ANTIBACTERIAL COMPOUNDS

RELATED APPLICATIONS

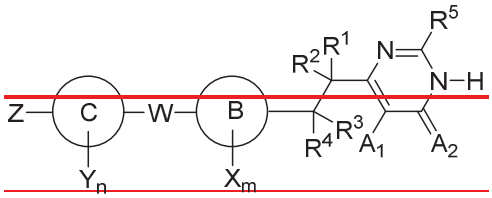
[0001] This application claims the benefit of U.S. Provisional Patent Application No. 62/734,173 filed on September 20, 2018, and U.S. Provisional Patent Application No. 62/767,313 filed on November 14, 2018, each of which is herein incoporated by reference in its entirety.

BACKGROUND

[0002] A need exists in the medicinal arts for the effective treatment of illness caused by bacterial infection.

SUMMARY

[0002a] In one aspect, the present disclosure refers to a compound, or a pharmaceutically acceptable salt, or solvate~~, or prodrug~~ thereof, having the structure of Formula (II):



~~Formula (I)~~

~~wherein,~~

~~n is 0-4;~~

~~m is 0-4;~~

~~A_1 is OH or SH;~~

~~A_2 is O or S;~~

~~R^1 and R^2 are each independently H or optionally substituted alkyl;~~

~~or R^1 and R^2 are taken together with the carbon to which they are attached to form $=C(R^{11})_2$, $=NR^{11}$, =O, or =S;~~

~~or R^1 and R^2 are taken together with the carbon to which they are attached to form an optionally substituted 3- to 6-membered carbocyclyl or optionally substituted 4- to 7-membered heterocyclyl containing 1 or 2 heteroatoms selected from O, N, and S;~~

~~R^3 is optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted aryl, optionally substituted aralkyl, optionally substituted heteroaryl, optionally substituted heteroaralkyl, optionally substituted heterocyclyl, optionally substituted heterocyclylalkyl, optionally substituted (C_0-C_4 alkylene) COR^{11}, optionally substituted (C_0-C_4 alkylene) CO_2R^{11}, optionally substituted (C_0-C_4 alkylene) $CON(R^{11})_2$, optionally substituted (C_0-C_4 alkylene) CN, optionally substituted (C_0-C_4 alkylene) OR^{11}, optionally substituted (C_0-C_4 alkylene) $N(R^{11})_2$, optionally substituted (C_0-C_4 alkylene) $N(R^{12})$ COR^{11}, optionally substituted (C_0-C_4 alkylene) $N(R^{12})$ CO_2R^{11}, optionally substituted (C_0-C_4 alkylene) $N(R^{12})$ $CON(R^{11})_2$, optionally substituted (C_0-C_4 alkylene) $N(R^{12})$ $SO_2N(R^{11})_2$, optionally substituted (C_0-C_4 alkylene) O-~~

~~$SO_2N(R^{11})_2$, optionally substituted (C_0-C_4 alkylene) $N(R^{11})$ PO(optionally substituted C_1-C_4 alkyl)$_2$, optionally substituted (C_0-C_4 alkylene) SO_2R^{11}, optionally substituted (C_0-C_4 alkylene) O SO_2R^{11}, optionally substituted (C_0-C_4 alkylene) $N(R^{12})$ SO_2R^{11}, optionally substituted (C_0-C_4 alkylene) $C(=N$-$OR^{11})(R^{11})$, or optionally substituted (C_0-C_4 alkylene) $OP(=O)(OR^{11})_2$;~~

~~R^4 is H or optionally substituted alkyl;~~

~~or R^3 and R^4 are taken together with the carbon to which they are attached to form $=C(R^{11})_2$, $=NR^{11}$, =O, or =S;~~

~~or R^3 and R^4 are taken together with the carbon to which they are attached to form an optionally substituted 3- to 6-membered carbocyclyl or optionally substituted 4- to 7-membered heterocyclyl containing 1 or 2 heteroatoms selected from O, N, and S;~~

~~R^5 is H, halogen, optionally substituted alkyl, hydroxyl, alkoxyl, cyano, amino, or nitro;~~

~~Ring B is aryl, carbocyclyl, heteroaryl, or heterocyclyl;~~

~~W is a bond, $-C\equiv C-$, bicyclo[1.1.1]pentanylene, $-C\equiv C-C\equiv C-$, $-CH=CH-$, or $-CH_2CH_2-$;~~

~~Ring C is aryl, carbocyclyl, heteroaryl, or heterocyclyl;~~

~~each X and Y is independently H, optionally substituted alkyl, halo, fluoroalkyl, cyano, nitro, $N(R^{13})_2$, or $-OR^{13}$;~~

~~or R^3 and one X are taken together with the intervening atoms to form an optionally substituted 5- to 7-membered carbocyclyl or optionally substituted 5- to 7-membered heterocyclyl containing 1 or 2 heteroatoms selected from O, N, and S;~~

~~Z is H, halo, nitro, or $-L-G$;~~

~~L is a bond or optionally substituted C_1-C_4 alkylene;~~

~~G is optionally substituted alkyl, optionally substituted alkenyl, optionally substituted alkynyl, optionally substituted carbocyclyl, optionally substituted aryl, optionally substituted heterocyclyl, optionally substituted heteroaryl, $-CN$, $-N(R^{13})_2$, $-OR^{13}$, $-COR^{13}$, $-CO_2R^{13}$, $-CON(R^{13})_2$, $-N(R^{14})$ $-COR^{13}$, $-SO_2R^{13}$, $-SO_2N(R^{13})_2$, $-N(R^{14})$ $-SO_2R^{13}$, $-N(R^{14})$ $-CON(R^{13})_2$, $-N(R^{14})$ $-CO_2R^{13}$, $-O-CON(R^{13})_2$, $-N(R^{14})$ $-SO_2N(R^{13})_2$, $-O-SO_2N(R^{13})_2$, $-N(R^{14})$ $-SO_2-OR^{13}$, or $-C(=N-OR^{14})(R^{13})$;~~

~~each R^{11} is independently H, optionally substituted alkyl, optionally substituted alkenyl, optionally substituted alkynyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted aryl, optionally substituted aralkyl, optionally substituted heterocyclyl, optionally substituted heterocyclylalkyl, optionally substituted heteroaryl, or optionally substituted heteroarylalkyl;~~

~~or two R^{11} on the same nitrogen atom are taken togehter with the nitrogen to which they are attached to form an optionally substituted N-heterocyclyl;~~

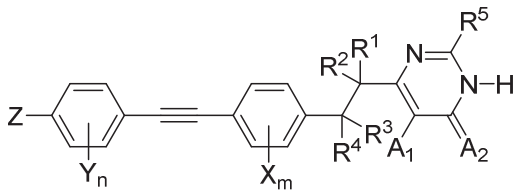
~~each R^{12} is independently H, optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted heterocyclyl, or optionally substituted heterocyclylalkyl;~~

~~each R^{13} is H, optionally substituted alkyl, optionally substituted alkenyl, optionally substituted alkynyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted aryl,~~

~~optionally substituted aralkyl, optionally substituted heterocyclyl, optionally substituted heterocyclylalkyl, optionally substituted heteroaryl, or optionally substituted heteroarylalkyl;~~

~~or two R^{13} on the same nitrogen atom are taken togehter with the nitrogen to which they are attached to form an optionally substituted N-heterocyclyl; and~~

~~each R^{14} is independently H, optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted heterocyclyl, or optionally substituted heterocyclylalkyl~~



Formula (II)

wherein,

n is 0 or 1;

m is 0 or 1;

A_1 is OH;

A_2 is O;

R^1 and R^2 are each H;

R^3 is optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted aryl, optionally substituted aralkyl, optionally substituted heteroaryl, optionally substituted heteroaralkyl, optionally substituted heterocyclyl, optionally substituted heterocyclylalkyl, optionally substituted (C_0-C_4 alkylene)-COR^{11}, optionally substituted (C_0-C_4 alkylene)-CO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-$CON(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-CN, optionally substituted (C_0-C_4 alkylene)-OR^{11}, optionally substituted (C_0-C_4 alkylene)-$N(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-COR^{11}, optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-CO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-$CON(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-$SO_2N(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-O-$SO_2N(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-$N(R^{11})$-PO(optionally substituted C_1-C_4 alkyl)$_2$, optionally substituted (C_0-C_4 alkylene)-SO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-O-SO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-SO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-$C(=N–OR^{11})(R^{11})$, or optionally substituted (C_0-C_4 alkylene)-$OP(=O)(OR^{11})_2$;

R^4 is H or optionally substituted alkyl;

R^5 is H;

each X and Y is independently H, optionally substituted alkyl, halo, fluoroalkyl, cyano, nitro, $-N(R^{13})_2$, or $-OR^{13}$;

Z is -L-G;

L is a bond or optionally substituted C_1-C_4 alkylene;

G is alkyl, carbocyclyl, aryl, heterocyclyl, heteroaryl, $-N(R^{13})_2$, $-OR^{13}$, -CN, $-COR^{13}$, $-CO_2R^{13}$, $-CON(R^{13})_2$, $-N(R^{14})-COR^{13}$, $-SO_2R^{13}$-, $-SO_2N(R^{13})_2$, or $-N(R^{14})-SO_2R^{13}$;

each R^{11} is independently H, optionally substituted alkyl, optionally substituted alkenyl, optionally substituted alkynyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted aryl, optionally substituted aralkyl, optionally substituted heterocyclyl, optionally substituted heterocyclylalkyl, optionally substituted heteroaryl, or optionally substituted heteroarylalkyl;

or two R^{11} on the same nitrogen atom are taken together with the nitrogen to which they are attached to form an optionally substituted N-heterocyclyl;

each R^{12} is independently H, optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted heterocyclyl, or optionally substituted heterocyclylalkyl;

each R^{13} is independently H, alkyl, carbocyclyl, carbocyclylalkyl, heterocyclyl, or heterocyclylalkyl; wherein each R^{13} is independently unsubstituted or substituted with $-R^b-OR^a$, $-R^b-C(O)OR^a$, or $-R^b-C(O)R^a$; wherein each R^a is independently hydrogen, alkyl which is optionally substituted with halogen, hydroxy, methoxy, or trifluoromethyl, or carbocyclyl which is optionally substituted with halogen, hydroxy, methoxy, or trifluoromethyl; and each R^b is independently a direct bond or a straight or branched alkylene;

two R^{13} groups are joined with the nitrogen to which they are attached join to form an N-heterocyclyl which is unsubstituted or substituted with alkyl, optionally substituted heterocyclyl, $-R^b-OR^a$, $-R^b-N(R^a)_2$, $-R^b-C(O)R^a$, $-R^b-CN$, or $-R^b-N(R^a)C(O)R^a$; wherein each R^a is independently hydrogen, alkyl which is optionally substituted with halogen, hydroxy, methoxy, or trifluoromethyl, or carbocyclyl which is optionally substituted with halogen, hydroxy, methoxy, or trifluoromethyl; and each R^b is independently a direct bond or a straight or branched alkylene; and

each R^{14} is independently H, unsubstituted alkyl, or unsubstituted heterocyclyl;

wherein

each carbocyclyl is a stable non-aromatic monocyclic, fused polycyclic, or bridged polycyclic hydrocarbon ring comprising from three to ten carbon atoms;

each carbocyclylalkyl is a $-R^c$-carbocyclyl, where R^c is a C_1-C_8 alkylene;

each heterocyclyl is a stable 3- to 18-membered non-aromatic monocyclic, fused bicyclic, bridged bicyclic, or spirocyclic bicyclic ring system comprising from two to twelve carbon atoms and from one to six heteroatoms selected from nitrogen, oxygen, and sulfur which is optionally oxidized; and

each heterocyclylalkyl is a $-R^c$- heterocyclyl, where R^c is a C_1-C_8 alkylene.

[0002b] In another aspect, the present disclosure refers to a compound selected from:

5-hydroxy-6-(3-hydroxy-3-methyl-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)butyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-hydroxy-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-hydroxy-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(3-((2-fluoroethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-((2-fluoroethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-((2-fluoroethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-hydroxy-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-hydroxy-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(morpholinomethyl) phenyl)ethynyl)phenyl)butyl)pyrimidin-4(3H)-one;

6-((R)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

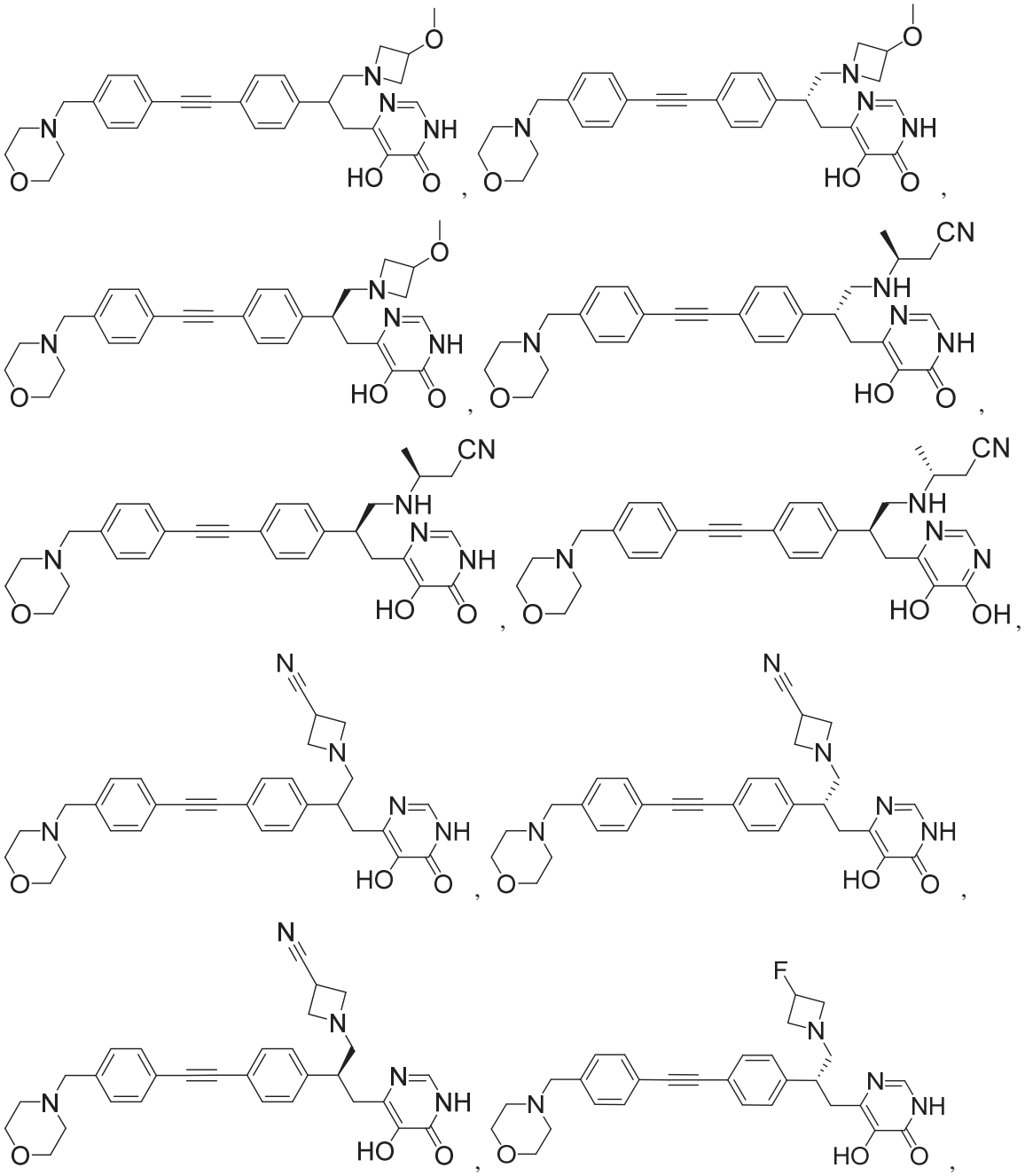
5-hydroxy-6-(3-(2-methyl-1H-imidazol-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

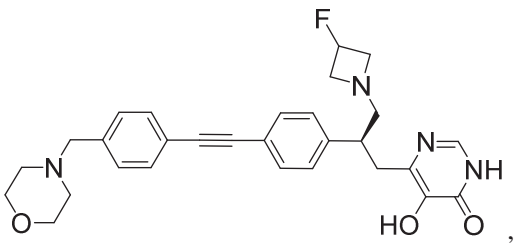
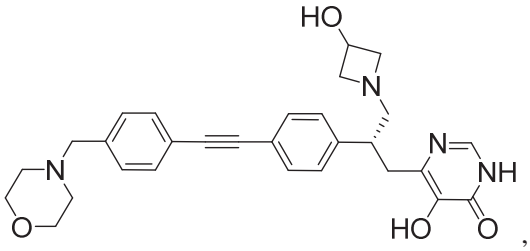
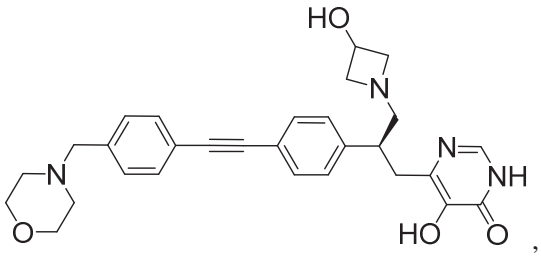
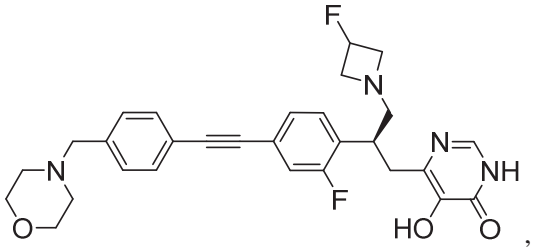
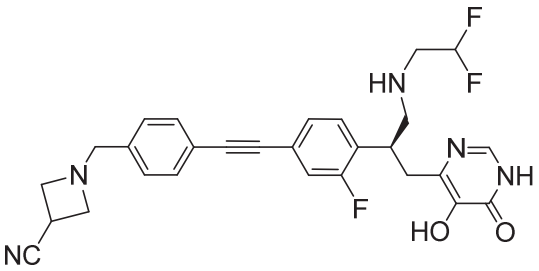
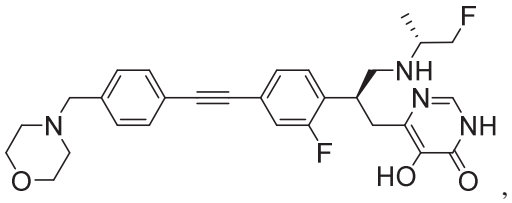
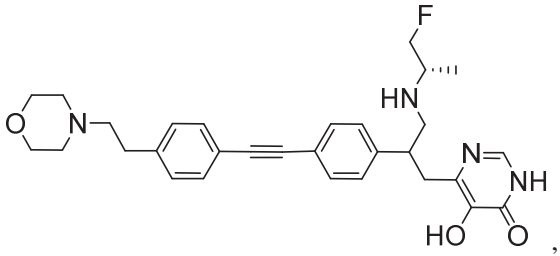
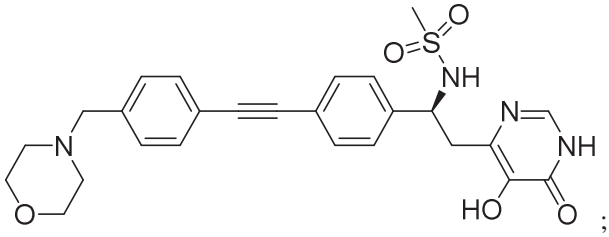
(S)-5-hydroxy-6-(3-(2-methyl-1H-imidazol-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one; and

(R)-5-hydroxy-6-(3-(2-methyl-1H-imidazol-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

or a pharmaceutically acceptable salt, or solvate, ~~or prodrug~~ thereof.

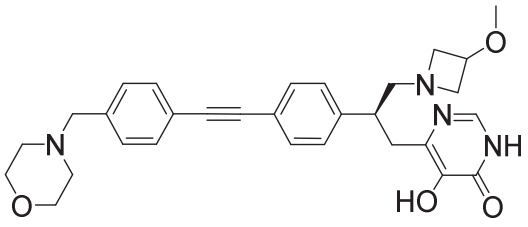
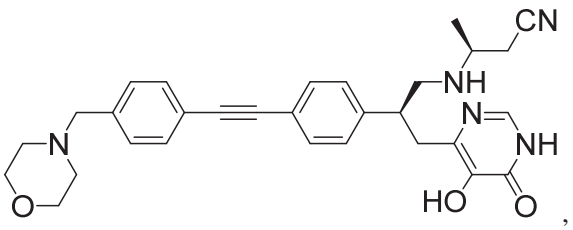
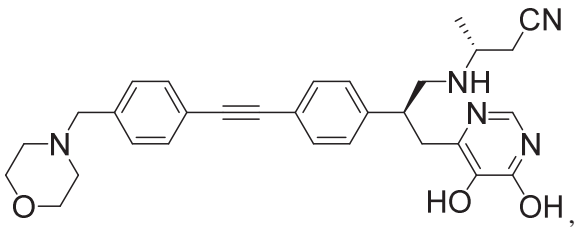
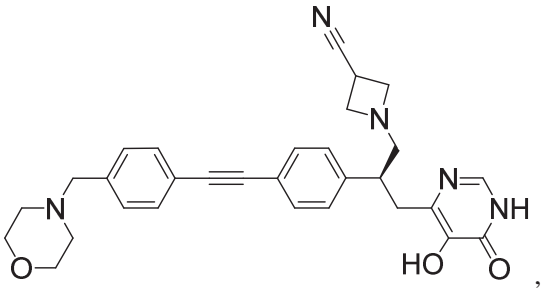
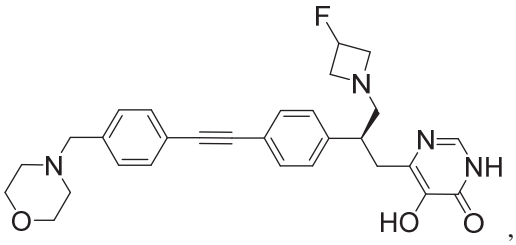
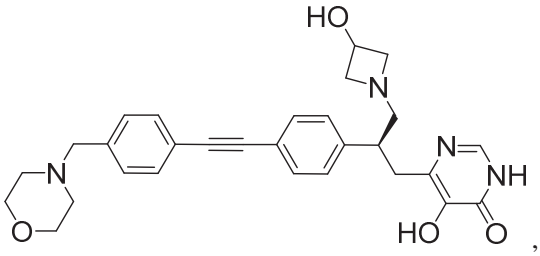
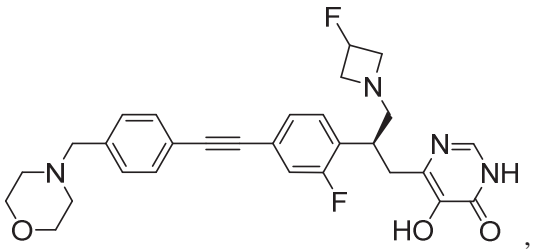
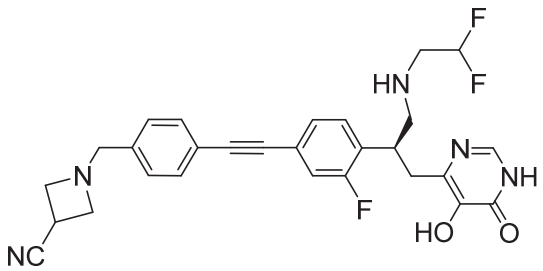
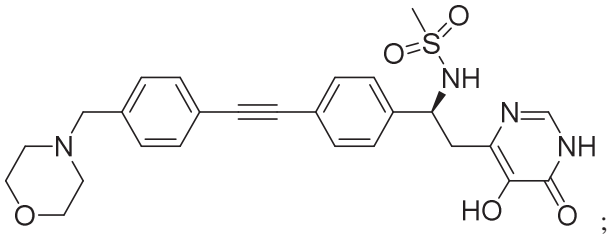
[0002c] In another aspect, the present disclosure refers to a compound selected from:




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or a pharmaceutically acceptable salt or, solvate, ~~or prodrug~~ thereof.

[0002d] In yet another aspect, the present disclosure refers to a compound selected from:


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or a pharmaceutically acceptable salt or, ~~,~~ solvate ~~, or prodrug~~ thereof.

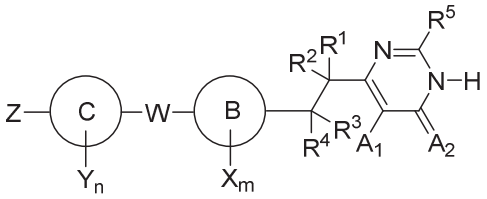
[0002e] In yet another aspect, the present disclosure refers to a pharmaceutical composition comprising a compound of any one of claims 1 - 21, or a pharmaceutically acceptable salt or, ~~,~~ solvate ~~, or prodrug~~ thereof, and a pharmaceutically acceptable excipient.

[0002f] In yet another aspect, the present disclosure refers to the use of a compound as described herein, or a pharmaceutically acceptable salt ~~,~~ or solvate ~~, or prodrug~~ thereof ~~, or the pharmaceutical composition as described herein~~, in the manufacture of a medicament for ~~treating~~ the treatment of a gram-negative bacterial infection in a patient in need thereof.

[0002g] In yet another aspect, the present disclosure refers to the compound as described herein, or a pharmaceutically acceptable salt or, solvate, or prodrug thereof, or the pharmaceutical composition as described herein, for use in therapy.

[0003] Provided herein are heterocyclic derivative compounds and pharmaceutical compositions comprising said compounds that are useful for inhibiting the growth of gram-negative bacteria. Furthermore, the subject compounds and compositions are useful for the treatment of bacterial infection, such as urinary tract infection and the like.

[0004] Provided herein is a compound, or a pharmaceutically acceptable salt, solvate, or prodrug thereof, having the structure of Formula (I):



Formula (I)

wherein,

n is 0-4;

m is 0-4;

A_1 is OH or SH;

A_2 is O or S;

R^1 and R^2 are each independently H or optionally substituted alkyl;

or R^1 and R^2 are taken together with the carbon to which they are attached to form $=C(R^{11})_2$, $=NR^{11}$, =O, or =S;

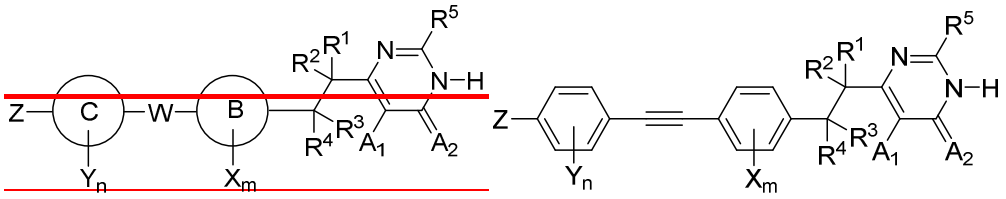
or R^1 and R^2 are taken together with the carbon to which they are attached to form an optionally substituted 3- to 6-membered carbocyclyl or optionally substituted 4- to 7-membered heterocyclyl containing 1 or 2 heteroatoms selected from O, N, and S;

R^3 is optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted aryl, optionally substituted aralkyl, optionally substituted heteroaryl, optionally substituted heteroaralkyl, optionally substituted heterocyclyl, optionally substituted heterocyclylalkyl, optionally substituted (C_0-C_4 alkylene)-COR^{11}, optionally substituted (C_0-C_4 alkylene)-CO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-$CON(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-CN, optionally

CLAIMS

We Claim:

1. A compound, or a pharmaceutically acceptable salt~~,~~ or solvate~~, or prodrug~~ thereof, having the structure of Formula (II):



Formula (II)

wherein,

n is 0~~–4~~ or 1;

m is 0~~–4~~ or 1;

A_1 is OH~~ or SH~~;

A_2 is O~~ or S~~;

R^1 and R^2 are each ~~independently~~ H~~ or optionally substituted alkyl~~;

~~or R^1 and R^2 are taken together with the carbon to which they are attached to form $=C(R^{11})_2$, $=NR^{11}$, =O, or =S;~~

~~or R^1 and R^2 are taken together with the carbon to which they are attached to form an optionally substituted 3- to 6-membered carbocyclyl or optionally substituted 4- to 7-membered heterocyclyl containing 1 or 2 heteroatoms selected from O, N, and S;~~

R^3 is optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted aryl, optionally substituted aralkyl, optionally substituted heteroaryl, optionally substituted heteroaralkyl, optionally substituted heterocyclyl, optionally substituted heterocyclylalkyl, optionally substituted (C_0-C_4 alkylene)-COR^{11}, optionally substituted (C_0-C_4 alkylene)-CO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-$CON(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-CN, optionally substituted (C_0-C_4 alkylene)-OR^{11}, optionally substituted (C_0-C_4 alkylene)-$N(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-COR^{11}, optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-CO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-$CON(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-$SO_2N(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-O-$SO_2N(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-$N(R^{11})$-PO(optionally substituted C_1-C_4 alkyl)$_2$, optionally substituted (C_0-C_4 alkylene)-SO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-O-SO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-SO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-C(=N–OR^{11})(R^{11}), or optionally substituted (C_0-C_4 alkylene)-$OP(=O)(OR^{11})_2$;

R^4 is H or optionally substituted alkyl;

~~or R^3 and R^4 are taken together with the carbon to which they are attached to form $=C(R^{11})_2$, $=NR^{11}$, =O, or =S;~~

~~or R^3 and R^4 are taken together with the carbon to which they are attached to form an optionally substituted 3- to 6-membered carbocyclyl or optionally substituted 4- to 7-membered heterocyclyl containing 1 or 2 heteroatoms selected from O, N, and S;~~

R^5 is H~~, halogen, optionally substituted alkyl, hydroxyl, alkoxyl, cyano, amino, or nitro~~;

~~Ring B is aryl, carbocyclyl, heteroaryl, or heterocyclyl;~~

~~W is a bond, C≡C , bicyclo[1.1.1]pentanylene, C≡C-C≡C , CH=CH , or CH_2CH_2 ;~~

~~Ring C is aryl, carbocyclyl, heteroaryl, or heterocyclyl;~~

each X and Y is independently H, optionally substituted alkyl, halo, fluoroalkyl, cyano, nitro, $-N(R^{13})_2$, or $-OR^{13}$;

~~or R^3 and one X are taken together with the intervening atoms to form an optionally substituted 5- to 7-membered carbocyclyl or optionally substituted 5- to 7-membered heterocyclyl containing 1 or 2 heteroatoms selected from O, N, and S;~~

Z is ~~H, halo, nitro, or~~ -L-G;

L is a bond or optionally substituted C_1-C_4 alkylene;

G is alkyl, carbocyclyl, aryl, heterocyclyl, heteroaryl, $-N(R^{13})_2$, $-OR^{13}$, -CN, $-COR^{13}$, $-CO_2R^{13}$, $-CON(R^{13})_2$, $-N(R^{14})-COR^{13}$, $-SO_2R^{13}$-, $-SO_2N(R^{13})_2$, or $-N(R^{14})-SO_2R^{13}$;

each R^{11} is independently H, optionally substituted alkyl, optionally substituted alkenyl, optionally substituted alkynyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted aryl, optionally substituted aralkyl, optionally substituted heterocyclyl, optionally substituted heterocyclylalkyl, optionally substituted heteroaryl, or optionally substituted heteroarylalkyl;

or two R^{11} on the same nitrogen atom are taken together with the nitrogen to which they are attached to form an optionally substituted N-heterocyclyl;

each R^{12} is independently H, optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted heterocyclyl, or optionally substituted heterocyclylalkyl;

each R^{13} is independently H, alkyl, carbocyclyl, carbocyclylalkyl, heterocyclyl, or heterocyclylalkyl; wherein each R^{13} is independently unsubstituted or substituted with $-R^b-OR^a$, $-R^b-C(O)OR^a$, or $-R^b-C(O)R^a$; wherein each R^a is independently hydrogen, alkyl which is optionally substituted with halogen, hydroxy, methoxy, or trifluoromethyl, or carbocyclyl which is optionally substituted with halogen, hydroxy, methoxy, or trifluoromethyl; and each R^b is independently a direct bond or a straight or branched alkylene;

two R^{13} groups are joined with the nitrogen to which they are attached join to form an N-heterocyclyl which is unsubstituted or substituted with alkyl, optionally substituted heterocyclyl, $-R^b-OR^a$, $-R^b-N(R^a)_2$, $-R^b-C(O)R^a$, $-R^b-CN$, or $-R^b-N(R^a)C(O)R^a$; wherein each R^a is independently hydrogen, alkyl

which is optionally substituted with halogen, hydroxy, methoxy, or trifluoromethyl, or carbocyclyl which is optionally substituted with halogen, hydroxy, methoxy, or trifluoromethyl; and each R^b is independently a direct bond or a straight or branched alkylene; and

each R^{14} is independently H, unsubstituted alkyl, or unsubstituted heterocyclyl;

wherein

each carbocyclyl is a stable non-aromatic monocyclic, fused polycyclic, or bridged polycyclic hydrocarbon ring comprising from three to ten carbon atoms;

each carbocyclylalkyl is a -R^c-carbocyclyl, where R^c is a C_1-C_8 alkylene;

each heterocyclyl is a stable 3- to 18-membered non-aromatic monocyclic, fused bicyclic, bridged bicyclic, or spirocyclic bicyclic ring system comprising from two to twelve carbon atoms and from one to six heteroatoms selected from nitrogen, oxygen, and sulfur which is optionally oxidized; and

each heterocyclylalkyl is a -R^c- heterocyclyl, where R^c is a C_1-C_8 alkylene.

~~optionally substituted alkyl, optionally substituted alkenyl, optionally substituted alkynyl, optionally substituted carbocyclyl, optionally substituted aryl, optionally substituted heterocyclyl, optionally substituted heteroaryl, -CN, -$N(R^{13})_2$, -OR^{13}, -COR^{13}, -CO_2R^{13}, -$CON(R^{13})_2$, -$N(R^{14})COR^{13}$, -SO_2R^{13}, -$SO_2N(R^{13})_2$, -$N(R^{14})SO_2R^{13}$, -$N(R^{14})CON(R^{13})_2$, -$N(R^{14})CO_2R^{13}$, -O-$CON(R^{13})_2$, -$N(R^{14})SO_2N(R^{13})_2$, -O-$SO_2N(R^{13})_2$, -$N(R^{14})SO_2$-OR^{13}, or -$C(=N$-$OR^{14})(R^{13})$;~~

~~each R^{11} is independently H, optionally substituted alkyl, optionally substituted alkenyl, optionally substituted alkynyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted aryl, optionally substituted aralkyl, optionally substituted heterocyclyl, optionally substituted heterocyclylalkyl, optionally substituted heteroaryl, or optionally substituted heteroarylalkyl;~~

~~or two R^{11} on the same nitrogen atom are taken togehter with the nitrogen to which they are attached to form an optionally substituted N-heterocyclyl;~~

~~each R^{12} is independently H, optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted heterocyclyl, or optionally substituted heterocyclylalkyl;~~

~~each R^{13} is H, optionally substituted alkyl, optionally substituted alkenyl, optionally substituted alkynyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted aryl, optionally substituted aralkyl, optionally substituted heterocyclyl, optionally substituted heterocyclylalkyl, optionally substituted heteroaryl, or optionally substituted heteroarylalkyl;~~

~~or two R^{13} on the same nitrogen atom are taken togehter with the nitrogen to which they are attached to form an optionally substituted N-heterocyclyl; and~~

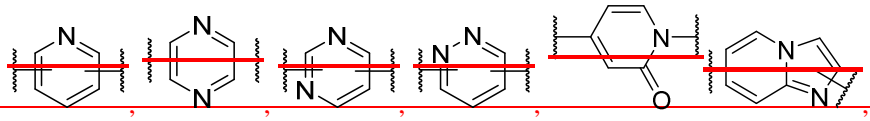
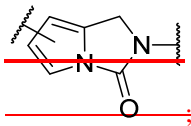
each R^{14} is independently H, optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted heterocyclyl, or optionally substituted heterocyclylalkyl.

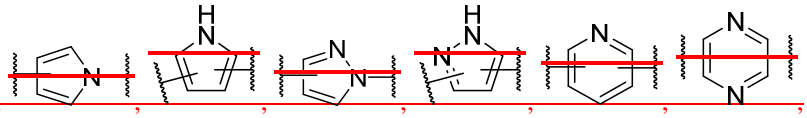
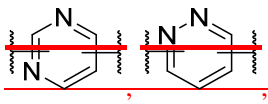
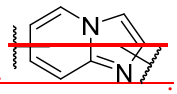
2. The compound of claim 1, or a pharmaceutically acceptable salt, solvate, or prodrug thereof, wherein:

Ring B is monocyclic or bicyclic aryl, monocyclic or bicyclic heteroaryl containing 1 or 2 heteroatoms selected from O, N, and S, or monocyclic or bicyclic 5- to 12-membered heterocyclyl containing 1 to 3 heteroatoms selected from O, N, and S; and

Ring C is phenyl, monocyclic 3- to 6-membered carbocyclyl, or monocyclic or bicyclic heteroaryl containing 1 or 2 heteroatoms selected from O, N, and S.

3. The compound of claim 1 or claim 2, or a pharmaceutically acceptable salt, solvate, or prodrug thereof, wherein:

Ring B is phenyl, , or ; and

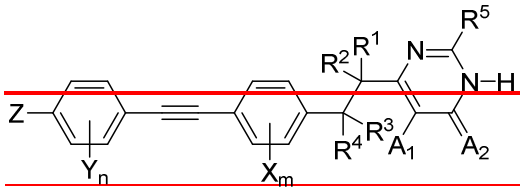
Ring C is phenyl, cyclohexyl, , , , or .

4. The compound of any one of claims 1 - 3, or a pharmaceutically acceptable salt, solvate, or prodrug thereof, wherein:

Ring B is phenyl; and

Ring C is phenyl.

5. The compound of any one of claims 1 - 4, or a pharmaceutically acceptable salt, solvate, or prodrug thereof, wherein the compound has the structure of Formula (II):



Formula (II).

6. The compound of any one of claims 1 - 5, or a pharmaceutically acceptable salt, solvate, or prodrug thereof, wherein:

each X and Y is independently H, optionally substituted alkyl, halo, or cyano;

~~n is 0 or 1; and~~

~~m is 0 or 1.~~

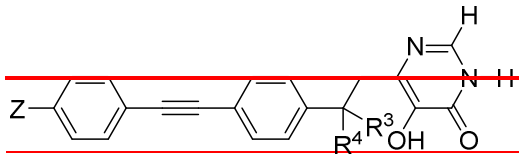
~~7. The compound of any one of claims 1 – 6, or a pharmaceutically acceptable salt, solvate, or prodrug thereof, wherein:~~

~~A_1 is OH;~~

~~A_2 is O; and~~

~~R^5 is H.~~

~~8. The compound of any one of claims 1 – 7, or a pharmaceutically acceptable salt, solvate, or prodrug thereof, wherein the compound has the structure of Formula (III):~~



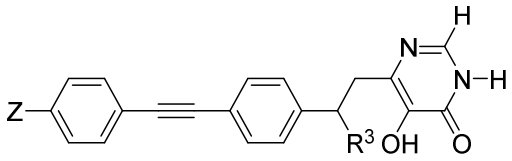
~~Formula (III).~~

2~~9~~. The compound of claim 1 ~~any one of claims 1 – 8~~, or a pharmaceutically acceptable salt~~,~~ or solvate~~, or prodrug~~ thereof, wherein:

each X and Y is independently H, F, or Cl; and

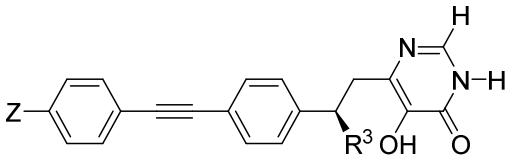
R^4 is H ~~or optionally substituted alkyl~~.

3~~10~~. The compound of ~~any one of claims 1 – 9~~ claim 1 or claim 2, or a pharmaceutically acceptable salt~~,~~ or solvate~~, or prodrug~~ thereof, wherein the compound has the structure of Formula (IV):

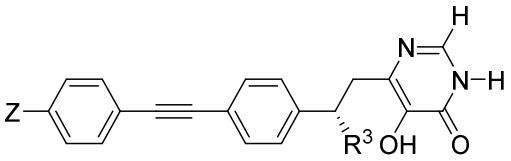


Formula (IV).

4~~11~~. The compound of ~~any one of claims 1 – 10~~ claim 1 or claim 2, or a pharmaceutically acceptable salt or~~,~~ solvate~~, or prodrug~~ thereof, wherein the compound has the structure of Formula (IVa) or Formula (IVb):



Formula (IVa)



Formula (IVb).

5~~12~~. The compound of any one of claims 1 - 4~~11~~, or a pharmaceutically acceptable salt or~~,~~ solvate~~, or prodrug~~ thereof, wherein:

R^3 is optionally substituted alkyl, optionally substituted (C_0-C_4 alkylene)-CO_2R^{11}, optionally substituted (C_0-C_4 alkylene)-$CON(R^{11})_2$, optionally substituted (C_0-C_4 alkylene)-OR^{11}, optionally substituted (C_0-C_4 alkylene)-$N(R^{11})_2$, or optionally substituted (C_0-C_4 alkylene)-$N(R^{12})$-SO_2R^{11}.

6~~13~~. The compound of any one of claims 1 - 5~~12~~, or a pharmaceutically acceptable salt or~~,~~ solvate ~~, or prodrug~~ thereof, wherein:

each R^{11} is H, optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted heterocyclyl, or optionally substituted heteroarylalkyl;

or two R^{11} groups are joined with the nitrogen to which they are attached join to form an optionally substituted N-heterocyclyl~~; and~~

~~each R^{12} is independently H or unsubstituted alkyl~~.

7~~14~~. The compound of claim 6~~13~~, or a pharmaceutically acceptable salt~~,~~ or solvate ~~, or prodrug~~ thereof, wherein:

each R^{11} is independently unsubstituted or substituted with halogen, -CN, $-R^b-OR^a$, $-R^b-C(O)R^a$, or $-R^b-S(O)_tR^a$; wherein t is 1 or 2; each R^a is independently hydrogen or alkyl which is optionally substituted with halogen, hydroxy, methoxy, or trifluoromethyl; and each R^b is independently a direct bond or a straight or branched alkylene;

or two R^{11} groups are joined with the nitrogen to which they are attached join to form an N-heterocyclyl which is optionally substituted with halogen, oxo, -CN, or $-R^b-OR^a$; wherein each R^a is independently hydrogen or alkyl which is optionally substituted with halogen, hydroxy, methoxy, or trifluoromethyl; and each R^b is independently a direct bond or a straight or branched alkylene; and

each R^{12} is independently H or unsubstituted alkyl.

~~15. The compound of claim 13 or claim 14, or a pharmaceutically acceptable salt, solvate, or prodrug thereof, wherein:~~

~~each R^{11} is independently unsubstituted or substituted with F, Cl, CN, OH, OMe, SO_2Me, or C(O)Me;~~

~~or two R^{11} groups are joined with the nitrogen to which they are attached join to form an N-heterocyclyl which is unsubstituted or substituted with CN, OH, or OMe; and~~

~~each R^{12} is independently H or unsubstituted alkyl.~~

8~~16~~. The compound of any one of claims 1 - 7~~15~~, or a pharmaceutically acceptable salt~~,~~ or solvate ~~, or prodrug~~ thereof, wherein:

~~Z is -L-G.~~

L is a bond;

or L is optionally substituted C_1-C_2 alkylene~~;~~.

~~G is optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted aryl, optionally substituted heterocyclyl, optionally substituted heteroaryl, $-N(R^{13})_2$, $-OR^{13}$, -CN, $-COR^{13}$, $-CO_2R^{13}$, $-CON(R^{13})_2$, $-N(R^{14})COR^{13}$, $-SO_2R^{13}$, $-SO_2N(R^{13})_2$, or $-N(R^{14})SO_2R^{13}$;~~

~~each R^{13} is independently H, optionally substituted alkyl, optionally substituted carbocyclyl, optionally substituted carbocyclylalkyl, optionally substituted heterocyclyl, or optionally substituted heterocyclylalkyl;~~

~~or two R^{13} groups are joined with the nitrogen to which they are attached join to form an optionally substituted N-heterocyclyl; and~~

~~each R^{14} is independently H, unsubstituted alkyl, or unsubstituted heterocyclyl.~~

~~17. The compound of any one of claims 1 - 16, or a pharmaceutically acceptable salt, solvate, or prodrug thereof, wherein:~~

~~L is optionally substituted C_1-C_2 alkylene;~~

~~G is optionally substituted alkyl, optionally substituted heterocyclyl, optionally substituted heteroaryl, $-N(R^{13})_2$, $-OR^{13}$, $-CON(R^{13})_2$, or $-N(R^{14})COR^{13}$;~~

~~each R^{13} is independently unsubstituted or substituted with -OH, -OMe, $-C(O)CH_2OH$, $-CH_2C(O)OH$, -C(O)OH, -C(O)-cyclopropyl;~~

~~or two R^{13} groups are joined with the nitrogen to which they are attached join to form an N-heterocyclyl which is unsubstituted or substituted with methyl, oxetanyl, morpholinyl, -OMe, $-CH_2OH$, $-NH_2$, $-CH_2NH_2$, $-C(O)CH_2OH$, -CN, $-CH_2CN$, $-CH_2NHC(O)CH_2OH$; and~~

~~each R^{14} is independently H, unsubstituted alkyl, or unsubstituted heterocyclyl.~~

~~18~~9. The compound of any one of claims 1 - ~~17~~8, or a pharmaceutically acceptable salt~~,~~ or solvate~~, or prodrug~~ thereof, wherein:

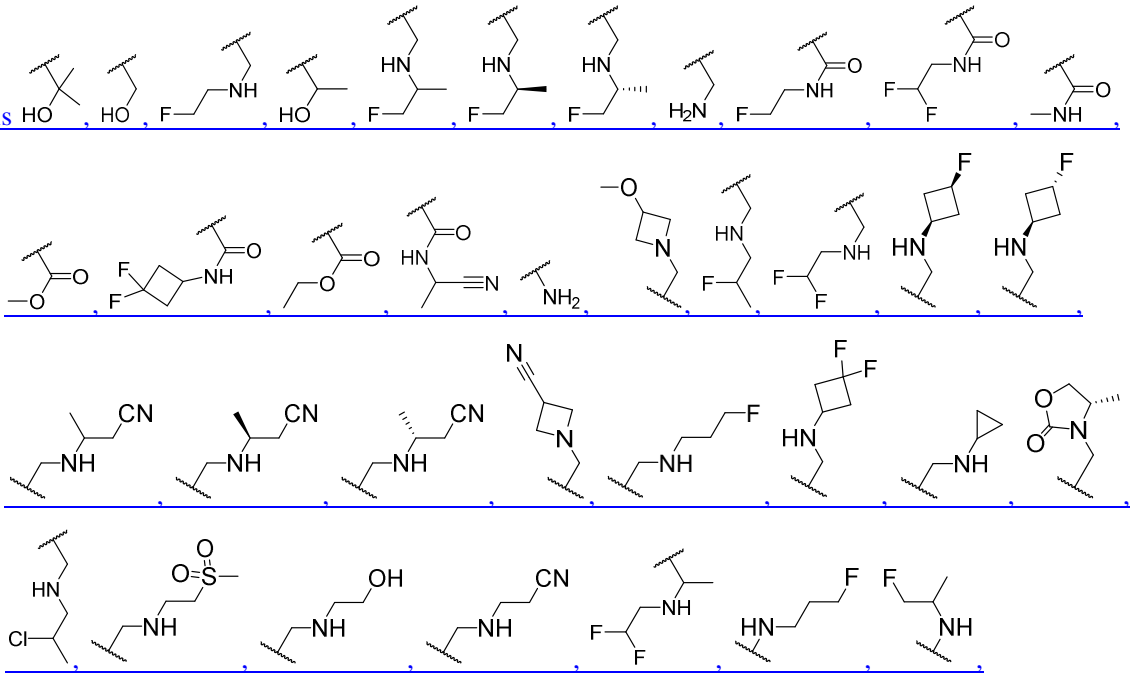
G is ~~optionally substituted~~ 4- to 6-membered monocyclic heterocyclyl.

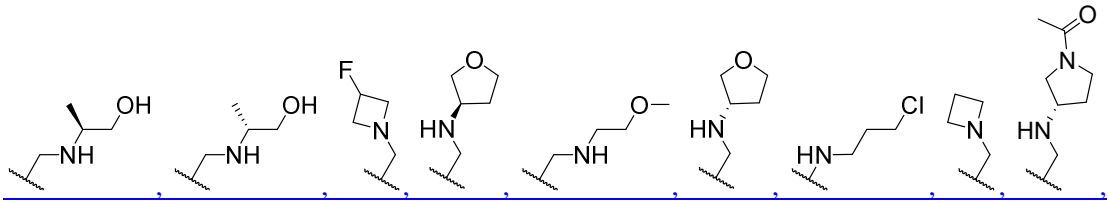
10. The compound of claim 1, or a pharmaceutically acceptable salt or solvate thereof, wherein;

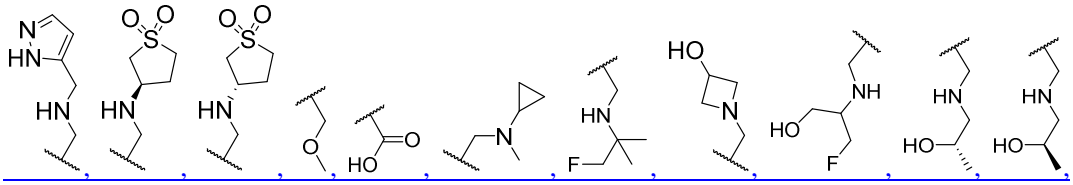
n is 0;

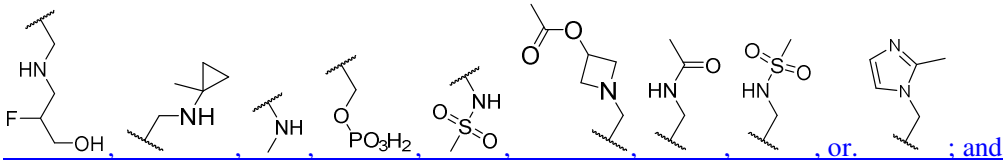
m is 0;

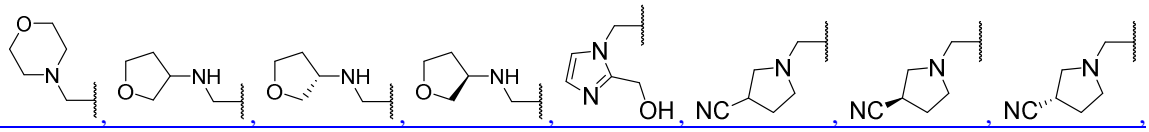
R^4 is H;

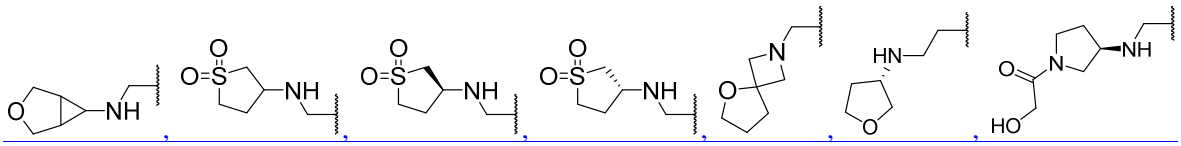
R^3 is 

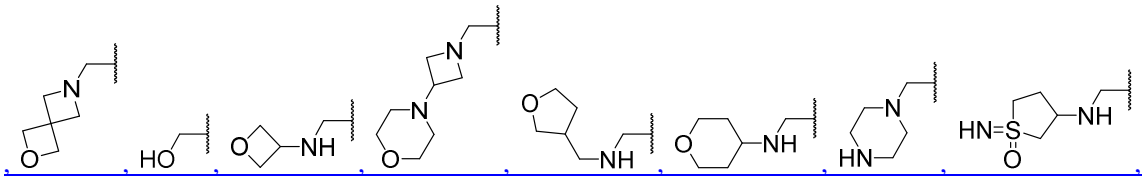


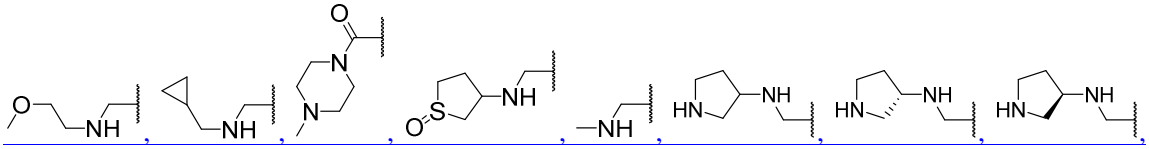


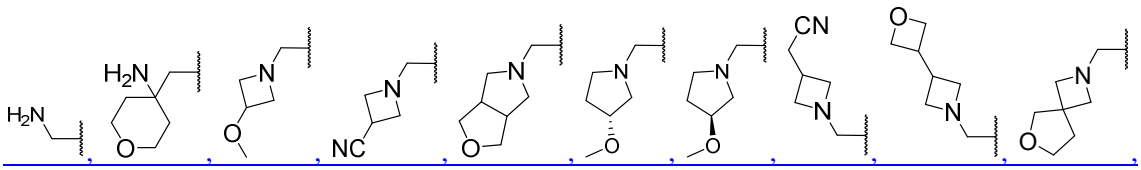

, or, ; and

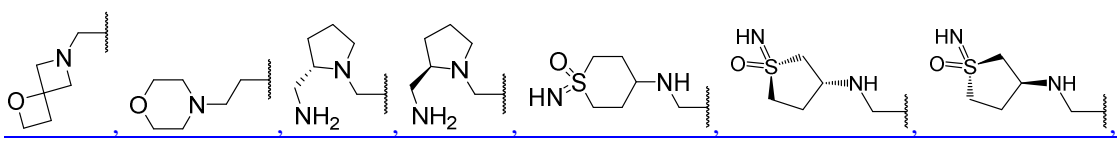
Z is 

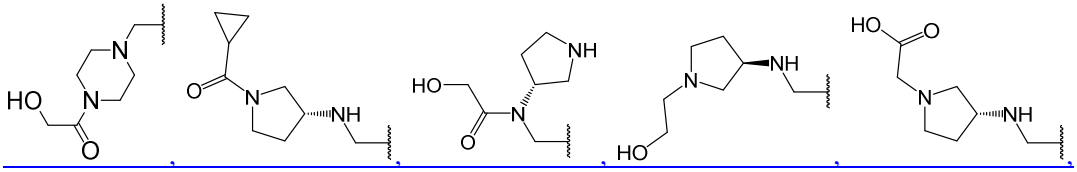


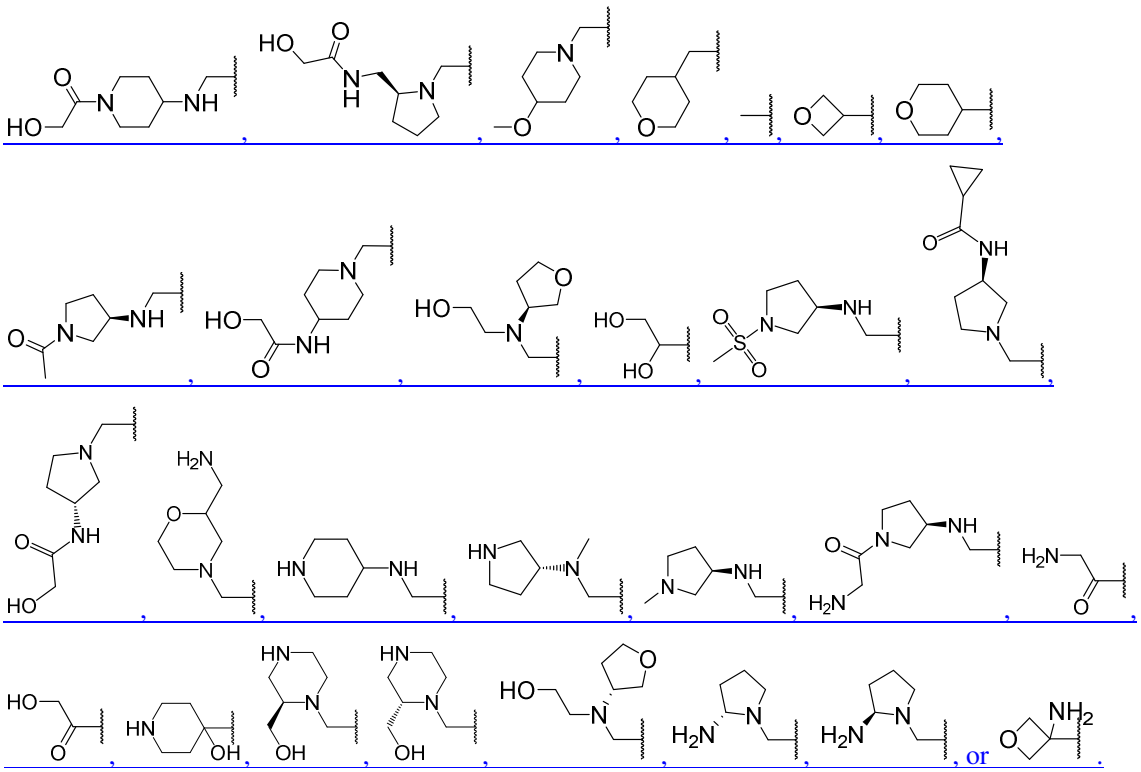












~~19~~11. A compound selected from:

5-hydroxy-6-(3-hydroxy-3-methyl-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)butyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-hydroxy-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-hydroxy-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(3-((2-fluoroethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-((2-fluoroethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-((2-fluoroethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-hydroxy-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-hydroxy-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)butyl)pyrimidin-4(3H)-one;

6-((R)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((R)-1-fluoropropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((R)-1-fluoropropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((R)-1-fluoropropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-amino-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

N-(2-fluoroethyl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propanamide;

(R)-N-(2-fluoroethyl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propanamide;

(S)-N-(2-fluoroethyl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propanamide;

N-(2,2-difluoroethyl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propanamide;

3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-N-methyl-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propanamide;

methyl 3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propanoate;

N-(3,3-difluorocyclobutyl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propanamide;

ethyl 3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(((tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propanoate;

methyl 3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(((tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propanoate;

N-(1-cyanoethyl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propanamide;

N-(2,2-difluoroethyl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-((2-(hydroxymethyl)-1H-imidazol-1-yl)methyl)phenyl)ethynyl)phenyl)propanamide;

N-(2-fluoroethyl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propanamide;

5-hydroxy-6-(3-hydroxy-2-methyl-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-methyl-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

1-(4-((4-((S)-1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

1-(4-((4-((R)-1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(3S)-1-(4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(3S)-1-(4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(S)-1-(4-((4-((S)-1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(S)-1-(4-((4-((R)-1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(3R)-1-(4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile ;

6-(2-(4-((4-(((3-oxabicyclo[3.1.0]hexan-6-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-(((1,1-dioxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-((((S)-1,1-dioxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-2-(4-((4-((((S)-1,1-dioxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-2-(4-((4-((((S)-1,1-dioxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-((((R)-1,1-dioxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-2-(4-((4-((((R)-1,1-dioxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-2-(4-((4-((((R)-1,1-dioxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-((5-oxa-2-azaspiro[3.4]octan-2-yl)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(2-(((S)-tetrahydrofuran-3-yl)amino)ethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((R)-1-(2-hydroxyacetyl)pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-hydroxy-2-(4-((4-((((R)-1-(2-hydroxyacetyl)pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-hydroxy-2-(4-((4-((((R)-1-(2-hydroxyacetyl)pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(2-(4-((4-((2-oxa-6-azaspiro[3.3]heptan-6-yl)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(hydroxymethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((oxetan-3-ylamino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

ethyl 3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-((oxetan-3-ylamino)methyl)phenyl)ethynyl)phenyl)propanoate;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((3-morpholinoazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((tetrahydrofuran-3-yl)methyl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(((tetrahydro-2H-pyran-4-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(piperazin-1-ylmethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-hydroxy-2-(4-((4-(piperazin-1-ylmethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-hydroxy-2-(4-((4-(piperazin-1-ylmethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(((1-imino-1-oxidotetrahydro-1H-1l6-thiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(((2-methoxyethyl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(2-(4-((4-(((cyclopropylmethyl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(4-methylpiperazine-1-carbonyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(((1-oxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((methylamino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((R)-pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(2-(4-((4-(aminomethyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-((4-aminotetrahydro-2H-pyran-4-yl)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-amino-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((R)-1-fluoropropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((R)-1-fluoropropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((R)-1-fluoropropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-(3-methoxyazetidin-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-(3-methoxyazetidin-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-(3-methoxyazetidin-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(3-((2-fluoropropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((cis-3-fluorocyclobutyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-((cis-3-fluorocyclobutyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-((cis-3-fluorocyclobutyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((trans-3-fluorocyclobutyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-((trans-3-fluorocyclobutyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-((trans-3-fluorocyclobutyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

1-(4-((4-(1-(((S)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)azetidine-3-carbonitrile;

1-(4-((4-((R)-1-(((S)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)azetidine-3-carbonitrile;

1-(4-((4-((S)-1-(((S)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)azetidine-3-carbonitrile;

1-(4-((4-(1-(((R)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)azetidine-3-carbonitrile;

1-(4-((4-((R)-1-(((R)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)azetidine-3-carbonitrile;

1-(4-((4-((S)-1-(((R)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)azetidine-3-carbonitrile;

6-(3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((tetrahydro-1H-furo[3,4-c]pyrrol-5(3H)-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2R)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((tetrahydro-1H-furo[3,4-c]pyrrol-5(3H)-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2S)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((tetrahydro-1H-furo[3,4-c]pyrrol-5(3H)-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(((R)-3-methoxypyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(((R)-3-methoxypyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(((R)-3-methoxypyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(((S)-3-methoxypyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(((S)-3-methoxypyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(((S)-3-methoxypyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(3R)-1-(4-((4-(1-(((S)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(3S)-1-(4-((4-(1-(((S)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(S)-1-(4-((4-((R)-1-(((S)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(S)-1-(4-((4-((S)-1-(((S)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

6-(3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((oxetan-3-ylamino)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

2-(1-(4-((4-(1-(((S)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)azetidin-3-yl)acetonitrile;

2-(1-(4-((4-((R)-1-(((S)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)azetidin-3-yl)acetonitrile;

2-(1-(4-((4-((S)-1-(((S)-1-fluoropropan-2-yl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)azetidin-3-yl)acetonitrile;

3-((3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)amino)butanenitrile;

(3S)-3-((3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)amino)butanenitrile;

(S)-3-(((S)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)amino)butanenitrile;

(S)-3-(((R)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)amino)butanenitrile;

(3R)-3-((3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)amino)butanenitrile;

(R)-3-(((S)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)amino)butanenitrile;

(R)-3-(((R)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)amino)butanenitrile;

6-(3-((2-fluoropropyl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

1-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)azetidine-3-carbonitrile;

(R)-1-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)azetidine-3-carbonitrile;

(S)-1-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)azetidine-3-carbonitrile;

6-(3-((3-fluoropropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-((3-fluoropropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-((3-fluoropropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((3,3-difluorocyclobutyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-((3,3-difluorocyclobutyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-((3,3-difluorocyclobutyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((2,2-difluoroethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-((2,2-difluoroethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-((2,2-difluoroethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(cyclopropylamino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-(cyclopropylamino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-(cyclopropylamino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(4S)-3-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-4-methyloxazolidin-2-one;

(S)-3-((S)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-4-methyloxazolidin-2-one;

(S)-3-((R)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-4-methyloxazolidin-2-one;

(3S)-1-(4-((4-(1-((2,2-difluoroethyl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

6-(2-(4-((4-((2-oxa-6-azaspiro[3.3]heptan-6-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((2-chloropropyl)amino)-2-(4-((4-(((R)-3-methoxypyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((3-(oxetan-3-yl)azetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-((6-oxa-2-azaspiro[3.4]octan-2-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-((2-(methylsulfonyl)ethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-((2-hydroxyethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

3-((3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)amino)propanenitrile;

6-(3-((2,2-difluoroethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)butyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-((1-oxa-6-azaspiro[3.3]heptan-6-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-2-(4-((4-((1-oxa-6-azaspiro[3.3]heptan-6-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-2-(4-((4-((1-oxa-6-azaspiro[3.3]heptan-6-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

3-((2-(4-((4-((3-(cyanomethyl)azetidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propyl)amino)propanenitrile;

6-(3-((2-fluoropropyl)amino)-2-(4-((4-(((R)-3-methoxypyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2R)-3-((2-fluoropropyl)amino)-2-(4-((4-(((R)-3-methoxypyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2S)-3-((2-fluoropropyl)amino)-2-(4-((4-(((R)-3-methoxypyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((2-fluoropropyl)amino)-2-(4-((4-(2-morpholinoethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-((3-fluoropropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(2-((3-fluoropropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(2-((3-fluoropropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-((1-fluoropropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2R)-2-((1-fluoropropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2S)-2-((1-fluoropropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)-2-methylpropyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-(((S)-1-hydroxypropan-2-yl)amino)-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(2-(4-((4-(((S)-2-(aminomethyl)pyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-2-(4-((4-(((S)-2-(aminomethyl)pyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-2-(4-((4-(((S)-2-(aminomethyl)pyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-(((R)-2-(aminomethyl)pyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(((1-imino-1-oxidohexahydro-1l6-thiopyran-4-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((1S,3R)-1-imino-1-oxidotetrahydro-1H-1l6-thiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((1R,3S)-1-imino-1-oxidotetrahydro-1H-1l6-thiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((4-(2-hydroxyacetyl)piperazin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(2-(4-((4-((((R)-1-(cyclopropanecarbonyl)pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

2-hydroxy-N-(4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)-N-((R)-pyrrolidin-3-yl)acetamide;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((R)-1-(2-hydroxyethyl)pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

2-((3R)-3-((4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)amino)pyrrolidin-1-yl)acetic acid;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(((1-(2-hydroxyacetyl)piperidin-4-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-hydroxy-2-(4-((4-(((1-(2-hydroxyacetyl)piperidin-4-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-hydroxy-2-(4-((4-(((1-(2-hydroxyacetyl)piperidin-4-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

2-hydroxy-N-(((2S)-1-(4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidin-2-yl)methyl)acetamide;

6-(3-(3-fluoroazetidin-1-yl)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-(3-fluoroazetidin-1-yl)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-(3-fluoroazetidin-1-yl)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(((R)-tetrahydrofuran-3-yl)amino)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(((R)-tetrahydrofuran-3-yl)amino)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(((R)-tetrahydrofuran-3-yl)amino)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-((2-methoxyethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-((2-methoxyethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-((2-methoxyethyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(((S)-tetrahydrofuran-3-yl)amino)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-(((S)-1-hydroxypropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-(((S)-1-hydroxypropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-(((S)-1-hydroxypropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-(((R)-1-hydroxypropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-(((R)-1-hydroxypropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-(((R)-1-hydroxypropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(2-((3-chloropropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-amino-2-(4-((4-((((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-amino-2-(4-((4-((4-methoxypiperidin-1-yl)methyl)phenyl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-(3-methoxyazetidin-1-yl)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-(3-methoxyazetidin-1-yl)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(3-(azetidin-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-(((1,1-dioxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2R)-2-(4-((4-(((1,1-dioxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2S)-2-(4-((4-(((1,1-dioxidotetrahydrothiophen-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((S)-1-acetylpyrrolidin-3-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((S)-1-acetylpyrrolidin-3-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((S)-1-acetylpyrrolidin-3-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((1H-pyrazol-5-yl)methyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((1H-pyrazol-5-yl)methyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((1H-pyrazol-5-yl)methyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((R)-1,1-dioxidotetrahydrothiophen-3-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((R)-1,1-dioxidotetrahydrothiophen-3-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((R)-1,1-dioxidotetrahydrothiophen-3-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((S)-1,1-dioxidotetrahydrothiophen-3-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-3-(((S)-1,1-dioxidotetrahydrothiophen-3-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-3-(((S)-1,1-dioxidotetrahydrothiophen-3-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-methoxy-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

N-(2,2-difluoroethyl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(6-((4-(morpholinomethyl)phenyl)ethynyl)-3-oxo-1H-pyrrolo[1,2-c]imidazol-2(3H)-yl)propanamide;

methyl 3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(6-((4-(morpholinomethyl)phenyl)ethynyl)-3-oxo-1H-pyrrolo[1,2-c]imidazol-2(3H)-yl)propanoate;

2-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)-6-((4-(morpholinomethyl)phenyl)ethynyl)-1,2-dihydro-3H-pyrrolo[1,2-c]imidazol-3-one;

5-hydroxy-6-(3-hydroxy-2-(4-((1-((tetrahydro-2H-pyran-4-yl)methyl)-1H-pyrazol-4-yl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((6-(morpholinomethyl)pyridin-3-yl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(5-((4-(morpholinomethyl)phenyl)ethynyl)pyridin-2-yl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((1-methyl-1H-pyrazol-4-yl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((E)-4-((((S)-tetrahydrofuran-3-yl)amino)methyl)styryl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(3-((2-fluoroethyl)amino)-2-(4-((1-methyl-1H-pyrazol-4-yl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-amino-2-(4-((1-methyl-1H-pyrazol-4-yl)ethynyl)phenyl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((1-(oxetan-3-yl)-1H-pyrazol-4-yl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((1-(tetrahydro-2H-pyran-4-yl)-1H-pyrazol-4-yl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

2-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)-6-(4-(2-morpholinoethyl)phenyl)-1,2-dihydro-3H-pyrrolo[1,2-c]imidazol-3-one;

5-hydroxy-6-((S)-3-hydroxy-2-(4-((trans-4-(morpholinomethyl)cyclohexyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-hydroxy-2-(4-((trans-4-(morpholinomethyl)cyclohexyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(3-((2-fluoroethyl)amino)-2-(4-((1-(tetrahydro-2H-pyran-4-yl)-1H-pyrazol-4-yl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((2-fluoroethyl)amino)-2-(4-((1-((tetrahydro-2H-pyran-4-yl)methyl)-1H-pyrazol-4-yl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((2-fluoroethyl)amino)-2-(4-(imidazo[1,5-a]pyridin-7-ylethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((2-fluoroethyl)amino)-2-(4-((1-(oxetan-3-yl)-1H-pyrazol-4-yl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-((5-((4-(morpholinomethyl)phenyl)ethynyl)-2,3-dihydro-1H-inden-1-yl)methyl)pyrimidin-4(3H)-one;

3-((5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)methyl)-6-((4-(morpholinomethyl)phenyl)ethynyl)isoindolin-1-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)-2-oxopyridin-1(2H)-yl)propyl)pyrimidin-4(3H)-one;

3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(6-(4-(2-morpholinoethyl)phenyl)-3-oxo-1H-pyrrolo[1,2-c]imidazol-2(3H)-yl)propanoic acid;

6-(2-([1,1'-biphenyl]-4-yl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-amino-2-(4'-(2-morpholinoethyl)-[1,1'-biphenyl]-4-yl)ethyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(3-fluoroazetidin-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-(3-fluoroazetidin-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-(3-fluoroazetidin-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one ;

(S)-5-hydroxy-6-(3-hydroxy-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-hydroxy-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-1-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)azetidine-3-carbonitrile;

(R)-1-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)azetidine-3-carbonitrile;

6-(3-(cyclopropyl(methyl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-(cyclopropyl(methyl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-(cyclopropyl(methyl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((1-fluoro-2-methylpropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-((1-fluoro-2-methylpropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-((1-fluoro-2-methylpropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(cyclopropylamino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-(cyclopropylamino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-(cyclopropylamino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(cyclopropylamino)-2-(2-fluoro-4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-(cyclopropylamino)-2-(2-fluoro-4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-(cyclopropylamino)-2-(2-fluoro-4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(cyclopropylamino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)-2-methylpropyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-(cyclopropylamino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)-2-methylpropyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-(cyclopropylamino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)-2-methylpropyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-(3-hydroxyazetidin-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-(3-hydroxyazetidin-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-(3-hydroxyazetidin-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-((((R)-1-acetylpyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-2-(4-((4-((((R)-1-acetylpyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-2-(4-((4-((((R)-1-acetylpyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-1-(4-((4-((R)-1-(3-cyanoazetidin-1-yl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(S)-1-(4-((4-((S)-1-(3-cyanoazetidin-1-yl)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-((3-fluoropropyl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-((3-fluoropropyl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-((3-fluoropropyl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(2-fluoro-4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-2-(2-fluoro-4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-2-(2-fluoro-4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(2-fluoro-4-((4-((tetrahydro-1H-furo[3,4-c]pyrrol-5(3H)-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2R)-2-(2-fluoro-4-((4-((tetrahydro-1H-furo[3,4-c]pyrrol-5(3H)-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2S)-2-(2-fluoro-4-((4-((tetrahydro-1H-furo[3,4-c]pyrrol-5(3H)-yl)methyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

2-hydroxy-N-(1-(4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)piperidin-4-yl)acetamide;

(R)-2-hydroxy-N-(1-(4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)piperidin-4-yl)acetamide;

(S)-2-hydroxy-N-(1-(4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)piperidin-4-yl)acetamide;

5-hydroxy-6-((S)-3-hydroxy-2-(4-((4-(((2-hydroxyethyl)((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-hydroxy-2-(4-((4-(((2-hydroxyethyl)((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(3-((1-fluoro-3-hydroxypropan-2-yl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-(((S)-2-hydroxypropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-(((S)-2-hydroxypropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-(((R)-2-hydroxypropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-(((R)-2-hydroxypropyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(3-((2-fluoro-3-hydroxypropyl)amino)-2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-((1-methylcyclopropyl)amino)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(2-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-((1-methylcyclopropyl)amino)propyl)pyrimidin-4(3H)-one;

6-(2-(4-((4-(1,2-dihydroxyethyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2R)-2-(4-((4-(1,2-dihydroxyethyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2S)-2-(4-((4-(1,2-dihydroxyethyl)phenyl)ethynyl)phenyl)-3-(((S)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-hydroxy-2-(4-((4-((((R)-1-(methylsulfonyl)pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-hydroxy-2-(4-((4-((((R)-1-(methylsulfonyl)pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((R)-1-(methylsulfonyl)pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

N-((3R)-1-(4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidin-3-yl)cyclopropanecarboxamide;

2-hydroxy-N-((3R)-1-(4-((4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidin-3-yl)acetamide;

6-(2-(4-((4-((2-(aminomethyl)morpholino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((piperidin-4-ylamino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((methyl((R)-pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-((((R)-1-methylpyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(2-(4-((4-((((R)-1-glycylpyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((1-glycylpiperidin-4-yl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((1-(2-hydroxyacetyl)piperidin-4-yl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(2-(methylamino)-2-(4-((4-((((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)ethyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(4-hydroxypiperidin-4-yl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(S)-5-hydroxy-6-(3-hydroxy-2-(4-((4-(4-hydroxypiperidin-4-yl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

(R)-5-hydroxy-6-(3-hydroxy-2-(4-((4-(4-hydroxypiperidin-4-yl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-hydroxy-2-(4-((4-(((R)-2-(hydroxymethyl)piperazin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-hydroxy-2-(4-((4-(((S)-2-(hydroxymethyl)piperazin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-hydroxy-2-(4-((4-(((2-hydroxyethyl)((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-hydroxy-2-(4-((4-(((2-hydroxyethyl)((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-hydroxy-2-(4-((4-(((2-hydroxyethyl)((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

6-(2-(4-((4-(1,2-dihydroxyethyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2S)-2-(4-((4-(1,2-dihydroxyethyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-((2R)-2-(4-((4-(1,2-dihydroxyethyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-hydroxy-2-(4-((4-((((R)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)ethyl)pyrimidine-4,5-diol;

N-(1-(4-((3-fluoro-4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)piperidin-4-yl)-2-hydroxyacetamide;

(R)-N-(1-(4-((3-fluoro-4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)piperidin-4-yl)-2-hydroxyacetamide;

(S)-N-(1-(4-((3-fluoro-4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)piperidin-4-yl)-2-hydroxyacetamide;

6-(2-(4-((4-((1-oxa-6-azaspiro[3.3]heptan-6-yl)methyl)phenyl)ethynyl)-2-fluorophenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(2-(4-((4-((1-oxa-6-azaspiro[3.3]heptan-6-yl)methyl)phenyl)ethynyl)-2-fluorophenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(2-(4-((4-((1-oxa-6-azaspiro[3.3]heptan-6-yl)methyl)phenyl)ethynyl)-2-fluorophenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

(3R)-1-(4-((3-fluoro-4-(1-hydroxy-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)phenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

6-(2-(4-((2-fluoro-4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(2-(4-((2-fluoro-4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(2-(4-((2-fluoro-4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((3-fluoro-4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(2-(4-((3-fluoro-4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(2-(4-((3-fluoro-4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(3-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(2-(3-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(2-(3-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-hydroxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-fluoroazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-fluoroazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-fluoroazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

1-(4-((4-(1-((2,2-difluoroethyl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)-3-fluorophenyl)ethynyl)benzyl)azetidine-3-carbonitrile;

(S)-1-(4-((4-(1-((2,2-difluoroethyl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)-3-fluorophenyl)ethynyl)benzyl)azetidine-3-carbonitrile;

(R)-1-(4-((4-(1-((2,2-difluoroethyl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)-3-fluorophenyl)ethynyl)benzyl)azetidine-3-carbonitrile;

6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(((R)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(((R)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(((R)-1-fluoropropan-2-yl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-1-(4-((4-(1-((2,2-difluoroethyl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)-3-fluorophenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(R)-1-(4-((4-((S)-1-((2,2-difluoroethyl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)-3-fluorophenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

(R)-1-(4-((4-((R)-1-((2,2-difluoroethyl)amino)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)propan-2-yl)-3-fluorophenyl)ethynyl)benzyl)pyrrolidine-3-carbonitrile;

6-(3-((2,2-difluoroethyl)amino)-2-(2-fluoro-4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-((2,2-difluoroethyl)amino)-2-(2-fluoro-4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-((2,2-difluoroethyl)amino)-2-(2-fluoro-4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((2,2-difluoroethyl)amino)-2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-((2,2-difluoroethyl)amino)-2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-((2,2-difluoroethyl)amino)-2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(((R)-2-hydroxypropyl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((S)-2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(((R)-2-hydroxypropyl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-((R)-2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(((R)-2-hydroxypropyl)amino)propyl)-5-hydroxypyrimidin-4(3H)-one;

5-hydroxy-6-(3-methoxy-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-methoxy-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-methoxy-2-(4-((4-((((S)-tetrahydrofuran-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl dihydrogen phosphate;

(S)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl dihydrogen phosphate;

(R)-3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl dihydrogen phosphate;

5-hydroxy-6-(3-methoxy-2-(4-((4-((((S)-pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-methoxy-2-(4-((4-((((S)-pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-methoxy-2-(4-((4-((((S)-pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-(3-methoxy-2-(4-((4-((((R)-pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((S)-3-methoxy-2-(4-((4-((((R)-pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

5-hydroxy-6-((R)-3-methoxy-2-(4-((4-((((R)-pyrrolidin-3-yl)amino)methyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

N-(2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

(R)-N-(2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

(S)-N-(2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

N-(2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-(4-hydroxypiperidin-4-yl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

(R)-N-(2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-(4-hydroxypiperidin-4-yl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

(S)-N-(2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-(4-hydroxypiperidin-4-yl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

N-(2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-(((S)-2-(hydroxymethyl)piperazin-1-yl)methyl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

N-((R)-2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-(((S)-2-(hydroxymethyl)piperazin-1-yl)methyl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

N-((S)-2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-(((S)-2-(hydroxymethyl)piperazin-1-yl)methyl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

6-(2-(4-((4-(((R)-3-aminopyrrolidin-1-yl)methyl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(2-(4-((4-(3-aminooxetan-3-yl)phenyl)ethynyl)phenyl)-3-hydroxypropyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(((S)-1-fluoropropan-2-yl)amino)-2-(4-((4-(2-morpholinoethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

N-(1-([1,1'-biphenyl]-4-yl)-2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)ethyl)methanesulfonamide;

(R)-N-(1-([1,1'-biphenyl]-4-yl)-2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)ethyl)methanesulfonamide;

(S)-N-(1-([1,1'-biphenyl]-4-yl)-2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)ethyl)methanesulfonamide;

N-(2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

(R)-N-(2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

(S)-N-(2-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-1-(4-((4-((3-methoxyazetidin-1-yl)methyl)phenyl)ethynyl)phenyl)ethyl)methanesulfonamide;

6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-methoxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-methoxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(2-(2-fluoro-4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)-3-(3-methoxyazetidin-1-yl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-((1-fluoro-2-methylpropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-((1-fluoro-2-methylpropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-((1-fluoro-2-methylpropan-2-yl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

6-(3-(cyclopropyl(methyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(S)-6-(3-(cyclopropyl(methyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

(R)-6-(3-(cyclopropyl(methyl)amino)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)-5-hydroxypyrimidin-4(3H)-one;

1-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)azetidin-3-yl acetate;

(S)-1-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)azetidin-3-yl acetate;

(R)-1-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)azetidin-3-yl acetate;

N-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)acetamide;

(S)-N-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)acetamide;

(R)-N-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)acetamide;

N-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)methanesulfonamide;

(S)-N-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)methanesulfonamide;

(R)-N-(3-(5-hydroxy-6-oxo-1,6-dihydropyrimidin-4-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)methanesulfonamide;

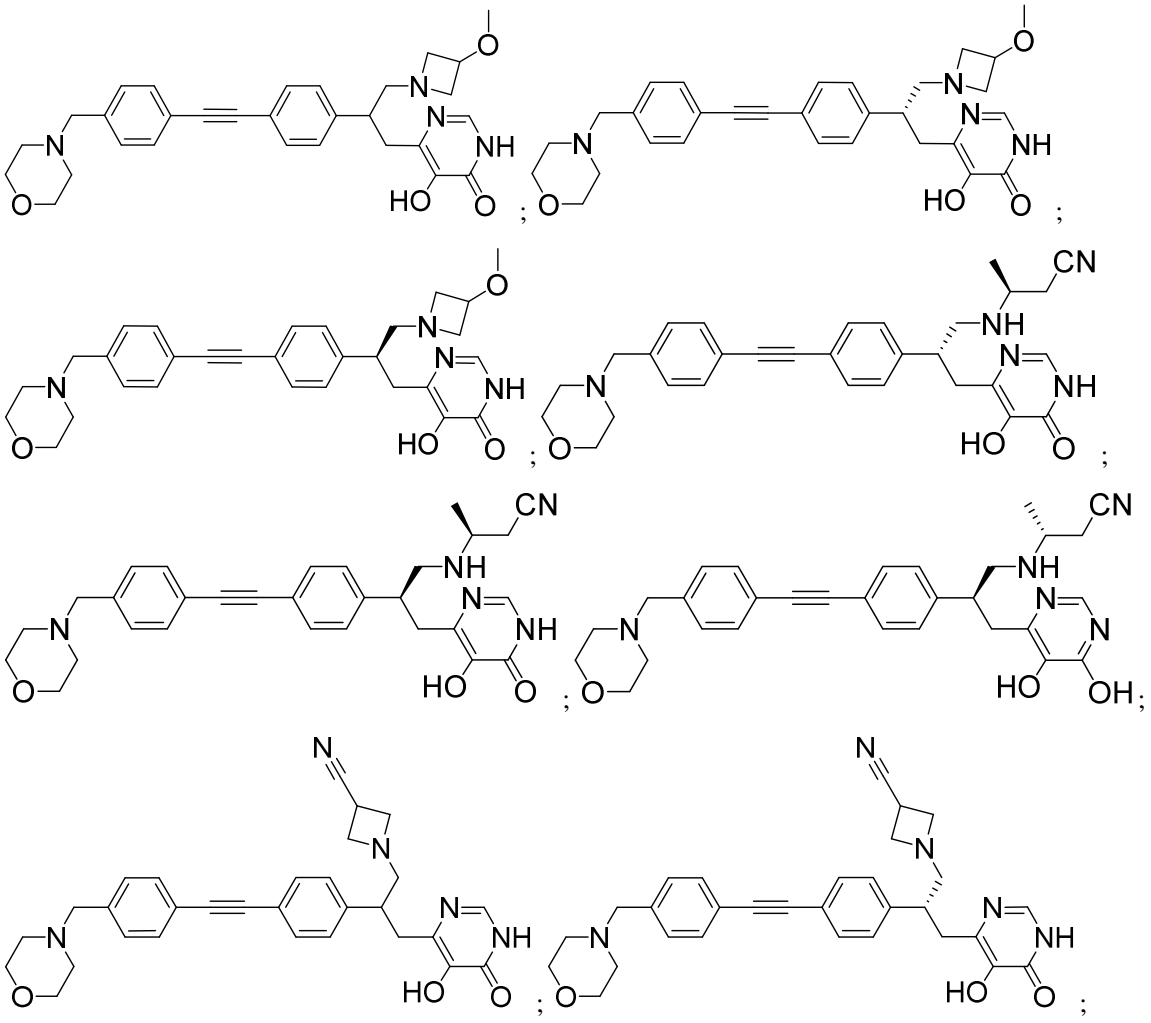
5-hydroxy-6-(3-(2-methyl-1H-imidazol-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

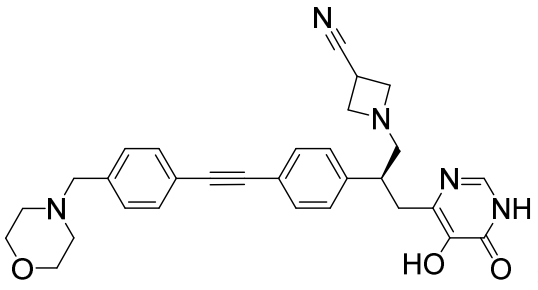
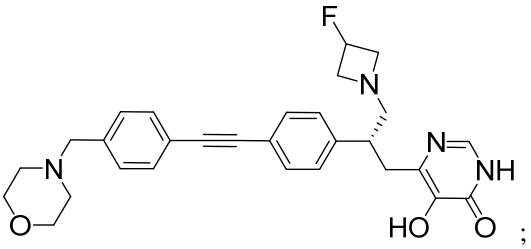
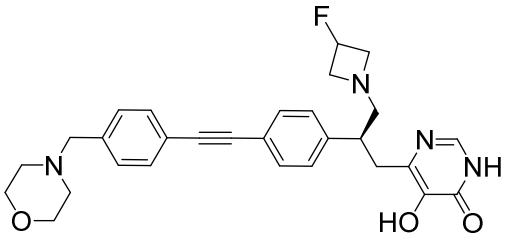
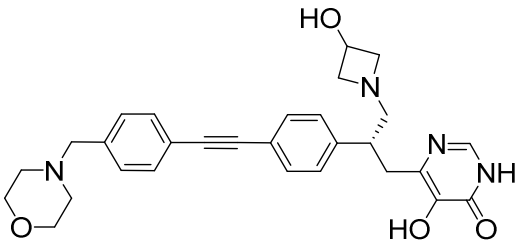
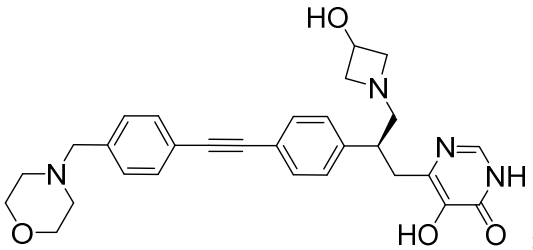
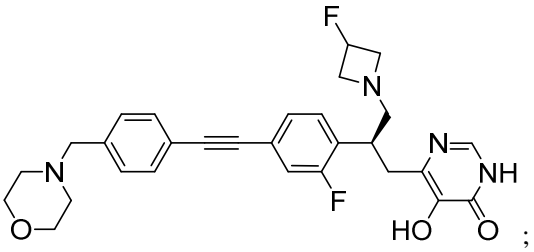
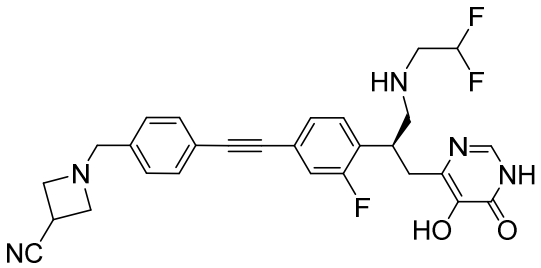
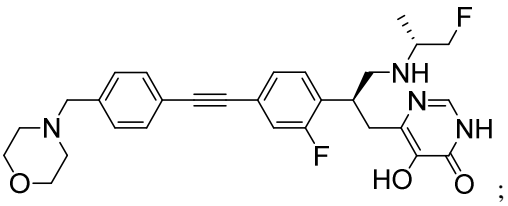
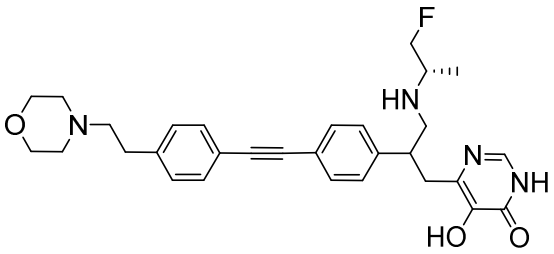
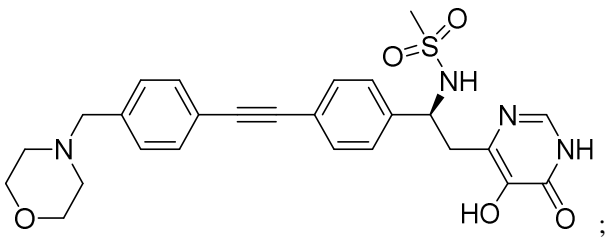
(S)-5-hydroxy-6-(3-(2-methyl-1H-imidazol-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one; and

(R)-5-hydroxy-6-(3-(2-methyl-1H-imidazol-1-yl)-2-(4-((4-(morpholinomethyl)phenyl)ethynyl)phenyl)propyl)pyrimidin-4(3H)-one;

or a pharmaceutically acceptable salt or, solvate, or prodrug thereof.

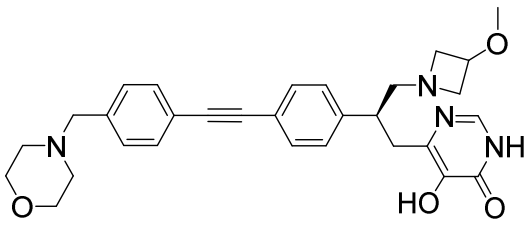
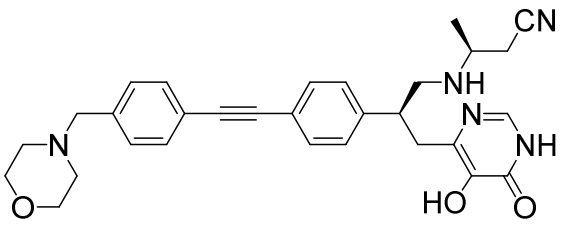
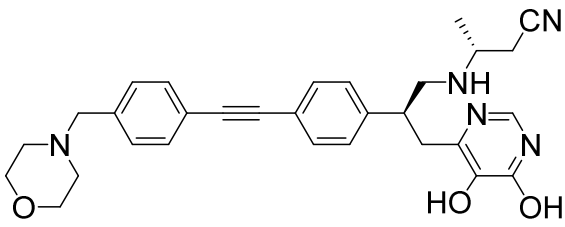
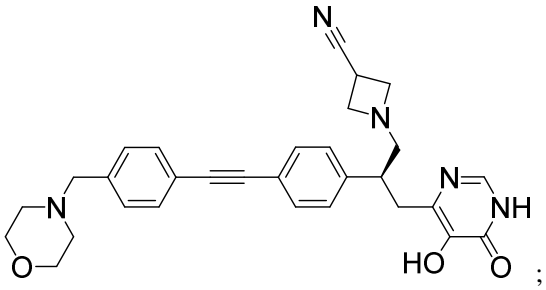
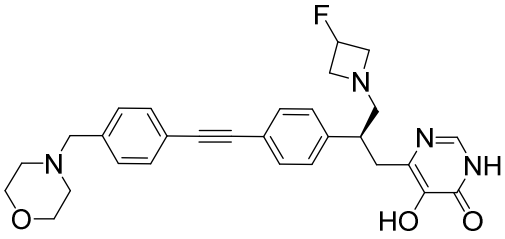
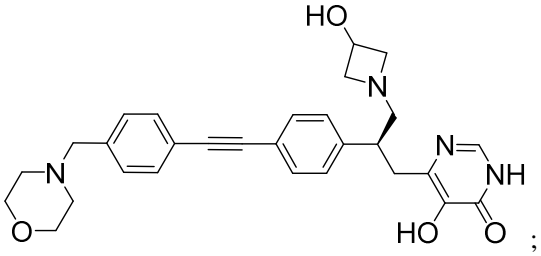
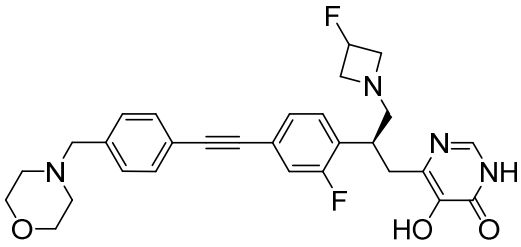
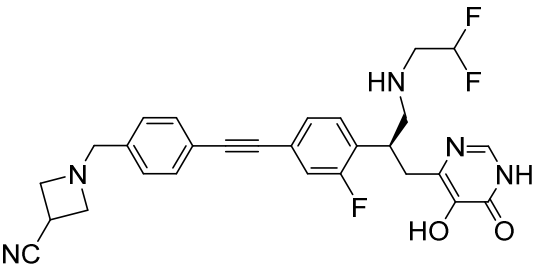
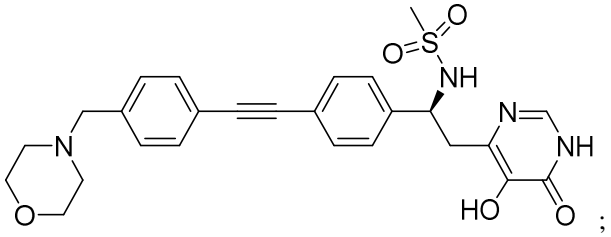
2012. A compound selected from:




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; and

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or a pharmaceutically acceptable salt~~,~~ or solvate~~, or prodrug~~ thereof.

~~21~~13. A compound selected from:


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;NC

; and

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or a pharmaceutically acceptable salt~~,~~ or solvate~~, or prodrug~~ thereof.

~~22~~14. A pharmaceutical composition comprising a compound of any one of claims 1 - ~~21~~13, or a pharmaceutically acceptable salt~~,~~ or solvate~~, or prodrug~~ thereof, and a pharmaceutically acceptable excipient.

~~23~~15. Use of a compound of any one of claims 1 - ~~21~~13, or a pharmaceutically acceptable salt or ~~,~~solvate~~, or prodrug~~ thereof, ~~or the pharmaceutical composition of claim 22,~~ in the manufacture of a medicament for the ~~treating~~ treatment of a gram-negative bacterial infection in a patient in need thereof.

~~24~~16. The use of claim ~~23~~15, wherein~~:~~

the gram-negative bacterial infection is selected from pneumonia, sepsis, cystic fibrosis, intra-abdominal infection, skin infection ~~and~~ urinary tract infection,~~;~~

~~or the gram-negative bacterial infection is selected from~~ chronic urinary tract infection, complicated urinary tract infection, cystitis, pyelonephritis, urethritis, recurrent urinary tract infections, bladder infections, urethral infections and kidney infections~~;~~.

~~or the gram-negative bacterial infection is chronic urinary tract infections;~~

~~or the gram-negative bacterial infection is complicated urinary tract infections.~~

~~25~~17. The compound of any one of claims 1 - ~~21~~13, or a pharmaceutically acceptable salt~~,~~ or solvate~~, or prodrug~~ thereof, or the pharmaceutical composition of claim ~~22~~14, for use in therapy.