COMPOUNDS FOR TREATING HUNTINGTON'S DISEASE

FIELD OF THE DISCLOSURE

An aspect of the present description relates to compounds, forms, and pharmaceutical compositions thereof and methods of using such compounds, forms, or compositions thereof useful for treating or ameliorating Huntington's disease. In particular, another aspect of the present description relates to substituted monocyclic and bicyclic heteroaryl compounds, forms and pharmaceutical compositions thereof and methods of using such compounds, forms, or compositions thereof for treating or ameliorating Huntington's disease.

BACKGROUND

Huntington's disease (HD) is a progressive, autosomal dominant neurodegenerative disorder of the brain, having symptoms characterized by involuntary movements, cognitive impairment, and mental deterioration. Death, typically caused by pneumonia or coronary artery disease, usually occurs 13 to 15 years after the onset of symptoms. The prevalence of HD is between three and seven individuals per 100,000 in populations of western European descent. In North America, an estimated 30,000 people have HD, while an additional 200,000 people are at risk of inheriting the disease from an affected parent. The disease is caused by an expansion of uninterrupted trinucleotide CAG repeats in the "mutant" huntingtin (Htt) gene, leading to production of HTT (Htt protein) with an expanded poly-glutamine (polyQ) stretch, also known as a "CAG repeat" sequence. There are no current small molecule therapies targeting the underlying cause of the disease, leaving a high unmet need for medications that can be used for treating or ameliorating HD. Consequently, there remains a need to identify and provide small molecule compounds for treating or ameliorating HD.

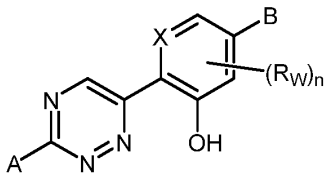
International Application Publication No. WO/2019/191229 A1 describes triazine compounds which may be suitable for treating or ameliorating Huntington's disease.

However, there remains a need to develop new compounds which are more potent, as measured by IC_{50}, and have improved pharmacokinetic properties (PK), as measured by such parameters as efflux transport, tissue exposure, absorption, distribution, metabolism, excretion, time to maximum plasma concentration (T_{max}), maximum concentration (C_{max}), concentration at 24 hours (C_{24}), area under the concentration-time curve (AUC), and terminal elimination half-life ($t_{1/2}$).

All other documents referred to herein are incorporated by reference into the present application as though fully set forth herein.

SUMMARY

An aspect of the present description includes compounds of Formula (I):



(I)

or a form thereof, wherein A, B, X, R_W, and n are as defined herein. B may be a monocyclic ring structure, and/or a bicyclic ring structure.

An aspect of the present description includes a method for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of a compound of Formula (I) or a form thereof.

An aspect of the present description includes a method for use of a compound of Formula (I) or a form or composition thereof for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the compound of Formula (I) or a form or composition thereof.

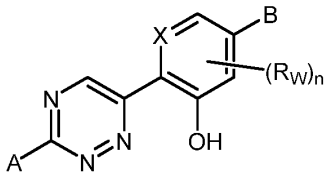
An aspect of the present description includes a use for a compound of Formula (I) or a form thereof for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the compound of Formula (I) or a form thereof.

An aspect of the present description includes a use for a compound of Formula (I) or a form thereof in the manufacture of a medicament for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the medicament.

An aspect of the present description includes a use for a compound of Formula (I) or a form thereof in a combination product with one or more therapeutic agents for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the compound of Formula (I) or a form thereof in combination with an effective amount of the one or more agents.

DETAILED DESCRIPTION

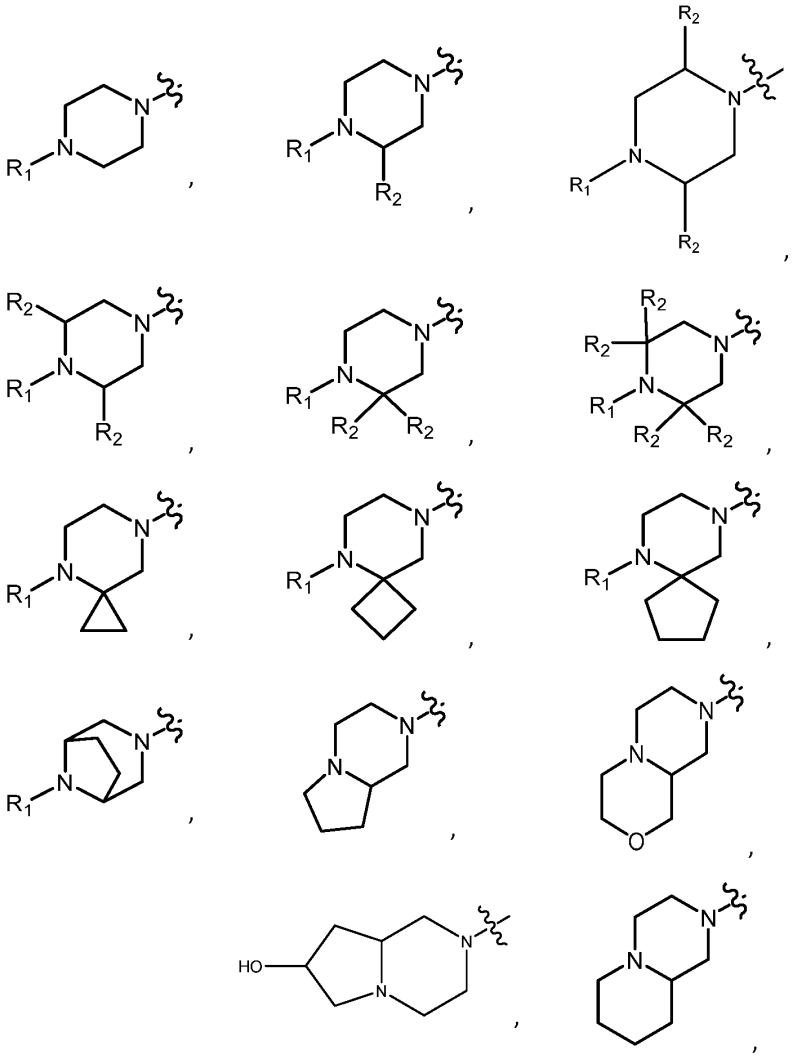
A compound of Formula (I) is disclosed:



(I)

or a salt, solvate, hydrate, ester, prodrug, enantiomer, stereoisomer, rotamer, tautomer, positional isomer, or racemate thereof, wherein:

A is selected from the group consisting of:



and any stereoisomer thereof;

R_1 is selected from the group consisting of hydrogen, C_{1-4}alkyl, and C_{3-6}cycloalkyl;

R_2 is independently selected from the group consisting of halogen, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, C_{1-4}alkoxy-C_{1-4}alkyl, C_{2-4}alkenyl, C_{3-6}cycloalkyl, phenyl, pyridinyl, and hetercyclyl, wherein heterocyclyl is a 3- to 6- membered carbon atom ring structure radical containing 1 or 2 heteroatom ring members selected from N, O, and S, and

wherein each instance of C_{1-4}alkyl, C_{3-6}cycloalkyl, phenyl, pyridnyl, and heterocyclyl is optionally substituted with one or two R_3 substituents;

R_3 is independently selected from the group consisting of halogen, hydroxyl, C_{1-4}alkyl, C_{1-4}alkoxy, and C_{3-6}cycloalkyl;

B is selected from the group consisting of:

phenyl optionally substituted with one or two independently selected R_4 substituents;

heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 heteroatoms selected from N, O, and S, optionally substituted with one R_4 substituent, or wherein heteroaryl is a 9- or 10- membered bicyclic aromatic carbon atom ring structure radical having 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S, optionally substituted with one or two independently selected R_4 substituents; and

heterocyclyl, wherein heterocyclyl is a 8- to 10- membered bicyclic aromatic carbon atom ring structure radical containing 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S, optionally substituted with one or two independently selected R_4 substituents;

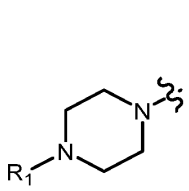
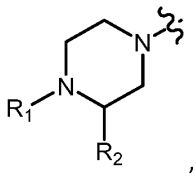
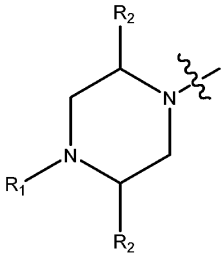
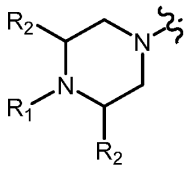
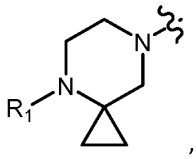
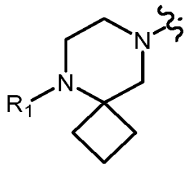
R_4 is selected from the group consisting of halogen, cyano, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, C_{1-4}alkoxy, deutero-C_{1-4}alkoxy, amino, C_{1-4}alkyl-amino, $(C_{1-4}alkyl)_2$-amino, C_{3-6}cycloalkyl, and heterocyclyl, wherein heterocyclyl is a 3- to 6- membered monocyclic carbon atom ring structure radical containing 1 or 2 heteroatom ring members independently selected from N, O, or S;

X is selected from the group consisting of CH, CF, and N;

R_w is selected from the group consisting of halogen, hydroxyl, cyano, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, amino, C_{1-4}alkyl-amino, $(C_{1-4}alkyl)_2$-amino, C_{1-4}alkoxy, and halo-C_{1-4}alkoxy; and

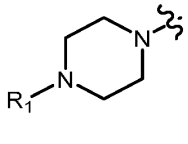
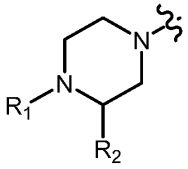
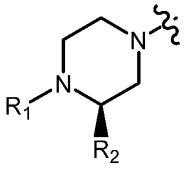
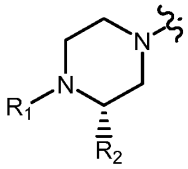
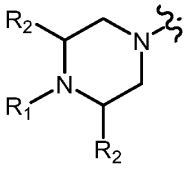
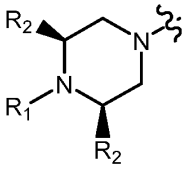
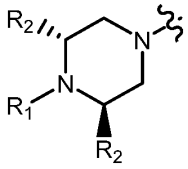
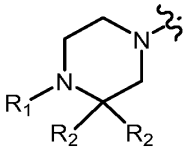
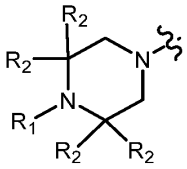
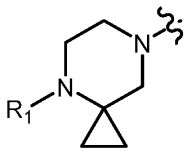
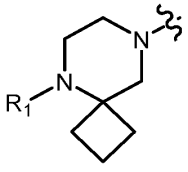
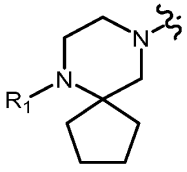
n is selected from the group consisting of 0 or 1.

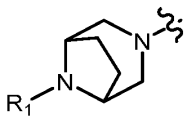
One aspect includes a compound of Formula (I), wherein **A** is selected from the group consisting of:


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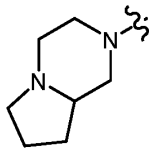
and any stereoisomer thereof.

A may be selected from the group consisting of:

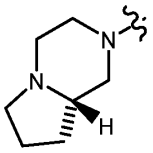

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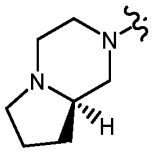
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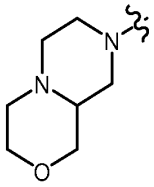
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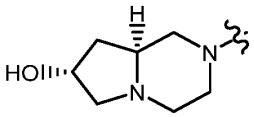
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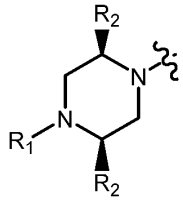
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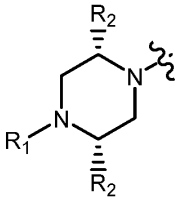
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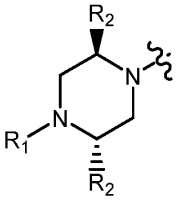
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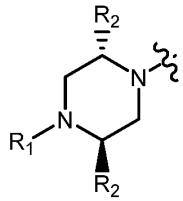
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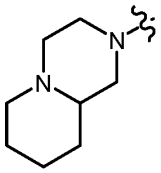
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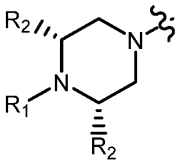
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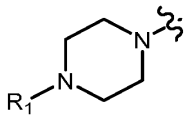


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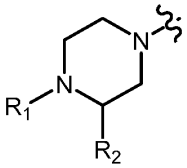


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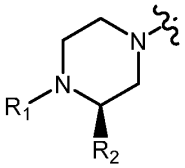
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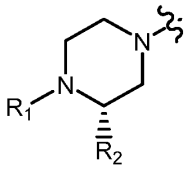
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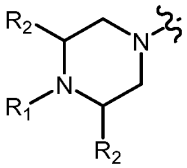
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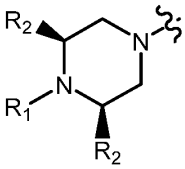
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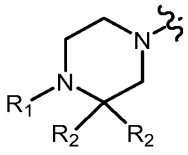
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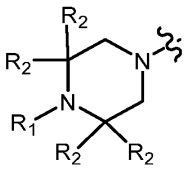
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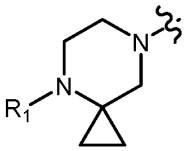
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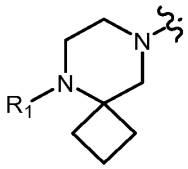
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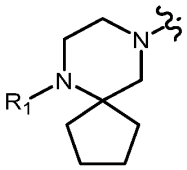
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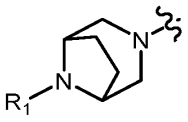
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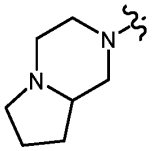
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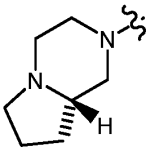
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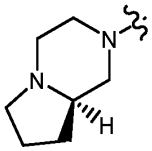
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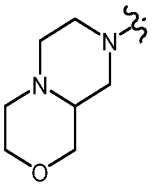
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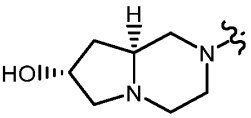
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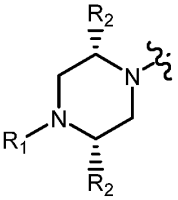
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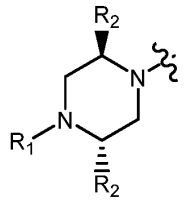
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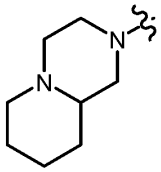
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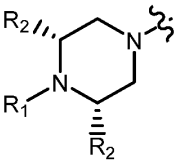
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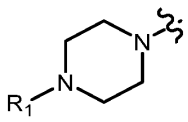


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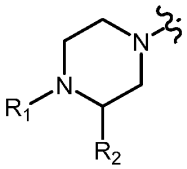


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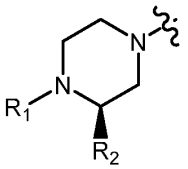
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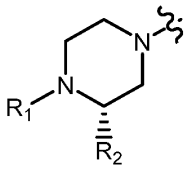
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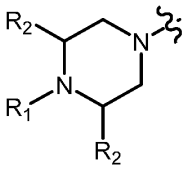
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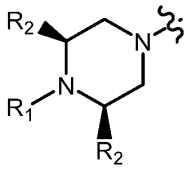
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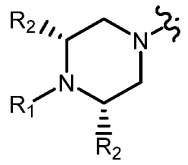
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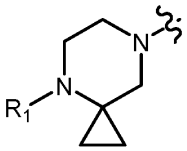
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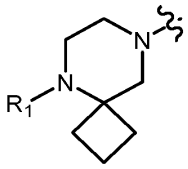
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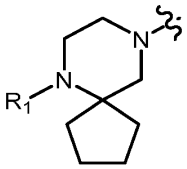
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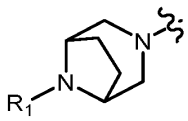
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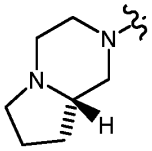
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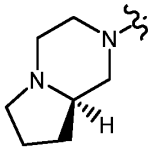
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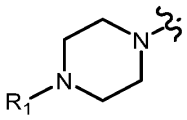


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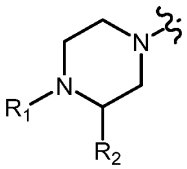
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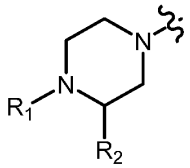


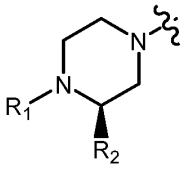
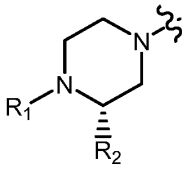
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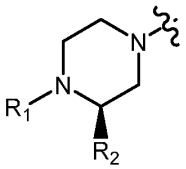


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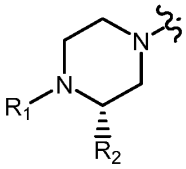
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A may be



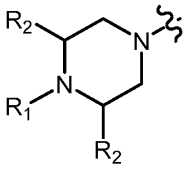
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A may be

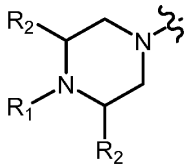


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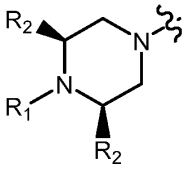
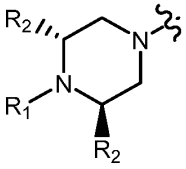
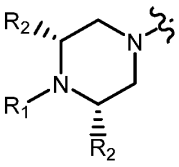
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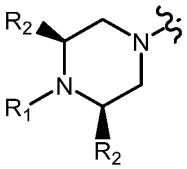
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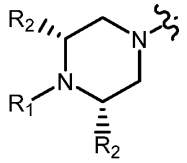

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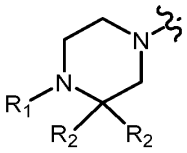
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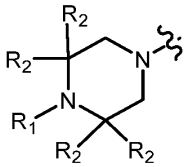
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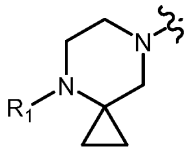
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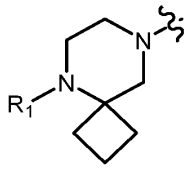
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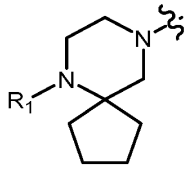
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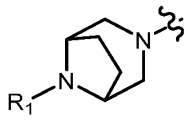
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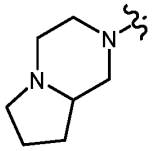
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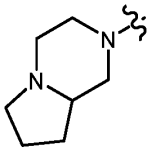


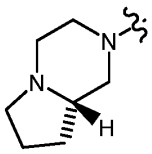
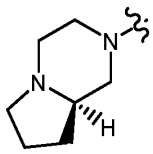
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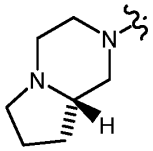


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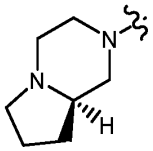
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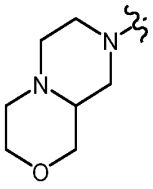
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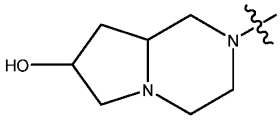


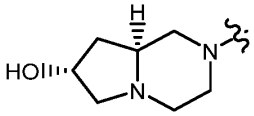
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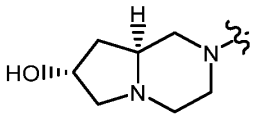


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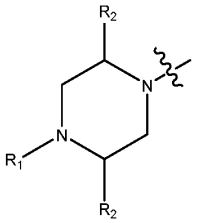
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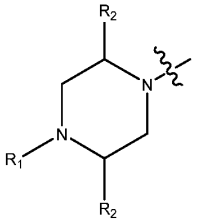
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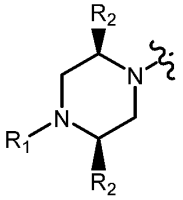
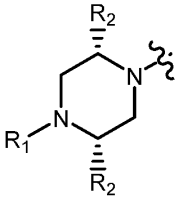
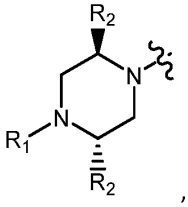
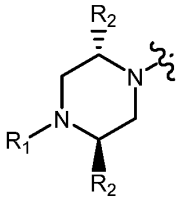
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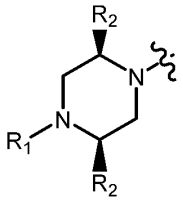
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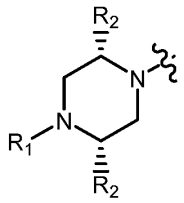
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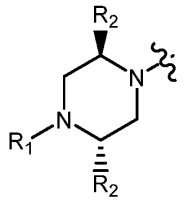
A19.

A may be



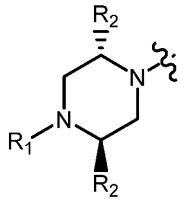
A20.

A may be



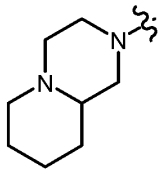
A21.

A may be



A22.

A may be



A23, or any stereoisomer thereof.

One aspect includes a compound of Formula (I), wherein R_1 is hydrogen or C_{1-4}alkyl. R_1 may be hydrogen. R_1 may be C_{1-4}alkyl selected from methyl, ethyl, propyl, isopropyl, butyl, and tert-butyl. R_1 may be methyl or ethyl. R_1 may be methyl. R_1 may be ethyl. One aspect includes a compound of Formula (I), wherein R_1 is C_{3-6}cycloalkyl selected from cyclopropyl, cyclobutyl, cyclopenyl, and cyclohexyl. R_1 may be cyclopropyl.

One aspect includes a compound of Formula (I), wherein R_2 is independently selected from the group consisting of C_{1-4}alkyl, halo-C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, C_{1-4}alkoxy-

C_{1-4}alkyl, C_{2-4}alkenyl, C_{3-6}cycloalkyl, phenyl, pyridinyl, and heterocyclyl, wherein heterocyclyl is a 3- to 6- membered carbon atom ring structure radical containing 1 or 2 heteroatom ring members selected from N, O, and S, and wherein each instance of C_{1-4}alkyl, C_{3-6}cycloalkyl, phenyl, pyridnyl, and heterocyclyl is optionally substituted with one or two R_3 substituents.

Another aspect includes a compound of Formula (I), wherein R_2 is independently selected from the group consisting of C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, C_{3-6}cycloalkyl, phenyl, pyridinyl, and heterocyclyl, wherein heterocyclyl is a 3- to 4- membered carbon atom ring structure radical containing 1 heteroatom ring members selected from N, and O, wherein each R_2 is optionally substituted with one R_3 substituent.

Another aspect includes a compound of Formula (I), wherein R_2 is independently selected from the group consisting of C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, and C_{3-6}cycloalkyl, wherein each R_2 is optionally substituted with one R_3 substituent.

R_2 may be C_{1-4}alkyl selected from the group consisting of methyl, ethyl, propyl, isopropyl, butyl and tert-butyl. R_2 may be methyl. R_2 may be ethyl. R_2 may be propyl. R_2 may be isopropyl. R_2 may be butyl. R_2 may be tert-butyl.

R_2 may be hydroxyl-C_{1-4}alkyl, wherein C_{1-4}alkyl is selected from the group consisting of methyl, ethyl, propyl, isopropyl, and butyl, partially or completely substituted with one or more hydroxyl groups where allowed by available valences. R_2 may be hydroxyl-C_{1-4}alkyl, wherein C_{1-4}alkyl is selected from methyl and isopropyl susbstitured with one hydroxyl group. R_2 may be hydroxymethyl. R_2 may be isopropyl substituted with one hydroxyl group. R_2 may be 2-hydroxypropan-2-yl.

R_2 may be halo-C_{1-4}alkyl wherein C_{1-4}alkyl is selected from the group consisting of methyl, ethyl, propyl, isopropyl, butyl, and tert-butyl, partially or completely substituted with one or more halogen atoms where allowed by available valences. R_2 may be difluoroethyl.

R_2 may be C_{1-4}alkoxy-C_{1-4}alkyl, wherein C_{1-4}alkyl is selected from the group consisting of methyl, ethyl, propyl, and butyl, partially or completely substituted with one or more C_{1-4}alkoxy groups selected from methoxy, ethoxy, propoxy, and butoxy where allowed by available valences. R_2 may be methoxymethyl.

R_2 may be C_{2-4}alkenyl selected from ethenyl, propenyl, and butenyl. R_2 may be ethenyl.

R_2 may be C_{3-6}cycloalkyl selected from cyclopropyl, cyclobutyl, cyclopenyl, and cyclohexyl optionally substituted with one or two R_3 substituents. R_2 may be C_{3-6}cycloalkyl selected from cyclopropyl and cyclobutyl optionally substituted with one or two R_3 substituents.

R_2 may be cyclopropyl, substituted with zero to two R_3 substituents. R_2 may be unsubstituted cyclopropyl. R_2 may be cyclopropyl unsubstituted or substituted with one R_3 substituent, wherein R_3 is halogen, hydroxyl, C_{1-4}alkyl, or C_{1-4}alkoxy. R_2 may be cyclopropyl unsubstituted or substituted with one R_3 substituent, wherein R_3 is halogen, hydroxyl, methyl, ethyl, methoxy, or ethoxy.

R_2 may be cyclobutyl optionally substituted with one or two R_3 substituents. R_2 may be unsubstitued cyclobutyl.

R_2 may be phenyl optionally substituted with one or two R_3 substituents. R_2 may be unsubstitued phenyl.

R_2 may be pyridinyl optionally substituted with one or two R_3 substituents. R_2 may be unsubstitued pyridinyl. R_2 may be unsubstitued pyridine-4-yl.

R_2 may be C_{1-4}alkyl, halo-C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, cyclopropyl, cyclobutyl, phenyl, or oxetanyl, optionally substituted with one or two R_3 substituents.

R_2 may be heterocyclyl selected from the group consisting of aziridinyl, oziranyl, thiiranyl, azetidinyl, oxetanyl, thietanyl, pyrrolidinyl, pyrazolinyl, tetrahyrofuranyl, thilolanyl, piperidinyl, piperazinyl, tetrahydro-2H-pyranyl, 1,4-dioxanyl, morpholinyl, and thianyl, wherein each instance of heterocyclyl is optionally substituted with one or two R_3 substituents.

R_2 may be oxetanyl optionally substituted with one or two R_3 substituents.

R_2 may be C_{1-4}alkyl, halo-C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, cyclopropyl, cyclobutyl, phenyl, or oxetanyl, optionally substituted with one or two R_3 substituents. R_2 may be unsubstituted C_{1-4}alkyl, unsubstituted halo-C_{1-4}alkyl, unsubstituted hydroxyl-C_{1-4}alkyl, unsubstituted cyclopropyl, unsubstituted cyclobutyl, unsubstituted phenyl, or unsubstituted oxetanyl.

One aspect includes a compound of Formula (I), wherein R_3 is independently selected from the group consisting of halogen, hydroxyl, C_{1-4}alkyl, C_{1-4}alkoxy, and C_{3-6}cycloalkyl. R_3 may be independently selected from the group consisting of halogen, hydroxyl, C_{1-4}alkyl, and C_{1-4}alkoxy.

R_3 may be halogen selected from the group consisting of bromo, chloro, fluoro, and iodo. R_3 may be fluoro.

R_3 may be hydroxyl.

R_3 may be C_{1-4}alkyl selected from the group consisting of methyl, ethyl, propyl, isopropyl, butyl, and tert-butyl. R_3 may be methyl or ethyl. R_3 may be methyl.

R_3 may be C_{1-4}alkoxy selected from the group consisting of methoxy, ethoxy, propoxy, isopropoxy, butoxy and tert-butoxy. R_3 may be methoxy. R_3 may be halogen, hydroxyl, C_{1-4}alkyl, or methoxy.

R_3 may be C_{3-6}cycloalkyl selected from the group consisting of cyclopropyl, cyclobutyl, cyclopenyl, and cyclohexyl. R_3 may be cyclopropyl.

One aspect includes a compound of Formula (I), wherein B is selected from the group consisting of:

phenyl optionally substituted with one or two independently selected R_4 substituents;

heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 heteroatoms selected from N, O, and S, optionally substituted with one R_4 substituent, or wherein heteroaryl is a 9- or 10- membered aromatic carbon atom ring structure radical having 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S, optionally substituted with one or two independently selected R_4 substituents; and

heterocyclyl, wherein heterocyclyl is a 8- to 10- membered bicyclic carbon atom ring structure radical containing 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S, optionally substituted with one or two independently selected R_4 substituents.

B may be phenyl optionally substituted with one or two independently selected R_4 substituents. B may be unsubstituted phenyl or phenyl substituted with one R_4 substituent.

One aspect includes a compound of Formula (I), wherein B is heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 heteroatoms selected from N, O, and S, optionally substituted with one R_4 substituent. B may be heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 N atoms, and optionally a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent. B may be heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 2 or 3 heteroatoms selected from N, O, and S, optionally substituted with one R_4 substituent. B may be heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 2 or 3 N atoms, and optionally a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of furanyl, thiophenyl, 1H-pyrazolyl, 1H-imidazolyl, isoxazolyl, 1,3-thiazolyl, 1,3-oxazolyl, tetrazolyl, 1H-1,2,3-triazolyl, 2H-1,2,3-triazolyl, 1,2,4-oxadiazolyl, 1,3,4-oxadiazolyl, 1,2,3-thiadiazolyl,

pyridinyl, pyrimidinyl, pyrazinyl, pyridazinyl, 1H-1,2,4-triazolyl, 1,2,4-thiadiazolyl, 1,3,4-oxadiazolyl, isothiazolyl, 1,2,5-thiadiazolyl, and 1,3,4-thiadiazolyl, optionally substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-pyrazolyl, 1H-imidazolyl, 1,3-thiazolyl, 1,3-oxazolyl, 1H-1,2,3-triazolyl, 2H-1,2,3-triazolyl, pyridinyl, pyrimidinyl, pyrazinyl, 1H-1,2,4-triazolyl, 1,2,4-thiadiazolyl, 1,3,4-oxadiazolyl, isothiazolyl, 1,2,5-thiadiazolyl, and 1,3,4-thiadiazolyl, optionally substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-pyrazolyl, 1H-imidazolyl, 1,3-thiazolyl, 1,3-oxazolyl, 1H-1,2,3-triazolyl, 2H-1,2,3-triazolyl, pyridinyl, pyrimidinyl, pyrazinyl, 1H-1,2,4-triazolyl, 1,2,4-thiadiazolyl, 1,3,4-oxadiazolyl, isothiazolyl, 1,2,5-thiadiazolyl, and 1,3,4-thiadiazolyl, substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of furan-2-yl, furan-3-yl, thiophen-2-yl, thiophen-3-yl, 1H-pyrazol-1-yl, 1H-pyrazol-3-yl, 1H-pyrazol-4-yl, 1H-pyrazol-5-yl, 1H-imidazol-1-yl, 1H-imidazol-4-yl, isoxazol-3-yl, isoxazol-4-yl, isoxazol-5-yl, 1,3-thiazol-2-yl, 1,3-thiazol-4-yl, 1,3-thiazol-5-yl, 1,3-oxazol-2-yl, 1,3-oxazol-4-yl, 1,3-oxazol-5-yl, tetrazol-5-yl, 1H-1,2,3-triazol-1-yl, 1H-1,2,3-triazol-4-yl, 1H-1,2,3-triazol-5-yl, 2H-1,2,3-triazol-2-yl, 2H-1,2,3-triazol-4-yl, 2H-1,2,3-triazol-5-yl, 1,2,4-oxadiazol-3-yl, 1,3,4-oxadiazol-2-yl, 1,2,3-thiadiazol-4-yl, 1,2,3-thiadiazol-5-yl, pyridin-2-yl, pyridin-3-yl, pyridin-4-yl, pyrimidin-4-yl, pyrimidin-2-yl, pyrazin-2-yl, pyridazin-3-yl, pyridazin-4-yl, 1H-1,2,4-triazol-3-yl, 1H-1,2,4-triazol-5-yl, 1,2,4-thiadiazol-3-yl, 1,2,4-thiadiazol-5-yl, 1,3,4-oxadiazol-2-yl, isothiazol-3-yl, isothiazol-4-yl, isothiazol-5-yl, 1,2,5-thiadiazol-3-yl, 1,3,4-thiadiazol-2-yl, optionally substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-pyrazol-4-yl, 1H-imidazol-1-yl, 1,3-thiazol-2-yl, 1,3-thiazol-4-yl, 1,3-oxazol-2-yl, 1H-1,2,3-triazol-4-yl, 1H-1,2,3-triazol-5-yl, 2H-1,2,3-triazol-2-yl, 2H-1,2,3-triazol-4-yl, pyridin-3-yl, pyridin-4-yl, pyrimidin-4-yl, pyrimidin-2-yl, pyrazin-2-yl, 1H-1,2,4-triazol-3-yl, 1,2,4-thiadiazol-3-yl, 1,2,4-thiadiazol-5-yl, 1,3,4-oxadiazol-2-yl, isothiazol-3-yl, isothiazol-4-yl, isothiazol-5-yl, 1,2,5-thiadiazol-3-yl, 1,3,4-thiadiazol-2-yl, optionally substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-pyrazol-4-yl, 1H-imidazol-1-yl, 1,3-thiazol-2-yl, 1,3-thiazol-4-yl, 1,3-oxazol-2-yl, 1H-1,2,3-triazol-4-yl, 1H-1,2,3-triazol-5-yl, 2H-1,2,3-triazol-2-yl, 2H-1,2,3-triazol-4-yl, pyridin-3-yl, pyridin-4-yl, pyrimidin-4-yl, pyrimidin-2-yl, pyrazin-2-yl, 1H-1,2,4-triazol-3-yl, 1,2,4-thiadiazol-3-yl, 1,2,4-thiadiazol-5-yl, 1,3,4-oxadiazol-2-yl, isothiazol-3-yl, isothiazol-4-yl, isothiazol-5-yl, 1,2,5-thiadiazol-3-yl, 1,3,4-thiadiazol-2-yl, substituted with one R_4 substituent.

One aspect includes a compound of Formula (I), wherein B is heteroaryl, wherein heteroaryl is a 9- or 10- membered aromatic carbon atom ring structure radical having 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S, optionally substituted with one or two independently selected R_4 substituents. B may be heteroaryl, wherein heteroaryl is a 9- or 10- membered bicyclic carbon atom ring structure radical containing at least one N atom ring member, optionally containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents. B may be heteroaryl, wherein heteroaryl is a 9- or 10- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N atoms, optionally containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents. B may be heteroaryl, wherein heteroaryl is a 9-membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally substituted with one or two independently selected R_4 substituents. B may be heteroaryl, wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents. B may be heteroaryl, wherein heteroaryl is a unsubstituted 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally containing a second heteroatom ring member selected from O or S. B may be heteroaryl, wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally containing a second heteroatom ring member selected from O or S, and is unsubstituted or substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-indazolyl, 2H-indazolyl, indolizinyl, benzofuranyl, benzothiophenyl, 1H-benzimidazolyl, 1,3-benzoxazolyl, 1,3-benzothiazolyl, 1,3-benzodioxolyl, 1,2,3-benzotriazolyl, 7H-purinyl, furo[3,2-b]pyridinyl, furo[3,2-c]pyridinyl, 1,3-oxazolo[5,4-b]pyridinyl, thieno[3,2-c]pyridinyl, thieno[2,3-d]pyrimidinyl, pyrrolo[1,2-a]pyrimidinyl, pyrrolo[1,2-a]pyrazinyl, pyrrolo[1,2-b]pyridazinyl, pyrazolo[1,5-a]pyridinyl, 1H-pyrrolo[2,3-b]pyridinyl, 5H-pyrrolo[2,3-b]pyrazinyl, 1H-pyrrolo[2,3-c]pyridinyl, 1H-pyrazolo[3,4-b]pyridinyl, 2H-pyrazolo[3,4-b]pyridinyl, 1H-pyrazolo[3,4-b]pyrazinyl, 1H-pyrazolo[3,4-c]pyridinyl, 1H-pyrazolo[4,3-b]pyridinyl, 2H-pyrazolo[4,3-b]pyridinyl, 2H-pyrazolo[4,3-c]pyridinyl, 1H-pyrazolo[4,3-d]pyrimidinyl, pyrazolo[1,5-a]pyrazinyl, imidazo[1,2-a]pyridinyl, imidazo[1,2-a]pyrimidinyl, imidazo[1,2-a]pyrazinyl, imidazo[1,2-b]pyridazinyl, imidazo[1,2-c]pyrimidin-2-yl, 1H-imidazo[4,5-b]pyridinyl, 3H-imidazo[4,5-b]pyridinyl,

imidazo[2,1-b][1,3]thiazolyl, imidazo[2,1-b][1,3,4]thiadiazolyl, [1,3]oxazolo[4,5-b]pyridinyl, [1,2,3]triazolo[1,5-a]pyridinyl, 1H-[1,2,3]triazolo[4,5-b]pyridinyl, 3H-[1,2,3]triazolo[4,5-b]pyridinyl, [1,2,4]triazolo[1,5-a]pyridinyl, [1,2,4]triazolo[1,5-a]pyrimidinyl, [1,2,4]triazolo[1,5-a]pyrazinyl, [1,2,4]triazolo[1,5-b]pyridazinyl, [1,2,4]triazolo[4,3-a]pyridinyl, tetrazolo[1,5-a]pyridinyl, tetrazolo[1,5-b]pyridazinyl, thiazolo[4,5-b]pyrazinyl, thiazolo[5,4-c]pyridinyl, quinolinyl, isoquinolinyl, [1,2,5]thiadiazolo[3,4-b]pyridinyl, 1H-pyrazolo[3,4-d]pyrimidinyl, 1H-pyrrolo[3,2-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-c]pyridinyl, 3H-[1,2,3]triazolo[4,5-b]pyridinyl, 3H-[1,2,3]triazolo[4,5-c]pyridinyl, benzo[c][1,2,5]thiadiazolyl, imidazo[1,5-a]pyridinyl, pyrazolo[1,5-a]pyrimidinyl, and thiazolo[5,4-b]pyridinyl, optionally substituted with one or two independently selected R_4 substituents.

B may be heteroaryl selected from the group consisting of 2H-indazolyl, 7H-purinyl, furo[3,2-b]pyridinyl, pyrazolo[1,5-a]pyridinyl, 2H-pyrazolo[3,4-b]pyridinyl, 1H-pyrazolo[4,3-b]pyridinyl, imidazo[1,2-a]pyridinyl, imidazo[1,2-a]pyrazinyl, imidazo[1,2-b]pyridazinyl, [1,2,3]triazolo[1,5-a]pyridinyl, 1H-[1,2,3]triazolo[4,5-b]pyridinyl, [1,2,4]triazolo[1,5-a]pyridinyl, [1,2,4]triazolo[1,5-a]pyrimidinyl, [1,2,4]triazolo[1,5-a]pyrazinyl, [1,2,4]triazolo[1,5-b]pyridazinyl, [1,2,4]triazolo[4,3-a]pyridinyl, thiazolo[4,5-b]pyrazinyl, thiazolo[5,4-c]pyridinyl, [1,2,5]thiadiazolo[3,4-b]pyridinyl, 1H-pyrazolo[3,4-d]pyrimidinyl, 1H-pyrrolo[3,2-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-c]pyridinyl, 3H-[1,2,3]triazolo[4,5-b]pyridinyl, 3H-[1,2,3]triazolo[4,5-c]pyridinyl, benzo[c][1,2,5]thiadiazolyl, imidazo[1,5-a]pyridinyl, pyrazolo[1,5-a]pyrimidinyl, and thiazolo[5,4-b]pyridinyl, optionally substituted with one or two independently selected R_4 substituents.

B may be heteroaryl selected from the group consisting of 2H-indazolyl, 7H-purinyl, furo[3,2-b]pyridinyl, pyrazolo[1,5-a]pyridinyl, 2H-pyrazolo[3,4-b]pyridinyl, 1H-pyrazolo[4,3-b]pyridinyl, imidazo[1,2-a]pyridinyl, imidazo[1,2-a]pyrazinyl, imidazo[1,2-b]pyridazinyl, [1,2,3]triazolo[1,5-a]pyridinyl, 1H-[1,2,3]triazolo[4,5-b]pyridinyl, [1,2,4]triazolo[1,5-a]pyridinyl, [1,2,4]triazolo[1,5-a]pyrimidinyl, [1,2,4]triazolo[1,5-a]pyrazinyl, [1,2,4]triazolo[1,5-b]pyridazinyl, [1,2,4]triazolo[4,3-a]pyridinyl, thiazolo[4,5-b]pyrazinyl, thiazolo[5,4-c]pyridinyl, [1,2,5]thiadiazolo[3,4-b]pyridinyl, 1H-pyrazolo[3,4-d]pyrimidinyl, 1H-pyrrolo[3,2-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-c]pyridinyl, 3H-[1,2,3]triazolo[4,5-b]pyridinyl, 3H-[1,2,3]triazolo[4,5-c]pyridinyl, benzo[c][1,2,5]thiadiazolyl, imidazo[1,5-a]pyridinyl, pyrazolo[1,5-a]pyrimidinyl, and thiazolo[5,4-b]pyridinyl, substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 2H-indazolyl, 7H-purinyl, furo[3,2-b]pyridinyl, pyrazolo[1,5-a]pyridinyl, 2H-pyrazolo[3,4-b]pyridinyl, 1H-pyrazolo[4,3-b]pyridinyl, imidazo[1,2-a]pyridinyl, imidazo[1,2-a]pyrazinyl, imidazo[1,2-b]pyridazinyl, [1,2,3]triazolo[1,5-a]pyridinyl, 1H-[1,2,3]triazolo[4,5-b]pyridinyl, [1,2,4]triazolo[1,5-a]pyridinyl, [1,2,4]triazolo[1,5-a]pyrimidinyl, [1,2,4]triazolo[1,5-a]pyrazinyl, [1,2,4]triazolo[1,5-b]pyridazinyl, [1,2,4]triazolo[4,3-a]pyridinyl, thiazolo[4,5-b]pyrazinyl, thiazolo[5,4-c]pyridinyl, [1,2,5]thiadiazolo[3,4-b]pyridinyl, 1H-pyrazolo[3,4-d]pyrimidinyl, 1H-pyrrolo[3,2-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-c]pyridinyl, 3H-[1,2,3]triazolo[4,5-b]pyridinyl, 3H-[1,2,3]triazolo[4,5-c]pyridinyl, benzo[c][1,2,5]thiadiazolyl, imidazo[1,5-a]pyridinyl, pyrazolo[1,5-a]pyrimidinyl, and thiazolo[5,4-b]pyridinyl, substituted with two independently selected R_4 substituents.

B may be heteroaryl selected from the group consisting of 1H-indazol-5-yl, 2H-indazol-5-yl, indolizin-2-yl,benzofuran-2-yl, benzofuran-5-yl, benzothiophen-2-yl, benzothiophen-3-yl, 1H-benzimidazol-2-yl, 1H-benzimidazol-5-yl, 1H-benzimidazol-6-yl, 1,3-benzoxazol-2-yl, 1,3-benzoxazol-5-yl, 1,3-benzoxazol-6-yl, 1,3-benzothiazol-2-yl, 1,3-benzothiazol-5-yl, 1,3-benzothiazol-6-yl, 1,3-benzodioxol-5-yl, 1,2,3-benzotriazol-5-yl, 7H-purin-2-yl, furo[3,2-b]pyridin-2-yl, furo[3,2-c]pyridin-2-yl, furo[2,3-c]pyridin-2-yl, 1,3-oxazolo[5,4-b]pyridine-5-yl, thieno[3,2-c]pyridin-2-yl, thieno[2,3-d]pyrimidin-6-yl, pyrrolo[1,2-a]pyrimidin-7-yl, pyrrolo[1,2-a]pyrazin-7-yl, pyrrolo[1,2-b]pyridazin-2-yl, pyrazolo[1,5-a]pyridin-2-yl, pyrazolo[1,5-a]pyridin-3-yl, pyrazolo[1,5-a]pyridin-5-yl, 1H-pyrrolo[2,3-b]pyridin-5-yl, 5H-pyrrolo[2,3-b]pyrazin-2-yl,1H-pyrrolo[2,3-c]pyridin-4-yl, 1H-pyrazolo[3,4-b]pyridin-5-yl, 1H-pyrazolo[3,4-b]pyridin-6-yl, 2H-pyrazolo[3,4-b]pyridin-5-yl, 1H-pyrazolo[3,4-b]pyrazin-5-yl, 1H-pyrazolo[3,4-c]pyridin-1-yl, 1H-pyrazolo[3,4-c]pyridin-5-yl, 1H-pyrazolo[4,3-b]pyridin-1-yl, 1H-pyrazolo[4,3-b]pyridin-5-yl, 1H-pyrazolo[4,3-b]pyridin-6-yl, 2H-pyrazolo[4,3-b]pyridin-5-yl, 2H-pyrazolo[4,3-c]pyridin-5-yl,1H-pyrazolo[4,3-d]pyrimidin-5-yl, pyrazolo[1,5-a]pyrazin-2-yl, imidazo[1,2-a]pyridin-2-yl, imidazo[1,2-a]pyridin-6-yl, imidazo[1,2-a]pyrimidin-2-yl, imidazo[1,2-a]pyrimidin-6-yl, imidazo[1,2-a]pyrazin-2-yl, imidazo[1,2-a]pyrazin-6-yl, imidazo[1,2-b]pyridazin-2-yl, imidazo[1,2-b]pyridazin-6-yl, imidazo[1,2-c]pyrimidin-2-yl, 1H-imidazo[4,5-b]pyridin-5-yl, 3H-imidazo[4,5-b]pyridin-5-yl, imidazo[2,1-b][1,3]thiazol-6-yl, imidazo[2,1-b][1,3,4]thiadiazol-6-yl, [1,3]oxazolo[4,5-b]pyridin-2-yl, [1,2,3]triazolo[1,5-a]pyridin-5-yl, [1,2,3]triazolo[1,5-a]pyridin-6-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 3H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, [1,2,4]triazolo[1,5-a]pyridin-6-yl, [1,2,4]triazolo[1,5-

a]pyrimidin-2-yl, [1,2,4]triazolo[1,5-a]pyrimidin-6-yl, [1,2,4]triazolo[1,5-a]pyrazin-6-yl, [1,2,4]triazolo[1,5-b]pyridazin-6-yl, [1,2,4]triazolo[4,3-a]pyridin-6-yl, tetrazolo[1,5-a]pyridin-7-yl, tetrazolo[1,5-b]pyridazin-7-yl, thiazolo[4,5-b]pyrazin-2-yl, thiazolo[5,4-c]pyridin-2-yl, tetrazolo[1,5-a]pyridin-7-yl, tetrazolo[1,5-b]pyridazin-7-yl, quinolin-6-yl, isoquinolin-6-yl, [1,2,5]thiadiazolo[3,4-b]pyridin-6-yl, 1H-pyrazolo[3,4-d]pyrimidin-1-yl, 1H-pyrrolo[3,2-b]pyridin-1-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 3H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, benzo[c][1,2,5]thiadiazol-5-yl, imidazo[1,5-a]pyridin-6-yl, imidazo[1,5-a]pyridin-7-yl, pyrazolo[1,5-a]pyrimidin-3-yl, thiazolo[5,4-b]pyridin-2-yl, optionally substituted with one or two independently selected R_4 substituents.

B may be heteroaryl selected from the group consisting of 1H-indazol-5-yl, , 7H-purin-2-yl, furo[3,2-b]pyridin-2-yl, pyrazolo[1,5-a]pyridin-3-yl, 2H-pyrazolo[3,4-b]pyridin-5-yl, 1H-pyrazolo[4,3-b]pyridin-1-yl, imidazo[1,2-a]pyridin-6-yl, imidazo[1,2-a]pyrazin-6-yl, imidazo[1,2-b]pyridazin-6-yl [1,2,3]triazolo[1,5-a]pyridin-6-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyrimidin-2-yl, [1,2,4]triazolo[1,5-a]pyrimidin-6-yl, [1,2,4]triazolo[1,5-a]pyrazin-6-yl, [1,2,4]triazolo[1,5-b]pyridazin-6-yl, [1,2,4]triazolo[4,3-a]pyridin-6-yl, thiazolo[4,5-b]pyrazin-2-yl, thiazolo[5,4-c]pyridin-2-yl, [1,2,5]thiadiazolo[3,4-b]pyridin-6-yl, 1H-pyrazolo[3,4-d]pyrimidin-1-yl, 1H-pyrrolo[3,2-b]pyridin-1-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 3H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, benzo[c][1,2,5]thiadiazol-5-yl, imidazo[1,5-a]pyridin-6-yl, imidazo[1,5-a]pyridin-7-yl, pyrazolo[1,5-a]pyrimidin-3-yl, thiazolo[5,4-b]pyridin-2-yl, optionally substituted with one or two independently selected R_4 substituents.

B may be heteroaryl selected from the group consisting of 1H-indazol-5-yl, , 7H-purin-2-yl, furo[3,2-b]pyridin-2-yl, pyrazolo[1,5-a]pyridin-3-yl, 2H-pyrazolo[3,4-b]pyridin-5-yl, 1H-pyrazolo[4,3-b]pyridin-1-yl, imidazo[1,2-a]pyridin-6-yl, imidazo[1,2-a]pyrazin-6-yl, imidazo[1,2-b]pyridazin-6-yl [1,2,3]triazolo[1,5-a]pyridin-6-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyrimidin-2-yl, [1,2,4]triazolo[1,5-a]pyrimidin-6-yl, [1,2,4]triazolo[1,5-a]pyrazin-6-yl, [1,2,4]triazolo[1,5-b]pyridazin-6-yl, [1,2,4]triazolo[4,3-a]pyridin-6-yl, thiazolo[4,5-b]pyrazin-2-yl, thiazolo[5,4-c]pyridin-2-yl, [1,2,5]thiadiazolo[3,4-b]pyridin-6-yl, 1H-pyrazolo[3,4-d]pyrimidin-1-yl, 1H-pyrrolo[3,2-b]pyridin-1-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl,

2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 3H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, benzo[c][1,2,5]thiadiazol-5-yl, imidazo[1,5-a]pyridin-6-yl, imidazo[1,5-a]pyridin-7-yl, pyrazolo[1,5-a]pyrimidin-3-yl, thiazolo[5,4-b]pyridin-2-yl, substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-indazol-5-yl, , 7H-purin-2-yl, furo[3,2-b]pyridin-2-yl, pyrazolo[1,5-a]pyridin-3-yl, 2H-pyrazolo[3,4-b]pyridin-5-yl, 1H-pyrazolo[4,3-b]pyridin-1-yl, imidazo[1,2-a]pyridin-6-yl, imidazo[1,2-a]pyrazin-6-yl, imidazo[1,2-b]pyridazin-6-yl [1,2,3]triazolo[1,5-a]pyridin-6-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyrimidin-2-yl, [1,2,4]triazolo[1,5-a]pyrimidin-6-yl, [1,2,4]triazolo[1,5-a]pyrazin-6-yl, [1,2,4]triazolo[1,5-b]pyridazin-6-yl, [1,2,4]triazolo[4,3-a]pyridin-6-yl, thiazolo[4,5-b]pyrazin-2-yl, thiazolo[5,4-c]pyridin-2-yl, [1,2,5]thiadiazolo[3,4-b]pyridin-6-yl, 1H-pyrazolo[3,4-d]pyrimidin-1-yl, 1H-pyrrolo[3,2-b]pyridin-1-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 3H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, benzo[c][1,2,5]thiadiazol-5-yl, imidazo[1,5-a]pyridin-6-yl, imidazo[1,5-a]pyridin-7-yl, pyrazolo[1,5-a]pyrimidin-3-yl, thiazolo[5,4-b]pyridin-2-yl, substituted with two independently selected R_4 substituents.

One aspect includes a compound of Formula (I), wherein B is heterocyclyl, wherein heterocyclyl is a 8- to 10- membered bicyclic carbon atom ring structure radical containing 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing at least one N, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally substituted with one or two independently selected R_4 substituents, or heterocyclyl is a 8 membered bicyclic carbon atom ring structure radical containing 2, or 3 N, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, and optionally containing a second heteroatom ring member selected from O or S, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a 9- membered bicyclic carbon atom ring

structure radical containing 2, 3 or 4 N, and optionally containing a second heteroatom ring member selected from O or S, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a unsubstituted 8- or 9-membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally containing a second heteroatom ring member selected from O or S. B may be heterocyclyl, wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally containing a second heteroatom ring member selected from O or S, and is unsubstituted or substituted with one R_4 substituent.

B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridinyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, optionally substituted with one or two independently selected R_4 substituents.

B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridinyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, substituted with two independently selected R_4 substituents.

B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridinyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, substituted with one R_4 substituent.

B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazol-7-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridin-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazol-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, optionally substituted with one or two independently selected R_4 substituents.

B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazol-7-yl, 5,6-

dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridin-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazol-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, substituted with one R_4 substituent.

B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazol-7-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridin-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazol-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, substituted with two independently selected R_4 substituents.

B may be selected from the group consisting of:

phenyl optionally substituted with one or two independently selected R_4 substituents;

heteroaryl,

wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing at least one N atom (or at least 2 N atoms), and optionally a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent, or

wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing at least 2 N atoms, optionally containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents; and

heterocyclyl,

wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing at least one N atom (or at least 2 N atoms), optionally containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents.

B may be selected from the group consisting of:

phenyl unsubstituted or substituted with one R_4 substituent;

heteroaryl,

wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 N atoms, and optionally, when the ring structure contains 1 or 2 N, a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent, or

wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N atoms, optionally, when the ring structure contains 2 or 3 N, containing

a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents; and

heterocyclyl,

wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, and optionally, when the ring structure contains 2 or 3 N, containing a second heteroatom ring member selected from O or S, optionally substituted with one or two independently selected R_4 substituents.B may be heteroaryl or heterocycl,

wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 N atoms, and optionally, when the ring structure contains 1 or 2 N, a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent, or

wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N atoms, optionally, when the ring structure contains 2 or 3 N, containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents; and

wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, and optionally, when the ring structure contains 2 or 3 N, containing a second heteroatom ring member selected from O or S, optionally substituted with one or two independently selected R_4 substituents.

B may be heteroaryl,

wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 N atoms, and optionally, when the ring structure contains 1 or 2 N, a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent, or

wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N atoms, optionally, when the ring structure contains 2 or 3 N, containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents.

One aspect includes a compound of Formula (I), wherein R_4 is selected from the group consisting of halogen, cyano, $C_{1\text{-}4}$alkyl, deutero-$C_{1\text{-}4}$alkyl, halo-$C_{1\text{-}4}$alkyl, $C_{1\text{-}4}$alkoxy, deutero-$C_{1\text{-}4}$alkoxy, amino, $C_{1\text{-}4}$alkyl-amino, $(C_{1\text{-}4}\text{alkyl})_2$-amino, $C_{3\text{-}6}$cycloalkyl, and heterocyclyl, wherein heterocyclyl is a 3- to 6- membered monocyclic carbon atom ring structure radical containing 1 or 2 heteroatom ring members independently selected from N, O, or S.

R_4 may be selected from the group consisting of halogen, cyano, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, C_{1-4}alkoxy, deutero-C_{1-4}alkoxy, C_{1-4}alkyl-amino, C_{3-6}cycloalkyl, and heterocylyl.

R_4 may be halogen selected from the group consisting of bromo, chloro, fluoro, and iodo. R_4 may be halogen selected from the group consisting of chloro and fluoro. R_4 may be chloro. R_4 may be fluoro.

R_4 may be cyano.

R_4 may be C_{1-4}alkyl selected from the group consisting of methyl, ethyl, propyl, isopropyl, butyl, and tert-butyl. R_4 may be methyl or ethyl. R_4 may be methyl. R_4 may be ethyl.

R_4 may be deutero-C_{1-4}alkyl wherein C_{1-4}alkyl is selected from the group consisting of methyl, ethyl, propyl, isopropyl, butyl, and tert-butyl partially or completely substituted with one or more deuterium atoms where allowed by available valences. R_4 may be ($^2H_3$)methyl.

R_4 may be halo-C_{1-4}alkyl wherein C_{1-4}alkyl is selected from the group consisting of methyl, ethyl, propyl, isopropyl, butyl, and tert-butyl, partially or completely substituted with one or more halogen atoms where allowed by available valences. R_4 may be halo-C_{1-4}alkyl selected from the group consisting of difluoromethyl and trifluoromethyl. R_4 may be difluoromethyl. R_4 may be trifluoromethyl.

R_4 may be C_{1-4}alkoxy selected from the group consisting of methoxy, ethoxy, propoxy, isopropoxy, butoxy and tert-butoxy. R_4 may be methoxy.

R_4 may be deutero- C_{1-4}alkoxy wherein C_{1-4}alkoxy is selected from the group consisting of methoxy, ethoxy, propoxy, isopropoxy, butoxy and tert-butoxy partially or completely substituted with one or more deuterium atoms where allowed by available valences. R_4 may be ($^2H_3$)methoxy.

R_4 may be C_{3-6}cycloalkyl selected from the group consisting of cyclopropyl, cyclobutyl, cyclopenyl, and cyclohexyl. R_4 may be cyclopropyl.

R_4 may be C_{1-4}alkyl-amino, wherein C_{1-4}alkyl is selected from the group consisting of methyl, ethyl, propyl, isopropyl, butyl, isobutyl, sec-butyl, and tert-butyl. R_4 may be methylamino.

R_4 may be heterocyclyl, wherein hetercylyl is a 3- to 6- membered monocyclic carbon atom ring structure radical containing 1 or 2 heteroatom ring members independently selected from N, O, or S. R_4 may be heterocyclyl selected from the group consisting of aziridinyl, oziranyl, thiiranyl, azetidinyl, oxetanyl, thietanyl, pyrrolidinyl, pyrazolinyl, tetrahyrofuranyl,

thilolanyl, piperidinyl, piperazinyl, tetrahydro-2H-pyranyl, 1,4-dioxanyl, morpholinyl, and thianyl. R_4 is azetidinyl.

R_4 may be selected from the group consisting of halogen, cyano, methyl, ethyl, ($^2H_3$)methyl, ($^2H_3$)ethyl, difluoromethyl, trifluoromethyl, methoxy, ethoxy, ($^2H_3$)methoxy, methylamino, ethylamino, cyclopropyl, and azetidinyl.

Certain aspects include a compound of Formula (I), wherein X is CH. In other aspects, X is CF. In other aspects, X is N.

One aspect includes a compound of Formula (I), wherein R_w is selected from the group consisting of halogen, hydroxyl, cyano, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, amino, C_{1-4}alkyl-amino, (C_{1-4}alkyl)$_2$-amino, C_{1-4}alkoxy, and halo-C_{1-4}alkoxy. R_w may be selected from the group consisting of halogen and C_{1-4}alkyl. R_w may be halogen selected from the group consisting of bromo, chloro, fluoro, and iodo. R_w may be fluoro. R_w may be C_{1-4}alkyl selected from methyl, ethyl, propyl, isopropyl, and tert-butyl. R_w may be methyl. R_w may be fluoro, chloro, bromo, methyl or ethyl.

Certain aspects include a compound of Formula (I), wherein n is 0. In other aspects, n is 1.

One aspect includes a compound of Formula (I), wherein n is 0, and R_1 is hydrogen or C_{1-4}alkyl. In one aspect, n is 0, and X is C.

One aspect includes a compound of Formula (I), wherein n is 0, R_2 is C_{1-4}alkyl, halo-C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, cyclopropyl, cyclobutyl, phenyl, or oxetanyl, optionally substituted with one or two R_3 substituents, and R_1 is hydrogen or C_{1-4}alkyl.

One aspect includes a compound of Formula (I), wherein A is A1, A2, A3, A4, A5 or A6, n is 0, and R_1 is hydrogen or C_{1-4}alkyl. In another aspect, A is A1, A2, A3, A4, A5 or A6, n is 0, X is C, and R_1 is hydrogen or C_{1-4}alkyl.

One aspect includes a compound of Formula (I), wherein:

n is 0;

X is C;

R_2 is C_{1-4}alkyl, halo-C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, cyclopropyl, cyclobutyl, phenyl, or oxetanyl, optionally substituted with one or two R_3 substituents;

R_4 is selected from the group consisting of halogen, cyano, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, C_{1-4}alkoxy, deutero-C_{1-4}alkoxy, C_{1-4}alkyl-amino, C_{3-6}cycloalkyl, and heterocylyl;

R_1 is hydrogen or C_{1-4}alkyl; and

B is selected from the group consisting of:

phenyl unsubstituted or substituted with one R_4 substituent;

heteroaryl,

wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 N atoms, and optionally, when the ring structure contains 1 or 2 N, a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent, or

wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N atoms, optionally, when the ring structure contains 2 or 3 N, containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents; and

heterocyclyl,

wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, and optionally, when the ring structure contains 2 or 3 N, containing a second heteroatom ring member selected from O or S, optionally substituted with one or two independently selected R_4 substituents.

One aspect includes a compound of Formula (I), wherein:

n is 0;

X is C;

R_2 is C_{1-4}alkyl, halo-C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, cyclopropyl, cyclobutyl, phenyl, or oxetanyl, optionally substituted with one or two R_3 substituents;

R_4 is selected from the group consisting of halogen, cyano, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, C_{1-4}alkoxy, deutero-C_{1-4}alkoxy, C_{1-4}alkyl-amino, C_{3-6}cycloalkyl, and heterocylyl;

R_1 is hydrogen or C_{1-4}alkyl;

A is A1-A24; and

B is selected from the group consisting of:

phenyl unsubstituted or substituted with one R_4 substituent;

heteroaryl,

wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 N atoms, and optionally, when the ring structure contains 1 or 2 N, a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent, or

wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N atoms, optionally, when the ring structure contains 2 or 3 N, containing

a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents; and

heterocyclyl,

wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, and optionally, when the ring structure contains 2 or 3 N, containing a second heteroatom ring member selected from O or S, optionally substituted with one or two independently selected R_4 substituents.

One aspect includes a compound of Formula (I), wherein:

n is 0;

X is C;

R_2 is $C_{1\text{-}4}$alkyl, halo-$C_{1\text{-}4}$alkyl, hydroxyl-$C_{1\text{-}4}$alkyl, cyclopropyl, cyclobutyl, phenyl, or oxetanyl, optionally substituted with one or two R_3 substituents;

R_4 is selected from the group consisting of halogen, cyano, methyl, ethyl, ($^2H_3$)methyl, ($^2H_3$)ethyl, difluoromethyl, trifluoromethyl, methoxy, ethoxy, ($^2H_3$)methoxy, methylamino, ethylamino, cyclopropyl, and azetidinyl;

R_1 is hydrogen or $C_{1\text{-}4}$alkyl;

A is A1-A6; and

B is selected from the group consisting of heteroaryl and heretocycl,

wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 N atoms, and optionally, when the ring structure contains 1 or 2 N, a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent, or

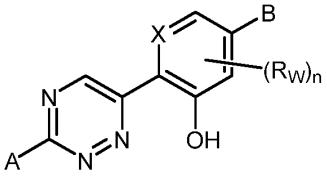
wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N atoms, optionally, when the ring structure contains 2 or 3 N, containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents; and

wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, and optionally, when the ring structure contains 2 or 3 N, containing a second heteroatom ring member selected from O or S, optionally substituted with one or two independently selected R_4 substituents.

In an embodiment thereof, B is unsubstituted. In another embodiment thereof, B is heteroaryl, unsubtituted or substituted with one R_4 substituent. In another embodiment thereof, B is the 9- membered bicyclic carbon atom ring structure radical. In another embodiment

thereof, B is the 5- or 6- membered monocyclic aromatic carbon atom ring structure radical. In an embodiment thereof, B is heterocycl, unsubtituted or substituted with one R_4 substituent.

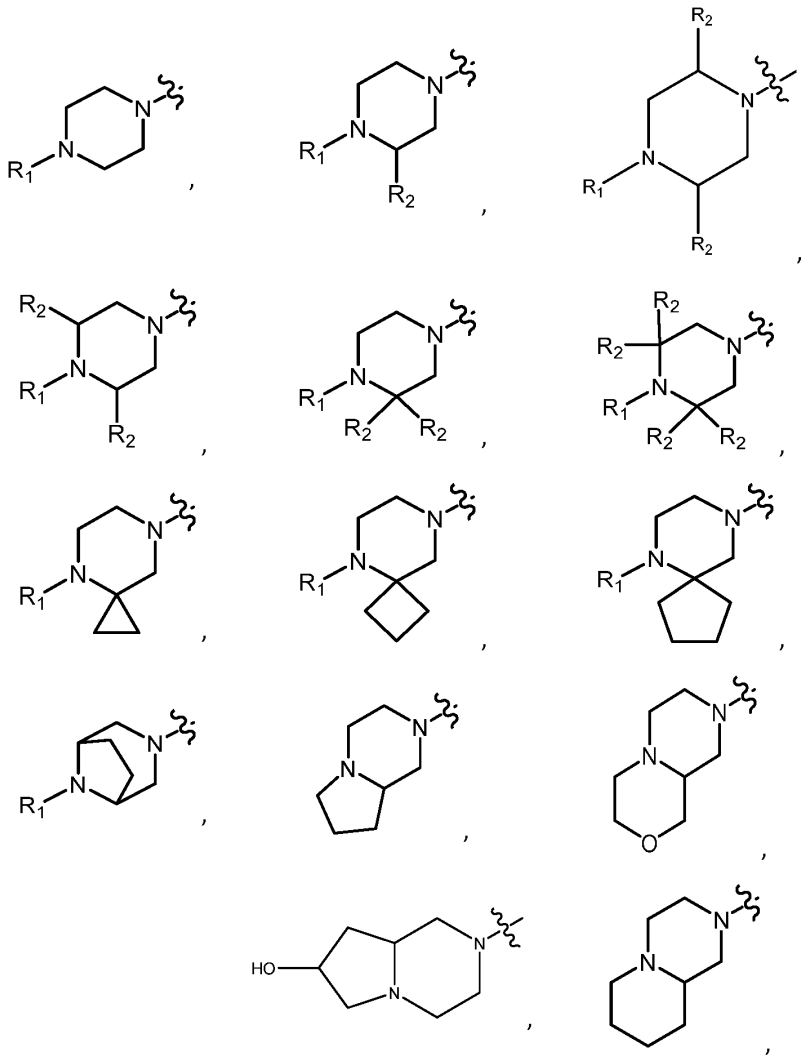
Another aspect includes a compound of Formula (I), or a form thereof:



(I)

wherein:

A is selected from the group consisting of:



and any stereoisomer thereof;

R_1 is selected from the group consisting of hydrogen, C_{1-4}alkyl, and C_{3-6}cycloalkyl;

R_2 is independently selected from the group consisting of halogen, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, C_{1-4}alkoxy-C_{1-4}alkyl, C_{2-4}alkenyl, C_{3-6}cycloalkyl, phenyl, pyridinyl, and hetercyclyl, wherein heterocyclyl is a 3- to 6-membered carbon atom ring structure radical containing 1 or 2 heteroatom ring members selected from N, O, and S, and

wherein each instance of C_{1-4}alkyl, C_{3-6}cycloalkyl, phenyl, pyridnyl, and heterocyclyl is optionally substituted with one or two R_3 substituents;

R_3 is independently selected from the group consisting of halogen, hydroxyl, C_{1-4}alkyl, C_{1-4}alkoxy, and C_{3-6}cycloalkyl;

B is selected from the group consisting of:

phenyl optionally substituted with one or two independently selected R_4 substituents; and

heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 heteroatoms selected from N, O, and S, optionally substituted with one R_4 substituent;

R_4 is selected from the group consisting of halogen, cyano, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, C_{1-4}alkoxy, deutero-C_{1-4}alkoxy, amino, C_{1-4}alkyl-amino, $(C_{1-4}alkyl)_2$-amino, C_{3-6}cycloalkyl, and heterocyclyl, wherein heterocyclyl is a 3- to 6- membered monocyclic carbon atom ring structure radical containing 1 or 2 heteroatom ring members independently selected from N, O, or S;

X is selected from the group consisting of CH, CF, and N;

R_w is selected from the group consisting of halogen, hydroxyl, cyano, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, amino, C_{1-4}alkyl-amino, $(C_{1-4}alkyl)_2$-amino, C_{1-4}alkoxy, and halo-C_{1-4}alkoxy; and

n is selected from the group consisting of 0 or 1;

wherein a form of the compound is selected from the group consisting of a salt, hydrate, solvate, and tautomer form thereof.

Except where provided, each of the terms and definitions for this aspect are the same as defined above.

In an aspect thereof, B may be a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 heteroatoms selected from N, O, and S. B may be selected from the group consisting of: phenyl optionally substituted with one or two independently selected R_4 substituents; and heteroaryl, wherein heteroaryl is a 5- or 6-membered monocyclic aromatic carbon atom ring structure radical containing at least one N

atom. B may be selected from the group consisting of: phenyl optionally substituted with one R_4 substituent; and heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 N atoms, and optionally, when the ring structure contains 1 or 2 N, a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent. B may be heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 N atoms, optionally substituted with one R_4 substituent.

One aspect includes a compound of Formula (I), wherein B is selected from the group consisting of:

phenyl optionally substituted with one or two independently selected R_4 substituents;

heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 heteroatoms selected from N, O, and S, optionally substituted with one R_4 substituent.

B may be phenyl optionally substituted with one or two independently selected R_4 substituents. B may be unsubstituted phenyl or phenyl substituted with one R_4 substituent.

One aspect includes a compound of Formula (I), wherein B is heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 heteroatoms selected from N, O, and S, optionally substituted with one R_4 substituent. B may be heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 1, 2, or 3 N atoms, and optionally a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent. B may be heteroaryl, wherein heteroaryl is a 5- or 6- membered monocyclic aromatic carbon atom ring structure radical containing 2 or 3 heteroatoms selected from N, O, and S, optionally substituted with one R_4 substituent. B may be heteroaryl, wherein heteroaryl is a 5- or 6-membered monocyclic aromatic carbon atom ring structure radical containing 2 or 3 N atoms, and optionally a second heteroatom selected from O, and S, optionally substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of furanyl, thiophenyl, 1H-pyrazolyl, 1H-imidazolyl, isoxazolyl, 1,3-thiazolyl, 1,3-oxazolyl, tetrazolyl, 1H-1,2,3-triazolyl, 2H-1,2,3-triazolyl, 1,2,4-oxadiazolyl, 1,3,4-oxadiazolyl, 1,2,3-thiadiazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyridazinyl, 1H-1,2,4-triazolyl, 1,2,4-thiadiazolyl, 1,3,4-oxadiazolyl, isothiazolyl, 1,2,5-thiadiazolyl, and 1,3,4-thiadiazolyl, optionally substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-pyrazolyl, 1H-imidazolyl, 1,3-thiazolyl, 1,3-oxazolyl, 1H-1,2,3-triazolyl, 2H-1,2,3-triazolyl, pyridinyl, pyrimidinyl, pyrazinyl, 1H-1,2,4-triazolyl, 1,2,4-thiadiazolyl, 1,3,4-oxadiazolyl, isothiazolyl, 1,2,5-thiadiazolyl, and 1,3,4-thiadiazolyl, optionally substituted with one R_4 substituent.

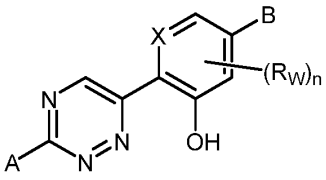
B may be heteroaryl selected from the group consisting of 1H-pyrazolyl, 1H-imidazolyl, 1,3-thiazolyl, 1,3-oxazolyl, 1H-1,2,3-triazolyl, 2H-1,2,3-triazolyl, pyridinyl, pyrimidinyl, pyrazinyl, 1H-1,2,4-triazolyl, 1,2,4-thiadiazolyl, 1,3,4-oxadiazolyl, isothiazolyl, 1,2,5-thiadiazolyl, and 1,3,4-thiadiazolyl, substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of furan-2-yl, furan-3-yl, thiophen-2-yl, thiophen-3-yl, 1H-pyrazol-1-yl, 1H-pyrazol-3-yl, 1H-pyrazol-4-yl, 1H-pyrazol-5-yl, 1H-imidazol-1-yl, 1H-imidazol-4-yl, isoxazol-3-yl, isoxazol-4-yl, isoxazol-5-yl, 1,3-thiazol-2-yl, 1,3-thiazol-4-yl, 1,3-thiazol-5-yl, 1,3-oxazol-2-yl, 1,3-oxazol-4-yl, 1,3-oxazol-5-yl, tetrazol-5-yl, 1H-1,2,3-triazol-1-yl, 1H-1,2,3-triazol-4-yl, 1H-1,2,3-triazol-5-yl, 2H-1,2,3-triazol-2-yl, 2H-1,2,3-triazol-4-yl, 2H-1,2,3-triazol-5-yl, 1,2,4-oxadiazol-3-yl, 1,3,4-oxadiazol-2-yl, 1,2,3-thiadiazol-4-yl, 1,2,3-thiadiazol-5-yl, pyridin-2-yl, pyridin-3-yl, pyridin-4-yl, pyrimidin-4-yl, pyrimidin-2-yl, pyrazin-2-yl, pyridazin-3-yl, pyridazin-4-yl, 1H-1,2,4-triazol-3-yl, 1H-1,2,4-triazol-5-yl, 1,2,4-thiadiazol-3-yl, 1,2,4-thiadiazol-5-yl, 1,3,4-oxadiazol-2-yl, isothiazol-3-yl, isothiazol-4-yl, isothiazol-5-yl, 1,2,5-thiadiazol-3-yl, 1,3,4-thiadiazol-2-yl, optionally substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-pyrazol-4-yl, 1H-imidazol-1-yl, 1,3-thiazol-2-yl, 1,3-thiazol-4-yl, 1,3-oxazol-2-yl, 1H-1,2,3-triazol-4-yl, 1H-1,2,3-triazol-5-yl, 2H-1,2,3-triazol-2-yl, 2H-1,2,3-triazol-4-yl, pyridin-3-yl, pyridin-4-yl, pyrimidin-4-yl, pyrimidin-2-yl, pyrazin-2-yl, 1H-1,2,4-triazol-3-yl, 1,2,4-thiadiazol-3-yl, 1,2,4-thiadiazol-5-yl, 1,3,4-oxadiazol-2-yl, isothiazol-3-yl, isothiazol-4-yl, isothiazol-5-yl, 1,2,5-thiadiazol-3-yl, 1,3,4-thiadiazol-2-yl, optionally substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-pyrazol-4-yl, 1H-imidazol-1-yl, 1,3-thiazol-2-yl, 1,3-thiazol-4-yl, 1,3-oxazol-2-yl, 1H-1,2,3-triazol-4-yl, 1H-1,2,3-triazol-5-yl, 2H-1,2,3-triazol-2-yl, 2H-1,2,3-triazol-4-yl, pyridin-3-yl, pyridin-4-yl, pyrimidin-4-yl, pyrimidin-2-yl, pyrazin-2-yl, 1H-1,2,4-triazol-3-yl, 1,2,4-thiadiazol-3-yl, 1,2,4-thiadiazol-5-yl, 1,3,4-oxadiazol-2-yl, isothiazol-3-yl, isothiazol-4-yl, isothiazol-5-yl, 1,2,5-thiadiazol-3-yl, 1,3,4-thiadiazol-2-yl, substituted with one R_4 substituent.

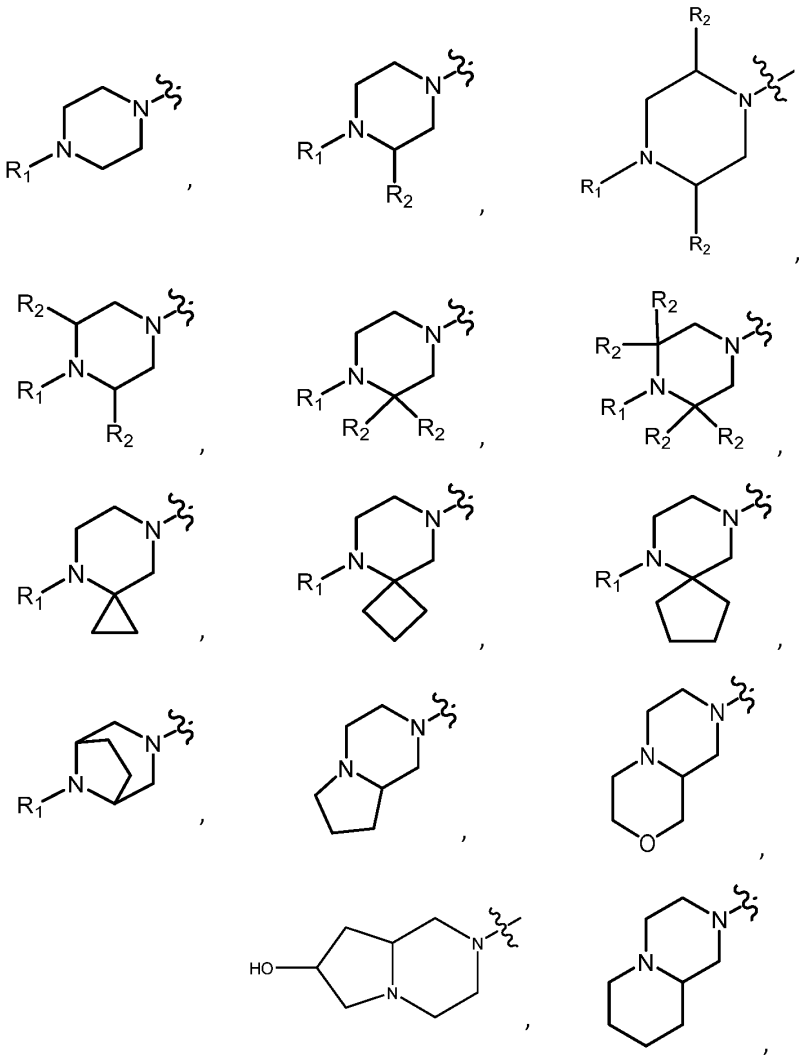
An aspect of the compound of Formula (I), or a form thereof, is:



(I)

wherein:

A is selected from the group consisting of:



and any stereoisomer thereof;

R_1 is selected from the group consisting of hydrogen, $C_{1\text{-}4}$alkyl, and $C_{3\text{-}6}$cycloalkyl;

R_2 is independently selected from the group consisting of halogen, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, hydroxyl-C_{1-4}alkyl, C_{1-4}alkoxy-C_{1-4}alkyl, C_{2-4}alkenyl, C_{3-6}cycloalkyl, phenyl, pyridinyl, and hetercyclyl, wherein heterocyclyl is a 3- to 6-membered carbon atom ring structure radical containing 1 or 2 heteroatom ring members selected from N, O, and S, and

wherein each instance of C_{1-4}alkyl, C_{3-6}cycloalkyl, phenyl, pyridnyl, and heterocyclyl is optionally substituted with one or two R_3 substituents;

R_3 is independently selected from the group consisting of halogen, hydroxyl, C_{1-4}alkyl, C_{1-4}alkoxy, and C_{3-6}cycloalkyl;

B is selected from the group consisting of:

heteroaryl, wherein heteroaryl is a 9- or 10- membered bicyclic aromatic carbon atom ring structure radical having 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S, optionally substituted with one or two independently selected R_4 substituents; and

heterocyclyl, wherein heterocyclyl is a 8- to 10- membered bicyclic aromatic carbon atom ring structure radical containing 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S, optionally substituted with one or two independently selected R_4 substituents;

R_4 is selected from the group consisting of halogen, cyano, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, C_{1-4}alkoxy, deutero-C_{1-4}alkoxy, amino, C_{1-4}alkyl-amino, $(C_{1-4}alkyl)_2$-amino, C_{3-6}cycloalkyl, and heterocyclyl, wherein heterocyclyl is a 3- to 6- membered monocyclic carbon atom ring structure radical containing 1 or 2 heteroatom ring members independently selected from N, O, or S;

X is selected from the group consisting of CH, CF, and N;

R_w is selected from the group consisting of halogen, hydroxyl, cyano, C_{1-4}alkyl, deutero-C_{1-4}alkyl, halo-C_{1-4}alkyl, amino, C_{1-4}alkyl-amino, $(C_{1-4}alkyl)_2$-amino, C_{1-4}alkoxy, and halo-C_{1-4}alkoxy; and

n is selected from the group consisting of 0 or 1;

wherein a form of the compound is selected from the group consisting of a salt, hydrate, solvate, and tautomer form thereof.

Except where provided, each of the terms and definitions for this aspect are the same as defined above.

B may be heteroaryl, wherein heteroaryl is a 9-or 10- membered bicyclic aromatic ring system having 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S.

B may be heteroaryl, wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N atoms, and optionally substituted with one or two independently selected R_4 substituents.

B may be heterocyclyl, wherein heterocyclyl is a 8- to 10- membered bicyclic aromatic carbon atom ring structure radical containing 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S.

B may be selected from the group consisting of:

heteroaryl, wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing at least 2 N atoms; and(

heterocyclyl, wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing at least one N atom.

B may be selected from the group consisting of:

heteroaryl, wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N atoms, optionally substituted with one or two independently selected R_4 substituents; and

heterocyclyl, wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally substituted with one or two independently selected R_4 substituents.

One aspect includes a compound of Formula (I), wherein B is heteroaryl, wherein heteroaryl is a 9- or 10- membered aromatic carbon atom ring structure radical having 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S, optionally substituted with one or two independently selected R_4 substituents. B may be heteroaryl, wherein heteroaryl is a 9- or 10- membered bicyclic carbon atom ring structure radical containing at least one N atom ring member, optionally containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents. B may be heteroaryl, wherein heteroaryl is a 9- or 10- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N atoms, optionally containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents. B may be heteroaryl, wherein heteroaryl is a 9-membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally substituted with one or two independently selected R_4 substituents. B may be heteroaryl, wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally containing a second heteroatom ring member selected from O or S, and optionally substituted with one or two independently selected R_4 substituents. B may be

heteroaryl, wherein heteroaryl is a unsubstituted 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally containing a second heteroatom ring member selected from O or S. B may be heteroaryl, wherein heteroaryl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally containing a second heteroatom ring member selected from O or S, and is unsubstituted or substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-indazolyl, 2H-indazolyl, indolizinyl, benzofuranyl, benzothiophenyl, 1H-benzimidazolyl, 1,3-benzoxazolyl, 1,3-benzothiazolyl, 1,3-benzodioxolyl, 1,2,3-benzotriazolyl, 7H-purinyl, furo[3,2-b]pyridinyl, furo[3,2-c]pyridinyl, 1,3-oxazolo[5,4-b]pyridinyl, thieno[3,2-c]pyridinyl, thieno[2,3-d]pyrimidinyl, pyrrolo[1,2-a]pyrimidinyl, pyrrolo[1,2-a]pyrazinyl, pyrrolo[1,2-b]pyridazinyl, pyrazolo[1,5-a]pyridinyl, 1H-pyrrolo[2,3-b]pyridinyl, 5H-pyrrolo[2,3-b]pyrazinyl, 1H-pyrrolo[2,3-c]pyridinyl, 1H-pyrazolo[3,4-b]pyridinyl, 2H-pyrazolo[3,4-b]pyridinyl, 1H-pyrazolo[3,4-b]pyrazinyl, 1H-pyrazolo[3,4-c]pyridinyl, 1H-pyrazolo[4,3-b]pyridinyl, 2H-pyrazolo[4,3-b]pyridinyl, 2H-pyrazolo[4,3-c]pyridinyl, 1H-pyrazolo[4,3-d]pyrimidinyl, pyrazolo[1,5-a]pyrazinyl, imidazo[1,2-a]pyridinyl, imidazo[1,2-a]pyrimidinyl, imidazo[1,2-a]pyrazinyl, imidazo[1,2-b]pyridazinyl, imidazo[1,2-c]pyrimidin-2-yl, 1H-imidazo[4,5-b]pyridinyl, 3H-imidazo[4,5-b]pyridinyl, imidazo[2,1-b][1,3]thiazolyl, imidazo[2,1-b][1,3,4]thiadiazolyl, [1,3]oxazolo[4,5-b]pyridinyl, [1,2,3]triazolo[1,5-a]pyridinyl, 1H-[1,2,3]triazolo[4,5-b]pyridinyl, 3H-[1,2,3]triazolo[4,5-b]pyridinyl, [1,2,4]triazolo[1,5-a]pyridinyl, [1,2,4]triazolo[1,5-a]pyrimidinyl, [1,2,4]triazolo[1,5-a]pyrazinyl, [1,2,4]triazolo[1,5-b]pyridazinyl, [1,2,4]triazolo[4,3-a]pyridinyl, tetrazolo[1,5-a]pyridinyl, tetrazolo[1,5-b]pyridazinyl, thiazolo[4,5-b]pyrazinyl, thiazolo[5,4-c]pyridinyl, quinolinyl, isoquinolinyl, [1,2,5]thiadiazolo[3,4-b]pyridinyl, 1H-pyrazolo[3,4-d]pyrimidinyl, 1H-pyrrolo[3,2-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-c]pyridinyl, 3H-[1,2,3]triazolo[4,5-b]pyridinyl, 3H-[1,2,3]triazolo[4,5-c]pyridinyl, benzo[c][1,2,5]thiadiazolyl, imidazo[1,5-a]pyridinyl, pyrazolo[1,5-a]pyrimidinyl, and thiazolo[5,4-b]pyridinyl, optionally substituted with one or two independently selected R_4 substituents.

B may be heteroaryl selected from the group consisting of 2H-indazolyl, 7H-purinyl, furo[3,2-b]pyridinyl, pyrazolo[1,5-a]pyridinyl, 2H-pyrazolo[3,4-b]pyridinyl, 1H-pyrazolo[4,3-b]pyridinyl, imidazo[1,2-a]pyridinyl, imidazo[1,2-a]pyrazinyl, imidazo[1,2-b]pyridazinyl, [1,2,3]triazolo[1,5-a]pyridinyl, 1H-[1,2,3]triazolo[4,5-b]pyridinyl, [1,2,4]triazolo[1,5-a]pyridinyl, [1,2,4]triazolo[1,5-a]pyrimidinyl, [1,2,4]triazolo[1,5-a]pyrazinyl,

[1,2,4]triazolo[1,5-b]pyridazinyl, [1,2,4]triazolo[4,3-a]pyridinyl, thiazolo[4,5-b]pyrazinyl, thiazolo[5,4-c]pyridinyl, [1,2,5]thiadiazolo[3,4-b]pyridinyl, 1H-pyrazolo[3,4-d]pyrimidinyl, 1H-pyrrolo[3,2-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-c]pyridinyl, 3H-[1,2,3]triazolo[4,5-b]pyridinyl, 3H-[1,2,3]triazolo[4,5-c]pyridinyl, benzo[c][1,2,5]thiadiazolyl, imidazo[1,5-a]pyridinyl, pyrazolo[1,5-a]pyrimidinyl, and thiazolo[5,4-b]pyridinyl, optionally substituted with one or two independently selected R_4 substituents.

B may be heteroaryl selected from the group consisting of 2H-indazolyl, 7H-purinyl, furo[3,2-b]pyridinyl, pyrazolo[1,5-a]pyridinyl, 2H-pyrazolo[3,4-b]pyridinyl, 1H-pyrazolo[4,3-b]pyridinyl, imidazo[1,2-a]pyridinyl, imidazo[1,2-a]pyrazinyl, imidazo[1,2-b]pyridazinyl, [1,2,3]triazolo[1,5-a]pyridinyl, 1H-[1,2,3]triazolo[4,5-b]pyridinyl, [1,2,4]triazolo[1,5-a]pyridinyl, [1,2,4]triazolo[1,5-a]pyrimidinyl, [1,2,4]triazolo[1,5-a]pyrazinyl, [1,2,4]triazolo[1,5-b]pyridazinyl, [1,2,4]triazolo[4,3-a]pyridinyl, thiazolo[4,5-b]pyrazinyl, thiazolo[5,4-c]pyridinyl, [1,2,5]thiadiazolo[3,4-b]pyridinyl, 1H-pyrazolo[3,4-d]pyrimidinyl, 1H-pyrrolo[3,2-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-c]pyridinyl, 3H-[1,2,3]triazolo[4,5-b]pyridinyl, 3H-[1,2,3]triazolo[4,5-c]pyridinyl, benzo[c][1,2,5]thiadiazolyl, imidazo[1,5-a]pyridinyl, pyrazolo[1,5-a]pyrimidinyl, and thiazolo[5,4-b]pyridinyl, substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 2H-indazolyl, 7H-purinyl, furo[3,2-b]pyridinyl, pyrazolo[1,5-a]pyridinyl, 2H-pyrazolo[3,4-b]pyridinyl, 1H-pyrazolo[4,3-b]pyridinyl, imidazo[1,2-a]pyridinyl, imidazo[1,2-a]pyrazinyl, imidazo[1,2-b]pyridazinyl, [1,2,3]triazolo[1,5-a]pyridinyl, 1H-[1,2,3]triazolo[4,5-b]pyridinyl, [1,2,4]triazolo[1,5-a]pyridinyl, [1,2,4]triazolo[1,5-a]pyrimidinyl, [1,2,4]triazolo[1,5-a]pyrazinyl, [1,2,4]triazolo[1,5-b]pyridazinyl, [1,2,4]triazolo[4,3-a]pyridinyl, thiazolo[4,5-b]pyrazinyl, thiazolo[5,4-c]pyridinyl, [1,2,5]thiadiazolo[3,4-b]pyridinyl, 1H-pyrazolo[3,4-d]pyrimidinyl, 1H-pyrrolo[3,2-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-b]pyridinyl, 2H-[1,2,3]triazolo[4,5-c]pyridinyl, 3H-[1,2,3]triazolo[4,5-b]pyridinyl, 3H-[1,2,3]triazolo[4,5-c]pyridinyl, benzo[c][1,2,5]thiadiazolyl, imidazo[1,5-a]pyridinyl, pyrazolo[1,5-a]pyrimidinyl, and thiazolo[5,4-b]pyridinyl, substituted with two independently selected R_4 substituents.

B may be heteroaryl selected from the group consisting of 1H-indazol-5-yl, 2H-indazol-5-yl, indolizin-2-yl,benzofuran-2-yl, benzofuran-5-yl, benzothiophen-2-yl, benzothiophen-3-yl, 1H-benzimidazol-2-yl, 1H-benzimidazol-5-yl, 1H-benzimidazol-6-yl, 1,3-benzoxazol-2-yl, 1,3-benzoxazol-5-yl, 1,3-benzoxazol-6-yl, 1,3-benzothiazol-2-yl, 1,3-benzothiazol-5-yl, 1,3-benzothiazol-6-yl, 1,3-benzodioxol-5-yl, 1,2,3-benzotriazol-5-yl,

7H-purin-2-yl, furo[3,2-b]pyridin-2-yl, furo[3,2-c]pyridin-2-yl, furo[2,3-c]pyridin-2-yl, 1,3-oxazolo[5,4-b]pyridine-5-yl, thieno[3,2-c]pyridin-2-yl, thieno[2,3-d]pyrimidin-6-yl, pyrrolo[1,2-a]pyrimidin-7-yl, pyrrolo[1,2-a]pyrazin-7-yl, pyrrolo[1,2-b]pyridazin-2-yl, pyrazolo[1,5-a]pyridin-2-yl, pyrazolo[1,5-a]pyridin-3-yl, pyrazolo[1,5-a]pyridin-5-yl, 1H-pyrrolo[2,3-b]pyridin-5-yl, 5H-pyrrolo[2,3-b]pyrazin-2-yl,1H-pyrrolo[2,3-c]pyridin-4-yl, 1H-pyrazolo[3,4-b]pyridin-5-yl, 1H-pyrazolo[3,4-b]pyridin-6-yl, 2H-pyrazolo[3,4-b]pyridin-5-yl, 1H-pyrazolo[3,4-b]pyrazin-5-yl, 1H-pyrazolo[3,4-c]pyridin-1-yl, 1H-pyrazolo[3,4-c]pyridin-5-yl, 1H-pyrazolo[4,3-b]pyridin-1-yl, 1H-pyrazolo[4,3-b]pyridin-5-yl, 1H-pyrazolo[4,3-b]pyridin-6-yl, 2H-pyrazolo[4,3-b]pyridin-5-yl, 2H-pyrazolo[4,3-c]pyridin-5-yl,1H-pyrazolo[4,3-d]pyrimidin-5-yl, pyrazolo[1,5-a]pyrazin-2-yl, imidazo[1,2-a]pyridin-2-yl, imidazo[1,2-a]pyridin-6-yl, imidazo[1,2-a]pyrimidin-2-yl, imidazo[1,2-a]pyrimidin-6-yl, imidazo[1,2-a]pyrazin-2-yl, imidazo[1,2-a]pyrazin-6-yl, imidazo[1,2-b]pyridazin-2-yl, imidazo[1,2-b]pyridazin-6-yl, imidazo[1,2-c]pyrimidin-2-yl, 1H-imidazo[4,5-b]pyridin-5-yl, 3H-imidazo[4,5-b]pyridin-5-yl, imidazo[2,1-b][1,3]thiazol-6-yl, imidazo[2,1-b][1,3,4]thiadiazol-6-yl, [1,3]oxazolo[4,5-b]pyridin-2-yl, [1,2,3]triazolo[1,5-a]pyridin-5-yl, [1,2,3]triazolo[1,5-a]pyridin-6-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 3H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, [1,2,4]triazolo[1,5-a]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyrimidin-2-yl, [1,2,4]triazolo[1,5-a]pyrimidin-6-yl, [1,2,4]triazolo[1,5-a]pyrazin-6-yl, [1,2,4]triazolo[1,5-b]pyridazin-6-yl, [1,2,4]triazolo[4,3-a]pyridin-6-yl, tetrazolo[1,5-a]pyridin-7-yl, tetrazolo[1,5-b]pyridazin-7-yl, thiazolo[4,5-b]pyrazin-2-yl, thiazolo[5,4-c]pyridin-2-yl, tetrazolo[1,5-a]pyridin-7-yl, tetrazolo[1,5-b]pyridazin-7-yl, quinolin-6-yl, isoquinolin-6-yl, [1,2,5]thiadiazolo[3,4-b]pyridin-6-yl, 1H-pyrazolo[3,4-d]pyrimidin-1-yl, 1H-pyrrolo[3,2-b]pyridin-1-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 3H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, benzo[c][1,2,5]thiadiazol-5-yl, imidazo[1,5-a]pyridin-6-yl, imidazo[1,5-a]pyridin-7-yl, pyrazolo[1,5-a]pyrimidin-3-yl, thiazolo[5,4-b]pyridin-2-yl, optionally substituted with one or two independently selected R_4 substituents.

B may be heteroaryl selected from the group consisting of 1H-indazol-5-yl, , 7H-purin-2-yl, furo[3,2-b]pyridin-2-yl, pyrazolo[1,5-a]pyridin-3-yl, 2H-pyrazolo[3,4-b]pyridin-5-yl, 1H-pyrazolo[4,3-b]pyridin-1-yl, imidazo[1,2-a]pyridin-6-yl, imidazo[1,2-a]pyrazin-6-yl, imidazo[1,2-b]pyridazin-6-yl [1,2,3]triazolo[1,5-a]pyridin-6-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyrimidin-2-yl, [1,2,4]triazolo[1,5-a]pyrimidin-6-yl, [1,2,4]triazolo[1,5-

a]pyrazin-6-yl, [1,2,4]triazolo[1,5-b]pyridazin-6-yl, [1,2,4]triazolo[4,3-a]pyridin-6-yl, thiazolo[4,5-b]pyrazin-2-yl, thiazolo[5,4-c]pyridin-2-yl, [1,2,5]thiadiazolo[3,4-b]pyridin-6-yl, 1H-pyrazolo[3,4-d]pyrimidin-1-yl, 1H-pyrrolo[3,2-b]pyridin-1-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 3H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, benzo[c][1,2,5]thiadiazol-5-yl, imidazo[1,5-a]pyridin-6-yl, imidazo[1,5-a]pyridin-7-yl, pyrazolo[1,5-a]pyrimidin-3-yl, thiazolo[5,4-b]pyridin-2-yl, optionally substituted with one or two independently selected R_4 substituents.

B may be heteroaryl selected from the group consisting of 1H-indazol-5-yl, , 7H-purin-2-yl, furo[3,2-b]pyridin-2-yl, pyrazolo[1,5-a]pyridin-3-yl, 2H-pyrazolo[3,4-b]pyridin-5-yl, 1H-pyrazolo[4,3-b]pyridin-1-yl, imidazo[1,2-a]pyridin-6-yl, imidazo[1,2-a]pyrazin-6-yl, imidazo[1,2-b]pyridazin-6-yl [1,2,3]triazolo[1,5-a]pyridin-6-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyrimidin-2-yl, [1,2,4]triazolo[1,5-a]pyrimidin-6-yl, [1,2,4]triazolo[1,5-a]pyrazin-6-yl, [1,2,4]triazolo[1,5-b]pyridazin-6-yl, [1,2,4]triazolo[4,3-a]pyridin-6-yl, thiazolo[4,5-b]pyrazin-2-yl, thiazolo[5,4-c]pyridin-2-yl, [1,2,5]thiadiazolo[3,4-b]pyridin-6-yl, 1H-pyrazolo[3,4-d]pyrimidin-1-yl, 1H-pyrrolo[3,2-b]pyridin-1-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 3H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, benzo[c][1,2,5]thiadiazol-5-yl, imidazo[1,5-a]pyridin-6-yl, imidazo[1,5-a]pyridin-7-yl, pyrazolo[1,5-a]pyrimidin-3-yl, thiazolo[5,4-b]pyridin-2-yl, substituted with one R_4 substituent.

B may be heteroaryl selected from the group consisting of 1H-indazol-5-yl, , 7H-purin-2-yl, furo[3,2-b]pyridin-2-yl, pyrazolo[1,5-a]pyridin-3-yl, 2H-pyrazolo[3,4-b]pyridin-5-yl, 1H-pyrazolo[4,3-b]pyridin-1-yl, imidazo[1,2-a]pyridin-6-yl, imidazo[1,2-a]pyrazin-6-yl, imidazo[1,2-b]pyridazin-6-yl [1,2,3]triazolo[1,5-a]pyridin-6-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 1H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyridin-6-yl, [1,2,4]triazolo[1,5-a]pyrimidin-2-yl, [1,2,4]triazolo[1,5-a]pyrimidin-6-yl, [1,2,4]triazolo[1,5-a]pyrazin-6-yl, [1,2,4]triazolo[1,5-b]pyridazin-6-yl, [1,2,4]triazolo[4,3-a]pyridin-6-yl, thiazolo[4,5-b]pyrazin-2-yl, thiazolo[5,4-c]pyridin-2-yl, [1,2,5]thiadiazolo[3,4-b]pyridin-6-yl, 1H-pyrazolo[3,4-d]pyrimidin-1-yl, 1H-pyrrolo[3,2-b]pyridin-1-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, 2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl, 3H-[1,2,3]triazolo[4,5-c]pyridin-6-yl, benzo[c][1,2,5]thiadiazol-5-yl, imidazo[1,5-a]pyridin-6-yl, imidazo[1,5-a]pyridin-7-yl, pyrazolo[1,5-a]pyrimidin-3-yl, thiazolo[5,4-b]pyridin-2-yl, substituted with two independently

selected R_4 substituents.

One aspect includes a compound of Formula (I), wherein B is heterocyclyl, wherein heterocyclyl is a 8- to 10- membered bicyclic carbon atom ring structure radical containing 1, 2, 3, or 4 heteroatom ring members independently selected from N, O, or S, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing at least one N, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally substituted with one or two independently selected R_4 substituents, or heterocyclyl is a 8 membered bicyclic carbon atom ring structure radical containing 2, or 3 N, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, and optionally containing a second heteroatom ring member selected from O or S, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, and optionally containing a second heteroatom ring member selected from O or S, optionally substituted with one or two independently selected R_4 substituents. B may be heterocyclyl, wherein heterocyclyl is a unsubstituted 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally containing a second heteroatom ring member selected from O or S. B may be heterocyclyl, wherein heterocyclyl is a 8- or 9- membered bicyclic carbon atom ring structure radical containing 2, 3 or 4 N, optionally containing a second heteroatom ring member selected from O or S, and is unsubstituted or substituted with one R_4 substituent.

B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridinyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, optionally substituted with one or two independently selected R_4 substituents.

B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazolyl, 5,6-dihydro-

4H-pyrrolo[1,2-b]pyrazolyl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridinyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, substituted with two independently selected R_4 substituents.

B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridinyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazolyl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazolyl, substituted with one R_4 substituent.

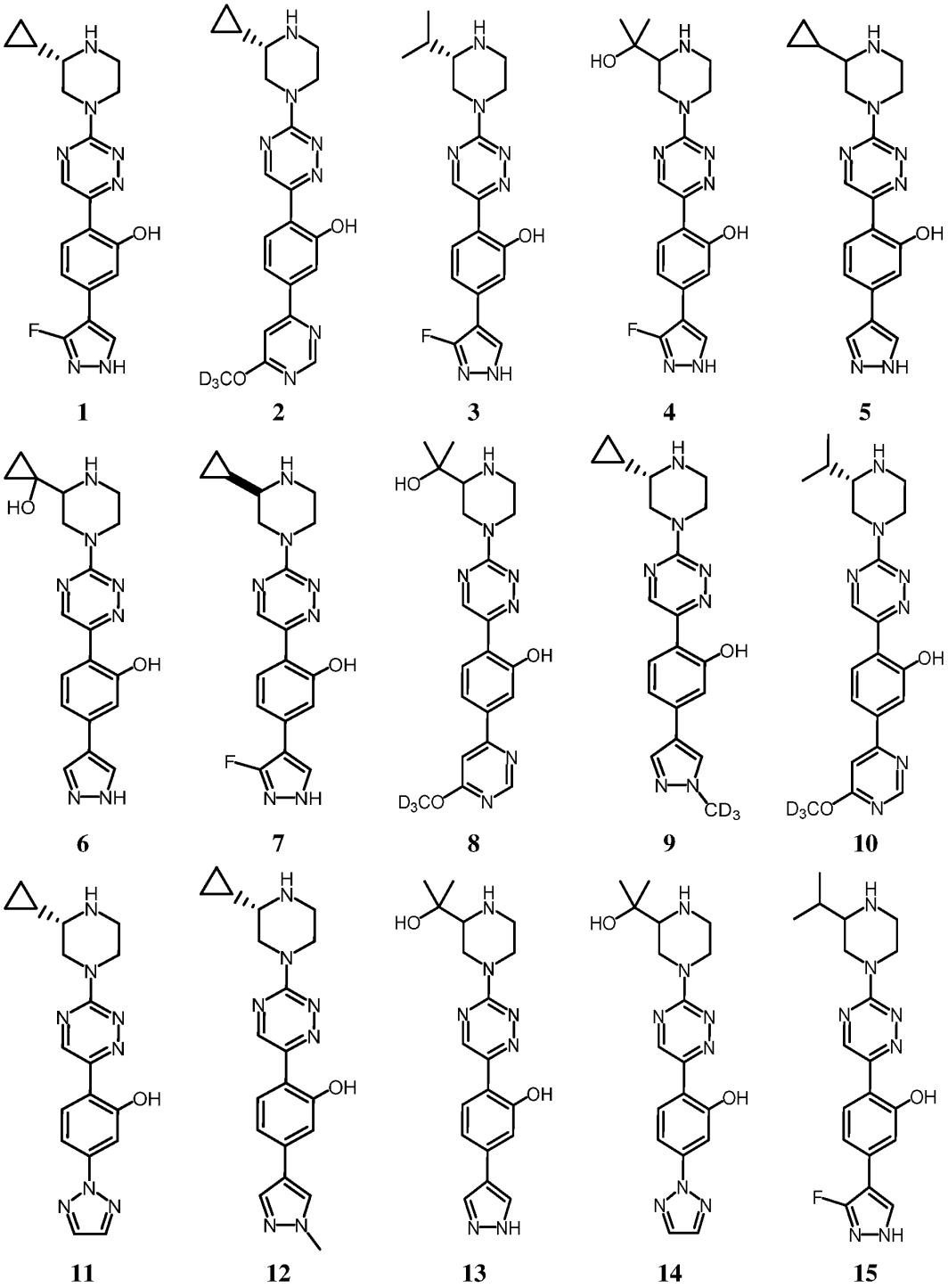
B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazol-7-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridin-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazol-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, optionally substituted with one or two independently selected R_4 substituents.

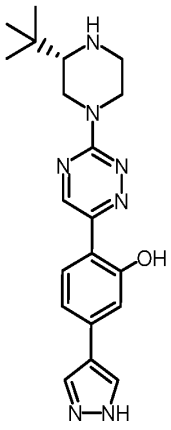
B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazol-7-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridin-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazol-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, substituted with one R_4 substituent.

B may be heterocyclyl selected from the group consisting of 5,6-dihydro-[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl, 2,3-dihydro-1H-imidazo[1,2-b]pyrazol-7-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 4,5,6,7-tetrahydropyrazolo[1,5-a]pyridin-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, 6,7-dihydro-5H-pyrrolo[1,2-a]imidazol-3-yl, 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, and 5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl, substituted with two independently selected R_4 substituents.

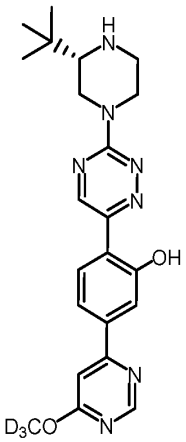
An aspect of the compound of Formula (I) or a form thereof includes a compound selected from the group consisting of, wherein "#" indicates that the compound is a racemic

mixture of enantiomers, and wherein "&" indicates that the compound may exist as the opposite enantiomer:

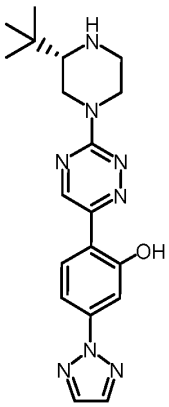




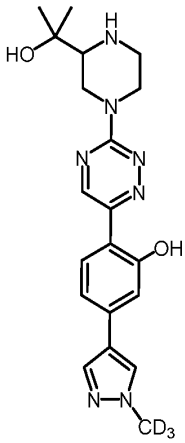
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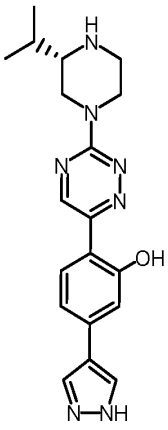
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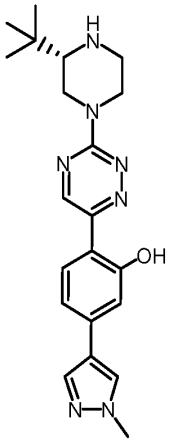
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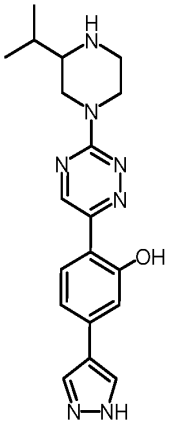
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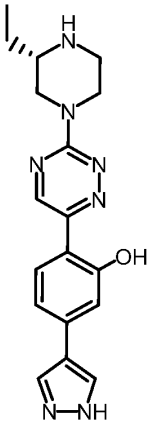
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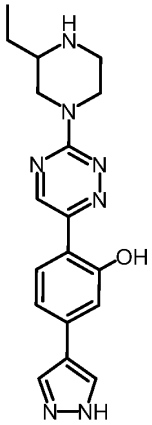
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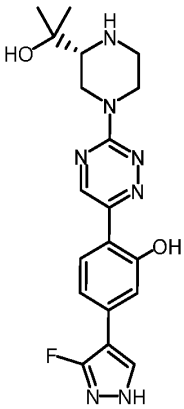
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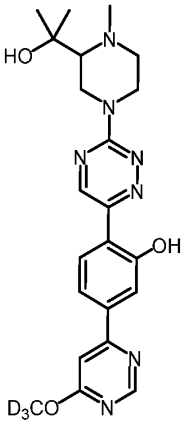
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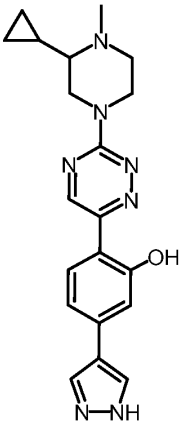
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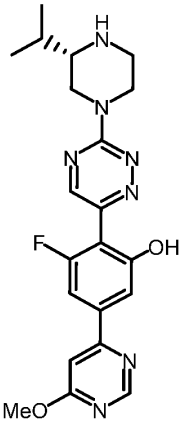
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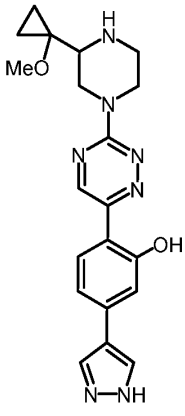
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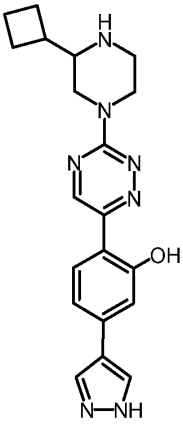
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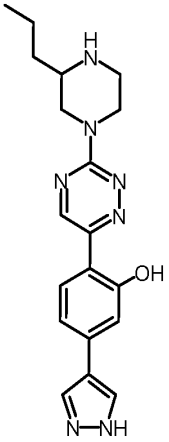
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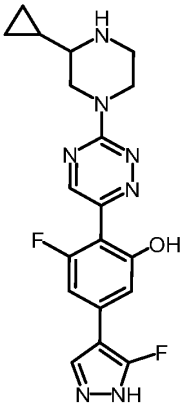


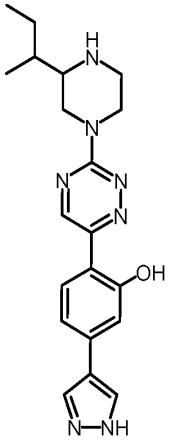
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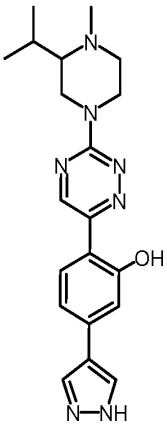


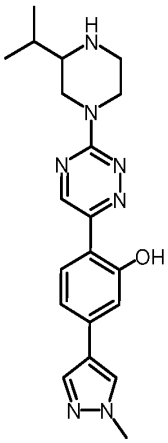
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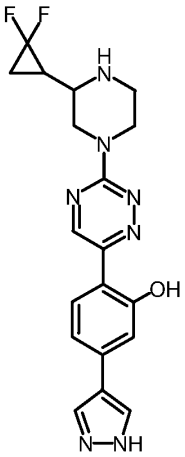

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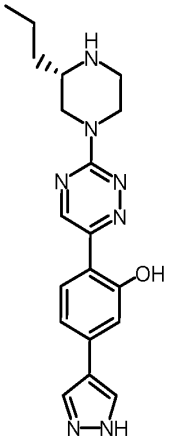

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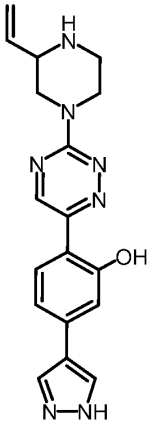

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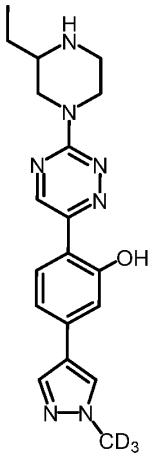

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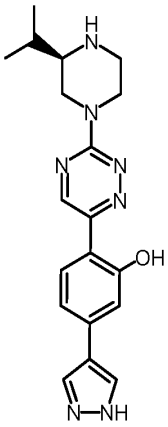

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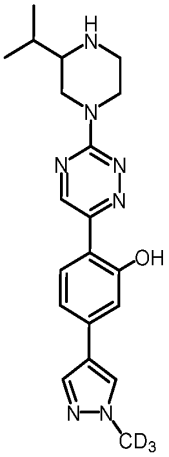

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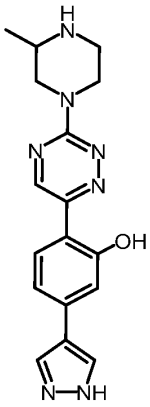

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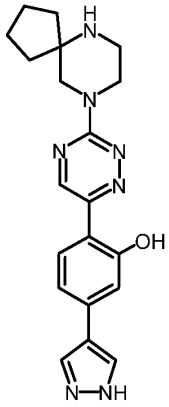

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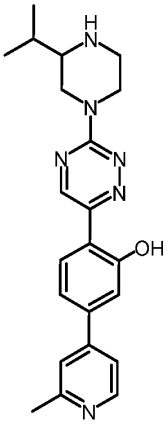

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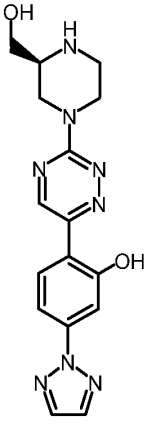

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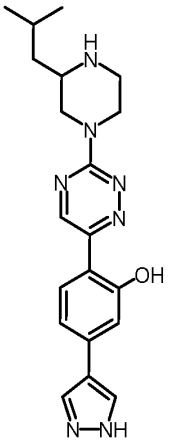

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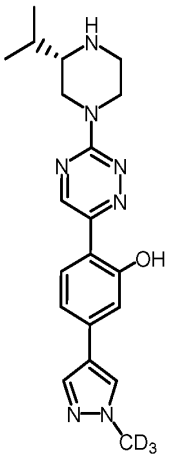

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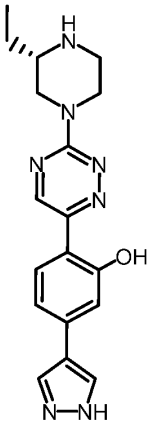

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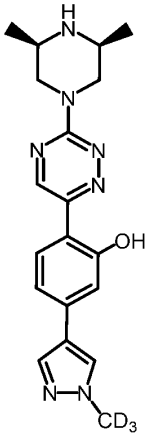
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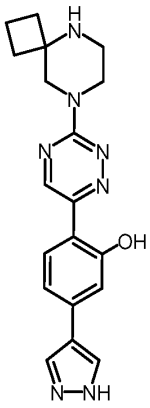
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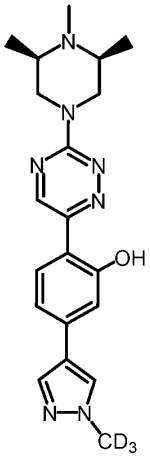
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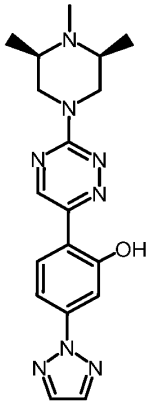
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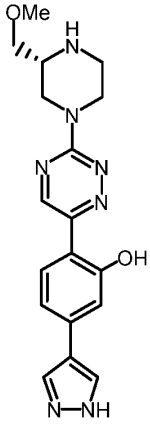
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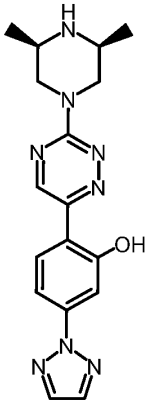
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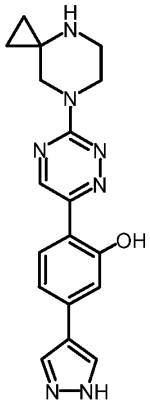
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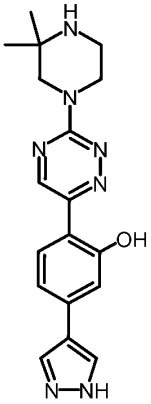
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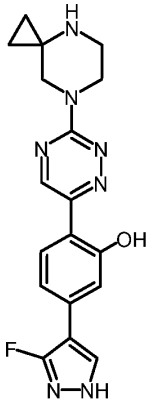
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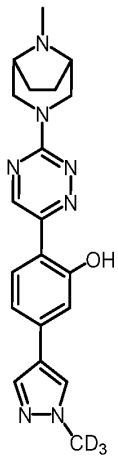
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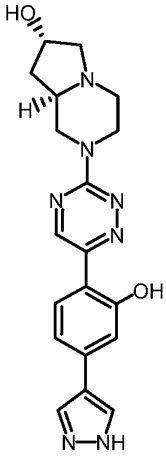
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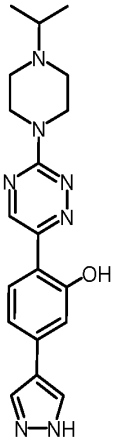
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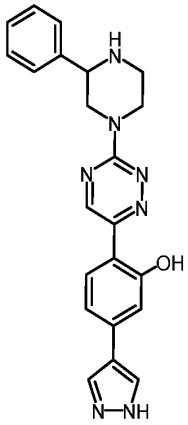
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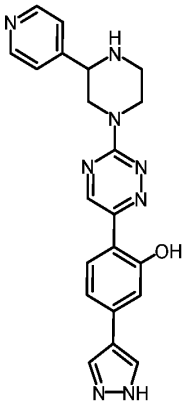
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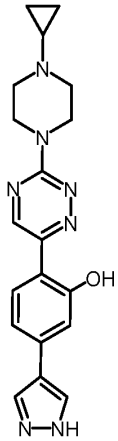
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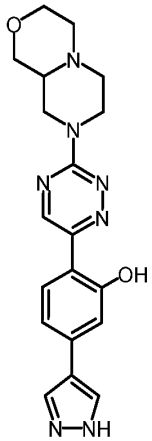
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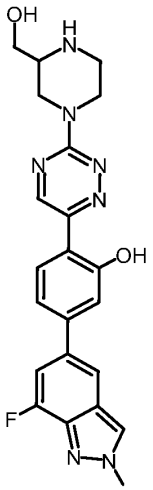
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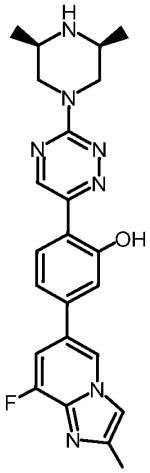
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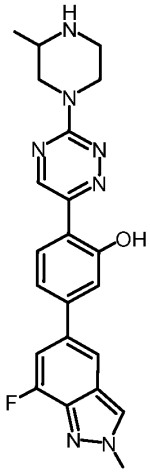
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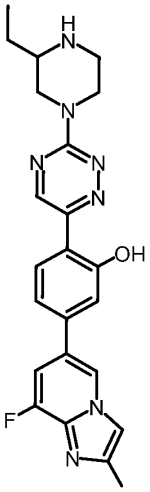
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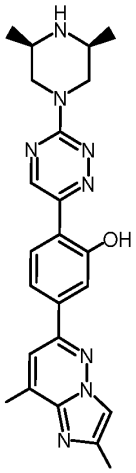
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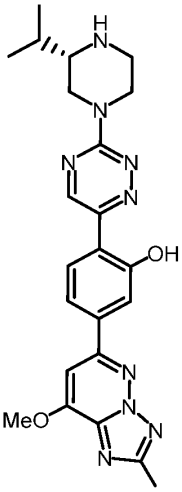
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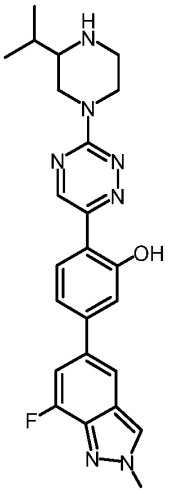
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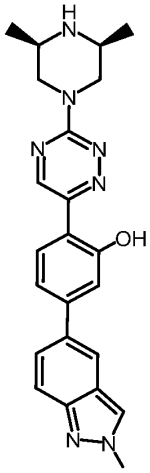
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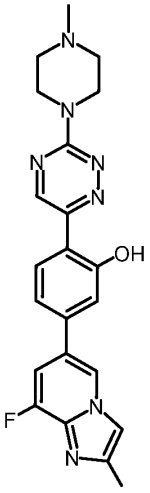
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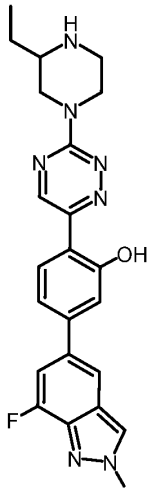
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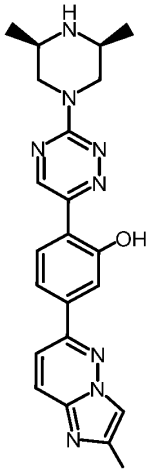
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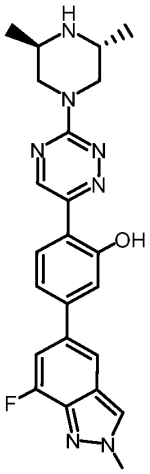
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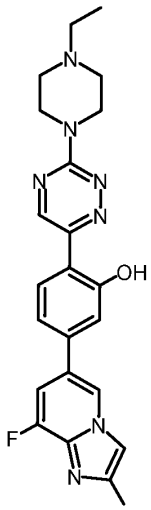
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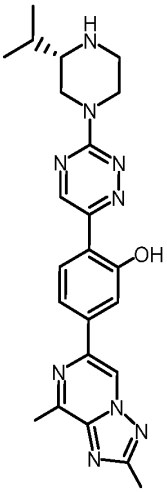
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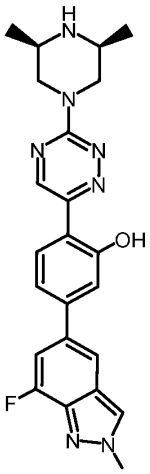
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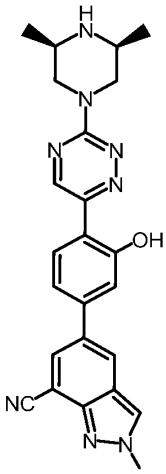
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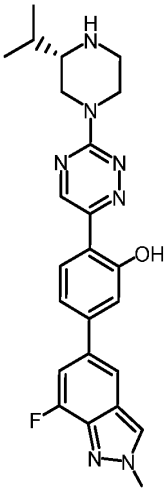
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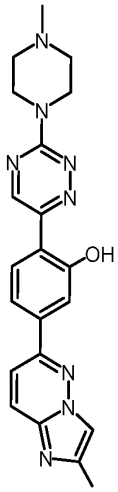
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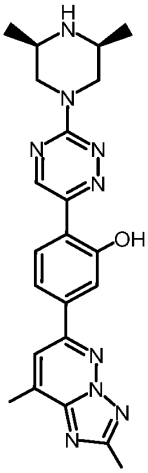
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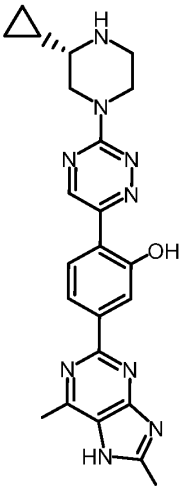
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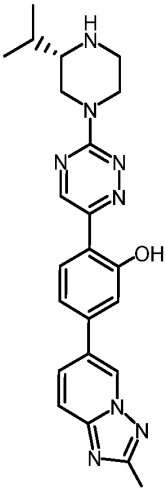
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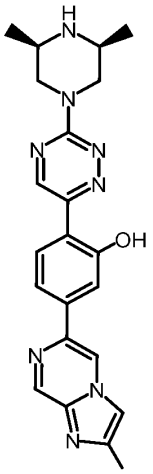
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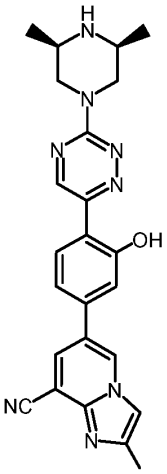
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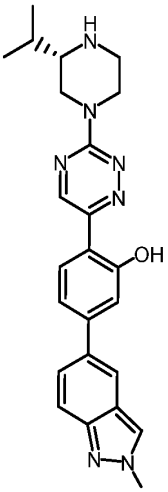
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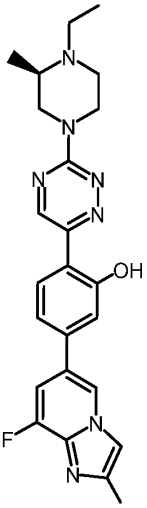
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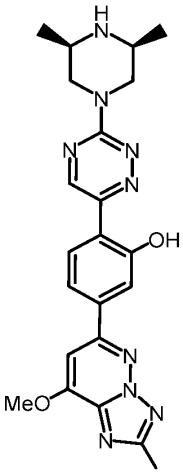
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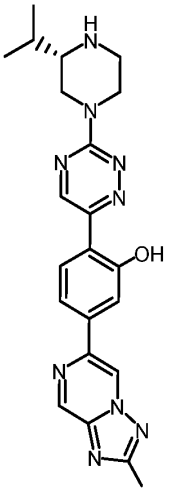
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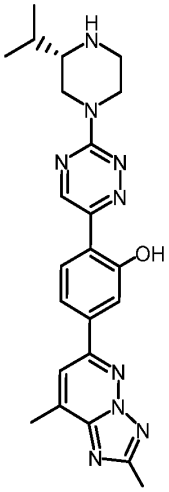
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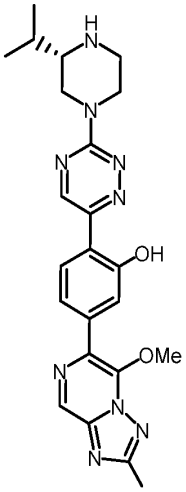
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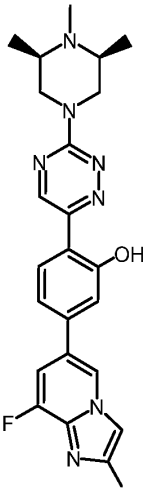
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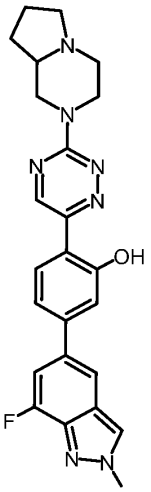
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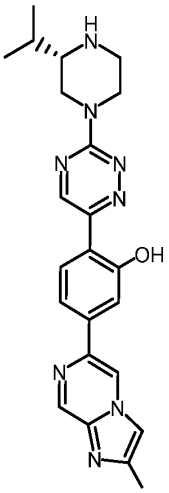
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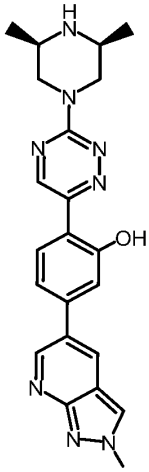
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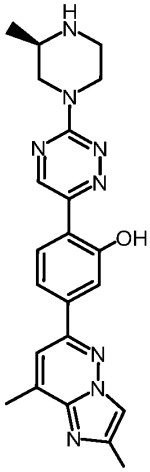
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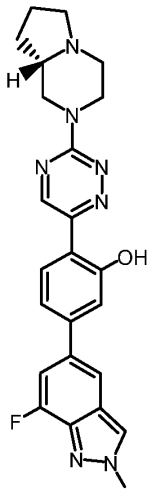
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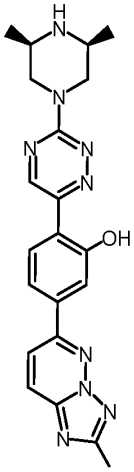
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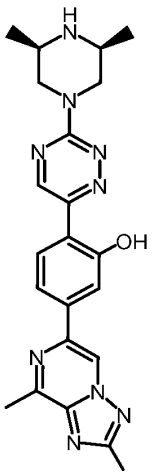
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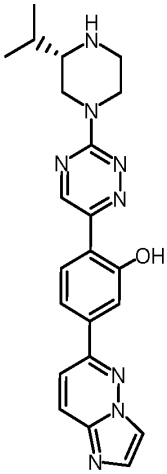
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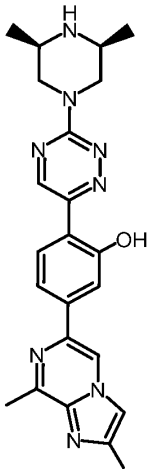
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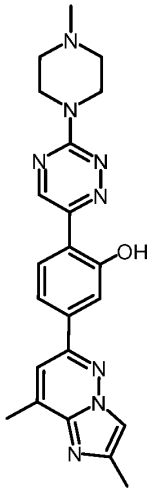
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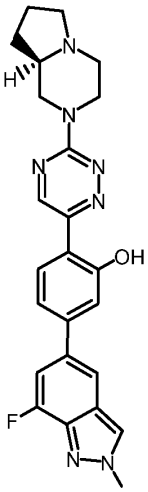
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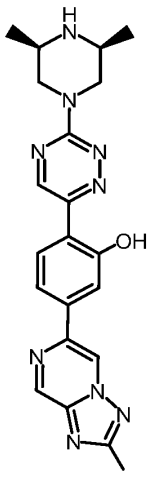
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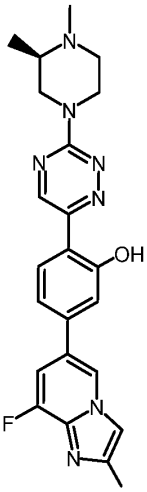
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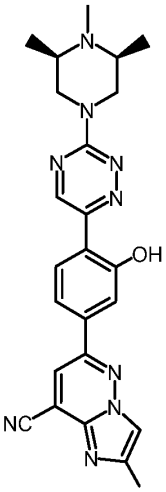
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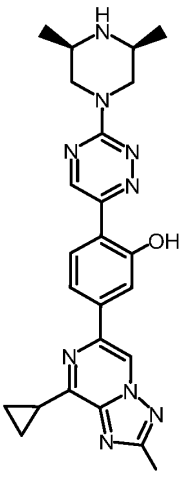
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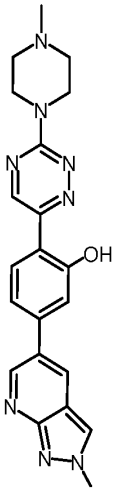
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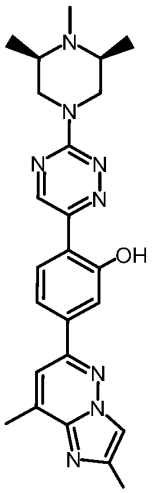
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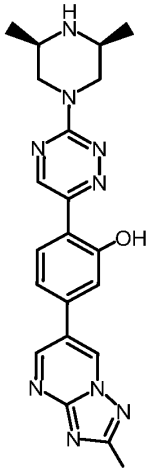
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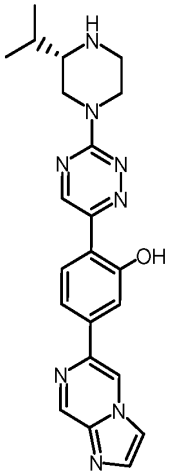
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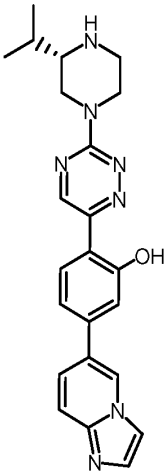
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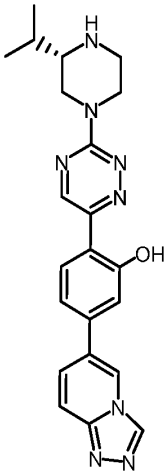
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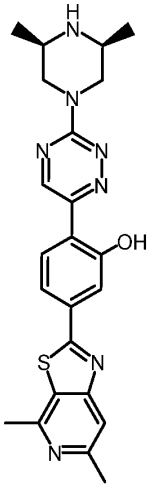
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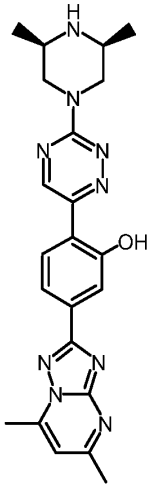
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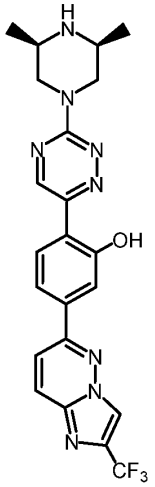
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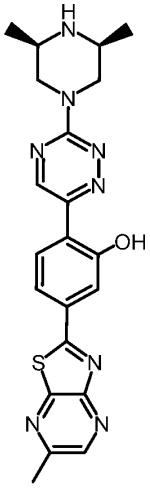
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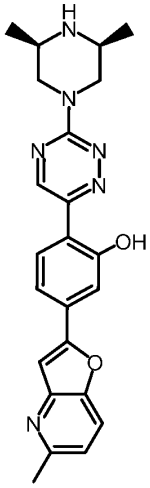
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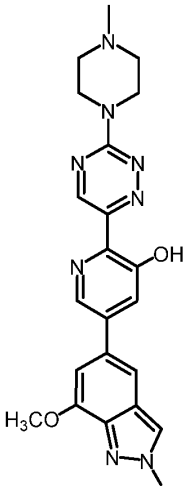
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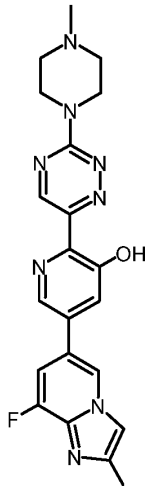
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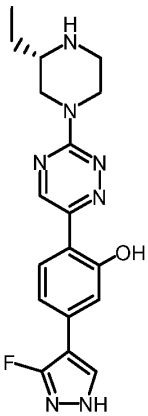
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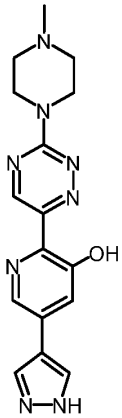
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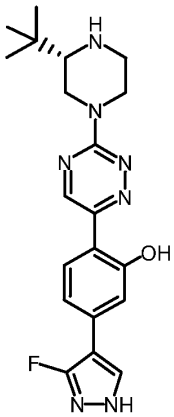
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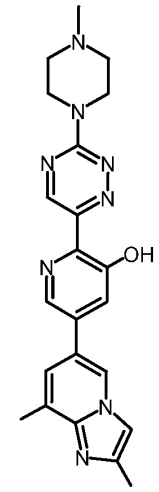
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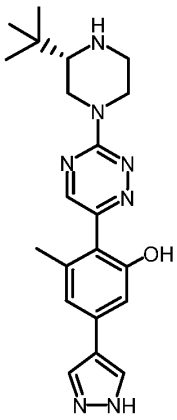
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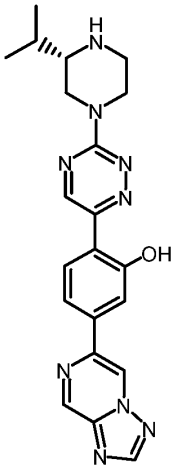
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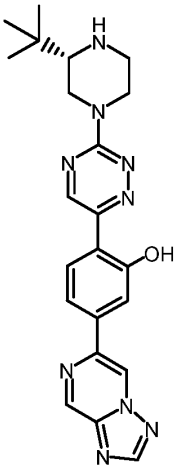
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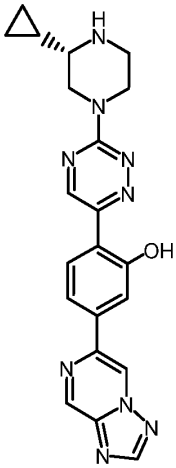
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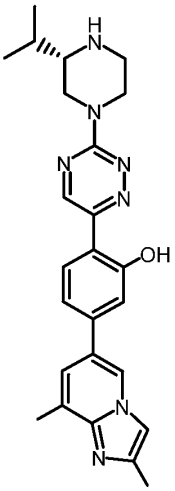
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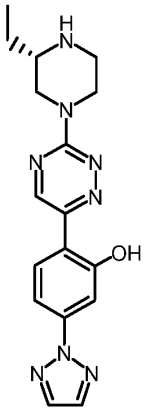
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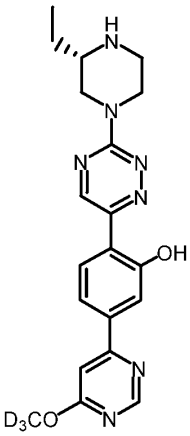
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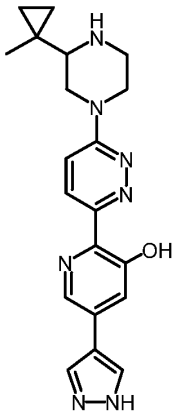
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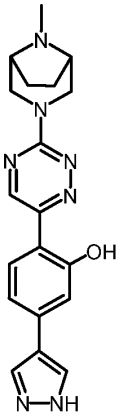
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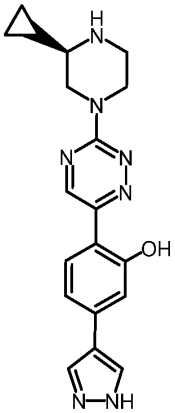
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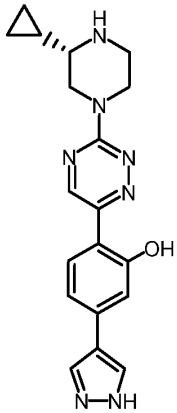
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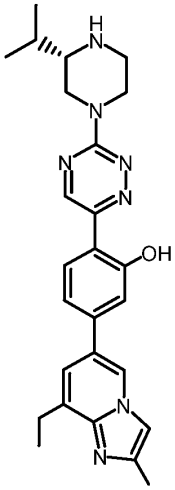
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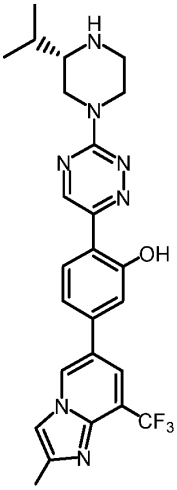
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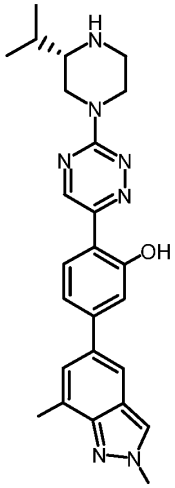
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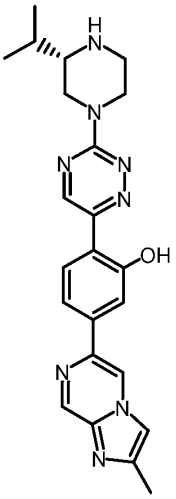
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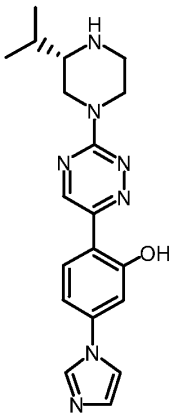
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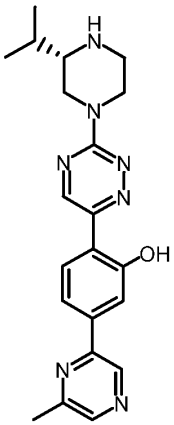
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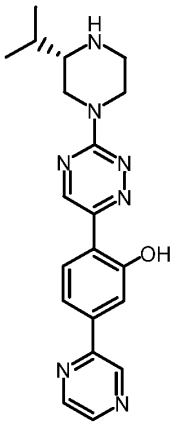
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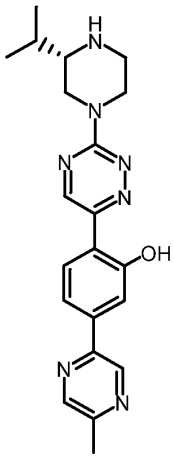
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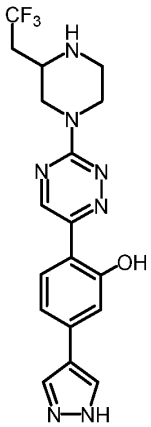
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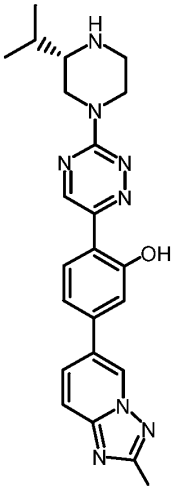
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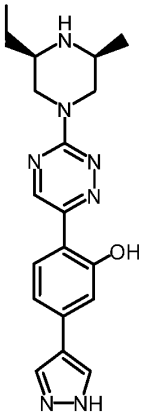
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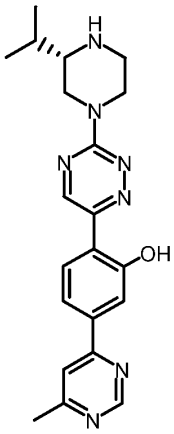
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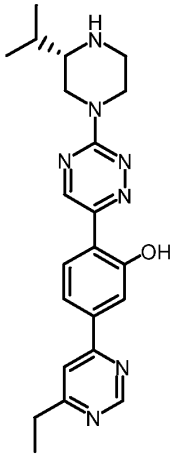
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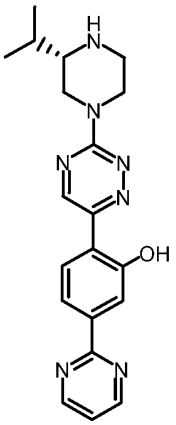
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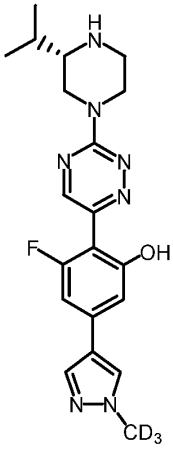
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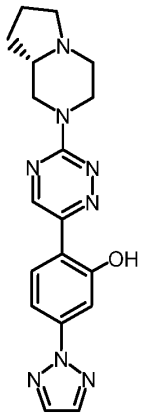
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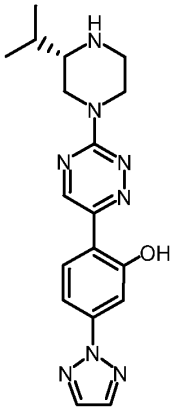
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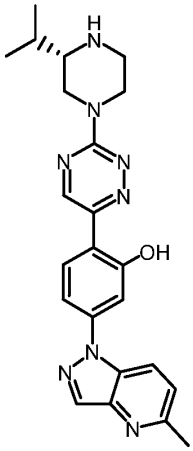
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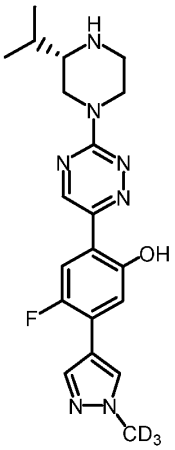
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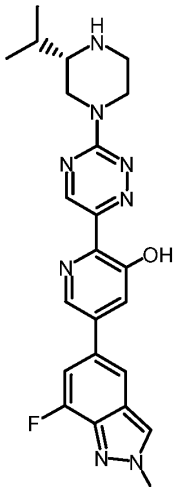
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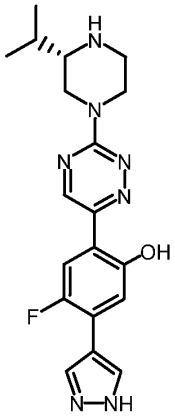
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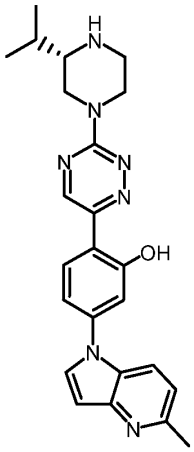
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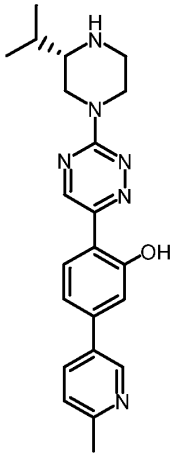
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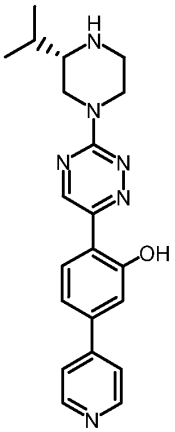
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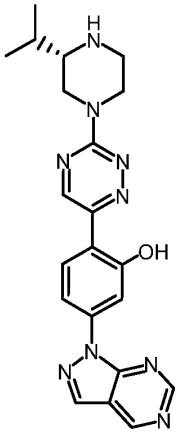
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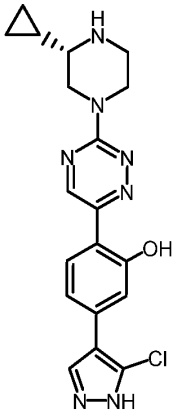
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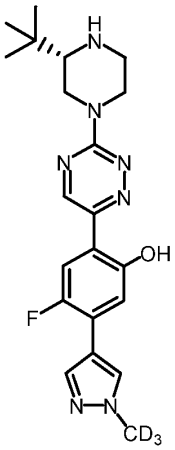
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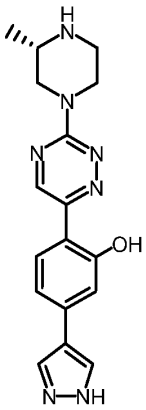
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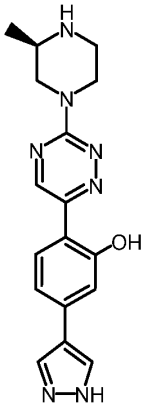
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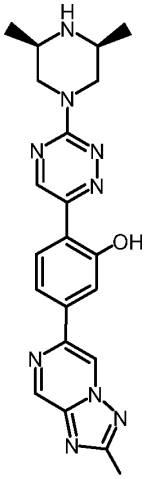
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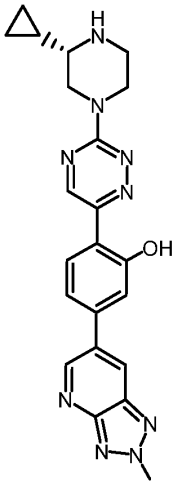
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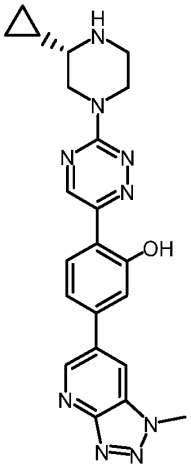
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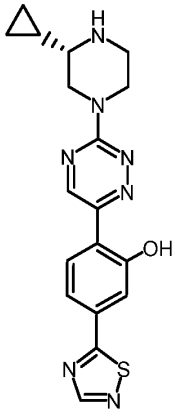
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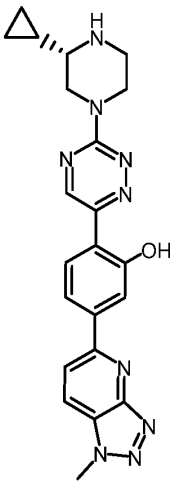
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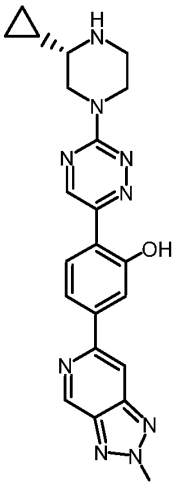
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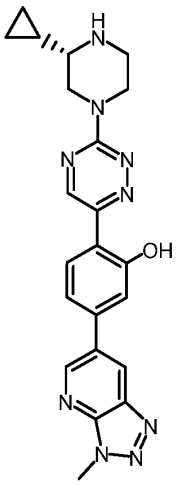
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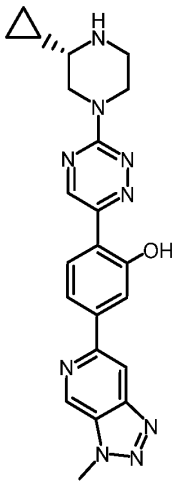
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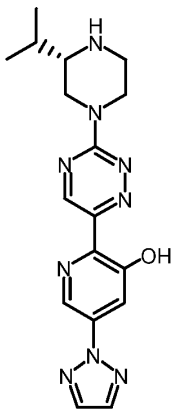
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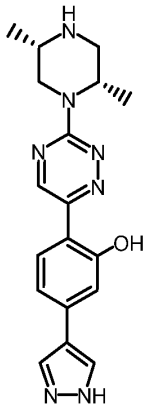
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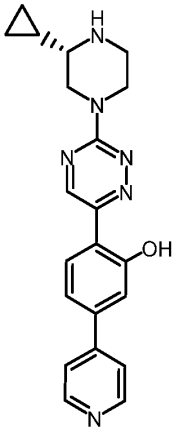
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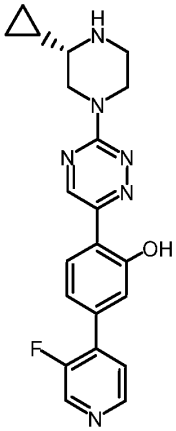
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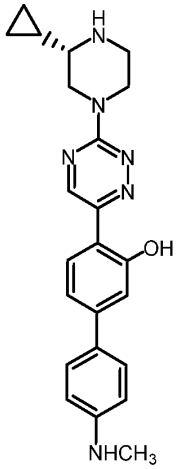
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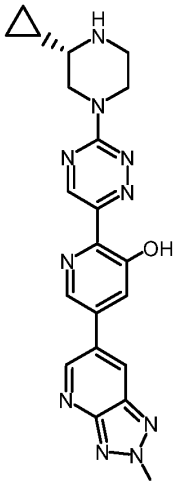
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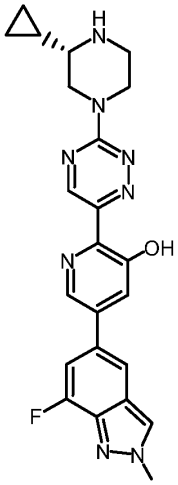
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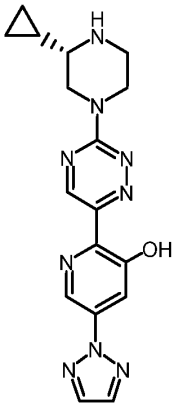
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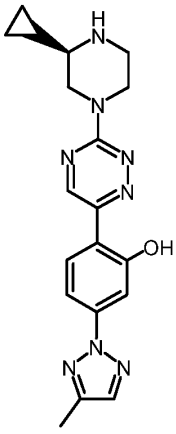
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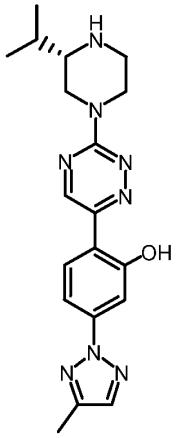
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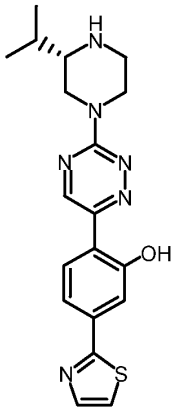
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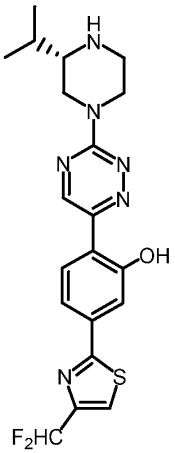
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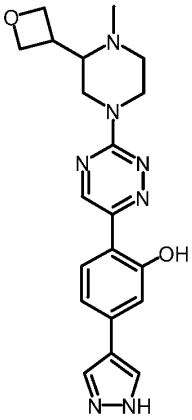
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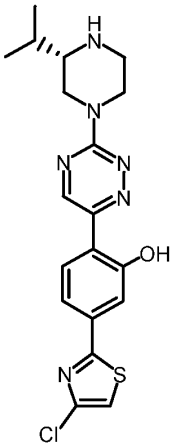
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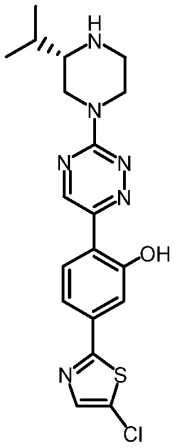
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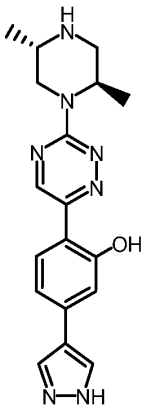
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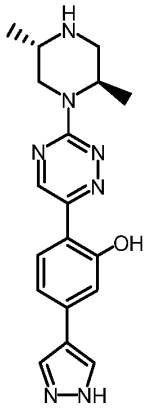
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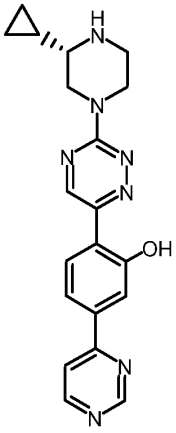
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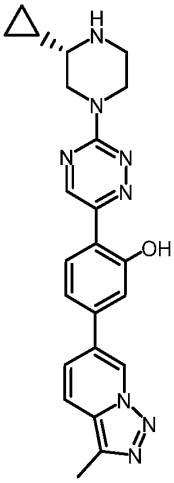
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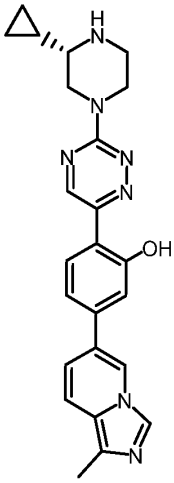
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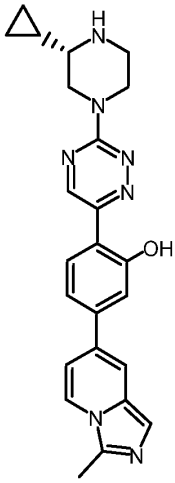
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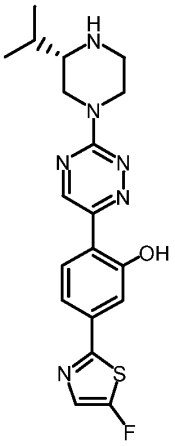
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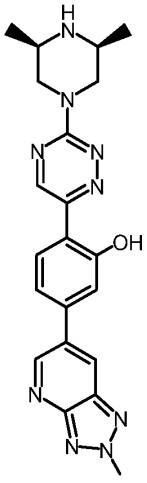
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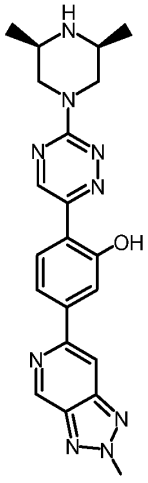
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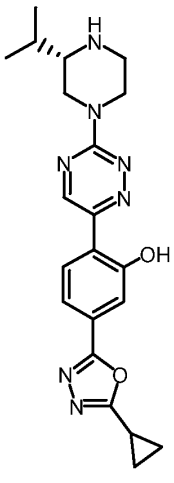
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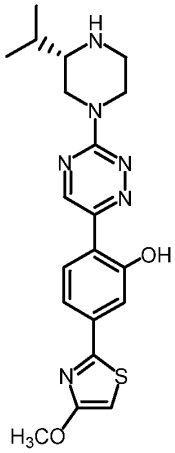
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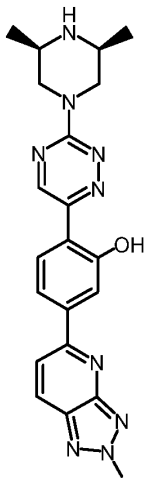
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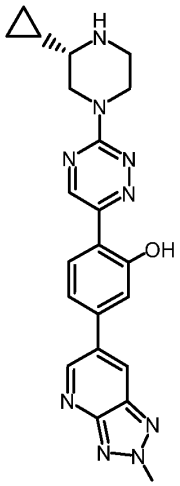
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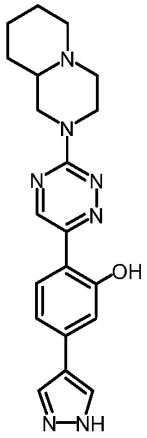
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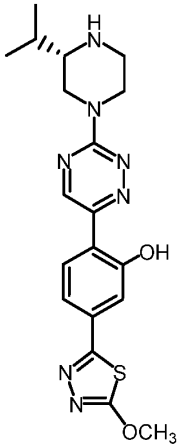
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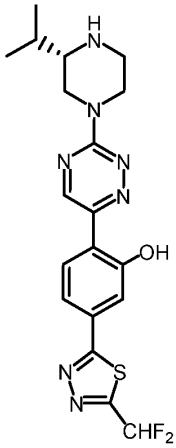
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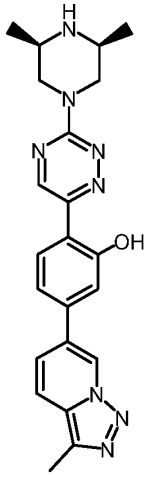
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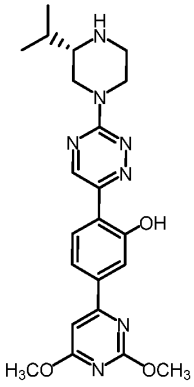
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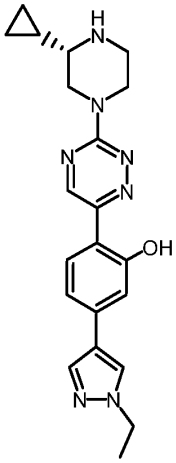
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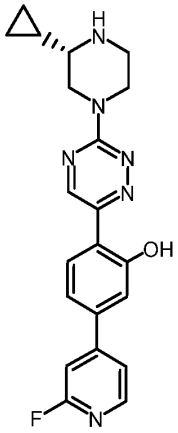
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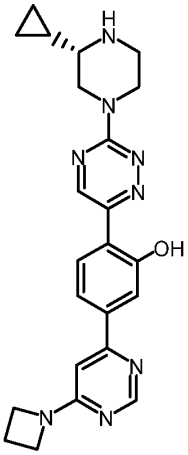
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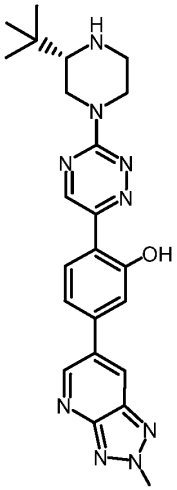
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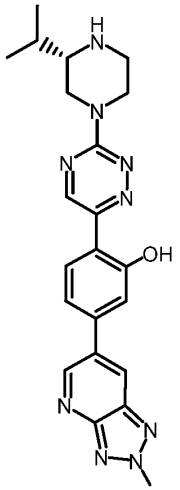
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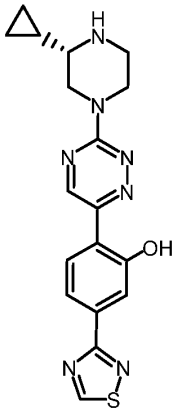
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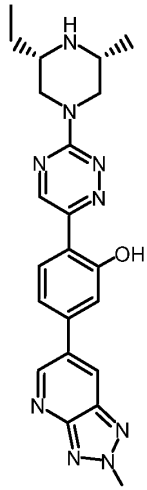
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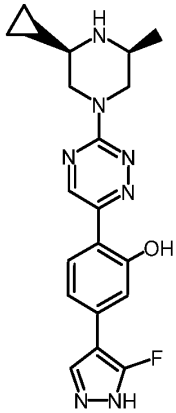
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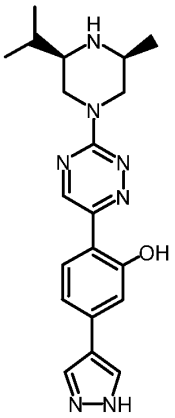
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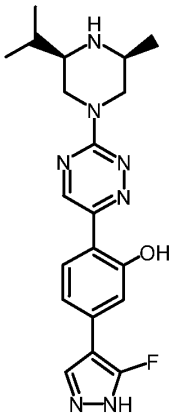
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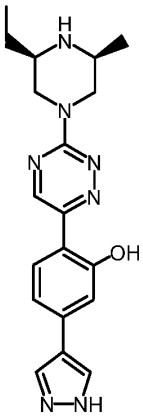
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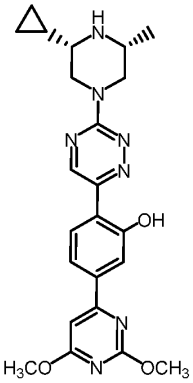
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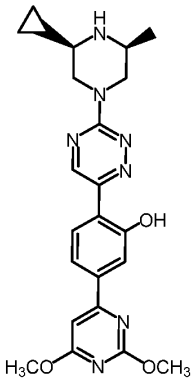
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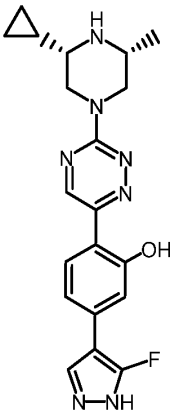
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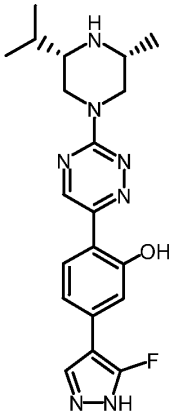
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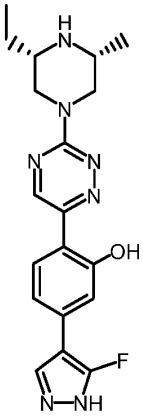
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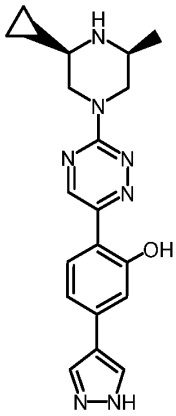
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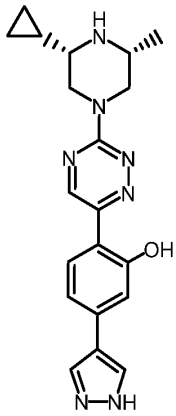
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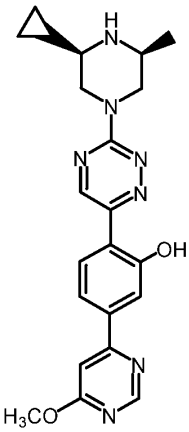
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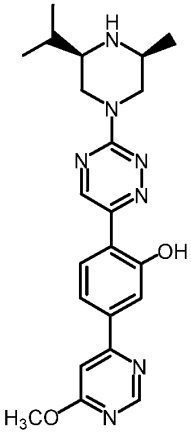
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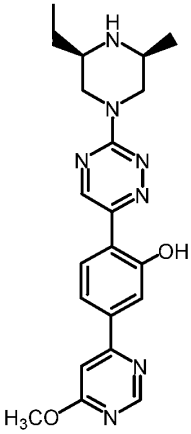
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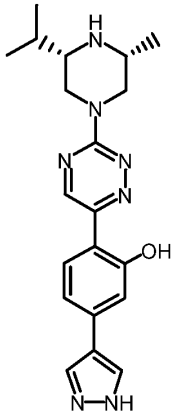
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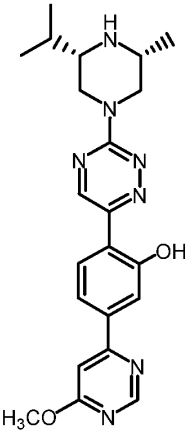
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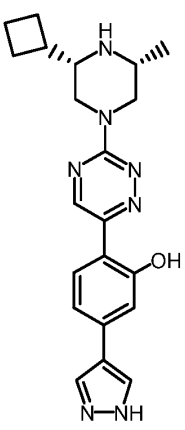
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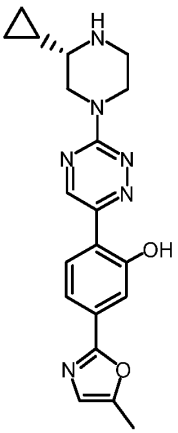
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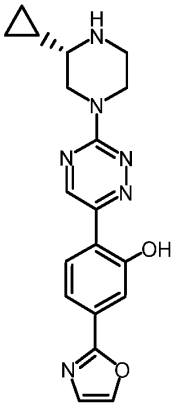
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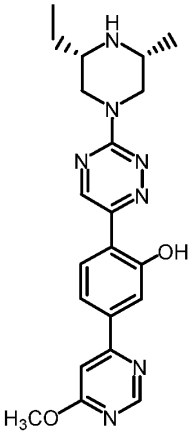
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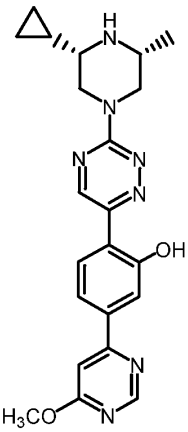
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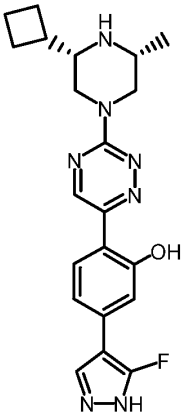
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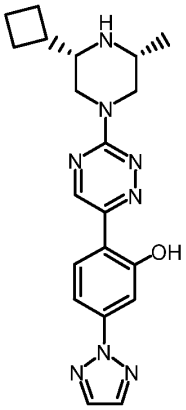
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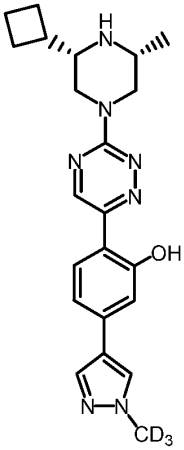
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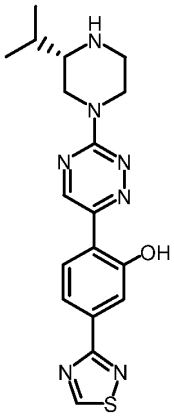
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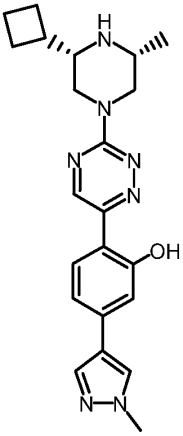
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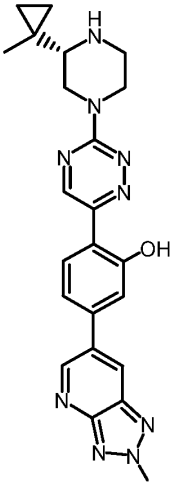
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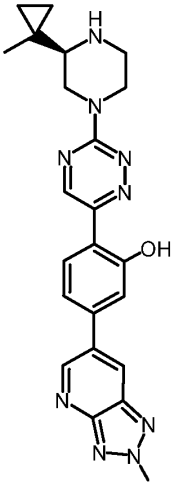
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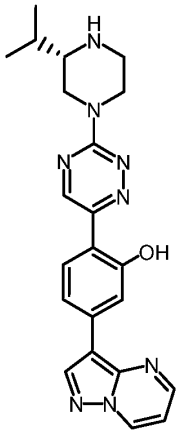
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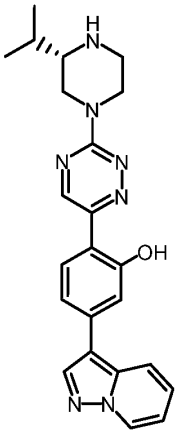
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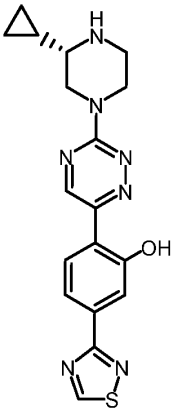
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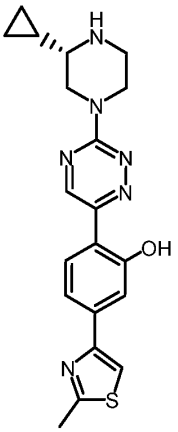
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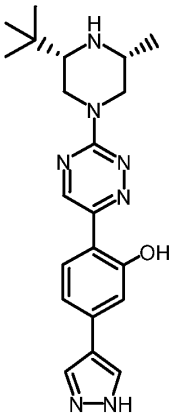
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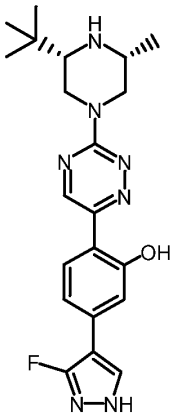
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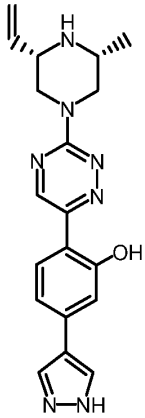
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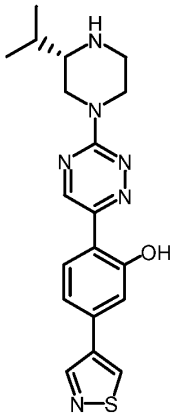
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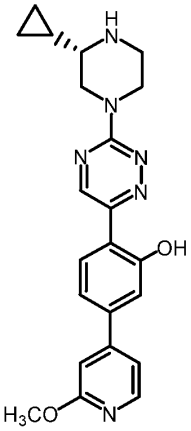
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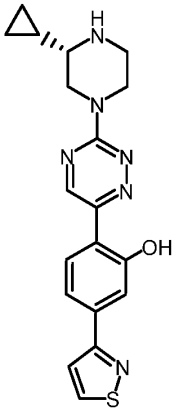
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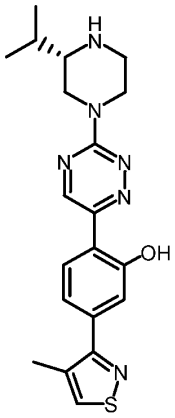
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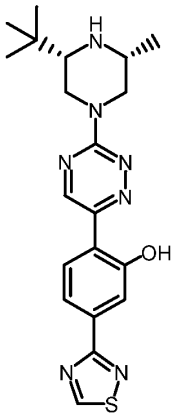
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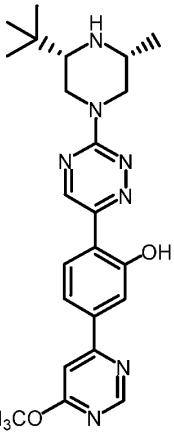
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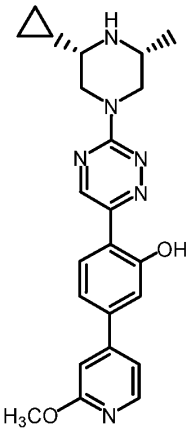
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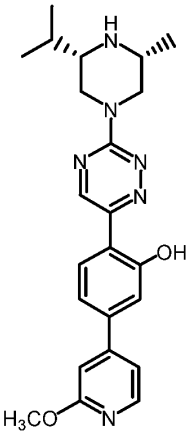
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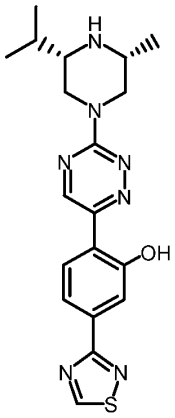
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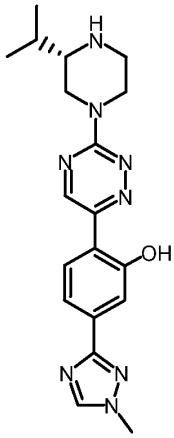
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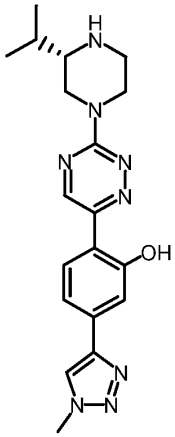
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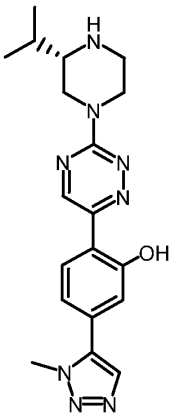
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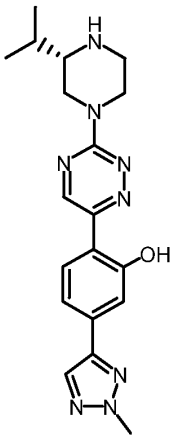
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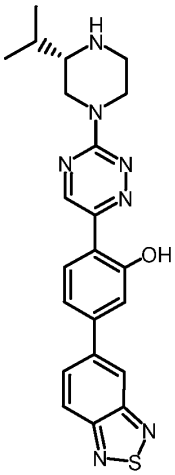
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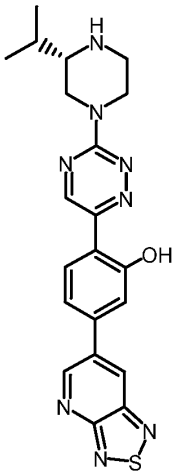
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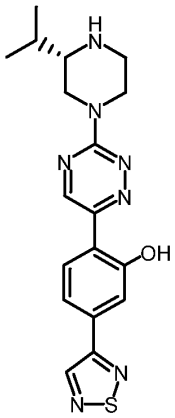
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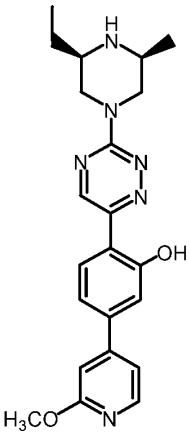
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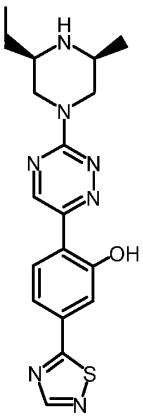
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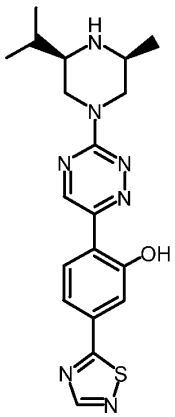
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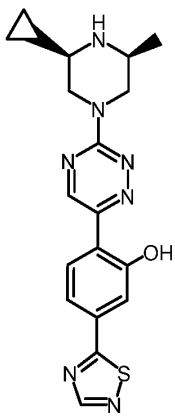
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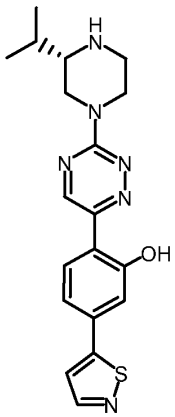
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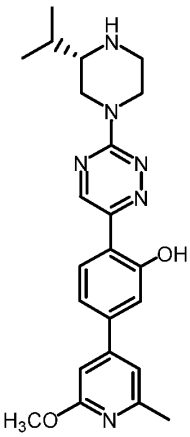
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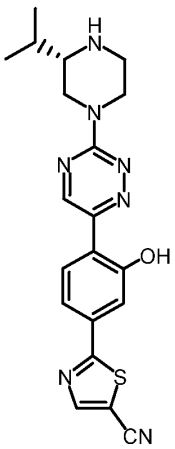
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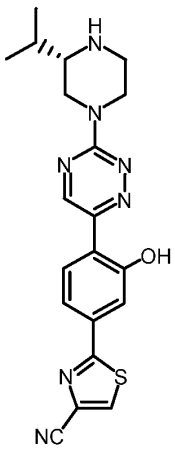
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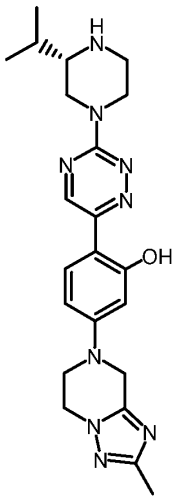
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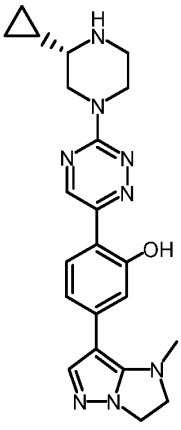
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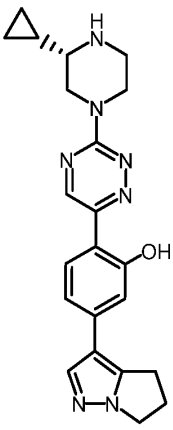
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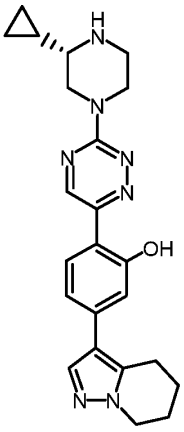
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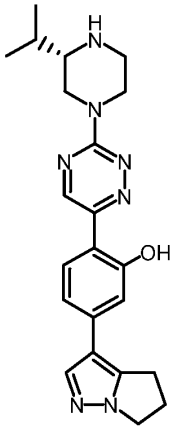
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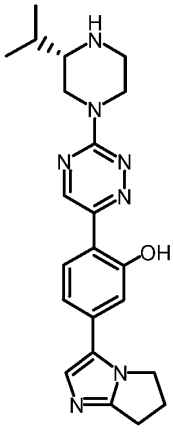
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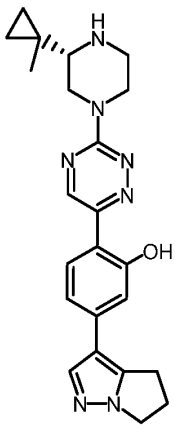
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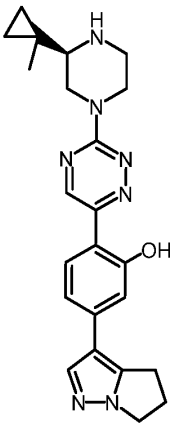
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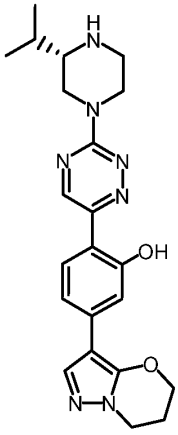
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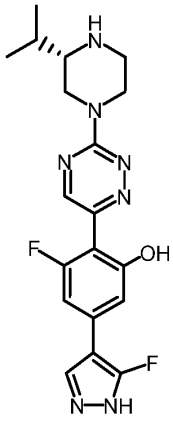
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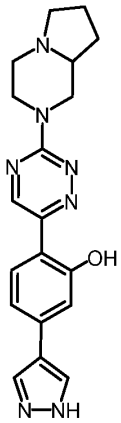
281



282



283, and



284;

wherein a form of the compound is selected from the group consisting of a salt, hydrate, ester, solvate, and tautomer form thereof.

An aspect the compound of Formula (I) or a form thereof (wherein compound number (#[1]) indicates that the salt form was isolated) includes a compound selected from the group consisting of:

Cpd	Name
1[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol
2[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-{6-[($^2H_3$)methyloxy]pyrimidin-4-yl}phenol
3[1]	5-(3-fluoro-1H-pyrazol-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
4[1]	5-(3-fluoro-1H-pyrazol-4-yl)-2-{3-[3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
5[1]	2-[3-(3-cyclopropylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol

Cpd	Name
6[1]	2-{3-[3-(1-hydroxycyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
7[1]	2-{3-[(3R)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol
8[1]	2-{3-[3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-{6-[($^2H_3$)methyloxy]pyrimidin-4-yl}phenol
9[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]phenol
10[1]	5-{6-[($^2H_3$)methyloxy]pyrimidin-4-yl}-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
11[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol
12[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-pyrazol-4-yl)phenol
13[1]	2-{3-[3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
14[1]	2-{3-[3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol
15[1]	5-(3-fluoro-1H-pyrazol-4-yl)-2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
16[1]	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
17[1]	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-{6-[($^2H_3$)methyloxy]pyrimidin-4-yl}phenol
18[1]	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol
19[1]	2-{3-[3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]phenol
20	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
21[1]	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-pyrazol-4-yl)phenol
22	2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
23	2-{3-[(3S)-3-ethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
24[1]	2-[3-(3-ethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
25[1]	5-(3-fluoro-1H-pyrazol-4-yl)-2-{3-[(3R)-3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
26[1]	2-{3-[3-(2-hydroxypropan-2-yl)-4-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-{6-[($^2H_3$)methyloxy]pyrimidin-4-yl}phenol
27[1]	2-[3-(3-cyclopropyl-4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol

Cpd	Name
28[1]	3-fluoro-5-(6-methoxypyrimidin-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
29[1]	2-{3-[3-(1-methoxycyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
30	2-[3-(3-cyclobutylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
31[1]	2-[3-(3-propylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
32[1]	2-[3-(3-cyclopropylpiperazin-1-yl)-1,2,4-triazin-6-yl]-3-fluoro-5-(5-fluoro-1H-pyrazol-4-yl)phenol
33[1]	2-{3-[3-(butan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
34	2-{3-[4-methyl-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
35[1]	5-(1-methyl-1H-pyrazol-4-yl)-2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
36[1]	2-{3-[3-(2,2-difluorocyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
37[1]	2-{3-[(3S)-3-propylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
38[1]	2-[3-(3-ethenylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
39[1]	2-[3-(3-ethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]phenol
40	2-{3-[(3R)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
41[1]	5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]-2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
42	2-[3-(3-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
43[1]	2-[3-(6,9-diazaspiro[4.5]decan-9-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
44[1]	5-(2-methylpyridin-4-yl)-2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
45[1]	2-{3-[(3S)-3-(hydroxymethyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol
46	2-{3-[3-(2-methylpropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
47[1]	5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
48	2-{3-[(3R)-3-ethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
49[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]phenol
50[1]	2-[3-(5,8-diazaspiro[3.5]nonan-8-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
51[1]	5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]-2-{3-[(3R,5S)-3,4,5-trimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol
52[1]	5-(2H-1,2,3-triazol-2-yl)-2-{3-[(3R,5S)-3,4,5-trimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol

Cpd	Name
53	2-{3-[(3R)-3-(methoxymethyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
54[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol
55[1]	2-[3-(4,7-diazaspiro[2.5]octan-7-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
56[1]	2-[3-(3,3-dimethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
57[1]	2-[3-(4,7-diazaspiro[2.5]octan-7-yl)-1,2,4-triazin-6-yl]-5-(3-fluoro-1H-pyrazol-4-yl)phenol
58[1]	2-[3-(8-methyl-3,8-diazabicyclo[3.2.1]octan-3-yl)-1,2,4-triazin-6-yl]-5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]phenol
59[1]	(7R,8aS)-2-{6-[2-hydroxy-4-(1H-pyrazol-4-yl)phenyl]-1,2,4-triazin-3-yl}octahydropyrrolo[1,2-a]pyrazin-7-ol
60	2-{3-[4-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
61[1]	2-[3-(3-phenylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
62[1]	5-(1H-pyrazol-4-yl)-2-{3-[3-(pyridin-4-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
63	2-[3-(4-cyclopropylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
64	2-[3-(hexahydropyrazino[2,1-c][1,4]oxazin-8(1H)-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
65[1]	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[3-(hydroxymethyl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
66[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)phenol
67[1]	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-[3-(3-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]phenol
68[1]	2-[3-(3-ethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)phenol
69[1]	5-(2,8-dimethylimidazo[1,2-b]pyridazin-6-yl)-2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol
70[1]	5-(8-methoxy-2-methyl[1,2,4]triazolo[1,5-b]pyridazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
71[1]	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
72	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-indazol-5-yl)phenol
73[1]	5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]phenol
74[1]	2-[3-(3-ethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(7-fluoro-2-methyl-2H-indazol-5-yl)phenol
75	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methylimidazo[1,2-b]pyridazin-6-yl)phenol

Cpd	Name
76	2-{3-[(3R,5R)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(7-fluoro-2-methyl-2H-indazol-5-yl)phenol
77[1]	2-[3-(4-ethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)phenol
78[1]	5-(2,8-dimethyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
79	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(7-fluoro-2-methyl-2H-indazol-5-yl)phenol
80[1]	5-(4-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-3-hydroxyphenyl)-2-methyl-2H-indazole-7-carbonitrile
81[1]	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
82	5-(2-methylimidazo[1,2-b]pyridazin-6-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]phenol
83[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2,8-dimethyl[1,2,4]triazolo[1,5-b]pyridazin-6-yl)phenol
84[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(6,8-dimethyl-7H-purin-2-yl)phenol
85[1]	5-(2-methyl[1,2,4]triazolo[1,5-a]pyridin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
86[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methylimidazo[1,2-a]pyrazin-6-yl)phenol
87	6-(4-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-3-hydroxyphenyl)-2-methylimidazo[1,2-a]pyridine-8-carbonitrile
88[1]	5-(2-methyl-2H-indazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
89[1]	2-{3-[(3R)-4-ethyl-3-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)phenol
90[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(8-methoxy-2-methyl[1,2,4]triazolo[1,5-b]pyridazin-6-yl)phenol
91[1]	5-(2-methyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
92[1]	5-(2,8-dimethyl[1,2,4]triazolo[1,5-b]pyridazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
93[1]	5-(8-methoxy-2-methyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
94[1]	5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)-2-{3-[(3R,5S)-3,4,5-trimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol
95[1]	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-[3-(hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl)-1,2,4-triazin-6-yl]phenol
96[1]	5-(2,8-dimethylimidazo[1,2-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol

Cpd	Name
97	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-pyrazolo[3,4-b]pyridin-5-yl)phenol
98[1]	5-(2,8-dimethylimidazo[1,2-b]pyridazin-6-yl)-2-{3-[(3R)-3-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol
99	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[(8aS)-hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl]-1,2,4-triazin-6-yl}phenol
100[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl[1,2,4]triazolo[1,5-b]pyridazin-6-yl)phenol
101[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2,8-dimethyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol
102[1]	5-(imidazo[1,2-b]pyridazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
103[1]	5-(2,8-dimethylimidazo[1,2-a]pyrazin-6-yl)-2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol
104[1]	5-(2,8-dimethylimidazo[1,2-b]pyridazin-6-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]phenol
105[1]	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[(8aR)-hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl]-1,2,4-triazin-6-yl}phenol
106[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol
107[1]	2-{3-[(3R)-3,4-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)phenol
108[1]	6-(3-hydroxy-4-{3-[(3R,5S)-3,4,5-trimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenyl)-2-methylimidazo[1,2-b]pyridazine-8-carbonitrile
109[1]	5-(8-cyclopropyl-2-methyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)-2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol
110	2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(2-methyl-2H-pyrazolo[3,4-b]pyridin-5-yl)phenol
111[1]	5-(2,8-dimethylimidazo[1,2-b]pyridazin-6-yl)-2-{3-[(3R,5S)-3,4,5-trimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol
112[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl[1,2,4]triazolo[1,5-a]pyrimidin-6-yl)phenol
113[1]	5-(imidazo[1,2-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
114[1]	5-(imidazo[1,2-a]pyridin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
115[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,2,4]triazolo[4,3-a]pyridin-6-yl)phenol
116[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(4,6-dimethyl[1,3]thiazolo[5,4-c]pyridin-2-yl)phenol
117[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5,7-dimethyl[1,2,4]triazolo[1,5-a]pyrimidin-2-yl)phenol

Cpd	Name
118[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-[2-(trifluoromethyl)imidazo[1,2-b]pyridazin-6-yl]phenol
119[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(6-methyl[1,3]thiazolo[4,5-b]pyrazin-2-yl)phenol
120[1]	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5-methylfuro[3,2-b]pyridin-2-yl)phenol
121[1]	5-(7-methoxy-2-methyl-2H-indazol-5-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]pyridin-3-ol
122[1]	5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]pyridin-3-ol
123[1]	2-{3-[(3S)-3-ethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol
124[1]	2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)pyridin-3-ol
125[1]	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol
126[1]	5-(2,8-dimethylimidazo[1,2-a]pyridin-6-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]pyridin-3-ol
127[1]	3-methyl-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
128[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol
129[1]	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol
130[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol
131[1]	5-(2,8-dimethylimidazo[1,2-a]pyridin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
132[1]	2-{3-[(3S)-3-ethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol
133[1]	2-{3-[(3S)-3-ethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-{6-[($^{2}H_3$)methyloxy]pyrimidin-4-yl}phenol
134[1]	2-{3-[3-(1-methylcyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
135[1]	2-[3-(3,8-diazabicyclo[3.2.1]octan-3-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol
136[1]	2-{3-[(3R)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
137[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
138[1]	5-(8-ethyl-2-methylimidazo[1,2-a]pyridin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
139[1]	5-[2-methyl-8-(trifluoromethyl)imidazo[1,2-a]pyridin-6-yl]-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol

Cpd	Name
140[1]	5-(2,7-dimethyl-2H-indazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
141[1]	5-(2-methylimidazo[1,2-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
142	5-(1H-imidazol-1-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
143	5-(6-methylpyrazin-2-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
144	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(pyrazin-2-yl)phenol
145	5-(5-methylpyrazin-2-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
146[1]	5-(1H-pyrazol-4-yl)-2-{3-[3-(2,2,2-trifluoroethyl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
147[1]	5-(2-methyl[1,2,4]triazolo[1,5-a]pyridin-7-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
148[1]	2-{3-[rac-(3S,5R)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
149[1]	5-(6-methylpyrimidin-4-yl)-2-{3-[(3RS)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
150	5-(6-ethylpyrimidin-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
151	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(pyrimidin-2-yl)phenol
152[1]	4-fluoro-5-[1-($^{2}H_3$)methyl-1H-pyrazol-4-yl]-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
153[1]	2-{3-[(8aS)-hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol
154[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol
155[1]	5-(5-methyl-1H-pyrazolo[4,3-b]pyridin-1-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
156[1]	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-4-fluoro-5-[1-($^{2}H_3$)methyl-1H-pyrazol-4-yl]phenol
157[1]	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}pyridin-3-ol
158[1]	4-fluoro-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
159[1]	5-(5-methyl-1H-pyrrolo[3,2-b]pyridin-1-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
160	5-(6-methylpyridin-3-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
161	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(pyridin-4-yl)phenol

Cpd	Name
162[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazolo[3,4-d]pyrimidin-1-yl)phenol
163[1]	5-(3-chloro-1H-pyrazol-4-yl)-2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol
164[1]	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-4-fluoro-5-(1H-pyrazol-4-yl)phenol
165[1]	2-{3-[(3S)-3-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
166[1]	2-{3-[(3R)-3-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
167	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol
168[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol
169[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol
170	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-5-yl)phenol
171[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-[1,2,3]triazolo[4,5-b]pyridin-5-yl)phenol
172[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl)phenol
173[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-methyl-3H-[1,2,3]triazolo[4,5-b]pyridin-5-yl)phenol
174[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-[1,2,3]triazolo[4,5-c]pyridin-6-yl)phenol
175[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)pyridin-3-ol
176[1]	2-{3-[(2S,5S)-2,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
177[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(pyridin-4-yl)phenol
178[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoropyridin-4-yl)phenol
179[1]	4-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-4'-(methylamino)[1,1'-biphenyl]-3-ol
180[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)pyridin-3-ol
181[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(7-fluoro-2-methyl-2H-indazol-5-yl)pyridin-3-ol
182[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)pyridin-3-ol
183[1]	2-{3-[(3R)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(4-methyl-2H-1,2,3-triazol-2-yl)phenol

Cpd	Name
184[1]	5-(4-methyl-2H-1,2,3-triazol-2-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
185	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,3-thiazol-2-yl)phenol
186	5-[4-(difluoromethyl)-1,3-thiazol-2-yl]-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
187[1]	2-{3-[4-methyl-3-(oxetan-3-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
188	5-(4-chloro-1,3-thiazol-2-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
189	5-(5-chloro-1,3-thiazol-2-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
190[1]	2-{3-[(2R,5S)-2,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol or 2-{3-[(2S,5R)-2,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
191[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,3]thiazolo[5,4-b]pyridin-2-yl)phenol
192	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(pyrimidin-4-yl)phenol
193[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-methyl[1,2,3]triazolo[1,5-a]pyridin-6-yl)phenol
194[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methylimidazo[1,5-a]pyridin-6-yl)phenol
195[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-methylimidazo[1,5-a]pyridin-7-yl)phenol
196	5-(5-fluoro-1,3-thiazol-2-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
197	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol
198	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl)phenol
199	5-(5-cyclopropyl-1,3,4-oxadiazol-2-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
200	5-(4-methoxy-1,3-thiazol-2-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
201	2-{3-[rac-(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl)phenol
202[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl)phenol
203	2-[3-(octahydro-2H-pyrido[1,2-a]pyrazin-2-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol

Cpd	Name
204	5-(5-methoxy-1,3,4-thiadiazol-2-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
205	5-[5-(difluoromethyl)-1,3,4-thiadiazol-2-yl]-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
206	2-{3-[rac-(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-methyl[1,2,3]triazolo[1,5-a]pyridin-6-yl)phenol
207	5-(2,6-dimethoxypyrimidin-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
208[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-ethyl-1H-pyrazol-4-yl)phenol
209	2-{3-[(3RS)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-fluoropyridin-4-yl)phenol
210	5-[6-(azetidin-1-yl)pyrimidin-4-yl]-2-{3-[(3RS)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol
211[1]	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol
212[1]	5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
213[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-3-yl)phenol
214	2-{3-[(3S,5R)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol
215	2-{3-[(3R,5S)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5-fluoro-1H-pyrazol-4-yl)phenol
216	2-{3-[(3S,5R)-3-methyl-5-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
217	5-(5-fluoro-1H-pyrazol-4-yl)-2-{3-[(3S,5R)-3-methyl-5-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
218	2-{3-[(3R,5S)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5-fluoro-1H-pyrazol-4-yl)phenol
219	2-{3-[(3S,5R)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2,6-dimethoxypyrimidin-4-yl)phenol
220	2-{3-[(3R,5S)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2,6-dimethoxypyrimidin-4-yl)phenol
221	2-{3-[(3S,5R)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5-fluoro-1H-pyrazol-4-yl)phenol
222	5-(5-fluoro-1H-pyrazol-4-yl)-2-{3-[(3R,5S)-3-methyl-5-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
223	2-{3-[(3S,5R)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5-fluoro-1H-pyrazol-4-yl)phenol
224	2-{3-[(3R,5S)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol

Cpd	Name
225	2-{3-[(3S,5R)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
226	2-{3-[(3R,5S)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(6-methoxypyrimidin-4-yl)phenol
227	5-(6-methoxypyrimidin-4-yl)-2-{3-[(3S,5R)-3-methyl-5-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
228	2-{3-[(3R,5S)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(6-methoxypyrimidin-4-yl)phenol
229	2-{3-[(3R,5S)-3-methyl-5-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
230	5-(6-methoxypyrimidin-4-yl)-2-{3-[(3R,5S)-3-methyl-5-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
231[1]	2-{3-[(3S,5R)-3-cyclobutyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
232[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5-methyl-1,3-oxazol-2-yl)phenol
233[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,3-oxazol-2-yl)phenol
234	2-{3-[(3S,5R)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(6-methoxypyrimidin-4-yl)phenol
235	2-{3-[(3S,5R)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(6-methoxypyrimidin-4-yl)phenol
236[1]	2-{3-[(3S,5R)-3-cyclobutyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol
237[1]	2-{3-[(3S,5R)-3-cyclobutyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol
238[1]	2-{3-[(3S,5R)-3-cyclobutyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-[1-($^{2}H_3$)methyl-1H-pyrazol-4-yl]phenol
239[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-3-yl)phenol
240[1]	2-{3-[(3S,5R)-3-cyclobutyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-pyrazol-3-yl)phenol
241[1]	2-{3-[(3S)-3-(1-methylcyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol
242[1]	2-{3-[(3R)-3-(1-methylcyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol
243[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(pyrazolo[1,5-a]pyrimidin-3-yl)phenol
244	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(pyrazolo[1,5-a]pyridin-3-yl)phenol
245[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5-methyl-1,2,4-thiadiazol-3-yl)phenol

Cpd	Name
246[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-1,3-thiazol-4-yl)phenol
247[1]	2-{3-[(3S,5R)-3-tert-butyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
248[1]	2-{3-[(3S,5R)-3-tert-butyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol
249[1]	2-{3-[(3S,5R)-3-ethenyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol
250[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2-thiazol-4-yl)phenol
251[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methoxypyridin-4-yl)phenol
252[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2-thiazol-3-yl)phenol
253	5-(4-methyl-1,2-thiazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
254[1]	2-{3-[(3S,5R)-3-tert-butyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-3-yl)phenol
255[1]	2-{3-[(3S,5R)-3-tert-butyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(6-methoxypyrimidin-4-yl)phenol
256	2-{3-[(3S,5R)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methoxypyridin-4-yl)phenol
257	5-(2-methoxypyridin-4-yl)-2-{3-[(3R,5S)-3-methyl-5-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
258	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-3-yl)phenol
259	5-(1-methyl-1H-1,2,4-triazol-3-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
260[1]	5-(1-methyl-1H-1,2,3-triazol-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
261[1]	5-(1-methyl-1H-1,2,3-triazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
262[1]	5-(2-methyl-2H-1,2,3-triazol-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
263[1]	5-(2,1,3-benzothiadiazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
264[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,2,5]thiadiazolo[3,4-b]pyridin-6-yl)phenol
265[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,5-thiadiazol-3-yl)phenol
266	2-{3-[(3S,5R)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methoxypyridin-4-yl)phenol or

Cpd	Name
	2-{3-[(3R,5S)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methoxypyridin-4-yl)phenol
267	2-{3-[(3S,5R)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-5-yl)phenol or 2-{3-[(3R,5S)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-5-yl)phenol
268	2-{3-[(3R,5S)-3-methyl-5-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-5-yl)phenol or 2-{3-[(3S,5R)-3-methyl-5-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-5-yl)phenol
269	2-{3-[(3S,5R)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-5-yl)phenol or 2-{3-[(3R,5S)-3-cyclopropyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-5-yl)phenol
270[1]	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2-thiazol-5-yl)phenol
271[1]	5-(2-methoxy-6-methylpyridin-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
272[1]	2-(3-hydroxy-4-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenyl)-1,3-thiazole-5-carbonitrile
273[1]	2-(3-hydroxy-4-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenyl)-1,3-thiazole-4-carbonitrile
274[1]	5-(2-methyl-5,6-dihydro[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
275[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-2,3-dihydro-1H-imidazo[1,2-b]pyrazol-7-yl)phenol
276[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl)phenol
277[1]	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(4,5,6,7-tetrahydropyrazolo[1,5-a]pyridin-3-yl)phenol or 2-{3-[(3R)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(4,5,6,7-tetrahydropyrazolo[1,5-a]pyridin-3-yl)phenol
278	5-(5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
279	5-(6,7-dihydro-5H-pyrrolo[1,2-a]imidazol-3-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
280	5-(5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl)-2-{3-[(3S)-3-(1-methylcyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
281	5-(5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl)-2-{3-[(3R)-3-(1-methylcyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol
282	5-(6,7-dihydro-5H-pyrazolo[5,1-b][1,3]oxazin-3-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol, and

Cpd	Name
283[1]	3-fluoro-5-(5-fluoro-1H-pyrazol-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol;

wherein a form of the compound is selected from the group consisting of a salt, hydrate, solvate, and tautomer form thereof.

Another aspect of the compound of Formula (I) or a form thereof is a compound salt selected from the group consisting of:

Cpd	Name
1	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol dihydrochloride
2	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-{6-[($^2H_3$)methyloxy]pyrimidin-4-yl}phenol dihydrochloride
3	5-(3-fluoro-1H-pyrazol-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
4	5-(3-fluoro-1H-pyrazol-4-yl)-2-{3-[3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
5	2-[3-(3-cyclopropylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol dihydrochloride
6	2-{3-[3-(1-hydroxycyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol diformate
7	2-{3-[(3R)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol dihydrochloride
8	2-{3-[3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-{6-[($^2H_3$)methyloxy]pyrimidin-4-yl}phenol dihydrochloride
9	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]phenol dihydrochloride
10	5-{6-[($^2H_3$)methyloxy]pyrimidin-4-yl}-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
11	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol dihydrochloride
12	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-pyrazol-4-yl)phenol dihydrochloride
13	2-{3-[3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride
14	2-{3-[3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol dihydrochloride
15	5-(3-fluoro-1H-pyrazol-4-yl)-2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
16	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride

Cpd	Name
17	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-{6-[($^2H_3$)methyloxy]pyrimidin-4-yl}phenol dihydrochloride
18	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol dihydrochloride
19	2-{3-[3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]phenol dihydrochloride
21	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-pyrazol-4-yl)phenol dihydrochloride
24	2-[3-(3-ethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol dihydrochloride
25	5-(3-fluoro-1H-pyrazol-4-yl)-2-{3-[(3R)-3-(2-hydroxypropan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
26	2-{3-[3-(2-hydroxypropan-2-yl)-4-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-{6-[($^2H_3$)methyloxy]pyrimidin-4-yl}phenol dihydrochloride
27	2-[3-(3-cyclopropyl-4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol dihydrochloride
28	3-fluoro-5-(6-methoxypyrimidin-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol formate
29	2-{3-[3-(1-methoxycyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol diformate
31	2-[3-(3-propylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol dihydrochloride
32	2-[3-(3-cyclopropylpiperazin-1-yl)-1,2,4-triazin-6-yl]-3-fluoro-5-(5-fluoro-1H-pyrazol-4-yl)phenol formate
33	2-{3-[3-(butan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride
35	5-(1-methyl-1H-pyrazol-4-yl)-2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
36	2-{3-[3-(2,2-difluorocyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride
38	2-[3-(3-ethenylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol dihydrochloride
39	2-[3-(3-ethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]phenol dihydrochloride
41	5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]-2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
43	2-[3-(6,9-diazaspiro[4.5]decan-9-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol dihydrochloride
44	5-(2-methylpyridin-4-yl)-2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
45	2-{3-[(3S)-3-(hydroxymethyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol dihydrochloride

Cpd	Name
47	5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
49	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]phenol dihydrochloride
50	2-[3-(5,8-diazaspiro[3.5]nonan-8-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol dihydrochloride
51	5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]-2-{3-[(3R,5S)-3,4,5-trimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
52	5-(2H-1,2,3-triazol-2-yl)-2-{3-[(3R,5S)-3,4,5-trimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
54	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol dihydrochloride
55	2-[3-(4,7-diazaspiro[2.5]octan-7-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol dihydrochloride
56	2-[3-(3,3-dimethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol dihydrochloride
57	2-[3-(4,7-diazaspiro[2.5]octan-7-yl)-1,2,4-triazin-6-yl]-5-(3-fluoro-1H-pyrazol-4-yl)phenol dihydrochloride
58	2-[3-(8-methyl-3,8-diazabicyclo[3.2.1]octan-3-yl)-1,2,4-triazin-6-yl]-5-[1-($^2H_3$)methyl-1H-pyrazol-4-yl]phenol dihydrochloride
59	(7R,8aS)-2-{6-[2-hydroxy-4-(1H-pyrazol-4-yl)phenyl]-1,2,4-triazin-3-yl}octahydropyrrolo[1,2-a]pyrazin-7-ol dihydrochloride
61	2-[3-(3-phenylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol dihydrochloride
62	5-(1H-pyrazol-4-yl)-2-{3-[3-(pyridin-4-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
65	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[3-(hydroxymethyl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
66	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)phenol dihydrochloride
67	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-[3-(3-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]phenol dihydrochloride
68	2-[3-(3-ethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)phenol dihydrochloride
69	5-(2,8-dimethylimidazo[1,2-b]pyridazin-6-yl)-2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
70	5-(8-methoxy-2-methyl[1,2,4]triazolo[1,5-b]pyridazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
71	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
73	5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]phenol dihydrochloride

Cpd	Name
74	2-[3-(3-ethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(7-fluoro-2-methyl-2H-indazol-5-yl)phenol dihydrochloride
77	2-[3-(4-ethylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)phenol dihydrochloride
78	5-(2,8-dimethyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
80	5-(4-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-3-hydroxyphenyl)-2-methyl-2H-indazole-7-carbonitrile dihydrochloride
81	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
83	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2,8-dimethyl[1,2,4]triazolo[1,5-b]pyridazin-6-yl)phenol dihydrochloride
84	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(6,8-dimethyl-7H-purin-2-yl)phenol diformate
85	5-(2-methyl[1,2,4]triazolo[1,5-a]pyridin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
86	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methylimidazo[1,2-a]pyrazin-6-yl)phenol dihydrochloride
88	5-(2-methyl-2H-indazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
89	2-{3-[(3R)-4-ethyl-3-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)phenol dihydrochloride
90	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(8-methoxy-2-methyl[1,2,4]triazolo[1,5-b]pyridazin-6-yl)phenol dihydrochloride
91	5-(2-methyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
92	5-(2,8-dimethyl[1,2,4]triazolo[1,5-b]pyridazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
93	5-(8-methoxy-2-methyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
94	5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)-2-{3-[(3R,5S)-3,4,5-trimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
95	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-[3-(hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl)-1,2,4-triazin-6-yl]phenol dihydrochloride
96	5-(2,8-dimethylimidazo[1,2-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
98	5-(2,8-dimethylimidazo[1,2-b]pyridazin-6-yl)-2-{3-[(3R)-3-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
100	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl[1,2,4]triazolo[1,5-b]pyridazin-6-yl)phenol dihydrochloride
101	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2,8-dimethyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol dihydrochloride

Cpd	Name
102	5-(imidazo[1,2-b]pyridazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
103	5-(2,8-dimethylimidazo[1,2-a]pyrazin-6-yl)-2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
104	5-(2,8-dimethylimidazo[1,2-b]pyridazin-6-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]phenol dihydrochloride
105	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[(8aR)-hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
106	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol dihydrochloride
107	2-{3-[(3R)-3,4-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)phenol dihydrochloride
108	6-(3-hydroxy-4-{3-[(3R,5S)-3,4,5-trimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenyl)-2-methylimidazo[1,2-b]pyridazine-8-carbonitrile dihydrochloride
109	5-(8-cyclopropyl-2-methyl[1,2,4]triazolo[1,5-a]pyrazin-6-yl)-2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
111	5-(2,8-dimethylimidazo[1,2-b]pyridazin-6-yl)-2-{3-[(3R,5S)-3,4,5-trimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
112	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl[1,2,4]triazolo[1,5-a]pyrimidin-6-yl)phenol dihydrochloride
113	5-(imidazo[1,2-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
114	5-(imidazo[1,2-a]pyridin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
115	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,2,4]triazolo[4,3-a]pyridin-6-yl)phenol dihydrochloride
116	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(4,6-dimethyl[1,3]thiazolo[5,4-c]pyridin-2-yl)phenol dihydrochloride
117	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5,7-dimethyl[1,2,4]triazolo[1,5-a]pyrimidin-2-yl)phenol dihydrochloride
118	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-[2-(trifluoromethyl)imidazo[1,2-b]pyridazin-6-yl]phenol dihydrochloride
119	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(6-methyl[1,3]thiazolo[4,5-b]pyrazin-2-yl)phenol dihydrochloride
120	2-{3-[(3R,5S)-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5-methylfuro[3,2-b]pyridin-2-yl)phenol dihydrochloride
121	5-(7-methoxy-2-methyl-2H-indazol-5-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]pyridin-3-ol hydrochloride
122	5-(8-fluoro-2-methylimidazo[1,2-a]pyridin-6-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]pyridin-3-ol hydrochloride
123	2-{3-[(3S)-3-ethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol dihydrochloride

Cpd	Name
124	2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)pyridin-3-ol hydrochloride
125	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol dihydrochloride
126	5-(2,8-dimethylimidazo[1,2-a]pyridin-6-yl)-2-[3-(4-methylpiperazin-1-yl)-1,2,4-triazin-6-yl]pyridin-3-ol hydrochloride
127	3-methyl-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol formate
128	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol dihydrochloride
129	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol dihydrochloride
130	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,2,4]triazolo[1,5-a]pyrazin-6-yl)phenol dihydrochloride
131	5-(2,8-dimethylimidazo[1,2-a]pyridin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
132	2-{3-[(3S)-3-ethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol dihydrochloride
133	2-{3-[(3S)-3-ethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-{6-[($^{2}H_3$)methyloxy]pyrimidin-4-yl}phenol dihydrochloride
134	2-{3-[3-(1-methylcyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol formate
135	2-[3-(3,8-diazabicyclo[3.2.1]octan-3-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol diformate
136	2-{3-[(3R)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride
137	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride
138	5-(8-ethyl-2-methylimidazo[1,2-a]pyridin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
139	5-[2-methyl-8-(trifluoromethyl)imidazo[1,2-a]pyridin-6-yl]-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
140	5-(2,7-dimethyl-2H-indazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
141	5-(2-methylimidazo[1,2-a]pyrazin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
146	5-(1H-pyrazol-4-yl)-2-{3-[3-(2,2,2-trifluoroethyl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol diformate
147	5-(2-methyl[1,2,4]triazolo[1,5-a]pyridin-7-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
148	2-{3-[rac-(3S,5R)-3-ethyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride

Cpd	Name
149	5-(6-methylpyrimidin-4-yl)-2-{3-[(3RS)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
152	4-fluoro-5-[1-($^{2}H_3$)methyl-1H-pyrazol-4-yl]-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol formate
153	2-{3-[(8aS)-hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol dihydrochloride
154	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol dihydrochloride
155	5-(5-methyl-1H-pyrazolo[4,3-b]pyridin-1-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
156	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-4-fluoro-5-[1-($^{2}H_3$)methyl-1H-pyrazol-4-yl]phenol formate
157	5-(7-fluoro-2-methyl-2H-indazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}pyridin-3-ol hydrochloride
158	4-fluoro-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol formate
159	5-(5-methyl-1H-pyrrolo[3,2-b]pyridin-1-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol formate
162	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazolo[3,4-d]pyrimidin-1-yl)phenol dihydrochloride
163	5-(3-chloro-1H-pyrazol-4-yl)-2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
164	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-4-fluoro-5-(1H-pyrazol-4-yl)phenol formate
165	2-{3-[(3S)-3-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride
166	2-{3-[(3R)-3-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride
168	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol dihydrochloride
169	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol dihydrochloride
171	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-[1,2,3]triazolo[4,5-b]pyridin-5-yl)phenol dihydrochloride
172	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-c]pyridin-6-yl)phenol dihydrochloride
173	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-methyl-3H-[1,2,3]triazolo[4,5-b]pyridin-5-yl)phenol dihydrochloride
174	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-[1,2,3]triazolo[4,5-c]pyridin-6-yl)phenol dihydrochloride
175	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)pyridin-3-ol dihydrochloride

Cpd	Name
176	2-{3-[(2S,5S)-2,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride
177	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(pyridin-4-yl)phenol dihydrochloride
178	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoropyridin-4-yl)phenol dihydrochloride
179	4-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-4'-(methylamino)[1,1'-biphenyl]-3-ol dihydrochloride
180	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)pyridin-3-ol trifluoroacetate
181	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(7-fluoro-2-methyl-2H-indazol-5-yl)pyridin-3-ol hydrochloride
182	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)pyridin-3-ol trifluoroacetate
183	2-{3-[(3R)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(4-methyl-2H-1,2,3-triazol-2-yl)phenol dihydrochloride
184	5-(4-methyl-2H-1,2,3-triazol-2-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
187	2-{3-[4-methyl-3-(oxetan-3-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol formate
190	2-{3-[(2R,5S)-2,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol or enantiomer trifluoroacetate
191	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,3]thiazolo[5,4-b]pyridin-2-yl)phenol trifluoroacetate
193	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-methyl[1,2,3]triazolo[1,5-a]pyridin-6-yl)phenol formate
194	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methylimidazo[1,5-a]pyridin-6-yl)phenol formate
195	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-methylimidazo[1,5-a]pyridin-7-yl)phenol formate
202	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-5-yl)phenol dihydrochloride
208	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-ethyl-1H-pyrazol-4-yl)phenol dihydrochloride
211	2-{3-[(3S)-3-tert-butylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol dihydrochloride
212	5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
213	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-3-yl)phenol trifluoroacetate
231	2-{3-[(3S,5R)-3-cyclobutyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride

Cpd	Name
232	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5-methyl-1,3-oxazol-2-yl)phenol trifluoroacetate
233	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,3-oxazol-2-yl)phenol trifluoroacetate
236	2-{3-[(3S,5R)-3-cyclobutyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol dihydrochloride
237	2-{3-[(3S,5R)-3-cyclobutyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2H-1,2,3-triazol-2-yl)phenol dihydrochloride
238	2-{3-[(3S,5R)-3-cyclobutyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-[1-($^{2}H_{3}$)methyl-1H-pyrazol-4-yl]phenol hydrochloride
239	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-3-yl)phenol trifluoroacetate
240	2-{3-[(3S,5R)-3-cyclobutyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-1H-pyrazol-3-yl)phenol hydrochloride
241	2-{3-[(3S)-3-(1-methylcyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol hydrochloride
242	2-{3-[(3R)-3-(1-methylcyclopropyl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)phenol hydrochloride
243	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(pyrazolo[1,5-a]pyrimidin-3-yl)phenol trifluoroacetate
245	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5-methyl-1,2,4-thiadiazol-3-yl)phenol trifluoroacetate
246	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methyl-1,3-thiazol-4-yl)phenol trifluoroacetate
247	2-{3-[(3S,5R)-3-tert-butyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride
248	2-{3-[(3S,5R)-3-tert-butyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(3-fluoro-1H-pyrazol-4-yl)phenol dihydrochloride
249	2-{3-[(3S,5R)-3-ethenyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1H-pyrazol-4-yl)phenol dihydrochloride
250	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2-thiazol-4-yl)phenol hydrochloride
251	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(2-methoxypyridin-4-yl)phenol trifluoroacetate
252	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2-thiazol-3-yl)phenol trifluoroacetate
254	2-{3-[(3S,5R)-3-tert-butyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,4-thiadiazol-3-yl)phenol hydrochloride
255	2-{3-[(3S,5R)-3-tert-butyl-5-methylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(6-methoxypyrimidin-4-yl)phenol hydrochloride
260	5-(1-methyl-1H-1,2,3-triazol-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride

Cpd	Name
261	5-(1-methyl-1H-1,2,3-triazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
262	5-(2-methyl-2H-1,2,3-triazol-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
263	5-(2,1,3-benzothiadiazol-5-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol hydrochloride
264	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-([1,2,5]thiadiazolo[3,4-b]pyridin-6-yl)phenol dihydrochloride
265	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2,5-thiadiazol-3-yl)phenol trifluoroacetate
270	2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1,2-thiazol-5-yl)phenol dihydrochloride
271	5-(2-methoxy-6-methylpyridin-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol dihydrochloride
272	2-(3-hydroxy-4-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenyl)-1,3-thiazole-5-carbonitrile hydrochloride
273	2-(3-hydroxy-4-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenyl)-1,3-thiazole-4-carbonitrile hydrochloride
274	5-(2-methyl-5,6-dihydro[1,2,4]triazolo[1,5-a]pyrazin-7(8H)-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol formate
275	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(1-methyl-2,3-dihydro-1H-imidazo[1,2-b]pyrazol-7-yl)phenol dihydrochloride
276	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(5,6-dihydro-4H-pyrrolo[1,2-b]pyrazol-3-yl)phenol dihydrochloride
277	2-{3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(4,5,6,7-tetrahydropyrazolo[1,5-a]pyridin-3-yl)phenol dihydrochloride or 2-{3-[(3R)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl}-5-(4,5,6,7-tetrahydropyrazolo[1,5-a]pyridin-3-yl)phenol or enantiomer dihydrochloride
283	3-fluoro-5-(5-fluoro-1H-pyrazol-4-yl)-2-{3-[(3S)-3-(propan-2-yl)piperazin-1-yl]-1,2,4-triazin-6-yl}phenol formate, and
284	2-[3-(hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl)-1,2,4-triazin-6-yl]-5-(1H-pyrazol-4-yl)phenol;

wherein a form of the compound is selected from the group consisting of a hydrate, solvate, and tautomer form thereof.

CHEMICAL DEFINITIONS

The chemical terms used above and throughout the description herein, unless specifically defined otherwise, shall be understood by one of ordinary skill in the art to have the following indicated meanings.

As used herein, the term "C_{1-44}alkyl" generally refers to saturated hydrocarbon radicals having from one to four carbon atoms in a straight or branched chain configuration, including, but not limited to, methyl, ethyl, n-propyl (also referred to as propyl or propanyl), isopropyl, n-butyl (also referred to as butyl or butanyl), isobutyl, sec-butyl, tert-butyl and the like. A C_{1-44}alkyl radical is optionally substituted with substituent species as described herein where allowed by available valences.

As used herein, the term "C_{2-4}alkenyl" generally refers to partially unsaturated hydrocarbon radicals having from two to four carbon atoms in a straight or branched chain configuration and one or more carbon-carbon double bonds therein, including, but not limited to, ethenyl (also referred to as vinyl), allyl, propenyl and the like. A C_{2-4}alkenyl radical is optionally substituted with substituent species as described herein where allowed by available valences.

As used herein, the term "C_{2-8}alkynyl" generally refers to partially unsaturated hydrocarbon radicals having from two to eight carbon atoms in a straight or branched chain configuration and one or more carbon-carbon triple bonds therein, including, but not limited to, ethynyl, propynyl, butynyl and the like. In certain aspects, C_{2-8}alkynyl includes, but is not limited to, C_{2-6}alkynyl, C_{2-4}alkynyl and the like. A C_{2-8}alkynyl radical is optionally substituted with substituent species as described herein where allowed by available valences.

As used herein, the term "C_{1-4}alkoxy" generally refers to saturated hydrocarbon radicals having from one to four carbon atoms in a straight or branched chain configuration of the formula: -O-C_{1-4}alkyl, including, but not limited to, methoxy, ethoxy, n-propoxy, isopropoxy, n-butoxy, isobutoxy, sec-butoxy, tert-butoxy and the like. A C_{1-44}alkoxy radical is optionally substituted with substituent species as described herein where allowed by available valences.

As used herein, the term "C_{3-6}cycloalkyl" generally refers to a saturated or partially unsaturated monocyclic, bicyclic or polycyclic hydrocarbon radical, including, but not limited to, cyclopropyl, cyclobutyl, cyclopentyl, and cyclohexyl. A C_{3-6}cycloalkyl radical is optionally substituted with substituent species as described herein where allowed by available valences.

As used herein, the term "aryl" generally refers to a monocyclic, bicyclic or polycyclic aromatic carbon atom ring structure radical, including, but not limited to, phenyl, naphthyl, anthracenyl, fluorenyl, azulenyl, phenanthrenyl and the like. An aryl radical is optionally substituted with substituent species as described herein where allowed by available valences.

As used herein, the term "heteroaryl" generally refers to a monocyclic, bicyclic or polycyclic aromatic carbon atom ring structure radical in which one or more carbon atom ring members have been replaced, where allowed by structural stability, with one or more heteroatoms, such as an O, S or N atom, including, but not limited to, furanyl, thienyl, pyrrolyl, pyrazolyl, imidazolyl, isoxazolyl, isothiazolyl, oxazolyl, 1,3-thiazolyl, triazolyl, oxadiazolyl, thiadiazolyl, tetrazolyl, pyridinyl, pyrimidinyl, pyrazinyl, pyridazinyl, triazinyl, indolyl, indazolyl, indolizinyl, isoindolyl, benzofuranyl, benzothienyl, benzoimidazolyl, 1,3-benzothiazolyl, 1,3-benzoxazolyl, purinyl, quinolinyl, isoquinolinyl, quinazolinyl, quinoxalinyl, 1,3-diazinyl, 1,2-diazinyl, 1,2-diazolyl, 1,4-diazanaphthalenyl, acridinyl, furo[3,2-*b*]pyridinyl, furo[3,2-*c*]pyridinyl, furo[2,3-*c*]pyridinyl, 6*H*-thieno[2,3-*b*]pyrrolyl, thieno[3,2-*c*]pyridinyl, thieno[2,3-*d*]pyrimidinyl, 1*H*-pyrrolo[2,3-*b*]pyridinyl, 1*H*-pyrrolo[2,3-*c*]pyridinyl, 1*H*-pyrrolo[3,2-*b*]pyridinyl, pyrrolo[1,2-*a*]pyrazinyl, pyrrolo[1,2-*b*]pyridazinyl, pyrazolo[1,5-*a*]pyridinyl, pyrazolo[1,5-*a*]pyrazinyl, imidazo[1,2-*a*]pyridinyl, 3*H*-imidazo[4,5-*b*]pyridinyl, imidazo[1,2-*a*]pyrimidinyl, imidazo[1,2-*c*]pyrimidinyl, imidazo[1,2-*b*]pyridazinyl, imidazo[1,2-*a*]pyrazinyl, imidazo[2,1-*b*][1,3]thiazolyl, imidazo[2,1-*b*][1,3,4]thiadiazolyl, [1,2,4]triazolo[1,5-*a*]pyridinyl, [1,2,4]triazolo[4,3-*a*]pyridinyl and the like. A heteroaryl radical is optionally substituted on a carbon or nitrogen atom ring member with substituent species as described herein where allowed by available valences.

In certain aspects, the nomenclature for a heteroaryl radical may differ, such as in non-limiting examples where furanyl may also be referred to as furyl, thiophenyl may also be referred to as thienyl, pyridinyl may also be referred to as pyridyl, benzothiophenyl may also be referred to as benzothienyl and 1,3-benzoxazolyl may also be referred to as 1,3-benzooxazolyl.

In certain other aspects, the term for a heteroaryl radical may also include other regioisomers, such as in non-limiting examples where the term pyrrolyl may also include 2*H*-pyrrolyl, 3*H*-pyrrolyl and the like, the term pyrazolyl may also include 1*H*-pyrazolyl and the like, the term imidazolyl may also include 1*H*-imidazolyl and the like, the term triazolyl may also include 1*H*-1,2,3-triazolyl and the like, the term oxadiazolyl may also include 1,2,4-oxadiazolyl, 1,3,4-oxadiazolyl and the like, the term tetrazolyl may also include 1*H*-tetrazolyl, 2*H*-tetrazolyl and the like, the term indolyl may also include 1*H*-indolyl and the like, the term indazolyl may also include 1*H*-indazolyl, 2*H*-indazolyl and the like, the term benzoimidazolyl may also include 1*H*-benzoimidazolyl and the term purinyl may also include 9*H*-purinyl and the like.

As used herein, the term "heterocyclyl" generally refers to a saturated or partially unsaturated monocyclic, bicyclic or polycyclic carbon atom ring structure radical in which one or more carbon atom ring members have been replaced, where allowed by structural stability, with a heteroatom, such as an O, S or N atom, including, but not limited to, oxiranyl, oxetanyl, azetidinyl, tetrahydrofuranyl, pyrrolinyl, pyrrolidinyl, pyrazolinyl, pyrazolidinyl, imidazolinyl, imidazolidinyl, isoxazolinyl, isoxazolidinyl, isothiazolinyl, isothiazolidinyl, oxazolinyl, oxazolidinyl, thiazolinyl, thiazolidinyl, triazolinyl, triazolidinyl, oxadiazolinyl, oxadiazolidinyl, thiadiazolinyl, thiadiazolidinyl, tetrazolinyl, tetrazolidinyl, pyranyl, dihydro-2*H*-pyranyl, thiopyranyl, 1,3-dioxanyl, 1,2,5,6-tetrahydropyridinyl, 1,2,3,6-tetrahydropyridinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, 1,4-diazepanyl, 1,3-benzodioxolyl, 1,4-benzodioxanyl, 2,3-dihydro-1,4-benzodioxinyl, hexahydropyrrolo[3,4-*b*]pyrrol-(1*H*)-yl, (3a*S*,6a*S*)-hexahydropyrrolo[3,4-*b*]pyrrol-(1*H*)-yl, (3a*R*,6a*R*)-hexahydropyrrolo[3,4-*b*]pyrrol-(1*H*)-yl, hexahydropyrrolo[3,4-*b*]pyrrol-(2*H*)-yl, (3a*S*,6a*S*)-hexahydropyrrolo[3,4-*b*]pyrrol-(2*H*)-yl, (3a*R*,6a*R*)-hexahydropyrrolo[3,4-*b*]pyrrol-(2*H*)-yl, hexahydropyrrolo[3,4-*c*]pyrrol-(1*H*)-yl, (3a*R*,6a*S*)-hexahydropyrrolo[3,4-*c*]pyrrol-(1*H*)-yl, (3a*R*,6a*R*)-hexahydropyrrolo[3,4-*c*]pyrrol-(1*H*)-yl, octahydro-5*H*-pyrrolo[3,2-*c*]pyridinyl, octahydro-6*H*-pyrrolo[3,4-*b*]pyridinyl, (4a*R*,7a*R*)-octahydro-6*H*-pyrrolo[3,4-*b*]pyridinyl, (4a*S*,7a*S*)-octahydro-6*H*-pyrrolo[3,4-*b*]pyridinyl, hexahydropyrrolo[1,2-*a*]pyrazin-(1*H*)-yl, (7*R*,8a*S*)-hexahydropyrrolo[1,2-*a*]pyrazin-(1*H*)-yl, (8a*S*)-hexahydropyrrolo[1,2-*a*]pyrazin-(1*H*)-yl, (8a*R*)-hexahydropyrrolo[1,2-*a*]pyrazin-(1*H*)-yl, (8a*S*)-octahydropyrrolo[1,2-*a*]pyrazin-(1*H*)-yl, (8a*R*)-octahydropyrrolo[1,2-*a*]pyrazin-(1*H*)-yl, hexahydropyrrolo[1,2-*a*]pyrazin-(2*H*)-one, octahydro-2*H*-pyrido[1,2-*a*]pyrazinyl, 3-azabicyclo[3.1.0]hexyl, (1*R*,5*S*)-3-azabicyclo[3.1.0]hexyl, 8-azabicyclo[3.2.1]octyl, (1*R*,5*S*)-8-azabicyclo[3.2.1]octyl, 8-azabicyclo[3.2.1]oct-2-enyl, (1*R*,5*S*)-8-azabicyclo[3.2.1]oct-2-enyl, 9-azabicyclo[3.3.1]nonyl, (1*R*,5*S*)-9-azabicyclo[3.3.1]nonyl, 2,5-diazabicyclo[2.2.1]heptyl, (1*S*,4*S*)-2,5-diazabicyclo[2.2.1]heptyl, 2,5-diazabicyclo[2.2.2]octyl, 3,8-diazabicyclo[3.2.1]octyl, (1*R*,5*S*)-3,8-diazabicyclo[3.2.1]octyl, 1,4-diazabicyclo[3.2.2]nonyl, azaspiro[3.3]heptyl, 2,6-diazaspiro[3.3]heptyl, 2,6-diazaspiro[3.4]octyl, 2,7-diazaspiro[3.5]nonyl, 5,8-diazaspiro[3.5]nonyl, 2,7-diazaspiro[4.4]nonyl, 6,9-diazaspiro[4.5]decyl, 7-azadispiro[5.1.5^8.3^6]hexadecanyl and the

like. A heterocyclyl radical is optionally substituted on a carbon or nitrogen atom ring member with substituent species as described herein where allowed by available valences.

In certain aspects, the nomenclature for a heterocyclyl radical may differ, such as in non-limiting examples where 1,3-benzodioxolyl may also be referred to as benzo[*d*][1,3]dioxolyl and 2,3-dihydro-1,4-benzodioxinyl may also be referred to as 2,3-dihydrobenzo[*b*][1,4]dioxinyl.

As used herein, the term "C_{1-4}alkoxy-C_{1-4}alkyl" refers to a radical of the formula: -C_{1-4}alkyl-O-C_{1-4}alkyl.

As used herein, the term "C_{1-4}alkyl-amino" refers to a radical of the formula: -NH-C_{1-4}alkyl.

As used herein, the term "(C_{1-4}alkyl)$_2$-amino" refers to a radical of the formula: -N(C_{1-4}alkyl)$_2$.

As used herein, the term "C_{1-4}alkyl-carbonyl" refers to a radical of the formula: -C(O)-C_{1-4}alkyl.

As used herein, the term "C_{1-4}alkyl-carbonyl-amino" refers to a radical of the formula: -NH-C(O)-C_{1-4}alkyl.

As used herein, the term "C_{1-4}alkyl-thio" refers to a radical of the formula: -S-C_{1-4}alkyl.

As used herein, the term "amino-C_{1-4}alkyl" refers to a radical of the formula: -C_{1-4}alkyl-NH_2.

As used herein, the term "deutero-C_{1-4}alkyl," refers to a radical of the formula: -C_{1-4}alkyl-deutero, wherein C_{1-4}alkyl is partially or completely substituted with one or more deuterium atoms where allowed by available valences.

As used herein, the term "halo" or "halogen" generally refers to a halogen atom radical, including fluoro, chloro, bromo and iodo.

As used herein, the term "halo-C_{1-4}alkoxy" refers to a radical of the formula: -O-C_{1-4}alkyl-halo, wherein C_{1-4}alkyl is partially or completely substituted with one or more halogen atoms where allowed by available valences.

As used herein, the term "halo-C_{1-4}alkyl" refers to a radical of the formula: -C_{1-4}alkyl-halo, wherein C_{1-4}alkyl is partially or completely substituted with one or more halogen atoms where allowed by available valences.

As used herein, the term "halo-C_{1-4}alkyl-amino" refers to a radical of the formula: -NH-C_{1-4}alkyl-halo.

As used herein, the term "hydroxy" refers to a radical of the formula: -OH.

As used herein, the term "hydroxy-C_{1-4}alkyl" refers to a radical of the formula: -C_{1-4}alkyl-OH, wherein C_{1-4}alkyl is partially or completely substituted with one or more hydroxy radicals where allowed by available valences.

As used herein, the term "substituent" means positional variables on the atoms of a core molecule that are substituted at a designated atom position, replacing one or more hydrogens on the designated atom, provided that the designated atom's normal valency is not exceeded, and that the substitution results in a stable compound. Combinations of substituents and/or variables are permissible only if such combinations result in stable compounds. A person of ordinary skill in the art should note that any carbon as well as heteroatom with valences that appear to be unsatisfied as described or shown herein is assumed to have a sufficient number of hydrogen atom(s) to satisfy the valences described or shown. In certain instances one or more substituents having a double bond (e.g., "oxo" or "=O") as the point of attachment may be described, shown or listed herein within a substituent group, wherein the structure may only show a single bond as the point of attachment to the core structure of Formula (I). A person of ordinary skill in the art would understand that, while only a single bond is shown, a double bond is intended for those substituents.

As used herein, the term "and the like," with reference to the definitions of chemical terms provided herein, means that variations in chemical structures that could be expected by one skilled in the art include, without limitation, isomers (including chain, branching or positional structural isomers), hydration of ring systems (including saturation or partial unsaturation of monocyclic, bicyclic or polycyclic ring structures) and all other variations where allowed by available valences which result in a stable compound.

For the purposes of this description, where one or more substituent variables for a compound of Formula (I) or a form thereof encompass functionalities incorporated into a compound of Formula (I), each functionality appearing at any location within the disclosed compound may be independently selected, and as appropriate, independently and/or optionally substituted.

As used herein, the terms "independently selected," or "each selected" refer to functional variables in a substituent list that may occur more than once on the structure of Formula (I), the pattern of substitution at each occurrence is independent of the pattern at any other occurrence. Further, the use of a generic substituent variable on any formula or structure for a compound described herein is understood to include the replacement of the generic substituent with species substituents that are included within the particular genus, *e.g.*, aryl

may be replaced with phenyl or naphthalenyl and the like, and that the resulting compound is to be included within the scope of the compounds described herein.

As used herein, the terms "each instance of" or "in each instance, when present," when used preceding a phrase such as "…C_{3-14}cycloalkyl, C_{3-14}cycloalkyl-C_{1-4}alkyl, aryl, aryl-C_{1-4}alkyl, heteroaryl, heteroaryl-C_{1-4}alkyl, heterocyclyl and heterocyclyl-C_{1-4}alkyl," are intended to refer to the C_{3-14}cycloalkyl, aryl, heteroaryl and heterocyclyl ring systems when each are present either alone or as a substituent.

As used herein, the term "optionally substituted" means optional substitution with the specified substituent variables, groups, radicals or moieties.

COMPOUND FORMS

As used herein, the term "form" means a compound of Formula (I) having a form selected from the group consisting of a free acid, free base, hydrate, solvate, ester, stereoisomer, and tautomer form thereof.

In certain aspects described herein, the form of the compound of Formula (I) is a free acid, free base or salt thereof.

In certain aspects described herein, the form of the compound of Formula (I)) is a salt thereof.

In certain aspects described herein, the form of the compound of Formula (I) is a tautomer thereof.

In certain aspects described herein, the form of the compound of Formula (I) is a pharmaceutically acceptable form.

In certain aspects described herein, the compound of Formula (I) or a form thereof is isolated for use.

As used herein, the term "isolated" means the physical state of a compound of Formula (I) or a form thereof after being isolated and/or purified from a synthetic process (*e.g.*, from a reaction mixture) or natural source or combination thereof according to an isolation or purification process or processes described herein or which are well known to the skilled artisan (*e.g.*, chromatography, recrystallization and the like) in sufficient purity to be characterized by standard analytical techniques described herein or well known to the skilled artisan.

As used herein, the term "protected" means that a functional group in a compound of Formula (I) or a form thereof is in a form modified to preclude undesired side reactions at the protected site when the compound is subjected to a reaction. Suitable protecting groups will

be recognized by those with ordinary skill in the art as well as by reference to standard textbooks such as, for example, T.W. Greene *et al*, *Protective Groups in organic Synthesis* (1991), Wiley, New York. Such functional groups include hydroxy, phenol, amino and carboxylic acid. Suitable protecting groups for hydroxy or phenol include trialkylsilyl or diarylalkylsilyl (e.g., t-butyldimethylsilyl, t-butyldiphenylsilyl or trimethylsilyl), tetrahydropyranyl, benzyl, substituted benzyl, methyl, methoxymethanol, and the like. Suitable protecting groups for amino, amidino and guanidino include t-butoxycarbonyl, benzyloxycarbonyl, and the like. Suitable protecting groups for carboxylic acid include alkyl, aryl or arylalkyl esters. In certain instances, the protecting group may also be a polymer resin, such as a Wang resin or a 2-chlorotrityl-chloride resin. Protecting groups may be added or removed in accordance with standard techniques, which are well-known to those skilled in the art and as described herein.

One or more compounds described herein may exist in unsolvated as well as solvated forms with pharmaceutically acceptable solvents such as water, ethanol, and the like, and the description herein is intended to embrace both solvated and unsolvated forms.

As used herein, the term “solvate” means a physical association of a compound described herein with one or more solvent molecules. This physical association involves varying degrees of ionic and covalent bonding, including hydrogen bonding. In certain instances the solvate will be capable of isolation, for example when one or more solvent molecules are incorporated in the crystal lattice of the crystalline solid. As used herein, “solvate” encompasses both solution-phase and isolatable solvates. Non-limiting examples of suitable solvates include ethanolates, methanolates, and the like.

As used herein, the term “hydrate” means a solvate wherein the solvent molecule is water.

The compounds of Formula (I) can form salts, which are intended to be included within the scope of this description. Reference to a compound of Formula (I) a form thereof herein is understood to include reference to salt forms thereof, unless otherwise indicated. The term "salt(s)", as employed herein, denotes acidic salts formed with inorganic and/or organic acids, as well as basic salts formed with inorganic and/or organic bases. In addition, when a compound of Formula (I)) or a form thereof contains both a basic moiety, such as, without limitation an amine moiety, and an acidic moiety, such as, but not limited to a carboxylic acid, zwitterions ("inner salts") may be formed and are included within the term "salt(s)" as used herein.

The term "pharmaceutically acceptable salt(s)", as used herein, means those salts of compounds described herein that are safe and effective (*i.e.*, non-toxic, physiologically acceptable) for use in mammals and that possess biological activity, although other salts are also useful. Salts of the compounds of the Formula (I) may be formed, for example, by reacting a compound of Formula (I) or a form thereof with an amount of acid or base, such as an equivalent amount, in a medium such as one in which the salt precipitates or in an aqueous medium followed by lyophilization.

Pharmaceutically acceptable salts include one or more salts of acidic or basic groups present in compounds described herein. Particular aspects of acid addition salts include, and are not limited to, acetate, ascorbate, benzoate, benzenesulfonate, bisulfate, bitartrate, borate, bromide, butyrate, chloride, citrate, camphorate, camphorsulfonate, ethanesulfonate, formate, fumarate, gentisinate, gluconate, glucaronate, glutamate, iodide, isonicotinate, lactate, maleate, methanesulfonate, naphthalenesulfonate, nitrate, oxalate, pamoate, pantothenate, phosphate, propionate, saccharate, salicylate, succinate, sulfate, tartrate, thiocyanate, toluenesulfonate (also known as tosylate), trifluoroacetate salts and the like. Certain particular aspects of acid addition salts include chloride or dichloride.

Additionally, acids which are generally considered suitable for the formation of pharmaceutically useful salts from basic pharmaceutical compounds are discussed, for example, by P. Stahl *et al*, Camille G. (eds.) *Handbook of Pharmaceutical Salts. Properties, Selection and Use.* (2002) Zurich: Wiley-VCH; S. Berge *et al*, *Journal of Pharmaceutical Sciences* (1977) 66(1) 1-19; P. Gould, *International J. of Pharmaceutics* (1986) 33, 201-217; Anderson *et al*, *The Practice of Medicinal Chemistry* (1996), Academic Press, New York; and in *The Orange Book* (Food & Drug Administration, Washington, D.C. on their website). These disclosures are incorporated herein by reference thereto.

Suitable basic salts include, but are not limited to, aluminum, ammonium, calcium, lithium, magnesium, potassium, sodium and zinc salts.

All such acid salts and base salts are intended to be included within the scope of pharmaceutically acceptable salts as described herein. In addition, all such acid and base salts are considered equivalent to the free forms of the corresponding compounds for purposes of this description.

Compounds of Formula (I) and forms thereof, may further exist in a tautomeric form. All such tautomeric forms are contemplated and intended to be included within the scope of the compounds of Formula (I) or a form thereof as described herein.

The compounds of Formula (I) or a form thereof may contain asymmetric or chiral centers, and, therefore, exist in different stereoisomeric forms. The present description is intended to include all stereoisomeric forms of the compounds of Formula (I) as well as mixtures thereof, including racemic mixtures.

The compounds described herein may include one or more chiral centers, and as such may exist as racemic mixtures (*R/S*) or as substantially pure enantiomers and diastereomers. The compounds may also exist as substantially pure (*R*) or (*S*) enantiomers (when one chiral center is present). In one particular aspect, the compounds described herein are (*S*) isomers and may exist as enantiomerically pure compositions substantially comprising only the (*S*) isomer. In another particular aspect, the compounds described herein are (*R*) isomers and may exist as enantiomerically pure compositions substantially comprising only the (*R*) isomer. As one of skill in the art will recognize, when more than one chiral center is present, the compounds described herein may also exist as a (*R,R*), (*R,S*), (*S,R*) or (*S,S*) isomer, as defined by *IUPAC* Nomenclature Recommendations.

As used herein, the term "chiral" refers to a carbon atom bonded to four nonidentical substituents. Stereochemical definitions and conventions used herein generally follow S. P. Parker, Ed., McGraw-Hill Dictionary of Chemical Terms (1984) McGraw-Hill Book Company, New York; and Eliel, E. and Wilen, S., "Stereochemistry of Organic Compounds", John Wiley & Sons, Inc., New York, 1994. In describing an optically active compound, the prefixes D and L, or *R* and *S*, are used to denote the absolute configuration of the molecule about its chiral center(s). The substituents attached to the chiral center under consideration are ranked in accordance with the Sequence Rule of Cahn, Ingold and Prelog. (Cahn et al. Angew. Chem. Inter. Edit. 1966, 5, 385; errata 511).

As used herein, the term "substantially pure" refers to compounds consisting substantially of a single isomer in an amount greater than or equal to 90%, in an amount greater than or equal to 92%, in an amount greater than or equal to 95%, in an amount greater than or equal to 98%, in an amount greater than or equal to 99%, or in an amount equal to 100% of the single isomer.

In one aspect of the description, a compound of Formula (I)) or a form thereof is a substantially pure (*S*) enantiomer form present in an amount greater than or equal to 90%, in an amount greater than or equal to 92%, in an amount greater than or equal to 95%, in an amount greater than or equal to 98%, in an amount greater than or equal to 99%, or in an amount equal to 100%.

In one aspect of the description, a compound of Formula (I) or a form thereof is a substantially pure (*R*) enantiomer form present in an amount greater than or equal to 90%, in an amount greater than or equal to 92%, in an amount greater than or equal to 95%, in an amount greater than or equal to 98%, in an amount greater than or equal to 99%, or in an amount equal to 100%.

As used herein, a "racemate" is any mixture of isometric forms that are not "enantiomerically pure", including mixtures such as, without limitation, in a ratio of about 50/50, about 60/40, about 70/30, or about 80/20.

In addition, the present description embraces all geometric and positional isomers. For example, if a compound of Formula (I) or a form thereof incorporates a double bond or a fused ring, both the cis- and trans-forms, as well as mixtures, are embraced within the scope of the description. Diastereomeric mixtures can be separated into their individual diastereomers on the basis of their physical chemical differences by methods well known to those skilled in the art, such as, for example, by chromatography and/or fractional crystallization. Enantiomers can be separated by use of chiral HPLC column or other chromatographic methods known to those skilled in the art. Enantiomers can also be separated by converting the enantiomeric mixture into a diastereomeric mixture by reaction with an appropriate optically active compound (*e.g.*, chiral auxiliary such as a chiral alcohol or Mosher's acid chloride), separating the diastereomers and converting (*e.g.*, hydrolyzing) the individual diastereomers to the corresponding pure enantiomers. Also, some of the compounds of Formula (I) may be atropisomers (*e.g.*, substituted biaryls) and are considered as part of this description.

All stereoisomers (for example, geometric isomers, optical isomers and the like) of the present compounds (including those of the salts, solvates, esters and prodrugs of the compounds as well as the salts, solvates and esters of the prodrugs), such as those which may exist due to asymmetric carbons on various substituents, including enantiomeric forms (which may exist even in the absence of asymmetric carbons), rotameric forms, atropisomers, and diastereomeric forms, are contemplated within the scope of this description, as are positional isomers (such as, for example, 4-pyridinyl and 3-pyridinyl). Individual stereoisomers of the compounds described herein may, for example, be substantially free of other isomers, or may be present in a racemic mixture, as described supra.

The use of the terms "salt", "solvate" and the like, is intended to equally apply to the salt and solvate of enantiomers, stereoisomers, rotamers, tautomers, positional isomers, or racemates of the instant compounds.

COMPOUND USES

In accordance with the intended scope of the present description, aspects of the present description include compounds that have been identified and have been demonstrated to be useful in selectively preventing, treating or ameliorating HD and have been provided for use for preventing, treating or ameliorating HD.

An aspect of the present description includes a method for preventing, treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of a compound of Formula (I) or a form thereof.

An aspect of the present description includes a method for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of a compound of Formula (I) or a form thereof.

An aspect of the present description includes a method for preventing HD in a subject in need thereof comprising, administering to the subject an effective amount of a compound of Formula (I) or a form thereof.

An aspect of the present description includes a method for treating HD in a subject in need thereof comprising, administering to the subject an effective amount of a compound of Formula (I) or a form thereof.

An aspect of the present description includes a method for ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of a compound of Formula (I) or a form thereof.

Another aspect of the present description includes a method for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of a compound salt of Formula (I) or a form thereof.

Another aspect of the present description includes a method for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of a compound or compound salt of Formula (I) or a form thereof, in a combination product, or as a combination therapy, with one or more therapeutic agents.

An aspect of the present description includes a method for use of a compound of Formula (I) or a form or composition thereof for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the compound of Formula (I) or a form or composition thereof.

Another aspect of the present description includes a method for use of a compound salt of Formula (I) or a form or composition thereof for treating or ameliorating HD in a subject in

need thereof comprising, administering to the subject an effective amount of the compound salt of Formula (I) or a form thereof.

An aspect of the present description includes a use for a compound of Formula (I) or a form thereof for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the compound of Formula (I) or a form thereof.

Another aspect of the present description includes a use for a compound salt of Formula (I) or a form thereof for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the compound salt of Formula (I) or a form thereof.

An aspect of the present description includes a use for a compound of Formula (I) or a form thereof in the manufacture of a medicament for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the medicament.

Another aspect of the present description includes a use for a compound salt of Formula (I)) or a form thereof in the manufacture of a medicament for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the medicament.

An aspect of the present description includes in vitro or in vivo use of the compound of Formula (I) or a form thereof having activity toward HD.

An aspect of the present description includes a use of the compound of Formula (I) or a form thereof in a combination therapy to provide additive or synergistic activity, thus enabling the development of a combination product for treating or ameliorating HD.

Another aspect of the present description includes a combination therapy comprising compounds described herein in combination with one or more known drugs or one or more known therapies may be used to treat HD regardless of whether HD is responsive to the known drug.

An aspect of the present description includes a use for a compound of Formula (I) or a form thereof in a combination product with one or more therapeutic agents for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the compound of Formula (I) or a form thereof in combination with an effective amount of the one or more agents.

Another aspect of the present description includes a use for a compound salt of Formula (I) or a form thereof in a combination product with one or more therapeutic agents

for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the compound salt of Formula (I) or a form thereof in combination with an effective amount of the one or more agents.

In an aspect of a use or method provided herein, compounds of Formula (I) or a form thereof used in combination with one or more additional agents can be administered to a subject or contacted with a subject or patient cell(s) prior to, concurrently with, or subsequent to administering to the subject or patient or contacting the cell with an additional agent(s). A compound(s) of Formula (I) or a form thereof and an additional agent(s) can be administered to a subject or contacted with a cell in single composition or different compositions. In a specific aspect, a compound(s) of Formula (I) or a form thereof is used in combination with gene therapy to inhibit HTT expression (using, *e.g.*, viral delivery vectors) or the administration of another small molecule HTT inhibitor. In another specific aspect, a compound(s) of Formula (I) or a form thereof are used in combination with cell replacement using differentiated non-mutant HTT stem cells. In another specific aspect, a compound(s) of Formula (I) or a form thereof are used in combination with cell replacement using differentiated HTT stem cells.

In one aspect, provided herein is the use of compounds of Formula (I) or a form thereof in combination with supportive standard of care therapies, including palliative care.

In one respect, for each of such aspects, the subject is treatment naive. In another respect, for each of such aspects, the subject is not treatment naive.

As used herein, the term "preventing" refers to keeping a disease, disorder or condition from occurring in a subject that may be predisposed to the disease, disorder and/or condition but has not yet been diagnosed as having the disease, disorder and/or condition.

As used herein, the term "treating" refers to inhibiting the progression of a disease, disorder or condition in a subject already exhibiting the symptoms of the disease, disorder and/or condition, *i.e.*, arresting the development of a disease, disorder and/or condition that has already affected the subject.

As used herein, the term "ameliorating" refers to relieving the symptoms of a disease, disorder or condition in a subject already exhibiting the symptoms of the disease, disorder and/or condition, *i.e.*, causing regression of the disease, disorder and/or condition that has already affected the subject.

As used herein, the term "subject" refers to an animal or any living organism having sensation and the power of voluntary movement, and which requires oxygen and organic food. Nonlimiting examples include members of the human, primate, equine, porcine, bovine,

murine, rattus, canine and feline specie. In certain aspects, the subject is a mammal or a warm-blooded vertebrate animal. In other aspects, the subject is a human. As used herein, the term “patient” may be used interchangeably with “subject” and “human”.

As used herein, the terms “effective amount” or "therapeutically effective amount" mean an amount of compound of Formula (I) or a form, composition or medicament thereof that achieves a target plasma concentration that is effective in treating or ameliorating HD as described herein and thus producing the desired therapeutic, ameliorative, inhibitory or preventative effect in a subject in need thereof. In one aspect, the effective amount may be the amount required to treat HD in a subject or patient, more specifically, in a human.

In another aspect, the concentration-biological effect relationships observed with regard to a compound of Formula (I) or a form thereof indicate a target plasma concentration ranging from approximately 0.001 μg/mL to approximately 50 μg/mL, from approximately 0.01 μg/mL to approximately 20 μg/mL, from approximately 0.05 μg/mL to approximately 10 μg/mL, or from approximately 0.1 μg/mL to approximately 5 μg/mL. To achieve such plasma concentrations, the compounds described herein may be administered at doses that vary, such as, for example, without limitation, from 1.0 ng to 10,000 mg.

In one aspect, the dose administered to achieve an effective target plasma concentration may be administered based upon subject or patient specific factors, wherein the doses administered on a weight basis may be in the range of from about 0.001 mg/kg/day to about 3500 mg/kg/day, or about 0.001 mg/kg/day to about 3000 mg/kg/day, or about 0.001 mg/kg/day to about 2500 mg/kg/day, or about 0.001 mg/kg/day to about 2000 mg/kg/day, or about 0.001 mg/kg/day to about 1500 mg/kg/day, or about 0.001 mg/kg/day to about 1000 mg/kg/day, or about 0.001 mg/kg/day to about 500 mg/kg/day, or about 0.001 mg/kg/day to about 250 mg/kg/day, or about 0.001 mg/kg/day to about 200 mg/kg/day, or about 0.001 mg/kg/day to about 150 mg/kg/day, or about 0.001 mg/kg/day to about 100 mg/kg/day, or about 0.001 mg/kg/day to about 75 mg/kg/day, or about 0.001 mg/kg/day to about 50 mg/kg/day, or about 0.001 mg/kg/day to about 25 mg/kg/day, or about 0.001 mg/kg/day to about 10 mg/kg/day, or about 0.001 mg/kg/day to about 5 mg/kg/day, or about 0.001 mg/kg/day to about 1 mg/kg/day, or about 0.001 mg/kg/day to about 0.5 mg/kg/day, or about 0.001 mg/kg/day to about 0.1 mg/kg/day, or from about 0.01 mg/kg/day to about 3500 mg/kg/day, or about 0.01 mg/kg/day to about 3000 mg/kg/day, or about 0.01 mg/kg/day to about 2500 mg/kg/day, or about 0.01 mg/kg/day to about 2000 mg/kg/day, or about 0.01 mg/kg/day to about 1500 mg/kg/day, or about 0.01 mg/kg/day to about 1000 mg/kg/day, or about 0.01 mg/kg/day to about 500 mg/kg/day, or about 0.01 mg/kg/day to about 250

mg/kg/day, or about 0.01 mg/kg/day to about 200 mg/kg/day, or about 0.01 mg/kg/day to about 150 mg/kg/day, or about 0.01 mg/kg/day to about 100 mg/kg/day, or about 0.01 mg/kg/day to about 75 mg/kg/day, or about 0.01 mg/kg/day to about 50 mg/kg/day, or about 0.01 mg/kg/day to about 25 mg/kg/day, or about 0.01 mg/kg/day to about 10 mg/kg/day, or about 0.01 mg/kg/day to about 5 mg/kg/day, or about 0.01 mg/kg/day to about 1 mg/kg/day, or about 0.01 mg/kg/day to about 0.5 mg/kg/day, or about 0.01 mg/kg/day to about 0.1 mg/kg/day, or from about 0.1 mg/kg/day to about 3500 mg/kg/day, or about 0.1 mg/kg/day to about 3000 mg/kg/day, or about 0.1 mg/kg/day to about 2500 mg/kg/day, or about 0.1 mg/kg/day to about 2000 mg/kg/day, or about 0.1 mg/kg/day to about 1500 mg/kg/day, or about 0.1 mg/kg/day to about 1000 mg/kg/day, or about 0.1 mg/kg/day to about 500 mg/kg/day, or about 0.1 mg/kg/day to about 250 mg/kg/day, or about 0.1 mg/kg/day to about 200 mg/kg/day, or about 0.1 mg/kg/day to about 150 mg/kg/day, or about 0.1 mg/kg/day to about 100 mg/kg/day, or about 0.1 mg/kg/day to about 75 mg/kg/day, or about 0.1 mg/kg/day to about 50 mg/kg/day, or about 0.1 mg/kg/day to about 25 mg/kg/day, or about 0.1 mg/kg/day to about 10 mg/kg/day, or about 0.1 mg/kg/day to about 5 mg/kg/day, or about 0.1 mg/kg/day to about 1 mg/kg/day, or about 0.1 mg/kg/day to about 0.5 mg/kg/day.

Effective amounts for a given subject may be determined by routine experimentation that is within the skill and judgment of a clinician or a practitioner skilled in the art in light of factors related to the subject. Dosage and administration may be adjusted to provide sufficient levels of the active agent(s) or to maintain the desired effect. Factors which may be taken into account include genetic screening, severity of the disease state, status of disease progression, general health of the subject, ethnicity, age, weight, gender, diet, time of day and frequency of administration, drug combination(s), reaction sensitivities, experience with other therapies, and tolerance/response to therapy.

The dose administered to achieve an effective target plasma concentration may be orally administered once (once in approximately a 24 hour period; i.e., "q.d."), twice (once in approximately a 12 hour period; i.e., "b.i.d." or "q.12h"), thrice (once in approximately an 8 hour period; i.e., "t.i.d." or "q.8h"), or four times (once in approximately a 6 hour period; i.e., "q.d.s.", "q.i.d." or "q.6h") daily.

In certain aspects, the dose administered to achieve an effective target plasma concentration may also be administered in a single, divided, or continuous dose for a patient or subject having a weight in a range of between about 40 to about 200 kg (which dose may be adjusted for patients or subjects above or below this range, particularly children under 40 kg). The typical adult subject is expected to have a median weight in a range of about 70 kg.

Long-acting pharmaceutical compositions may be administered every 2, 3 or 4 days, once every week, or once every two weeks depending on half-life and clearance rate of the particular formulation.

The compounds and compositions described herein may be administered to the subject via any drug delivery route known in the art. Nonlimiting examples include oral, ocular, rectal, buccal, topical, nasal, sublingual, transdermal, subcutaneous, intramuscular, intraveneous (bolus and infusion), intracerebral, and pulmonary routes of administration.

In another aspect, the dose administered may be adjusted based upon a dosage form described herein formulated for delivery at about 0.02, 0.025, 0.03, 0.05, 0.06, 0.075, 0.08, 0.09, 0.10, 0.20, 0.25, 0.30, 0.50, 0.60, 0.75, 0.80, 0.90, 1.0, 1.10, 1.20, 1.25, 1.50, 1.75, 2.0, 3.0, 5.0, 10, 20, 30, 40, 50, 100, 150, 200, 250, 300, 400, 500, 1000, 1500, 2000, 2500, 3000 or 4000 mg/day.

For any compound, the effective amount can be estimated initially either in cell culture assays or in relevant animal models, such as a mouse, guinea pig, chimpanzee, marmoset or tamarin animal model. Relevant animal models may also be used to determine the appropriate concentration range and route of administration. Such information can then be used to determine useful doses and routes for administration in humans. Therapeutic efficacy and toxicity may be determined by standard pharmaceutical procedures in cell cultures or experimental animals, *e.g.*, ED_{50} (the dose therapeutically effective in 50% of the population) and LD_{50} (the dose lethal to 50% of the population). The dose ratio between therapeutic and toxic effects is therapeutic index, and can be expressed as the ratio, LD_{50}/ED_{50}. In certain aspects, the effective amount is such that a large therapeutic index is achieved. In further particular aspects, the dosage is within a range of circulating concentrations that include an ED_{50} with little or no toxicity. The dosage may vary within this range depending upon the dosage form employed, sensitivity of the patient, and the route of administration.

In one aspect, provided herein are methods for modulating the amount of HTT (huntingtin protein), comprising contacting a human cell with a compound of Formula (I) or a form thereof. In a specific aspect, provided herein are methods for modulating the amount of HTT, comprising contacting a human cell with a compound of Formula (I)) or a form thereof that modulates the expression of HTT. The human cell can be contacted with a compound of Formula (I)) or a form thereof *in vitro*, or *in vivo*, *e.g.*, in a non-human animal or in a human. In a specific aspect, the human cell is from or in a human. In another specific aspect, the human cell is from or in a human with HD. In another specific aspect, the human cell is from or in a human with HD, caused by a CAG repeat in the Htt gene, resulting in a loss of HTT

expression and/or function. In another aspect, the human cell is from a human with HD. In another aspect, the human cell is in a human with HD. In one aspect, the compound is a form of the compound of Formula (I).

In a specific aspect, provided herein is a method for enhancing the inhibition of mutant HTT transcribed from the Htt gene, comprising contacting a human cell with a compound of Formula (I) or a form thereof. The human cell can be contacted with a compound of Formula (I) or a form thereof *in vitro*, or *in vivo*, *e.g.*, in a non-human animal or in a human. In a specific aspect, the human cell is from or in a human. In another specific aspect, the human cell is from or in a human with HD. In another specific aspect, the human cell is from or in a human with HD, caused by a CAG repeat in the Htt gene, resulting in a loss of wild-type "normal" HTT expression and/or function. In another aspect, the human cell is from a human with HD. In another aspect, the human cell is in a human with HD. In one aspect, the compound is a form of the compound of Formula (I).

In another aspect, provided herein is a method for modulating the inhibition of mutant HTT transcribed from the Htt gene, comprising administering to a non-human animal model for HD a compound of Formula (I) or a form thereof. In a specific aspect, provided herein is a method for modulating the inhibition of mutant HTT transcribed from the Htt gene, comprising administering to a non-human animal model for HD a compound of Formula (I) or a form thereof. In a specific aspect, the compound is a form of the compound of Formula (I).

In another aspect, provided herein is a method for decreasing the amount of mutant HTT, comprising contacting a human cell with a compound of Formula (I) or a form thereof. In a specific aspect, provided herein is a method for decreasing the amount of mutant HTT, comprising contacting a human cell with a compound of Formula (I) that inhibits the transcription of mutant *HTT* (huntingtin mRNA) from the Htt gene. In another specific aspect, provided herein is a method for decreasing the amount of HTT, comprising contacting a human cell with a compound of Formula (I) that inhibits the expression of mutant HTT transcribed from the Htt gene. The human cell can be contacted with a compound of Formula (I) or a form thereof *in vitro*, or *in vivo*, *e.g.*, in a non-human animal or in a human. In a specific aspect, the human cell is from or in a human. In another specific aspect, the human cell is from or in a human with HD. In another specific aspect, the human cell is from or in a human with HD, caused by a CAG repeat in the Htt gene, resulting in a loss of HTT expression and/or function. In another aspect, the human cell is from a human with HD. In another aspect, the human cell is in a human with HD. In one aspect, the compound is a form of the compound of Formula (I)).

In certain aspects, treating or ameliorating HD with a compound of Formula (I) or a form thereof (alone or in combination with an additional agent) has a therapeutic effect and/or beneficial effect. In a specific aspect, treating HD with a compound of Formula (I) or a form thereof (alone or in combination with an additional agent) results in one, two or more of the following effects: (i) reduces or ameliorates the severity of HD; (ii) delays onset of HD; (iii) inhibits the progression of HD; (iv) reduces hospitalization of a subject; (v) reduces hospitalization length for a subject; (vi) increases the survival of a subject; (vii) improves the quality of life for a subject; (viii) reduces the number of symptoms associated with HD; (ix) reduces or ameliorates the severity of a symptom(s) associated with HD; (x) reduces the duration of a symptom associated with HD; (xi) prevents the recurrence of a symptom associated with HD; (xii) inhibits the development or onset of a symptom of HD; and/or (xiii) inhibits of the progression of a symptom associated with HD.

METABOLITES

Another aspect included within the scope of the present description are the use of *in vivo* metabolic products of the compounds described herein. Such products may result, for example, from the oxidation, reduction, hydrolysis, amidation, esterification and the like of the administered compound, primarily due to enzymatic processes. Accordingly, the description includes the use of compounds produced by a process comprising contacting a compound described herein with a mammalian tissue or a mammal for a period of time sufficient to yield a metabolic product thereof.

Such products typically are identified by preparing a radio-labeled isotopologue (*e.g.*, ^{14}C or ^{3}H) of a compound described herein, administering the radio-labeled compound in a detectable dose (*e.g.*, greater than about 0.5 mg/kg) to a mammal such as a rat, mouse, guinea pig, dog, monkey or human, allowing sufficient time for metabolism to occur (typically about 30 seconds to about 30 hours), and identifying the metabolic conversion products from urine, bile, blood or other biological samples. The conversion products are easily isolated since they are "radiolabeled" by virtue of being isotopically-enriched (others are isolated by the use of antibodies capable of binding epitopes surviving in the metabolite). The metabolite structures are determined in conventional fashion, *e.g.*, by MS or NMR analysis. In general, analysis of metabolites may be done in the same way as conventional drug metabolism studies well-known to those skilled in the art. The conversion products, so long as they are not otherwise found *in vivo*, are useful in diagnostic assays for therapeutic dosing of the compounds described herein even if they possess no biological activity of their own.

PHARMACEUTICAL COMPOSITIONS

In accordance with the intended scope of the present description, aspects of the present description include compounds that have been identified and have been demonstrated to be useful in selectively preventing, treating or ameliorating HD and have been provided for use as one or more pharmaceutical compositions for preventing, treating or ameliorating HD.

An aspect of the present description includes a use for a compound of Formula (I) or a form thereof in the preparation of a pharmaceutical composition for treating or ameliorating HD in a subject in need thereof comprising, administering to the subject an effective amount of the compound of Formula (I) or a form thereof in admixture with one or more pharmaceutically acceptable excipients.

An aspect of the present description includes a use for a pharmaceutical composition of the compound of Formula (I) or a form thereof in the preparation of a kit for treating or ameliorating HD in a subject in need thereof comprising, the pharmaceutical composition of the compound of Formula (I) or a form thereof and instructions for administering the pharmaceutical composition.

As used herein, the term "composition" means a product comprising the specified ingredients in the specified amounts, as well as any product which results, directly or indirectly, from combination of the specified ingredients in the specified amounts.

The pharmaceutical composition may be formulated to achieve a physiologically compatible pH, ranging from about pH 3 to about pH 11. In certain aspects, the pharmaceutical composition is formulated to achieve a pH of from about pH 3 to about pH 7. In other aspects, the pharmaceutical composition is formulated to achieve a pH of from about pH 5 to about pH 8.

The term "pharmaceutically acceptable excipient" refers to an excipient for administration of a pharmaceutical agent, such as the compounds described herein. The term refers to any pharmaceutical excipient that may be administered without undue toxicity. Pharmaceutically acceptable excipients may be determined in part by the particular composition being administered, as well as by the particular mode of administration and/or dosage form. Nonlimiting examples of pharmaceutically acceptable excipients include carriers, solvents, stabilizers, adjuvants, diluents, *etc.* Accordingly, there exists a wide variety of suitable formulations of pharmaceutical compositions for the instant compounds described herein (*see, e.g.*, Remington's Pharmaceutical Sciences).

Suitable excipients may be carrier molecules that include large, slowly metabolized macromolecules such as proteins, polysaccharides, polylactic acids, polyglycolic acids,

polymeric amino acids, amino acid copolymers, and inactive antibodies. Other exemplary excipients include antioxidants such as ascorbic acid; chelating agents such as EDTA; carbohydrates such as dextrin, hydroxyalkylcellulose, hydroxyalkylmethylcellulose (*e.g.*, hydroxypropylmethylcellulose, also known as HPMC), stearic acid; liquids such as oils, water, saline, glycerol and ethanol; wetting or emulsifying agents; pH buffering substances; and the like. Liposomes are also included within the definition of pharmaceutically acceptable excipients.

The pharmaceutical compositions described herein may be formulated in any form suitable for the intended use described herein. Suitable formulations for oral administration include solids, liquid solutions, emulsions and suspensions, while suitable inhalable formulations for pulmonary administration include liquids and powders. Alternative formulations include syrups, creams, ointments, tablets, and lyophilized solids which can be reconstituted with a physiologically compatible solvent prior to administration.

When intended for oral use for example, tablets, troches, lozenges, aqueous or oil suspensions, non-aqueous solutions, dispersible powders or granules (including micronized particles or nanoparticles), emulsions, hard or soft capsules, syrups or elixirs may be prepared. Compositions intended for oral use may be prepared according to any method known to the art for the manufacture of pharmaceutical compositions, and such compositions may contain one or more agents including sweetening agents, flavoring agents, coloring agents, and preserving agents, in order to provide a palatable preparation.

Pharmaceutically acceptable excipients suitable for use in conjunction with tablets include, for example, inert diluents, such as celluloses, calcium or sodium carbonate, lactose, calcium or sodium phosphate; disintegrating agents, such as croscarmellose sodium, cross-linked povidone, maize starch, or alginic acid; binding agents, such as povidone, starch, gelatin or acacia; and lubricating agents, such as magnesium stearate, stearic acid, or talc. Tablets may be uncoated or may be coated by known techniques including microencapsulation to delay disintegration and adsorption in the gastrointestinal tract and thereby provide a sustained action over a longer period. For example, a time delay material such as glyceryl monostearate or glyceryl distearate alone or with a wax may be employed.

Formulations for oral use may be also presented as hard gelatin capsules where the active ingredient is mixed with an inert solid diluent, for example celluloses, lactose, calcium phosphate, or kaolin, or as soft gelatin capsules wherein the active ingredient is mixed with non-aqueous or oil medium, such as glycerin, propylene glycol, polyethylene glycol, peanut oil, liquid paraffin, or olive oil.

In other aspects, pharmaceutical compositions described herein may be formulated as suspensions comprising a compound of Formula (I) or a form thereof in admixture with one or more pharmaceutically acceptable excipients suitable for the manufacture of a suspension. In yet other aspects, pharmaceutical compositions described herein may be formulated as dispersible powders and granules suitable for preparation of a suspension by the addition of one or more excipients.

Excipients suitable for use in connection with suspensions include suspending agents, such as sodium carboxymethylcellulose, methylcellulose, hydroxypropyl methylcelluose, sodium alginate, polyvinylpyrrolidone, gum tragacanth, gum acacia, dispersing or wetting agents such as a naturally occurring phosphatide (*e.g.*, lecithin), a condensation product of an alkylene oxide with a fatty acid (*e.g.*, polyoxyethylene stearate), a condensation product of ethylene oxide with a long chain aliphatic alcohol (*e.g.*, heptadecaethyleneoxycethanol), a condensation product of ethylene oxide with a partial ester derived from a fatty acid and a hexitol anhydride (*e.g.*, polyoxyethylene sorbitan monooleate); and thickening agents, such as carbomer, beeswax, hard paraffin, or cetyl alcohol. The suspensions may also contain one or more preservatives such as acetic acid, methyl and/or n-propyl p-hydroxy-benzoate; one or more coloring agents; one or more flavoring agents; and one or more sweetening agents such as sucrose or saccharin.

The pharmaceutical compositions described herein may also be in the form of oil-in-water emulsions. The oily phase may be a vegetable oil, such as olive oil or arachis oil, a mineral oil, such as liquid paraffin, or a mixture of these. Suitable emulsifying agents include naturally-occurring gums, such as gum acacia and gum tragacanth; naturally occurring phosphatides, such as soybean lecithin, esters or partial esters derived from fatty acids; hexitol anhydrides, such as sorbitan monooleate; and condensation products of these partial esters with ethylene oxide, such as polyoxyethylene sorbitan monooleate. The emulsion may also contain sweetening and flavoring agents. Syrups and elixirs may be formulated with sweetening agents, such as glycerol, sorbitol or sucrose. Such formulations may also contain a demulcent, a preservative, a flavoring or a coloring agent.

Additionally, the pharmaceutical compositions described herein may be in the form of a sterile injectable preparation, such as a sterile injectable aqueous emulsion or oleaginous suspension. Such emulsion or suspension may be formulated according to the known art using those suitable dispersing or wetting agents and suspending agents which have been mentioned above. The sterile injectable preparation may also be a sterile injectable solution or suspension in a non-toxic parenterally acceptable diluent or solvent, such as a solution in 1,2-

propanediol. The sterile injectable preparation may also be prepared as a lyophilized powder. Among the acceptable vehicles and solvents that may be employed are water, Ringer's solution and isotonic sodium chloride solution. In addition, sterile fixed oils may be employed as a solvent or suspending medium. For this purpose any bland fixed oil may be employed including synthetic mono- or di-glycerides. In addition, fatty acids such as oleic acid may likewise be used in the preparation of injectables.

The compounds described herein may be substantially insoluble in water and sparingly soluble in most pharmaceutically acceptable protic solvents and vegetable oils, but generally soluble in medium-chain fatty acids (*e.g.*, caprylic and capric acids) or triglycerides and in propylene glycol esters of medium-chain fatty acids. Thus, contemplated in the description are compounds which have been modified by substitutions or additions of chemical or biochemical moieties which make them more suitable for delivery (*e.g.*, increase solubility, bioactivity, palatability, decrease adverse reactions, *etc.*), for example by esterification, glycosylation, PEGylation, etc.

In certain aspects, the compound described herein is formulated for oral administration in a lipid-based composition suitable for low solubility compounds. Lipid-based formulations can generally enhance the oral bioavailability of such compounds. As such, pharmaceutical compositions described herein may comprise a effective amount of a compound of Formula (I) or a form thereof, together with at least one pharmaceutically acceptable excipient selected from medium chain fatty acids or propylene glycol esters thereof (*e.g.*, propylene glycol esters of edible fatty acids such as caprylic and capric fatty acids) and pharmaceutically acceptable surfactants, such as polysorbate 20 or 80 (also referred to as Tween® 20 or Tween® 80, respectively) or polyoxyl 40 hydrogenated castor oil.

In other aspects, the bioavailability of low solubility compounds may be enhanced using particle size optimization techniques including the preparation of nanoparticles or nanosuspensions using techniques known to those skilled in the art. The compound forms present in such preparations include amorphous, partially amorphous, partially crystalline or crystalline forms.

In alternative aspects, the pharmaceutical composition may further comprise one or more aqueous solubility enhancer(s), such as a cyclodextrin. Nonlimiting examples of cyclodextrin include hydroxypropyl, hydroxyethyl, glucosyl, maltosyl and maltotriosyl derivatives of α-, β-, and γ-cyclodextrin, and hydroxypropyl-β-cyclodextrin (HPBC). In certain aspects, the pharmaceutical composition further comprises HPBC in a range of from about 0.1% to about 20%, from about 1% to about 15%, or from about 2.5% to about 10%.

The amount of solubility enhancer employed may depend on the amount of the compound in the composition.

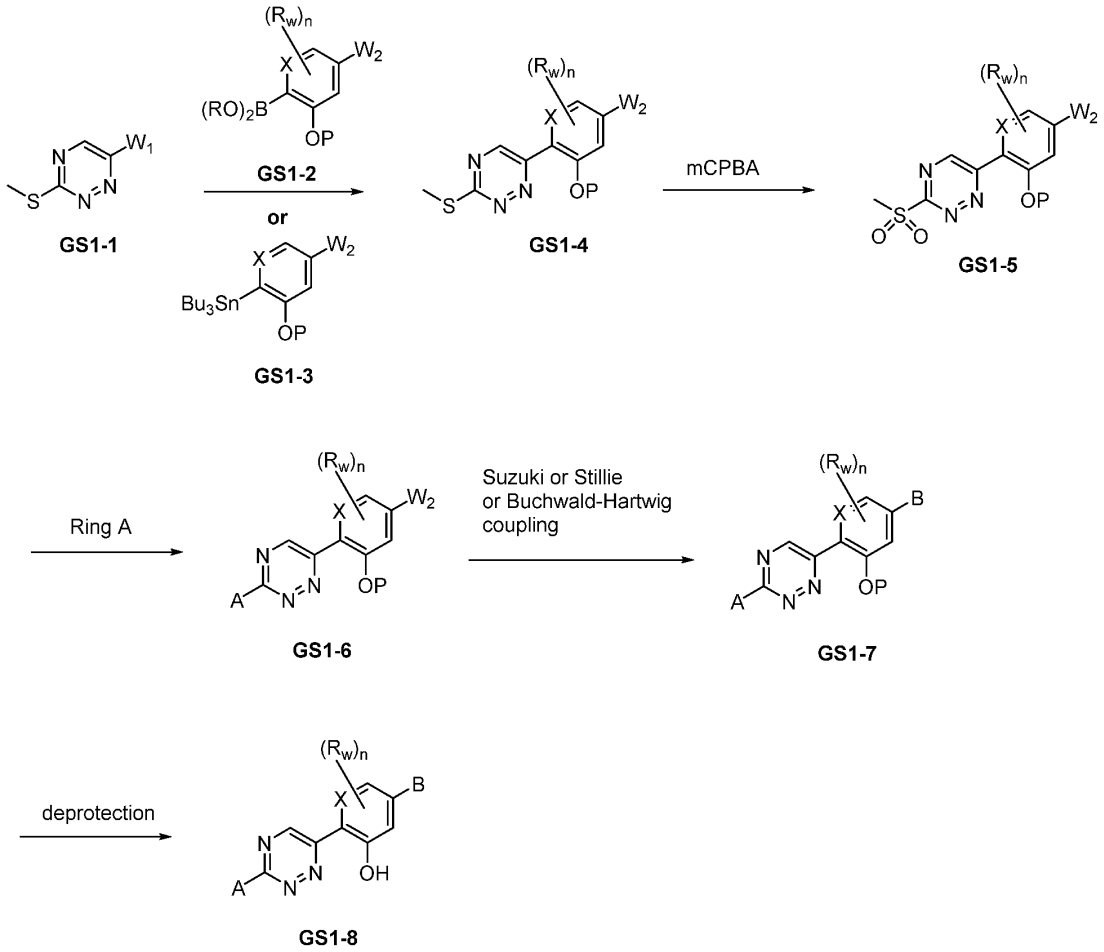
PREPARATION OF COMPOUNDS

GENERAL SYNTHETIC METHODS

As disclosed herein, general methods for preparing the compounds of Formula (I) or a form thereof as described herein are available via standard, well-known synthetic methodology. Many of the starting materials are commercially available or, when not available, can be prepared using the routes described below using techniques known to those skilled in the art. The synthetic schemes provided herein comprise multiple reaction steps, each of which is intended to stand on its own and can be carried out with or without any preceding or succeeding step(s). In other words, each of the individual reaction steps of the synthetic schemes provided herein in isolation is contemplated.

Compounds of Formula **GS1-8**, wherein A, B, X, R_W, and n are as defined for Formula (I) may be prepared as described in General Scheme 1 below.

General Scheme 1



Compound **GS1-1** (where W_1 is bromine, chlorine and the like) is converted to Compound **GS1-4** by a Suzuki coupling with an aryl-boronic acid (or pinacol boronic ester) **GS1-2** (where W_2 is bromine, chlorine and the like; R_1 is hydrogen, fluorine, chlorine, hydroxy, methoxy, aryl or heteroaryl; and P is a protecting group such as MOM and the like) in the presence of a catalyst (such as $Pd(dppf)Cl_2$ and the like) and base (such as aqueous K_2CO_3 and the like) in a suitable solvent (such as 1,4-dioxane and the like). Alternatively, Compound **GS1-1** is converted to Compound **GS1-4** by a Stille coupling with an aryl-stanane **GS1-3** (where W_2 is bromine, chlorine and the like; and P is a protecting group such as MOM and the like) in the presence of a catalyst (such as $Pd(PPh_3)_2Cl_2$ and the like) and additive (such as CuI and the like) in a suitable solvent (such as 1,4-dioxane and the like). Compound **GS1-4** is converted to Compound **GS1-5** by treatment with an oxidizing agent (such as

mCPBA or oxone and the like) in a suitable solvent (such as dichloromethane and the like). Compound **GS1-5** is converted to Compound **GS1-6** by a nucleophilic substitution with a piperazine Ring **A** in the presence of a suitable base (such as Et_3N and the like) in a suitable solvent (such as DMF and the like). Compound **GS1-6** is converted to Compound **GS1-7** by a Suzuki coupling with an aryl- or heteroaryl-boronic acid (or pinacol boronic ester) in the presence of a catalyst (such as $Pd(dppf)Cl_2$ and the like) and a base (such as aqueous K_2CO_3 and the like) in a suitable solvent (such as 1,4-dioxane and the like). Alternatively, Compound **A7** is converted to Compound **GS1-7** by a Stille coupling with an aryl- or heteroaryl-stannane in the presence of a catalyst (such as $Pd_2(dba)_3$ and the like), a ligand (such as X-Phos and the like) and a base (such as CsF and the like) in a suitable solvent (such as 1,4-dioxane and the like). Alternatively, Compound **GS1-6** is converted to Compound **GS1-7** by treatment with pinacolatodiboron and a base (such as KOAc and the like) in the presence of a catalyst (such as $Pd(dppf)Cl_2$ and the like) in an appropriate solvent (such as 1,4-dioxane and the like), followed by addition of an aryl- or heteroaryl-halide. Alternatively, Compound **GS1-6** is converted to Compound **GS1-7** by a Buchwald-Hartwig coupling with a heteroaryl or amine in the presence of a catalyst (such as $Pd_2(dba)_3$ and the like), a ligand (such as tBuX-Phos and the like) and a base (such as K_3PO_4 and the like) in a suitable solvent (such as 1,4-dioxane and the like). Compound **GS1-7** is converted to Compound **GS1-8** upon treatment with conditions appropriate to the removal of the protecting groups (such as HCl in dioxane for a MOM protecting group) in a suitable solvent (such as dioxane and the like).

SPECIFIC SYNTHETIC EXAMPLES

To describe in more detail and assist in understanding, the following non-limiting examples are offered to more fully illustrate the scope of compounds described herein and are not to be construed as specifically limiting the scope thereof. Such variations of the compounds described herein that may be now known or later developed, which would be within the purview of one skilled in the art to ascertain, are considered to fall within the scope of the compounds as described herein and hereinafter claimed. These examples illustrate the preparation of certain compounds. Those of skill in the art will understand that the techniques described in these examples represent techniques, as described by those of ordinary skill in the art, that function well in synthetic practice, and as such constitute preferred modes for the practice thereof. However, it should be appreciated that those of skill in the art should, in light of the present disclosure, appreciate that many changes can be made in the specific methods

that are disclosed and still obtain a like or similar result without departing from the spirit and scope of the present description.

Other than in the following examples of the embodied compounds, unless indicated to the contrary, all numbers expressing quantities of ingredients, reaction conditions, experimental data, and so forth used in the specification and claims are to be understood as being modified by the term "about." Accordingly, all such numbers represent approximations that may vary depending upon the desired properties sought to be obtained by a reaction or as a result of variable experimental conditions. Therefore, within an expected range of experimental reproducibility, the term "about" in the context of the resulting data, refers to a range for data provided that may vary according to a standard deviation from the mean. As well, for experimental results provided, the resulting data may be rounded up or down to present data consistently, without loss of significant figures. At the very least, and not as an attempt to limit the application of the doctrine of equivalents to the scope of the claims, each numerical parameter should be construed in light of the number of significant digits and rounding techniques used by those of skill in the art.

While the numerical ranges and parameters setting forth the broad scope of the present description are approximations, the numerical values set forth in the examples set forth below are reported as precisely as possible. Any numerical value, however, inherently contains certain errors necessarily resulting from the standard deviation found in their respective testing measurements.

The starting materials used in the examples provided are commercially available or can be prepared according to methods known to one skilled in the art or can be prepared by the proceedures disclosed herein. Structural confirmation was preformed using analytical techniques known to those skilled in the art such as ^{1}H or ^{13}C nuclear magnetic resonance spectroscopy or mass spectrometry.

Compounds of Formula (I) were synthesized using proceedures similar to those described in International Application Publication No. WO/2019/191229 A1, filed on March 27, 2019, and claiming priority to United States Provisional Application U.S. 62/648,699 filed on March 27, 2018, by substituting the appropriate starting materials, reagents and reaction conditions.

COMPOUND EXAMPLES

As used above, and throughout the present description, the following abbreviations, unless otherwise indicated, shall be understood to have the following meanings:

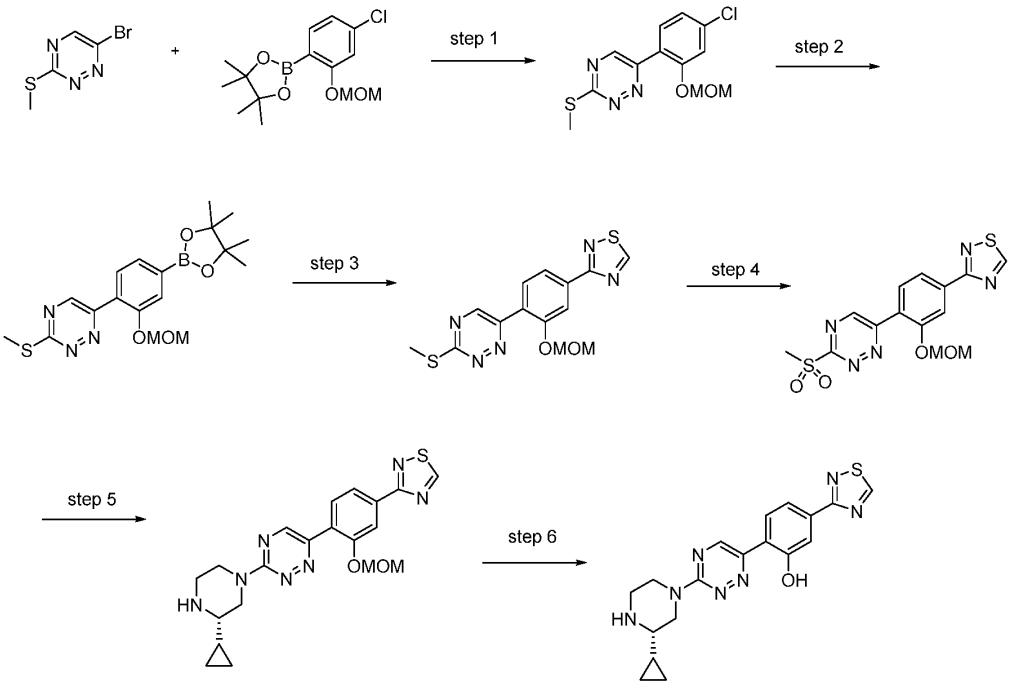
Abbreviation	Meaning
Δ	heating (chemistry) or deletion (biology)
AcOH or HOAc	acetic acid
Ar	argon
ACN or CH_3CN	acetonitrile
aq.	aqueous
atm	atmosphere(s)
BBr_3	boron tribromide
B_2pin_2	bis(pinacolato)diboron
Boc	tert-butoxy-carbonyl
Boc_2O	di-tert-butyl dicarbonate
t-Bu	tert-butyl
t-BuOK or KOtBu	postassium tert-butoxide
BuOH or *n*-BuOH	*n*-butanol
t-BuXPhos	2-di-tert-butylphosphino-2′,4′,6′-triisopropylbiphenyl
°C	degrees Centigrade
Celite® or Celite	diatomaceous earth
mCPBA	meta-chloroperoxybenzoic acid
CuI	copper(I) iodide
d/h/hr/hrs/min/s	day(d)/hour(h, hr or hrs)/minute(min)/second(s)
DCM or CH_2Cl_2	dichloromethane
DIPEA	N, N-diisopropylethylamine
DMAP	4-dimethylaminopyridine
DMF	dimethylformamide
DMSO	dimethylsulfoxide
EtOAc	ethyl acetate
EtOH	ethanol
Et_2O	diethyl ether
equiv	equivalents
H_2	hydrogen
HBr	hydrobromic acid
HCl	hydrochloric acid
H_2SO_4	sulfuric acid

Abbreviation	Meaning
K_2CO_3	potassium carbonate
K_3PO_4	tripotassium phosphate
KOAc	potassium acetate
KOH	potassium hydroxide
LC/MS, LCMS or LC-MS	liquid chromatographic mass spectroscopy
LiO*t*-Bu	lithium tert-butoxide
LiOH	lithium hydroxide
MeOH	methanol
$MeSO_3H$	methanesulfonic acid
$MgSO_4$	magnesium sulfate
mL	milliliter
MOM	methoxymethyl
MS	mass spectroscopy
NBS	N-bromosuccinimide
NEt_3	triethylamine
NH_4Cl	ammonium chloride
NH_4OAc	ammonium acetate
Na_2CO_3	sodium carbonate
NaH	sodium hydride
$NaHCO_3$	sodium bicarbonate
NaOH	sodium hydroxide
Na_2SO_4	sodium sulfate
N_2	nitrogen
NH_4Cl	ammoniuim chloride
NMP	N-methylpyrrolidone
NMR	nuclear magnetic resonance
Pd	palladium
Pd/C	palladium on carbon
$PdCl_2(PPh_3)_2$	bis(triphenylphosphine)palladium(II) dichloride
$Pd_2(dba)_3$	tris(dibenzylideneacetone)dipalladium(0)
Pd(dppf)Cl_2 or Pd(dppf)Cl_2-CH_2Cl_2	[1,1'-bis(diphenylphosphino)ferrocene]dichloropalladium(II), complex with dichloromethane
PhMe	toluene
Psi	pounds per square inch pressure

Abbreviation	Meaning
QPhos	1,2,3,4,5-pentaphenyl-1′-(di-*tert*-butylphosphino)ferrocene
Rt or rt	room temperature
S-Phos, SPhos or Sphos	2-dicyclohexylphosphino-2',6'-dimethoxybiphenyl
S-Phos G_2	chloro(2-dicyclohexylphosphino-2',6'-dimethoxy-1,1'-biphenyl)(2'-amino-1,1'-biphenyl-2-yl) palladium(II)
TBAF	tetrabutylammonium fluoride
TBS	tert-butyldimethylsilyl
TEA, Et_3N or NEt_3	triethylamine
Tf	trifluoromethane sulfonyl or triflate
TFA	trifluoroacetic acid
THF	tetrahydrofuran
THP	tetrahydropyranyl
TIPS	tiisopropylsilane
TLC	thin layer chromatography
UPLC	Ulta performance liquid chromatography
XPhos Pd G4	ligand for classic cross-coupling reactions (CAS Number 1599466-81-5)

Example 1

Preparation of Compound 213



Step 1: To a dry round bottom flask were added: 6-bromo-3-methylsulfanyl-1,2,4-triazine (7.0 g, 34.0 mmol), 2-[4-chloro-2-(methoxymethoxy)phenyl]-4,4,5,5-tetramethyl-1,3,2-dioxaborolane (10 g, 33.5 mmol), K_2CO_3 (9.5 g, 68 mmol), $PddppfCl_2$ (3.8 g, 5.1 mmol). The mixture was degassed with argon for 10 min, then dioxane (100 mL) and water (20 mL) were added and the reaction was heated at 90 ºC for 16 h (overnight). Reaction was cooled down to rt, partitioned between EtOAc and water, organic parts were dried over Na_2SO_4, concentrated, purified by silica-gel column chromatography eluting with a gradient 0-20 % EtOAc in pentanes. Provided 6-[4-chloro-2-(methoxymethoxy)phenyl]-3-methylsulfanyl-1,2,4-triazine (4.5 g, 44% yield) as yellow solid. MS *m/z* 298.1, 300.1 $[M+H]^+$.

Step 2: A suspension of 6-[4-chloro-2-(methoxymethoxy)phenyl]-3-methylsulfanyl-1,2,4-triazine (350 mg, 1.18 mmol) was added 4,4,5,5-tetramethyl-2-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)-1,3,2-dioxaborolane (750 mg, 2.9 mmol), KOAc (575 mg, 5.86 mmol), X Phos Pd G4 (175 mg, 0.20 mmol) in dry dioxane (5 mL) was sparged with argon for 10 minutes, then heated to 90 °C under argon atmosphere for 2 h, after which complete conversion to 6-(2-(methoxymethoxy)-4-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)phenyl)-3-(methylthio)-1,2,4-triazine was observed. The reaction mixture was then cooled to room temperature and used directly in the next step. MS *m/z* 390.4 $[M+H]^+$.

Step 3: To the mixture from Step 2 above were added: 3-bromo-1,2,4-thiadiazole (0.15 g, 0.91 mmol), (2-dicyclohexylphosphino-2',4',6'-triisopropyl-1,1'-biphenyl)[2-(2'-amino-1,1'-biphenyl)]palladium(II) methanesulfonate (0.16 g, 0.181 mmol) and potassium carbonate (0.38 g, 2.72 mmol). The reaction was degassed via bubbling of nitrogen for 5 min and continued under nitrogen. 1,4-Dioxane (3 mL) and water (3 mL) were added to the mixture and the reaction was stirred at 90 °C for 2h. The reaction was cooled to room temperature, diluted with water and extracted with EtOAc. Combined organic layers were dried over $MgSO_4$, filtered and concentrated under reduced pressure. The residue was purified by column chromatography eluting with a gradient (0-100%) EtOAc/hexanes to afford 3-[3-(methoxymethoxy)-4-(3-methylsulfanyl-1,2,4-triazin-6-yl)phenyl]-1,2,4-thiadiazole (0.2 g, 60% yield). MS m/z 348.2 $[M+H]^+$.

Step 4: To a solution of 3-[3-(methoxymethoxy)-4-(3-methylsulfanyl-1,2,4-triazin-6-yl)phenyl]-1,2,4-thiadiazole (0.2 g, 0.6 mmol) in CH_2Cl_2 (5 mL) was added 3-chloroperoxybenzoic acid (0.2 g, 1.0 mmol) portion-wise. The reaction mixture was stirred at room temperature for 16 h. Solvent was removed under reduced pressure, the reminder was purified by column chromatography eluting with a gradient (0-10%) CH_2Cl_2/MeOH to afford 3-[3-(methoxymethoxy)-4-(3-methylsulfonyl-1,2,4-triazin-6-yl)phenyl]-1,2,4-thiadiazole (0.1 g, 50% yield); MS m/z 380.3 $[M+H]^+$.

Step 5: To a solution of 3-[3-(methoxymethoxy)-4-(3-methylsulfonyl-1,2,4-triazin-6-yl)phenyl]-1,2,4-thiadiazole (0.1 g, 0.3 mmol) in ACN (3 mL) were added (2S)-2-cyclopropylpiperazine (0.07 g, 0.6 mmol) and N,N-diisopropylethylamine (0.2 mL, 1 mmol). The reaction mixture was stirred at 80 °C for 30 min. Solvent was removed under reduced pressure, the reminder was purified by column chromatography eluting with a gradient (0-20%) CH_2Cl_2/MeOH to afford 3-[4-[3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl]-3-(methoxymethoxy)phenyl]-1,2,4-thiadiazole (0.07 g, 60% yield). MS m/z 426.5 $[M+H]^+$.

Step 6: To a solution of 3-[4-[3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl]-3-(methoxymethoxy)phenyl]-1,2,4-thiadiazole (0.07 g, 0.2 mmol) in CH_2Cl_2 (1 mL) and 2 drops of MeOH was added HCl (4 mol/L) in 1,4-dioxane (0.1 mL, 0.4 mmol). The reaction was stirred for 1 h until UPLC showed complete consumption of the starting material. The solvents were removed under reduced pressure, the reminder was purified by prep-HPLC eluting with a gradient 5-40% ACN in water (containing 0.1% formic acid) to provide 2-[3-[(3S)-3-cyclopropylpiperazin-1-yl]-1,2,4-triazin-6-yl]-5-(1,2,4-thiadiazol-3-yl)phenol (55 mg, 90% ield). MS m/z 382.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 10.99 (s, 1H), 10.39 (s, 1H), 9.16 (s, 1H), 9.10 (s, 1H), 8.00 (d, J = 8.1 Hz, 1H), 7.95 (d, J = 1.7 Hz, 1H), 7.87 (dd, J =

8.1, 1.6 Hz, 1H), 4.83 – 4.67 (m, 2H), 3.59 – 3.35 (m, 3H), 3.18 – 3.07 (m, 1H), 2.77 – 2.63 (m, 1H), 1.04 (tt, J = 8.4, 3.9 Hz, 1H), 0.67 (ddp, J = 17.3, 8.5, 4.2 Hz, 2H), 0.56 (dq, J = 9.9, 4.7 Hz, 1H), 0.45 (dq, J = 8.8, 4.5 Hz, 1H).

Using the procedure described for Example 1, above, additional compounds described herein may be prepared by substituting the appropriate starting material, suitable reagents and reaction conditions, obtaining compounds such as those selected from:

Cpd	Data
1	MS m/z 382.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.05 (s, 1H), 7.98 (s, 1H), 7.83 (d, J=8.7 Hz, 1H), 7.18-7.23 (m, 2H), 4.80 (br d, J=12.4 Hz, 1H), 4.64 (d, J=13.7 Hz, 1H), 3.12-3.22 (m, 2H), 3.01 (dd, J=12.9, 10.9 Hz, 1H), 2.79-2.89 (m, 1H), 1.94-2.03 (m, 1H), 0.82-0.95 (m, 1H), 0.57-0.67 (m, 2H), 0.31-0.43 (m, 2H); 3Hs not observed (2 NHs and OH).
2	MS m/z 409.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.10 (s, 1H), 8.81 (s, 1H), 7.98 (d, J=8.2 Hz, 1H), 7.70 (s, 1H), 7.68 (br d, J=8.2 Hz, 1H), 7.33 (s, 1H), 4.83 (br d, J=13.3 Hz, 1H), 4.72 (d, J=13.4 Hz, 1H), 3.12-3.24 (m, 2H), 3.02 (dd, J=12.8, 10.7 Hz, 1H), 2.83 (td, J=12.2, 2.5 Hz, 1H), 1.98 (td, J=9.2, 2.7 Hz, 1H), 0.83-0.95 (m, 1H), 0.57-0.67 (m, 2H), 0.30-0.43 (m, 2H); 2Hs not observed (NH and OH).
3	MS *m/z* 384.1 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.93 (s, 1H), 7.86 (s, 1H), 7.72 (d, J = 8.6 Hz, 1H), 7.11 – 7.07 (m, 2H), 4.73 – 4.52 (m, 2H), 3.09 – 2.95 (m, 2H), 2.77 – 2.65 (m, 2H), 2.34 (ddd, J = 10.4, 7.0, 3.0 Hz, 1H), 1.62 (h, J = 6.9 Hz, 1H), 0.95 (dd, J = 6.9, 1.9 Hz, 6H); 3Hs not observed (2 NHs and OH).
4	MS m/z 400.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.07 (s, 1H), 7.98 (d, J=1.8 Hz, 1H), 7.85 (d, J=8.5 Hz, 1H), 7.16-7.26 (m, 2H), 4.91-4.95 (m, 1H), 4.75 (d, J=13.3 Hz, 1H), 3.20 (d, J=13.0 Hz, 1H), 3.08 (td, J=12.4, 2.5 Hz, 1H), 2.81-2.94 (m, 2H), 2.66 (br d, J=10.1 Hz, 1H), 1.31 (d, J=12.4 Hz, 6H); 4Hs not observed (2 NHs and 2OHs).
5	MS m/z 364.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.79 (d, J = 10.5 Hz, 1H), 9.43 (d, J = 10.8 Hz, 1H), 9.14 (s, 1H), 8.10 (s, 2H), 7.87 (d, J = 8.0 Hz, 1H), 7.21-7.27 (m, 2H), 4.61-4.80 (m, 2H), 3.45-3.60 (m, 2H), 3.41 (dt, J = 12.8, 2.9 Hz, 1H), 2.99-3.11 (m, 1H), 2.61-2.72 (m, 1H), 1.10 (ddt, J = 13.0, 8.0, 4.4 Hz, 1H), 0.56-0.73 (m, 3H), 0.38-0.47 (m, 1H); 1H not observed.
6	MS m/z 380.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.08-9.15 (s, 1H), 8.31-8.43 (s, 2H), 8.01 (s, 2H), 7.78-7.87 (m, 1H), 7.21-7.25 (m, 1H), 7.18-7.21 (m, 1H), 4.97-5.09 (m, 2H), 3.47-3.59 (m, 2H), 3.37-3.46 (m, 1H), 3.23-3.31 (m, 1H), 2.87-2.96 (m, 1H), 0.92-1.05 (m, 2H), 0.76-0.88 (m, 2H); 2Hs not observed (NH and OH).

Cpd	Data
7	MS m/z 382.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.06 (br s, 1H), 7.98 (br s, 1H), 7.84 (d, J=7.8 Hz, 1H), 7.21 (br s, 2H), 4.80 (br d, J=12.5 Hz, 1H), 4.69 (d, J=12.5 Hz, 1H), 3.11-3.23 (m, 2H), 3.00 (br dd, J=10.6, 9.8 Hz, 1H), 2.73-2.90 (m, 1H), 1.87-2.09 (m, 1H), 0.78-0.97 (m, 1H), 0.55-0.71 (m, 2H), 0.27-0.48 (m, 2H); 3Hs not observed (2 NHs and OH).
8	MS m/z 427.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.12 (s, 1H), 8.81 (s, 1H), 7.95-8.08 (m, 1H), 7.63-7.78 (m, 2H), 7.34 (s, 1H), 4.95-5.07 (m, 1H), 4.03-4.14 (m, 1H), 3.12-3.21 (m, 2H), 2.91-3.08 (m, 2H), 2.71-2.89 (m, 1H), 1.34 (br d, J=13.9 Hz, 6H); 3Hs not observed (NH and 2OHs).
9	MS m/z 381.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.66 (d, J = 10.5 Hz, 1H), 9.36 (d, J = 10.8 Hz, 1H), 9.12 (s, 1H), 8.17 (s, 1H), 7.83-7.89 (m, 2H), 7.15-7.22 (m, 2H), 4.60-4.82 (m, 2H), 3.38-3.57 (m, 3H), 3.07 (q, J = 11.3 Hz, 1H), 2.63-2.71 (m, 1H), 1.05-1.15 (m, 1H), 0.55-0.73 (m, 3H), 0.39-0.47 (m, 1H).
10	MS m/z 411.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.10 (br s, 1H), 8.81 (s, 1H), 7.98 (br s, 1H), 7.61-7.77 (m, 2H), 7.33 (s, 1H), 4.64-4.78 (m, 2H), 3.12-3.22 (m, 2H), 2.81-2.95 (m, 2H), 2.38-2.57 (m, 1H), 1.72-1.82 (m, 1H), 0.95-1.18 (m, 6H); 2Hs not observed (NH and OH).
12	MS m/z 378.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.05 (s, 1H), 8.01 (s, 1H), 7.86 (s, 1H), 7.81 (d, J=8.1 Hz, 1H), 7.19 (br d, J=8.4 Hz, 1H), 7.16 (s, 1H), 4.79 (d, J=13.3 Hz, 1H), 4.68 (br d, J=13.3 Hz, 1H), 3.95 (s, 3H), 3.08-3.21 (m, 2H), 3.00 (dd, J=13.5, 12.1 Hz, 1H), 2.82 (td, J=12.8, 2.5 Hz, 1H), 1.96 (br td, J=8.5, 1.8 Hz, 1H), 0.81-0.96 (m, 1H), 0.54-0.70 (m, 2H), 0.30-0.42 (m, 2H); 2Hs not observed (NH and OH).
13	MS m/z 382.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.47 - 8.40 (m, 1H), 8.35 (br s, 1H), 7.70 (d, J = 8.1 Hz, 1H), 7.38 - 7.19 (m, 2H), 6.03 (br s, 1H), 5.20 - 4.98 (m, 1H), 4.39 - 4.25 (m, 1H), 3.78 - 3.38 (m, 5H), 1.48 - 1.37 (m, 6H); 4Hs not observed (2 NHs and 2OHs).
15	MS *m/z* 384.1 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.97 (s, 1H), 7.91 – 7.85 (m, 1H), 7.74 – 7.70 (m, 1H), 7.12 – 7.02 (m, 2H), 4.84 (d, J = 13.0 Hz, 1H), 4.74 – 4.69 (m, 1H), 3.30 – 3.25 (m, 1H), 3.20 – 3.16 (m, 1H), 2.98 (td, J = 12.9, 9.6 Hz, 2H), 2.77 (ddd, J = 11.2, 7.5, 3.3 Hz, 1H), 1.80 (h, J = 6.9 Hz, 1H), 1.02 (dd, J = 6.7, 1.9 Hz, 6H); 3Hs not observed (2 NHs and OH).
16	MS m/z 380.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.13 (s, 1H), 8.07 (s, 2H), 7.84 (d, J=8.1 Hz, 1H), 7.26 (d, J=8.5 Hz, 1H), 7.22 (s, 1H), 5.13 (br d, J=14.2 Hz, 1H), 5.00 (d, J=14.3 Hz, 1H), 3.57 (d, J=12.8 Hz, 1H), 3.45 (td, J=13.5, 3.1 Hz, 1H), 3.23-3.33 (m, 2H), 3.18 (dd, J=11.3, 2.1 Hz, 1H), 1.19 (s, 9H); 3Hs not observed (2 NHs and OH).
17	MS m/z 425.6 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.07 (s, 1H), 8.79 (s, 1H), 7.95 (d, J=8.2 Hz, 1H), 7.67 (s, 1H), 7.64 (d, J=8.2 Hz, 1H), 7.29 (s, 1H), 4.91 (br d, J=13.0 Hz, 1H), 4.76 (br d, J=13.0 Hz, 1H), 3.19 (br d, J=12.1 Hz, 1H), 3.08 (td, J=12.7, 2.9 Hz, 1H), 2.78-2.92 (m, 2H), 2.46 (br d, J=9.3 Hz, 1H), 1.06 (s, 9H); 2Hs not observed (NH and OH).

Cpd	Data
19	MS m/z 399.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.07 (s, 1H), 8.02 (s, 1H), 7.86 (s, 1H), 7.83 (d, J=7.9 Hz, 1H), 7.20 (dd, J=8.4, 1.4 Hz, 1H), 7.17 (d, J=1.4 Hz, 1H), 4.93 (br d, J=12.1 Hz, 1H), 4.78 (d, J=13.2 Hz, 1H), 3.24 (d, J=13.1 Hz, 1H), 3.11 (td, J=12.1, 3.5 Hz, 1H), 2.92 (t, J=12.1 Hz, 2H), 2.73 (br d, J=11.4 Hz, 1H), 1.34 (s, 3H), 1.31 (s, 3H); 3Hs not observed (NH and 2OHs).
20	MS m/z 366.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.01 (s, 1H), 8.10 - 7.92 (m, 2H), 7.79 (d, J = 8.1 Hz, 1H), 7.18 (s, 2H), 4.85 - 4.72 (m, 1H), 4.69 - 4.61 (m, 1H), 3.17 - 3.06 (m, 2H), 2.88 - 2.77 (m, 2H), 2.48 - 2.42 (m, 1H), 1.73 (qd, J = 6.8, 13.7 Hz, 1H), 1.06 (dd, J = 3.4, 6.8 Hz, 6H); 3Hs not observed (2 NHs and OH).
21	MS m/z 394.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.05 (s, 1H), 8.02 (s, 1H), 7.86 (s, 1H), 7.82 (d, J=8.2 Hz, 1H), 7.19 (d, J=8.2 Hz, 1H), 7.17 (s, 1H), 4.89-4.93 (m, 1H), 4.73 (d, J=12.8 Hz, 1H), 3.96 (s, 3H), 3.18 (br d, J=12.7 Hz, 1H), 3.07 (td, J=12.7, 2.5 Hz, 1H), 2.85 (dd, J=12.7, 10.2 Hz, 2H), 2.45 (br d, J=10.2 Hz, 1H), 1.06 (s, 9H); 2Hs not observed (NH and OH).
22	MS m/z 366.6 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.04 (s, 1H), 8.01 (br s, 2H), 7.81 (d, J = 8.2 Hz, 1H), 7.26 - 7.15 (m, 2H), 4.81 (d, J = 13.0 Hz, 1H), 4.68 (d, J = 13.7 Hz, 1H), 3.19 - 3.08 (m, 2H), 2.90 - 2.79 (m, 2H), 2.52 - 2.43 (m, 1H), 1.80 - 1.68 (m, 1H), 1.07 (d, J = 6.7 Hz, 6H); 3Hs not observed (2 NHs and OH).
23	MS m/z 352.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.02 (s, 1H), 8.06 - 7.92 (m, 2H), 7.78 (d, J = 8.1 Hz, 1H), 7.21 - 7.16 (m, 2H), 4.78 - 4.64 (m, 2H), 3.15 (s, 2H), 2.94 - 2.85 (m, 1H), 2.84 - 2.75 (m, 1H), 2.74 - 2.65 (m, 1H), 1.56 (s, 2H), 1.06 (t, J = 7.6 Hz, 3H); 3Hs not observed (2 NHs and OH).
24	MS m/z 352.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.61 (s, 2H), 7.73 (d, J = 8.2 Hz, 1H), 7.43 - 7.29 (m, 2H), 6.04 (s, 1H), 4.42 - 4.14 (m, 2H), 3.82 - 3.40 (m, 5H), 1.92 - 1.77 (m, 2H), 1.24 - 1.10 (m, 3H); 3Hs not observed (2 NHs and OH).
25	MS m/z 400.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.06 (s, 1H), 7.98 (d, J=1.4 Hz, 1H), 7.84 (d, J=8.7 Hz, 1H), 7.16-7.26 (m, 2H), 4.91 (d, J=13.4 Hz, 1H), 4.75 (d, J=13.4 Hz, 1H), 3.21 (d, J=12.8 Hz, 1H), 3.08 (td, J=12.8, 3.1 Hz, 1H), 2.82-2.93 (m, 2H), 2.67 (d, J=11.0 Hz, 1H), 1.31 (d, J=12.8 Hz, 6H); 4Hs not observed (2 NHs and 2OHs).
26	MS m/z 441.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.11 (s, 1H), 8.81 (s, 1H), 7.98 (d, J=8.2 Hz, 1H), 7.71 (d, J=2.0 Hz, 1H), 7.68 (dd, J=8.1, 2.0 Hz, 1H), 7.34 (s, 1H), 4.61 (dd, J=13.8, 4.3 Hz, 1H), 4.25-4.38 (m, 1H), 3.37-3.51 (m, 2H), 3.09-3.26 (m, 1H), 2.74-2.85 (m, 1H), 2.59 (s, 3H), 2.46 (dd, J=8.9, 4.3 Hz, 1H), 1.30 (d, J=6.4 Hz, 6H); 2Hs not observed (2OHs).
27	MS m/z 378.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.14 (d, J = 10.9 Hz, 1H), 8.96 (s, 1H), 8.09 (s, 2H), 7.87 (dd, J = 10.1, 4.0 Hz, 1H), 7.15-7.37 (m, 3H), 4.75-4.84 (m, 2H), 3.45-3.64 (m, 3H), 3.08-3.15 (m, 1H), 2.94 (s, 3H), 2.60-2.73 (m, 1H), 1.08-1.17 (m, 1H), 0.82-0.89 (m, 1H), 0.69-0.80 (m, 1H), 0.66-0.63 (m, 1H), 0.31-0.39 (m, 1H).

Cpd	Data
28	MS *m/z* 426.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.82 (s, 2H), 8.51 (s, 1H), 7.57 (s, 1H), 7.52 (d, J=12.1 Hz, 1H), 7.35 (s, 1H), 5.00 (d, J=13.3 Hz, 1H), 4.81 - 4.80 (m, 1H), 4.07 (m, 3H), 3.37-3.39 (m, 2H), 3.07-3.13 (m, 2H), 2.86-2.90 (m, 1H), 1.89-1.93 (m, 1H), 1.13 (d, J=6.85 Hz, 6H); 2Hs not observed (NH and OH), 1H from formic acid salt.
29	MS m/z 394.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.11 (s, 1H), 8.31 (s, 1H), 8.02 (s, 2H), 7.77-7.89 (m, 1H), 7.16-7.28 (m, 1H), 4.95-5.05 (m, 2H), 3.48-3.56 (m, 1H), 3.35-3.45 (m, 2H), 3.36 (s, 3H), 3.25-3.31 (m, 2H), 1.10-1.26 (m, 2H), 0.81-0.94 (m, 2H); 3Hs not observed (2 NHs and OH).
30	MS m/z 378.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.03 (s, 1H), 8.07 - 7.88 (m, 2H), 7.79 (d, J = 8.1 Hz, 1H), 7.22 - 7.16 (m, 2H), 4.72 - 4.63 (m, 2H), 3.24 - 3.13 (m, 2H), 2.97 - 2.87 (m, 1H), 2.85 - 2.72 (m, 2H), 2.47 - 2.37 (m, 1H), 2.19 - 2.09 (m, 2H), 2.05 - 1.84 (m, 4H); 3Hs not observed (2 NHs and OH).
31	MS m/z 366.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.13 (s, 1H), 8.14 (s, 2H), 7.87 (d, J = 8.5 Hz, 1H), 7.27 - 7.23 (m, 2H), 4.80 - 4.65 (m, 2H), 3.60 - 3.49 (m, 1H), 3.41 - 3.25 (m, 3H), 3.16 - 3.04 (m, 1H), 1.76 - 1.61 (m, 2H), 1.53 - 1.36 (m, 2H), 0.94 (t, J = 7.2 Hz, 3H); 3Hs not observed (2 NHs and OH).
32	MS *m/z* 400.3 $[M+H]^+$; $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.71-8.69 (m, 1H), 8.42 (s, 1H), 7.91 (d, J=1.6 Hz, 1H), 6.97-7.0 (m, 1H), 6.90 (d, J=12.4 Hz, 1H), 4.79-4.82 (m, 1H), 4.69 - 4.72 (m, 1H), 3.26-3.31 (m, 2H), 3.07-3.16 (m, 1H), 2.89-2.95 (m, 1H), 2.18-2.22 (m, 1H), 0.84-0.87 (m, 1H), 0.58-0.62 (m, 2H), 0.36-0.39 (m, 2H); 3Hs not observed (2 NHs and OH), 1H from formic acid salt.
33	MS m/z 380.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.14 (s, 1H), 8.14 (s, 2H), 7.87 (d, J = 8.5 Hz, 1H), 7.26 - 7.24 (m, 2H), 4.83 - 4.72 (m, 2H), 3.57 - 3.48 (m, 1H), 3.40 - 3.32 (m, 1H), 3.31 - 3.22 (m, 2H), 3.17 (s, 1H), 1.93 - 1.84 (m, 1H), 1.62 - 1.51 (m, 1H), 1.39 - 1.29 (m, 1H), 1.07 - 1.03 (m, 3H), 0.93 (t, J = 7.3 Hz, 3H); 3Hs not observed (2 NHs and OH).
34	MS m/z 380.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.04 (s, 1H), 8.13 - 7.88 (m, 2H), 7.81 (d, J = 8.1 Hz, 1H), 7.24 - 7.17 (m, 2H), 4.74 - 4.61 (m, 2H), 3.22 (d, J = 2.6 Hz, 1H), 3.05 - 2.88 (m, 2H), 2.45 - 2.39 (m, 1H), 2.39 - 2.36 (m, 3H), 2.31 - 2.23 (m, 1H), 2.07 - 2.00 (m, 1H), 1.10 (d, J = 6.9 Hz, 3H), 1.02 (d, J = 6.9 Hz, 3H); 2Hs not observed (NH and OH).
35	MS *m/z* 380.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.93 (s, 1H), 7.90 (s, 1H), 7.74 (s, 1H), 7.69 (d, J = 8.2 Hz, 1H), 7.07 (dd, J = 8.1, 1.8 Hz, 1H), 7.04 (d, J = 1.7 Hz, 1H), 4.71 (d, J = 12.7 Hz, 1H), 4.58 (d, J = 14.9 Hz, 1H), 3.83 (s, 3H), 3.09 – 2.96 (m, 2H), 2.80 – 2.70 (m, 2H), 2.39 (ddd, J = 10.4, 7.1, 3.0 Hz, 1H), 1.64 (h, J = 6.9 Hz, 1H), 0.96 (dd, J = 6.8, 2.0 Hz, 6H); 2Hs not observed (NH and OH).

Cpd	Data
36	MS m/z 400.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.23 (s, 0.5H), 8.49 (s, 1H), 8.39 (s, 1H), 7.94 (d, J=8.2 Hz, 0.5H), 7.71 (d, J=8.2 Hz, 0.5H), 7.27-7.36 (m, 2H), 6.02 (s, 0.5H), 4.80-4.87 (m, 1H), 4.12-4.33 (m, 1H), 3.42-3.88 (m, 5H), 2.05-2.23 (m, 1H), 1.85-2.00 (m, 1H), 1.65-1.83 (m, 1H); 3Hs not observed (2 NHs and OH).
37	MS m/z 366.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.13 (s, 1H), 8.02 (s, 2H), 7.85 - 7.82 (m, 1H), 7.25 - 7.19 (m, 2H), 5.00 - 4.88 (m, 2H), 3.58 - 3.49 (m, 2H), 3.37 (s, 1H), 3.32 - 3.23 (m, 2H), 1.81 - 1.70 (m, 2H), 1.65 - 1.51 (m, 2H), 1.07 (t, J = 7.3 Hz, 3H); 3Hs not observed (2 NHs and OH).
38	MS m/z 350.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.11 (s, 0.5 H), 8.35 (s, 1H), 8.26 (s, 1H), 7.82 (d, J=8.2 Hz, 0.5H), 7.59 (d, J=8.2 Hz, 0.5H), 7.14-7.26 (m, 2H), 5.83-5.95 (m, 1.5H), 5.49-5.69 (m, 2H), 4.80-4.87 (m, 1H), 3.91-4.09 (m, 2H), 3.31-3.69 (m, 4H); 3Hs not observed (2 NHs and OH).
39	MS m/z 369.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.13 (s, 1H), 8.04 (s, 1H), 7.87 (s, 1H), 7.83 (d, J=8.2 Hz, 1H), 7.22 (d, J=7.9 Hz, 1H), 7.18 (s, 1H), 4.93-5.01 (m, 2H), 3.48-3.61 (m, 2H), 3.24-3.38 (m, 3H), 1.73-1.92 (m, 2H), 1.16 (t, J=7.6 Hz, 3H); 2 Hs not observed (NH and OH).
40	MS m/z 366.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.01 (s, 1H), 8.10 - 7.92 (m, 2H), 7.79 (d, J = 8.1 Hz, 1H), 7.18 (s, 2H), 4.85 - 4.72 (m, 1H), 4.69 - 4.61 (m, 1H), 3.17 - 3.06 (m, 2H), 2.88 - 2.77 (m, 2H), 2.48 - 2.42 (m, 1H), 1.73 (qd, J = 6.8, 13.7 Hz, 1H), 1.06 (dd, J = 3.4, 6.8 Hz, 6H); 3Hs not observed (2 NHs and OH).
41	MS *m/z* 383.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.13 (s, 1H) 8.01 (s, 1H), 7.87-7.78 (m, 2H) 7.22-7.17 (m, 1H) 7.16 (s, 1H) 5.10-5.01 (m, 1H) 4.98-4.92 (m, 1H) 3.56 - 3.12 (m, 4H), 2.25-2.13 (m, 1H), 2.06-1.95 (m, 1H), 1.17 (d, J=6.9 Hz, 6H); 2Hs not observed (NH and OH).
42	MS m/z 338.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.09 (s, 1H), 7.96 (s, 2H), 7.81 (d, J = 8.1 Hz, 1H), 7.25 - 7.14 (m, 2H), 4.95 - 4.86 (m, 2H), 3.59 - 3.42 (m, 3H), 3.31 - 3.20 (m, 2H), 1.46 (d, J = 6.6 Hz, 3H); 3Hs not observed (2 NHs and OH).
43	MS m/z 378.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.05 (s, 1H), 7.94-8.13 (m, 2H), 7.82 (d, J=8.2 Hz, 1H), 7.23 (dd, J=8.2, 1.8 Hz, 1H), 7.20 (d, J=1.5 Hz, 1H), 3.94 (dd, J=6.8, 5.1 Hz, 2H), 3.79-3.83 (m, 2H), 3.02 (dd, J=6.8, 4.6 Hz, 2H), 1.71-1.85 (m, 6H), 1.57-1.65 (m, 2H); 3Hs not observed (2 NHs and OH).
44	MS m/z 391.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.15 (s, 1H), 8.82 (d, J = 6.4 Hz, 1H), 8.31 (s, 1H), 8.19 - 8.15 (m, 1H), 8.09 (d, J = 8.1 Hz, 1H), 7.66 - 7.60 (m, 2H), 4.94 - 4.69 (m, 2H), 3.62 - 3.50 (m, 1H), 3.44 - 3.37 (m, 1H), 3.37 - 3.26 (m, 1H), 3.16 - 3.08 (m, 2H), 2.82 (s, 3H), 2.14 - 2.01 (m, 1H), 1.08 (t, J = 6.9 Hz, 6H); 2Hs not observed (NH and OH).

Cpd	Data
46	MS m/z 380.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.10 (s, 1H), 7.95 (s, 2H), 7.81 (d, J = 8.1 Hz, 1H), 7.25 - 7.14 (m, 2H), 5.02 - 4.85 (m, 2H), 3.61 - 3.47 (m, 2H), 3.43 - 3.36 (m, 1H), 3.28 (s, 2H), 1.94 - 1.82 (m, 1H), 1.72 - 1.56 (m, 2H), 1.05 (d, J = 6.6 Hz, 6H); 3Hs not observed (2 NHs and OH).
47	MS m/z 383.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.06 (s, 1H), 8.02 (s, 1H), 7.86 (s, 1H), 7.82 (d, J=8.1 Hz, 1H), 7.19 (d, J=8.2 Hz, 1H), 7.16 (s, 1H), 4.83 (d, J=12.8 Hz, 1H), 4.70 (d, J=14.0 Hz, 1H), 3.11-3.21 (m, 2H), 2.79-2.93 (m, 2H), 2.44-2.58 (m, 1H), 1.76 (h, J=6.9 Hz, 1H), 1.08 (d, J=6.9 Hz, 6H); 2Hs not observed (NH and OH).
48	MS m/z 352.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.02 (s, 1H), 8.06 - 7.92 (m, 2H), 7.78 (d, J = 8.1 Hz, 1H), 7.21 - 7.16 (m, 2H), 4.78 - 4.64 (m, 2H), 3.15 (s, 2H), 2.94 - 2.85 (m, 1H), 2.84 - 2.75 (m, 1H), 2.74 - 2.65 (m, 1H), 1.56 (s, 2H), 1.06 (t, J = 7.6 Hz, 3H); 3Hs not observed (2 NHs and OH).
49	MS *m/z* 369.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.13 (s, 1H), 8.01 (s, 1H), 7.85 (s, 1H), 7.84 (d, J=8.1 Hz, 1H), 7.20 (d, J=6.6 Hz, 1H), 7.16 (d, J=1.5 Hz, 1H), 5.05 (dd, J=14.3, 2.7 Hz, 2H), 3.43-3.52 (m, 2H), 3.04 (dd, J=14.5, 11.6 Hz, 2H), 1.44 (d, J=6.6 Hz, 6H); 2Hs not observed (NH and OH).
50	MS m/z 364.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.05 (s, 1H), 8.07 (br s, 1H), 7.93 (br s, 1H), 7.82 (d, J=8.2 Hz, 1H), 7.22 (d, J=8.1 Hz, 1H), 7.19 (s, 1H), 3.93 (s, 2H), 3.85-3.91 (m, 2H), 2.87-2.96 (m, 2H), 2.00-2.09 (m, 2H), 1.94 (td, J=14.2, 7.2 Hz, 4H); 3Hs not observed (2 NHs and OH).
51	MS *m/z* 383.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.03 (s, 1H), 7.99 (s, 1H), 7.84 (s, 1H), 7.76 (d, J=7.0 Hz, 1H), 7.09-7.19 (m, 2H), 4.64 (br d, J=12.8 Hz, 2H), 2.80-2.91 (m, 2H), 2.30 – 2.38 (m, 4H), 1.23 (d, J=6.1 Hz, 6H); 2Hs not observed (NH and OH).
53	MS m/z 368.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.98 (s, 1H), 7.95 (s, 2H), 7.79 - 7.73 (m, 1H), 7.18 - 7.15 (m, 2H), 4.65 (t, J = 14.2 Hz, 2H), 3.51 - 3.41 (m, 2H), 3.41 - 3.40 (m, 3H), 3.19 - 3.10 (m, 2H), 3.04 - 2.97 (m, 1H), 2.96 - 2.82 (m, 2H); 3Hs not observed (2 NHs and OH).
55	MS m/z 350.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.11 (s, 1H), 8.14 (s, 2H), 7.87 (d, J = 7.9 Hz, 1H), 7.27 - 7.22 (m, 2H), 4.23 - 4.14 (m, 2H), 4.03 (s, 2H), 3.35 - 3.29 (m, 1H), 1.19 - 1.02 (m, 3H), 0.97 - 0.92 (m, 2H); 3Hs not observed (2 NHs and OH).
56	MS m/z 352.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.09-9.18 (m, 1H), 8.02-8.08 (m, 2H), 7.85 (d, J=8.2 Hz, 1H), 7.26 (dd, J=8.1, 1.7 Hz, 1H), 7.22 (d, J=1.5 Hz, 1H), 4.23 (br t, J=5.3 Hz, 2H), 4.06 (s, 2H), 3.44 (br d, J=5.5 Hz, 1H), 3.40-3.47 (m, 1H),1.50 (s, 6H); 3Hs not observed (2 NHs and OH).

Cpd	Data
57	MS m/z 368.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.05 (s, 1H), 7.98 (s, 1H), 7.85 (d, J=8.5 Hz, 1H), 7.16-7.26 (m, 2H), 3.97-4.06 (m, 2H), 3.66-3.72 (m, 1H), 3.02-3.13 (m, 2H), 0.84-0.98 (m, 1H), 0.63-0.78 (m, 4H); 3Hs not observed (2 NHs and OH).
58	MS *m/z* 381.2$[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.03 (s, 1H), 7.99 (s, 1H), 7.84 (s, 1H), 7.80 (d, J=8.1 Hz, 1H), 7.17 (br d, J=8.2 Hz, 1H), 7.14 (s, 1H), 4.36 (br d, J=12.5 Hz, 2H), 3.38 (s, 2H), 3.27 (s, 1H), 2.41 (s, 3H), 2.06-2.18 (m, 3H), 1.70 (br d, J=7.9 Hz, 2H); 1H not observed (OH).
59	MS m/z 380.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.07 (s, 1H), 8.09 (br s, 1H), 7.95 (br s, 1H), 7.84 (d, J=8.2 Hz, 1H), 7.24 (dd, J=8.2, 1.5 Hz, 1H), 7.21 (d, J=1.5 Hz, 1H), 4.93 (dd, J=13.7, 1.5 Hz, 1H), 4.81 (dd, J=13.7, 1.8 Hz, 1H), 4.43-4.51 (m, 1H), 3.54 (dd, J=9.8, 6.7 Hz, 1H), 3.12-3.24 (m, 2H), 2.84 (t, J=11.4 Hz, 1H), 2.39-2.55 (m, 2H), 2.18-2.27 (m, 1H), 1.82-1.91 (m, 2H); 3Hs not observed (2 NHs and OH).
60	MS m/z 366.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.11 (s, 1H), 8.00 - 7.94 (m, 2H), 7.85 - 7.81 (m, 1H), 7.73 (s, 1H), 7.25 - 7.16 (m, 2H), 5.05 (d, J = 13.9 Hz, 2H), 3.69 - 3.52 (m, 5H), 3.28 - 3.15 (m, 2H), 1.46 (d, J = 6.6 Hz, 6H); 1H not observed (NH or OH).
61	MS m/z 400.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.16 (s, 1H), 8.13 (s, 2H), 7.88 (d, J = 7.9 Hz, 1H), 7.73 (d, J = 7.2 Hz, 2H), 7.57 - 7.45 (m, 3H), 7.28 - 7.21 (m, 2H), 4.92 - 4.79 (m, 2H), 4.57 (br s, 1H), 3.79 - 3.61 (m, 2H), 3.54 - 3.47 (m, 1H), 3.36 - 3.24 (m, 1H); 3Hs not observed (2 NHs and OH).
62	MS m/z 401.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.17 (s, 1H), 9.15 - 9.00 (m, 2H), 8.49 - 8.33 (m, 2H), 8.15 (s, 2H), 7.95 - 7.77 (m, 1H), 7.26 (s, 2H), 5.02 - 4.78 (m, 3H), 3.89 - 3.70 (m, 2H), 3.66 - 3.55 (m, 1H), 3.36 - 3.25 (m, 1H); 3Hs not observed (2 NHs and OH).
63	MS m/z 364.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.07 (s, 1H), 8.28 - 7.91 (m, 2H), 7.85 (d, J = 8.1 Hz, 1H), 7.26 - 7.17 (m, 2H), 3.81 (br s, 4H), 2.66 (br s, 4H), 1.69 (br s, 1H), 0.47 (d, J = 5.8 Hz, 2H), 0.39 (d, J = 2.0 Hz, 2H); 2Hs not observed (NH and OH).
64	MS m/z 380.4 $[M+H]^+$
96	MS m/z 394.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.11 (s, 1H), 8.31 (s, 1H), 8.02 (s, 2H), 7.77-7.89 (m, 1H), 7.16-7.28 (m, 1H), 4.95-5.05 (m, 2H), 3.48-3.56 (m, 1H), 3.35-3.45 (m, 2H), 3.36 (s, 3H), 3.25-3.31 (m, 2H), 1.10-1.26 (m, 2H), 0.81-0.94 (m, 2H); 3Hs not observed (2 NHs and OH).
123	MS m/z 370.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.09 (s, 1H), 8.23 - 8.20 (m, 1H), 7.88 (d, J = 8.1 Hz, 1H), 7.25 - 7.22 (m, 1H), 7.20 (d, J = 8.2 Hz, 1H), 4.79 - 4.63 (m, 2H), 3.60 - 3.49 (m, 1H), 3.42 - 3.34 (m, 1H), 3.34 - 3.27 (m, 1H), 3.27 - 3.18 (m, 1H), 3.16 - 3.04 (m, 1H), 1.85 - 1.64 (m, 2H), 1.03 (t, J = 7.5 Hz, 3H); 3Hs not observed (2 NHs and OH).

Cpd	Data
125	MS m/z 398.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.06 (s, 1H), 7.98 (s, 1H), 7.85 (br d, J=8.4 Hz, 1H), 7.18-7.28 (m, 2H), 4.90 (d, J=13.2 Hz, 1H), 4.74 (d, J=13.3 Hz, 1H), 3.18 (d, J=12.5 Hz, 1H), 3.07 (t, J=12.5 Hz, 1H), 2.85 (t, J=11.6 Hz, 2H), 2.46 (d, J=11.6 Hz, 1H), 1.06 (s, 9H); 3Hs not observed (2 NHs and OH).
127	MS *m/z* 380.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.40 (s, 1H), 8.36 (s, 1H), 7.85 (s, 2H), 6.99 (s, 1H), 6.88 (s, 1H), 4.90 (d, J=13.2 Hz, 1H), 4.81 - 4.80 (m, 1H), 3.28-3.31 (m, 2H), 3.00-3.07 (m, 2H), 2.81-2.84 (m, 1H), 2.12, (s, 3H), 1.80-1.84 (m, 1H), 1.02 (dd, J=5.85, 0.85 Hz, 6H); 3Hs not observed (2 NHs and OH), 1H from formic acid salt.
133	MS m/z 397.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.21 (s, 1H), 9.00 (s, 1H), 8.09 (d, J=8.1 Hz, 1H), 7.61-7.70 (m, 2H), 7.51 (s, 1H), 4.99 (d, J=13.6 Hz, 1H), 4.91 (d, J=16.0 Hz, 1H), 3.53-3.61 (m, 2H), 3.47 (br s, 1H), 3.34-3.38 (m, 2H), 1.78-1.86 (m, 2H), 1.18 (t, J=7.2 Hz, 3H); 2Hs not observed (NH and OH).
134	MS *m/z* 378.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.09 (s, 1H), 8.02 (s, 2H), 7.82 (d, J = 8.2 Hz, 1H), 7.23 (dd, J = 8.2, 1.7 Hz, 1H), 7.19 (s, 1H), 4.99 – 4.93 (m, 2H), 3.50 – 3.44 (m, 1H), 3.19 (td, J = 12.5, 3.6 Hz, 1H), 2.60 (dd, J = 11.6, 3.2 Hz, 1H), 1.23 (s, 3H), 0.76 – 0.66 (m, 2H), 0.65 – 0.55 (m, 2H); 5Hs not observed (2 NHs and OH; CH_2 signal overlaps with MeOH signal).
135	MS m/z 350.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.10-9.17 (m, 1H), 8.33 (s, 1H), 8.02 (s, 2H), 7.80-7.90 (m, 1H), 7.19-7.30 (m, 1H), 4.68-4.78 (m, 2H), 4.21-4.30 (m, 2H), 3.42-3.53 (m, 2H), 2.13-2.22 (m, 2H), 2.00-2.10 (m, 2H); 3Hs not observed (2 NHs and OH).
136	MS m/z 364.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.78 (d, J = 10.5 Hz, 1H), 9.44 (d, J = 10.8 Hz, 1H), 9.16 (s, 1H), 8.13 (s, 2H), 7.87 (d, J = 8.0 Hz, 1H), 7.23-7.29 (m, 2H), 4.59-4.72 (m, 2H), 3.45-3.61 (m, 2H), 3.34-3.54 (m, 1H), 2.99-3.14 (m, 1H), 2.66-2.71 (m, 1H), 1.06-1.12 (m, 1H), 0.55-0.72 (m, 3H), 0.38-0.46 (m, 1H); 1H not observed (OH or NHs).
137	MS m/z 364.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.76 (d, J = 10.0 Hz, 1H), 9.42 (d, J = 11.0 Hz, 1H), 9.18 (s, 1H), 8.17 (s, 2H), 7.88 (d, J = 8.0 Hz, 1H), 7.26-7.31 (m, 2H), 4.57-4.71 (m, 2H), 3.48-3.63 (m, 2H), 3.34-3.59 (m, 1H), 2.97-3.13 (m, 1H), 2.64-2.71 (m, 1H), 1.09-1.18 (m, 1H), 0.57-0.73 (m, 3H), 0.37-0.46 (m, 1H); 1H not observed (OH or NHs).
143	MS m/z 392.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.10 (s, 1H), 9.05 (s, 1H), 8.54 (s, 1H), 7.99 (d, J = 8.1 Hz, 1H), 7.83 (s, 1H), 7.74 (d, J = 8.2 Hz, 1H), 4.92 - 4.71 (m, 2H), 3.57 - 3.48 (m, 1H), 3.43 - 3.36 (m, 1H), 3.33 - 3.24 (m, 1H), 3.16 - 3.09 (m, 2H), 2.59 (s, 3H), 2.12 - 2.01 (m, 1H), 1.11 - 1.06 (m, 6H); 2Hs not observed (NH and OH).
144	MS m/z 378.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.26 (s, 1H), 9.12 (s, 1H), 8.77 - 8.74 (m, 1H), 8.65 (d, J = 2.3 Hz, 1H), 8.01 (d, J = 8.2 Hz, 1H), 7.84 - 7.82 (m, 1H), 7.76 (dd, J = 1.5, 8.2 Hz, 1H), 4.91 - 4.73 (m, 2H), 3.48 - 3.37 (m, 2H), 3.28 (dd, J = 11.3, 13.9 Hz, 1H), 3.20 - 3.09 (m, 2H), 2.09-2.02 (m, 1H), 1.11 - 1.04 (m, 6H); 2Hs not observed (NH and OH).

Cpd	Data
145	MS m/z 392.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.11 (s, 2H), 8.64 (s, 1H), 7.99 (d, J = 8.1 Hz, 1H), 7.81 - 7.78 (m, 1H), 7.71 (dd, J = 1.4, 8.2 Hz, 1H), 4.85 (d, J = 13.4 Hz, 1H), 4.81 - 4.73 (m, 1H), 3.52 (s, 1H), 3.43 - 3.35 (m, 1H), 3.27 (d, J = 14.0 Hz, 1H), 3.17 (s, 2H), 2.55 (s, 3H), 2.11 - 2.01 (m, 1H), 1.10 - 1.04 (m, 6H); 2Hs not observed (NH and OH).
146	MS *m/z* 406.1 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.02 (s, 1H), 8.00 (s, 2H), 7.76 (d, J = 8.1 Hz, 1H), 7.17 (d, J = 1.7 Hz, 1H), 7.15 (s, 1H), 4.90 – 4.81 (m, 1H), 4.74 – 4.64 (m, 1H), 3.48 – 3.41 (m, 1H), 3.19 – 3.05 (m, 2H), 2.71 – 2.51 (m, 2H); 5Hs not observed (2 NHs and OH; CH_2 signal overlaps with MeOH signal).
148	MS m/z 366.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.14 (s, 1H), 8.14 (s, 2H), 7.87 (d, J = 8.2 Hz, 1H), 7.30 - 7.19 (m, 2H), 4.94 - 4.78 (m, 2H), 3.42 - 3.32 (m, 1H), 3.24 - 3.18 (m, 2H), 3.17 - 3.10 (m, 1H), 1.89 - 1.79 (m, 1H), 1.77 - 1.66 (m, 1H), 1.39 (d, J = 6.4 Hz, 3H), 1.09 - 0.98 (m, 3H); 3Hs not observed (2 NHs and OH).
149	MS m/z 392.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.14 (s, 1H), 9.13 - 9.11 (m, 1H), 8.05 - 7.99 (m, 2H), 7.91 (s, 1H), 7.79 (d, J = 8.2 Hz, 1H), 4.90 - 4.73 (m, 2H), 3.56 - 3.47 (m, 1H), 3.43 - 3.37 (m, 1H), 3.28 (dd, J = 11.2, 13.8 Hz, 1H), 3.17 - 3.10 (m, 2H), 2.57 (s, 3H), 2.11 - 1.99 (m, 1H), 1.11 - 1.04 (m, 6H); 2Hs not observed (NH and OH).
150	MS m/z 406.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.15 (s, 1H), 9.10 (s, 1H), 8.01 (d, J = 8.1 Hz, 1H), 7.99 - 7.97 (m, 1H), 7.90 (s, 1H), 7.82 - 7.78 (m, 1H), 4.86 - 4.65 (m, 2H), 3.49 - 3.40 (m, 1H), 3.18 - 2.98 (m, 2H), 2.98 - 2.87 (m, 1H), 2.83 (q, J = 7.6 Hz, 2H), 1.97 - 1.84 (m, 1H), 1.30 (t, J = 7.6 Hz, 3H), 1.08 - 1.01 (m, 7H); 2Hs not observed (NH and OH).
151	MS m/z 378.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.07 (s, 1H), 8.93 (d, J = 4.9 Hz, 2H), 8.08 (s, 1H), 8.02 - 7.97 (m, 2H), 7.48 (t, J = 4.8 Hz, 1H), 4.82 - 4.62 (m, 2H), 3.48 - 3.41 (m, 1H), 3.28 - 3.18 (m, 2H), 3.07 - 2.88 (m, 2H), 1.89 - 1.78 (m, 1H), 1.02 (dd, J = 3.2, 6.7 Hz, 6H); 2Hs not observed (NH and OH).
152	MS *m/z* 401.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.09 (s, 1H), 8.52 (br. s, 1H), 8.07 (d, J=1.45 Hz, 1H), 7.91 (s, 1H), 7.67 (d, J=12.4 Hz, 1H), 7.21 (d, J=6.55 Hz, 1H), 4.96 (d, J=13.2 Hz, 1H), 4.81 - 4.83 (m, 1H), 3.28-3.31 (m, 2H), 3.05-3.11 (m, 2H), 2.84-2.88 (m, 1H), 1.89-1.93 (m, 1H), 1.12 (d, J=6.7 Hz, 6H); 2Hs not observed (NH and OH), 1H from formic acid salt.
156	MS *m/z* 415.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.92 (s, 1H), 8.43 (s, 1H), 7.94 (d, J=1.45 Hz, 1H), 7.79 (s, 1H), 7.54 (d, J=12.4 Hz, 1H), 7.07 (d, J=6.55 Hz, 1H), 4.83 (d, J=13.2 Hz, 1H), 4.68 (d, J=13.5 Hz, 1H), 3.13-3.16 (m, 1H), 3.05-3.11 (m, 1H), 2.79-2.87 (m, 2H), 2.50-2.52 (m, 1H), 0.96 (s, 9H); 2Hs not observed (NH and OH), 1H from formic acid salt.

Cpd	Data
158	MS *m/z* 384.3, $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.97 (s, 1H), 8.40 (s, 1H), 7.95 (br. s, 2H), 7.58 (d, J=12.1 Hz, 1H), 7.14 (d, J=6.55 Hz, 1H), 4.85 (d, J=13.2 Hz, 1H), 4.69 - 4.72 (m, 1H), 3.26-3.30 (m, 1H), 3.15-3.18 (m, 1H), 2.92-2.99 (m, 2H), 2.70-2.73 (m, 1H), 1.74-1.79 (m, 1H), 1.00 (d, J=6.8 Hz, 6H); 3Hs not observed (2 NHs and OH), 1H from formic acid salt.
160	MS m/z 391.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.14 (s, 1H), 8.74 - 8.68 (m, 1H), 8.04 (dd, J = 2.3, 8.1 Hz, 1H), 7.95 (d, J = 7.9 Hz, 1H), 7.43 (d, J = 8.1 Hz, 1H), 7.28 - 7.25 (m, 2H), 5.07 (br d, J = 13.1 Hz, 1H), 4.97 (br d, J = 14.3 Hz, 1H), 3.60 - 3.51 (m, 2H), 3.31 - 3.22 (m, 2H), 3.13 (ddd, J = 3.2, 7.4, 10.9 Hz, 1H), 2.62 (s, 3H), 2.16 - 2.05 (m, 1H), 1.19 (dd, J = 4.3, 6.8 Hz, 6H); 2Hs not observed (NH and OH).
161	MS m/z 377.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.04 (s, 1H), 8.60 (d, J = 5.6 Hz, 2H), 7.93 (d, J = 8.1 Hz, 1H), 7.73 (d, J = 6.0 Hz, 2H), 7.36 - 7.32 (m, 2H), 4.80 (br d, J = 12.8 Hz, 1H), 4.67 (br d, J = 13.7 Hz, 1H), 3.17 - 3.06 (m, 2H), 2.88 - 2.76 (m, 2H), 2.44 (ddd, J = 2.8, 7.2, 10.3 Hz, 1H), 1.72 (qd, J = 6.8, 13.6 Hz, 1H), 1.06 (d, J = 6.7 Hz, 6H); 2Hs not observed (NH and OH).
163	MS m/z 398.1 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.79 (s, 1H), 7.76 (s, 1H), 7.59 (d, J=8.2 Hz, 1H), 6.92-7.07 (m, 2H), 4.53 (br d, J=13.1 Hz, 1H), 4.36 (d, J=14.6 Hz, 1H), 2.84-2.94 (m, 2H), 2.73 (dd, J=13.1, 10.7 Hz, 1H), 2.50-2.60 (m, 1H), 1.65-1.70 (m, 1H), 0.53-0.67 (m, 1H), 0.28-0.37 (m, 2H), 0.05-0.16 (m, 2H); 3Hs not observed (2 NHs and OH).
164	MS *m/z* 398.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.96 (s, 1H), 8.41 (s, 1H), 7.95 (br. s, 2H), 7.57 (d, J=12.1 Hz, 1H), 7.13 (d, J=6.55 Hz, 1H), 4.89 (d, J=13.2 Hz, 1H), 4.69 - 4.72 (m, 1H), 3.13-3.16 (m, 1H), 3.05-3.11 (m, 1H), 2.86-3.00 (m, 2H), 2.62-2.65 (m, 1H), 0.99 (s, 9H); 3Hs not observed (2 NHs and OH), 1H from formic acid salt.
165	MS m/z 338.0 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.16 (s, 1H), 8.10 (s, 2H), 7.87 (d, J=8.2 Hz, 1H), 7.27 (dd, J=8.2, 1.8 Hz, 1H), 7.24 (d, J=1.8 Hz, 1H), 4.89-4.95 (m, 2H), 4.17-4.28 (m, 2H), 3.40-3.66 (m, 3H), 1.46 (d, J=6.0 Hz, 3H); 3 Hs not observed (2 NHs and OH).
166	MS m/z 338.0 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.18 (s, 1H), 8.14-8.11 (m, 2H), 7.89 (d, J=8.1 Hz, 1H), 7.29 (br d, J=7.3 Hz, 1H), 7.25 (br d, J=6.7 Hz, 1H), 4.91-4.98 (m, 2H), 4.09-4.36 (m, 2H), 3.42-3.73 (m, 3H), 1.46 (br d, J=5.8 Hz, 3H); 3 Hs not observed (2 NHs and OH).
170	MS m/z 382.0 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.14 (s, 1H), 8.81 (s, 1H), 8.04 (d, J = 8.1 Hz, 1H), 7.62-7.70 (m, 2H), 4.93 (d, J = 13.7 Hz, 1H), 4.75-4.87 (m, 1H), 3.37 (t, J = 12.3 Hz, 2H), 3.26 (dd, J = 13.8, 10.6 Hz, 1H), 3.00-3.09 (m, 1H), 2.35 (d, J = 10.8 Hz, 1H), 0.89-1.04 (m, 1H), 0.68-0.78 (m, 2H), 0.45-0.51 (m, 2H); 2Hs not observed (OH and NH).
177	MS m/z 375.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.21 (s, 1H), 8.92 - 8.86 (m, 2H), 8.43 (d, J = 6.3 Hz, 2H), 8.17 - 8.11 (m, 1H), 7.67 - 7.53 (m, 2H), 5.06 - 4.97 (m, 1H), 4.96 - 4.89 (m, 1H), 3.64 - 3.49 (m, 3H), 3.29 - 3.21 (m, 1H), 2.76 - 2.66 (m, 1H), 1.16 - 1.06 (m, 1H), 0.89 - 0.76 (m, 2H), 0.66 - 0.51 (m, 2H); 2Hs not observed (NH and OH).

Cpd	Data
178	MS m/z 393.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.09 (s, 1H), 8.79 (d, J = 2.3 Hz, 1H), 8.59 (d, J = 5.0 Hz, 1H), 8.00 (d, J = 8.1 Hz, 1H), 7.80 - 7.71 (m, 1H), 7.38 (s, 1H), 7.30 (d, J = 8.1 Hz, 1H), 4.80 - 4.67 (m, 2H), 3.64 - 3.49 (m, 2H), 3.46 - 3.36 (m, 1H), 3.07 (br d, J = 11.7 Hz, 1H), 2.72 - 2.61 (m, 1H), 1.17 - 1.10 (m, 1H), 0.71 - 0.59 (m, 3H), 0.46 - 0.40 (m, 1H); 2Hs not observed (NH and OH).
179	MS m/z 403.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.11 (s, 1H), 7.92 (d, J = 8.1 Hz, 1H), 7.70 (br d, J = 8.2 Hz, 2H), 7.34 - 7.21 (m, 4H), 4.79 - 4.65 (m, 2H), 3.60 - 3.47 (m, 3H), 3.46 - 3.40 (m, 1H), 3.07 (br d, J = 9.8 Hz, 1H), 2.87 (s, 3H), 2.69 - 2.62 (m, 1H), 0.71 - 0.59 (m, 3H), 0.47 - 0.39 (m, 1H); 2Hs not observed (NH and OH).
185	MS m/z 383.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.10 (s, 1H), 7.99 – 7.93 (m, 1H), 7.92 (d, J = 3.3 Hz, 1H), 7.66 (d, J = 3.2 Hz, 1H), 7.58 (d, J = 7.3 Hz, 2H), 4.88-4.91 (m, 1H), 4.75 (d, J = 13.4 Hz, 1H), 3.13-3.24 (m, 2H), 2.91 (dd, J = 13.5, 10.3 Hz, 2H), 2.56 (t, J = 8.0 Hz, 1H), 1.72-1.83 (m, 1H), 1.09 (d, J = 6.8 Hz, 6H); 2Hs not observed (OH and NH).
186	MS m/z 433.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.09 (s, 1H), 7.94-7.99 (m, 2H), 7.57-7.63 (m, 2H), 6.77-7.03 (m, 1H), 4.86-4.93 (m, 1H), 4.74 (d, J = 13.3 Hz, 1H), 3.12-3.23 (m, 2H), 2.87-2.92 (m, 2H), 2.52-2.56 (m, 1H), 1.77 (h, J = 6.9 Hz, 1H), 1.09 (d, J = 6.8 Hz, 6H); 2Hs not observed (OH and NH).
187	MS *m/z* 394.1 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: ^1H NMR (DMSO-d_6) δ: 12.93 (br s, 1H), 11.31 (br s, 1H), 9.09 (s, 1H), 7.96-8.19 (m, 2H), 7.82-7.89 (m, 1H), 7.17-7.27 (m, 2H), 4.68 (dd, J=8.2, 5.8 Hz, 1H), 4.51-4.64 (m, 3H), 4.36 (d, J=12.8 Hz, 1H), 4.26 (d, J=12.5 Hz, 1H), 3.19-3.35 (m, 2H), 2.88 (dd, J=13.0, 9.3 Hz, 1H), 2.81 (dt, J=11.5, 3.1 Hz, 1H), 2.54 (dd, J=9.5, 2.7 Hz, 1H), 2.29 (td, J=11.5, 4.0 Hz, 1H), 2.11 (s, 3H).
188	MS m/z 417.3, 419.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.15 (s, 1H), 9.00 (s, 1H), 7.97 (d, J = 8.1 Hz, 1H), 7.84 (s, 1H), 7.56 (d, J = 1.8 Hz, 1H), 7.51 (dd, J = 8.1, 1.8 Hz, 1H), 4.65 (d, J = 12.6 Hz, 1H), 4.56 (d, J = 12.6 Hz, 1H), 2.97-3.09 (m, 2H), 2.62-2.83 (m, 2H), 2.29-2.42 (m, 1H), 1.66 (h, J = 6.7 Hz, 1H), 0.97 (d, J = 6.8 Hz, 6H); 1H not observed (OH or NH).
189	MS m/z 417.3, 419.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.22 (s, 1H), 9.00 (s, 1H), 7.94-8.02 (m, 2H), 7.52 (d, J = 1.8 Hz, 1H), 7.46 (dd, J = 8.1, 1.8 Hz, 1H), 4.64 (d, J = 12.5 Hz, 1H), 4.55 (d, J = 12.5 Hz, 1H), 2.95-3.08 (m, 2H), 2.61-2.81 (m, 2H), 2.33-2.36 (m, 1H), 1.65 (h, J = 6.8 Hz, 1H), 0.97 (d, J = 6.8 Hz, 6H)); 1H not observed (OH or NH).
192	MS m/z 376.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.22 - 9.19 (m, 1H), 9.08 (s, 1H), 8.80 (d, J = 5.5 Hz, 1H), 7.99 - 7.96 (m, 2H), 7.81 - 7.77 (m, 1H), 7.75 (dd, J = 1.7, 8.2 Hz, 1H), 4.82 - 4.79 (m, 1H), 4.79 - 4.64 (m, 1H), 3.22 - 3.12 (m, 2H), 3.02 (dd, J = 10.6, 13.2 Hz, 1H), 2.84 (dt, J = 3.4, 12.2 Hz, 1H), 2.01 - 1.93 (m, 1H), 0.94 - 0.80 (m, 1H), 0.61 (td, J = 4.0, 7.6 Hz, 2H), 0.41 - 0.31 (m, 2H); 2Hs not observed (NH and OH).

Cpd	Data
196	MS m/z 401.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.19 (s, 1H), 9.00 (s, 1H), 7.95 (d, J = 8.1 Hz, 1H), 7.79 (d, J = 3.1 Hz, 1H), 7.49 (d, J = 1.8 Hz, 1H), 7.42 (dd, J = 8.1, 1.8 Hz, 1H), 4.64 (d, J = 12.6 Hz, 1H), 4.55 (d, J = 12.6 Hz, 1H), 2.96-3.08 (m, 2H), 2.65-2.79 (m, 2H), 2.32-2.37 (m, 1H), 1.65 (h, J = 6.7 Hz, 1H), 0.97 (d, J = 6.8 Hz, 6H); 1H not observed (OH or NH).
199	MS m/z 408.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.20 (s, 1H), 9.00 (s, 1H), 8.02 (d, J = 8.1 Hz, 1H), 7.57 (d, J = 1.6 Hz, 1H), 7.54 (dd, J = 8.1, 1.7 Hz, 1H), 4.67 (d, J = 12.5 Hz, 1H), 4.57 (d, J = 12.7 Hz, 1H), 3.00-3.10 (m, 2H), 2.69-2.83 (m, 2H), 2.29-2.37 (m, 1H), 1.58-1.73 (m, 1H), 1.07-1.31 (m, 5H), 0.98 (d, J = 6.9, 6H); 1H not observed (OH or NH).
200	MS m/z 413.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.11 (s, 1H), 9.00 (s, 1H), 7.94 (d, J = 8.1 Hz, 1H), 7.52 (d, J = 1.8 Hz, 1H), 7.46 (dd, J = 8.2, 1.8 Hz, 1H), 6.67 (s, 1H), 4.65 (d, J = 12.7 Hz, 1H), 4.56 (d, J = 12.6 Hz, 1H), 3.89 (s, 3H), 2.98-3.09 (m, 2H), 2.62-2.83 (m, 2H), 2.37 (s, 1H), 1.66 (h, J = 6.8 Hz, 1H), 0.98 (d, J = 6.9, 6H); 1H not observed (OH or NH).
203	MS m/z 378.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.02 (s, 1H), 8.09 - 7.88 (m, 2H), 7.79 (d, J = 8.2 Hz, 1H), 7.23 - 7.15 (m, 2H), 4.75 (d, J = 13.4 Hz, 1H), 4.64 (d, J = 13.1 Hz, 1H), 3.23 (dt, J = 3.1, 12.9 Hz, 1H), 2.97 - 2.90 (m, 2H), 2.82 (dd, J = 10.8, 13.3 Hz, 1H), 2.35 (dt, J = 3.2, 12.1 Hz, 1H), 2.22 - 2.06 (m, 2H), 1.86 (br d, J = 12.7 Hz, 1H), 1.79 - 1.64 (m, 3H), 1.47 - 1.30 (m, 2H); 2Hs not observed (NH and OH).
204	MS m/z 414.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.15 (s, 1H), 8.95-9.12 (m, 1H), 7.97 (d, J = 8.2 Hz, 1H), 7.24-7.63 (m, 3H), 4.73 (d, J = 13.1 Hz, 1H), 4.64 (d, J = 13.0 Hz, 1H), 4.20 (s, 3H), 3.17 (s, 3H), 2.81-3.02 (m, 2H), 1.76 (s, 1H), 1.00 (d, J = 6.8 Hz, 6H).
205	MS m/z 434.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.33 (s, 1H), 9.03 (s, 1H), 8.04 (d, J = 8.1 Hz, 1H), 7.51-7.81 (m, 3H), 4.66 (d, J = 12.6 Hz, 1H), 4.56 (d, J = 12.7 Hz, 1H), 2.97-3.09 (m, 2H), 2.77 (dd, J = 12.6, 10.5 Hz, 1H), 2.62-2.71 (m, 1H), 2.28-2.41 (m, 1H), 1.65 (h, J = 6.7 Hz, 1H), 0.97 (d, J = 6.8, 6H); 1H not observed (OH or NH).
207	MS m/z 438.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.21 (s, 1H), 9.05 (s, 1H), 7.97 (d, J = 8.2 Hz, 1H), 7.81 (d, J = 1.8 Hz, 1H), 7.73 (dd, J = 8.3, 1.8 Hz, 1H), 7.11 (s, 1H), 4.65 (d, J = 12.6 Hz, 1H), 4.55 (d, J = 12.6 Hz, 1H), 4.01 (s, 3H), 3.96 (s, 3H), 2.90-3.10 (m, 2H), 2.76 (dd, J = 12.7, 10.4 Hz, 1H), 2.61-2.72 (m, 1H), 2.33-2.40 (m, 1H), 1.65 (h, J = 6.7 Hz, 1H), 0.97 (d, J = 6.8 Hz, 6H); 1H not observed (OH or NH).
208	MS m/z 392.3 $[M+H]^+$.
209	MS m/z 393.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.13 (s, 1H), 8.28 (d, J = 5.3 Hz, 1H), 7.99 (d, J = 8.1 Hz, 1H), 7.63 (d, J = 5.3 Hz, 1H), 7.43 - 7.34 (m, 3H), 5.00 - 4.78 (m, 1H), 3.43 (d, J = 6.7 Hz, 2H), 3.37 - 3.29 (m, 2H), 3.11 (dt, J = 3.5, 12.9 Hz, 1H), 2.45 (dt, J = 3.1, 10.0 Hz, 1H), 1.02 (tdd, J = 4.5, 8.5, 12.9 Hz, 1H), 0.81 - 0.71 (m, 2H), 0.58 - 0.46 (m, 2H); 2Hs not observed (NH and OH).

Cpd	Data
210	MS m/z 431.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.04 (s, 1H), 8.46 (s, 1H), 7.89 (d, J = 8.2 Hz, 1H), 7.57 - 7.51 (m, 2H), 6.68 (s, 1H), 4.79 (d, J = 12.1 Hz, 1H), 4.68 (d, J = 13.1 Hz, 1H), 4.20 (t, J = 7.6 Hz, 4H), 3.20 - 3.10 (m, 2H), 2.98 (dd, J = 10.9, 12.7 Hz, 1H), 2.85 - 2.77 (m, 1H), 2.50 (quin, J = 7.6 Hz, 2H), 1.97 - 1.90 (m, 1H), 0.86 (dt, J = 4.1, 8.5 Hz, 1H), 0.64 - 0.57 (m, 2H), 0.41 - 0.31 (m, 2H); 2Hs not observed (NH and OH).
215	MS m/z 396.0 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ 9.03 (s, 1H), 7.98 (s, 1H), 7.82 (d, J=8.8 Hz, 1H), 7.20 (d,J=6.8 Hz, 2H), 4.83 (d, J=13.2 Hz, 1H), 4.70 (d, J=12.8 Hz, 1H), 2.84 (t, J=12.0 Hz, 2H), 2.66 (t, J=12.0, 1H), 1.94 (t, J=9.2 Hz, 1H), 1.21 (d, J=6.4 Hz, 3H), 0.85 (q, J=3.2 Hz, 1H), 0.60 (d, J=8.0 Hz, 2H), 0.36 (q, J=14.8 Hz, 2H); 3Hs not observed (2 NHs and OH).
216	MS m/z: 380.0 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ: 9.03 (s, 1H), 8.01(s, 2H), 7.80(d, J= 8.4 Hz, 1H), 7.22-7.18(m, 2H), 4.89-4.84 (m, 1H), 4.71-4.68 (m, 1H), 2.93-2.81 (m, 1H), 2.69-2.59 (m, 2H), 2.49-2.46 (m, 1H), 1.73-1.71 (m, 1H), 1.21 (d, J= 6.4 Hz, 3H), 1.07 (d, J= 6.2 Hz, 6H); 3Hs not observed (2 NHs and OH).
217	MS m/z: 398.0 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ 9.03 (s, 1H), 7.95 (d, J=2.0 Hz, 1H), 7.80 (d, J=8.4 Hz, 1H), 7.18 (d, J=6.8 Hz, 2H), 4.76 (t, J=14.4 Hz, 1H), 4.60 (s, 1H), 2.97 (s, 1H), 2.69 (m, 3H), 1.76 (q, J=6.8 Hz, 1H), 1.24 (d, J=6.4 Hz, 3H), 1.08 (t, J=6.4 Hz, 6H); 3Hs not observed (2 NHs and OH).
218	MS m/z: 384.0 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ 9.03 (s, 1H), 7.97 (d, J=1.6 Hz, 1H), 7.82 (d, J=8.8 Hz, 1H), 7.20 (d,J=6.4 Hz, 2H), 4.83 (d, J=12.8 Hz, 1H), 4.73 (d, J=12.8 Hz, 1H), 2.91 (m, 1H), 2.63 (q, J=11.2 Hz, 3H), 1.55 (m, 2H), 1.20 (d, J=6.0 Hz, 3H), 1.07 (t, J=7.6 Hz, 3H); 3Hs not observed (2 NHs and OH).
219	MS m/z: 450.1 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ: 9.08 (s, 1H), 7.93 (d, J = 7.6 Hz, 1H), 7.71 (s, 1H), 7.66 (d, J = 9.2 Hz, 1H), 6.96 (s, 1H), 4.75 (d, J = 12.8 Hz, 1H), 4.10 (s, 3H), 4.01 (s, 3H), 2.90-2.84 (m, 2H), 2.68 (t, J = 11.8 Hz, 1H), 1.98 (t, J = 11.2 Hz, 1H), 1.31 (s, 1H), 1.22 (d, J = 5.2 Hz, 3H), 0,91-0.86 (m, 1H), 0.61 (d, J = 7.6 Hz, 2H), 0.36 (d, J = 14.8 Hz, 2H); 2Hs not observed (NH and OH).
220	MS m/z: 450.1 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ: 9.08 (s, 1H), 7.93 (d, J = 7.6 Hz, 1H), 7.71 (s, 1H), 7.66 (d, J = 9.2 Hz, 1H), 6.96 (s, 1H), 4.75 (d, J = 12.4 Hz, 1H), 4.10 (s, 3H), 4.01 (s, 3H), 2.90-2.84 (m, 2H), 2.68 (t, J = 12.0 Hz, 1H), 1.98 (t, J = 11.2 Hz, 1H), 1.31 (s, 1H), 1.22 (d, J = 6.4 Hz, 3H), 0,90-0.83 (m, 1H), 0.61 (d, J = 8.0 Hz, 2H), 0.42-0.33 (m, 2H); 3Hs not observed (2 NHs and OH).
221	MS m/z: 396.0 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ: 7.45 (s, 1H), 6.40 (d, J=2.0 Hz, 1H), 6.23 (t, J=6.8 Hz, 1H), 5.61 (d, J=6.4 Hz, 2H), 4.82 (t, J=1.2 Hz, 1H), 4.67 (d, J=12.8 Hz, 1H), 2.82 (m, 2H), 2.63 (q, J=2.4 Hz, 1H), 1.94 (m, 1H), 1.19 (d, J=6.4 Hz, 3H), 0.82 (m, 1H), 0.58 (m, 2H), 0.35 (m, 2H); 3Hs not observed (2 NHs and OH).

Cpd	Data
222	MS m/z: 398.0 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ: 9.03 (s, 1H), 7.97 (d, J=2.0 Hz, 1H), 7.82 (d, J=8.8 Hz, 1H), 7.20 (d,J=6.8 Hz, 2H), 4.86 (s, 1H), 4.69 (t, J=12.0 Hz, 1H), 2.88 (m, 1H), 2.66 (m, 2H), 2.50 (m, 1H), 1.72 (m, 1H), 1.21 (d, J=6.4 Hz, 3H), 1.08 (q, J=0.8 Hz, 6H); 3Hs not observed (2 NHs and OH).
223	MS m/z: 384. $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ 9.05 (s, 1H), 7.98 (s, 1H), 7.83 (d, J=8.8 Hz, 1H), 7.21 (d,J=6.8 Hz, 2H), 4.84 (d, J=13.6 Hz, 1H), 4.74 (d, J=13.2 Hz, 1H), 2.91 (s, 1H), 2.63 (q, J=12.4 Hz, 3H), 1.56 (s, 2H), 1.21 (d, J=6.0 Hz, 3H), 1.07 (t, J=7.2 Hz, 3H); 3Hs not observed (2 NHs and OH).
224	MS m/z: 378.1 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ: 9.04 (s, 1 H), 8.10-7.89 (m, 2H), 7.80 (d, J= 8.0 Hz, 1 H), 7.18(dd, J= 1.6 Hz, 1H), 7.18(d, J= 1.6 Hz, 1H), 4.87-4.82 (m, 1H), 4.72-4.66 (m, 1H), 2.86-2.84 (m, 2H), 2.71-2.58 (m, 1H), 2.02-1.92 (m, 1H), 1.21(d, J= 6.0 Hz, 3H), 0.91-0.80(m, 1H), 0.60-0.59 (m, 2H), 0.41-0.27 (m, 2H); 3Hs not observed (2 NHs and OH).
225	MS m/z: 378. $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ: 9.04 (s, 1H), 8.09-7.91(m, 2H), 7.08(d, J= 8.0 Hz, 1H), 7.21(dd, J= 1.8 Hz, 1H), 7.18(d, J= 1.6 Hz, 1H), 4.85-4.82 (m, 1H), 4.72-4.68 (m, 1H), 2.84-2.81(m, 2H), 2.72-2.62(m, 1H), 1.99-1.92(m, 1H), 1.21(d, J= 6.4 Hz, 3H), 0.90-0.78(m, 1H), 0.60-0.58 (m, 2H), 0.38-0.31 (m, 2H); 3Hs not observed (2 NHs and OH).
226	MS m/z: 420.1 $[M+H]^+$; ^1H NMR (400 MHz, DMSO-d_6) δ: 11.17 (s, 1H), 9.04 (s, 1H), 8.87 (s, 1H), 7.96 (d, J = 8.4 Hz, 1H), 7.82 (d, J = 1.6 Hz, 1H), 7.75 (q, J = 8.0 Hz, 1H), 7.47 (s, 1H), 4.65 (q, J = 11.2 Hz, 2H), 3.98 (s, 3H), 2.54-2.76 (m, 4H), 1.93 (d, J = 19.6 Hz, 1H), 1.08 (d, J = 6.4 Hz, 3H), 0.76-0.79 (m, 1H), 0.44 (d, J = 4.8 Hz, 2H), 0.24-0.31 (m, 2H).
227	MS m/z: 422.1 $[M+H]^+$; ^1H NMR (400 MHz, DMSO-d_6) δ: 11.21 (s, 1H), 9.05 (s, 1H), 8.89 (d, J=0.8 Hz, 1H), 7.98 (d, J = 8.0 Hz, 1H), 7.84 (d, J = 1.6 Hz, 1H), 7.76 (q, J= 8.0 Hz, 1H), 7.49 (d, J=1.2 Hz, 1H), 4.67 (q, J = 36.0 Hz, 2H), 4.01 (d, J=7.6 Hz, 3H), 2.75-2.77 (m, 1H), 2.57-2.66 (m, 2H), 2.42-2.46 (m, 2H), 1.64-1.69 (m, 1H), 1.10 (d, J = 6.4 Hz, 3H), 1.00 (d, J = 3.6 Hz, 3H), 0.98 (d, J=3.6 Hz, 3H).
228	MS m/z: 408.2 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ: 8.97 (s, 1H), 8.75 (s, 1H), 7.85 (d, J=8.4 Hz, 1H), 7.57 (t, J = 6.0 Hz, 2H), 7.17 (s, 1H), 4.75 (q, J= 48.0 Hz, 2H), 4.02 (s, 3H), 2.88-2.93 (m, 1H), 2.57-2.71 (m, 3H), 1.48-1.55 (m, 2H), 1.18 (d, J = 6.4 Hz, 3H), 1.03 (d, J = 7.6 Hz, 3H); 2Hs not observed (NH and OH).
229	MS m/z: 380.0 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.06 (s, 1 H), 8.13-7.92(m, 2H), 7.83(d, J= 8.4 Hz, 1H), 7.23(dd, J= 1.0 Hz, 1H), 7.20 (s, 1H), 4.87-4.86 (m, 1H), 4.71-4.70 (m, 1H), 2.93-2.82 (m, 1H), 2.69-2.64 (m, 2H), 2.53-2.48 (m, 1H), 1.77-1.70 (m, 1H), 1.22(d, J= 3.0 Hz, 3H), 1.09-1.07(m, 6H); 3Hs not observed (2 NHs and OH).

Cpd	Data
230	MS m/z: 422.0 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ: 9.02 (s, 1H), 8.80 (s, 1H), 7.90 (d, J=8.4 HZ, 1H), 7.62 (q, J = 14.8 Hz, 2H), 7.22 (s, 1H), 4.92 (q, J= 12.4 Hz, 1H), 4.76 (d, J= 11.6 Hz, 1H), 4.07 (s, 3H), 2.94 (s, 1H), 2.65-2.76 (m, 2H), 2.55-2.56 (m, 1H), 1.73-1.75 (m, 1H), 1.24 (d, J = 6.0 Hz, 3H), 1.08 (d, J = 6.4 Hz, 6H); 2Hs not observed (NH and OH).
231	MS *m/z* 392.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.31 (s, 1H), 9.08 (s, 1H), 8.22 (s, 1H), 7.93 (s, 1H), 7.84 (d, J = 8.0 Hz, 1H), 7.22 (d, J = 8.0 Hz, 1H), 7.21 (s, 1H), 4.69-4.52 (m, 2H), 4.12-4.08 (m, 1H), 3.17 (d, J = 5.0 Hz, 3H), 2.08-1.78 (m, 6H), 1.28-1.00 (m, 4H); 2Hs not observed (2 NHs).
232	MS m/z 379.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ 11.01 (s, 1H), 9.14 (s, 1H), 9.07 (s, 1H), 7.96 (d, J = 8.1 Hz, 1H), 7.58 (d, J = 1.7 Hz, 1H), 7.53 (dd, J = 8.1, 1.6 Hz, 1H), 7.03 (s, 1H), 4.81 – 4.65 (m, 2H), 3.45 (ddt, J = 17.6, 14.2, 6.9 Hz, 3H), 3.15 – 3.07 (m, 1H), 2.73 – 2.64 (m, 1H), 2.41 (s, 3H), 1.04 (tp, J = 8.7, 4.7 Hz, 1H), 0.67 (ttd, J = 13.4, 8.7, 4.7 Hz, 2H), 0.56 (dq, J = 9.9, 4.7 Hz, 1H), 0.44 (dq, J = 8.6, 4.5 Hz, 1H).
233	MS m/z 365.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ 9.05 (s, 1H), 8.26 (s, 1H), 7.98 (d, J = 8.1 Hz, 1H), 7.63 (d, J = 1.7 Hz, 1H), 7.58 (dd, J = 8.1, 1.6 Hz, 1H), 7.42 (s, 1H), 4.76 – 4.60 (m, 2H), 3.42 – 3.21 (m, 3H), 2.96 (td, J = 12.9, 3.5 Hz, 1H), 2.43 (td, J = 9.9, 3.3 Hz, 1H), 1.01 – 0.92 (m, 1H), 0.60 (dtt, J = 13.1, 8.6, 4.0 Hz, 2H), 0.47 (dq, J = 9.6, 4.5 Hz, 1H), 0.41 – 0.34 (m, 1H); 2Hs not observed (NH and OH).
234	MS m/z: 408.1 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4 and $CDCl_3$) δ: 9.05 (s, 1H), 8.81 (d, J=0.8 Hz, 1H), 7.93 (d, J=8.4 Hz, 1H), 7.62-7.65 (m, 2H), 7.24 (d, J=0.8 Hz, 1H), 4.86 (d, J= 12.8 Hz, 1H), 4.78 (d, J= 13.6 Hz, 1H), 4.07 (s, 3H), 2.93-2.97 (m, 1H), 2.63-2.75 (m, 3H), 1.53-1.59 (m, 2H), 1.23 (d, J = 6.4 Hz, 3H), 1.08 (t, J = 7.6 Hz, 3H); 2Hs not observed (NH and OH).
235	MS m/z: 420.0 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4 and $CDCl_3$) δ: 9.06 (s, 1H), 8.79 (s, 1H), 7.93 (d, J=8.0 Hz, 1H), 7.61-7.65 (m, 2H), 7.25 (d, J=0.8 Hz, 1H), 4.87 (q, J= 16.0 Hz, 2H), 4.06 (s, 3H), 4.92 (t, J=3.0 Hz, 2H), 2.68-2.74 (m, 1H), 1.98-2.04 (m, 1H), 1.23 (q, J = 16.0 Hz, 3H), 0.83-0.89 (m, 1H), 0.60-0.63 (m, 2H), 0.36-0.37 (m, 2H); 2Hs not observed (NH and OH).
236	MS *m/z* 410.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.02 (s, 1H), 7.97 (s, 1H), 7.82 (d, J = 8.5 Hz, 1H), 7.20 (d, J = 8.5 Hz, 1H), 7.19 (s, 1H), 4.73-4.69 (m, 2H), 2.95-2.86 (m, 1H), 2.78-2.73 (m, 1H), 2.62 (t, J = 11.5 Hz, 1H), 2.52 (t, J = 13.0 Hz, 1H), 2.41-2.32 (m, 1H), 2.17-2.10 (m, 2H), 2.03-1.86 (m, 4H), 1.20 (d, J = 6.0 Hz, 3H); 3Hs not observed (2 NHs and OH).
238	MS *m/z* 409.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.03 (s, 1H), 7.99 (s, 1H), 7.84 (s, 1H), 7.80 (d, J = 8.5 Hz, 1H), 7.17 (d, J = 8.0 Hz, 1H), 7.14 (s, 1H), 4.77-4.67 (m, 2H), 2.92-2.86 (m, 1H), 2.76 (t, J = 9.5 Hz, 1H), 2.63 (t, J = 13.0 Hz, 1H), 2.53 (t, J = 13.0 Hz, 1H), 2.41-2.32 (m, 1H), 2.16-2.09 (m, 2H), 2.02-1.85 (m, 4H), 1.20 (d, J = 6.5 Hz, 3H); 2Hs not observed (NH and OH).

Cpd	Data
239	MS m/z 384.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ 11.00 (s, 1H), 10.39 (s, 1H), 9.10 (s, 1H), 8.01 (d, J = 8.1 Hz, 1H), 7.95 (s, 1H), 7.88 (d, J = 1.5 Hz, 1H), 4.83 (dd, J = 38.0, 13.7 Hz, 2H), 3.49 – 3.38 (m, 2H), 3.21 (tt, J = 19.0, 11.0 Hz, 3H), 1.98 (h, J = 6.8 Hz, 1H), 1.07 (dd, J = 6.9, 4.3 Hz, 6H); 1 H not observed (NH or OH).
240	MS *m/z* 406.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.06 (s, 1H), 7.85 (d, J = 8.5 Hz, 1H), 7.63 (s, 1H), 7.40 (d, J = 8.5 Hz, 1H), 7.39 (s, 1H), 6.66 (s, 1H), 4.75 (d, J = 13.5 Hz, 2H), 3.96 (s, 3H), 2.96-2.90 (m, 1H), 2.82-2.76 (m, 1H), 2.66 (t, J = 13.0 Hz, 1H), 2.55 (t, J = 13.0 Hz, 1H), 2.42-2.33 (m, 1H), 2.18-2.11 (m, 2H), 2.04-1.86 (m, 4H), 1.22 (d, J = 6.5 Hz, 3H); 2Hs not observed (NH and OH).
245	MS m/z 396.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ 10.95 (s, 1H), 9.09 (s, 1H), 7.98 (d, J = 8.1 Hz, 1H), 7.89 (d, J = 1.5 Hz, 1H), 7.82 (dd, J = 8.2, 1.6 Hz, 1H), 4.80 – 4.68 (m, 2H), 3.52 – 3.32 (m, 4H), 3.11 (dq, J = 14.2, 8.1 Hz, 1H), 2.80 (s, 3H), 2.70 (t, J = 6.2 Hz, 1H), 1.05 (tp, J = 8.8, 4.8 Hz, 1H), 0.68 (dtq, J = 17.2, 8.6, 4.6, 4.0 Hz, 2H), 0.56 (dq, J = 9.8, 4.6 Hz, 1H), 0.45 (dq, J = 8.7, 4.5 Hz, 1H); 1H not observed (NH or OH).
246	MS m/z 395.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ 9.11 (s, 1H), 7.98 (s, 1H), 7.91 (d, J = 8.2 Hz, 1H), 7.63 (d, J = 1.6 Hz, 1H), 7.55 (dd, J = 8.3, 1.6 Hz, 1H), 4.78 – 4.66 (m, 2H), 3.45 (td, J = 14.1, 13.3, 6.2 Hz, 3H), 3.11 (dtd, J = 12.8, 8.8, 4.4 Hz, 1H), 2.74 (s, 3H), 2.71 – 2.64 (m, 1H), 1.04 (tt, J = 8.3, 3.9 Hz, 1H), 0.67 (dtq, J = 17.2, 8.6, 4.4 Hz, 2H), 0.56 (dq, J = 10.0, 4.7 Hz, 1H), 0.44 (dq, J = 8.7, 4.5 Hz, 1H); 2Hs not observed (NH and OH).
247	MS *m/z* 394.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.02 (s, 1H), 8.10-7.90 (s, 2H), 7.80 (d, J = 8.5 Hz, 1H), 7.21 (d, J = 8.5 Hz, 1H), 7.19 (s, 1H), 4.86 (d, J = 12.5 Hz, 1H), 4.68 (d, J = 12.5 Hz, 1H), 2.93-2.84 (m, 1H), 2.76 (t, J = 11.5 Hz, 1H), 2.62 (t, J = 12.5 Hz, 1H), 2.50 (t, J = 13.0 Hz, 1H), 1.21 (d, J = 6.5 Hz, 3H), 1.06 (s, 9H); 3Hs not observed (2 NHs and OH).
248	MS m/z 412.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.00 (s, 1H), 7.95 (s, 1H), 7.80 (d, J = 9.0 Hz, 1H), 7.18 (d, J = 9.0 Hz, 1H), 7.17 (s, 1H), 4.68 (d, J = 13.0 Hz, 1H), 3.74 (t, J = 5.5 Hz, 1H), 3.67 (q, J = 5.0 Hz, 2H), 3.58 (t, J = 5.0 Hz, 1H), 2.94-2.86 (m, 1H), 2.77 (t, J = 12.0 Hz, 1H), 2.63 (t, J = 12.5 Hz, 1H), 2.52 (dd, J = 10.5, 2.0 Hz, 1H), 1.22 (d, J = 6.5 Hz, 3H), 1.05 (s, 9H).
249	MS *m/z* 364.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.03 (s, 1H), 8.10-7.90 (s, 2H), 7.80 (d, J = 8.0 Hz, 1H), 7.21 (d, J = 8.0 Hz, 1H), 7.18 (s, 1H), 5.94-5.86 (m, 1H), 5.40 (d, J = 17.5 Hz, 1H), 5.28 (d, J = 11.0 Hz, 1H), 4.77-4.70 (m, 2H), 3.42-3.39 (m, 1H), 2.98-2.91 (m, 1H), 2.77 (t, J = 12.5 Hz, 1H), 2.65 (t, J = 12.5 Hz, 1H), 1.21 (d, J = 6.5 Hz, 3H); 3Hs not observed (2 NHs and OH).
250	MS m/z 383.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.23 (br s, 1H), 9.40 (s, 1H), 9.15 (s, 1H), 9.12 (br s, 1H), 9.06 (s, 1H), 7.96 (d, J = 8.1 Hz, 1H), 7.44 (dd, J = 8.2, 1.8 Hz, 1H), 7.40 (s, 1H), 4.85 (d, J = 13.8 Hz, 1H), 4.77 (d, J = 14.1 Hz, 1H), 3.55 – 3.33 (m, 2H), 3.32 – 3.06 (m, 3H), 2.12 – 1.88 (m, 1H), 1.07 (app t, J = 6.0 Hz, 6H).

Cpd	Data
251	MS m/z 405.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ 9.12 (s, 1H), 8.27 (d, J = 5.3 Hz, 1H), 7.97 (d, J = 8.1 Hz, 1H), 7.40 (dd, J = 8.1, 1.8 Hz, 1H), 7.35 (d, J = 1.8 Hz, 1H), 7.30 (dd, J = 5.3, 1.6 Hz, 1H), 7.09 (d, J = 1.5 Hz, 1H), 4.81 – 4.65 (m, 2H), 3.52 – 3.38 (m, 3H), 3.18 (s, 3H), 3.15 – 3.05 (m, 1H), 2.75 – 2.63 (m, 1H), 1.04 (tq, J = 8.7, 4.3 Hz, 1H), 0.68 (dddd, J = 15.3, 12.6, 8.2, 4.4 Hz, 2H), 0.56 (dq, J = 9.8, 4.6 Hz, 1H), 0.44 (dq, J = 8.5, 4.5 Hz, 1H); 2Hs not observed (NH and OH).
252	MS m/z 381.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ 11.04 (s, 1H), 9.19 (d, J = 4.7 Hz, 1H), 9.12 (s, 1H), 8.00 – 7.93 (m, 2H), 7.71 (d, J = 1.6 Hz, 1H), 7.65 (dd, J = 8.2, 1.7 Hz, 1H), 4.81 – 4.65 (m, 2H), 3.51 – 3.40 (m, 3H), 3.16 – 3.04 (m, 1H), 2.70 (d, J = 9.9 Hz, 1H), 1.04 (ddt, J = 13.0, 8.6, 4.5 Hz, 1H), 0.67 (dtt, J = 21.3, 8.6, 4.2 Hz, 2H), 0.56 (dq, J = 9.8, 4.6 Hz, 1H), 0.45 (dq, J = 8.4, 4.5 Hz, 1H); 1H not observed (NH or OH).
253	MS m/z 397.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.18 (br s, 1H), 9.01 (s, 1H), 8.48 (s, 1H), 7.97 (d, J = 7.9 Hz, 1H), 7.18 – 7.12 (m, 2H), 4.65 (d, J = 12.6 Hz, 1H), 4.55 (d, J = 12.7 Hz, 1H), 3.16 – 2.96 (m, 2H), 2.75 (dd, J = 12.6, 10.5 Hz, 1H), 2.68 (td, J = 11.6, 3.3 Hz, 1H), 2.40 (s, 3H), 2.37-2.33 (m, 1H), 1.65 (q, J = 6.7 Hz, 1H), 0.97 (dd, J = 6.8, 2.2 Hz, 6H); 1 H not observed (NH or OH).
254	MS *m/z* 344.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.37 (s, 1H), 8.27 (s, 1H), 7.15 (d, J = 8.5 Hz, 1H), 7.12 (s, 1H), 7.12 (d, J = 8.5 Hz, 1H), 4.09 (d, J = 13.0 Hz, 1H), 3.91 (d, J = 13.0 Hz, 1H), 2.93 (t, J = 5.0 Hz, 1H), 2.87-2.84 (m, 1H), 2.77 (t, J = 5.0 Hz, 1H), 2.11-2.04 (m, 1H), 1.98 (t, J = 12.5 Hz, 1H), 1.83 (t, J = 13.0 Hz, 1H), 1.70 (d, J = 11.0 Hz, 1H), 0.39 (d, J = 4.0 Hz, 2H), 0.25 (s, 9H).
255	MS *m/z* 344.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.19 (s, 1H), 9.09 (s, 1H), 8.87 (s, 1H), 8.49 (s, 1H), 7.97 (d, J = 8.0 Hz, 1H), 7.88 (s, 1H), 7.75 (d, J = 8.0 Hz, 1H), 7.45 (s, 1H), 4.95 (d, J = 13.5 Hz, 1H), 4.80 (d, J = 13.5 Hz, 1H), 3.98 (s, 3H), 3.40-3.25 (m, 3H), 3.17-3.09 (m, 1H), 1.50 (d, J = 6.0 Hz, 3H), 1.12 (s, 9H).
256	MS m/z: 419.0 $[M+H]^+$; ^1H NMR (400 MHz, DMSO-d_6) δ: 11.25 (m, 1H), 9.03 (s, 1H), 8.24 (d, J=5.2 Hz, 1H), 7.94 (d,J=8.4 Hz, 1H), 7.33 (m, 3H), 7.08 (d, J=0.8 Hz, 1H), 4.63 (q, J=34.4 Hz, 2H), 3.90 (s, 3H), 2.74 (q, J=1.6 Hz, 2H), 2.55 (d, J=12.4 Hz, 1H), 1.93 (m, 1H), 1.07 (d, J=6.4 Hz, 3H), 0.77 (m, 1H), 0.45 (m, 2H), 0.27 (m, 3H).
257	MS m/z: 421.0 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ 11.28 (s, 1H), 9.03 (s, 1H), 8.24 (d, J=5.2 Hz, 1H), 7.95 (d, J=8.4 Hz, 1H), 7.33 (m, 3H), 7.08 (d, J=0.8 Hz, 1H), 4.64 (q, J=25.6 Hz, 2H), 3.87 (s, 3H), 2.74 (m, 1H), 2.59 (q, J=12.4 Hz, 1H), 2.43 (m, 2H), 1.64 (m, 1H), 1.07 (t, J=6.4 Hz, 3H), 0.97 (m, 6H); 1 H not observed.
258	MS m/z 398.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 10.21 (s, 1H), 9.20 (s, 1H), 8.02 (d, J=8.2 Hz, 1H), 7.98 (s, 1H), 7.97 (br s, 1H), 5.22 (br d, J=11.9 Hz, 1H), 5.09 (d, J=14.0 Hz, 1H), 3.57 (d, J=11.9 Hz, 1H), 3.35-3.48 (m, 2H), 3.19-3.29 (m, 2H), 1.25 (s, 9H); 2 Hs not observed (NH and OH).

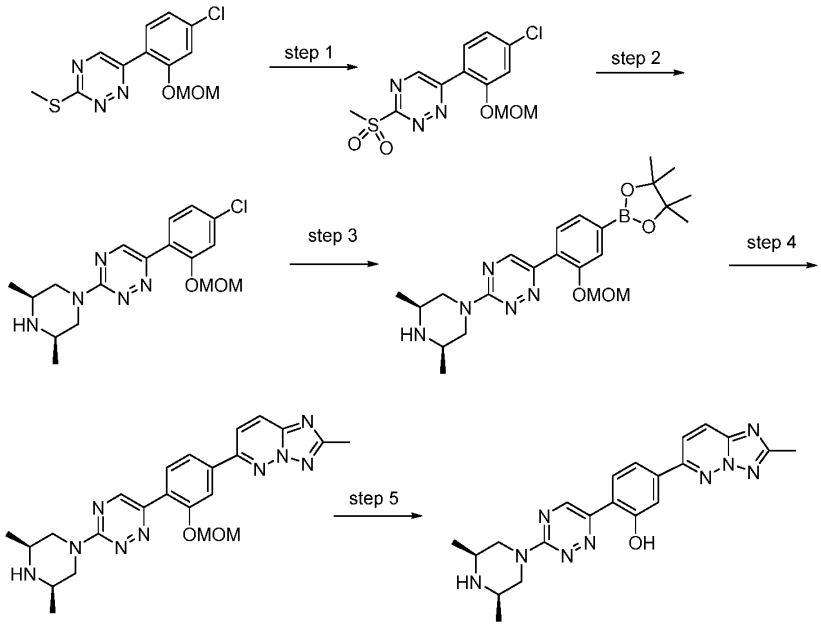
Cpd	Data
259	MS m/z 381.4 $[M+H]^+$; ^1H NMR (500 MHz, $CDCl_3$) δ: 11.35 (br s, 1H), 9.45 (br s, 1H), 9.13 (s, 1H), 8.62 (s, 1H), 7.98 (d, J = 8.1 Hz, 1H), 7.74 (s, 1H), 7.72 – 7.60 (m, 1H), 4.91 (d, J = 13.9 Hz, 1H), 4.83 (d, J = 14.2 Hz, 1H), 4.00 (s, 3H), 3.58-3.52 (m, 2H), 3.50 – 3.42 (m, 2H), 3.35-3.32 (m, 1H), 2.10 (q, J = 6.8 Hz, 1H), 1.13 (t, J = 6.3 Hz, 6H).
260	MS m/z 381.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.03 (br s, 1H), 9.27 (br s, 1H), 9.15 (br s, 1H), 8.56 (s, 1H), 7.92 (d, J = 8.0 Hz, 1H), 7.54 (s, 1H), 7.42 (d, J = 8.1 Hz, 1H), 4.85 (d, J = 13.9 Hz, 1H), 4.77 (d, J = 14.1 Hz, 1H), 4.11 (s, 3H), 3.54 – 3.34 (m, 2H), 3.28-3.23 (m, 1H), 3.15 (d, J = 10.4 Hz, 2H), 2.14 – 1.94 (m, 1H), 1.08 (t, J = 6.1 Hz, 6H).
261	MS m/z 381.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.17 (br s, 1H), 9.21-9.36 (m, 1H), 9.10 (s, 1H), 7.99 (d, J = 8.3 Hz, 1H), 7.97 (s, 1H), 7.25-7.23 (m, 2H), 4.86 (d, J = 13.9 Hz, 1H), 4.78 (d, J = 14.5 Hz, 1H), 4.13 (s, 3H), 3.55 – 3.36 (m, 3H), 3.30-3.25 (m 1H), 3.17-3.15 (m, 1H), 2.04 (h, J = 6.8 Hz, 1H), 1.08 (t, J = 6.5 Hz, 6H).
262	MS m/z 381.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.17 (br s, 1H), 9.36 (br s, 1H), 9.10 (s, 1H), 8.30 – 7.74 (m, 2H), 7.25-7.23 (m, J = 7.9 Hz, 2H), 4.86 (d, J = 13.9 Hz, 1H), 4.78 (d, J = 14.5 Hz, 1H), 4.13 (s, 3H), 3.49 (t, J = 13.0 Hz, 1H), 3.42-3.39 (m, 2H), 3.29-3.25 (m, 1H), 3.15-3.13 (m, 1H), 2.04 (q, J = 6.8 Hz, 1H), 1.08 (t, J = 6.5 Hz, 6H).
265	MS m/z 384.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.21 (br s, 1H), 9.47 (s, 1H), 9.12 (s, 1H), 8.74 (br s, 1H), 8.03 (d, J = 8.4 Hz, 1H), 7.77-7.77 (m, 2H), 4.83 (d, J = 13.7 Hz, 1H), 4.75 (d, J = 13.9 Hz, 1H), 3.35-3.32 (m, 2H), 3.18-3.10 (m, 3H), 1.94-1.92 (m, 1H), 1.05 (dd, J = 7.0, 3.9 Hz, 6H).
266	MS m/z: 407.1 $[M+H]^+$; ^1H NMR (400 MHz, methanol-d_4) δ: 9.07 (s, 1H), 8.18 (d, J=6.0 Hz, 1H), 7.94 (d, J=8.0 Hz, 1H), 7.30-7.33 (m, 2H), 7.26 (q, J=8.0 Hz, 1H), 7.07 (d, J=0.8 Hz, 1H), 4.83-4.89 (m, 1H), 4.74 (d, J=11.6 Hz, 1H), 3.96 (s, 3H), 2.85-2.89 (m, 1H), 2.58-2.67 (m, 3H), 1.52-1.57 (m, 2H), 1.19 (d, J = 6.4 Hz, 3H), 1.06 (t, J=7.4 Hz, 3H); 2Hs not observed (NH and OH).
267	MS m/z: 384.0 $[M+H]^+$; ^1H NMR (400 MHz, DMSO-d_6) δ: 9.00 (d, J=9.6 Hz, 2H), 8.02 (d, J=8.0 Hz, 1H), 7.62-7.66 (m, 2H), 4.66 (q, J=24.0 Hz, 2H), 2.75 (d, J=6.8 Hz, 1H), 2.52-2.57 (m, 2H), 1.38-1.45 (m, 2H), 1.23 (s, 1H), 1.07 (d, J = 6.0 Hz, 3H), 0.95 (t, J=7.4 Hz, 3H); 2Hs not observed (NH and OH).
268	MS m/z: 397.9 $[M+H]^+$; ^1H NMR (400 MHz, DMSO-d_6) δ: 11.28 (s, 1H), 8.99 (d, J=9.6 Hz, 2H), 8.02 (d, J=8.0 Hz, 1H), 7.64 (t, J=12.6 Hz, 2H), 4.67 (q, J=7.8 Hz, 2H), 2.75 (s, 1H), 2.56-2.65 (m, 2H), 2.41-2.49 (m, 1H), 1.64-1.66 (m, 1H), 1.23(s, 1H), 1.08 (d, J = 6.0 Hz, 3H), 0.97 (t, J=5.2 Hz, 6H).
269	MS m/z: 396.1 $[M+H]^+$; ^1H NMR (400 MHz, DMSO-d_6) δ: 11.24 (s, 1H), 9.04 (s, 1H), 8.99 (s, 1H), 8.02 (d, J=8.0 Hz, 1H), 7.65 (d, J=16.4 Hz, 2H), 4.70 (d, J=24.0 Hz, 2H), 2.85 (s, 2H), 1.22 (s, 2H), 1.14 (s, 3H), 0.81 (s, 1H), 0.50 (s, 2H), 0.31 (d, J = 19.6 Hz, 2H); 1 H not observed (NH or OH).

Cpd	Data
270	MS *m/z* 383.1 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.59 (s, 1H), 9.34 (s, 1H), 9.08 (s, 1H), 8.62 (s, 1H), 7.96 (d, J = 8.5 Hz, 1H), 7.80 (s, 1H), 7.37 (s, 2H), 4.84 (d, J = 13.9 Hz, 1H), 4.76 (d, J = 14.1 Hz, 1H), 3.51 (t, J = 12.6 Hz, 1H), 3.39 (d, J = 12.8 Hz, 1H), 3.28 (dd, J = 13.8, 11.0 Hz, 1H), 3.17 – 3.09 (m, 2H), 2.06 (h, J = 6.8 Hz, 1H), 1.07 (d, J = 6.8 Hz, 6H).
271	MS *m/z* 421.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.72 (s, 1H), 9.43 (s, 1H), 9.12 (s, 1H), 7.99 (d, J = 8.0 Hz, 1H), 7.46 – 7.40 (m, 2H), 7.31 (s, 1H), 7.06 (s, 1H), 4.85 – 4.76 (m, 2H), 3.98 (s, 3H), 3.59 – 3.50 (m, 1H), 3.43 – 3.35 (m, 1H), 3.30 (dd, J = 14.0, 11.1 Hz, 1H), 3.13 (d, J = 10.6 Hz, 2H), 2.52 (s, 3H), 2.08 (h, J = 6.9 Hz, 1H), 1.08 (d, J = 7.0 Hz, 6H).
272	MS m/z 408.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.25 (s, 1H), 8.69 (s, 1H), 7.91 (d, J=7.9 Hz, 1H), 7.38 (s, 1H), 7.26 (br d, J=7.9 Hz, 1H), 4.62 (d, J=12.2 Hz, 1H), 4.52 (br d, J=11.9 Hz, 1H), 2.91-3.04 (m, 2H), 2.71 (dd, J=12.5, 10.5 Hz, 1H), 2.64 (td, J=11.6, 3.6 Hz, 1H), 2.31 (ddd, J=10.0, 6.7, 3.4 Hz, 1H), 1.63 (dq, J=13.4, 6.7 Hz, 1H), 0.96 (dd, J=6.7, 1.2 Hz, 6H); 2 Hs not observed (NH and OH).
273	MS m/z 408.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.10 (s, 1H), 8.72 (s, 1H), 7.98 (d, J=7.6 Hz, 1H), 7.68 (s, 1H), 7.62 (d, J=8.5 Hz, 1H), 4.84 (d, J=13.4 Hz, 1H), 4.76 (d, J=14.2 Hz, 1H), 3.49 (t, J=12.4 Hz, 1H), 3.39 (d, J=12.5 Hz, 1H), 3.26 (dd, J=13.1, 11.0 Hz, 1H), 3.06-3.18 (m, 2H), 2.04 (dq, J=14.0, 6.7 Hz, 1H), 1.08 (t, J=6.7 Hz, 6H); 2 Hs not observed (NH and OH).
275	MS m/z 419.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.75 (s, 1H), 7.53 (d, J=7.9 Hz, 1H), 7.28 (s, 1H), 6.72 (dd, J=8.2, 1.5 Hz, 1H), 6.70 (s, 1H), 4.53 (d, J=12.8 Hz, 1H), 4.42 (br d, J=12.8 Hz, 1H), 3.93 (dd, J=8.5, 8.0 Hz, 2H), 3.64 (dd, J=8.5, 8.0 Hz, 2H), 2.86-2.99 (m, 2H), 2.76 (dd, J=12.8, 10.7 Hz, 1H), 2.64 (s, 3H), 2.59 (td, J=12.5, 3.2 Hz, 1H), 1.74 (td, J=11.0, 3.7 Hz, 1H), 0.56-0.72 (m, 1H), 0.35-0.44 (m, 2H), 0.02-0.20 (m, 2H); 2 Hs not observed (NH and OH).
276	MS m/z 404.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.11 (s, 1H), 7.93 (s, 1H), 7.82 (d, J=8.2 Hz, 1H), 7.15 (d, J=8.2 Hz, 1H), 7.11 (s, 1H), 4.95 (dd, J=14.5, 2.5 Hz, 1H), 4.86-4.89 (m, 1H), 4.14-4.23 (m, 2H), 3.61-3.77 (m, 1H), 3.52-3.59 (m, 1H), 3.48 (dd, J=14.0, 10.7 Hz, 1H), 3.14-3.22 (m, 3H), 2.72-2.81 (m, 2H), 2.67 (td, J=10.2, 3.4 Hz, 1H), 1.05-1.15 (m, 1H), 0.75-0.88 (m, 2H), 0.51-0.69 (m, 2H); 2 Hs not observed (NH and OH).
277	MS m/z 418.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.13 (s, 1H), 7.86 (d, J=8.2 Hz, 1H), 7.74 (s, 1H), 7.08 (d, J=1.7 Hz, 2H), 4.92-5.03 (m, 1H), 4.83-4.86 (m, 1H), 4.19 (t, J=6.1 Hz, 2H), 3.36-3.56 (m, 3H), 3.12-3.26 (m, 1H), 3.04 (t, J=6.3 Hz, 2H), 2.52-2.69 (m, 1H), 2.06-2.19 (m, 2H), 1.93-2.00 (m, 2H), 0.97-1.16 (m, 1H), 0.76-0.92 (m, 2H), 0.38-0.65 (m, 2H); 2Hs not observed (NH and OH).
278	MS m/z 406.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.22 (s, 1H), 9.04 (s, 1H), 7.90 (s, 1H), 7.84 (d, J = 8.0 Hz, 1H), 7.08-7.15 (m, 2H), 4.63 (d, J = 12.7 Hz, 1H), 4.53 (d, J = 12.6 Hz, 1H), 4.11 (t, J = 7.3 Hz, 2H), 2.91-3.19 (m, 4H), 2.60-2.81 (m, 4H), 2.37 (s, 1H), 1.63-1.68 (m, 1H), 0.98 (d, J = 6.9 Hz, 6H); 1H not observed (OH or NH).

Cpd	Data
279	MS m/z 406.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.23 (s, 1H), 9.03 (s, 1H), 7.87 (d, J = 8.0 Hz, 1H), 7.30 (s, 1H), 7.16 (d, J = 8.6 Hz, 2H), 4.63 (d, J = 12.7 Hz, 1H), 4.54 (d, J = 12.7 Hz, 1H), 4.22 (t, J = 7.0 Hz, 2H), 2.89-3.13(m, 2H), 2.67-2.89 (m, 4H), 2.55-2.65 (m, 2H), 2.35-2.43 (m, 1H), 1.66 (h, J = 6.8 Hz, 1H), 0.98 (d, J = 6.9 Hz, 6H); 1H not observed (OH or NH).
280	MS m/z 418.1 $[M+H]^+$; ^1H NMR (400 MHz, $CDCl_3$) δ: 12.31 (s, 1H), 8.81 (s, 1H), 7.85 (s, 1H), 7.61 (d, J = 8.3 Hz, 1H), 7.14 (d, J = 1.6 Hz, 1H), 7.07 (dd, J = 8.2, 1.7 Hz, 1H), 4.74-4.65 (m, 2H), 4.19 (t, J = 7.3 Hz, 2H), 3.22-3.17 (m, 1H), 3.14 (t, J = 12 Hz, 2H), 3.08 (td, J = 11.7, 3.1 Hz, 1H), 3.00 (t, J = 12 Hz, 1H), 2.85 (td, J = 11.7, 3.0 Hz, 1H), 2.76 – 2.67 (m, 2H), 1.95 (dd, J = 10.9, 2.8 Hz, 1H), 1.25 (s, 1H), 1.12 (s, 3H), 0.52-0.45 (m, 1H), 0.42-0.32 (m, 3H).
281	MS m/z 418.1 $[M+H]^+$; ^1H NMR (400 MHz, $CDCl_3$) δ: 12.31 (s, 1H), 8.81 (s, 1H), 7.85 (s, 1H), 7.61 (d, J = 8.3 Hz, 1H), 7.14 (d, J = 1.6 Hz, 1H), 7.07 (dd, J = 8.2, 1.7 Hz, 1H), 4.74-4.65 (m, 2H), 4.19 (t, J = 7.3 Hz, 2H), 3.22-3.17 (m, 1H), 3.14 (t, J = 12 Hz, 2H), 3.08 (td, J = 11.7, 3.1 Hz, 1H), 3.00 (t, J = 12 Hz, 1H), 2.85 (td, J = 11.7, 3.0 Hz, 1H), 2.76 – 2.67 (m, 2H), 1.95 (dd, J = 10.9, 2.8 Hz, 1H), 1.25 (s, 1H), 1.12 (s, 3H), 0.52-0.45 (m, 1H), 0.42-0.32 (m, 3H).
282	MS m/z 422.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.01 (s, 1H), 9.09 (s, 1H), 7.83 (d, J = 8.2 Hz, 1H), 7.77 (s, 1H), 7.24 (d, J = 1.8 Hz, 1H), 7.20 (dd, J = 8.3, 1.6 Hz, 1H), 4.80 (d, J = 13.6 Hz, 1H), 4.68-4.75 (m, 1H), 4.46 (t, J = 5.2 Hz, 2H), 4.15 (t, J = 6.1 Hz, 2H), 3.38 (s, 2H), 2.93-3.20 (m, 3H), 2.23-2.28 (m, 2H), 1.80-2.01 (m, 1H), 1.05 (d, J = 6.8 Hz, 6H); 1H not observed (OH or NH).
283	MS *m/z* 402.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.71 (s, 1H), 8.36 (s, 1H), 7.91 (d, J=1.6 Hz, 1H), 6.96 (s, 1 H), 6.90 (d, J=12.4 Hz, 1H), 4.89 (d, J=13.3 Hz, 1H), 4.81 - 4.80 (m, 1H), 3.26-3.31 (m, 2H), 2.99-3.06 (m, 2H), 2.80-2.84 (m, 1H), 1.80-1.83 (m, 1H), 1.02 (d, J=6.8 Hz, 6H); 3Hs not observed (2 NHs and OH), 1H from formic acid salt.
284	MS m/z 364.3 $[M+H]^+$

Example 2

Preparation of Compound 100



Step 1: To a stirred solution of 6-(4-chloro-2-(methoxymethoxy)phenyl)-3-(methylthio)-1,2,4-triazine (2.0 g, 6.7 mmol) in CH_2Cl_2 (35 mL) was added mCPBA (4.6 g, 20 mmol) portionwise and the reaction was allowed to stir at rt for 5h. It was then quenched with saturated aqueous $NaHCO_3$. Organic layers were dried over $MgSO_4$ and concentrated. The residue was purified by silica gel column chromatography eluting with a gradient EtOAc/hexanes (0-100% EtOAc) to afford 6-(4-chloro-2-(methoxymethoxy)phenyl)-3-(methylsulfonyl)-1,2,4-triazine (1.75 g, 79% yield) as a tan solid. ^1H NMR (500 MHz, $CDCl_3$) δ: 9.40 (s, 1H), 8.06 (d, J=8.5 Hz, 1H), 7.40 (d, J=1.2 Hz, 1H), 7.27 (dd, J=8.5, 1.2 Hz, 1H), 5.32 (s, 2H), 3.56 (s, 3H), 3.52 (s, 3H).

Step 2: To a stirred solution of 6-(4-chloro-2-(methoxymethoxy)phenyl)-3-(methylsulfonyl)-1,2,4-triazine (900 mg, 2.73 mmol) in ACN (10 mL) were added cis-2,6-dimethylpiperazine (400 mg, 3.5 mmol) and DIPEA (1.0 mL, 5.73 mmol). The reaction mixture was heated at 50 ºC for 1h until UPLC showed complete conversion to the desired product. Solvent was removed under reduced pressure, the residue was purified by silica gel column chromatography eluting with a gradient CH_2Cl_2/MeOH (0-20% MeOH) to afford 6-(4-chloro-2-(methoxymethoxy)phenyl)-3-(cis-3,5-dimethylpiperazin-1-yl)-1,2,4-triazine (993 mg, 70.5% yield) as a yellowish solid. MS *m/z* 364.2, 366.2 $[M+H]^+$.

Step 3: A suspension of 6-(4-chloro-2-(methoxymethoxy)phenyl)-3-(cis-3,5-dimethylpiperazin-1-yl)-1,2,4-triazine (650 mg, 1.79mmol), 4,4,5,5-tetramethyl-2-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)-1,3,2-dioxaborolane (780 mg, 2.68 mmol), KOAc (525 mg, 5.35 mmol), X Phos Pd G4 (80 mg, 0.09 mmol) in dry dioxane (12 mL) was sparged with argon for 10 minutes, then heated to 90 °C under argon atmosphere for 2 h, after which complete conversion to 3-(cis-3,5-dimethylpiperazin-1-yl)-6-(2-(methoxymethoxy)-4-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)phenyl)-1,2,4-triazine was observed. The reaction mixture was then cooled to room temperature, concentrated and purified by silica gel column chromatography eluting with a gradient CH_2Cl_2/MeOH (0-15% MeOH) to afford 3-(cis-3,5-dimethylpiperazin-1-yl)-6-(2-(methoxymethoxy)-4-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)phenyl)-1,2,4-triazine (800 mg, 98% yield) as a brown crystalline solid. MS *m/z* 456.5 $[M+H]^+$.

Step 4: A mixture of 3-(cis-3,5-dimethylpiperazin-1-yl)-6-(2-(methoxymethoxy)-4-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)phenyl)-1,2,4-triazine (60 mg, 0.13 mmol), 6-bromo-2-methyl-[1,2,4]triazolo[1,5-b]pyridazine (42.0 mg, 0.20 mmol), [1,1′-Bis(diphenylphosphino)ferrocene]dichloropalladium(II) (9.5 mg, 0.013 mmol), and aqueous potassium carbonate (0.2 mL, 2 M) in dioxane (1 mL) was degassed with argon for 10 minutes, then heated to 90 ºC for 1h. The mixture was cooled to room temperature and purified directly by silica gel column chromatography, eluting with a gradient 0-20% methanol in dichloromethane, to afford 6-[4-[3-[cis-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl]-3-(methoxymethoxy)phenyl]-2-methyl-[1,2,4]triazolo[1,5-b]pyridazine (50 mg, 83% yield). MS m/z 462.4 $[M+H]^+$.

Step 5: 6-[4-[3-[cis-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl]-3-(methoxymethoxy)phenyl]-2-methyl-[1,2,4]triazolo[1,5-b]pyridazine (50 mg, 0.11 mmol) was dissolved in methanol (2 mL) and HCl (4 mol/L) in 1,4-dioxane (0.2 mL, 0.8 mmol) was added. The reaction was stirred at room temperature for 2h, concentrated and purified by silica gel column chromatography, eluting with a gradient CH_2Cl_2/MeOH (10-30% MeOH) to afford 2-[3-[cis-3,5-dimethylpiperazin-1-yl]-1,2,4-triazin-6-yl]-5-(2-methyl-[1,2,4]triazolo[1,5-b]pyridazin-6-yl)phenol;dihydrochloride as a yellow solid (30 mg, 47% yield). MS m/z 418.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.90 (s, 1H), 9.57 (s, 1H), 9.11 (s, 1H), 8.41 (d, J = 9.5 Hz, 1H), 8.18 (d, J = 9.0 Hz, 1H), 8.04 (d, J = 8.5 Hz, 1H), 7.80 (s, 1H), 7.71 (d, J = 8.0 Hz, 1H), 4.85 (d, J = 14.0 Hz, 2H), 3.35-3.38 (m, 2H), 3.19-3.23 (m, 2H), 3.17 (s, 3H), 1.38 (d, J = 6.0 Hz, 6H).

Using the procedure described for Example 2, above, additional compounds described herein may be prepared by substituting the appropriate starting material, suitable reagents and reaction conditions, obtaining compounds such as those selected from:

Cpd	Data
54	MS m/z 444.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.19 (s, 1H), 8.31 (br s, 1H), 8.07 - 8.09 (m, 1H), 7.95 - 7.97 (m, 1H), 3.53 - 3.57 (m, 4H), 3.44 - 3.53 (m, 2H), 3.19 - 3.26 (m, 1H), 2.65 - 2.68 (m, 6H), 1.06 - 1.11 (m, 1H), 0.81 - 0.85 (m, 2H), 0.57 - 0.60 (m, 2H); 3 Hs not observed (2 NHs and OH).
65	MS m/z 436.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.45 (s, 1H), 7.86 (s, 1H), 7.73 (d, J = 8.2 Hz, 1H), 7.39 (d, J = 12.7 Hz, 1H), 7.32 (d, J = 8.4 Hz, 1H), 7.27 (s, 1H), 6.05 (s, 1H), 4.41 - 4.20 (m, 5H), 4.01 - 3.90 (m, 1H), 3.76-3.59 (m, 4H), 1.40 (t, J = 6.8 Hz, 2H); 3Hs not observed (NH and 2 OHs).
66	MS m/z 434.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.19 (s, 1H), 8.92 (s, 1H), 8.01-8.11 (m, 3H), 7.34-7.43 (m, 2H), 5.08 (br d, J=14.0 Hz, 2H), 3.45-3.55 (m, 2H), 3.13 (dd, J=14.2, 11.7 Hz, 2H), 2.61 (s, 3H), 1.48 (d, J=6.4 Hz, 6H); 2Hs not observed (NH and OH).
67	MS m/z 420.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.49 - 8.44 (m, 1H), 7.88 (s, 1H), 7.74 (d, J = 8.2 Hz, 1H), 7.41 (d, J = 12.5 Hz, 1H), 7.33 (d, J = 8.4 Hz, 1H), 7.29 (s, 1H), 6.05 (s, 1H), 4.30 (s, 3H), 4.34-4.3 (m, 2H), 4.25 - 4.19 (m, 1H), 3.85 - 3.39 (m, 4H), 1.47-1.51 (m, 3H); 2Hs not observed (NH and OH).
68	MS m/z 434.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.08 (s, 1H), 8.58 (s, 1H), 7.93 (d, J=7.9 Hz, 1H), 7.78 (s, 1H), 7.42 (d, J=11.9 Hz, 1H), 7.29 (d, J=7.6 Hz, 1H), 7.26 (s, 1H), 4.76 (q, J=13.5 Hz, 2H), 3.12-3.24 (m, 2H), 2.88 (td, J=12.5, 3.9 Hz, 1H), 2.82 (dd, J=13.0, 10.5 Hz, 1H), 2.63-2.73 (m, 1H), 2.43 (s, 3H), 1.57 (quin, J=7.5 Hz, 2H), 1.07 (t, J=7.5 Hz, 3H); 2Hs not observed (NH and OH).
69	MS m/z 431.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.06 (s, 1H), 7.92 (d, J=8.5 Hz, 1H), 7.90 (s, 1H), 7.57-7.63 (m, 2H), 7.53 (s, 1H), 4.77 (d, J=12.6 Hz, 2H), 2.86-3.05 (m, 2H), 2.71 (t, J=12.6 Hz, 2H), 2.67 (s, 3H), 2.49 (s, 3H), 1.24 (d, J=6.4 Hz, 6H); 2Hs not observed (NH and OH).
70	MS m/z 462.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.11 (s, 1H), 8.01 (br d, J=8.9 Hz, 1H), 7.72 (s, 1H), 7.71 (s, 1H), 7.49 (s, 1H), 4.84-4.86 (m, 1H), 4.74 (d, J=13.4 Hz, 1H), 4.28 (s, 3H), 3.12-3.23 (m, 2H), 2.79-2.95 (m, 2H), 2.62 (s, 3H), 2.50-2.56 (m, 1H), 1.77 (td, J=13.7, 7.0 Hz, 1H), 1.09 (d, J=6.7 Hz, 6H); 2 Hs not observed (NH and OH).
71	MS m/z 448.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.43 (s, 1H), 7.86 (s, 1H), 7.75 (d, J = 8.4 Hz, 1H), 7.29 (s, 3H), 6.06 (s, 1H), 4.40 - 4.18 (m, 5H), 3.84 - 3.42 (m, 5H), 2.19 - 2.04 (m, 1H), 1.28 - 1.14 (m, 6H); 2Hs not observed (NH and OH).

Cpd	Data
72	MS m/z 416.6 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.07 (s, 1H), 8.20 (s, 1H), 7.95 (s, 1H), 7.87 (d, J = 7.9 Hz, 1H), 7.69 (d, J = 9.0 Hz, 1H), 7.66 - 7.62 (m, 1H), 7.31 - 7.27 (m, 2H), 4.87 (br d, J = 13.6 Hz, 2H), 4.24 (s, 3H), 3.16 (br s, 2H), 2.83 (t, J = 12.5 Hz, 2H), 1.32 (d, J = 6.4 Hz, 6H); 2Hs not observed (NH and OH).
73	MS m/z 420.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.21 (s, 1H), 8.91 (s, 1H), 8.07 (s, 1H), 7.97-8.03 (m, 2H), 7.35-7.42 (m, 2H), 4.95 – 5.11 (m, 2H), 3.31-3.76 (m, 6H), 3.02 (s, 3H), 2.66 (s, 3H); 1H not observed (OH).
74	MS m/z 434.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.45 (s, 1H), 7.88 (s, 1H), 7.75 (d, J = 8.2 Hz, 1H), 7.41 (d, J = 12.7 Hz, 1H), 7.34 (d, J = 8.2 Hz, 1H), 7.29 (s, 1H), 6.06 (s, 1H), 4.34-4.20 (m, 5H), 3.83 - 3.41 (m, 5H), 1.94 - 1.76 (m, 2H), 1.24 - 1.10 (m, 3H); 2Hs not observed (NH and OH).
75	MS m/z 417.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.07 (s, 1H), 7.93 (s, 3H), 7.69 (d, J = 9.5 Hz, 1H), 7.65 - 7.60 (m, 2H), 4.78 (d, J = 13.1 Hz, 2H), 2.96 (br s, 2H), 2.67 (t, J = 12.1 Hz, 2H), 2.50 (s, 3H), 1.22 (d, J = 6.4 Hz, 6H); 2Hs not observed (NH and OH).
76	MS m/z 434.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.42 (s, 1H), 7.86 (s, 1H), 7.74 (d, J = 7.3 Hz, 1H), 7.42 - 7.23 (m, 3H), 6.05 (d, J = 4.3 Hz, 1H), 4.31 (s, 3H), 4.17 - 3.91 (m, 4H), 3.83 - 3.70 (m, 2H), 1.52 - 1.44 (m, 3H), 1.42 - 1.37 (m, 3H); 2Hs not observed (NH and OH).
77	MS m/z 434.4 $[M+H]^+$
78	MS m/z 446.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.44 (s, 1H), 9.33-9.41 (m, 1H), 9.17-9.23 (m, 1H), 9.13 (s, 1H), 7.98 (d, J=8.2 Hz, 1H), 7.84 (s, 1H), 7.74 (d, J=8.2 Hz, 1H), 4.85 (d, J=13.5 Hz, 1H), 4.78 (d, J=13.5 Hz, 1H), 3.45-3.55 (m, 1H), 3.41 (br d, J=12.5 Hz, 1H), 3.27 (dd, J=14.3, 10.7 Hz, 1H), 3.10-3.21 (m, 2H), 2.86 (s, 3H), 2.58 (s, 3H), 2.04 (sxt, J=6.4 Hz, 1H), 1.08 (t, J=6.1 Hz, 6H).
79	MS m/z 434.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.43 - 8.40 (m, 1H), 7.86 (s, 1H), 7.74 (d, J = 8.1 Hz, 1H), 7.41 - 7.30 (m, 2H), 7.29 (s, 1H), 6.06 (s, 1H), 4.40 - 4.32 (m, 1H), 4.28 (s, 3H), 4.26 - 4.18 (m, 1H), 3.73 - 3.59 (m, 2H), 3.52 - 3.39 (m, 2H), 1.54 - 1.45 (m, 6H); 2 Hs not observed (NH and OH).
80	MS m/z 441.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 8.53 - 8.49 (m, 1H), 8.31 (s, 1H), 8.09 (s, 1H), 7.73 (d, J = 8.4 Hz, 1H), 7.31 (d, J = 8.2 Hz, 1H), 7.27 - 7.24 (m, 1H), 6.05 (s, 1H), 4.41 - 4.34 (m, 1H), 4.31 (s, 3H), 4.29 - 4.21 (m, 1H), 3.73-3.63 (m, 2H), 3.54 - 3.38 (m, 2H), 1.48-1.50 (m, 6H); 2 Hs not observed (NH and OH).
81	MS m/z 448.2 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.05 (s, 1H), 8.31 (d, J = 2.4 Hz, 1H), 7.86 (d, J = 8.1 Hz, 1H), 7.77 (s, 1H), 7.35 - 7.23 (m, 3H), 4.76 - 4.66 (m, 1H), 4.26 (s, 3H), 3.23 - 3.09 (m, 2H), 2.94 - 2.81 (m, 2H), 2.57 - 2.49 (m, 1H), 1.81 - 1.71 (m, 1H), 1.08 (d, J = 6.9 Hz, 6H); 3Hs not observed (NH, OH and 1 CH overlapped with solvent peak).

Cpd	Data
82	MS m/z 403.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 7.81 (s, 1H), 7.65 (d, J = 2.1 Hz, 1H), 7.15 (d, J = 2.3 Hz, 1H), 7.05 (s, 1H), 6.66 (d, J = 8.7 Hz, 1H), 6.05 - 6.02 (m, 2H), 3.04 (s, 3H), 2.70-2.80 (m, 4H), 1.35-1.46 (m, 4H), 1.18 (s, 3H); 1H not observed (OH).
83	MS m/z 432.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.49 (d, J = 10.7 Hz, 1H), 9.05-9.08 (m, 2H), 8.02 (s, 1H), 7.98 (d, J = 8.2 Hz, 1H), 7.70 (d, J = 1.8 Hz, 1H), 7.64 (dd, J = 8.2, 1.8 Hz, 1H), 4.81 (d, J = 13.9 Hz, 2H), 3.35 (s, 2H), 2.96-3.13 (m, 2H), 2.61 (s, 3H), 2.50 (s, 3H), 1.30 (d, J = 6.4 Hz, 6H).
85	MS m/z 431.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.32 (br s, 1H), 9.22 (s, 1H), 9.15 (s, 1H), 9.07 (br s, 1H), 7.95-8.05 (m, 2H), 7.84 (d, J=9.2 Hz, 1H), 7.42 (br dd, J=7.9, 1.3 Hz, 1H), 7.38 (s, 1H), 4.86 (br d, J=13.7 Hz, 1H), 4.79 (br d, J=14.0 Hz, 1H), 3.35-3.51 (m, 2H), 3.22-3.30 (m, 2H), 3.17 (s, 3H), 3.09-3.16 (m, 1H), 1.97-2.13 (m, 1H), 1.07 (dd, J=7.3, 4.5 Hz, 6H).
86	MS m/z 417.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.63 (d, J = 10.1 Hz, 1H), 9.40 (d, J = 1.5 Hz, 1H), 9.29 (s, 1H), 9.24 (d, J = 9.5 Hz, 1H), 9.13 (s, 1H), 8.12 (s, 1H), 8.02 (d, J = 8.2 Hz, 1H), 7.78 (d, J = 1.7 Hz, 1H), 7.65 (dd, J = 8.2, 1.8 Hz, 1H), 4.86 (d, J = 13.7 Hz, 2H), 3.35-3.43 (m, 1H), 3.05-3.19 (m, 3H), 2.53 (s, 3H), 1.37 (d, J = 6.4 Hz, 6H).
87	MS m/z 441.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.16 (s, 1H), 9.09 - 9.06 (m, 1H), 8.27 - 8.22 (m, 1H), 8.01 - 7.95 (m, 1H), 7.90 - 7.85 (m, 1H), 7.36 - 7.30 (m, 2H), 4.68 - 4.55 (m, 2H), 3.60 - 3.57 (m, 2H), 2.86 - 2.73 (m, 2H), 2.41 (s, 3H), 1.11 - 1.04 (m, 6H); 2Hs not observed (NH and OH).
88	MS m/z 430.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.71 (br s, 1H), 9.42 (br s, 1H), 9.12 (s, 1H), 8.43 (s, 1H), 8.00 (s, 1H), 7.94 (d, J=8.2 Hz, 1H), 7.70 (d, J=8.5 Hz, 1H), 7.57 (dd, J=8.9, 1.8 Hz, 1H), 7.29-7.37 (m, 2H), 4.85 (br d, J=13.4 Hz, 1H), 4.77 (br d, J=14.0 Hz, 1H), 3.53 (td, J=12.5, 1.2 Hz, 1H), 3.40 (br d, J=12.5 Hz, 1H), 3.29 (dd, J=13.7, 11.3 Hz, 1H), 3.07-3.19 (m, 2H), 2.45 (s, 3H), 2.08 (dq, J=13.6, 6.8 Hz, 1H), 1.08 (t, J=6.8 Hz, 6H).
89	MS m/z 448.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.06 (s, 1H), 8.57 (s, 1H), 7.92 (br d, J=7.9 Hz, 1H), 7.78 (d, J=2.0 Hz, 1H), 7.41 (d, J=11.6 Hz, 1H), 7.28 (br d, J=7.9 Hz, 1H), 7.25 (s, 1H), 4.48 (br d, J=13.4 Hz, 1H), 4.43 (br d, J=12.8 Hz, 1H), 3.44-3.57 (m, 1H), 3.22 (dd, J=14.0, 8.5 Hz, 1H), 3.08 (dt, J=12.2, 3.0 Hz, 1H), 2.98 (dquin, J=13.7, 6.7 Hz, 1H), 2.62-2.72 (m, 1H), 2.54-2.61 (m, 1H), 2.47-2.53 (m, 1H), 2.46 (s, 3H), 1.21 (d, J=6.1 Hz, 3H), 1.15 (t, J=7.2 Hz, 3H); 1 H not observed (OH).
90	MS m/z 448.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.61 (d, J = 10.4 Hz, 1H), 9.22 (d, J = 10.8 Hz, 1H), 9.13 (s, 1H), 8.04 (d, J = 8.1 Hz, 1H), 7.73-7.82 (m, 2H), 7.55 (s, 1H), 4.87 (d, J = 13.9 Hz, 2H), 4.22 (s, 3H), 3.37-3.42 (m, 2H), 3.11-3.17 (m, 2H), 2.52 (s, 3H), 1.36 (d, J = 6.5 Hz, 6H).
91	MS m/z 432.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.29 (s, 1H), 9.25 (s, 1H), 9.19 (s, 1H), 8.00 (d, J=7.9 Hz, 1H), 7.76 (s, 1H), 7.71 (br d, J=8.2 Hz, 1H), 5.10 (d, J=13.7 Hz, 1H), 4.99 (d, J=14.0 Hz, 1H), 3.44-3.61 (m, 2H), 3.25-3.32 (m, 2H), 3.14-3.22 (m, 1H), 2.67 (s, 3H), 2.07 (spt, J=6.8 Hz, 1H), 1.20 (d, J=6.8 Hz, 6H); 2Hs not observed (NH and OH).

Cpd	Data
92	MS m/z 446.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.16 (s, 1H), 8.02 (br d, J=7.6 Hz, 1H), 7.99 (s, 1H), 7.67-7.73 (m, 2H), 5.03 (d, J=13.7 Hz, 1H), 4.92 (d, J=13.7 Hz, 1H), 3.37-3.49 (m, 2H), 3.11-3.27 (m, 2H), 2.95-3.03 (m, 1H), 2.74 (s, 3H), 2.64 (s, 3H), 1.88-2.02 (m, 1H), 1.16 (d, J=6.4 Hz, 6H); 2 Hs not observed (NH and OH).
93	MS m/z 462.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.17 (s, 1H), 8.83 (s, 1H), 7.96 (br d, J=8.5 Hz, 1H), 7.74 (s, 1H), 7.68 (d, J=8.2 Hz, 1H), 5.09 (d, J=13.7 Hz, 1H), 4.98 (d, J=14.0 Hz, 1H), 4.27-4.36 (m, 2H), 3.70 (s, 3H), 3.43-3.62 (m, 1H), 3.24-3.32 (m, 1H), 3.05-3.20 (m, 1H), 2.62 (s, 3H), 2.02-2.16 (m, 1H), 1.19 (d, J=6.8 Hz, 6H); 2 Hs not observed (NH and OH).
94	MS m/z 448.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.10 (br s, 1H), 9.16 (s, 1H), 9.14 (s, 1H), 8.20 (br s, 1H), 8.17 (s, 1H), 8.04 (d, J=8.2 Hz, 1H), 7.36-7.44 (m, J=2.1 Hz, 2H), 4.83 (d, J=10.7 Hz, 2H), 3.43 (s, 3H), 3.32-3.40 (m, 2H), 2.82 (d, J=4.6 Hz, 2H), 2.53 (s, 3H), 1.47 (d, J=5.5 Hz, 6H).
95	MS m/z 446.4 $[M+H]^+$
96	MS m/z 445.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.20 (s, 1H), 9.19 (s, 1H), 8.18 (s, 1H), 8.02 (d, J=8.2 Hz, 1H), 7.76 (br s, 1H), 7.71 (br d, J=8.2 Hz, 1H), 5.10 (d, J=14.6 Hz, 1H), 4.99 (d, J=14.6 Hz, 1H), 3.50-3.63 (m, 2H), 3.27-3.31 (m, 2H), 3.18-3.27 (m, 1H), 2.99 (s, 3H), 2.66 (s, 3H), 2.10 (qd, J=14.3, 6.7 Hz, 1H), 1.20 (dd, J=6.7, 2.7 Hz, 6H); 2Hs not observed (NH and OH).
97	MS m/z 417.4 $[M+H]^+$
98	MS m/z 417.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.07 (s, 1H), 9.27 (br s, 1H), 9.11 (s, 1H), 8.07 (s, 1H), 8.01 (d, J=7.9 Hz, 1H), 7.71 (s, 1H), 7.64 (br d, J=8.2 Hz, 1H), 7.62 (s, 1H), 4.69-4.80 (m, 2H), 3.40-3.52 (m, 2H), 3.22-3.30 (m, 2H), 3.16 (td, J=12.8, 3.9 Hz, 1H), 2.62 (s, 3H), 2.42 (s, 3H), 1.33 (d, J=6.5 Hz, 3H).
99	MS m/z 446.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.08 (s, 1H), 8.55 (d, J = 2.4 Hz, 1H), 7.94 (d, J = 8.1 Hz, 1H), 7.87 (s, 1H), 7.42 - 7.36 (m, 1H), 7.33 (d, J = 8.2 Hz, 1H), 7.31 - 7.29 (m, 1H), 4.83 (d, J = 2.6 Hz, 2H), 4.23 (s, 3H), 3.22 - 2.99 (m, 3H), 2.80 (br s, 1H), 2.26 - 2.04 (m, 2H), 2.03 - 1.82 (m, 2H), 1.82 - 1.64 (m, 2H), 1.53 - 1.35 (m, 1H); 1H not observed (OH).
101	MS m/z 432.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.68 (d, J = 10.4 Hz, 1H), 9.43 (s, 1H), 9.30 (d, J = 10.8 Hz, 1H), 9.12 (s, 1H), 7.98 (d, J = 8.2 Hz, 1H), 7.84 (d, J = 1.7 Hz, 1H), 7.73 (dd, J = 8.2, 1.7 Hz, 1H), 4.82-4.89 (m, 2H), 3.38-3.42 (m, 2H), 3.12-3.19 (m, 2H), 2.86 (s, 3H), 2.58 (s, 3H), 1.37 (d, J = 6.5 Hz, 6H).
102	MS m/z 417.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.23 (br s, 1H), 9.36 (br s, 1H), 9.12 (s, 1H), 8.63 (s, 1H), 8.46 (br d, J=9.5 Hz, 1H), 8.19 (s, 1H), 8.15 (br d, J=9.5 Hz, 1H), 8.07 (d, J=8.2 Hz, 1H), 7.80 (d, J=1.2 Hz, 1H), 7.73 (dd, J=8.5, 1.8 Hz, 1H), 4.86 (br d, J=13.7 Hz, 1H), 4.78 (d, J=14.3 Hz, 1H), 3.49-3.59 (m, 1H), 3.41 (d, J=12.5 Hz, 1H), 3.30 (dd, J=14.0, 11.3 Hz, 1H), 3.07-3.17 (m, 2H), 2.07 (dq, J=13.4, 6.4 Hz, 1H), 1.08 (t, J=6.4 Hz, 6H).
103	MS m/z 431.5 $[M+H]^+$

Cpd	Data
104	MS m/z 417.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.15-9.25 (m, 1H), 8.08 (br s, 1H), 8.02 (d, J=7.0 Hz, 1H), 7.86 (br s, 1H), 7.63-7.76 (m, 2H), 3.40-3.82 (m, 4H), 3.37 (s, 3H), 3.01 (s, 3H), 2.70-2.76 (m, 4H), 2.56 (s, 3H), 1 H not obsrved (OH)
105	MS m/z 446.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.08 (s, 1H), 8.55 (d, J = 2.4 Hz, 1H), 7.94 (d, J = 8.1 Hz, 1H), 7.87 (s, 1H), 7.42 - 7.36 (m, 1H), 7.33 (d, J = 8.2 Hz, 1H), 7.31 - 7.29 (m, 1H), 4.83 (d, J = 2.6 Hz, 2H), 4.23 (s, 3H), 3.22 - 2.99 (m, 3H), 2.80 (br s, 1H), 2.26 - 2.04 (m, 2H), 2.03 - 1.82 (m, 2H), 1.82 - 1.64 (m, 2H), 1.53 - 1.35 (m, 1H); 1H not observed (OH).
106	MS m/z 418.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.59-9.63 (m, 2H), 9.38 (d, J = 1.4 Hz, 1H), 9.18-9.24 (m, 1H), 9.14 (s, 1H), 8.00 (d, J = 8.2 Hz, 1H), 7.85 (d, J = 1.7 Hz, 1H), 7.76 (dd, J = 8.2, 1.8 Hz, 1H), 4.86 (d, J = 13.7 Hz, 2H), 3.39-3.41 (m, 2H), 3.13 (dd, J = 14.1, 11.5 Hz, 2H), 2.60 (s, 3H), 1.36 (d, J = 6.4 Hz, 6H).
107	MS m/z 434.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 8.72-8.78 (m, 1H), 8.20 (s, 1H), 7.94-8.02 (m, 1H), 7.86 (d, J=7.3 Hz, 1H), 7.78 (s, 1H), 7.55 (t, J=7.6 Hz, 1H), 7.39 (d, J=12.5 Hz, 1H), 4.50 (dd, J=27.2, 13.4 Hz, 2H), 3.21-3.27 (m, 1H), 2.80-2.91 (m, 2H), 2.51 (s, 3H), 2.26 (s, 3H), 2.21 (td, J=11.6, 3.4 Hz, 1H), 2.09-2.16 (m, 1H), 1.10 (d, J=6.1 Hz, 3H); 1H not observed (OH).
108	MS m/z 456.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.16 (br s, 1H), 8.19 (s, 1H), 8.11 (s, 1H), 8.01 (br d, J=7.0 Hz, 1H), 7.57-7.70 (m, 2H), 5.02 (d, J=8.9 Hz, 2H), 3.43-3.53 (m, 4H), 2.99 (s, 3H), 2.56 (s, 3H), 1.58 (d, J=6.0 Hz, 6H); 1H not observed (OH).
109	MS m/z 458.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.60 (d, J = 10.6 Hz, 1H), 9.34 (s, 1H), 9.21 (d, J = 10.5 Hz, 1H), 9.12 (s, 1H), 7.96 (d, J = 8.2 Hz, 1H), 7.80 (d, J = 1.8 Hz, 1H), 7.70 (dd, J = 8.2, 1.7 Hz, 1H), 4.82-4.89 (m, 2H), 3.40 (s, 2H), 3.13 (dd, J = 14.1, 11.5 Hz, 2H), 2.81 (td, J = 8.2, 4.2 Hz, 1H), 2.58 (s, 3H), 1.41 – 1.32 (m, 8H), 1.30 – 1.25 (m, 2H).
110	MS m/z 403.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.09 (s, 1H), 7.97 - 7.91 (m, 3H), 7.68 (d, J = 9.5 Hz, 1H), 7.65 - 7.61 (m, 2H), 4.03 (br s, 4H), 2.69 (br s, 4H), 2.52 - 2.49 (m, 3H), 2.46 (s, 3H); 1H not observed (OH).
111	MS m/z 445.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.23 (s, 1H), 8.33 (s, 1H), 8.26 (s, 1H), 8.11 (d, J=7.9 Hz, 1H), 7.68-7.84 (m, 2H), 5.08 (d, J=13.4 Hz, 2H), 3.44-3.57 (m, 2H), 3.03 (s, 3H), 2.76-2.87 (m, 2H), 2.68 (s, 3H), 2.05 (s, 3H), 1.57 (d, J=6.0 Hz, 6H); 1 H not observed (OH).
112	MS m/z 418.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.68 (d, J = 2.4 Hz, 1H), 9.63 (d, J = 10.3 Hz, 1H), 9.21-9.29 (m, 1H), 9.18 (d, J = 2.4 Hz, 1H), 9.16 (s, 1H), 8.03 (d, J = 8.1 Hz, 1H), 7.47 (dd, J = 8.1, 1.9 Hz, 1H), 7.44 (d, J = 1.9 Hz, 1H), 4.86 (d, J = 13.8 Hz, 2H), 3.41 (s, 2H), 3.09-3.19 (m, 2H), 2.55 (s, 3H), 1.36 (d, J = 6.4 Hz, 6H).

Cpd	Data
113	MS m/z 417.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.28 (s, 1H), 9.23 (s, 1H), 9.21 (s, 1H), 8.26 (s, 1H), 8.06 (s, 1H), 8.03 (d, J=8.2 Hz, 1H), 7.76 (s, 1H), 7.71 (d, J=7.6 Hz, 1H), 5.11 (d, J=13.7 Hz, 1H), 5.01 (d, J=14.0 Hz, 1H), 3.58 (d, J=12.5 Hz, 1H), 3.45-3.53 (m, 1H), 3.27-3.32 (m, 2H), 3.17-3.24 (m, 1H), 2.06 (dq, J=13.4, 6.8 Hz, 1H), 1.20 (d, J=6.8 Hz, 6H); 2Hs not observed (NH and OH).
114	MS m/z 416.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.19 (s, 1H), 9.08 (s, 1H), 8.19 (s, 1H), 8.10 (dd, J=9.5, 0.9 Hz, 1H), 8.03 (d, J=8.2 Hz, 1H), 7.93 (d, J=1.2 Hz, 1H), 7.91 (d, J=9.5 Hz, 1H), 7.39 (dd, J=8.2, 1.2 Hz, 1H), 7.37 (s, 1H), 5.09 (br d, J=13.4 Hz, 1H), 4.98 (d, J=14.9 Hz, 1H), 3.50-3.62 (m, 2H), 3.27-3.32 (m, 2H), 3.17-3.26 (m, 1H), 2.10 (qd, J=13.7, 6.9 Hz, 1H), 1.20 (dd, J=6.9, 3.8 Hz, 6H); 2 Hs not observed (NH and OH).
115	MS m/z 448.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.23 (s, 1H), 9.05 (s, 1H), 8.81 (s, 1H), 7.92 (d, J=7.9 Hz, 1H), 7.82 (s, 2H), 7.29 (s, 1H), 7.26 (s, 1H), 4.82 (d, J=12.5 Hz, 1H), 4.69 (d, J=14.3 Hz, 1H), 3.09-3.20 (m, 2H), 2.80-2.91 (m, 2H), 2.44-2.54 (m, 1H), 1.75 (dq, J=13.7, 7.0 Hz, 1H), 1.07 (dd, J=6.7, 3.4 Hz, 6H); 2Hs not observed (NH and OH).
116	MS m/z 448.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.58 (s, 1H), 9.22 (s, 1H), 9.03 (s, 1H), 8.18 (s, 1H), 8.03 (d, J = 8.1 Hz, 1H), 7.86 (d, J = 1.8 Hz, 1H), 7.76 (dd, J = 8.2, 1.8 Hz, 1H), 4.81 (d, J = 13.9 Hz, 2H), 3.34 (s, 2H), 3.09 – 3.03 (m, 2H), 2.95 (s, 3H), 2.72 (s, 3H), 1.30 (d, J = 6.5 Hz, 6H).
117	MS m/z 432.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.90 (s, 1H), 9.58 (s, 1H), 8.96-9.14 (m, 1H), 7.90-8.03 (m, 1H), 7.80-7.86 (m, 1H), 7.21 (s, 1H), 6.93-7.05 (m, 1H), 4.85 (d, J = 14.0 Hz, 2H), 3.35-3.38 (m, 2H), 3.19-3.23 (m, 2H), 2.90 (s, 3H), 2,61 (s, 3H), 1.38 (t, J = 5.9 Hz, 6H).
118	MS m/z 471.4 $[M+H]^+$
119	MS m/z 435.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.21 (s, 1H), 9.49 (s, 1H), 9.10 (s, 1H), 8.73 (s, 1H), 8.07 (d, J = 8.1 Hz, 1H), 7.84 (d, J = 1.8 Hz, 1H), 7.76 (dd, J = 8.2, 1.8 Hz, 1H), 4.89 (d, J = 13.8 Hz, 2H), 3.42 (s, 2H), 3.11 (dd, J = 14.1, 11.5 Hz, 2H), 2.69 (s, 3H), 1.36 (d, J = 6.5 Hz, 6H).
120	MS m/z 417.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.70 (d, J = 10.1 Hz, 1H), 9.34 (d, J = 11.1 Hz, 1H), 9.13 (s, 1H), 8.47 (d, J = 8.3 Hz, 1H), 8.06 (d, J = 8.2 Hz, 1H), 7.78 (s, 1H), 7.74 (dd, J = 8.2, 1.7 Hz, 1H), 7.70 (d, J = 1.8 Hz, 1H), 7.56 (d, J = 8.6 Hz, 1H), 4.86 (d, J = 13.8 Hz, 2H), 3.37-3.42 (m, 2H), 3.11-3.20 (m, 2H), 2.74 (s, 3H), 1.37 (d, J = 6.5 Hz, 6H).
128	MS m/z 418.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.75 (d, J = 1.5 Hz, 1H), 9.54 (d, J = 1.4 Hz, 1H), 9.45 (d, J = 10.6 Hz, 1H), 9.25 (d, J = 10.9 Hz, 1H), 9.15 (s, 1H), 8.80 (s, 1H), 8.01 (d, J = 8.2 Hz, 1H), 7.88 (d, J = 1.7 Hz, 1H), 7.79 (dd, J = 8.2, 1.8 Hz, 1H), 4.72-4.92 (m, 2H), 3.51 (td, J = 13.2, 3.1 Hz, 1H), 3.38-3.45 (m, 1H), 3.28 (dd, J = 14.1, 11.1 Hz, 1H), 3.14-3.17 (m, 2H), 2.05 (h, J = 6.8 Hz, 1H), 1.08 (dd, J = 6.9, 5.5 Hz, 6H).

Cpd	Data
129	MS m/z 432.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.75 (d, J = 1.4 Hz, 1H), 9.54 (d, J = 1.4 Hz, 1H), 9.34 (d, J = 10.7 Hz, 1H), 9.15 (s, 1H), 8.93 (d, J = 11.2 Hz, 1H), 8.80 (s, 1H), 8.01 (d, J = 8.2 Hz, 1H), 7.88 (d, J = 1.7 Hz, 1H), 7.79 (dd, J = 8.2, 1.8 Hz, 1H), 4.96 (d, J = 13.5 Hz, 1H), 4.84 (d, J = 14.0 Hz, 1H), 3.45-3.55 (m, 1H), 3.42 (d, J = 12.3 Hz, 1H), 3.17-3.29 (m, 3H), 1.11 (s, 9H).
130	MS m/z 416.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.73-9.81 (m, 2H), 9.54 (d, J = 1.4 Hz, 1H), 9.43 (d, J = 10.4 Hz, 1H), 9.15 (s, 1H), 8.80 (s, 1H), 8.00 (d, J = 8.2 Hz, 1H), 7.88 (d, J = 1.7 Hz, 1H), 7.79 (dd, J = 8.2, 1.8 Hz, 1H), 4.67-4.87 (m, 2H), 3.47-3.61 (m, 2H), 3.39-3.46 (m, 1H), 3.07 (dt, J = 15.8, 8.5 Hz, 1H), 2.60-2.75 (m, 1H), 1.06-1.17 (m, 1H), 0.73 – 0.57 (m, 3H), 0.41-0.46 (m, 1H).
131	MS m/z 444.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.54 (br s, 1H), 9.32 (br s, 1H), 9.16 (s, 1H), 9.14 (s, 1H), 8.12 (s, 2H), 8.04 (d, J=8.2 Hz, 1H), 7.40 (br s, 1H), 7.38 (s, 1H), 4.85 (d, J=14.0 Hz, 1H), 4.77 (d, J=12.8 Hz, 1H), 3.48-3.57 (m, 1H), 3.41 (d, J=13.7 Hz, 1H), 3.24-3.34 (m, 1H), 3.07-3.20 (m, 2H), 2.68 (s, 3H), 2.55 (s, 3H), 2.06 (dq, J=13.7, 6.8 Hz, 1H), 1.08 (t, J=6.8 Hz, 6H).
138	MS m/z 458.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.11 (s, 1H), 8.54 (s, 1H), 7.92 (d, J=8.2 Hz, 1H), 7.66 (s, 1H), 7.41 (s, 1H), 7.29 (br d, J=8.2 Hz, 1H), 7.25 (s, 1H), 3.25-3.50 (m, 4H), 3.11-3.18 (m, 2H), 3.03 (q, J=7.4 Hz, 2H), 2.95 (br t, J=7.5 Hz, 1H), 2.46 (s, 3H), 1.97 (dq, J=14.0, 6.7 Hz, 1H), 1.42 (t, J=7.4 Hz, 3H), 1.15 (dd, J=6.7, 2.7 Hz, 6H); 2Hs not observed (NH and OH).
139	MS m/z 498.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.13 (s, 1H), 9.07 (s, 1H), 8.16 (s, 1H), 7.98 (s, 1H), 7.96 (s, 1H), 7.36 (dd, J=8.2, 1.8 Hz, 1H), 7.32 (s, 1H), 5.06 (d, J=14.0 Hz, 1H), 4.96 (d, J=14.0 Hz, 1H), 3.43-3.64 (m, 2H), 3.28-3.32 (m, 1H), 3.27 (d, J=4.0 Hz, 1H), 3.18 (ddd, J=10.4, 7.9, 3.7 Hz, 1H), 2.54 (s, 3H), 2.09 (dq, J=13.7, 6.7 Hz, 1H), 1.19 (dd, J=6.7, 3.7 Hz, 6H); 2 Hs not observed (NH and OH).
140	MS m/z 444.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.02 (s, 1H), 8.22 (s, 1H), 7.84 (d, J=8.2 Hz, 1H), 7.78 (s, 1H), 7.39 (s, 1H), 7.26 (dd, J=8.2, 1.2 Hz, 1H), 7.24 (s, 1H), 4.77 (br d, J=12.2 Hz, 1H), 4.65 (d, J=13.7 Hz, 1H), 4.24 (s, 3H), 3.09-3.17 (m, 2H), 2.73-2.89 (m, 2H), 2.64 (s, 3H), 2.43 (ddd, J=11.3, 6.8, 2.8 Hz, 1H), 1.72 (dq, J=13.7, 6.8 Hz, 1H), 1.05 (dd, J=6.8, 3.7 Hz, 6H); 2 Hs not observed (NH and OH).
141	MS m/z 431.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.41 (s, 1H), 9.36 (s, 1H), 9.21 (s, 1H), 8.23 (s, 1H), 8.04 (d, J=8.2 Hz, 1H), 7.77 (s, 1H), 7.74 (br d, J=8.2 Hz, 1H), 5.09 (d, J=13.7 Hz, 1H), 4.99 (d, J=15.0 Hz, 1H), 3.35-3.64 (m, 4H), 3.16-3.25 (m, 1H), 2.68 (s, 3H), 2.02-2.18 (m, 1H), 1.20 (dd, J=6.7, 3.1 Hz, 6H); 2 Hs not observed (NH and OH).
147	MS m/z 431.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.08-9.18 (m, 1H), 8.31-8.37 (m, 1H), 8.00-8.06 (m, 1H), 7.87-7.95 (m, 1H), 7.51-7.58 (m, 2H), 6.06-6.14 (m, 1H), 4.97-5.14 (m, 1H), 4.19-4.41 (m, 2H), 3.69-3.82 (m, 1H), 3.54-3.65 (m, 1H), 3.36-3.53 (m, 2H), 2.79 (s, 3H), 2.04-2.18 (m, 1H), 1.17-1.24 (m, 6H); 2Hs not observed (NH and OH).

Cpd	Data
167	MS m/z 418.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.13 (s, 1H), 9.63 (d, J = 1.4 Hz, 1H), 9.38 (d, J = 1.4 Hz, 1H), 9.20 (d, J = 10.0 Hz, 1H), 9.15 (s, 1H), 8.00 (d, J = 8.2 Hz, 1H), 7.84 (d, J = 1.8 Hz, 1H), 7.76 (dd, J = 8.2, 1.8 Hz, 1H), 4.90 (d, J = 13.9 Hz, 2H), 3.45 (s, 2H), 3.03 (dd, J = 14.2, 11.4 Hz, 2H), 2.60 (s, 3H), 1.33 (d, J = 6.4 Hz, 6H).
168	MS m/z 430.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.34 (s, 1H), 9.12 (d, J = 2.0 Hz, 1H), 9.10 (s, 1H), 8.68 (d, J = 2.0 Hz, 1H), 8.00 (d, J = 2.0 Hz, 1H), 7.45 (d, J = 8.0 Hz, 1H), 7.42 (s, 1H), 4.67 (d, J = 12.5 Hz, 1H), 4.58 (s, 3H), 3.16 (t, J = 10.5 Hz, 2H), 3.06 (t, J = 11.5 Hz, 1H), 2.78 (t, J = 8.5 Hz, 1H), 2.13 (t, J = 8.5 Hz, 1H), 0.93-0.84 (m, 1H), 0.56-0.47 (m, 2H), 0.42-0.35 (m, 1H), 0.32-0.28 (m, 1H).
169	MS m/z 430.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.27 (s, 1H), 9.47 (s, 1H), 9.28 (s, 1H), 9.16 (s, 1H), 9.04 (d, J = 2.0 Hz, 1H), 8.70 (d, J = 2.0 Hz, 1H), 8.04 (d, J = 8.0 Hz, 1H), 7.49 (d, J = 9.0 Hz, 1H), 7.47 (s, 1H), 4.76 (d, J = 13.5 Hz, 1H), 4.70 (d, J = 14.5 Hz, 1H), 4.41 (s, 3H), 3.55-3.43 (m, 1H), 3.12-3.07 (m, 2H), 2.71-2.63 (m, 1H), 1.11-1.05 (m, 1H), 0.71-0.64 (m, 1H), 0.60-0.56 (m, 1H), 0.46-0.41 (m, 1H), 0.35-0.27 (m, 1H).
171	MS m/z 430.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.04 (s, 1H), 9.89 (s, 1H), 9.49 (s, 1H), 9.12 (s, 1H), 8.50 (d, J = 8.5 Hz, 1H), 8.23 (d, J = 8.5 Hz, 1H), 8.01 (d, J = 8.0 Hz, 1H), 7.94 (d, J = 1.5 Hz, 1H), 7.81 (dd, J = 8.0, 1.5 Hz, 1H), 4.76 (d, J = 14.0 Hz, 1H), 4.70 (d, J = 14.0 Hz, 1H), 4.38 (s, 3H), 3.60-3.49 (m, 2H), 3.42 (d, J = 10.5 Hz, 1H), 3.10-3.04 (m, 1H), 2.67-2.63 (m, 1H), 1.14-1.07 (m, 1H), 0.68-0.58 (m, 3H), 0.45-0.41 (m, 1H); 1H is from HCl salt.
172	MS m/z 430.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.13 (s, 1H), 9.57 (s, 1H), 9.08 (s, 1H), 8.47 (d, J = 1.5 Hz, 1H), 7.97 (d, J = 8.5 Hz, 1H), 7.88 (d, J = 1.5 Hz, 1H), 7.79 (dd, J = 8.5, 2.0 Hz, 1H), 4.66 (d, J = 12.5 Hz, 1H), 4.61 (s, 3H), 4.57 (d, J = 12.5 Hz, 1H), 3.14 (t, J = 9.0 Hz, 2H), 3.02 (t, J = 11.5 Hz, 1H), 2.75 (t, J = 11.5 Hz, 1H), 2.09 (t, J = 8.0 Hz, 1H), 0.90-0.82 (m, 1H), 0.54-0.46 (m, 2H), 0.38-0.34 (m, 1H), 0.31-0.27 (m, 1H); 1H not observed (NH or OH).
173	MS m/z 430.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.13 (s, 1H), 10.04 (s, 1H), 9.59 (s, 1H), 9.11 (s, 1H), 8.63 (d, J = 9.0 Hz, 1H), 8.08 (d, J = 9.0 Hz, 1H), 7.95 (s, 1H), 7.84 (d, J = 10.0 Hz, 1H), 4.76 (d, J = 13.0 Hz, 1H), 4.69 (d, J = 13.0 Hz, 1H), 4.36 (s, 3H), 3.62-3.51 (m, 2H), 3.40 (d, J = 12.5 Hz, 1H), 3.08-3.01 (m, 1H), 2.67-2.61 (m, 1H), 1.17-1.09 (m, 1H), 0.69-0.64 (m, 3H), 0.44-0.40 (m, 1H).
174	MS m/z 430.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.08 (s, 1H), 9.80 (s, 1H), 9.53 (s, 1H), 9.43 (s, 1H), 9.15 (s, 1H), 8.53 (s, 1H), 8.02 (d, J = 8.5 Hz, 1H), 7.97 (d, J = 1.5 Hz, 1H), 7.83 (dd, J = 8.5, 1.5 Hz, 1H), 4.76 (d, J = 16.0 Hz, 1H), 4.69 (d, J = 13.5 Hz, 1H), 4.40 (s, 3H), 3.58-3.41 (m, 3H), 3.09-3.03 (m, 1H), 2.70-2.63 (m, 1H), 1.14-1.07 (m, 1H), 0.72-0.58 (m, 3H), 0.45-0.41 (m, 1H); 1H from HCl salt.

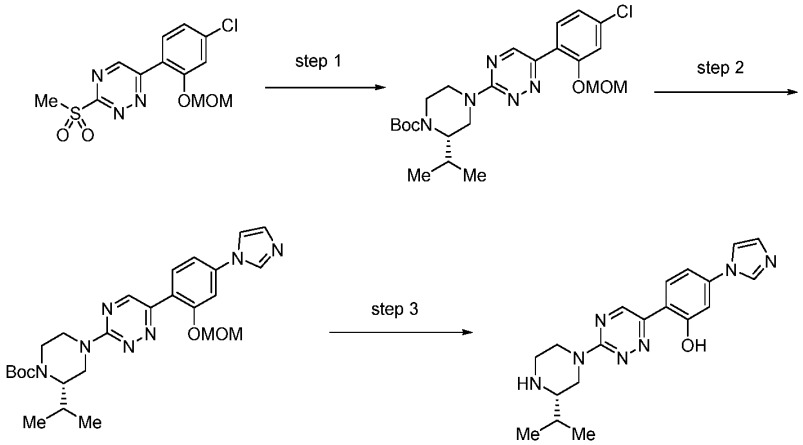
Cpd	Data
176	MS m/z 352.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 7.70 (s, 1H), 6.65 (br s, 2H), 6.46 (d, J = 8.2 Hz, 1H), 5.87 (dd, J = 8.1, 1.8 Hz, 1H), 5.84 (d, J = 1.7 Hz, 1H), 3.59 (t, J = 6.0 Hz, 1H), 3.25 (dd, J = 12.8, 2.6 Hz, 1H), 1.84 – 1.66 (m, 2H), 1.61 – 1.48 (m, 2H), 0.01 (d, J = 6.9 Hz, 3H), 0.10 (d, J = 6.0 Hz, 3H); 3Hs not observed (2 NHs and OH).
190	MS m/z 352.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ 9.54 (br s, 1H), 9.11 (s, 1H), 8.09 (s, 2H), 7.86 (d, J = 8.0 Hz, 1H), 7.32 (s, 1H), 7.27 – 7.19 (m, 2H), 7.12 (s, 1H), 5.05 (t, J = 7.2 Hz, 1H), 4.50 (d, J = 14.3 Hz, 1H), 3.77 (s, 1H), 3.59 (dd, J = 14.5, 3.7 Hz, 1H), 3.4-3.42 (m, 1H), 3.22 – 3.11 (m, 1H), 1.39 (d, J = 6.9 Hz, 3H), 1.32 (d, J = 6.8 Hz, 3H).
191	MS m/z 434.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.14 (s, 1H), 9.11 (s, 1H), 8.76-8.99 (m, 1H), 8.66 (dd, J = 4.6, 1.5 Hz, 1H), 8.49 (dd, J = 8.2, 1.5 Hz, 1H), 8.06 (d, J = 8.1 Hz, 1H), 7.80 (d, J = 1.8 Hz, 1H), 7.73 (dd, J = 8.1, 1.8 Hz, 1H), 7.65 (dd, J = 8.2, 4.6 Hz, 1H), 4.88 (d, J = 13.5 Hz, 1H), 4.80 (d, J = 14.0 Hz, 1H), 3.38-3.48 (m, 2H), 3.16-3.28 (m, 3H), 1.98 (h, J = 6.8 Hz, 1H), 1.07 (d, J = 6.9 Hz, 6H).
193	MS m/z 429.4 $[M+H]^+$; ^1H NMR (DMSO-d6) δ: 11.28-11.58 (m, 1H), 9.33 (s, 1H), 9.10 (s, 1H), 8.21 (s, 1H), 8.02 (d, J=9.5 Hz, 1H), 7.99 (d, J=8.2 Hz, 1H), 7.68 (dd, J=9.5, 1.2 Hz, 1H), 7.46 (dd, J=8.1, 1.7 Hz, 1H), 7.41 (d, J=1.5 Hz, 1H), 4.64 (br d, J=12.2 Hz, 1H), 4.55 (br d, J=11.3 Hz, 1H), 3.00-3.13 (m, 2H), 2.90 (br dd, J=12.7, 10.5 Hz, 1H), 2.62-2.71 (m, 1H), 2.58 (s, 3H), 1.90-1.98 (m, 1H), 0.78-0.85 (m, 1H), 0.42-0.51 (m, 2H), 0.23-0.34 (m, 2H).
194	MS m/z 428.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.14-11.41 (m, 1H), 9.09 (s, 1H), 8.63 (s, 1H), 8.31 (s, 1H), 8.16 (s, 1H), 7.96 (d, J=7.9 Hz, 1H), 7.61 (d, J=9.5 Hz, 1H), 7.30 (dd, J=8.1, 1.7 Hz, 1H), 7.27 (d, J=1.5 Hz, 1H), 7.01 (dd, J=9.5, 1.2 Hz, 1H), 4.67 (br d, J=12.2 Hz, 1H), 4.58 (br d, J=12.8 Hz, 1H), 3.10-3.21 (m, 2H), 2.97-3.09 (m, 1H), 2.71-2.83 (m, 1H), 2.44 (s, 3H), 2.12 (br t, J=8.4 Hz, 1H), 0.81-0.94 (m, 1H), 0.47-0.60 (m, 2H), 0.26-0.41 (m, 2H).
195	MS m/z 428.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.08 (s, 1H), 8.20 (s, 2H), 8.17 (d, J=7.6 Hz, 1H), 7.95 (d, J=8.2 Hz, 1H), 7.88 (s, 1H), 7.31-7.41 (m, 3H), 7.04 (dd, J=7.5, 1.7 Hz, 1H), 4.64 (br d, J=12.2 Hz, 1H), 4.49-4.59 (m, 1H), 3.04-3.11 (m, 1H), 2.92 (br dd, J=12.7, 10.5 Hz, 1H), 2.64-2.73 (m, 2H), 2.63 (s, 3H), 1.92-2.02 (m, 1H), 0.78-0.89 (m, 1H), 0.42-0.53 (m, 2H), 0.23-0.36 (m, 2H).
197	MS m/z 418.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.13 (d, J = 2.1 Hz, 1H), 9.08 (s, 1H), 8.69 (d, J = 2.1 Hz, 1H), 8.01 (d, J = 8.2 Hz, 1H), 7.45 (dd, J = 1.7, 8.1 Hz, 1H), 7.41 (d, J = 1.8 Hz, 1H), 4.59 (s, 5H), 2.78 (ddd, J = 3.1, 6.5, 10.1 Hz, 2H), 2.56 - 2.52 (m, 2H), 1.07 (d, J = 6.1 Hz, 6H); 2Hs not observed (NH and OH).
198	MS m/z 418.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.57 (s, 1H), 9.06 (s, 1H), 8.47 (s, 1H), 7.97 (d, J = 8.2 Hz, 1H), 7.90 - 7.87 (m, 1H), 7.80 - 7.77 (m, 1H), 4.65 - 4.57 (m, 5H), 2.78 (dd, J = 3.1, 6.3 Hz, 2H), 2.58 - 2.52 (m, 2H), 1.11 - 1.03 (m, 6H); 2Hs not observed (NH and OH).

Cpd	Data
201	MS m/z 418.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.06 (s, 1H), 8.53 (d, J = 9.0 Hz, 1H), 8.12 (d, J = 8.9 Hz, 1H), 8.01 (d, J = 8.2 Hz, 1H), 7.90 - 7.86 (m, 1H), 7.82 - 7.77 (m, 1H), 4.70 - 4.60 (m, 5H), 2.91 - 2.78 (m, 2H), 2.63 - 2.54 (m, 2H), 1.12 - 1.07 (m, 6H); 2Hs not observed (NH and OH).
202	MS m/z 430.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.15 (s, 1H), 9.06 (s, 1H), 8.52 (d, J = 9.0 Hz, 1H), 8.11 (d, J = 9.0 Hz, 1H), 8.00 (d, J = 8.5 Hz, 1H), 7.87 (s, 1H), 7.78 (d, J = 9.5 Hz, 1H), 4.65 (d, J = 9.5 Hz, 1H), 4.56 (s, 3H), 4.55 (d, J = 12.0 Hz, 1H), 4.15-4.03 (m, 1H), 3.07 (t, J = 11.5 Hz, 2H), 2.92 (t, J = 11.0 Hz, 1H), 2.71-2.65 (m, 1H), 1.97 (t, J = 8.5 Hz, 1H), 0.85-0.78 (m, 1H), 0.51-0.43 (m, 2H), 0.32-0.24 (m, 2H).
206	MS m/z 417.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.32 (s, 1H), 9.08 (s, 1H), 8.03 - 7.97 (m, 2H), 7.68 (d, J = 9.3 Hz, 1H), 7.45 (dd, J = 1.6, 8.2 Hz, 1H), 7.39 (d, J = 1.5 Hz, 1H), 4.60 (d, J = 11.9 Hz, 2H), 2.77 (ddd, J = 3.1, 6.4, 10.0 Hz, 2H), 2.58 (s, 3H), 1.09 - 1.03 (m, 7H); 3Hs not observed (NH, OH and 1 CH overlapped with solvent peak).
211	MS *m/z* 446.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.22 (s, 1H), 9.33 (d, J = 9.0 Hz, 1H), 9.15 (s, 1H), 9.12 (s, 1H), 8.92 (d, J = 9.0 Hz, 1H), 8.68 (s, 1H), 8.01 (d, J = 8.0 Hz, 1H), 7.46 (d, J = 8.0 Hz, 1H), 7.45 (s, 1H), 4.95 (d, J = 13.5 Hz, 1H), 4.83 (d, J = 14.0 Hz, 1H), 4.58 (s, 3H), 3.49 (t, J = 12.5 Hz, 2H), 3.27-3.18 (m, 3H), 1.10 (s, 9H); 1H from HCl salt.
212	MS *m/z* 432.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.42 (s, 1H), 9.14 (s, 1H), 9.08 (s, 1H), 8.67 (d, J = 2.0 Hz, 1H), 8.00 (d, J = 2.0 Hz, 1H), 7.47 (d, J = 7.5 Hz, 1H), 7.42 (s, 1H), 4.91-4.48 (m 2H), 4.61 (s, 3H), 3.54-3.46 (m, 3H), 3.11 (t, J = 11.5 Hz, 1H), 2.70 (t, J = 9.0 Hz, 1H), 2.44-2.35 (m, 1H), 1.09-1.02 (m, 1H), 0.99 (d, J = 6.0 Hz, 6H).
214	MS *m/z* 432.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.27 (s, 1H), 9.61 (s, 1H), 9.49 (s, 1H), 9.14 (s, 1H), 9.13 (d, J = 8.5 Hz, 1H), 8.68 (s, 1H), 8.01 (d, J = 8.5 Hz, 1H), 4.93 (d, J = 12.5 Hz, 1H), 4.85 (d, J = 12.5 Hz, 1H), 4.58 (s, 3H), 3.39-3.37 (m, 1H), 3.25-3.13 (m, 4H), 1.87-1.79 (m, 1H), 1.75-1.68 (m, 1H), 1.39 (d, J = 6.0 Hz, 3H), 1.04 (d, J = 7.5 Hz, 3H).
241	MS m/z 444.0 $[M+H]^+$; ^1H NMR (400 MHz, CDCl3) δ : 12.46 (s, 1H), 9.10 (d, J = 2.2 Hz, 1H), 8.87 (s, 1H), 8.38 (d, J = 2.2 Hz, 1H), 7.77 (d, J = 8.3 Hz, 1H), 7.39 (d, J = 1.9 Hz, 1H), 7.27-7.24 (m, 1H), 4.78-4.70 (m, 2H), 4.60 (s, 3H), 3.23-3.18 (m, 1H), 3.11 (td, J = 11.7, 3.1 Hz, 1H), 3.03 (t, J = 12, 1H), 2.87 (td, J = 11.7, 3.1 Hz, 1H), 1.96 (dd, J = 10.8, 2.9 Hz, 1H), 1.13 (s, 3H), 0.53-0.46 (m, 1H), 0.43 – 0.32 (m, 3H); 1H not observed (NH or OH).
242	MS m/z 444.0 $[M+H]^+$; ^1H NMR (400 MHz, $CDCl_3$) δ: 12.46 (s, 1H), 9.10 (d, J = 2.2 Hz, 1H), 8.87 (s, 1H), 8.38 (d, J = 2.2 Hz, 1H), 7.77 (d, J = 8.3 Hz, 1H), 7.39 (d, J = 1.9 Hz, 1H), 7.27-7.24 (m, 1H), 4.78-4.70 (m, 2H), 4.60 (s, 3H), 3.23-3.18 (m, 1H), 3.11 (td, J = 11.7, 3.1 Hz, 1H), 3.03 (t, J = 12 Hz, 1H), 2.87 (td, J = 11.7, 3.1 Hz, 1H), 1.96 (dd, J = 10.8, 2.9 Hz, 1H), 1.13 (s, 3H), 0.53-0.46 (m, 1H), 0.43 – 0.32 (m, 3H); 1H not observed (NH or OH).

Cpd	Data
243	MS m/z 417.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.09 (s, 1H), 9.22 (dd, J = 7.0, 1.7 Hz, 1H), 9.15 (s, 1H), 8.94 – 8.86 (m, 1H), 8.81 (s, 1H), 8.78 (s, 1H), 8.73-8.72 (m, 1H), 7.96 – 7.92 (m, 2H), 7.74 (d, J = 8.2, 1H), 7.17 (dd, J = 7.1, 4.1 Hz, 1H), 4.85 (d, J = 12.5 Hz, 1H), 4.78 (d, J = 13.9 Hz, 1H), 3.56 – 3.26 (m, 2H), 3.22-3.18 (m, 3H), 1.98 (q, J = 6.7 Hz, 1H), 1.07 (dd, J = 6.9, 3.9 Hz, 6H); 1H from HCl salt.
244	MS m/z 416.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.21 (s, 1H), 9.06 (s, 1H), 8.77 (d, J = 7.0 Hz, 1H), 8.43 (s, 1H), 8.01 (d, J = 9.0 Hz, 1H), 7.94 (d, J = 8.5 Hz, 1H), 7.39 (dd, J = 9.0, 6.8 Hz, 1H), 7.33-7.32 (m, 2H), 7.00 (t, J = 6.9 Hz, 1H), 4.64 (d, J = 11.4 Hz, 1H), 4.54 (d, J = 12.6 Hz, 1H), 3.21 – 2.90 (m, 2H), 2.86 – 2.61 (m, 2H), 2.37-2.33 (m, 2H), 1.72 – 1.62 (m, 1H), 0.98 (d, J = 6.8 Hz, 5H); 1 H not observed (NH or OH).
263	MS m/z 434.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.16 (br s, 1H), 9.16 (s, 1H), 9.04 (br s, 2H), 8.37 (s, 1H), 8.23 (d, J = 9.1 Hz, 1H), 8.09 (d, J = 9.7 Hz, 1H), 8.03 (d, J = 8.1 Hz, 1H), 7.51 (d, J = 8.2 Hz, 1H), 7.47 (s, 1H), 4.87 (d, J = 13.7 Hz, 1H), 4.80 (d, J = 14.1 Hz, 1H), 3.51 – 3.44 (m, 2H), 3.31 – 3.00 (m, 3H), 2.07 – 1.97 (m, 1H), 1.08 (dd, J = 6.8, 4.6 Hz, 6H); 1H from HCl salt.
264	MS m/z 435.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.29 (br s, 1H), 9.52 (d, J = 2.3 Hz, 1H), 9.44 (br s, 1H), 9.25 (br s, 1H), 9.17 (s, 1H), 8.82 (d, J = 2.3 Hz, 1H), 8.06 (d, J = 8.1 Hz, 1H), 7.59 (d, J = 8.0 Hz, 1H), 7.56 (s, 1H), 4.87 (d, J = 14.2 Hz, 1H), 4.79 (d, , J = 14.3 Hz, 1H), 3.61 – 3.39 (m, 3H), 3.29 (dd, J = 14.0, 11.1 Hz, 1H), 3.16-3.13 (m, 1H), 2.06 (h, J = 6.8 Hz, 1H), 1.09 (t, J = 6.0 Hz, 6H); 1H from HCl salt.

Example 3

Preparation of Compound 142



Step 1: To a solution of 6-(4-chloro-2-(methoxymethoxy)phenyl)-3-(methylsulfonyl)-1,2,4-triazine (1.0g, 3.2 mmol) and (*S*)-1-Boc-2-Isopropylpiperazine (873 mg, 3.81 mmol) in ACN (6 mL) was added DIPEA (1.1 mL, 6.4 mmol). The mixture was heated at 70 for 3 h

until UPLC showed complete consumption of the starting material. The reaction mixture was then cooled to room temperature, concentrated and purified by silica gel column chromatography eluting with a gradient CH_2Cl_2/MeOH (0-15% MeOH) to afford tert-butyl (S)-4-(6-(4-chloro-2-(methoxymethoxy)phenyl)-1,2,4-triazin-3-yl)-2-isopropylpiperazine-1-carboxylate (1.02 g, 67% yield) as an off white foam. MS m/z 478.4 $[M+H]^+$.

Step 2: A mixture of $Pd_2(dba)_3$ (10.0 mg, 0.01 mmol), Me4tButylXphos (10 mg, 0.01 mmol), 1,4-dioxane (0.2 mL) and toluene (0.8 mL) was purged with Ar and then heated at 120 ºC for 10 minutes. The reaction was cooled down to rt and then transferred into the vial containing tert-butyl (S)-4-(6-(4-chloro-2-(methoxymethoxy)phenyl)-1,2,4-triazin-3-yl)-2-isopropylpiperazine-1-carboxylate (100 mg, 0.21 mmol), K_3PO_4 (90 mg, 0.42 mmol) and imidazole (28 mg, 0.42 mmol). The combined mixture was then purged with Ar and then heated at 120 ºC for 4 h. The reaction mixture was then cooled to room temperature, concentrated and purified by silica gel column chromatography eluting with a gradient CH_2Cl_2/MeOH (0-10% MeOH) to afford tert-butyl (S)-4-(6-(4-(1H-imidazol-1-yl)-2-(methoxymethoxy)phenyl)-1,2,4-triazin-3-yl)-2-isopropylpiperazine-1-carboxylate (76 mg, 71% yield) as a yellow solid. MS m/z 510.4 $[M+H]^+$.

Step 3: To a solution of tert-butyl (S)-4-(6-(4-(1H-imidazol-1-yl)-2-(methoxymethoxy)phenyl)-1,2,4-triazin-3-yl)-2-isopropylpiperazine-1-carboxylate (76 mg, 0.15 mmol) in methanol (2 mL) was added HCl (4 mol/L) in 1,4-dioxane (0.2 mL, 0.8 mmol). The reaction was stirred at room temperature for 2h, concentrated and purified by silica gel column chromatography, eluting with a gradient CH_2Cl_2/MeOH/NH_4OH (10-30% MeOH/NH_4OH) to afford (S)-5-(1H-imidazol-1-yl)-2-(3-(3-isopropylpiperazin-1-yl)-1,2,4-triazin-6-yl)phenol (30 mg, 47% yield).

MS m/z 366.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.14 (s, 1H), 8.22 (s, 1H), 8.01 (d, J = 8.4 Hz, 1H), 7.61 (s, 1H), 7.22 - 7.17 (m, 3H), 5.08 (br d, J = 13.6 Hz, 1H), 5.02 - 4.93 (m, 1H), 3.59 - 3.45 (m, 2H), 3.31 - 3.23 (m, 2H), 3.18 - 3.11 (m, 1H), 2.11 - 2.02 (m, 1H), 1.19 (d, J = 6.9 Hz, 6H); 2Hs not observed (NH and OH).

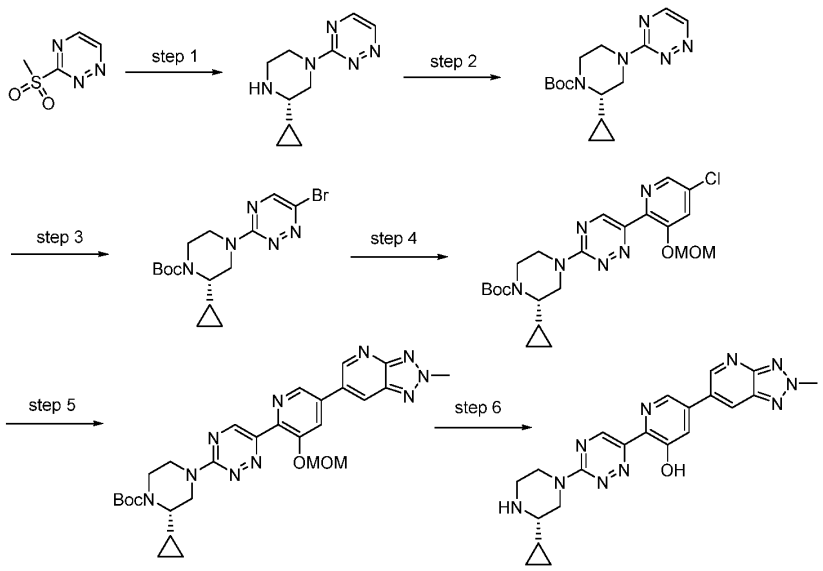
Using the procedure described for Example 3, above, additional compounds described herein may be prepared by substituting the appropriate starting material, suitable reagents and reaction conditions, obtaining compounds such as those selected from:

Cpd	Data
159	MS m/z 430.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.18 (s, 1H), 8.35 (br s, 1H), 8.07 (d, J=8.5 Hz, 1H), 8.04 (d, J=8.5 Hz, 1H), 7.86 (d, J=3.4 Hz, 1H), 7.19-7.25 (m, 2H), 6.79 (d, J=3.4 Hz, 1H), 5.07 (br d, J=13.7 Hz, 1H), 4.97 (br d, J=14.3 Hz, 1H), 3.49-3.55 (m, 1H), 3.40-3.48 (m, 1H), 3.20-3.30 (m, 2H), 3.11 (ddd, J=10.6, 7.4, 3.1 Hz, 1H), 2.67 (s, 3H), 1.96-2.08 (m, 1H), 1.18 (d, J=7.0 Hz, 6H); 2Hs not observed (NH and OH).
54	MS *m/z* 353.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.13 (s, 1 H), 8.01 (s, 1 H), 7.85 (s, 1 H), 7.84 (d, J=8.09 Hz, 1 H), 7.20 (d, J=6.56 Hz, 1 H), 7.16 (d, J=1.53 Hz, 1H), 5.05 (dd, J=14.34, 2.75 Hz, 2H), 3.48 (m, 2H), 3.04 (dd, J=14.50, 11.60 Hz, 2H), 1.44 (d, J=6.56 Hz, 6H); 2Hs not observed (NH and OH).
52	MS *m/z* 367.4 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.08 (s, 1H), 7.97 (d, J=9.3 Hz, 1H), 7.96 (s, 2H), 7.70-7.73 (m, 2H), 4.69 (br d, J=14.6 Hz, 2H), 2.90 (dd, J=13.4, 11.1 Hz, 2H), 2.34-2.41 (m, 5H), 1.26 (s, 3H), 1.25 (s, 3H); 1H not observed (OH).
45	MS *m/z* 355.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.08 (s, 1H), 7.97 (d, J=9.0 Hz, 1H), 7.96 (s, 2H), 7.71 (s, 2H), 4.79 (br d, J=11.4 Hz, 1H), 4.67 (d, J=12.8 Hz, 1H), 3.57-3.69 (m, 2H), 3.14-3.22 (m, 2H), 2.85-2.96 (m, 3H); 3Hs not observed (NH and 2 OHs).
274	MS m/z 436.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.05 (s, 1H), 8.45-8.60 (s, 1H), 7.73-7.83 (m, 1H), 6.70-6.79 (m, 1H), 6.59-6.66 (m, 1H), 4.89-4.95 (m, 1H), 4.74-4.82 (m, 1H), 4.56 (s, 2H), 4.21-4.29 (m, 2H), 3.88-3.97 (m, 2H), 3.34-3.38 (m, 1H), 3.21-3.30 (m, 1H), 2.97-3.10 (m, 2H), 2.74-2.86 (m, 1H), 2.38 (s, 3H), 1.82-1.93 (m, 1H), 1.05-1.18 (m, 6H); 2Hs not observed (NH and OH).
155	MS m/z 431.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.59 (br d, J=9.5 Hz, 1H), 9.34 (br d, J=8.9 Hz, 1H), 9.08 (s, 1H), 8.60 (s, 1H), 8.52 (d, J=8.9 Hz, 1H), 8.04 (d, J=8.5 Hz, 1H), 7.59 (d, J=9.2 Hz, 1H), 7.56 (d, J=2.1 Hz, 1H), 7.47 (dd, J=8.5, 2.1 Hz, 1H), 4.86 (br d, J=13.4 Hz, 1H), 4.78 (br d, J=14.0 Hz, 1H), 3.47-3.56 (m, 1H), 3.39-3.45 (m, 1H), 3.29 (dd, J=14.0, 11.3 Hz, 1H), 3.09-3.18 (m, 2H), 2.73 (s, 3H), 2.01-2.12 (m, 1H), 1.04-1.12 (m, 6H).
237	MS m/z 393.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.30 (s, 1H), 8.98 (s, 1H), 8.14 (s, 1H), 7.99 (d, J = 8.5 Hz, 1H), 7.68 (s, 1H), 7.63 (d, J = 8.5 Hz, 1H), 4.69-4.51 (m, 2H), 2.76-2.44 (m, 4H), 2.29-2.21 (m, 1H), 2.04-1.98 (m, 2H), 1.91-1.76 (m, 4H), 1.09 (d, J = 5.0 Hz, 3H); 2Hs not observed (NH and OH).

Cpd	Data
162	MS m/z 418.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.76 (br, d, J=9.46 Hz, 1H), 9.56 (s, 1H), 9.46 (br, d, J=9.54 Hz, 1H), 9.25 (s, 1H), 9.08 (s, 1H), 8.76 (s, 1H), 8.03 - 8.06 (m, 3H), 7.93 (dd, J=8.7, 1.98 Hz, 1H), 4.77 (br d, J=14.04 Hz, 2H), 4.85 (br, d, J=13.43 Hz, 1H), 3.52 - 3.57 (m, 1H), 3.38 - 3.41 (m, 1H), 3.27 - 3.30 (m, 2H), 2.05 - 2.12 (m, 1H), 1.07 - 1.10 (t, J=6.82 Hz, 6H); 1H from HCl salt.
183	MS m/z 379.5 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.77 (br s, 1H), 9.42 - 9.43 (m, 1H), 9.05 (s, 1H), 7.98 (d, J=8.54 Hz, 1H), 7.92 (s, 1H), 7.68 (d, J=2.14 Hz, 1H), 7.58 (dd, J=8.54, 2.14 Hz, 1H), 4.68 - 4.77 (m, 2H), 3.41 - 3,58 (m, 2H), 3.03 - 3.10 (m, 1H), 2.65 - 2.69 (m, 1H), 2.38 (s, 2H), 2.27 (s, 3H), 1.05 -1.14 (m, 1H), 0.60 - 0.68 (m, 2H), 0.42 - 0.44 (m, 1H).
184	MS m/z 381.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.45 (s, 1H), 9.27 (s, 1H), 9.05 (s, 1H), 7.98 (d, J=8.54 Hz, 1H), 7.92 (s, 1H), 7.68 - 7.70 (m, 1H), 7.58 (dd, J=8.54, 2.14 Hz, 1H), 4.76 - 4.86 (m, 2H) 3.44 – 3.53 (m, 3H), 3.39 - 3.42 (m, 1H), 3.24 - 3.29 (m, 1H), 2.38 (s, 3H), 2.02 - 2.27 (m, 1H), 1.06 - 1.09 (m, 6H).
11	MS m/z 365.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.08 (s, 1H), 7.99 (d, J=9.2 Hz, 1H), 7.96 (s, 2H), 7.67-7.74 (m, 2H), 4.82 (br d, J=13.5 Hz, 1H), 4.71 (d, J=13.6 Hz, 1H), 3.11-3.23 (m, 2H), 3.01 (dd, J=12.9, 10.2 Hz, 1H), 2.82 (td, J=12.9, 3.5 Hz, 1H), 1.96 (td, J=10.2, 3.5 Hz, 1H), 0.83-0.96 (m, 1H), 0.58-0.65 (m, 2H), 0.31-0.42 (m, 2H); 2Hs not observed (NH and OH).
14	MS m/z 383.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.09 (s, 1H), 8.00 (d, J=9.0 Hz, 1H), 7.96 (s, 2H), 7.70-7.75 (m, 2H), 4.95 (d, J=13.5 Hz, 1H), 4.79 (d, J=13.5 Hz, 1H), 3.17-3.27 (m, 1H), 3.11 (td, J=13.2, 3.7 Hz, 1H), 2.86-2.96 (m, 2H), 2.64-2.75 (m, 1H), 1.32 (d, J=13.0 Hz, 6H); 3Hs not observed (NH and 2OHs).
18	MS m/z 381.5 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.06 (s, 1H), 7.89-8.03 (m, 3H), 7.66 (s, 2H), 4.91 (d, J=12.7 Hz, 1H), 4.74 (br d, J=12.7 Hz, 1H), 3.57-3.80 (m, 1H), 3.00-3.22 (m, 2H), 2.85 (br t, J=11.1 Hz, 1H), 2.45 (d, J=11.1 Hz, 1H), 1.06 (s, 9H); 2Hs not observed (NH and OH).
132	MS m/z 353.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.09 (s, 1H), 7.99 (d, J=9.0 Hz, 1H), 7.96 (s, 2H), 7.69-7.74 (m, 2H), 4.81 (br d, J=12.8 Hz, 1H), 4.74 (d, J=12.9 Hz, 1H), 3.16-3.25 (m, 2H), 2.90-2.97 (m, 1H), 2.83-2.89 (m, 1H), 2.72-2.81 (m, 1H), 1.59 (quin, J=7.4 Hz, 2H), 1.08 (t, J=7.4 Hz, 3H); 2Hs not observed (NH and OH).
153	MS m/z 365.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.09 (s, 1H), 7.98-8.02 (m, 1H), 7.96 (s, 2H), 7.68-7.75 (m, 2H), 4.93 (d, J=11.1 Hz, 1H), 4.84 (d, J=14.3 Hz, 1H), 3.14-3.28 (m, 3H), 2.85-2.93 (m, 1H), 2.29 (d, J=9.0 Hz, 2H), 2.13-2.23 (m, 1H), 1.83-2.05 (m, 3H), 1.51-1.63 (m, 1H); 1H not observed (OH).
154	MS m/z 367.3 $[M+H]^+$; ^1H NMR (500 MHz, methanol-d_4) δ: 9.17 (s, 1H), 7.99-8.05 (m, 1H), 7.97 (s, 2H), 7.73-7.76 (m, 2H), 5.04 (d, J=14.6 Hz, 1H), 4.94 (d, J=11.4 Hz, 1H), 3.50 (d, J=12.2 Hz, 1H), 3.46 (br d, J=9.6 Hz, 1H), 3.21-3.29 (m, 2H), 3.12-3.20 (m, 1H), 1.96-2.08 (m, 1H), 1.18 (d, J=6.9 Hz, 6H); 2Hs not observed (NH and OH).

Example 4

Preparation of Compound 180



Step 1: To a flask containing a stirbar 3-(methylsulfonyl)-1,2,4-triazine (3.3 g, 21.0 mmol, 1.0 equiv.) and the HCl salt of (*S*)-2-cyclopropylpiperazine (5.0 g, 25.1 mmol, 1.2 equiv.) was added dmf (100 mL) followed by DIPEA (11 mL, 63.0 mmol, 3.0 equiv.) and was allowed to stir for 12 h at 22 ˚C. After, the solvent was removed and the crude oil was purified by silica gel chromatography using a gradient from 100% CH_2Cl_2 up to 20% MeOH/CH_2Cl_2 to afford (*S*)-3-(3-cyclopropylpiperazin-1-yl)-1,2,4-triazine (4.1 g, 20.1 mmol, 96% yield) as a brown solid. MS m/z 206.3 $[M+H]^+$.

Step 2: To a flask containing (*S*)-3-(3-cyclopropylpiperazin-1-yl)-1,2,4-triazine (4.1 g, 20.1 mmol, 1.0 equiv.) was added Boc_2O (6.0 g, 27.0 mmol, 1.3 equiv.), NEt_3 (8.6 mL, 68.0 mmol, 3.0 equiv.) and CH_2Cl_2 (100 mL). Next, DMAP (0.5 g, 4.0 mmol, 0.2 equiv.) was added and the mixture was allowed to stir at 22 ˚C for 12 h. The solvent was removed and the crude oil was purified by silica gel chromatography using a gradient from 100% hexanes up to 50% EtOAc/hexanes to afford *tert*-butyl (*S*)-2-cyclopropyl-4-(1,2,4-triazin-3-yl)piperazine-1-carboxylate (3.4 g, 11.3 mmol, 55% yield) as an oil. MS m/z 306.3 $[M+H]^+$.

Step 3: To a flask containing *tert*-butyl (*S*)-2-cyclopropyl-4-(1,2,4-triazin-3-yl)piperazine-1-carboxylate (3.4 g, 11.3 mmol, 1.0 equiv.) was added acetonitrile (90 mL) and water (30 mL) and was added NBS (2.4 g, 13.0 mmol, 1.2 equiv.) and the mixture was allowed to stir for 12 h at 22 ˚C. Next, the mixture was diluted with EtOAc and water. The organic layer was separated, dried over $MgSO_4$, filtered and concentrated. Purified by silica

gel chromatography using a gradient from 100% hexanes up to 50% EtOAc/hexanes to afford *tert*-butyl (*S*)-4-(6-bromo-1,2,4-triazin-3-yl)-2-cyclopropylpiperazine-1-carboxylate (2.9 g, 7.5 mmol, 67% yield) as an orange oil. MS m/z 384.1, 386.1 $[M+H]^+$.

Step 4: To a flask containing *tert*-butyl (*S*)-4-(6-bromo-1,2,4-triazin-3-yl)-2-cyclopropylpiperazine-1-carboxylate (1.0 g, 2.6 mmol, 1.0 equiv.), tributyl(4-chloro-2-(methoxymethoxy)phenyl)stannane (1.4 g, 3.0 mmol, 1.2 equiv.), $PdCl_2(PPh_3)_2$ (0.2 g, 0.3 mmol, 0.1 equiv.), CuI (0.1 g, 0.5 mmol, 0.2 equiv.) was added dioxane (20 mL) and purged with Ar. The flask was allowed to stir for 2 h at 100 °C under Ar. Once cooled, the mixture was filtered and the filtrate was concentrated and purified by silica gel chromatography using a gradient from 100% hexanes up to 50% EtOAc/hexanes to afford *tert*-butyl (*S*)-4-(6-(5-chloro-3-(methoxymethoxy)pyridin-2-yl)-1,2,4-triazin-3-yl)-2-cyclopropylpiperazine-1-carboxylate (0.4 g, 0.9 mmol, 34% yield) as a yellow oil. MS m/z 477.1, 479.1 $[M+H]^+$.

Step 5: To a vial containing *tert*-butyl (*S*)-4-(6-(5-chloro-3-(methoxymethoxy)pyridin-2-yl)-1,2,4-triazin-3-yl)-2-cyclopropylpiperazine-1-carboxylate (0.05 g, 0.1 mmol, 1.0 equiv.) was added (2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)boronic acid (0.04 g, 0.2 mmol, 1.5 equiv.), K_2CO_3 (0.05 g, 0.4 mmol, 3.0 equiv.), XPhos Pd G_3 (0.01 g, 0.01 mmol, 0.1 equiv.) dioxane (1 mL) and water (1 mL). The mixture was allowed to stir at 100 °C for 1 h. Once cooled, the mixture was purified by silica gel chromatography using a gradient from 100% hexanes to 50% EtOAc/hexanes to afford *tert*-butyl (*S*)-2-cyclopropyl-4-(6-(3-(methoxymethoxy)-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)pyridin-2-yl)-1,2,4-triazin-3-yl)piperazine-1-carboxylate (0.02 g, 0.03 mmol, 28% yield) as a yellow solid. MS m/z 475.4 $[M+H]^+$.

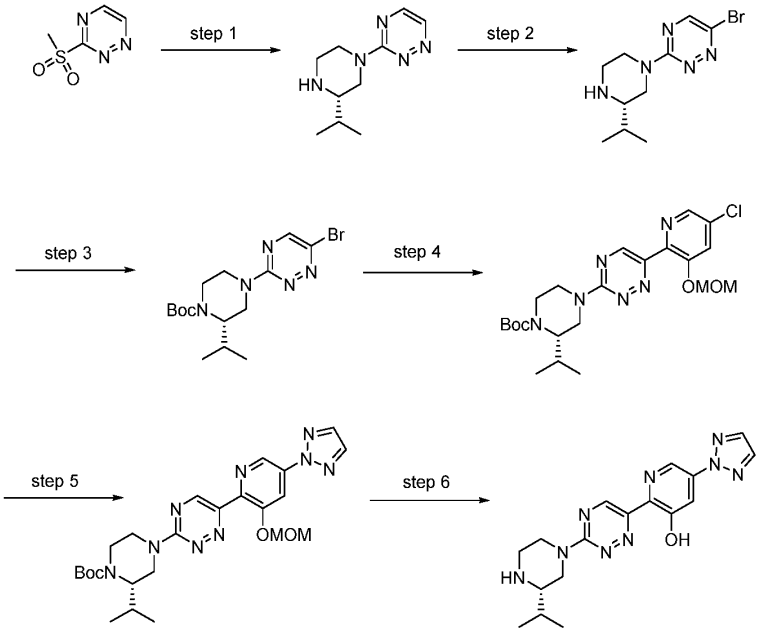
Step 6: To a vial containing *tert*-butyl (*S*)-2-cyclopropyl-4-(6-(3-(methoxymethoxy)-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)pyridin-2-yl)-1,2,4-triazin-3-yl)piperazine-1-carboxylate (0.02 g, 0.03 mmol, 1.0 equiv.) was dissolved in MeOH (1 mL) and was added 4.0 M HCl/dioxane. The mixture was allowed to stir for 1 h at 22 °C. Next, the mixture was concentrated and purified by silica gel chromatography using a gradient from 100% CH_2Cl_2 up to 15% MeOH/CH_2Cl_2 to afford the HCl salt of (*S*)-2-(3-(3-cyclopropylpiperazin-1-yl)-1,2,4-triazin-6-yl)-5-(2-methyl-2H-[1,2,3]triazolo[4,5-b]pyridin-6-yl)pyridin-3-ol (0.01 g, 0.02 mmol, 58% yield) as a white solid upon drying. MS m/z 431.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 12.02 (s, 1H), 9.45 (s, 1H), 9.25 (s, 1H), 9.16 (s, 1H), 8.90 (s, 1H), 8.78 (s, 1H), 7.99 (s, 1H), 4.77 (d, J = 14.0 Hz, 1H), 4.72 (d, J = 15.0 Hz, 1H), 4.60 (s, 3H), 3.54-3.46 (m, 3H), 3.11 (t, J = 11.5 Hz, 1H), 2.70 (t, J = 9.0 Hz, 1H), 1.09-1.02 (m, 1H), 0.72-0.64 (m, 2H), 0.57-0.54 (m, 1H), 0.46-0.43 (m, 1H).

Using the procedure described for Example 4, above, additional compounds described herein may be prepared by substituting the appropriate starting material, suitable reagents and reaction conditions, obtaining compounds such as those selected from:

Cpd	Data
181	MS m/z 447.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 12.02 (s, 1H), 11.13 (s, 1H), 9.36 (s, 1H), 8.65 (d, J = 2.0 Hz, 1H), 8.58 (d, J = 2.0 Hz, 1H), 8.03 (d, J = 1.0 Hz, 1H), 7.75 (d, J = 2.0 Hz, 1H), 7.55 (d, J = 14.5 Hz, 1H), 4.68 (d, J = 12.0 Hz, 1H), 4.60 (d, J = 12.5 Hz, 1H), 4.23 (s, 3H), 3.25-3.08 (m, 3H), 2.79 (t, J = 11.5 Hz, 1H), 2.23-2.12 (m, 1H), 0.93-0.85 (m, 1H), 0.56-0.48 (m, 2H), 0.39-0.36 (m, 1H), 0.33-0.30 (m, 1H).
157	MS m/z 449.5 $[M+H]^+$; ^1H (500 MHz, DMSO-d_6) δ: 12.07 (s, 1H), 9.34 (s, 1H), 8.64 (d, J = 2.0 Hz, 1H), 8.58 (d, J = 2.0 Hz, 1H), 8.03 (s, 1H), 7.77 (d, J = 2.0 Hz, 1H), 7.55 (d, J = 14.0 Hz, 1H), 4.67-4.55 (m, 2H), 4.23 (s, 3H), 3.05 (t, J = 11.5 Hz, 2H), 2.81 (t, J = 11.5 Hz, 1H), 2.70 (t, J = 12.0 Hz, 1H), 2.43-2.36 (m, 1H), 1.70-1.63 (m, 1H), 0.97 (d, J = 7.0 Hz, 6H); 1H not observed (NH or OH).
121	MS m/z 433.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 12.04 (s, 1H), 9.35 (s, 1H), 8.65 (s, 1H), 8.38 (s, 1H), 7.77 (s, 1H), 7.69 (s, 1H), 6.98 (s, 1H), 4.17 (s, 3H), 4.02 (s, 3H), 3.90 (t, J = 5.0 Hz, 4H), 2.45 (t, J = 5.0 Hz, 4H), 2.25 (s, 3H).
122	MS m/z 421.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 11.55 (s, 1H), 9.41 (s, 1H), 8.98 (s, 1H), 8.66 (s, 1H), 7.85 (s, 1H), 7.84 (d, J = 12.0 Hz, 1H), 7.82 (s, 1H), 4.80 (m, 4H), 3.16 (s, 3H), 2.78 (m, 4H), 2.39 (s, 3H).
124	MS m/z 339.2 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 13.21 (s, 1H), 12.00 (s, 1H), 9.31 (s, 1H), 8.57 (s, 1H), 8.41 (s, 1H), 8.10 (s, 1H), 7.67 (s, 1H), 3.88 (s, 2H), 3.30 (s, 2H), 2.49 (s, 3H), 2.45 (s, 2H), 2.24 (s, 2H).
126	MS m/z 417.4 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 12.04 (s, 1H), 11.59 (s, 1H), 9.42 (s, 1H), 9.20 (s, 1H), 8.69 (s, 1H), 7.99 (s, 1H), 7.87 (s, 1H), 4.87-4.76 (m, 2H), 3.66-3.48 (m, 4H), 3.22-3.10 (m, 2H), 2.79 (s, 3H), 2.64 (s, 3H), 2.52 (s, 3H).

Example 5

Preparation of Compound 175



Step 1: To a solution of 3-(methylsulfonyl)-1,2,4-triazine (3.0 g, 18.8 mmol) and the HCl salt of (*S*)-2-isopropylpiperazine (4.5 g, 22.6 mmol) in DMF (100 mL) was added DIPEA (33 mL, 187 mmol). The reaction mixture was stirred at rt for 12 h until UPLC showed complete conversion. The solvent was removed, and the crude oil was purified by silica gel chromatography eluting with a gradient CH_2Cl_2/MeOH (0-20% MeOH) to afford (*S*)-3-(3-isopropylpiperazin-1-yl)-1,2,4-triazine (1.8 g, 46% yield) as a brown solid. MS m/z 208.3 $[M+H]^+$.

Step 2: To a solution of (*S*)-3-(3-isopropylpiperazin-1-yl)-1,2,4-triazine (1.8 g, 8.7 mmol) in water (100 mL) and MeOH (25 mL) was added bromine (0.7 mL, 13.0 mmol) dropwise and the mixture was allowed to stir for 1 h. Saturated aqueous sodium thiosulfate was added followed by water and EtOAc. The aqueous layer was then extracted with EtOAc 3 times. The organic layer was washed with water, brine and dried over $MgSO_4$, filtered and concentrated. The crude oil was purified by silica gel chromatography eluting with a gradient CH_2Cl_2/MeOH (0-10% MeOH) to afford (*S*)-6-bromo-3-(3-isopropylpiperazin-1-yl)-1,2,4-triazine (1.7 g, 69% yield) as a brown solid. MS m/z 286.1, 288.1 $[M+H]^+$.

Step 3: To a flask containing (*S*)-6-bromo-3-(3-isopropylpiperazin-1-yl)-1,2,4-triazine (1.7 g, 6.0 mmol), Et_3N (2.5 mL, 18.0 mmol) and Boc_2O (1.7 g, 7.8 mmol) was added CH_2Cl_2 (100 mL). DMAP (0.2 g, 1.2 mmol) was added and the mixture was allowed to stir at rt for 12

h. The solvent was removed, and the crude oil was purified by silica gel chromatography eluting with a gradient hexanes/EtOAc (0-100% EtOAc) to afford *tert*-butyl (*S*)-4-(6-bromo-1,2,4-triazin-3-yl)-2-isopropylpiperazine-1-carboxylate (2.1 g, 90% yield) as a white solid. MS m/z 386.1,388.1 $[M+H]^+$.

Step 4: To a vial containing *tert*-butyl (*S*)-4-(6-bromo-1,2,4-triazin-3-yl)-2-isopropylpiperazine-1-carboxylate (1.3 g, 3.4 mmol) were added tributyl(4-chloro-2-(methoxymethoxy)phenyl)stannane (1.9 g, 4.1 mmol), CuI (0.1 g, 0.7 mmol) and $PdCl_2(PPh_3)_2$ (0.2 g, 0.3 mmol) followed by dioxane (10 mL). The mixture was purged with Ar and stirred at 100 ˚C for 2h. The crude mixture was filtered and concentrated. The residual oil was purified by silica gel chromatography eluting with a gradient hexanes/EtOAc (0-100% EtOAc) to afford *tert*-butyl (*S*)-4-(6-(5-chloro-3-(methoxymethoxy)pyridin-2-yl)-1,2,4-triazin-3-yl)-2-isopropylpiperazine-1-carboxylate (0.8 g, 51% yield) as a colorless oil. MS m/z 479.2, 481.2 $[M+H]^+$.

Step 5: To a vial containing tBuXPhos (0.01 g, 0.01 mmol), $Pd_2(dba)_3$ (0.01 g, 0.01 mmol) was added toluene (1 mL) and the mixture was purged with Ar. The reaction was heated to 110 ˚C for 10 min, then cooled to rt. The catalyst solution was added to a vial containing *tert*-butyl (*S*)-4-(6-(5-chloro-3-(methoxymethoxy)pyridin-2-yl)-1,2,4-triazin-3-yl)-2-isopropylpiperazine-1-carboxylate (0.06 g, 0.1 mmol), triazole (0.01 g, 0.15 mmol), K_3PO_4 (0.05 g, 0.2 mmol) in toluene (5 mL). The reaction mixture was allowed to stir at 110 ˚C for 4 h. The crude mixture was filtered, concentrated and purified eluting with a gradient hexanes/EtOAc (0-50% EtOAc) to afford *tert*-butyl (*S*)-2-isopropyl-4-(6-(3-(methoxymethoxy)-5-(2H-1,2,3-triazol-2-yl)pyridin-2-yl)-1,2,4-triazin-3-yl)piperazine-1-carboxylate (0.04 g, 72% yield) as a colorless oil. MS m/z 512.5 $[M+H]^+$

Step 6: To a vial containing *tert*-butyl (*S*)-2-isopropyl-4-(6-(3-(methoxymethoxy)-5-(2H-1,2,3-triazol-2-yl)pyridin-2-yl)-1,2,4-triazin-3-yl)piperazine-1-carboxylate (0.04 g, 0.08 mmol) and a stirbar was added MeOH (2 mL). 4.0 M HCl/dioxane (2 mL) was added and the mixture was stirred at rt for 1 h. The solvent was removed and the crude solid was purified by silica gel chromatography using a gradient CH_2Cl_2/MeOH (0-10% MeOH) to afford (*S*)-2-(3-(3-isopropylpiperazin-1-yl)-1,2,4-triazin-6-yl)-5-(2H-1,2,3-triazol-2-yl)pyridin-3-ol (0.02 g, 73% yield) as a white solid upon drying. MS *m/z* 368.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 9.26 (s, 1H), 8.90 (d, J = 2.0 Hz, 1H), 8.24 (s, 2H), 7.93 (d, J = 2.0 Hz, 1H), 4.65 (d, J = 11.0 Hz, 1H), 4.57 (d, J = 11.0 Hz, 1H), 3.05 (t, J = 11.0 Hz, 2H), 2.82 (t, J = 11.5 Hz, 1H), 2.70 (t, J = 10.5 Hz, 1H), 2.39-2.36 (m, 1H), 1.71-1.60 (m, 1H), 0.97 (d, J = 6.5 Hz, 6H); 2Hs not observed (NH and OH).

Using the procedure described for Example 5, above, additional compounds described herein may be prepared by substituting the appropriate starting material, suitable reagents and reaction conditions, obtaining compounds such as:

Cpd	Data
182	MS m/z 366.3 $[M+H]^+$; ^1H NMR (500 MHz, DMSO-d_6) δ: 12.14 (s, 1H), 9.34 (s, 1H), 9.09 (s, 2H), 8.94 (s, 1H), 8.26 (s, 2H), 7.97 (s, 1H), 4.77 (d, J = 14.5 Hz, 1H), 4.71 (d, J = 14.0 Hz, 1H), 3.53-3.44 (m, 3H), 3.17-3.08 (m, 1H), 2.74-2.68 (m, 1H), 1.06-1.00 (m, 1H), 0.72-0.63 (m, 2H), 0.57-0.53 (m, 1H), 0.47-0.42 (m, 1H); 1H from HCl salt.

BIOLOGICAL EXAMPLES

The following *in vitro* biological examples demonstrate the usefulness of the compounds of the present description for treating Huntington's disease.

To describe in more detail and assist in understanding the present description, the following non-limiting biological examples are offered to more fully illustrate the scope of the description and are not to be construed as specifically limiting the scope thereof. Such variations of the present description that may be now known or later developed, which would be within the purview of one skilled in the art to ascertain, are considered to fall within the scope of the present description and as hereinafter claimed.

Compounds of Formula (I) were tested using the Meso Scale Discovery (MSD) Assay provided in International Application No. PCT/US2016/066042, filed on December 11, 2016 and claiming priority to United States Provisional Application U.S. 62/265,652 filed on December 10, 2015, the entire contents of which are incorporated herein by reference.

The Endogenous Huntingtin Protein assay used in Biological Example 1 was developed using the ELISA-based MSD electrochemiluminescence assay platform.

Biological Example 1

Endogenous Huntingtin Protein Assay

Meso Scale Discovery (MSD) 96-well or 384-well plates were coated overnight at 4°C with MW1 (expanded polyglutamine) or MAB2166 monoclonal antibody (for capture) at a concentration of 1 µg/mL in PBS (30 µL per well). Plates were then washed three times with 300 µL wash buffer (0.05% Tween-20 in PBS) and blocked (100 µL blocking buffer; 5% BSA in PBS) for 4-5 hours at room temperature with rotational shaking and then washed three times with wash buffer.

Samples (25 µL) were transferred to the antibody-coated MSD plate and incubated overnight at 4°C. After removal of the lysates, the plate was washed three times with wash buffer, and 25 µL of #5656S (Cell signaling; rabbit monoclonal) secondary antibody (diluted to 0.25 µg/mL in 0.05% Tween-20 in blocking buffer) was added to each well and incubated with shaking for 1Hour at room temperature. Following incubation with the secondary antibody, the wells were rinsed with wash buffer after which 25 µL of goat anti-rabbit SULFO TAG secondary detection antibody (required aspect of the MSD system) (diluted to 0.25 µg/mL in 0.05% Tween-20 in blocking buffer) was added to each well and incubated with shaking for 1 hour at room temperature. After rinsing three times with wash buffer, 150 µL of read buffer T with surfactant (MSD) were added to each empty well, and the plate was imaged on a SI 6000 imager (MSD) according to manufacturers' instructions provided for 96- or 384-well plates. The resulting IC_{50} values (nM) for compounds tested are provided in Tables 2 and 3.

Table 2. IC_{50} (nM) Values for Compunds 1-284

Cpd	IC50 (nM)	Cpd	IC50 (nM)	Cpd	IC50 (nM)
1	2.1	**95**	8.3	**189**	461.2
2	3.9	**96**	8.6	**190**	166.4
3	4.4	**97**	9.8	**191**	621.0
4	4.5	**98**	10.2	**192**	34.2
5	6	**99**	10.5	**193**	4.8
6	6.7	**100**	10.9	**194**	19.3
7	6.9	**101**	13.0	**195**	9.1
8	8.7	**102**	13.4	**196**	491.5
9	9.5	**103**	13.4	**197**	11.6
10	9.5	**104**	14.6	**198**	12.5
11	9.6	**105**	15.3	**199**	525.7
12	10.9	**106**	16.5	**200**	30.1
13	11.4	**107**	16.8	**201**	11.1
14	12.2	**108**	19.8	**202**	3.7
15	12.8	**109**	22.1	**203**	42.5
16	13.2	**110**	24.9	**204**	162.5
17	14.1	**111**	27.5	**205**	903.6
18	15.1	**112**	34.0	**206**	33.7
19	17.1	**113**	36.0	**207**	10.0
20	17.1	**114**	54.4	**210**	975.4

21	20.8
22	22.3
23	22.7
24	23.4
25	24.6
26	24.9
27	25.1
28	26.2
29	28.5
30	33.3
31	34.3
32	40.2
33	41.4
34	42
35	46.1
36	47.7
37	50.7
38	56.2
39	63.7
40	68.2
41	109.8
42	110.9
43	116.2
44	137.7
45	141.5
46	150.3
47	154.5
48	166.9
49	256.1
50	312.2
51	329.9
52	335.5
53	435.1
54	439.8
55	547.9
56	654.2
57	668.3
58	678.1

115	179.5
116	678.5
117	inactive
118	inactive
119	inactive
120	inactive
121	16.1
122	41.2
123	11.2
124	608.6
125	7.3
126	100.0
127	672.3
128	26.7
129	18.6
130	12.0
131	7.6
132	38.9
133	13.0
134	14.9
135	184.7
136	50.5
137	7.5
138	19.9
139	18.0
140	5.8
141	2.3
142	60.3
143	634.4
144	466.3
145	479.5
146	538.8
147	38.8
148	53.4
149	90.7
150	138.7
151	204.4
152	15.0

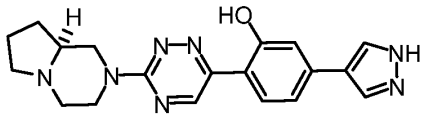
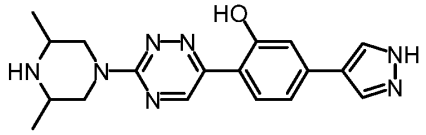
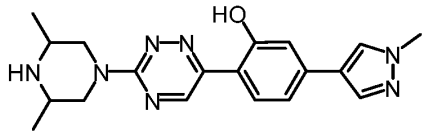
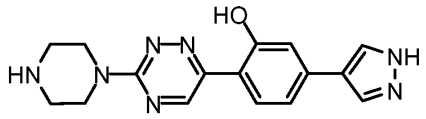
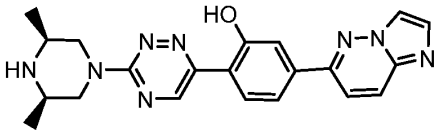
212	3.3
213	7.7
214	36.6
215	113.5
216	218.4
217	67.9
218	73.8
219	13.7
220	88.3
221	4.4
222	9.4
223	7.0
224	232.1
225	13.4
226	130.1
227	37.3
228	73.1
229	18.2
230	11.3
231	21.9
232	147.1
233	17.7
234	8.7
235	8.1
236	9.5
237	103.5
238	55.1
239	12.4
240	171.1
241	5.7
242	106.0
243	218.0
244	381.5
245	11.5
246	28.4
247	201.2
248	74.9
249	318.9

59	875.6
60	1007.2
61	1437.5
62	2390.1
63	inactive
64	inactive
65	1.6
66	3.7
67	2.1
68	2.2
69	3.6
70	3.8
71	4.0
72	4.1
73	4.1
74	4.3
75	4.3
76	4.3
77	4.9
78	5.1
79	5.2
80	5.3
81	5.4
82	5.5
83	5.6
83	5.6
84	5.9
85	6.0
86	6.0
87	6.4
88	6.5
89	6.7
90	6.8
91	7.3
92	7.9
94	8.3

153	438.4
154	18.7
155	19.5
156	12.5
157	7.8
158	5.6
159	62.2
160	788.6
161	60.5
162	25.4
163	4.2
164	4.0
165	123.6
166	191.7
167	14.2
168	1.7
169	354.7
170	21.5
171	14.1
172	4.0
173	21.3
174	607.3
175	235.6
176	619.2
177	29.0
178	299.9
179	113.2
180	27.6
181	6.2
182	173.1
183	18.8
184	49.0
185	158.1
186	77.7
187	401.2
188	105.4

250	140.0
251	18.6
252	16.5
253	375.5
254	555.6
255	340.7
256	12.4
257	19.8
258	10.8
259	337.8
260	417.3
261	992.7
262	94.2
263	214.1
264	125.1
265	291.7
266	19.4
267	145.0
268	124.5
269	53.9
270	29.2
271	943.4
272	356.5
273	645.7
274	291.0
275	755.2
276	3.7
277	24.8
278	8.3
279	5.2
280	4.5
281	198.2
282	131.0
283	7.3
284	91.4

Table 3. IC_{50} (nM) Values for Selected Compounds from WO/2019/191229 A1

WO/2019/191229 A1	IC_{50} (nM)
Cpd 116 	91.4
Cpd 262 	72.8
Cpd 263 	352.1
Cpd 328 	106.3
Cpd 385 	33.1

In Table 2, where the compounds are reported as "inactive," the results were above the detection limit of the assay, and the compounds are considered to be inactive.

Without regard to whether a document cited herein was specifically and individually indicated as being incorporated by reference, all documents referred to herein are incorporated by reference into the present application for any and all purposes to the same extent as if each individual reference was fully set forth herein.

Having now fully described the subject matter of the claims, it will be understood by those having ordinary skill in the art that the same can be performed within a wide range of equivalents without affecting the scope of the subject matter or particular aspects described herein. It is intended that the appended claims be interpreted to include all such equivalents.