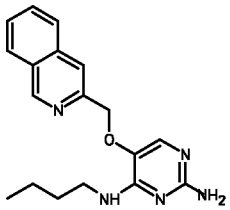
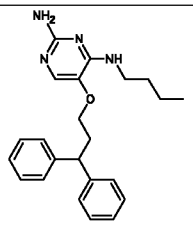
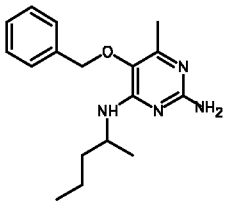
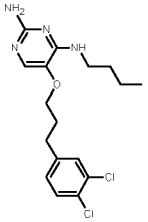
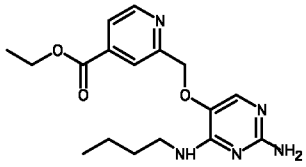
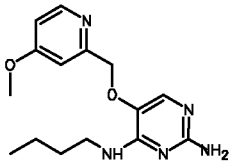
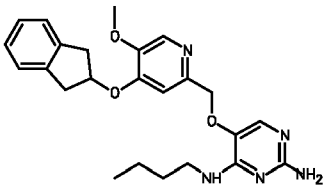
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
131		1.700	>25	0.806	ND
132		1.470	>25	0.634	ND
133		1.500	3.090	0.585	ND
134		2.010	2.110	0.935	ND
135		3.230	1.970	3.190	ND
136		2.000	2.030	0.275	ND
137		0.757	1.760	22.760	ND

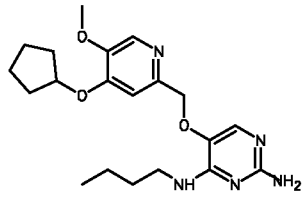
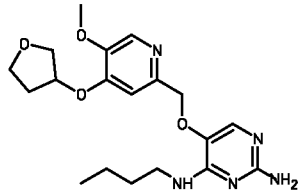
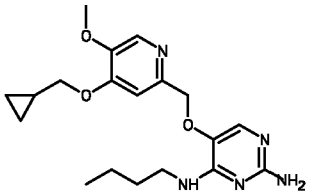
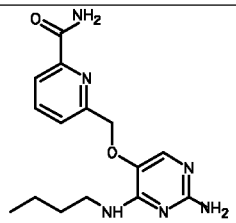
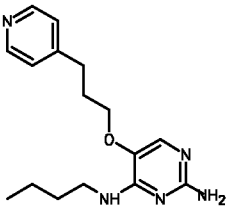
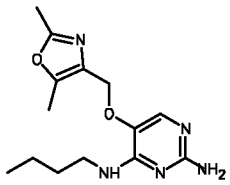
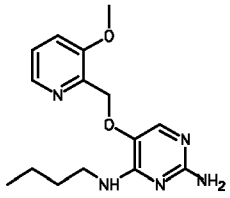
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
138		1.040	1.050	0.570	ND
139		0.025	0.286	0.009	ND
140		0.617	2.250	0.175	ND
141		4.360	0.704	0.733	ND
142		>25	2.370	19.680	ND
143		1.810	0.880	0.443	ND
144		13.010	20.790	1.320	ND

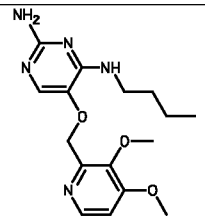
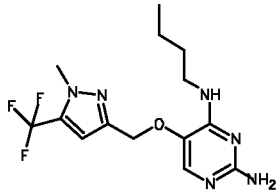
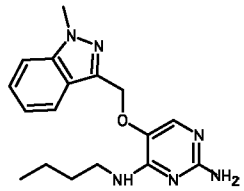
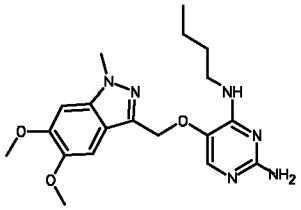
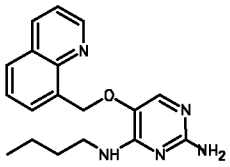
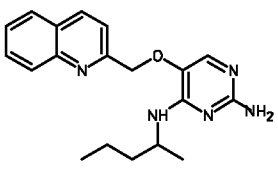
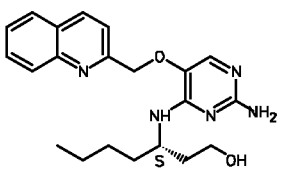
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
145		2.140	1.920	0.632	ND
146		1.230	2.300	0.707	ND
147		2.520	4.340	0.746	ND
148		0.691	4.850	0.634	ND
149		2.080	5.850	0.703	ND
150		1.180	4.310	0.689	ND
151		0.431	1.860	0.188	ND

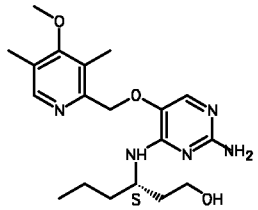
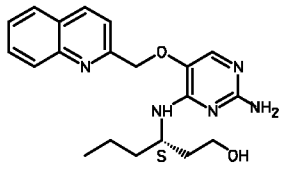
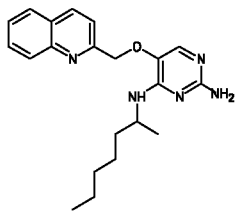
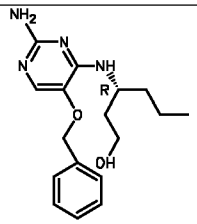
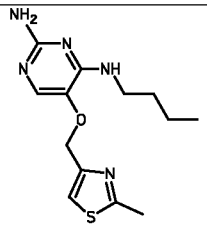
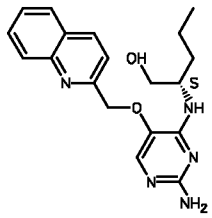
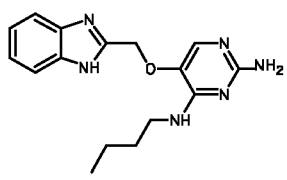
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
152		5.410	>25	3.350	ND
153		10.640	ND	3.430	ND
154		3.710	2.960	3.020	ND
155		2.660	4.560	3.440	ND
156		0.828	2.060	0.697	ND
157		0.333	1.110	0.162	ND
158		3.080	>25	3.310	ND

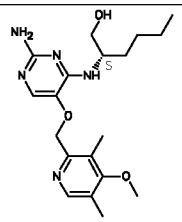
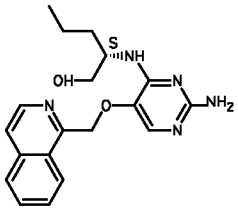
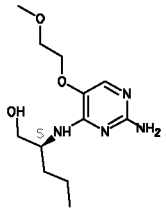
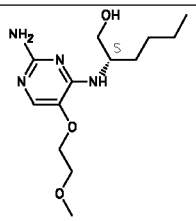
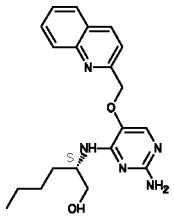
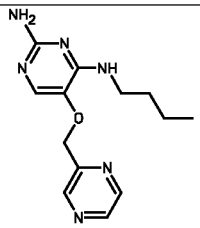
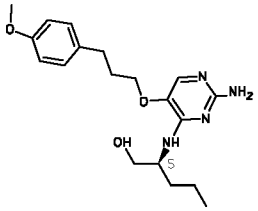
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
159		0.159	1.080	0.018	ND
160		0.756	2.710	0.634	ND
161		0.672	3.480	0.629	ND
162		11.850	>25	ND	ND
163		0.573	2.500	0.728	ND
164		0.606	23.030	0.769	ND
165		0.683	1.800	0.187	ND

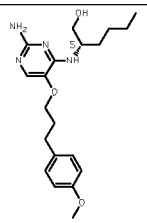
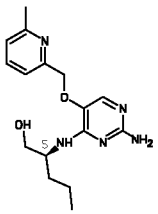
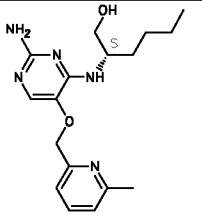
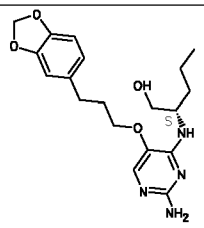
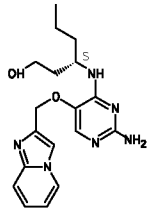
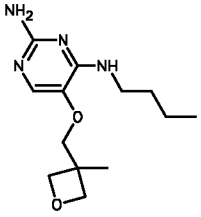
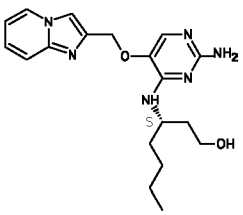
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
166		0.128	0.980	0.046	ND
167		1.200	>25	0.764	ND
168		>25	14.900	11.740	ND
169		5.110	>25	3.130	ND
170		0.319	1.750	2.630	ND
171		0.396	1.060	0.158	ND
172		0.187	2.000	0.045	ND

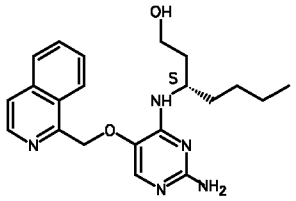
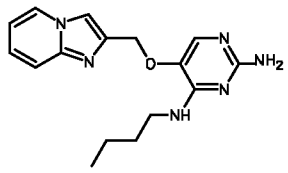
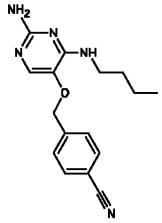
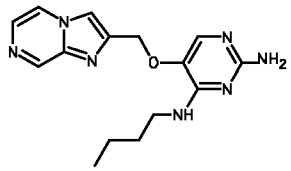
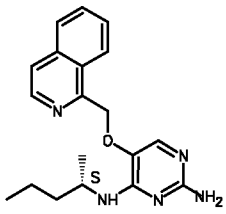
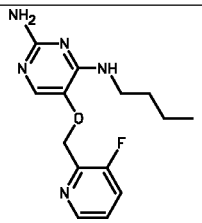
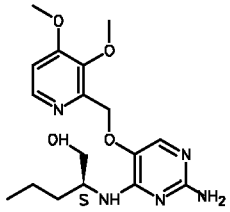
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
173		0.222	2.550	0.086	ND
174		0.447	2.610	0.052	ND
175		0.367	2.480	0.167	ND
176		0.868	0.463	0.173	ND
177		0.795	0.819	0.197	ND
178		0.810	0.410	0.302	ND
179		0.078	0.142	0.021	ND

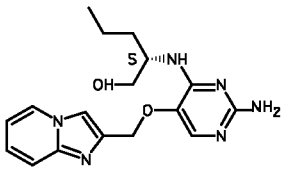
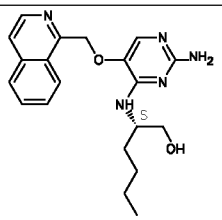
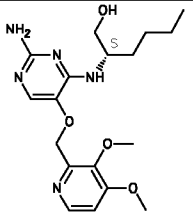
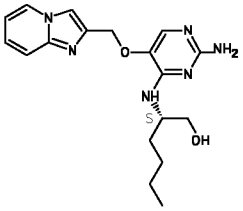
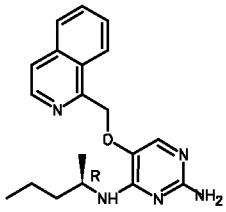
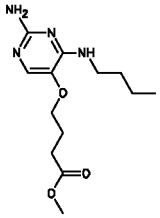
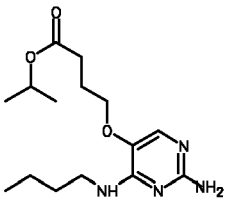
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
180		0.135	0.524	0.047	ND
181		0.146	1.210	0.096	ND
182		0.014	0.178	0.007	ND
183		0.056	1.580	0.023	ND
184		0.157	1.650	0.053	ND
185		0.743	2.340	0.488	ND
186		0.122	0.680	0.065	ND

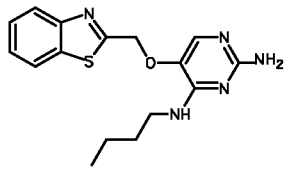
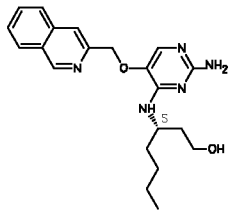
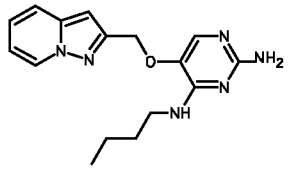
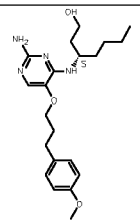
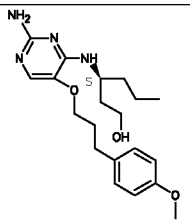
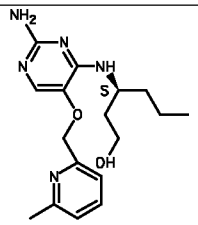
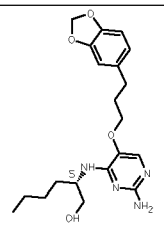
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
187		0.074	0.178	0.022	ND
188		0.237	0.530	0.086	ND
189		11.990	>25	17.570	ND
190		8.620	4.330	3.230	ND
191		0.286	0.743	0.066	ND
192		0.080	0.220	0.044	ND
193		0.032	0.654	0.017	ND

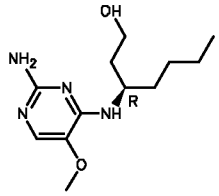
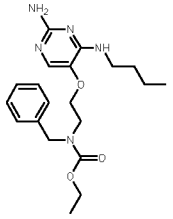
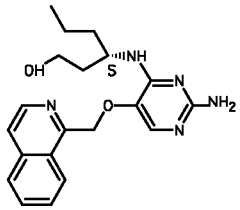
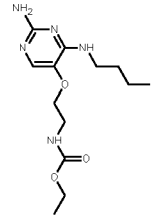
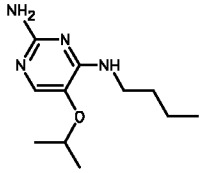
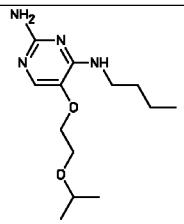
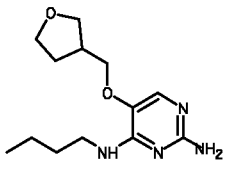
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
194		0.031	0.164	0.019	ND
195		0.003	0.056	0.003	ND
196		1.900	0.469	0.687	ND
197		2.650	0.624	0.767	ND
198		0.076	0.511	0.089	ND
199		0.512	2.280	0.218	ND
200		0.253	0.181	0.200	ND

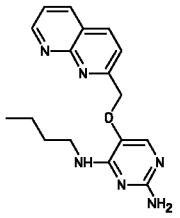
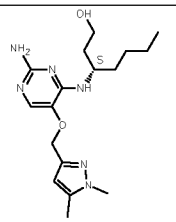
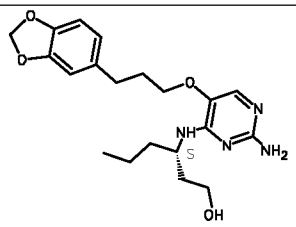
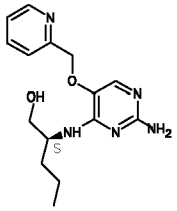
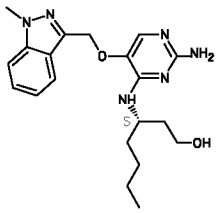
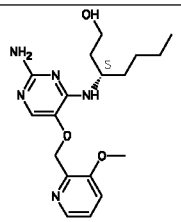
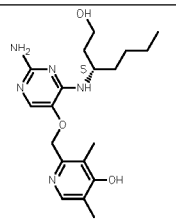
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
201		0.566	0.647	0.758	ND
202		0.164	0.089	0.049	ND
203		0.124	0.160	0.054	ND
204		0.791	0.791	0.493	ND
205		0.369	1.110	0.047	ND
206		>25	9.450	>23.81	ND
207		0.177	1.450	0.063	ND

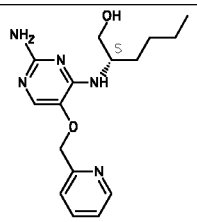
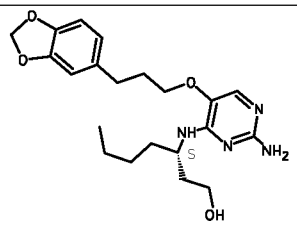
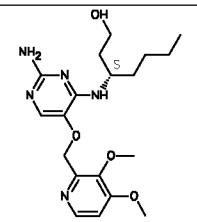
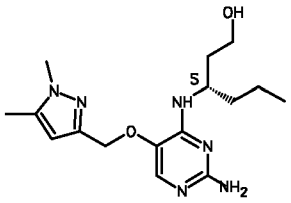
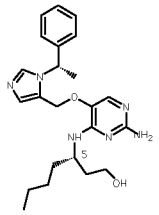
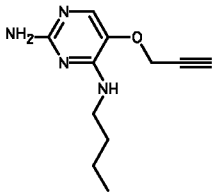
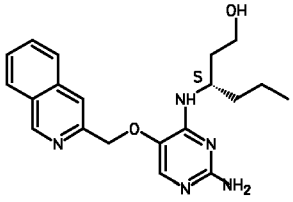
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
208		0.001	0.093	0.000	ND
209		0.074	0.667	0.076	ND
210		0.686	0.896	0.237	ND
211		0.208	1.040	0.097	ND
212		0.007	0.148	0.005	ND
213		0.225	0.207	ND	0.032
214		0.134	0.593	ND	0.027

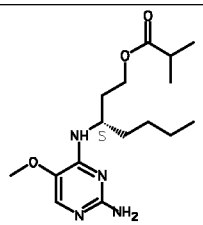
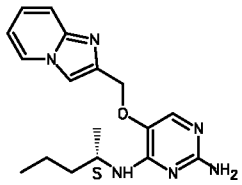
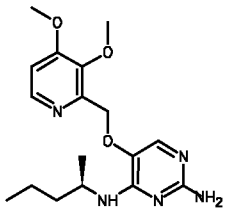
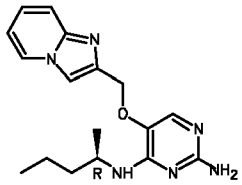
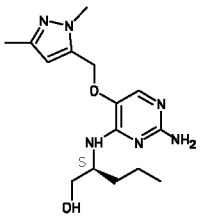
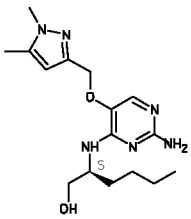
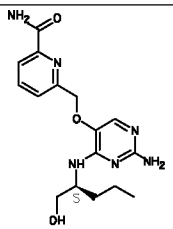
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
215		0.171	0.300	ND	0.029
216		0.008	0.111	ND	0.002
217		0.106	0.433	ND	0.007
218		0.154	0.352	ND	0.032
219		0.125	1.640	ND	0.029
220		1.940	1.450	ND	1.720
221		0.654	0.859	ND	0.496

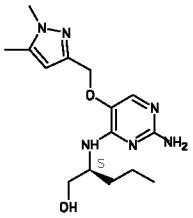
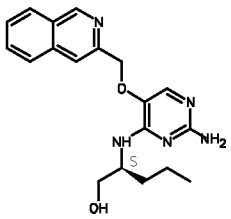
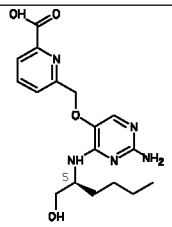
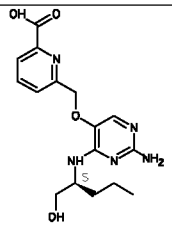
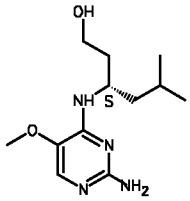
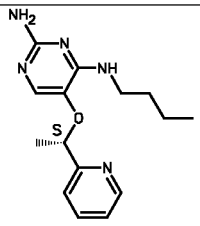
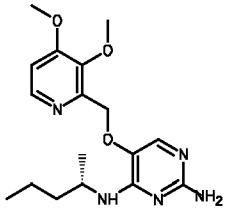
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
222		0.277	2.600	ND	0.106
223		0.100	1.400	ND	0.033
224		0.206	1.270	ND	0.037
225		0.385	2.180	ND	0.129
226		0.274	1.050	ND	0.036
227		0.170	0.717	ND	0.030
228		1.410	1.510	ND	0.112

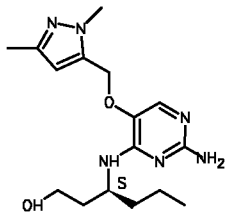
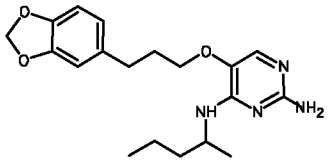
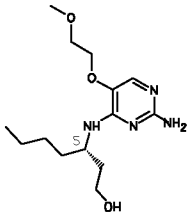
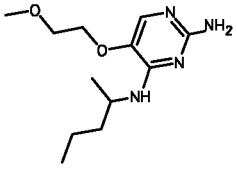
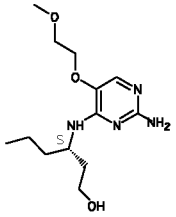
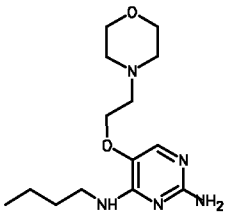
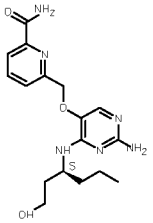
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
229		>25	8.850	ND	7.790
230		0.593	3.470	ND	0.138
231		0.001	0.052	ND	<0.001
232		0.967	1.680	ND	0.110
233		10.070	0.662	ND	0.503
234		1.850	1.170	ND	0.464
235		>25	10.410	ND	1.890

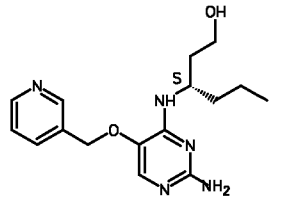
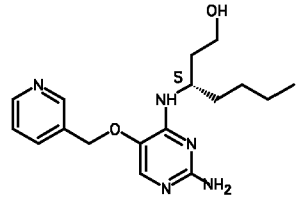
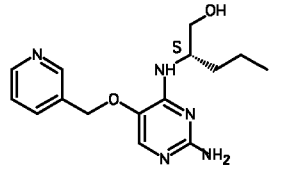
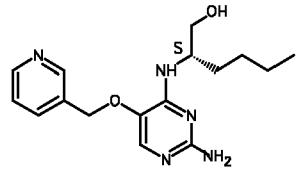
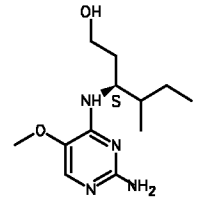
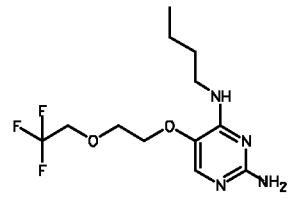
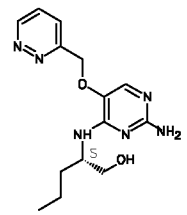
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
236		0.241	0.333	ND	0.031
237		0.156	1.830	ND	0.051
238		0.234	1.920	ND	0.091
239		0.464	0.247	ND	0.145
240		0.008	0.442	ND	0.005
241		0.008	0.304	ND	0.004
242		5.400	3.010	ND	0.006

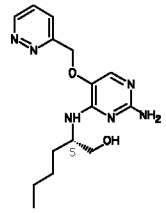
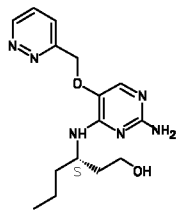
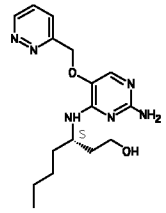
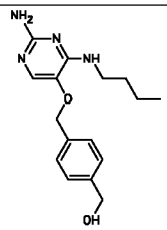
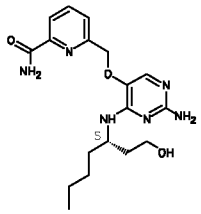
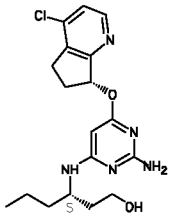
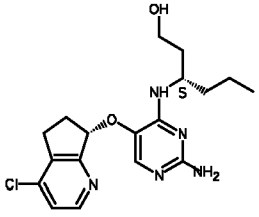
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
243		0.343	0.103	ND	0.190
244		0.202	1.400	ND	0.104
245		0.040	0.507	ND	0.011
246		0.157	1.150	ND	0.048
247		12.390	8.240	ND	3.200
248		2.120	0.654	ND	0.529
249		0.039	0.172	ND	0.036

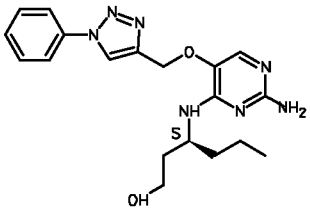
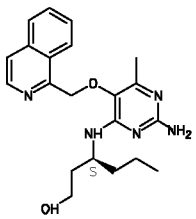
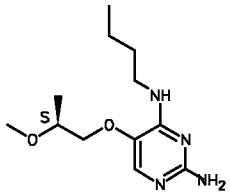
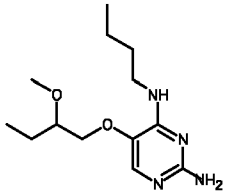
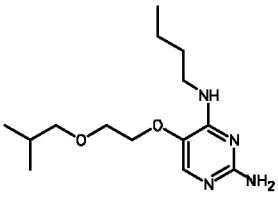
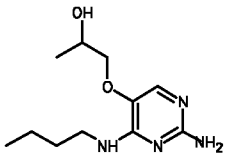
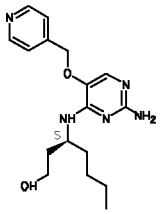
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
250		0.724	3.250	ND	0.580
251		0.164	0.556	ND	0.087
252		4.020	>25	ND	1.310
253		12.760	>25	ND	6.230
254		1.770	0.467	ND	0.364
255		0.552	0.515	ND	0.315
256		1.630	0.100	ND	0.039

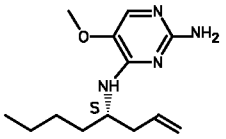
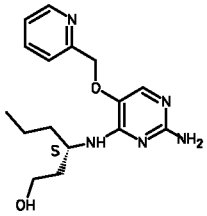
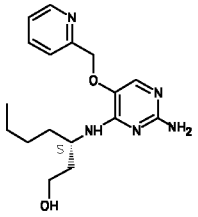
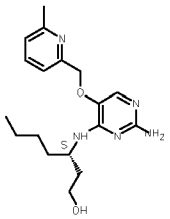
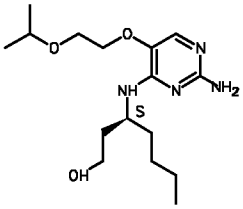
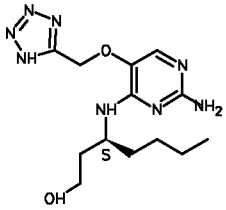
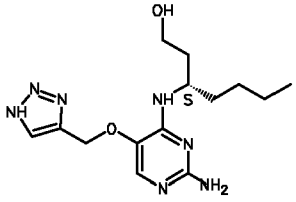
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
257		0.697	0.444	ND	0.304
258		0.074	0.153	ND	0.060
259		6.980	3.150	ND	1.250
260		4.850	2.830	ND	>8.14
261		10.790	2.300	ND	7.460
262		0.658	0.168	ND	0.363
263		0.049	0.159	ND	0.056

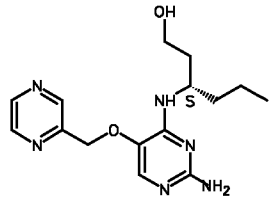
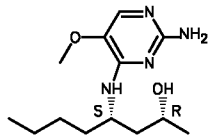
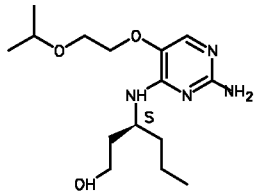
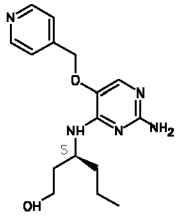
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
264		0.215	0.489	ND	0.087
265		0.752	3.700	ND	0.591
266		1.070	1.890	ND	0.557
267		4.880	0.719	ND	1.710
268		2.100	1.800	ND	1.170
269		24.550	8.280	ND	9.750
270		0.109	0.131	ND	0.006

	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
271		0.261	0.511	ND	0.088
272		0.207	1.110	ND	0.084
273		1.100	0.516	ND	0.552
274		1.140	0.303	ND	0.357
275		10.380	2.600	ND	2.230
276		3.260	0.707	ND	0.652
277		1.180	0.438	ND	0.135

	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
278		0.880	0.270	ND	0.136
279		0.724	1.100	ND	0.034
280		0.392	1.450	ND	0.038
281		0.472	1.630	ND	0.191
282		0.133	0.502	ND	0.005
283		17.180	8.320	ND	4.350
284		9.910	10.300	ND	2.150

	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
285		0.266	0.530	ND	0.036
286		0.620	7.660	ND	0.128
287		15.530	5.380	ND	3.560
288		>25	10.200	ND	14.890
289		0.875	0.930	ND	0.647
290		>25	9.110	ND	12.460
291		0.120	0.407	ND	0.106

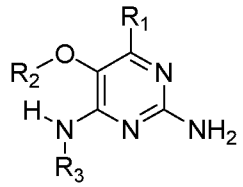
	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
292		0.913	3.082	ND	ND
293		0.197	0.530	ND	0.088
294		0.133	0.521	ND	0.042
295		0.047	0.430	ND	0.034
296		0.664	2.540	ND	0.310
297		2.810	>25	ND	2.540
298		0.394	2.840	ND	0.058

	Structure	TLR 7 wt LEC	TLR 8 wt LEC	PBMC HUH-7 EC_{50}	HEK-ISREluc LEC
299		0.211	0.433	ND	0.159
300		1.740	0.720	ND	0.475
301		1.170	2.640	ND	0.534
302		0.385	0.487	ND	0.070

ND = Not done.

Claims

1. A compound of formula (I)



or a pharmaceutically acceptable salt, tautomer(s), solvate or polymorph thereof, wherein

R_1 is hydrogen, C_{1-4}alkyl, cyclopropyl or C_{1-6}alkoxy, halogen, hydroxyl, trifluoromethyl.

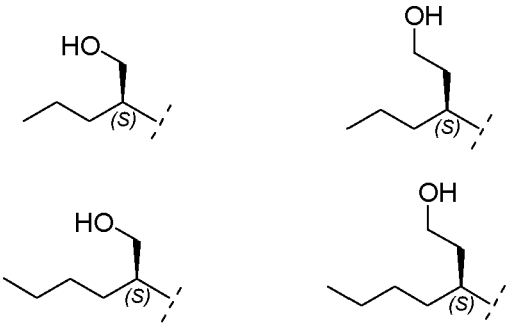
R_2 is C_{1-8}alkyl, (C_{1-4})alkoxy-(C_{1-4})alkyl, C_{3-7}cycloalkyl, C_{4-7}heterocycle, aromatic, bicyclic heterocycle, arylalkyl, heteroaryl, heteroarylalkyl each of which is optionally substituted by one or more substituents independently selected from halogen, hydroxyl, amino, C_{1-6}alkyl, di-(C_{1-6})alkylamino, C_{1-6}alkylamino, C_{1-6}alkyl, C_{1-6}alkoxy, C_{3-6}cycloalkyl, carboxylic acid, carboxylic ester, carboxylic amide, heterocycle, aryl, alkenyl, alkynyl, arylalkyl, heteroaryl, heteroarylalkyl, nitrile.

R_3 is C_{4-8}alkyl, C_{4-8}alkoxy, C_{2-6}alkenyl or C_{2-6} alkynyl, each of which is optionally substituted by one or more substituents independently selected from halogen, hydroxyl, amino, C_{1-3}alkyl, C_{1-3}alkoxy or C_{3-6} cycloalkyl, nitrile.

2. A compound of formula (I) according to claim 1 wherein R_3 is butyl or pentyl and wherein R_2 and R_1 are as specified above.

3. A compound of formula (I) according to claim 1 wherein R_3 is C_{4-8}alkyl substituted with hydroxyl, and wherein R_2 and R_1 are as specified above.

4. A compound of formula (I) according to claim 3 wherein R_3, when being C_{4-8}alkyl substituted with hydroxyl, is one of the following

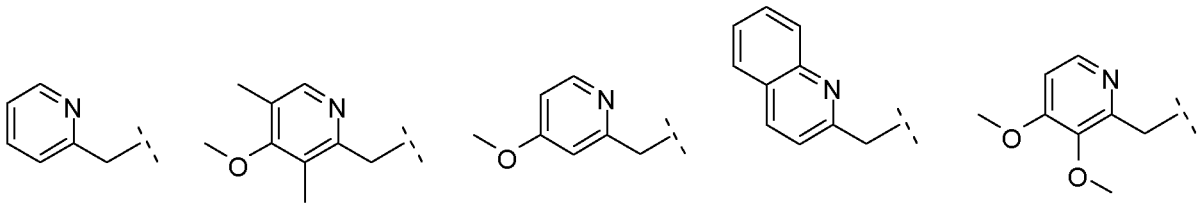


5. A compound of formula (I) according to claim 1 wherein R_1 is hydrogen or -CH_3 and wherein R_2 and R_3 are as specified above.

6. A compound of formula (I) according to claim 1 wherein wherein R_2 is arylalkyl or heteroarylalkyl, substituted with C_{1-3}alkyl, hydroxyl, alkoxy, nitrile, heterocycle or ester and wherein R_1 and R_3 are as specified above.

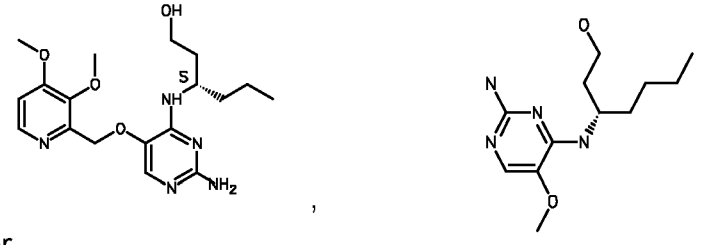
7. A compound of formula (I) according to claim 1 wherein R_2 is C_{1-3}alkyl substituted by aryl, heterocycle, or heteroaryl which is further substituted by C_{1-3}alkyl, alkoxy, carboxylic ester or carboxylic amide and wherein R_1 and R_3 are as specified above.

8. A compound of formula (I) according to claim 1 wherein R_2 is

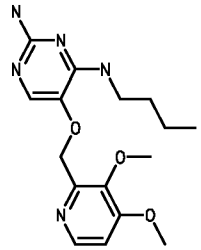


each of which can be optionally further substituted with C_{1-6}alkyl, hydroxyl, C_{1-6}alkoxy, nitrile,heterocycle or ester and wherein R_1 and R_3 are as specified above.

9. A compound according to claim 1 having the formula:



or



10. A pharmaceutical composition comprising a compound of formula (I) or a pharmaceutically acceptable salt, tautomer(s), solvate or polymorph thereof according to one of claims 1-9 together with one or more pharmaceutically acceptable excipients, diluents or carriers.

11. A compound of formula (I) or a pharmaceutically acceptable salt, tautomer(s), solvate or polymorph thereof according to one of claims 1-9, or a pharmaceutical composition according to claim 10 for use as a medicament.

12. A compound of formula (I) or a pharmaceutically acceptable salt, tautomer(s), solvate or polymorph thereof according to one of claims 1-9, or a pharmaceutical composition according to claim 10 for use in the treatment of a disorder or disease in which the modulation of TLR7 and /or TLR8 is involved.