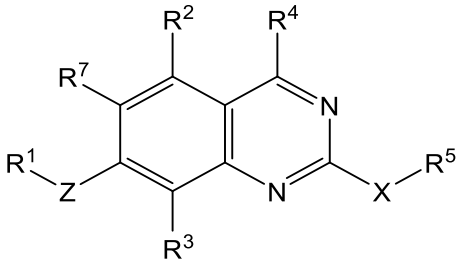
WHAT IS CLAIMED IS:

1. A compound of formula (I):



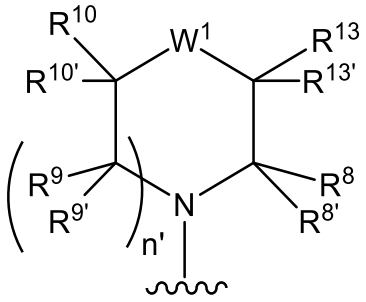
(I);

or a pharmaceutically acceptable salt thereof, wherein:

Z is a bond, O, $NR^e$ or $CR^eR^f$, wherein $R^e$ and $R^f$ are independently hydrogen or $C_1$-$C_3$alkyl;

$R^1$ is aryl or heteroaryl, wherein the aryl and the heteroaryl are optionally substituted with one, two, three, four, or five substituents independently selected from the group consisting of $C_2$-$C_4$alkenyl, $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, $C_2$-$C_4$alkynyl, $C_2$-$C_4$alkynyloxy, amino, amino$C_1$-$C_3$alkyl, cyano, cyano$C_1$-$C_3$alkoxy, $C_3$-$C_8$cycloalkyl optionally substituted with one, two, or three halo groups, halo, halo$C_1$-$C_3$alkyl, halo$C_1$-$C_3$alkoxy, hydroxy, hydroxy$C_1$-$C_3$alkyl, heteroaryl, heterocyclyl, and phenyl, wherein the heteroaryl, heterocyclyl, and phenyl are optionally substituted with one, two, or three substituents independently selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, halo, and halo$C_1$-$C_3$alkyl;

$R^2$, $R^3$, and $R^7$ are independently selected from the group consisting of hydrogen, $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, cyano, halo, halo$C_1$-$C_3$alkyl, -C(O)$NH_2$, -C(O)NH($C_1$-$C_3$alkyl), -C(O)N($C_1$-$C_3$alkyl)$_2$, and hydroxy;

$R^4$ is $-NHR^{50}$ or  ;

wherein

$R^{50}$ is a five membered ring optionally containing one or two heteroatoms independently selected from the group consisting of nitrogen and oxygen, wherein the ring is optionally substituted with one or two groups independently selected from $C_1$-$C_3$alkyl and oxo;

n′ is 0, 1, 2, or 3;

$R^8$, $R^{8'}$, $R^9$, $R^{9'}$, $R^{10}$, $R^{10'}$, $R^{13}$, and $R^{13'}$ are each independently selected from the group consisting of hydrogen, $C_1$-$C_3$alkoxy, $C_1$-$C_6$alkoxy$C_1$-$C_6$alkyl, $C_1$-$C_3$alkyl, $C_3$-$C_6$cycloalkyl, cyano, halo, halo$C_1$-$C_3$alkyl, hydroxy, and hydroxy$C_1$-$C_3$alkyl; or

$R^8$ and $R^9$, together form a $C_1$-$C_3$alkylene; or

$R^8$ and $R^{10}$ together form $C_1$-$C_3$alkylene; or

$R^8$ and $R^{13}$ together form $C_1$-$C_3$alkylene; or

$R^9$ and $R^{13}$ together form $C_1$-$C_3$alkylene; or

$R^{10}$ and $R^{13}$ together form $C_1$-$C_3$alkylene; or

$R^8$ and $R^{8'}$, together with the atoms to which they are attached, form a three-, four- or five-membered ring optionally containing an oxygen atom or $SO_2$ group, wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy; or

$R^9$ and $R^{9'}$, together with the atoms to which they are attached, form a three-, four- or five-membered ring optionally containing an oxygen atom or $SO_2$ group, wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy; or

$R^{10}$ and $R^{10'}$, together with the atoms to which they are attached, form a three-, four- or five- membered ring optionally containing an oxygen atom or $SO_2$ group, wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy; or

$R^{13}$ and $R^{13'}$, together with the atoms to which they are attached, form a three-, four- or five- membered ring optionally containing an oxygen atom or $SO_2$ group, wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy;

$W^1$ is $CR^{11}R^{12}$, $NR^{17}$, $NR^{15''}C(O)$, $N(C(O)(CH_2)_nOR^{15})$, O, $SO_2$, $SO_2NR^{15'}$, or $P(O)CH_3$; wherein

n is 0 or 1;

$R^{11}$ and $R^{12}$ are independently selected from the group consisting of hydrogen, $C_1$-$C_3$alkoxy, $C_1$-$C_6$alkoxy$C_1$-$C_6$alkyl, $C_1$-$C_3$alkyl, cyano, dimethylphosphino; dimethylsulfonamide, halo, hydroxy, and methylsulfonyl; or

$R^{11}$ and $R^{12}$, together with the atoms to which they are attached, form a four- to six-membered ring optionally containing an oxygen atom or $SO_2$ group, wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy; or

$R^{11}$ and $R^{13}$, together with the atoms to which they are attached, form a three-, four- or five-membered ring optionally containing one or two heteroatoms independently selected from nitrogen and oxygen, wherein the ring optionally contains one or two double bonds, and wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy; or

$R^{13}$ and $R^{15}$, together form $CH_2$;

$R^{15'}$ is $C_1$-$C_3$alkyl or $C_1$-$C_6$alkoxy$C_1$-$C_6$alkyl;

$R^{15''}$ is hydrogen or $C_1$-$C_3$alkyl;

$R^{17}$ is selected from the group consisting of $C_1$-$C_3$alkylcarbonyl, $C_3$-$C_6$cycloalkylcarbonyl, halo$C_1$-$C_3$alkylcarbonyl, methylsulfonyl, and tetrahydropyranylcarbonyl, wherein the $C_3$-$C_6$cycloalkyl and the tetrahydropyranyl are optionally substituted with one or two substituents independently selected from the group consisting of cyano, halo, and hydroxy;

X is O or $NR^{16}$, wherein $R^{16}$ is hydrogen or $C_1$-$C_3$alkyl;

$R^5$ is selected from the group consisting of hydrogen, $C_1$-$C_6$alkoxy$C_1$-$C_6$alkyl, $C_1$-$C_6$alkyl, aryl, aryl$C_1$-$C_6$alkyl, carboxy$C_1$-$C_6$alkyl, $C_3$-$C_6$cycloalkyl, $C_3$-$C_6$cycloalkyl$C_1$-$C_6$alkyl, di($C_1$-$C_3$alkyl)amino$C_2$-$C_6$alkyl, halo$C_1$-$C_6$alkyl, heteroaryl, heteroaryl$C_1$-$C_6$alkyl, heterocyclyl, heterocyclyl$C_1$-$C_6$alkyl, hydroxy$C_1$-$C_6$alkyl, $NR^aR^b$-C(O)-$C_1$-$C_6$alkyl), $NR^aR^b$$C_1$-$C_6$alkyl, wherein the aryl, the aryl part of the aryl$C_1$-$C_6$alkyl, the $C_3$-$C_6$cycloalkyl, the cycloalkyl part of the $C_3$-$C_6$cycloalkyl$C_1$-$C_6$alkyl, the heteroaryl, the heteroaryl part of the heteroaryl$C_1$-$C_6$alkyl, the heterocyclyl, the heterocyclyl part of the heterocyclyl$C_1$-$C_6$alkyl, are optionally substituted with one, two, three, or four groups independently selected from $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, deuterated $C_1$-$C_3$alkyl, $C_3$-$C_6$cycloalkyl, ($C_1$-$C_6$alkyl)amino, ($C_1$-$C_6$alkyl)amino$C_1$-$C_3$alkyl, amino, amino$C_1$-$C_3$alkyl, carboxy, cyano, di($C_1$-$C_6$alkyl)amino, di($C_1$-$C_6$alkyl)amino$C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkoxy, halo$C_1$-$C_3$alkyl, heterocyclyl, heterocyclyl$C_1$-$C_3$alkyl, hydroxy, hydroxy$C_1$-$C_3$alkyl, nitro, and oxo; wherein the heterocyclyl and the heterocyclyl part of the heterocyclyl$C_1$-$C_3$alkyl is further optionally substituted with one, two, or three groups independently selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, halo, and halo$C_1$-$C_3$alkyl; or

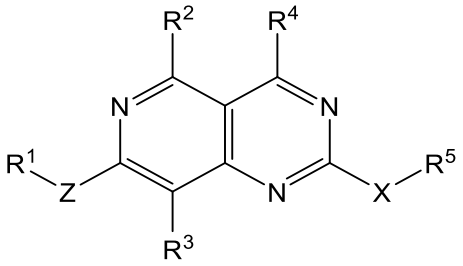
$R^5$ and $R^{16}$, together with the nitrogen atom to which they are attached, form a heterocyclic group optionally substituted with one, two, three, four, or five groups independently selected from the group consisting of one, two, three, or four groups independently selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkoxyalkyl, $C_1$-$C_3$alkyl, amino, amino$C_1$-$C_3$alkyl, hydroxy, and hydroxy$C_1$-$C_3$alkyl; and

one of $R^a$ and $R^b$ is selected from the group consisting of hydrogen and $C_1$-$C_3$alkyl and the other is selected from the group consisting of hydrogen, $C_1$-$C_3$alkyl, $C_1$-$C_3$alkoxycarbonyl, $C_1$-$C_3$alkylcarbonyl, aryl$C_1$-$C_6$alkyl, $C_3$-$C_6$cycloalkyl, and $C_3$-$C_6$cycloalkyl$C_1$-$C_6$alkyl.

2. The compound of claim 1, or a pharmaceutically acceptable salt thereof, wherein $R^7$ is chloro.

3. The compound of claim 1, or a pharmaceutically acceptable salt thereof, wherein $R^7$ is hydrogen.

4. A compound of formula (II):



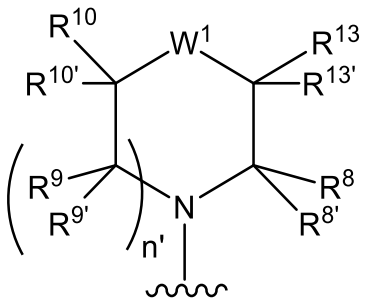
(II);

or a pharmaceutically acceptable salt thereof, wherein:

Z is a bond, O, $NR^e$ or $CR^eR^f$, wherein $R^e$ and $R^f$ are independently hydrogen or $C_1$-$C_3$alkyl;

$R^1$ is aryl or heteroaryl, wherein the aryl and the heteroaryl are optionally substituted with one, two, three, four, or five substituents independently selected from the group consisting of $C_2$-$C_4$alkenyl, $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, $C_2$-$C_4$alkynyl, $C_2$-$C_4$alkynyloxy, amino, amino$C_1$-$C_3$alkyl, cyano, cyano$C_1$-$C_3$alkoxy, $C_3$-$C_8$cycloalkyl optionally substituted with one, two, or three halo groups, halo, halo$C_1$-$C_3$alkyl, halo$C_1$-$C_3$alkoxy, hydroxy, hydroxy$C_1$-$C_3$alkyl, heteroaryl, heterocyclyl, and phenyl, wherein the heteroaryl, heterocyclyl, and phenyl are optionally substituted with one, two, or three substituents independently selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, halo, and halo$C_1$-$C_3$alkyl;

$R^2$ and $R^3$ are independently selected from the group consisting of hydrogen, $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, cyano, halo, halo$C_1$-$C_3$alkyl, -C(O)$NH_2$, -C(O)NH($C_1$-$C_3$alkyl), -C(O)N($C_1$-$C_3$alkyl)$_2$, and hydroxy;

$R^4$ is $-NHR^{50}$ or  ;

wherein

$R^{50}$ is a five membered ring optionally containing one or two heteroatoms independently selected from the group consisting of nitrogen and oxygen, wherein the ring is optionally substituted with one or two groups independently selected from $C_1$-$C_3$alkyl and oxo;

n′ is 0, 1, 2, or 3;

$R^8$, $R^{8'}$, $R^9$, $R^{9'}$, $R^{10}$, $R^{10'}$, $R^{13}$, and $R^{13'}$ are each independently selected from the group consisting of hydrogen, $C_1$-$C_3$alkoxy, $C_1$-$C_6$alkoxy$C_1$-$C_6$alkyl, $C_1$-$C_3$alkyl, $C_3$-$C_6$cycloalkyl, cyano, cyano$C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, hydroxy, and hydroxy$C_1$-$C_3$alkyl; or

$R^8$ and $R^9$, together form a $C_1$-$C_3$alkylene; or

$R^8$ and $R^{10}$ together form $C_1$-$C_3$alkylene; or

$R^8$ and $R^{13}$ together form $C_1$-$C_3$alkylene; or

$R^9$ and $R^{13}$ together form $C_1$-$C_3$alkylene; or

$R^{10}$ and $R^{13}$ together form $C_1$-$C_3$alkylene; or

$R^8$ and $R^{8'}$, together with the atoms to which they are attached, form a three-, four- or five-membered ring optionally containing an oxygen atom or $SO_2$ group, wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy; or

$R^9$ and $R^{9'}$, together with the atoms to which they are attached, form a three-, four- or five-membered ring optionally containing an oxygen atom or $SO_2$ group, wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy; or

$R^{10}$ and $R^{10'}$, together with the atoms to which they are attached, form a three-, four- or five- membered ring optionally containing an oxygen atom or $SO_2$ group, wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy; or

$R^{13}$ and $R^{13'}$, together with the atoms to which they are attached, form a three-, four- or five- membered ring optionally containing an oxygen atom or $SO_2$ group, wherein the ring is

optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy;

$W^1$ is $CR^{11}R^{12}$, $NR^{17}$, $NR^{15''}C(O)$, $N(C(O)(CH_2)_nOR^{15})$, O, $SO_2$, $SO_2NR^{15'}$, or $P(O)CH_3$; wherein

n is 0 or 1;

$R^{11}$ and $R^{12}$ are independently selected from the group consisting of hydrogen, $C_1$-$C_3$alkoxy, $C_1$-$C_6$alkoxy$C_1$-$C_6$alkyl, $C_1$-$C_3$alkyl, cyano, dimethylphosphino; dimethylsulfonamide, halo, hydroxy, and methylsulfonyl; or

$R^{11}$ and $R^{12}$, together with the atoms to which they are attached, form a four- or five-membered ring optionally containing an oxygen atom or $SO_2$ group, wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy; or

$R^{11}$ and $R^{13}$, together with the atoms to which they are attached, form a three-, four- or five-membered ring optionally containing one or two heteroatoms independently selected from nitrogen and oxygen, wherein the ring optionally contains one or two double bonds, and wherein the ring is optionally substituted with two groups selected from the group consisting of $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, and hydroxy; or

$R^{13}$ and $R^{15}$, together form $CH_2$;

$R^{15'}$ is $C_1$-$C_3$alkyl or $C_1$-$C_6$alkoxy$C_1$-$C_6$alkyl;

$R^{15''}$ is hydrogen or $C_1$-$C_3$alkyl;

$R^{17}$ is selected from the group consisting of $C_1$-$C_3$alkylcarbonyl, $C_3$-$C_6$cycloalkylcarbonyl, halo$C_1$-$C_3$alkylcarbonyl, methylsulfonyl, and tetrahydropyranylcarbonyl, wherein the $C_3$-$C_6$cycloalkyl and the tetrahydropyranyl are optionally substituted with one or two substituents independently selected from the group consisting of cyano, halo, and hydroxy;

X is O or $NR^{16}$, wherein $R^{16}$ is hydrogen or $C_1$-$C_3$alkyl;

$R^5$ is selected from the group consisting of hydrogen, $C_1$-$C_6$alkoxy$C_1$-$C_6$alkyl, $C_1$-$C_6$alkyl, aryl, aryl$C_1$-$C_6$alkyl, carboxy$C_1$-$C_6$alkyl, $C_3$-$C_6$cycloalkyl, $C_3$-$C_6$cycloalkyl$C_1$-$C_6$alkyl, di($C_1$-$C_3$alkyl)amino$C_2$-$C_6$alkyl, halo$C_1$-$C_6$alkyl, heteroaryl, heteroaryl$C_1$-$C_6$alkyl, heterocyclyl, heterocyclyl$C_1$-$C_6$alkyl, hydroxy$C_1$-$C_6$alkyl, $NR^aR^b$-C(O)-$C_1$-$C_6$alkyl), $NR^aR^b$$C_1$-$C_6$alkyl, wherein the aryl, the aryl part of the aryl$C_1$-$C_6$alkyl, the $C_3$-$C_6$cycloalkyl, the cycloalkyl part of the $C_3$-$C_6$cycloalkyl$C_1$-$C_6$alkyl, the heteroaryl, the heteroaryl part of the heteroaryl$C_1$-$C_6$alkyl, the heterocyclyl, the heterocyclyl part of the heterocyclyl$C_1$-$C_6$alkyl, are optionally substituted with one, two, three, or four groups independently selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, deuterated $C_1$-$C_3$alkyl, $C_3$-$C_6$cycloalkyl, ($C_1$-$C_6$alkyl)amino, ($C_1$-$C_6$alkyl)amino$C_1$-$C_3$alkyl, amino, amino$C_1$-$C_3$alkyl, carboxy, cyano, di($C_1$-$C_6$alkyl)amino, di($C_1$-$C_6$alkyl)amino$C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkoxy, halo$C_1$-$C_3$alkyl, heterocyclyl, heterocyclyl$C_1$-$C_3$alkyl, hydroxy, hydroxy$C_1$-$C_3$alkyl, nitro, and oxo; wherein the heterocyclyl and the heterocyclyl part of the heterocyclyl$C_1$-$C_3$alkyl is further optionally substituted with one, two, or three groups independently selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, halo, and halo$C_1$-$C_3$alkyl; or

$R^5$ and $R^{16}$, together with the nitrogen atom to which they are attached, form a heterocyclic group optionally substituted with one, two, three, four, or five groups independently selected from the group consisting of one, two, three, or four groups independently selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkoxyalkyl, $C_1$-$C_3$alkyl, amino, amino$C_1$-$C_3$alkyl, hydroxy, and hydroxy$C_1$-$C_3$alkyl; and

one of $R^a$ and $R^b$ is selected from hydrogen and $C_1$-$C_3$alkyl and the other is selected from the group consisting of hydrogen, $C_1$-$C_3$alkyl, $C_1$-$C_3$alkoxycarbonyl, $C_1$-$C_3$alkylcarbonyl, aryl$C_1$-$C_6$alkyl, $C_3$-$C_6$cycloalkyl, and $C_3$-$C_6$cycloalkyl$C_1$-$C_6$alkyl.

5. The compound of any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein $R^4$ is –$NHR^{50}$, and $R^{50}$ is a five membered ring optionally containing one or two heteroatoms independently selected from the group consisting of nitrogen and oxygen, wherein the ring is optionally substituted with one or two groups independently selected from $C_1$-$C_3$alkyl and oxo.

6. The compound of any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein $W^1$ is $CR^{11}R^{12}$.

7. The compound of any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein $W^1$ is $NR^{17}$.

8. The compound of any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein $W^1$ is $NR^{15''}(C(O)$.

9. The compound of any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein $W^1$ is $N(C(O)(CH_2)_nOR^{15})$.

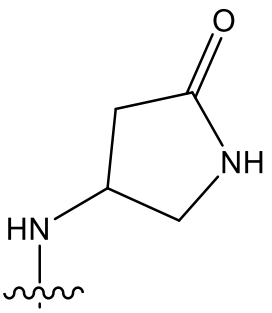
10. The compound of any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein $W^1$ is O.

11. The compound of any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein $W^1$ is $SO_2$.

12. The compound of any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein $W^1$ is $SO_2NR^{15'}$.

13. The compound of any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein $W^1$ is $P(O)CH_3$.

14. The compound of any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein $R^4$ is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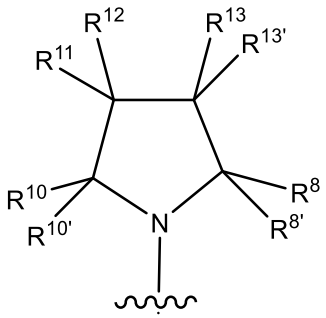
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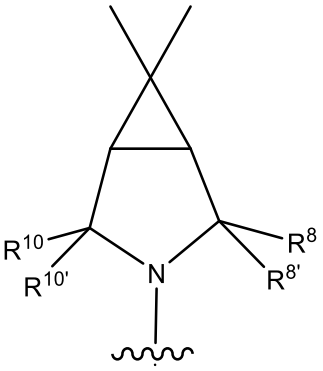
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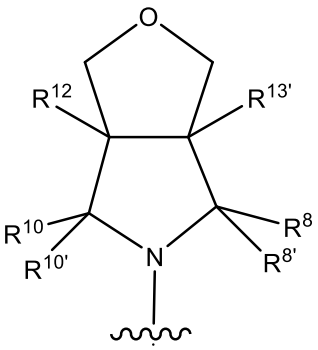
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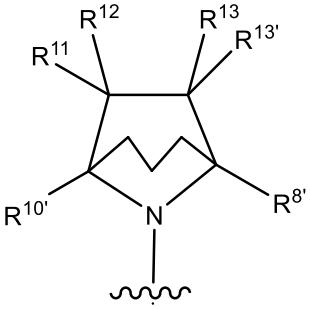
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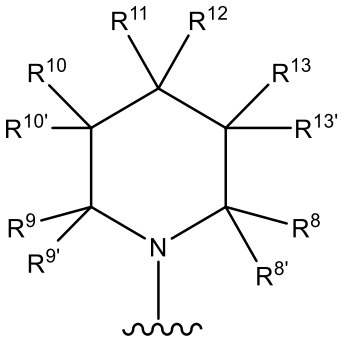
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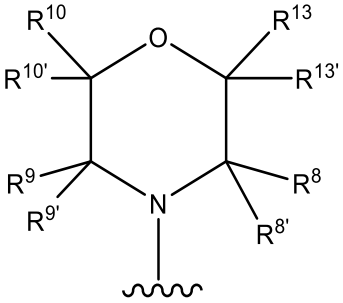
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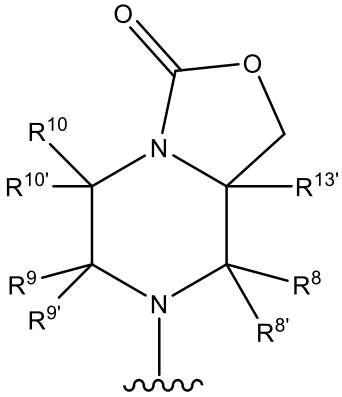
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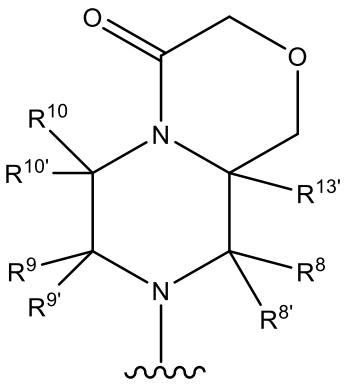
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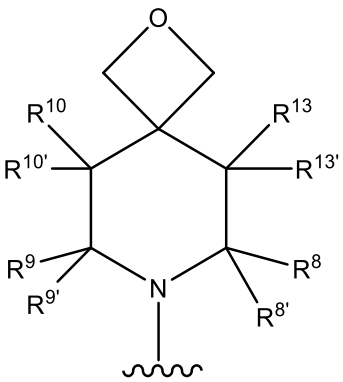
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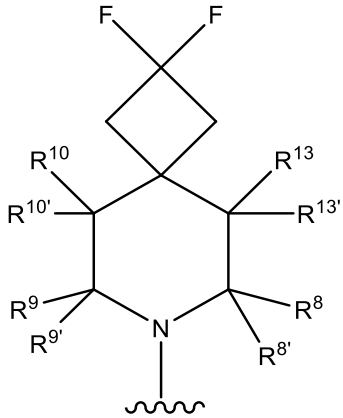
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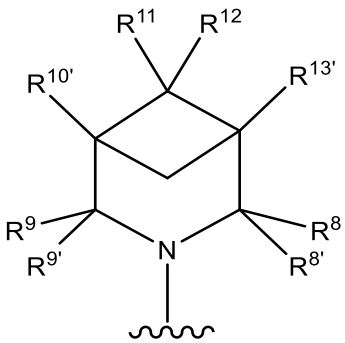
R11
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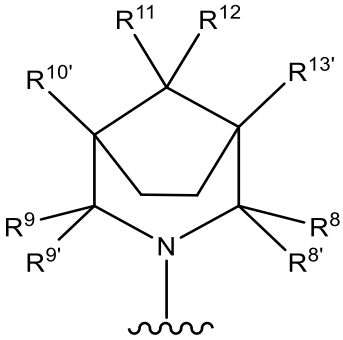
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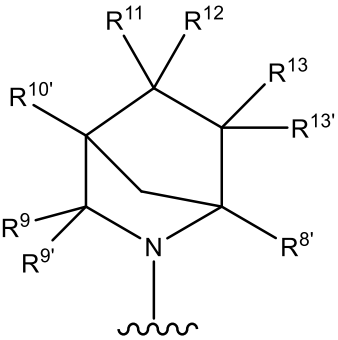
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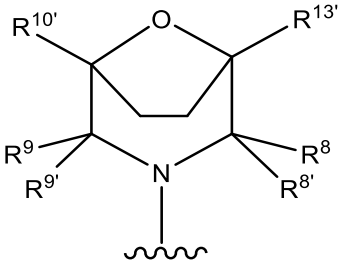
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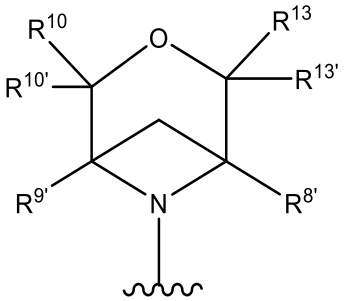
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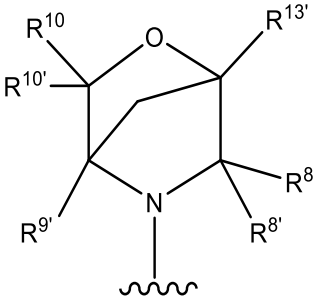
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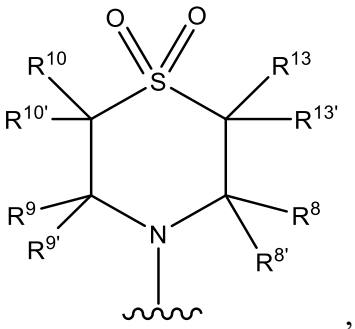
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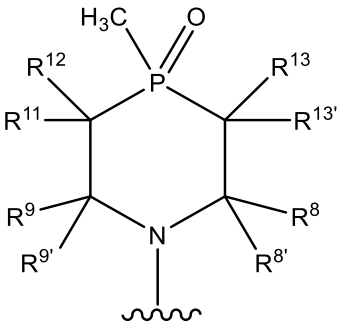
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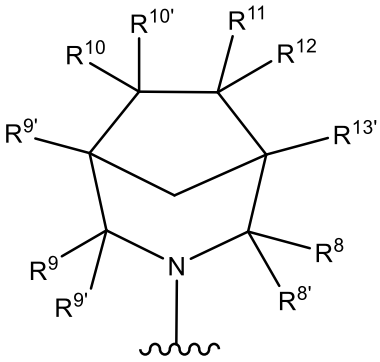
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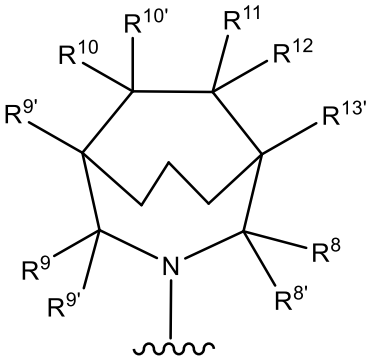
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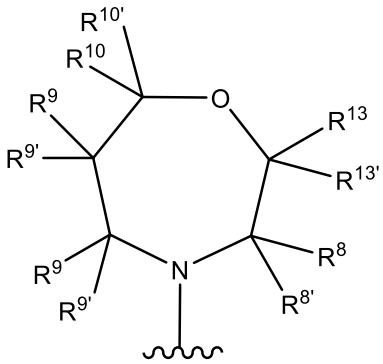
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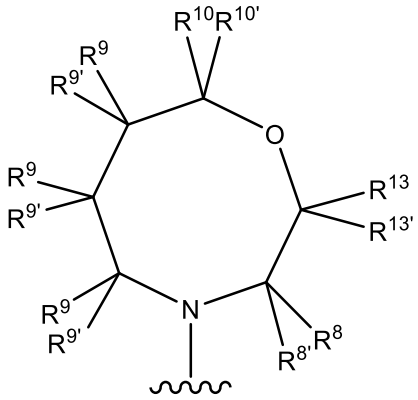
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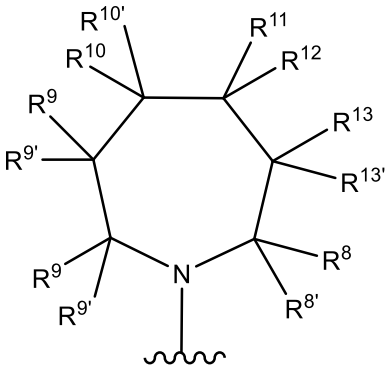
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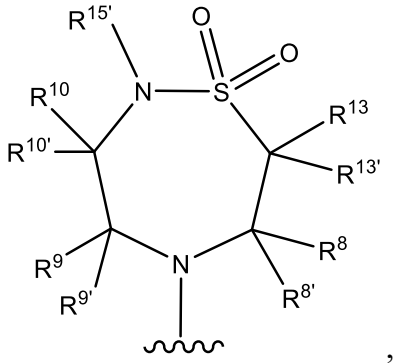

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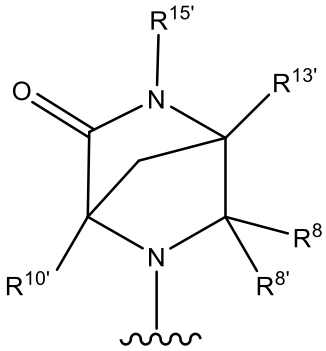

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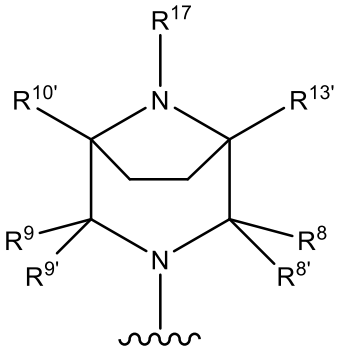

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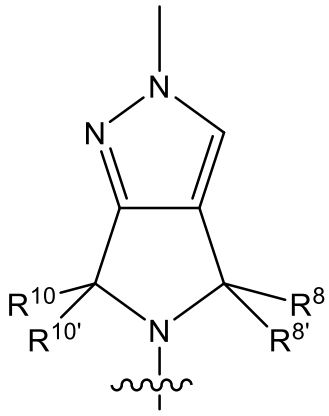
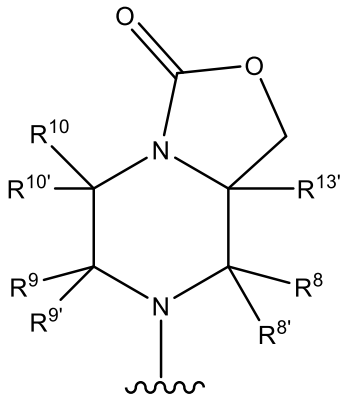


, wherein ~~~ represents the point of attachment to the core of formula (I) or (II).

15. The compound of claim 4 or 14, or a pharmaceutically acceptable salt thereof, wherein $R^4$ is



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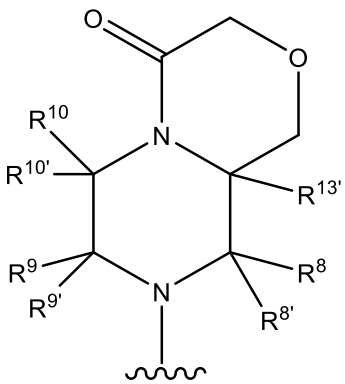

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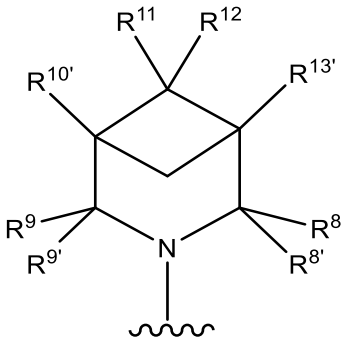

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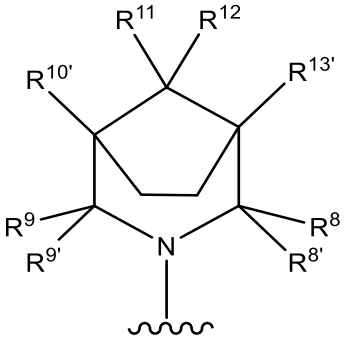

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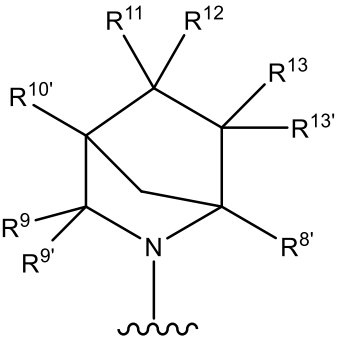

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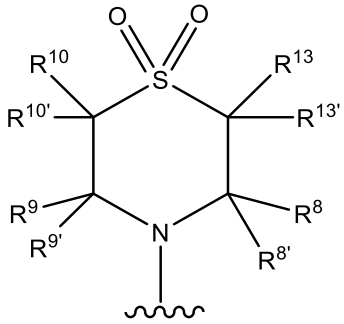

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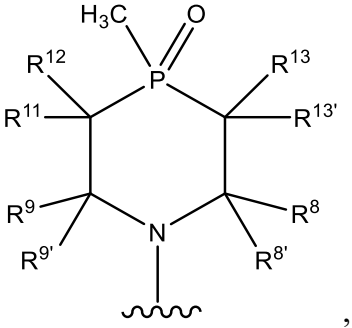

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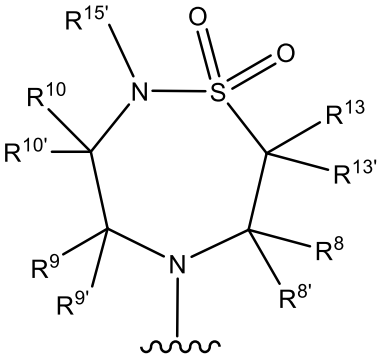

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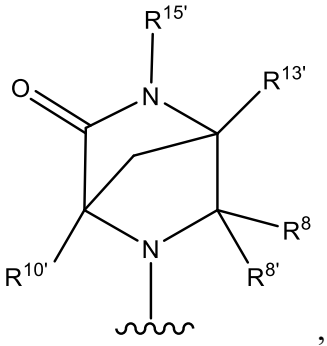

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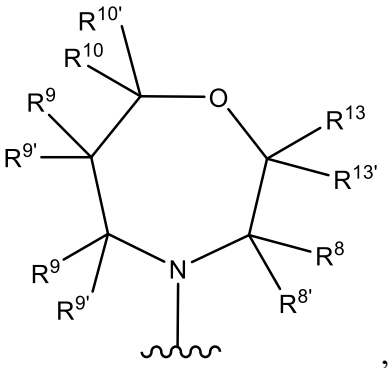

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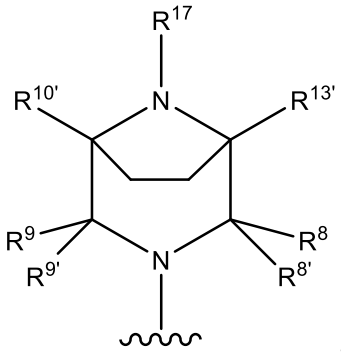


,

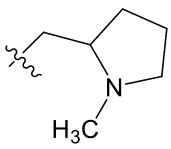
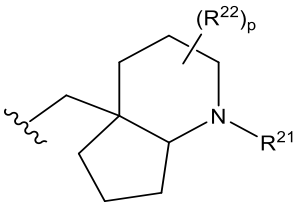
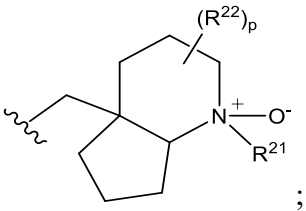
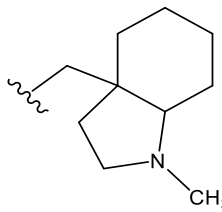
wherein ~~~ represents the point of attachment to the core of formula (II).

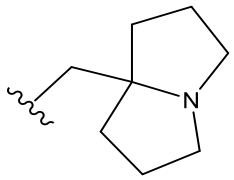
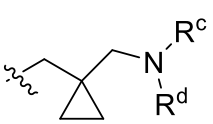
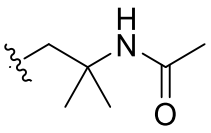
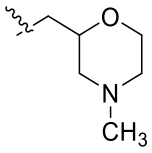
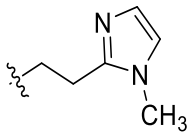
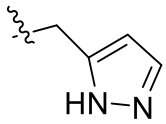
16. The compound of any one of claims 1 to 15, or a pharmaceutically acceptable salt thereof, wherein $R^2$ is hydrogen.

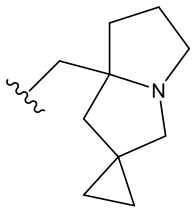
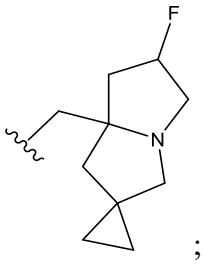
17. The compound of any one of claims 1 to 16, or a pharmaceutically acceptable salt thereof, wherein $R^3$ is halo.

18. The compound of any one of claims 1 to 17, or a pharmaceutically acceptable salt thereof, wherein X is O.

19. The compound of any one of claims 1 to 18, or a pharmaceutically acceptable salt thereof, wherein $R^5$ is selected from the group consisting of

 ;  ;  ;  ;

 ;  ;  ;  ;  ;  ;

 ; and  ;

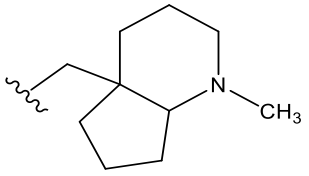
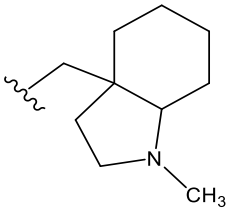
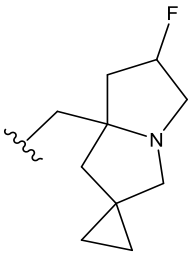
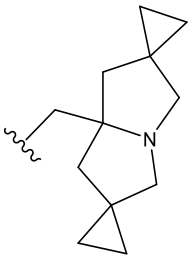
wherein each ring is optionally substituted with 1, 2, or 3 groups independently selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkoxy$C_1$-$C_3$alkyl, $C_1$-$C_3$alkyl, deuterated $C_1$-$C_3$alkyl, $C_3$-$C_6$cycloalkyl, benzyl, halo, halo$C_1$-$C_3$alkyl, hydroxy, hydroxy$C_1$-$C_3$alkyl, and oxo; and wherein

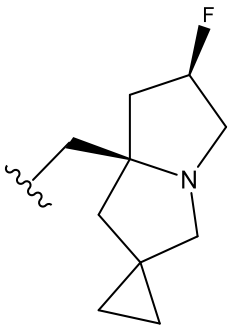
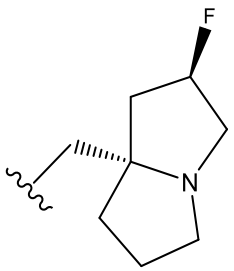
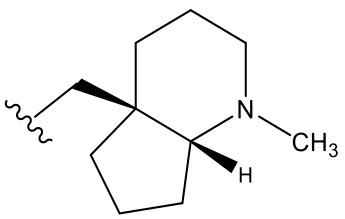
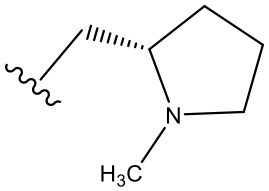
$R^c$ and $R^d$, together with the nitrogen atom to which they are attached, form a five- to ten-membered ring monocyclic or bicyclic ring optionally containing one additional heteroatom selected from nitrogen, oxygen, and sulfur, wherein the ring is optionally substituted with one, two, or three groups independently selected from $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkoxy$C_1$-$C_3$alkyl, $C_1$-$C_3$alkyl, benzyl, halo, halo$C_1$-$C_3$alkyl, hydroxy, hydroxy$C_1$-$C_3$alkyl, and oxo; or

one of $R^c$ and $R^d$ is selected from hydrogen and $C_1$-$C_3$alkyl and the other is selected from hydrogen, $C_1$-$C_3$alkyl, $C_1$-$C_3$alkoxycarbonyl, and $C_1$-$C_3$alkylcarbonyl.

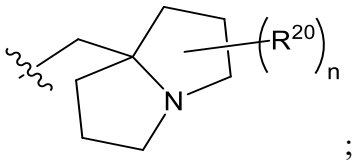
20. The compound of any one of claims 1 to 19, or a pharmaceutically acceptable salt thereof, wherein $R^5$ is –($C_1$-$C_3$alkyl)-$R^6$, wherein $R^6$ is a three- to six-membered monocyclic ring system, an eight- or nine-membered bicyclic fused saturated ring system, a ten-membered tricyclic saturated ring system, or a twelve-membered tetracyclic saturated ring system, wherein each ring system optionally contains one or more nitrogen, oxygen and/or sulfur atoms, and wherein each ring system is optionally substituted with one to four groups independently selected from the group consisting of $C_1$-$C_3$alkyl, halo, oxo, and (4- to 6-membered heterocyclyl)$C_1$-$C_3$alkyl; wherein the

heterocyclyl part of the (4- to 6-membered heterocyclyl)$C_1$-$C_3$alkyl is further optionally substituted with a halo group.

21. The compound of claim 19 or 20, or a pharmaceutically acceptable salt thereof, wherein $R^5$ is  ;  ;  ;  ; or  , and ~~~ represents the point of attachment to X.

22. The compound of any one of claims 1 to 21, or a pharmaceutically acceptable salt thereof, wherein $R^5$ is  ;  ;  ; or  ; and ~~~ represents the point of attachment to X.

23. The compound of any one of claims 1 to 20, or a pharmaceutically acceptable salt thereof, wherein $R^5$ is



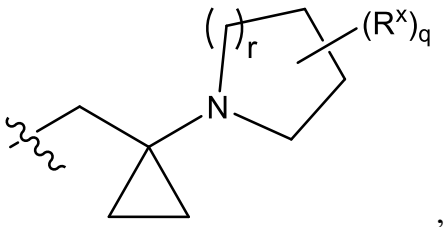
;

wherein n is 0, 1, or 2;

each $R^{20}$ is halo; and

~~~ represents the point of attachment to X.

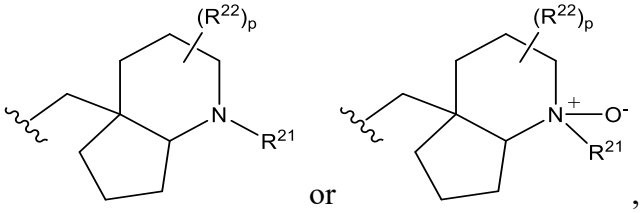
24. The compound of any one of claims 1 to 20, or a pharmaceutically acceptable salt thereof, wherein $R^5$ is




,

wherein r is 1 or 2; q is 0, 1, or 2; $R^x$ is selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkoxy$C_1$-$C_3$alkyl, $C_1$-$C_3$alkyl, benzyl, halo, halo$C_1$-$C_3$alkyl, hydroxy, hydroxy$C_1$-$C_3$alkyl, and oxo, and ~~~ denotes the point of attachment to X.

25. The compound of any one of claims 1 to 20, or a pharmaceutically acceptable salt thereof, wherein $R^5$ is

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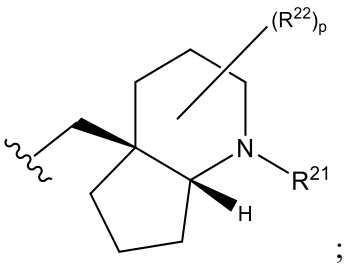
wherein $R^{21}$ is selected from the group consisting of $C_1$-$C_3$alkyl, deuterated $C_1$-$C_3$alkyl, and $C_3$-$C_6$cycloalkyl, $R^{22}$ is halo; p is 0 or 1; and ~~~ denotes the point of attachment to X.

26. The compound of any one of claims 1 to 20, or a pharmaceutically acceptable salt thereof, wherein $R^5$ is




; ;
~~~

wherein

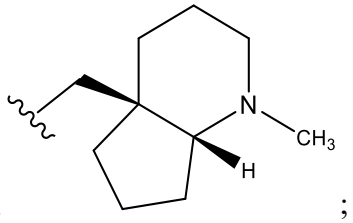
~~~ represents the point of attachment to X;

p is 0 or 1;

$R^{21}$ is selected from the group consisting of $C_1$-$C_3$alkyl, deuterated $C_1$-$C_3$alkyl, and $C_3$-$C_6$cycloalkyl; and

$R^{22}$ is halo; p is 0 or 1; and ~~~ denotes the point of attachment to X.

27. The compound of any one of claims 1 to 20, or a pharmaceutically acceptable salt thereof, wherein $R^5$ is




; ;

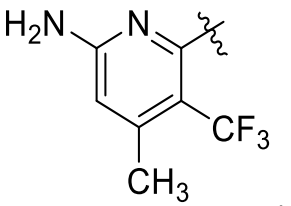
wherein

~~~ represents the point of attachment to X.

28. The compound of any one of claims 1 to 27, or a pharmaceutically acceptable salt thereof, wherein Z is a bond.

29. The compound of any one of claims 1 to 28, wherein $R^1$ is a monocyclic heteroaryl ring containing one, two, or three nitrogen atoms, wherein the ring is optionally substituted with one, two, three, four, or five substituents independently selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, $C_2$-$C_4$alkenyl, $C_2$-$C_4$alkynyl, amino, amino$C_1$-$C_3$alkyl, cyano, $C_3$-$C_4$cycloalkyl, halo, halo$C_1$-$C_3$alkyl, halo$C_1$-$C_3$alkoxy, hydroxy, and hydroxy$C_1$-$C_3$alkyl.

30. The compound of any one of claims 1 to 29, or a pharmaceutically acceptable salt thereof, wherein $R^1$ is

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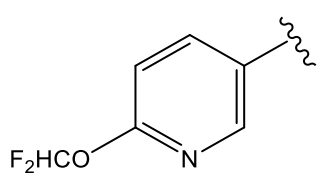


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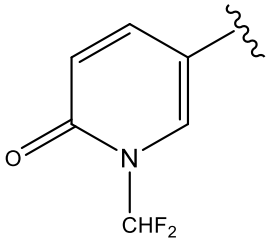



;

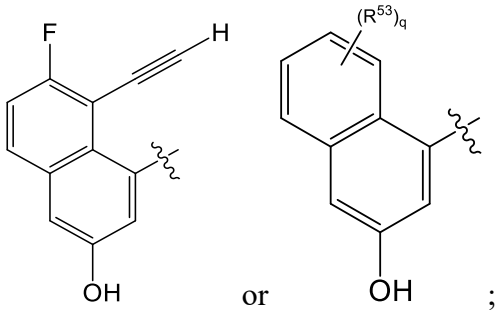
wherein ~~~ denotes the point of attachment to the core of formula (I) or (II).

31. The compound of any one of claims 1 to 28, or a pharmaceutically acceptable salt thereof, wherein $R^1$ is $C_6$-$C_{10}$aryl optionally substituted with one, two, three, four, or five substituents independently selected from the group consisting of $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, $C_2$-$C_4$alkenyl, $C_2$-$C_4$alkynyl, amino, amino$C_1$-$C_3$alkyl, cyano, $C_3$-$C_5$cycloalkyl, halo, halo$C_1$-$C_3$alkyl, hydroxy, and hydroxy$C_1$-$C_3$alkyl.

32. The compound of any one of claims 1 to 28, or a pharmaceutically acceptable salt thereof, wherein $R^1$ is aryl or heteroaryl, wherein the aryl and heteroaryl are optionally substituted with one, two, three, four, or five substituents independently selected from the group consisting of $C_2$-$C_4$alkenyl, $C_1$-$C_3$alkoxy, $C_1$-$C_3$alkyl, $C_2$-$C_4$alkynyl, $C_2$-$C_4$alkynyloxy, amino, cyano, cyano$C_1$-$C_3$alkoxy, $C_3$-$C_5$cycloalkyl optionally substituted with one or two halo groups, halo, halo$C_1$-$C_3$alkyl, halo$C_1$-$C_3$alkoxy, 4- to 6-membered heterocyclyl, and hydroxy.

33. The compound of any one of claims 1 to 27 and 32, or a pharmaceutically acceptable salt thereof, wherein $R^1$ is naphthyl, wherein the naphthyl is substituted with one, two, or three groups independently selected from the group consisting of $C_2$-$C_4$alkynyl, halo, and hydroxy.

34. The compound of any one of claims 1 to 29 and 31 to 33, or a pharmaceutically acceptable salt thereof, wherein $R^1$ is




;

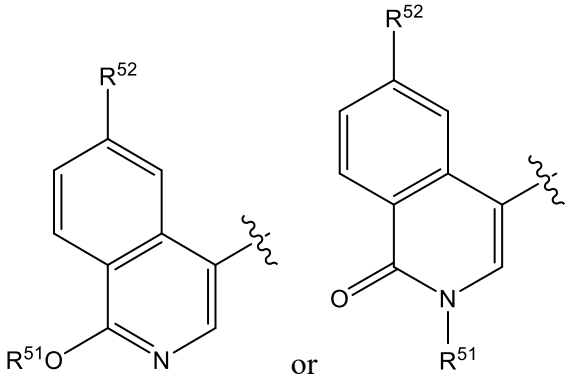
wherein $R^{53}$ is $C_1$-$C_3$alkyl, halo, halo$C_1$-$C_3$alkyl, halo$C_1$-$C_3$alkoxy, a 4- to 6-membered heterocyclyl, or hydroxy; q is an integer of 0 to 4; and ~~~ denotes the point of attachment to the core of formula (I) or (II).

35. The compound of any one of claims 1 to 28, and 32, or a pharmaceutically acceptable salt thereof, wherein $R^1$ is phenyl, wherein the phenyl is substituted with one, two, or three groups

independently selected from the group consisting of $C_1$-$C_3$alkyl, $C_3$-$C_5$cycloalkyl, heterocyclyl, and hydroxy.

36. The compound of any one of claims 1 to 28, and 32 or a pharmaceutically acceptable salt thereof, wherein $R^1$ is isoquinolinyl, wherein the isoquinolinyl is substituted with one, two, or three groups independently selected from the group consisting of halo$C_1$-$C_3$alkyl and halo$C_1$-$C_3$alkoxy.

37. The compound of claim 36, or a pharmaceutically acceptable salt thereof, wherein $R^1$ is



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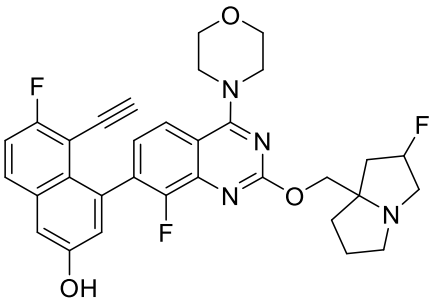
wherein $R^{51}$ is halo$C_1$-$C_3$alkyl; $R^{52}$ is hydrogen or halo$C_1$-$C_3$alkyl; and

 denotes the point of attachment to the core of formula (I) or (II).

38. The compound of any one of claims 1 to 28, and 32, or a pharmaceutically acceptable salt thereof, wherein $R^1$ is indolyl substituted with halo$C_1$-$C_3$alkyl.

39. The compound of any one of claims 1 to 28, and 32, or a pharmaceutically acceptable salt thereof, wherein $R^1$ is indazolyl substituted with one, two, or three substituents selected from $C_1$-$C_3$alkyl and halo.

40. The compound of claim 1 that is selected from the group consisting of: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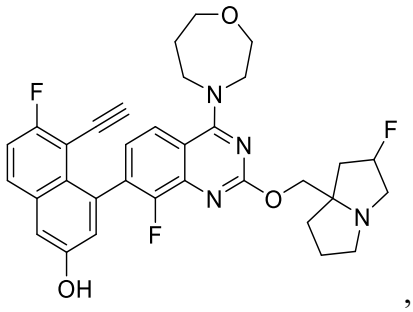
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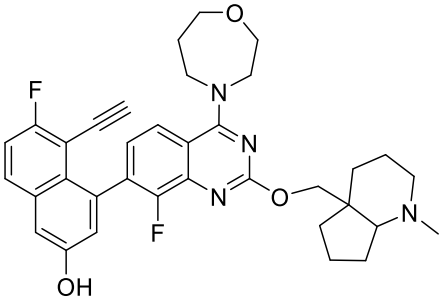
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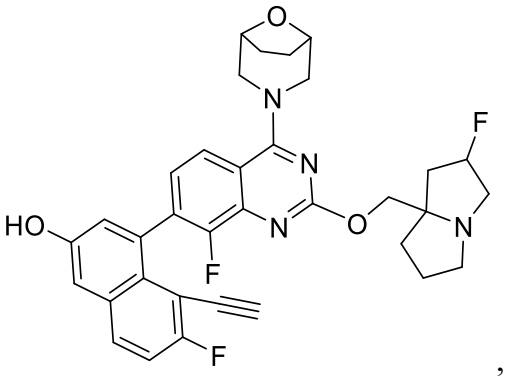
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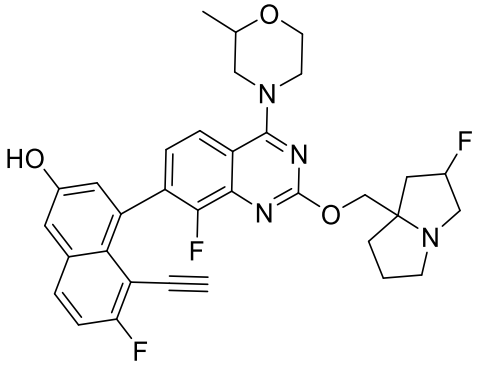
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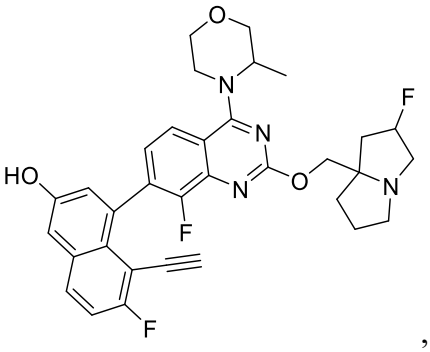
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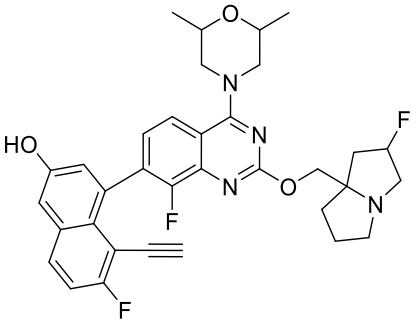
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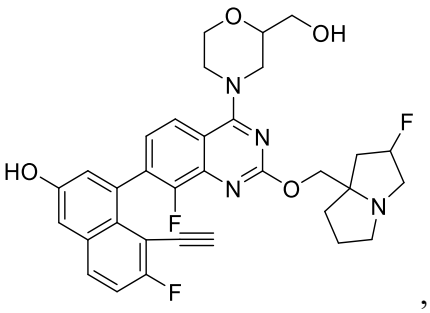
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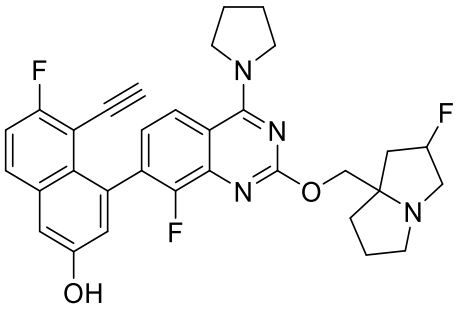
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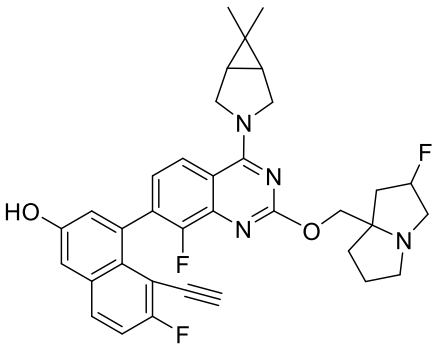
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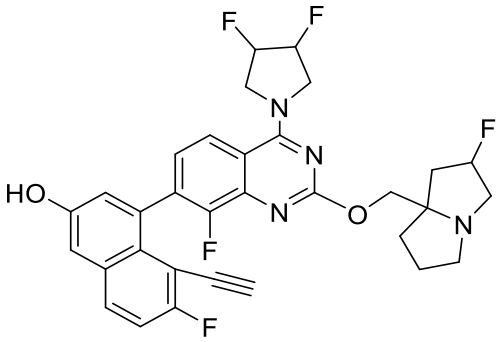
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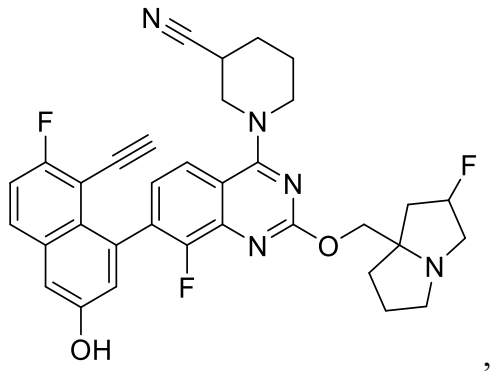
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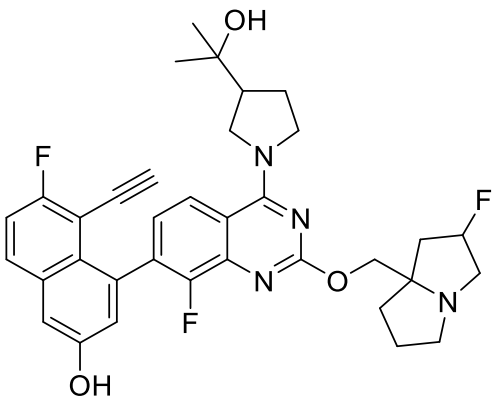
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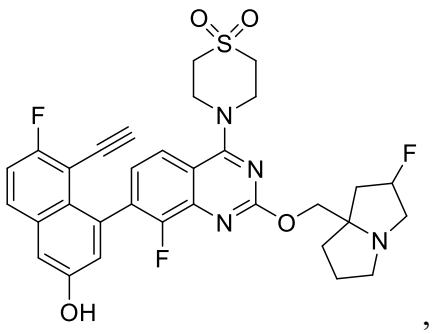
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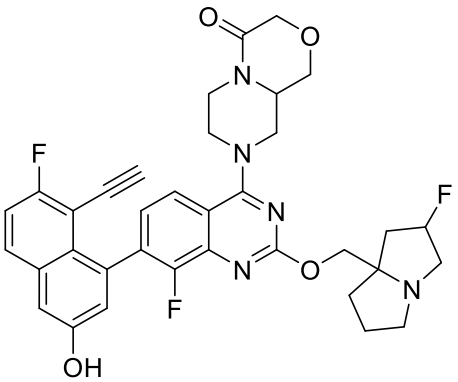
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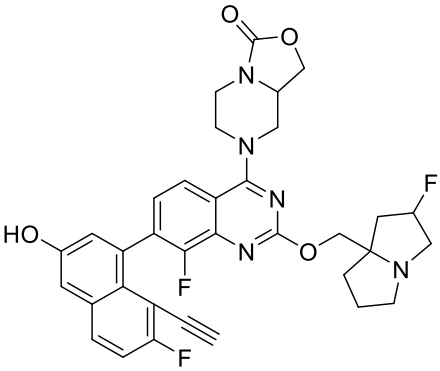
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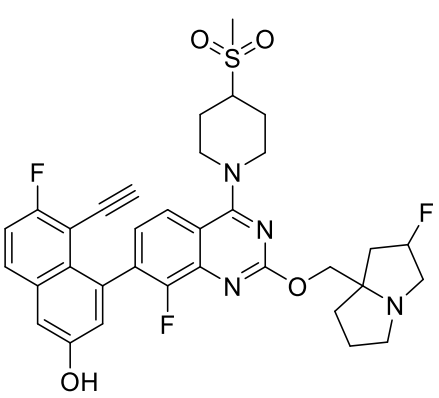
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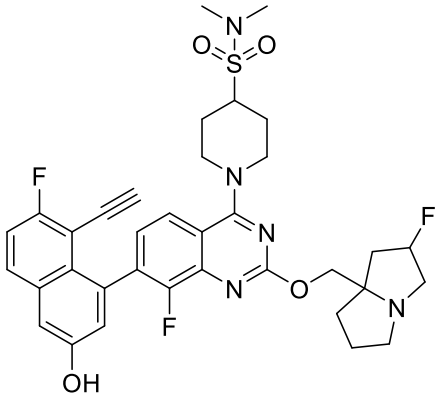
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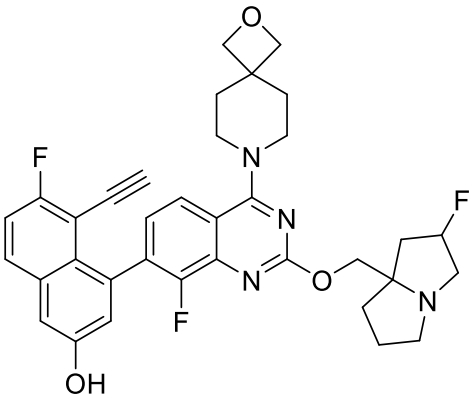
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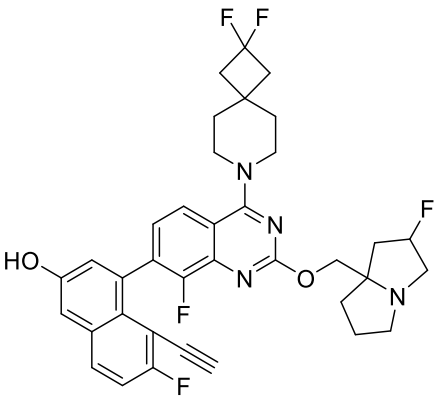
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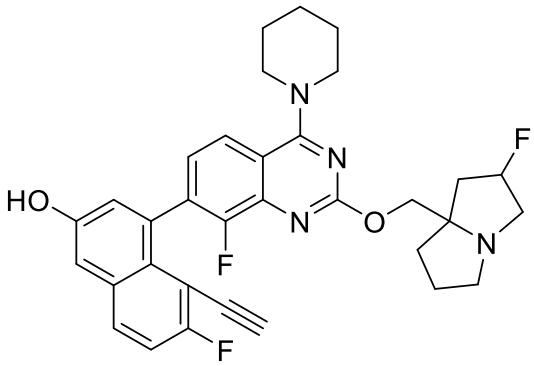
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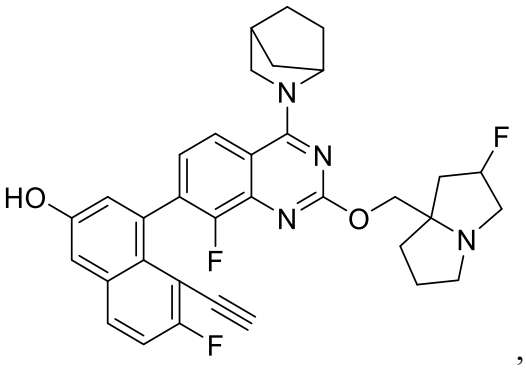
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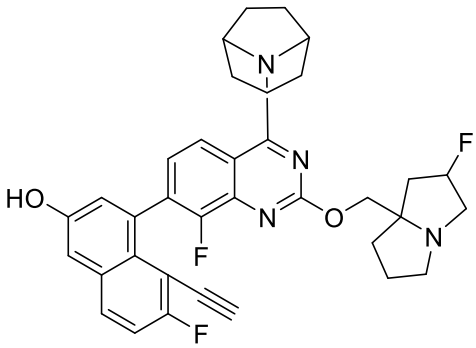
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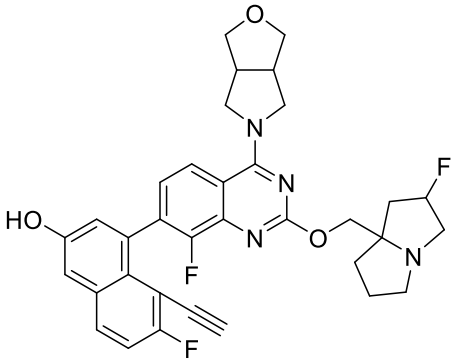
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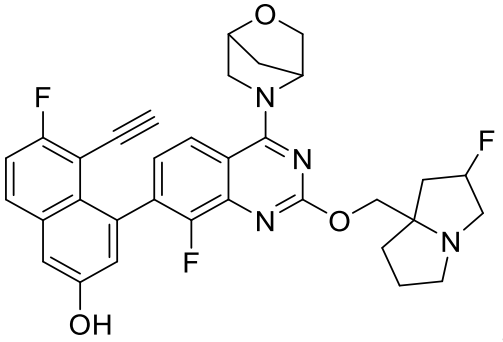
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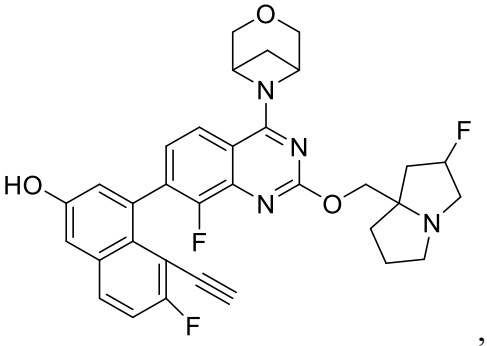
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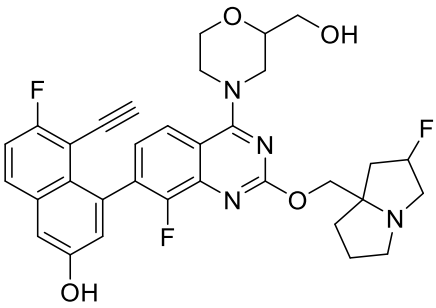
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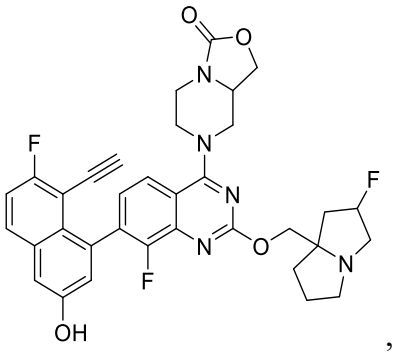
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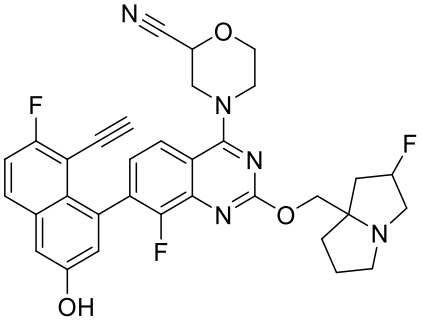
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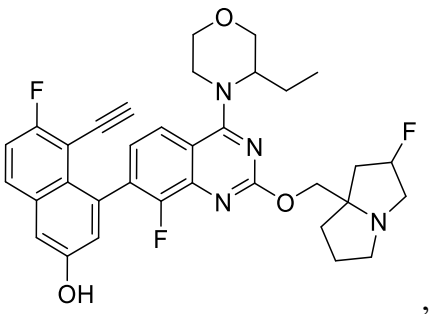
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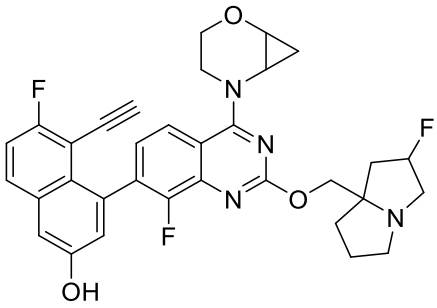
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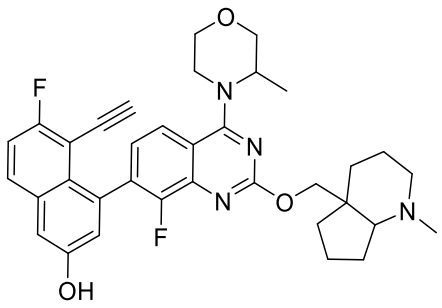
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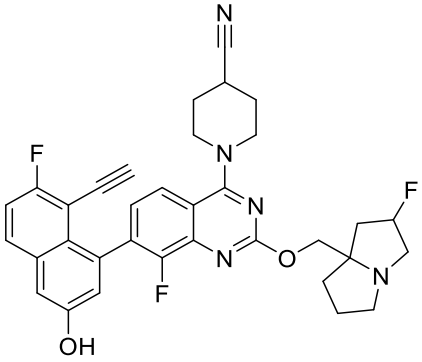
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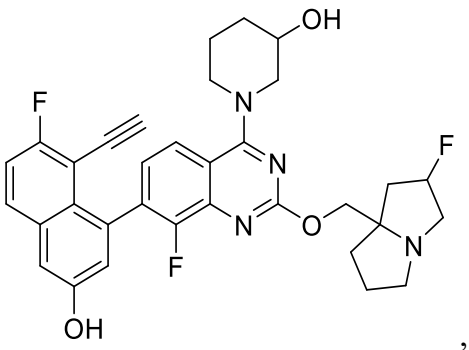
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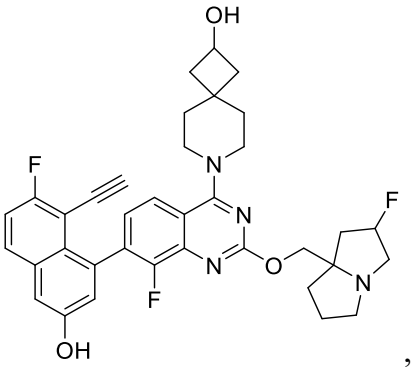
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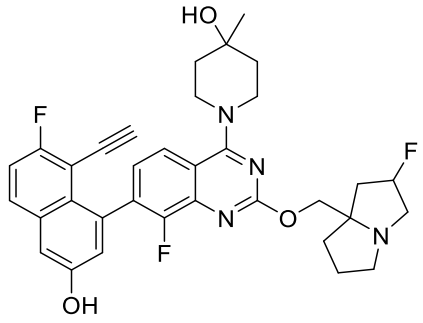
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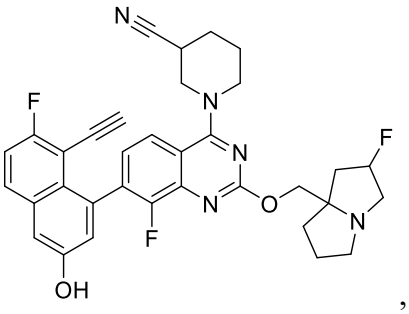
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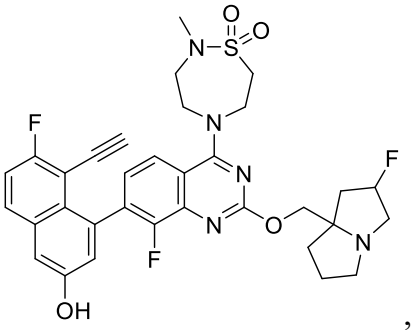
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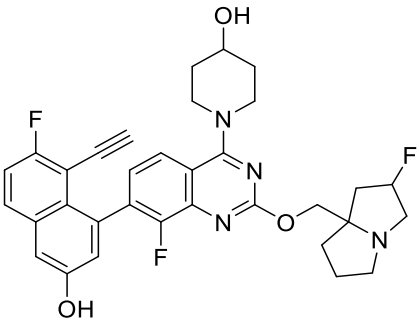
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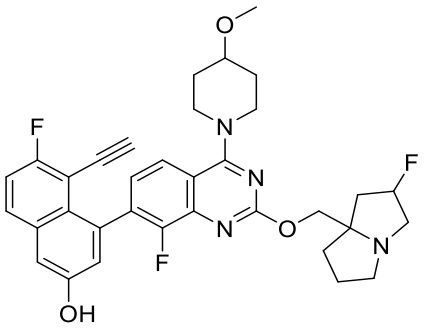
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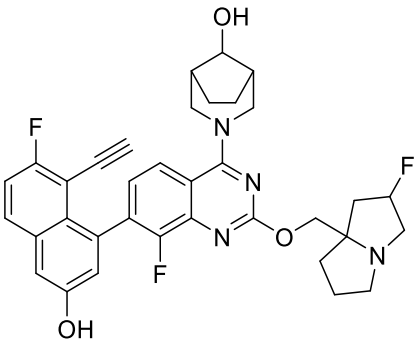
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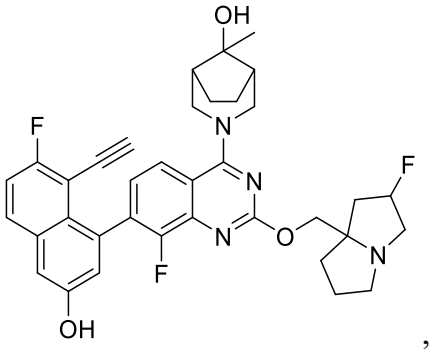
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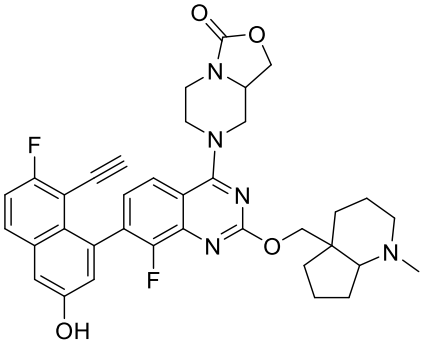
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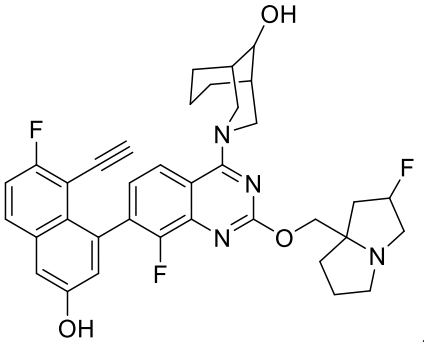
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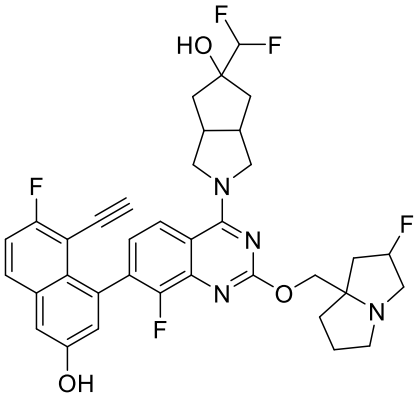
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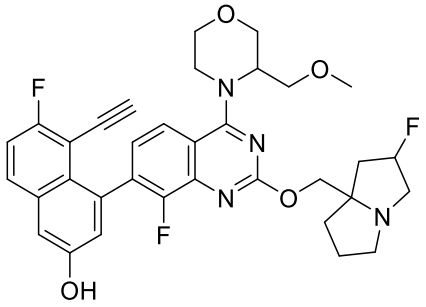
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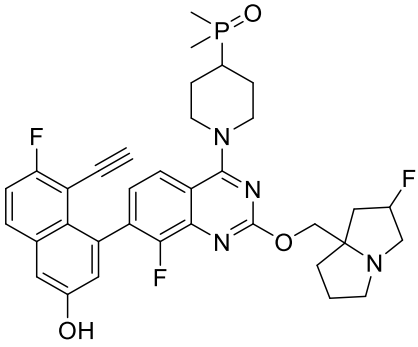
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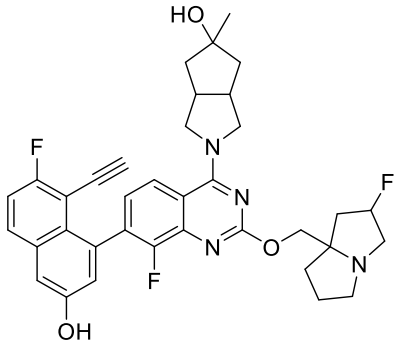
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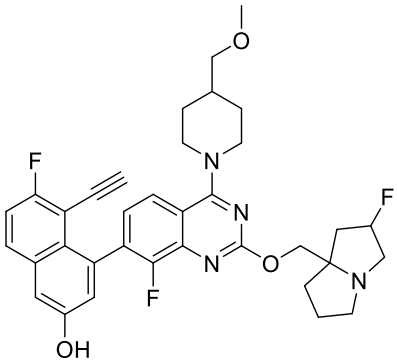
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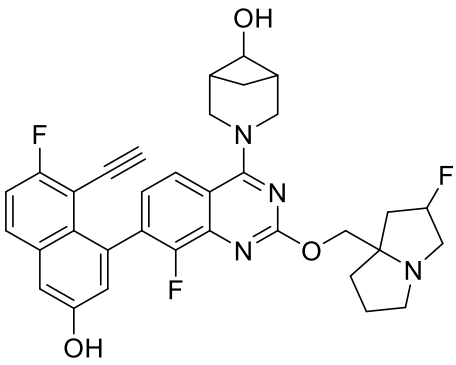
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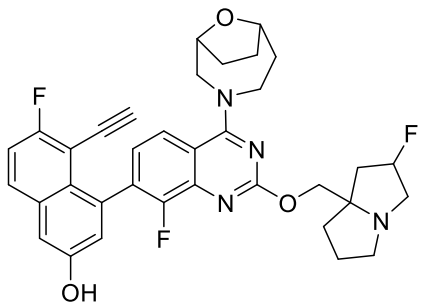
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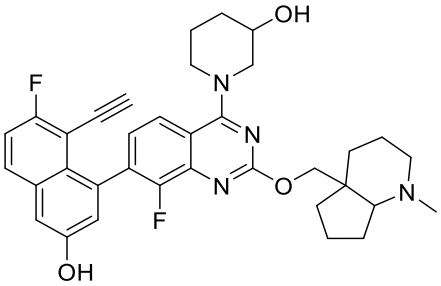
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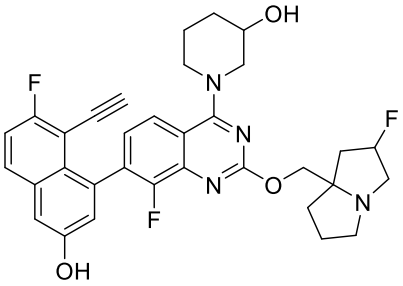
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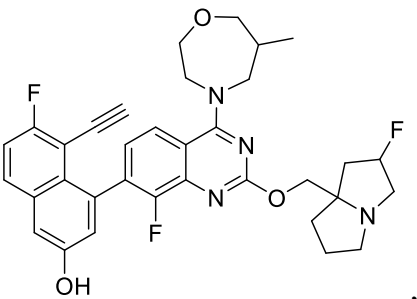
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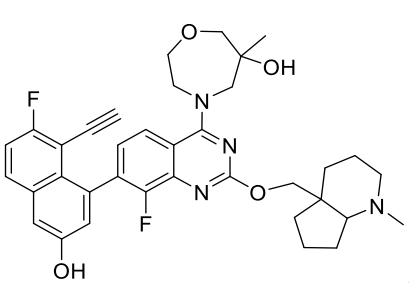
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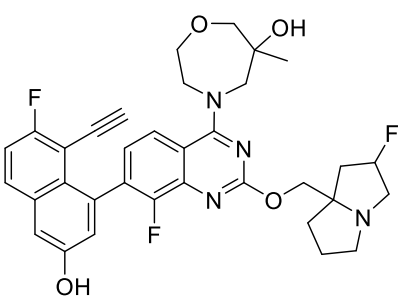

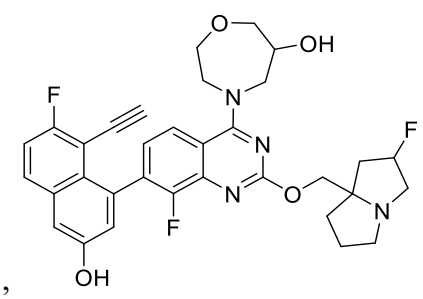
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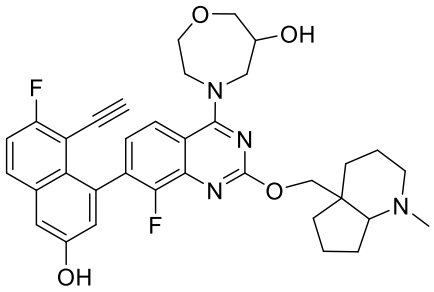
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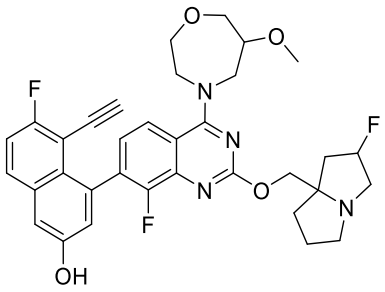
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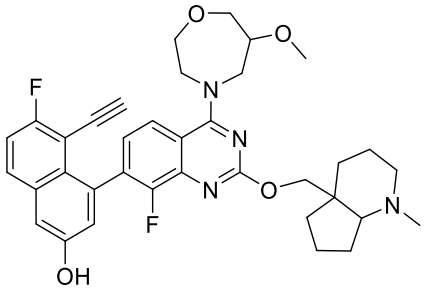
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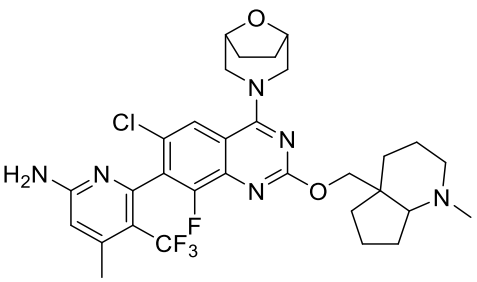
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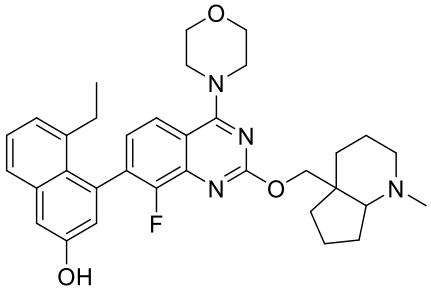
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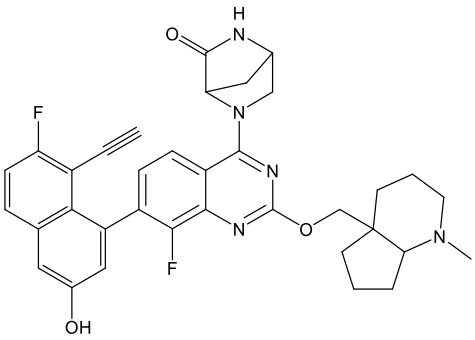
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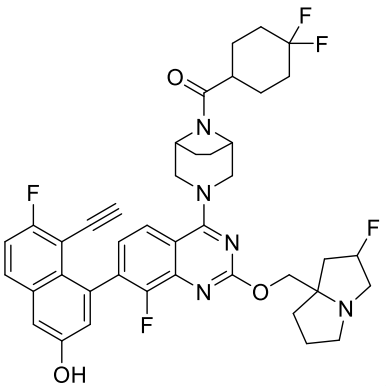
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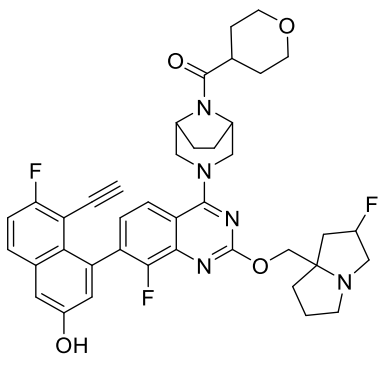
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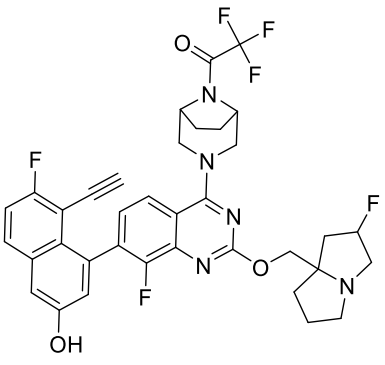
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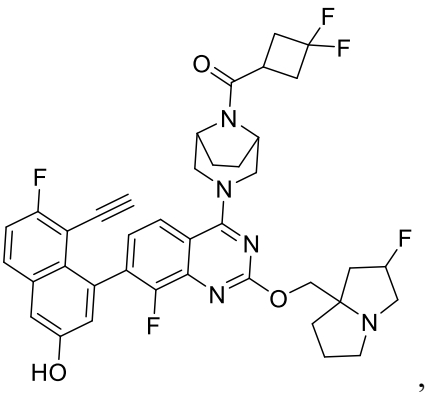
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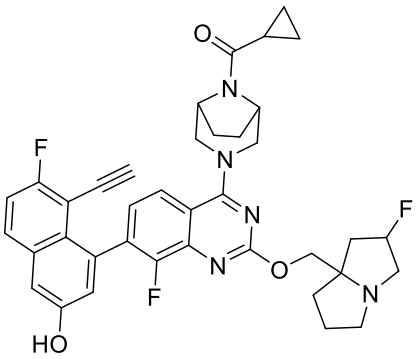
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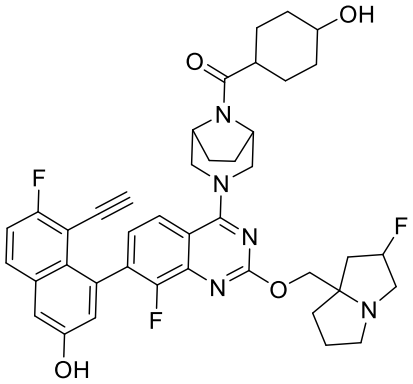
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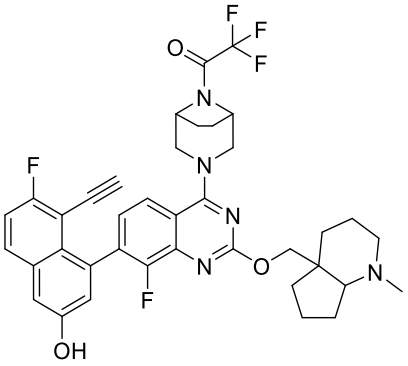
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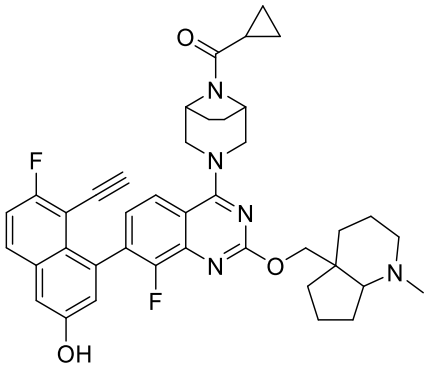
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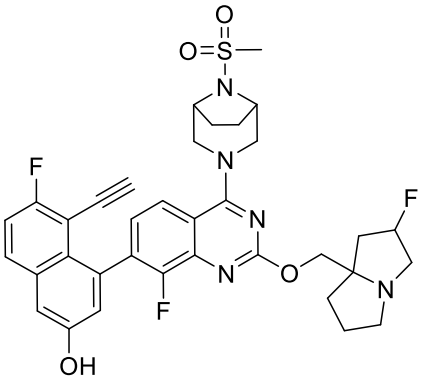
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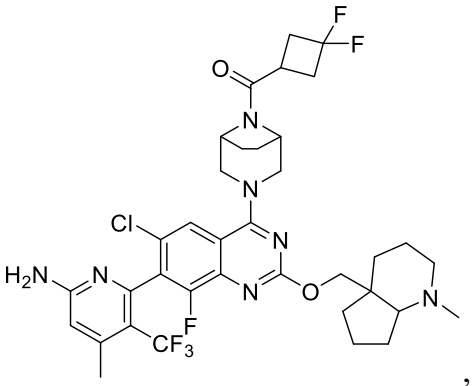
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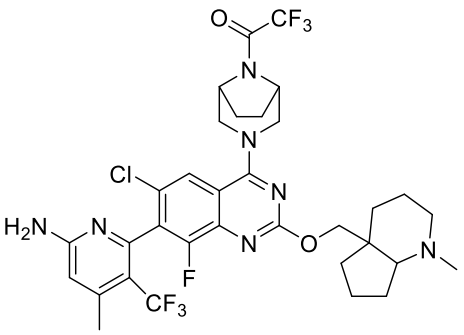
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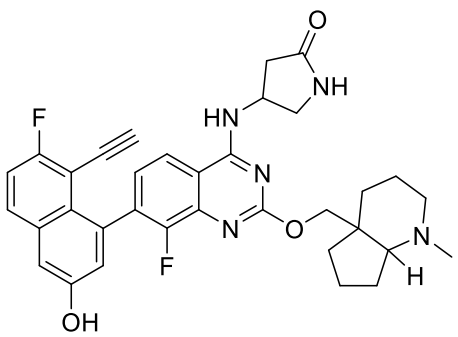
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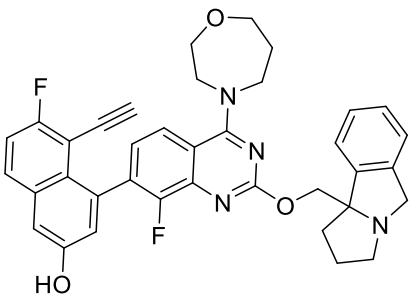
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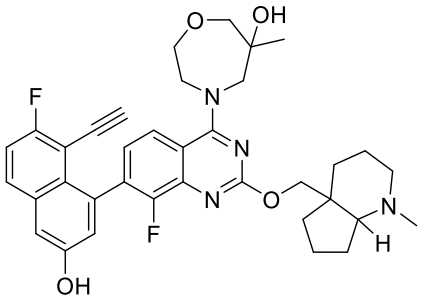
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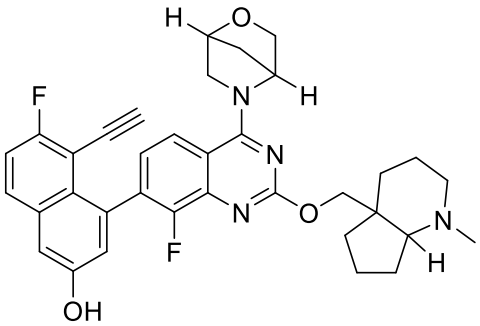
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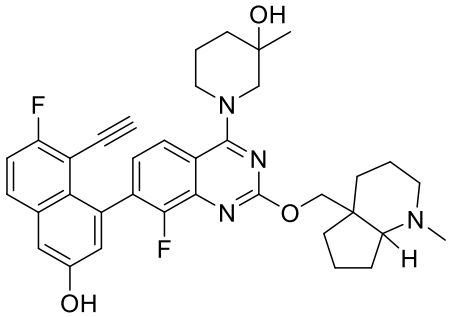
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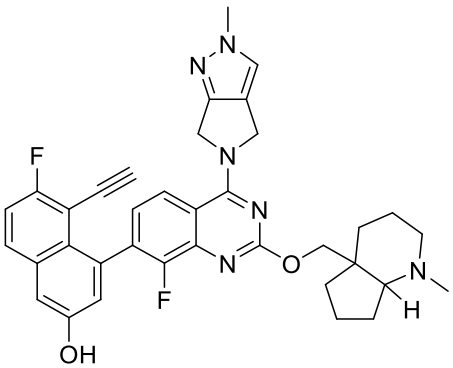
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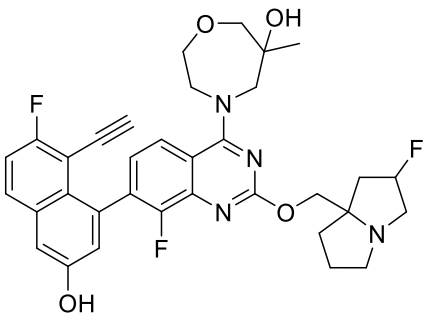
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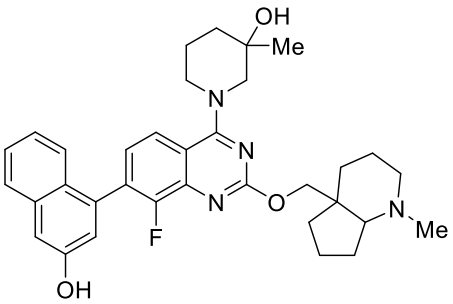
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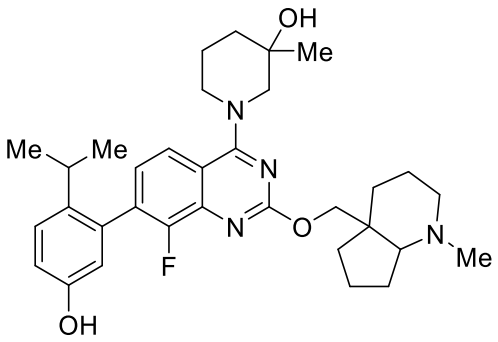
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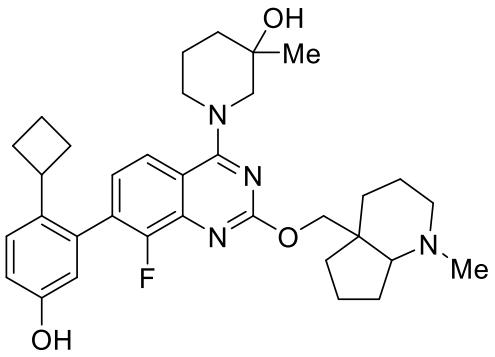
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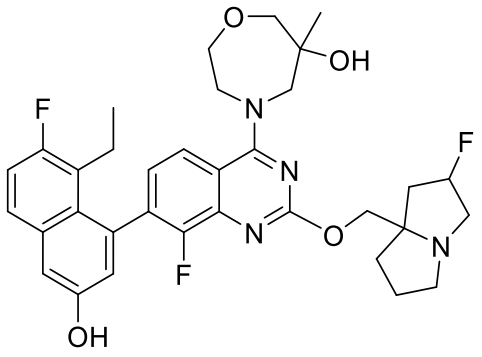
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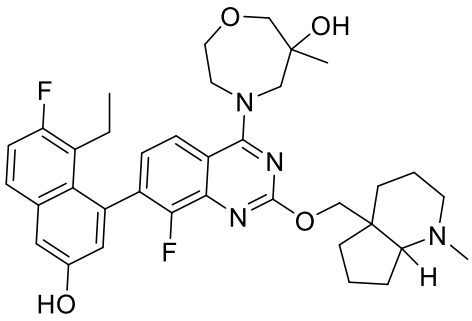
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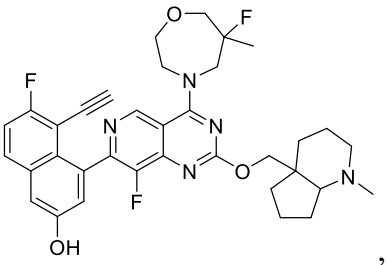
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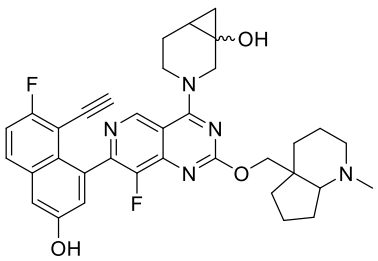
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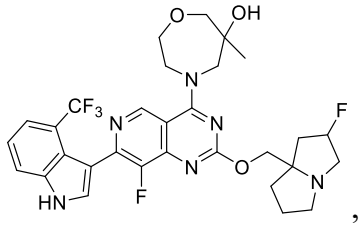
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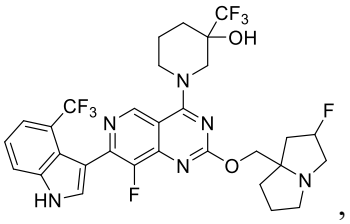
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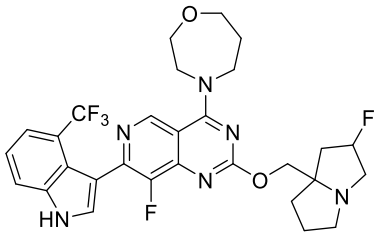
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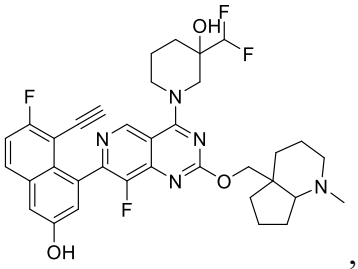
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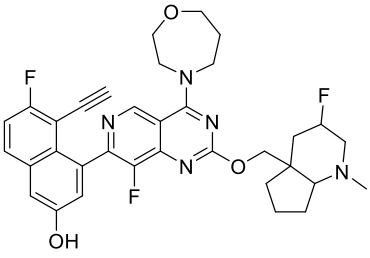
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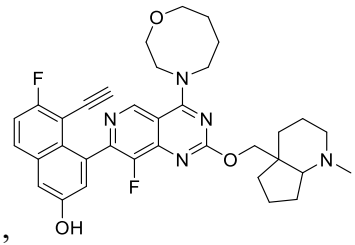
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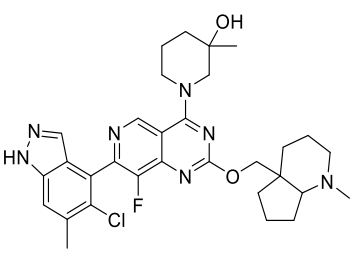
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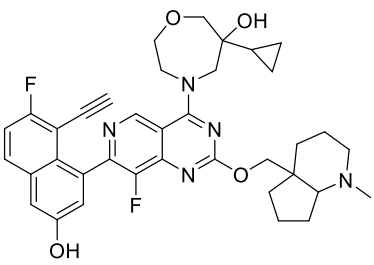
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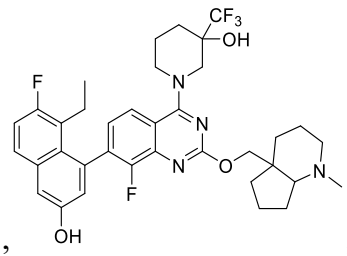
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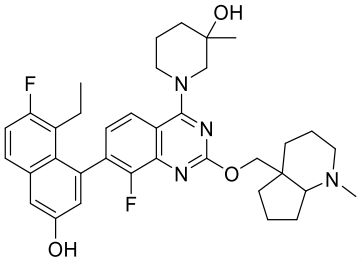
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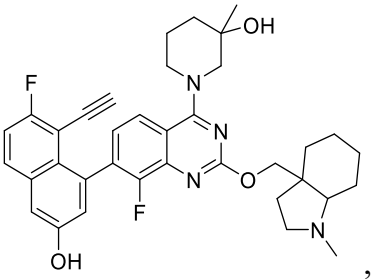
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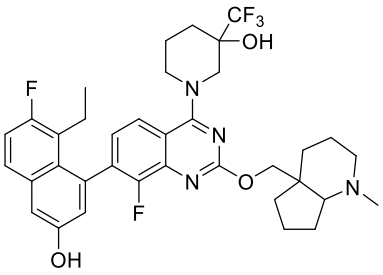
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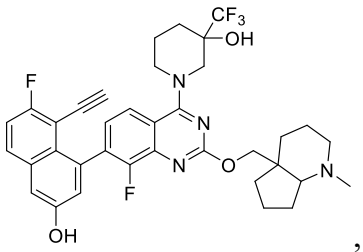
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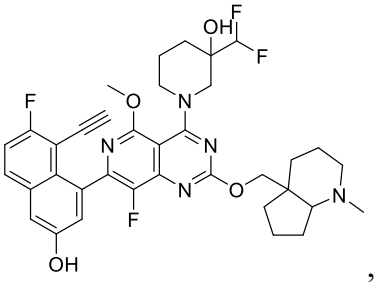
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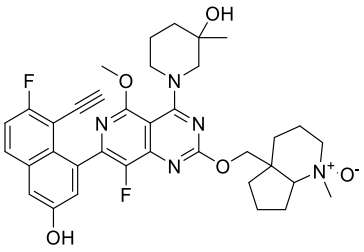
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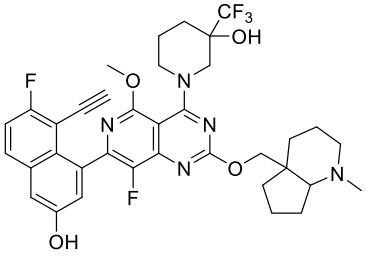
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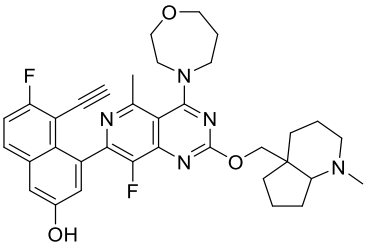
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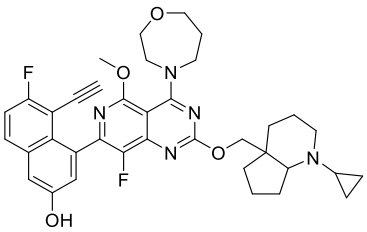
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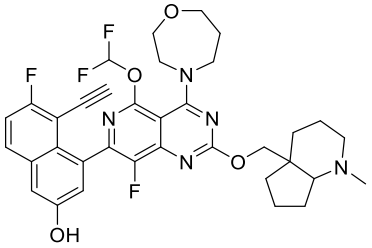
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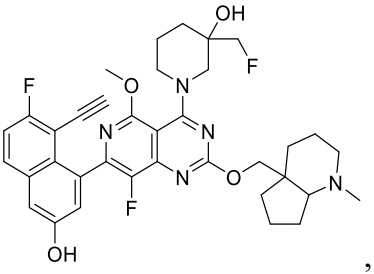
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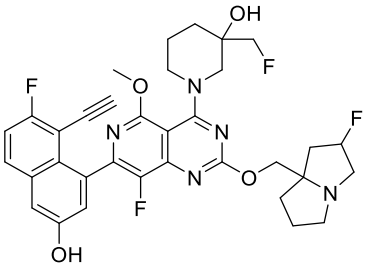
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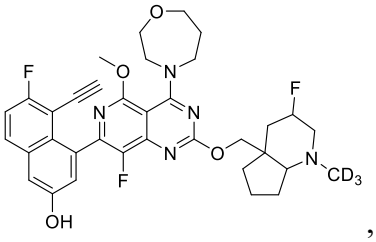
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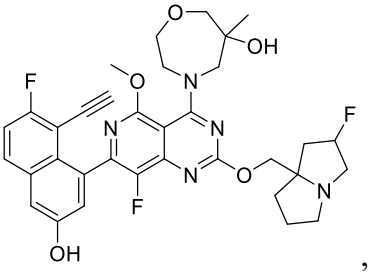
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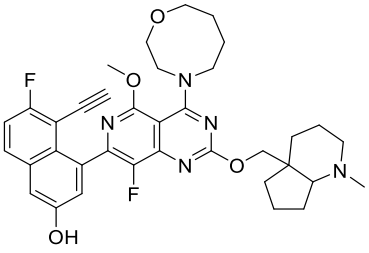
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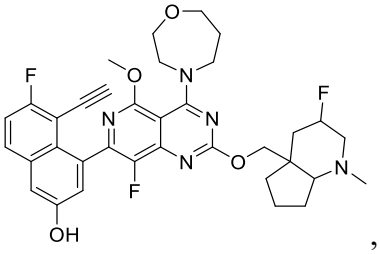
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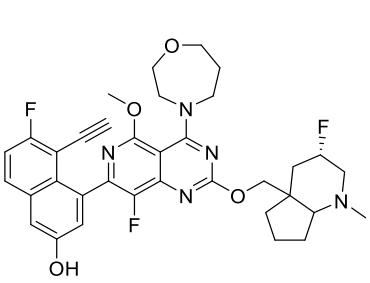
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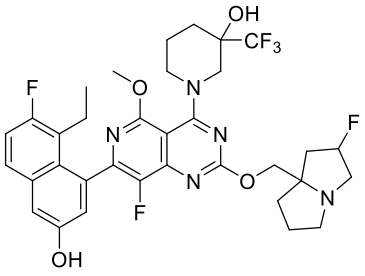
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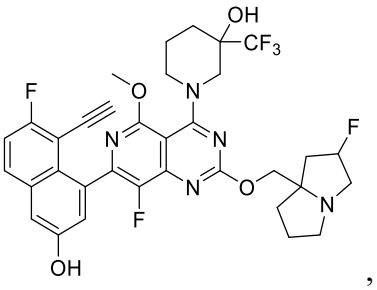
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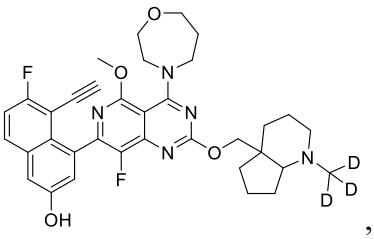
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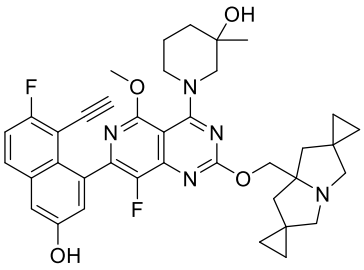
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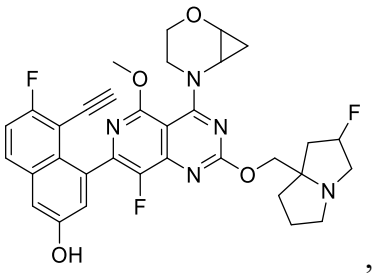
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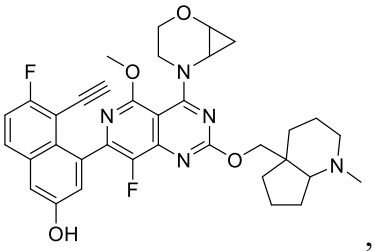
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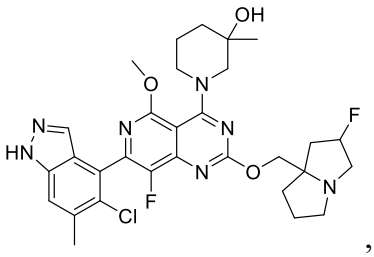
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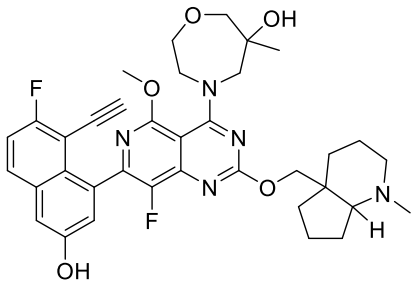
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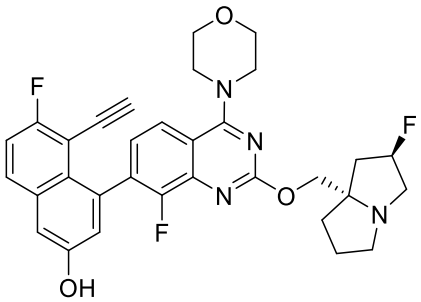
or a pharmaceutically acceptable salt thereof.

41. The compound of claim 40 that is selected from the group consisting of: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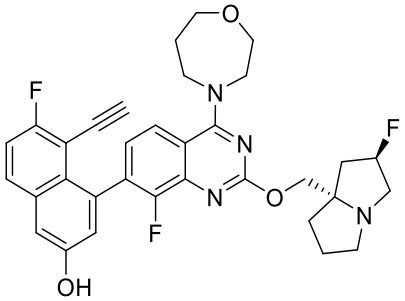
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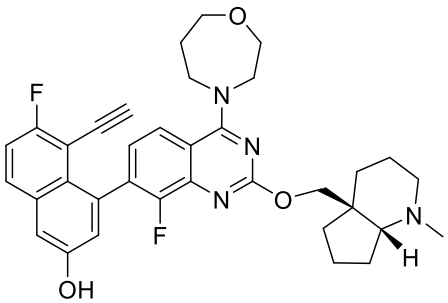
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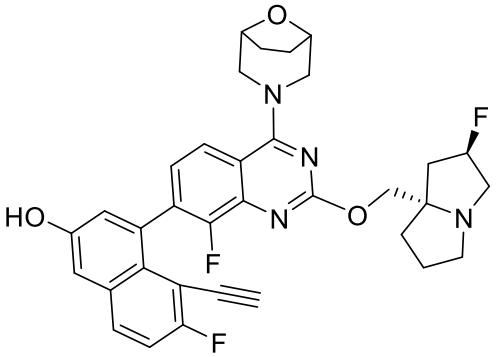
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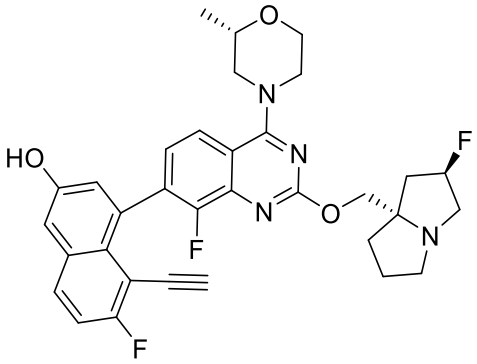
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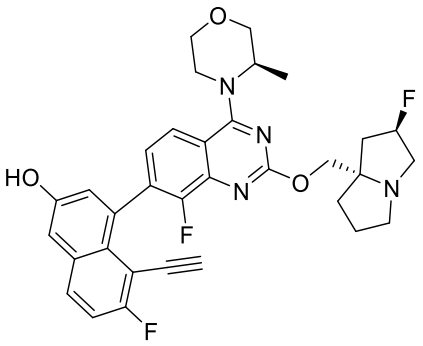
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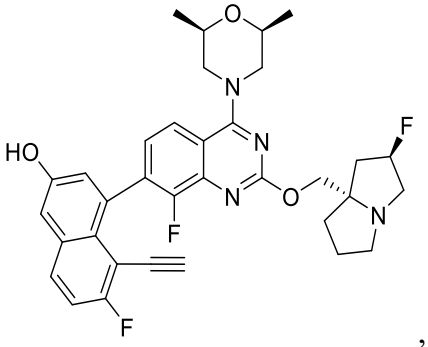
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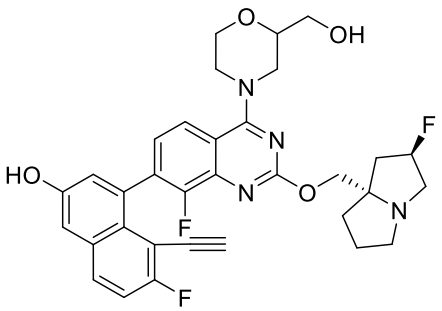
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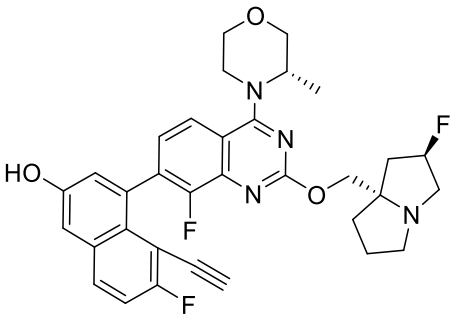
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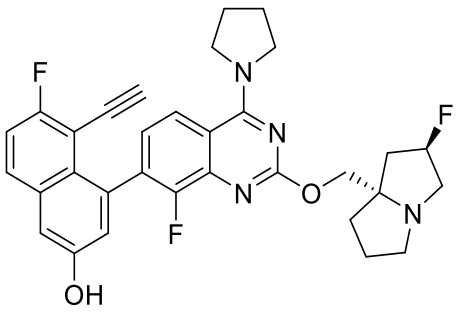
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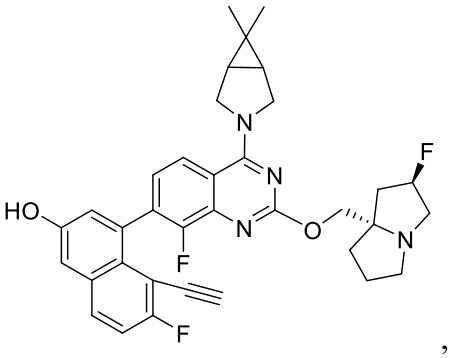
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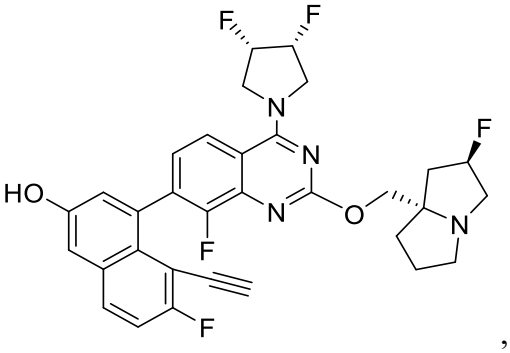
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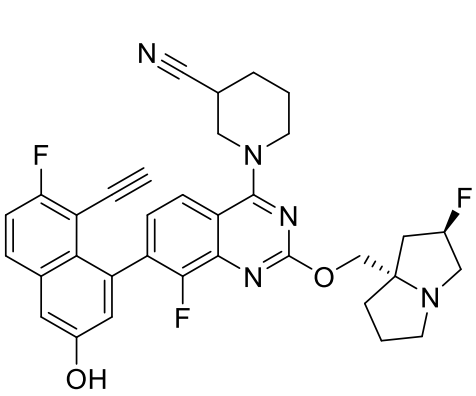
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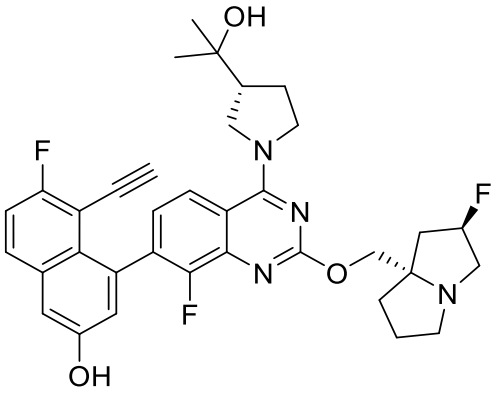
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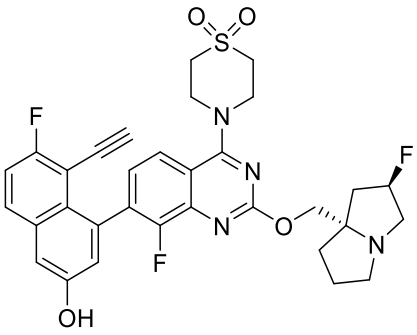
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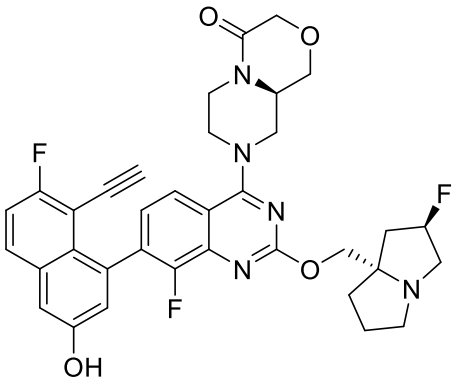
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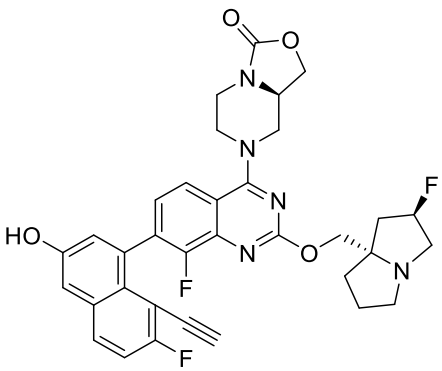
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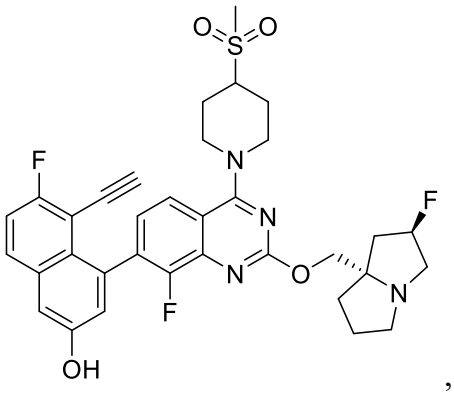
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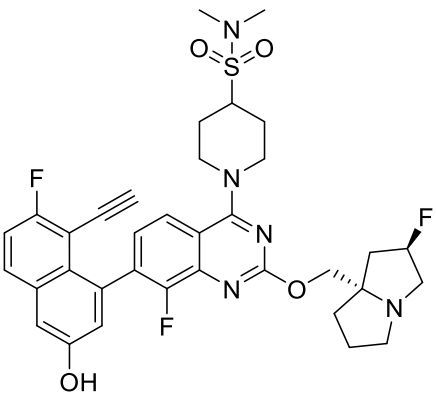
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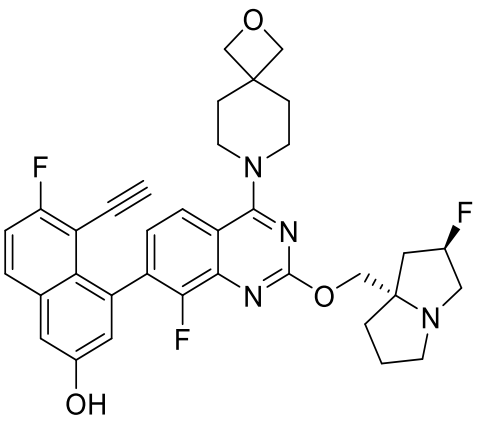
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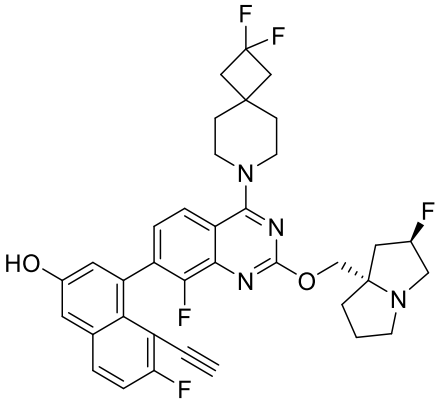
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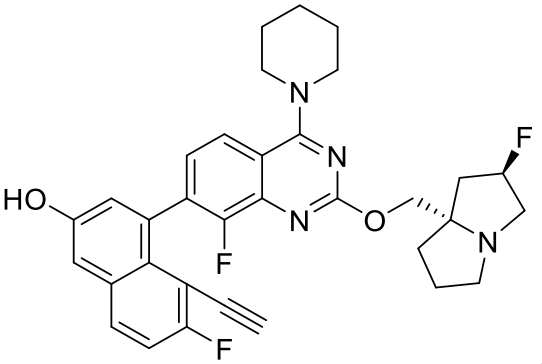
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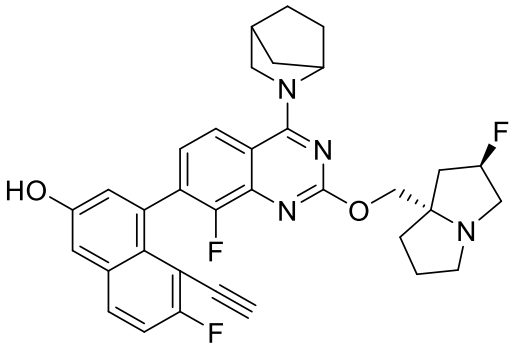
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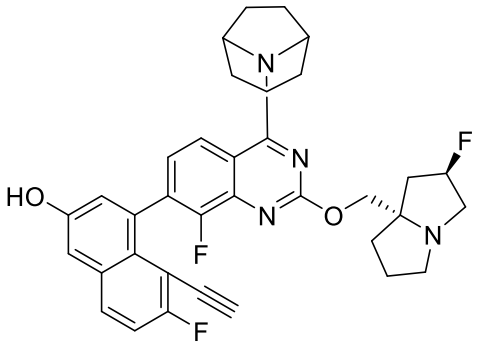
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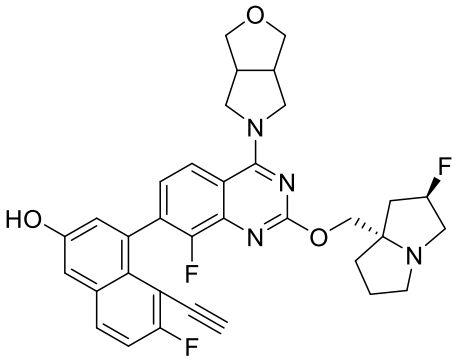
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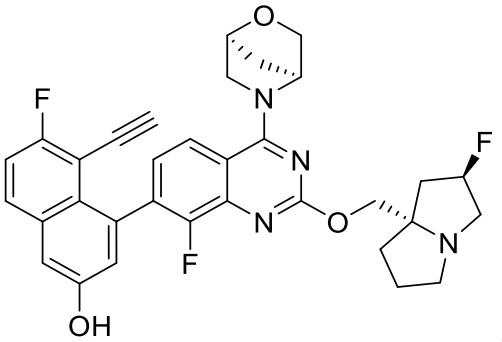
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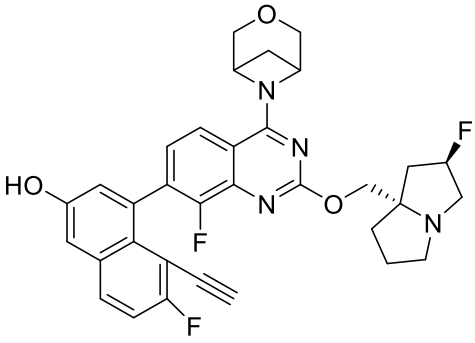
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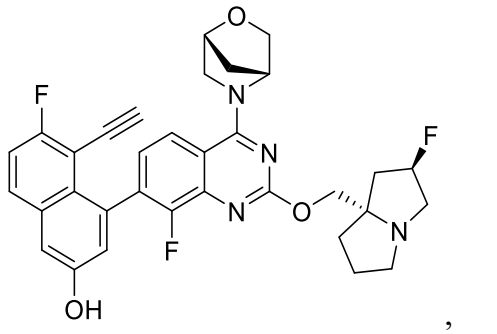
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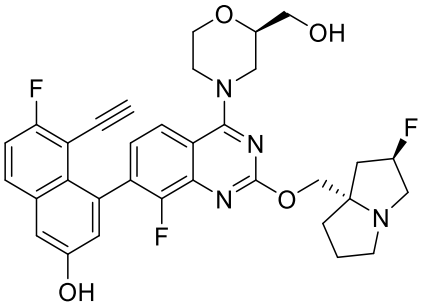
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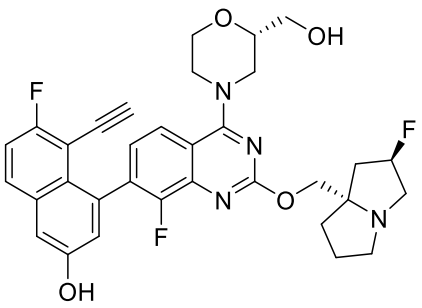
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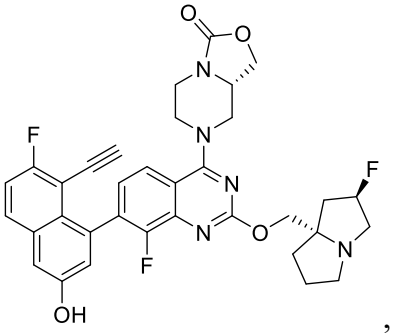
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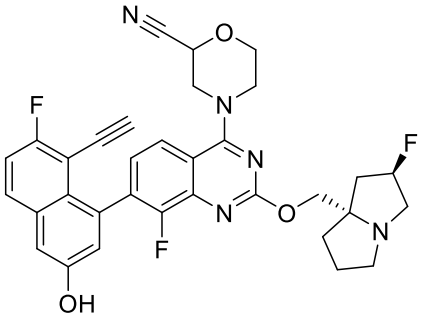
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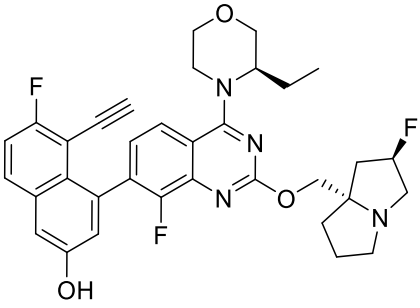
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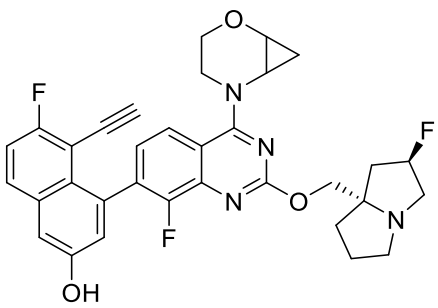
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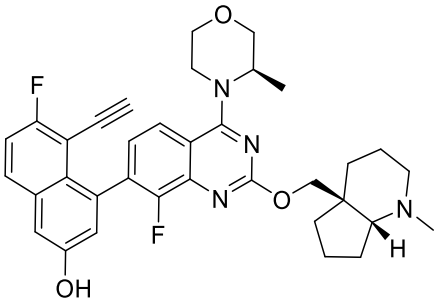
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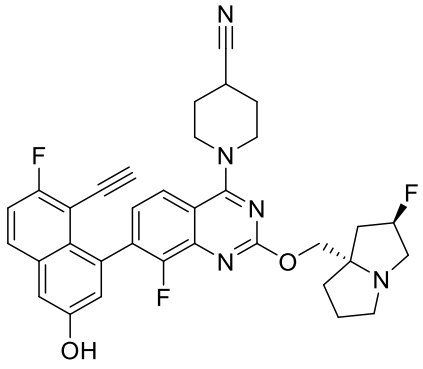
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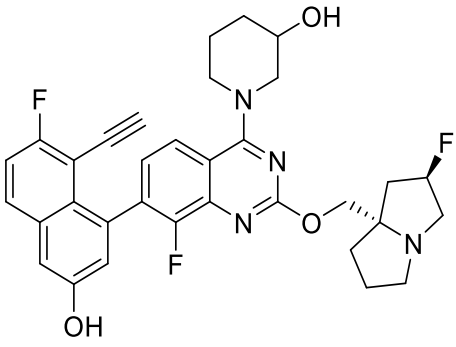
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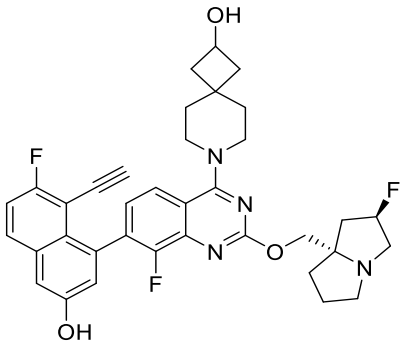
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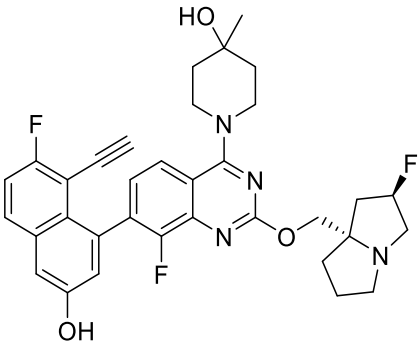


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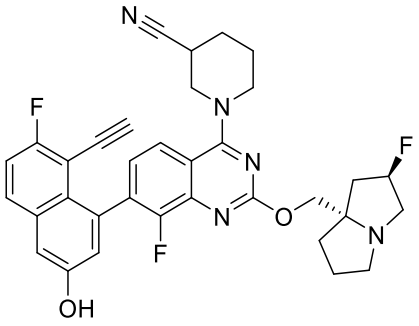


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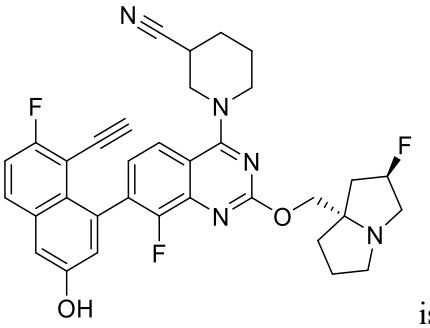


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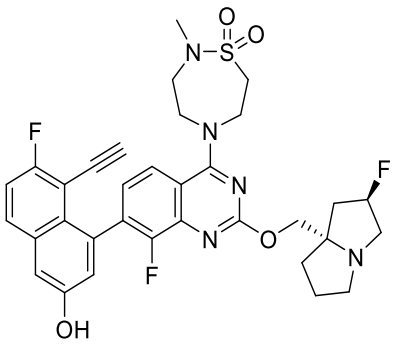


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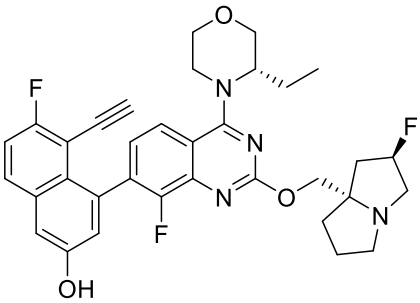


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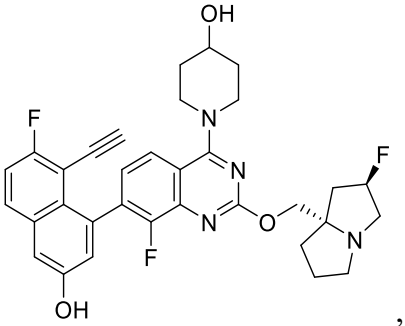


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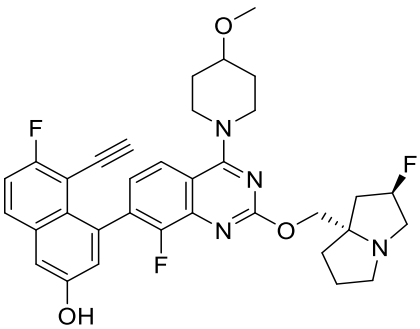


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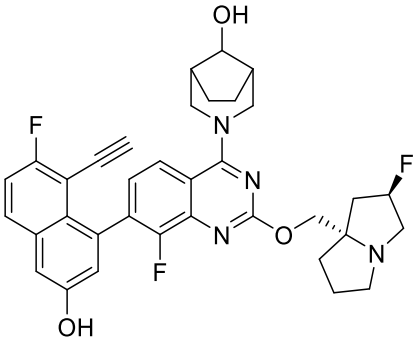


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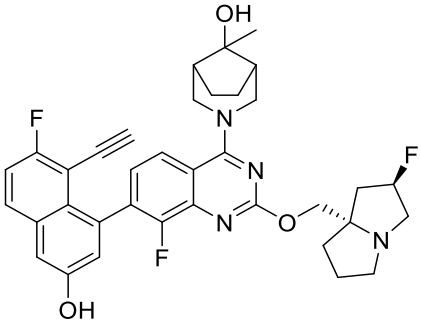
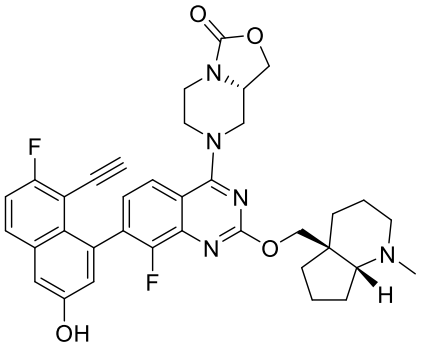
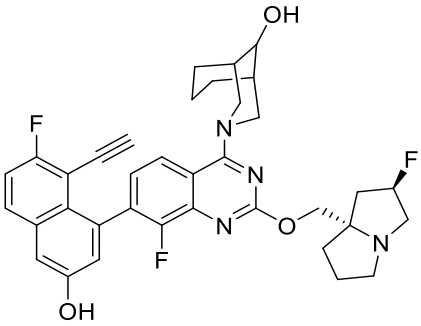
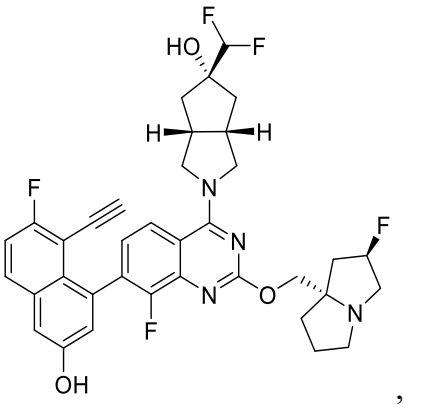
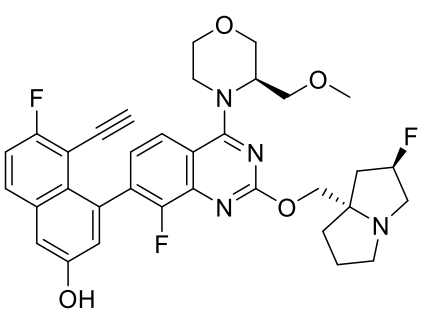
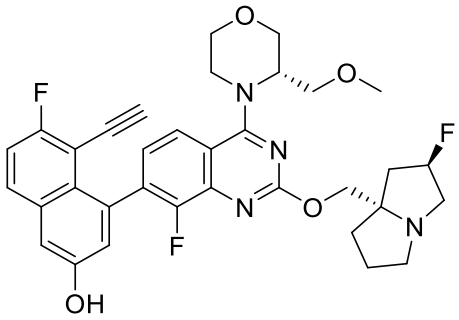
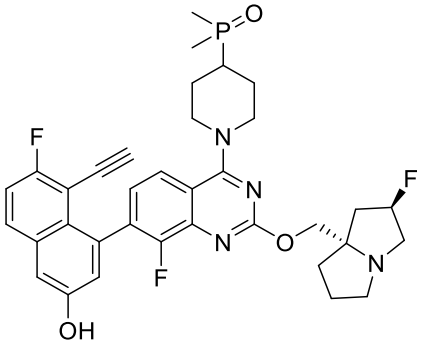
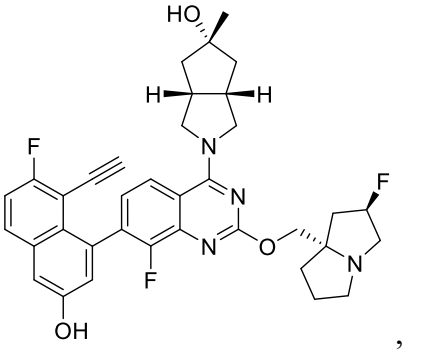

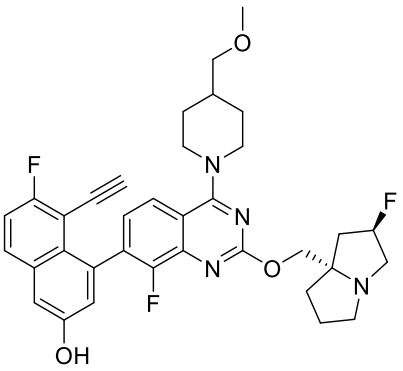
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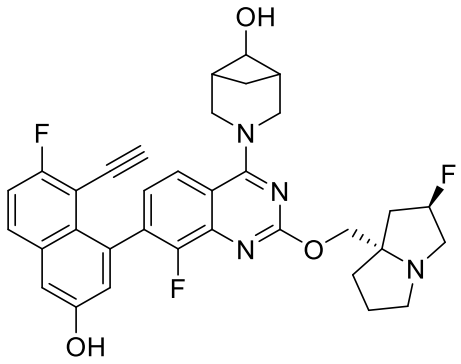
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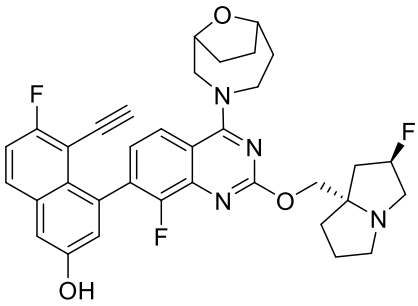
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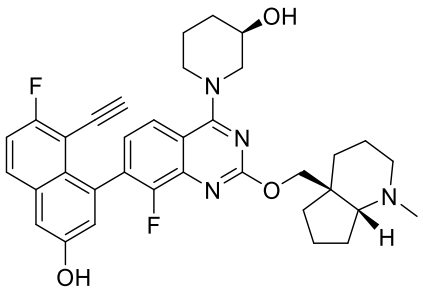
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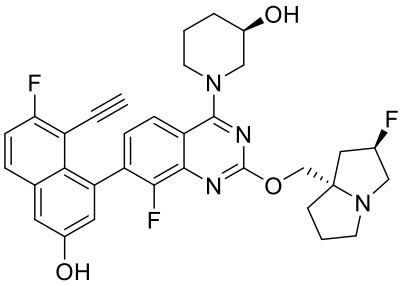
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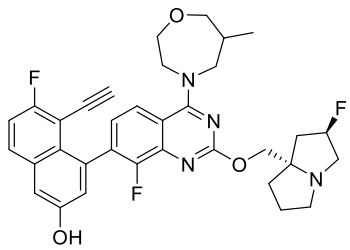
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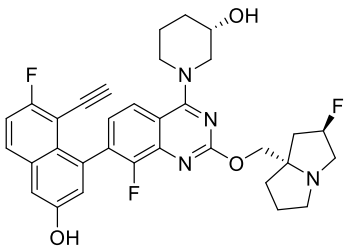
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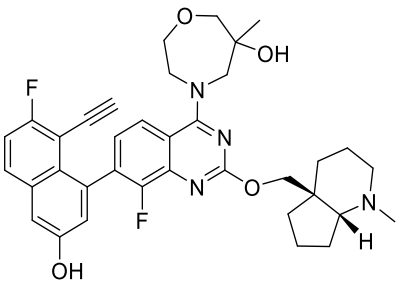
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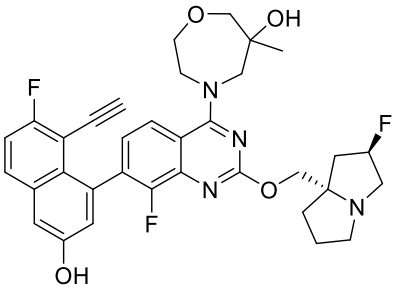
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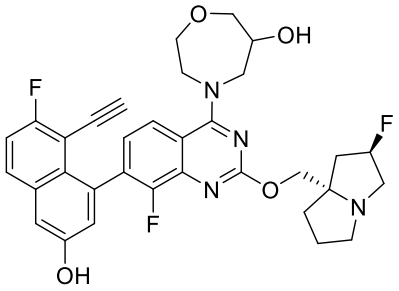
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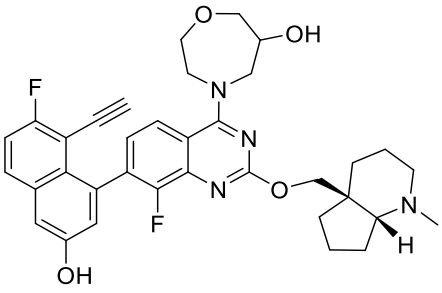
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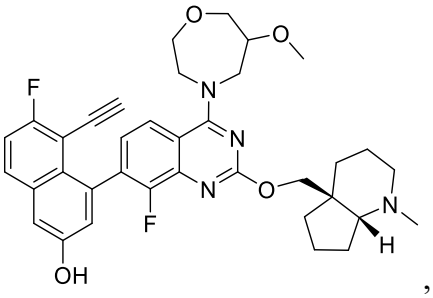
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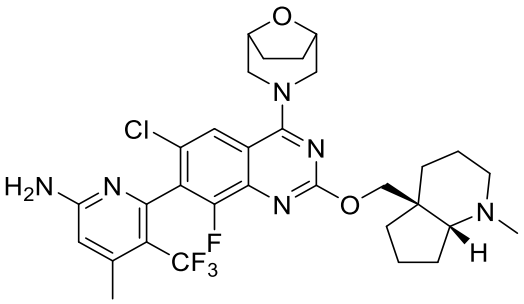
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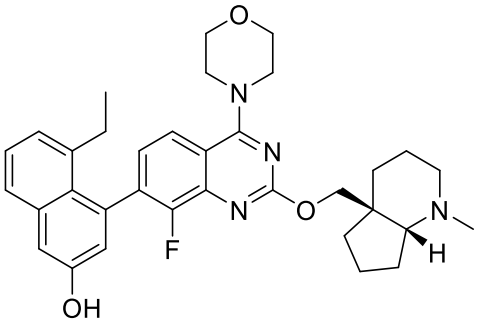
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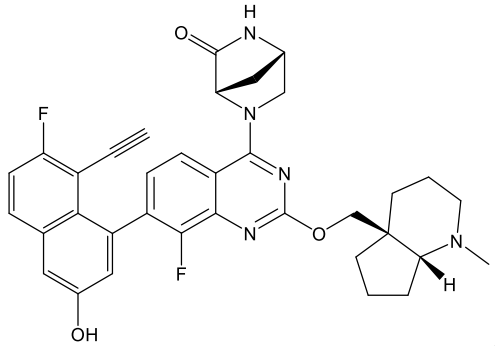
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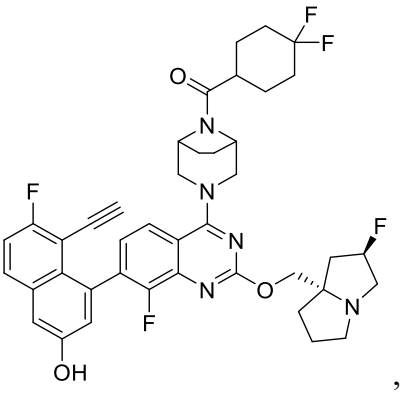
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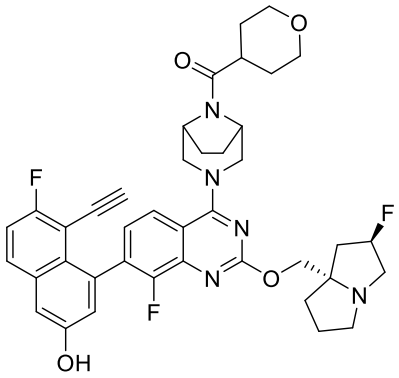
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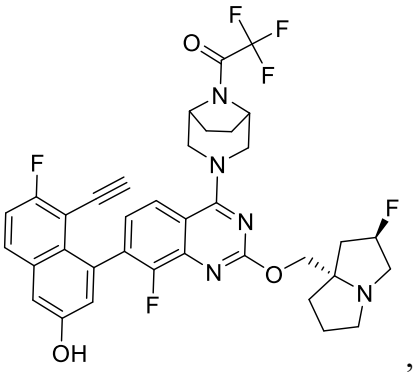
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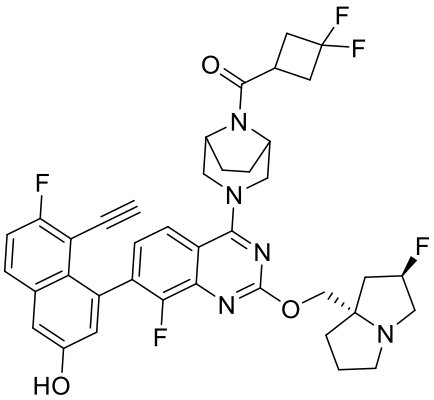
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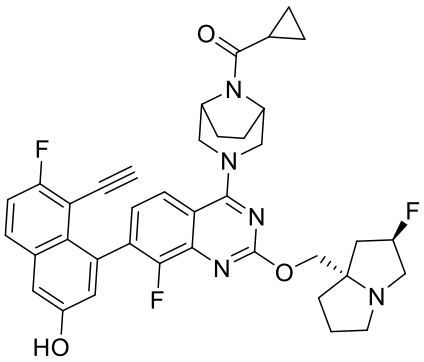
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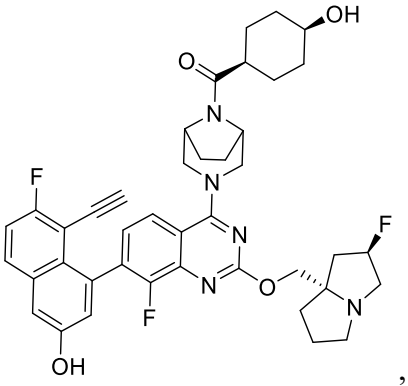
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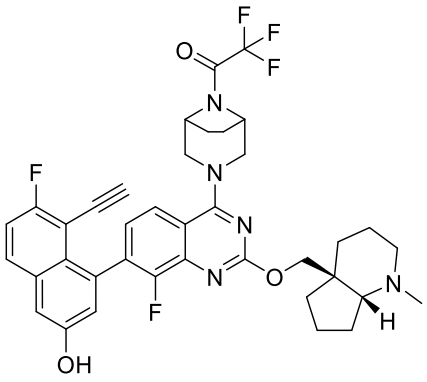
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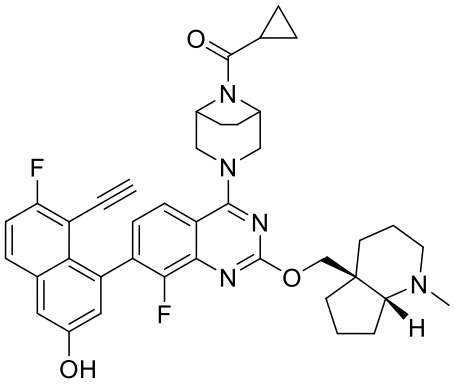
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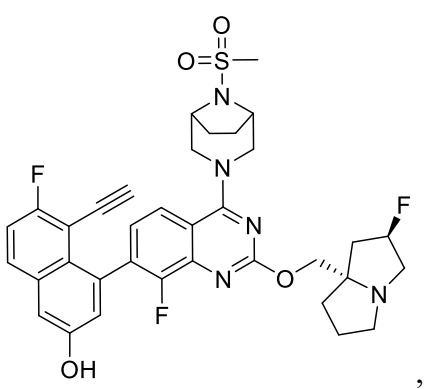
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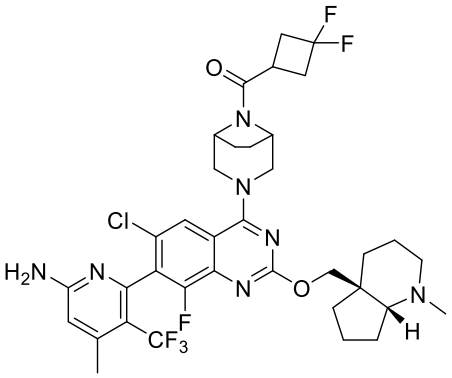
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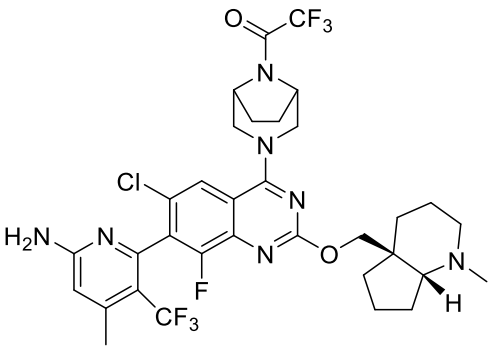
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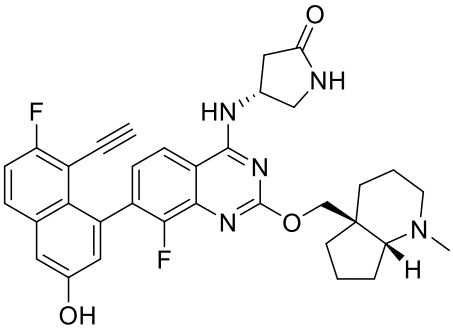
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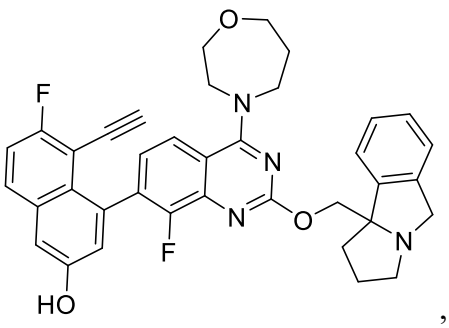
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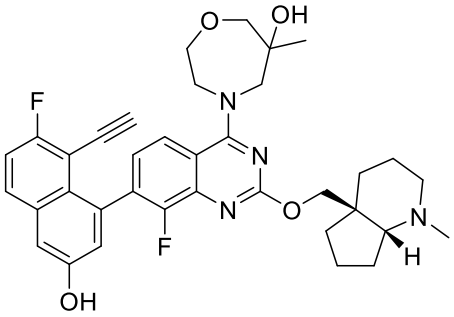
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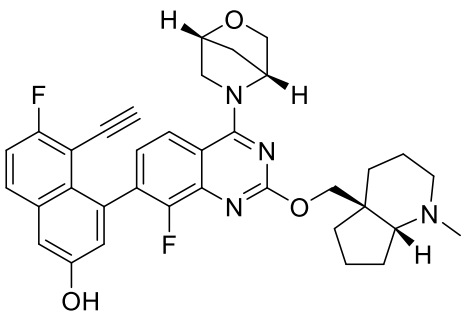
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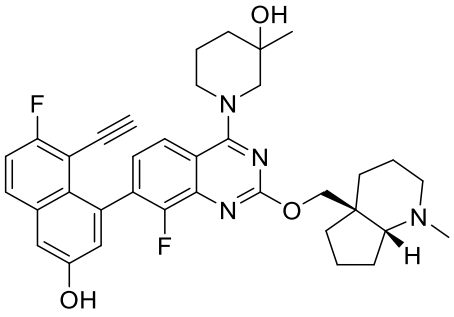
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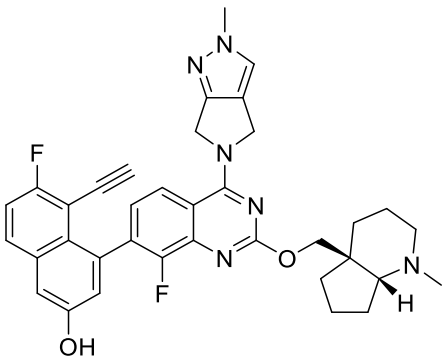
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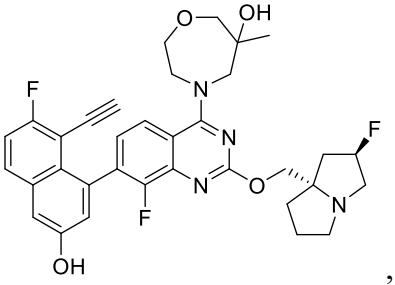
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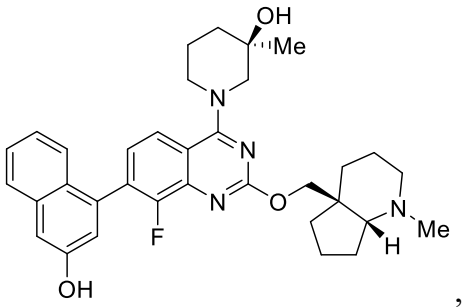
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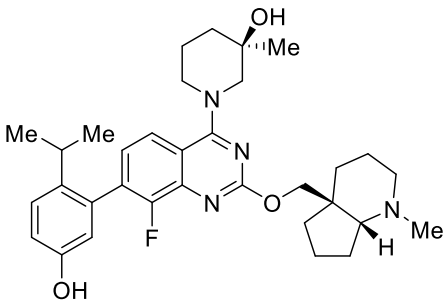
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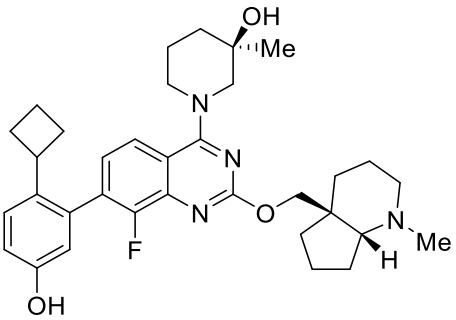
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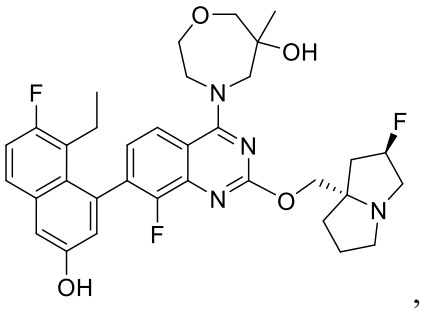
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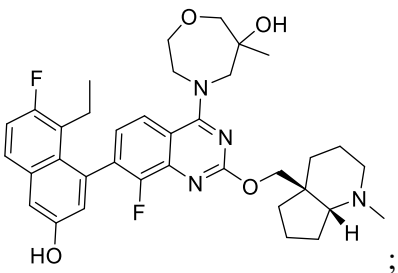
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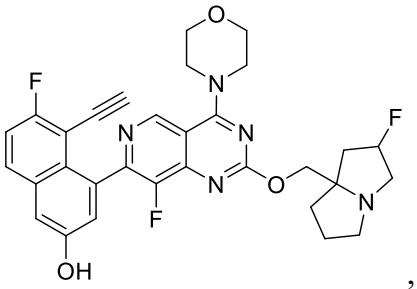
or a pharmaceutically acceptable salt thereof.

42. The compound of claim 4 that is selected from the group consisting of: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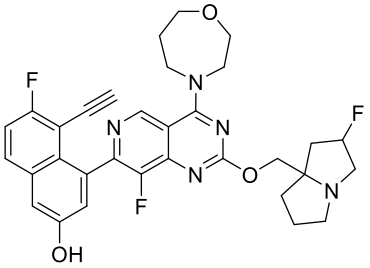
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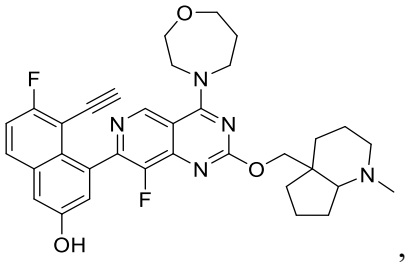
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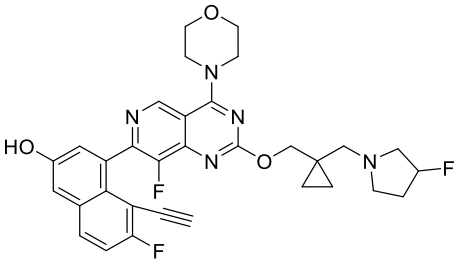
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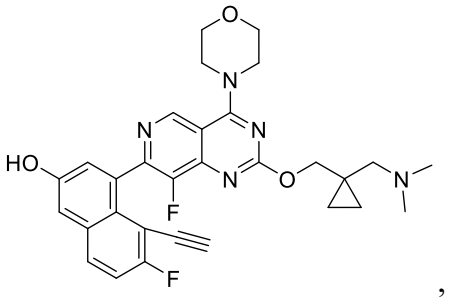
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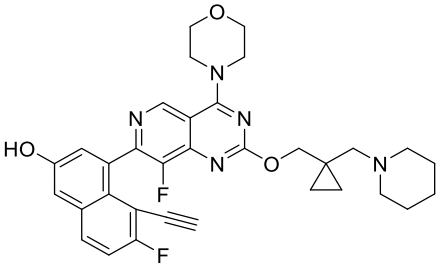
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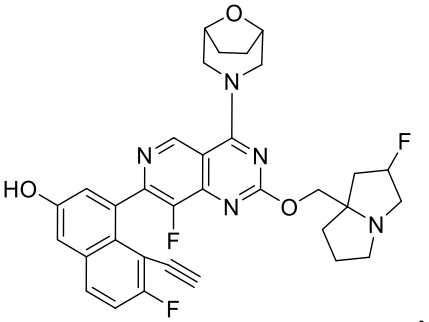
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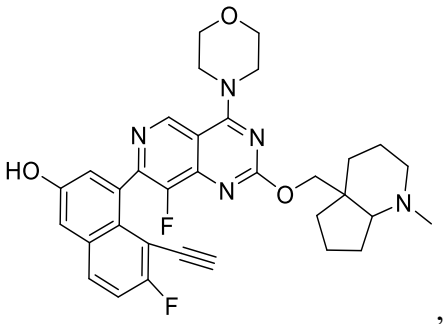
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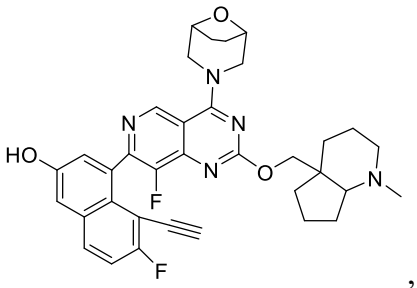
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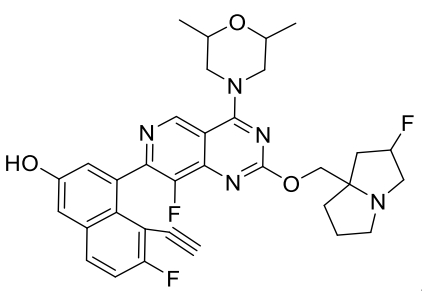
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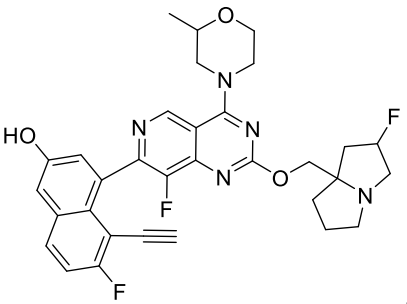
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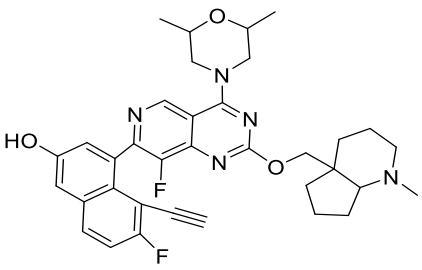
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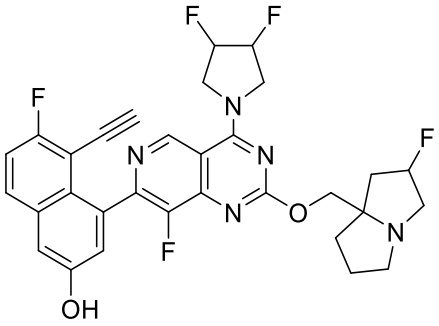
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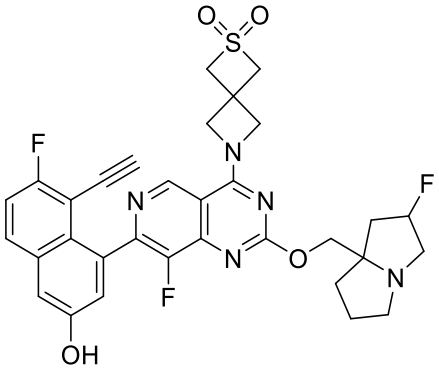
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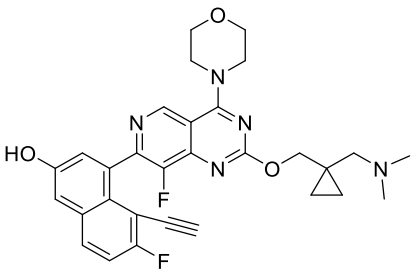
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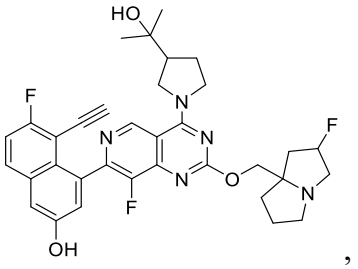
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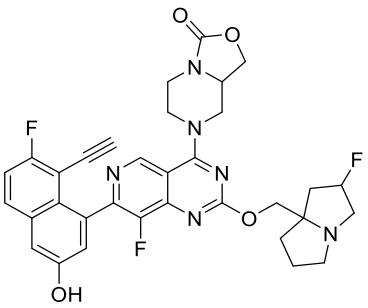
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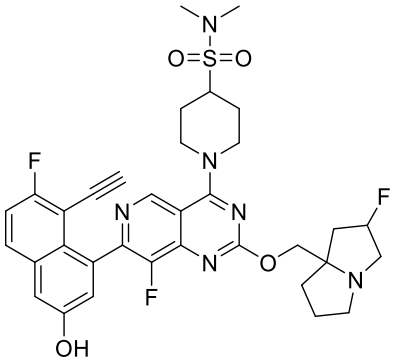
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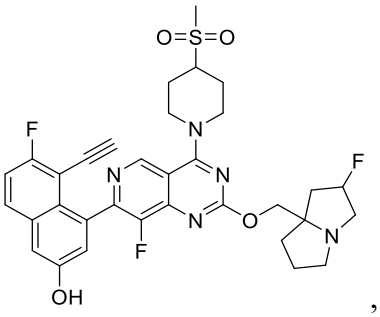
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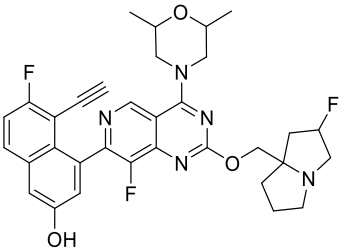
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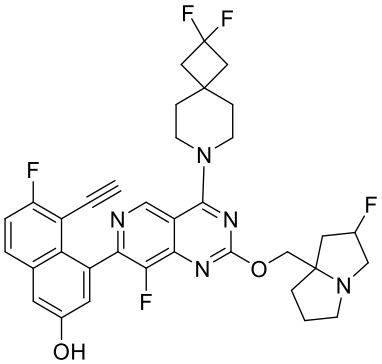
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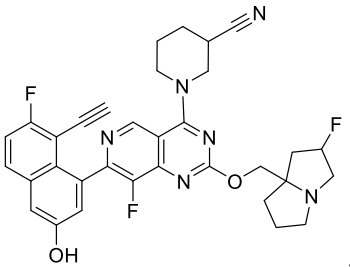
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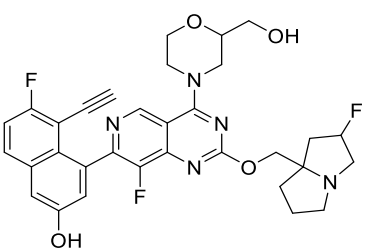
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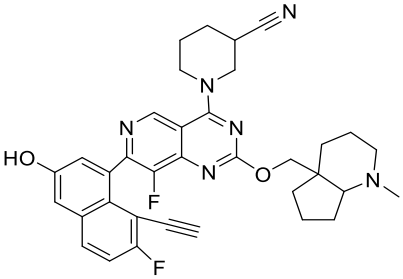
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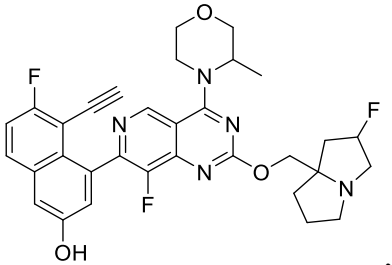
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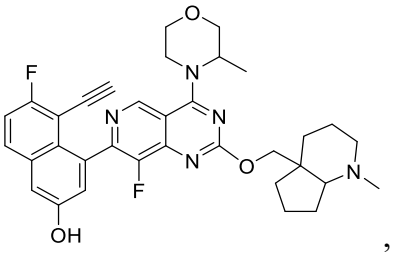
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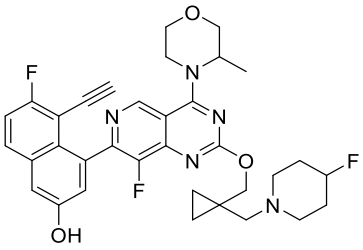
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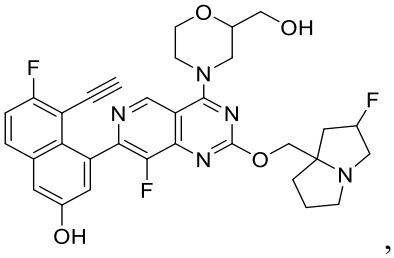
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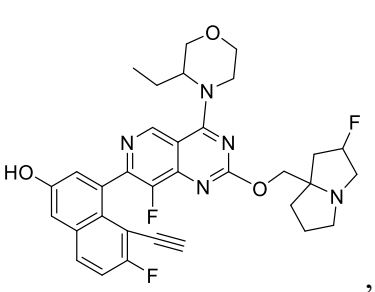
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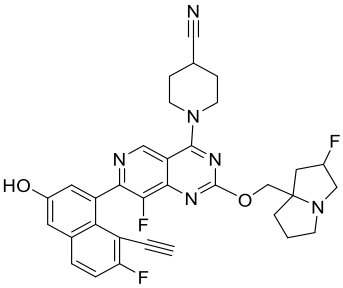
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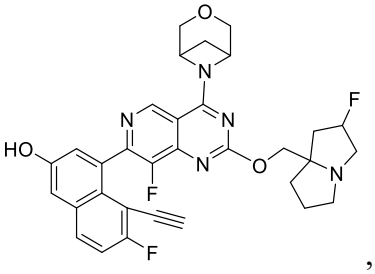
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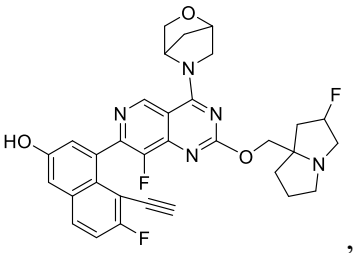
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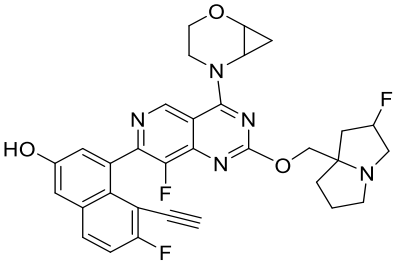
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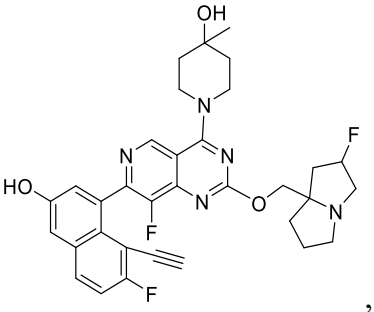
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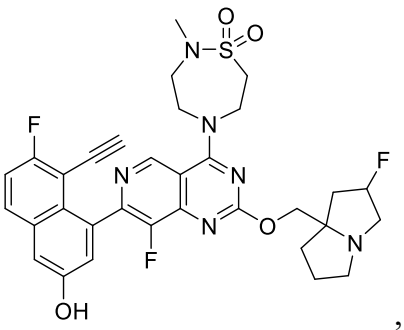
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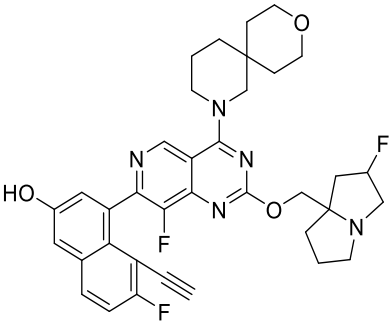
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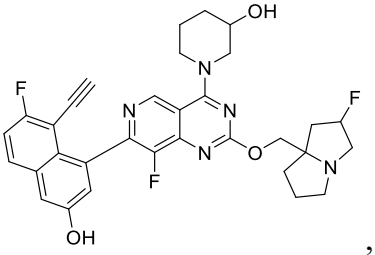
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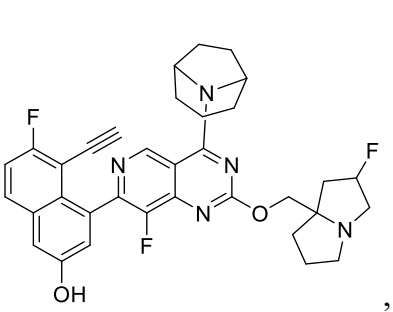
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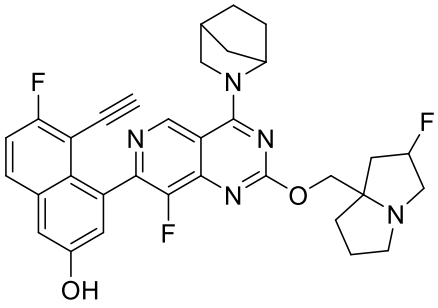
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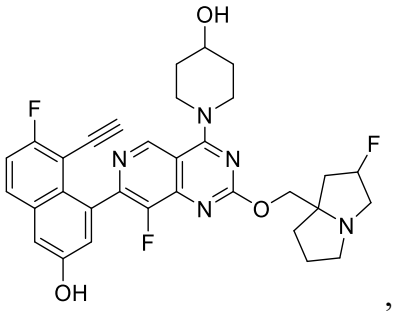
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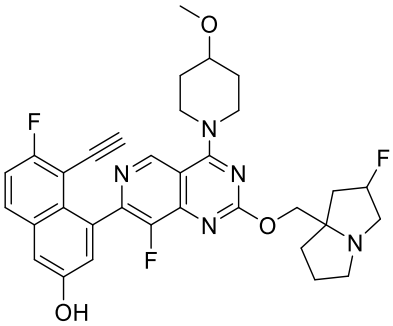
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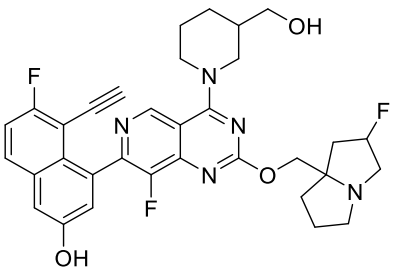
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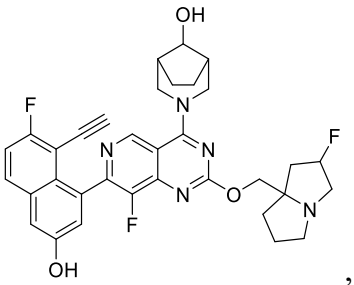
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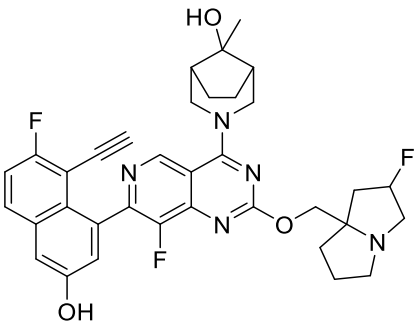
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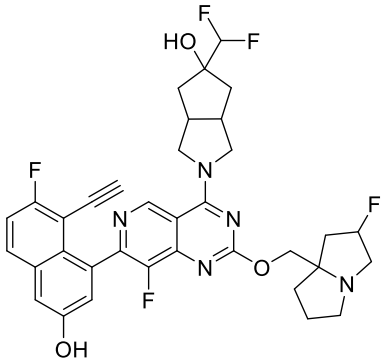
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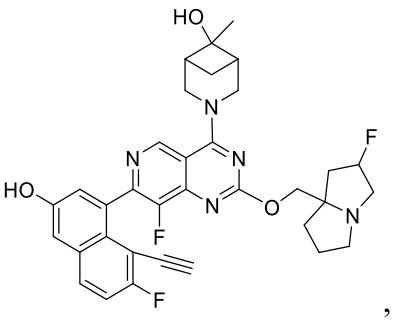
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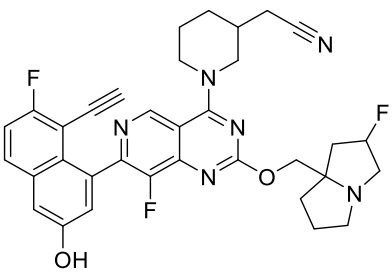
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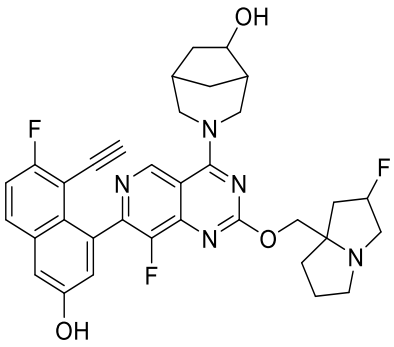
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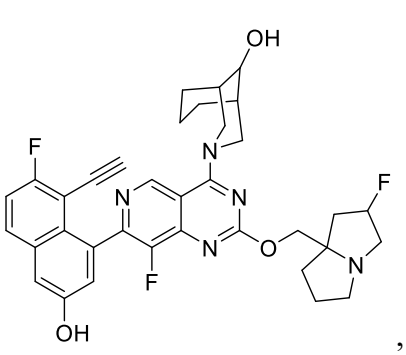
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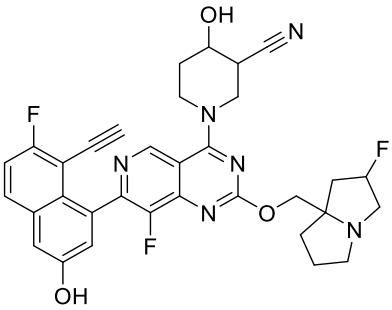
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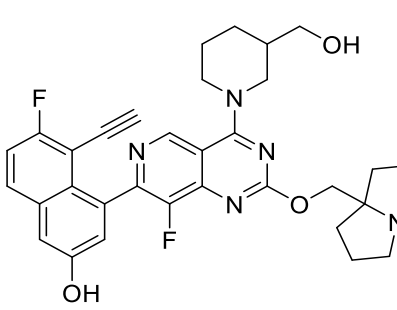
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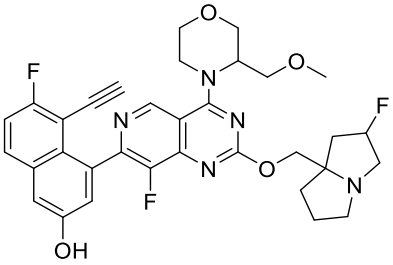
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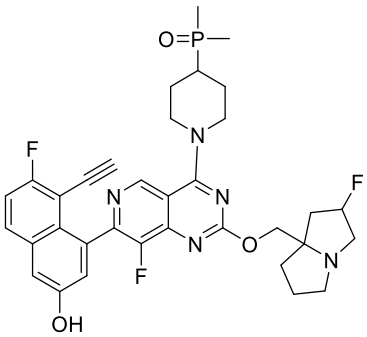
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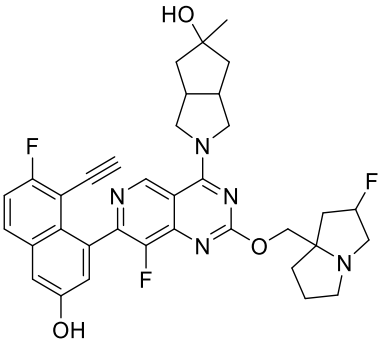
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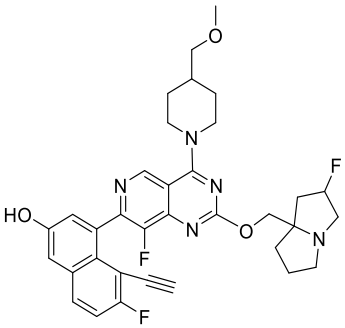
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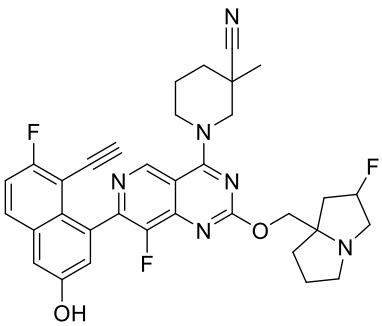
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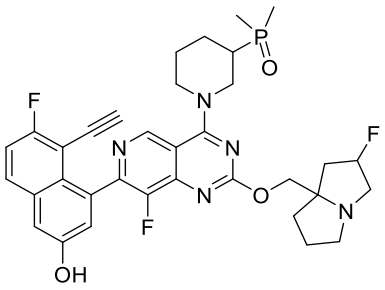
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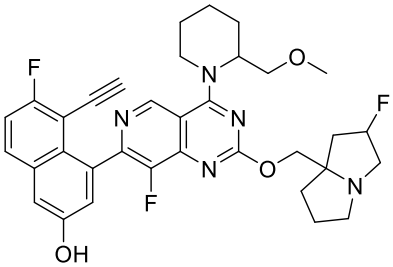
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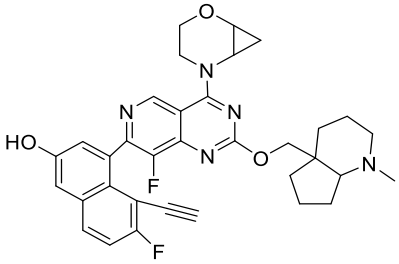
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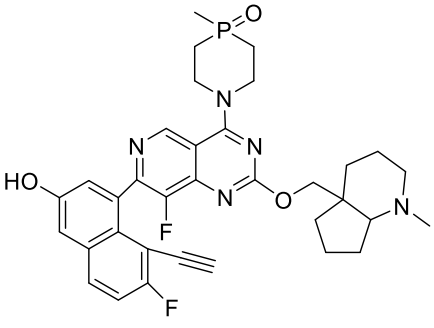
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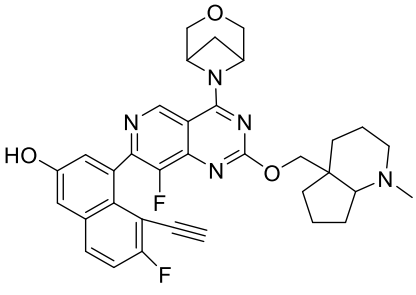
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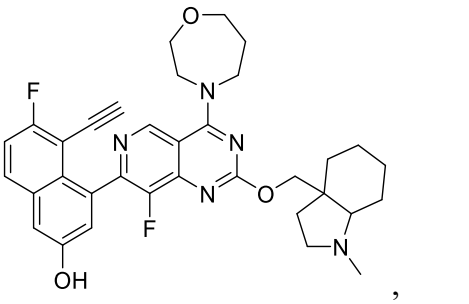
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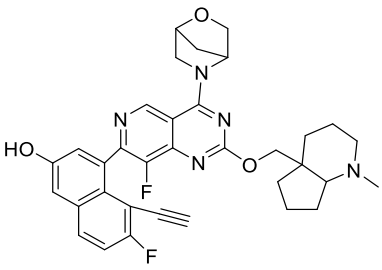
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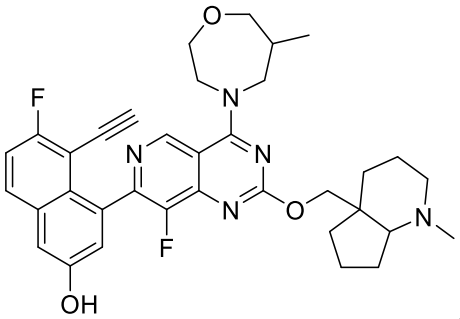
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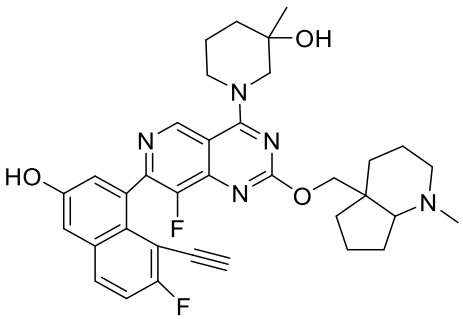
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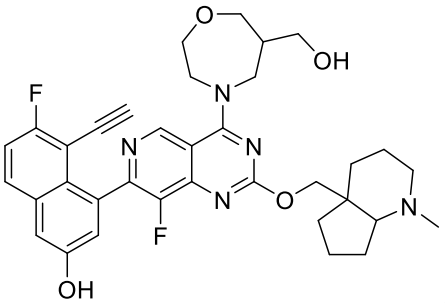
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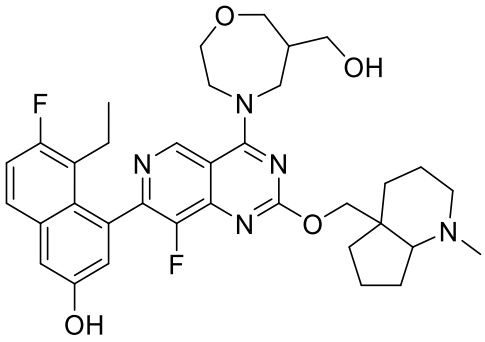
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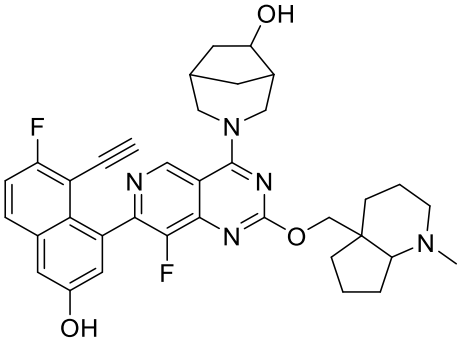
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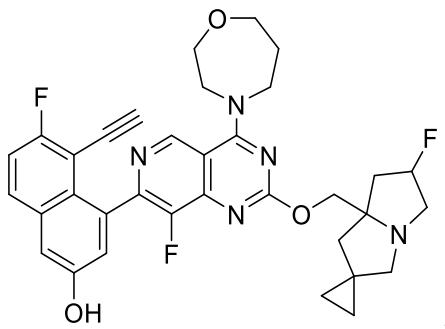
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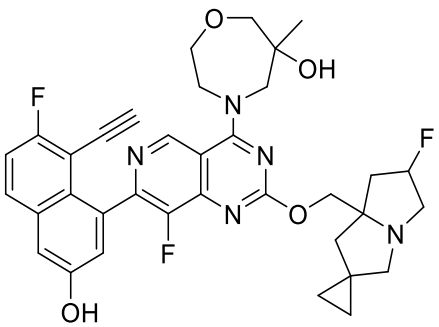
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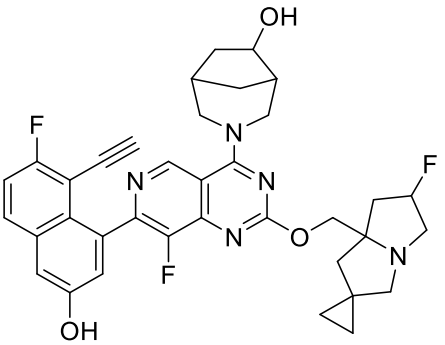
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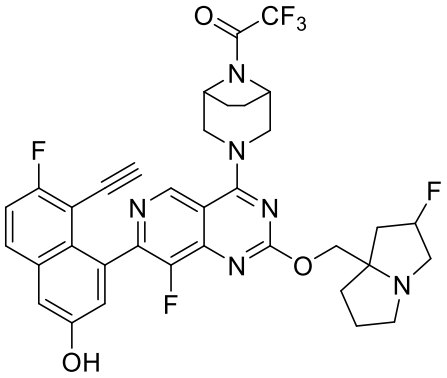
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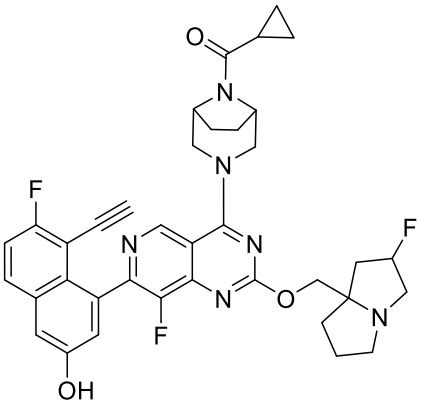
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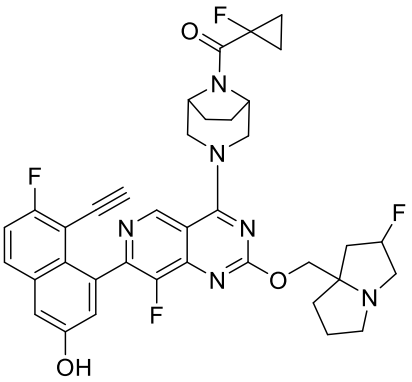
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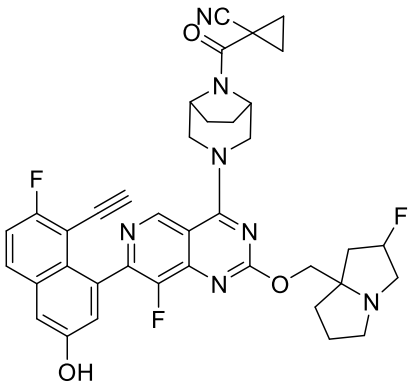
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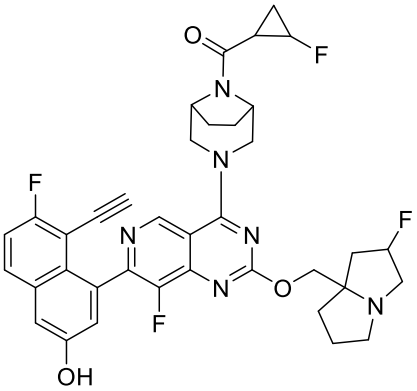
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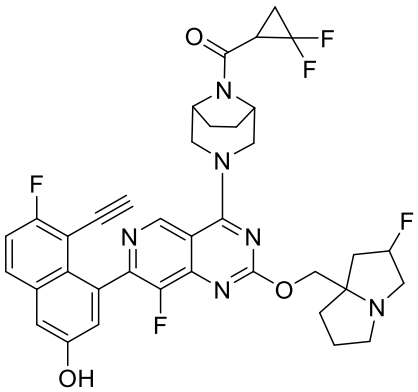
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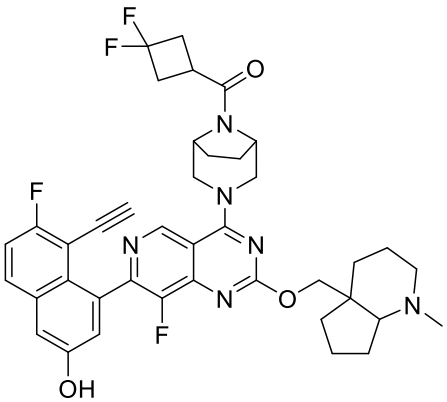
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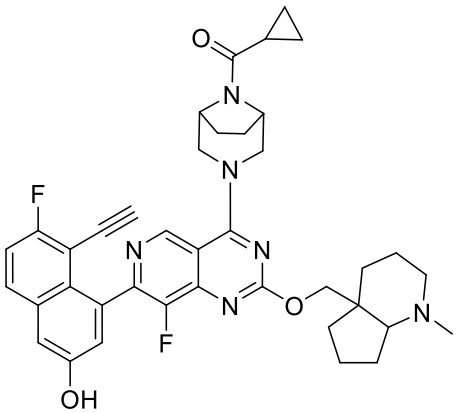
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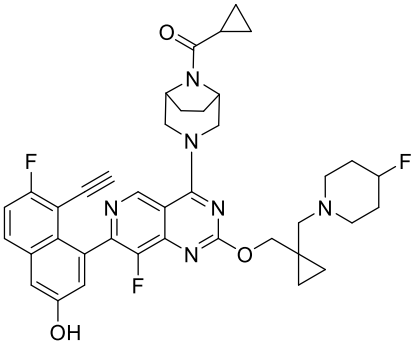
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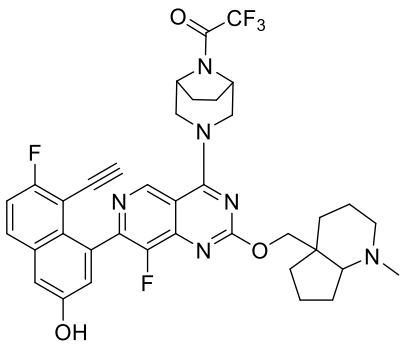
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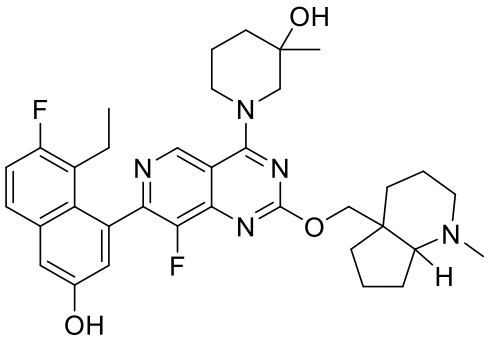
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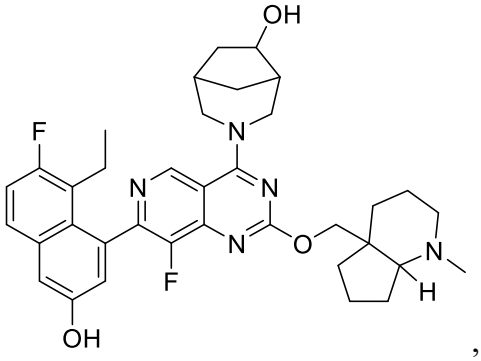
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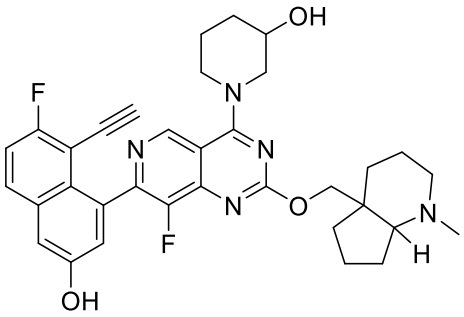
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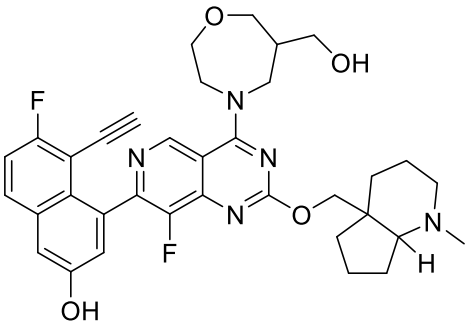
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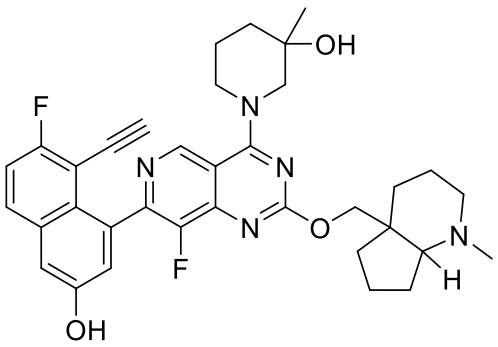
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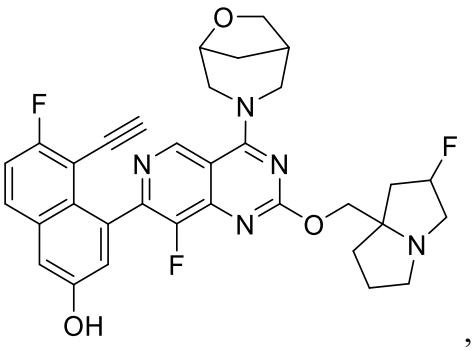
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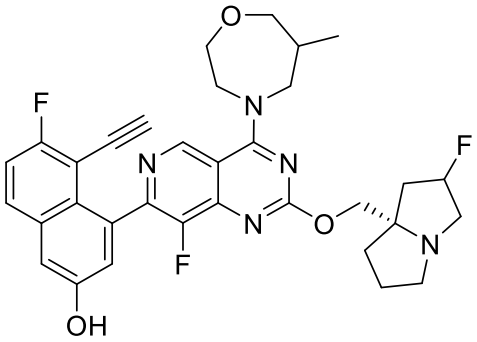
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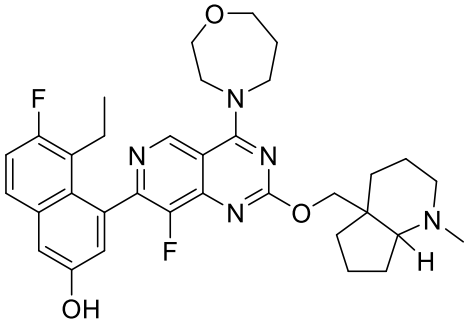
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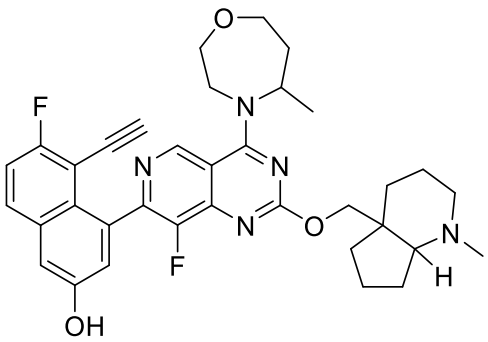
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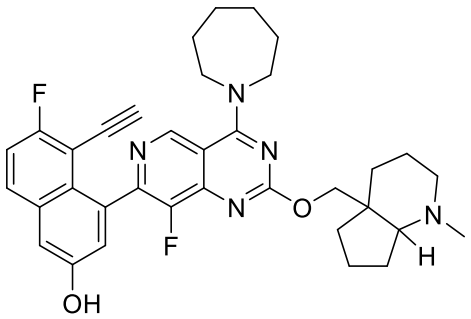
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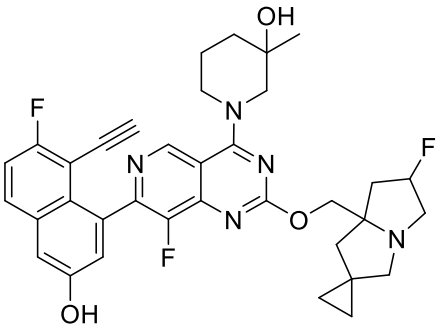
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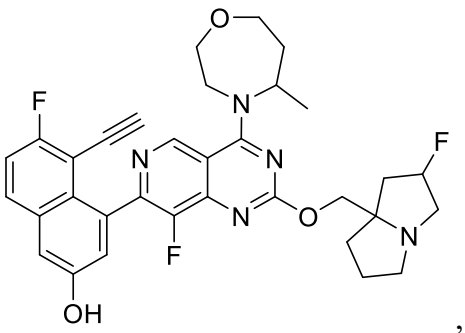
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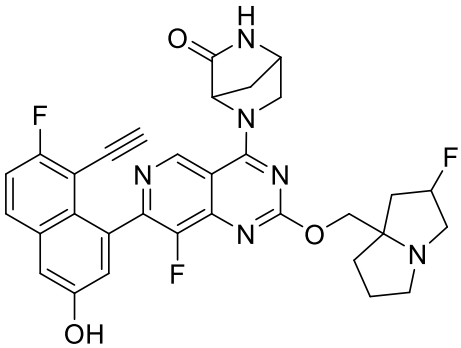
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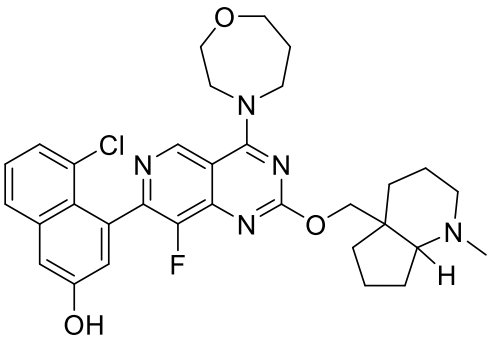
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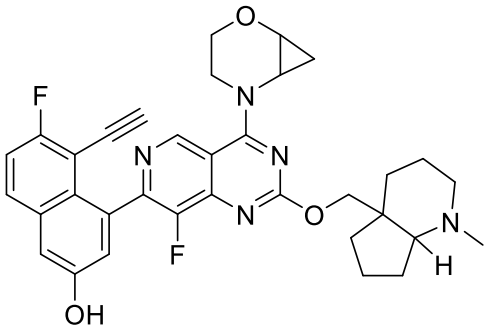
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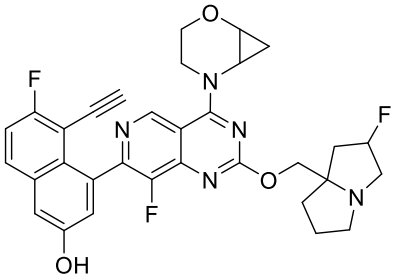
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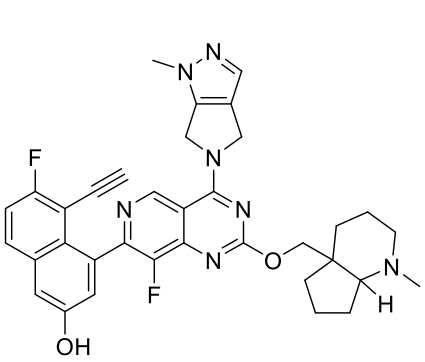
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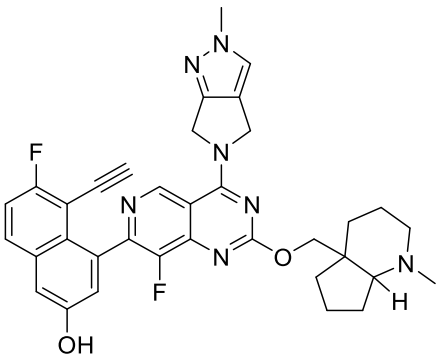
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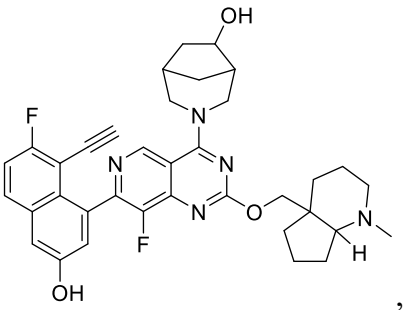
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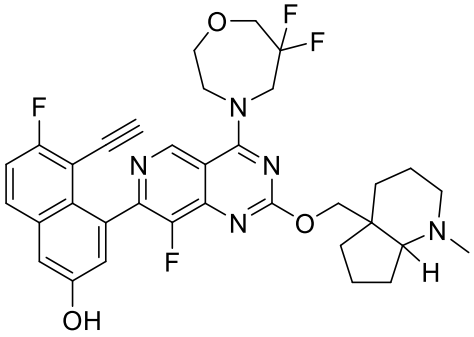
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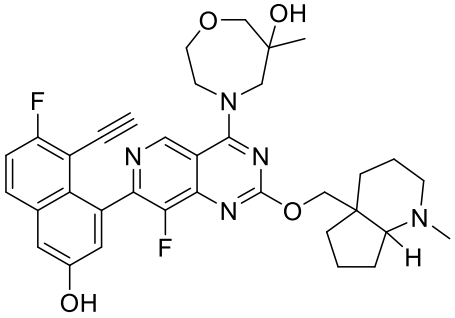
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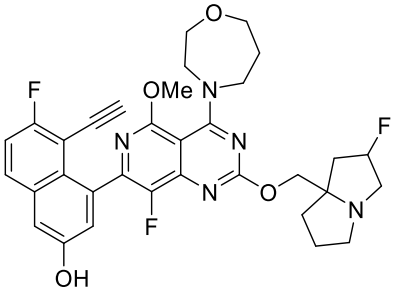
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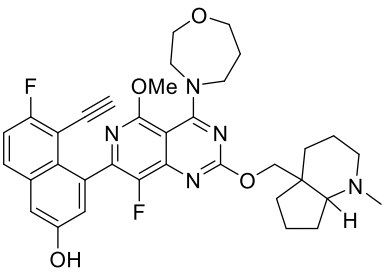
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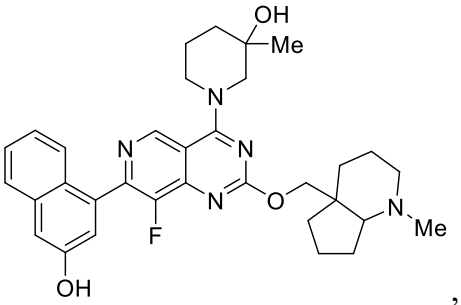
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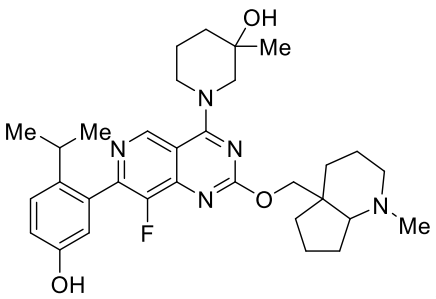
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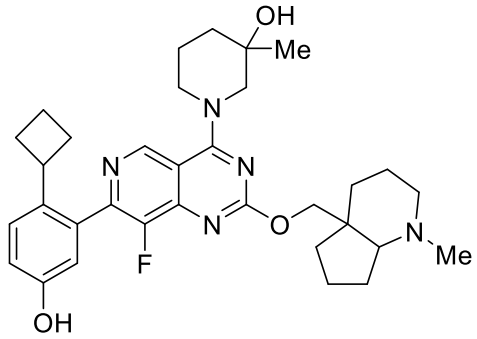
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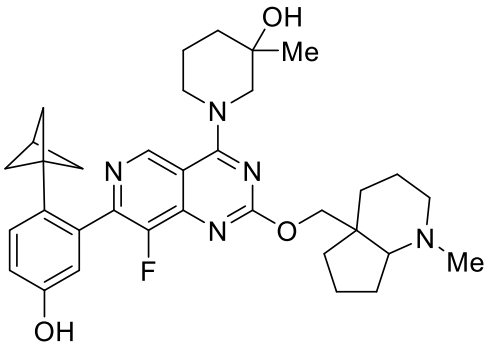
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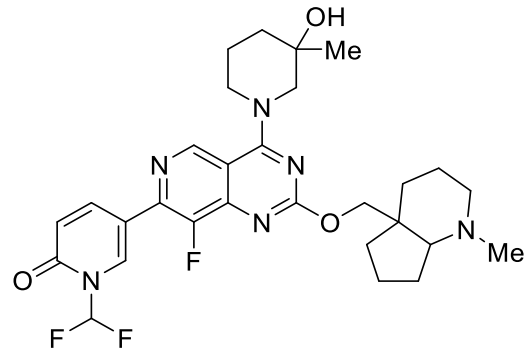
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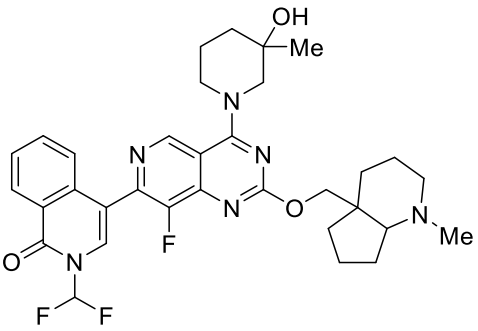
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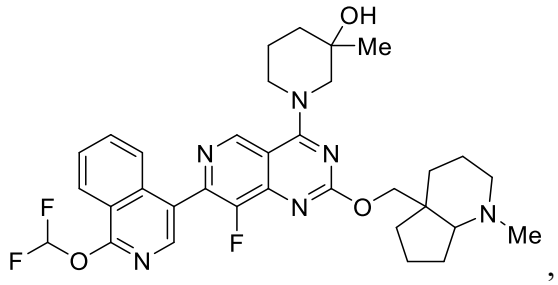
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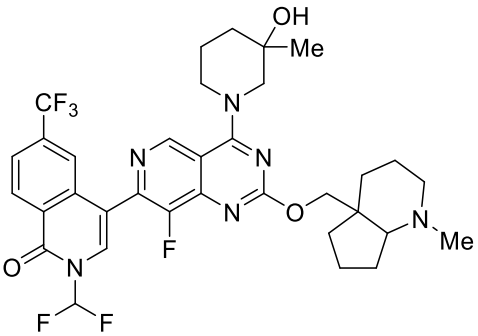
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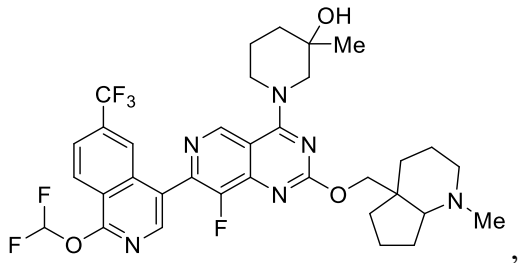
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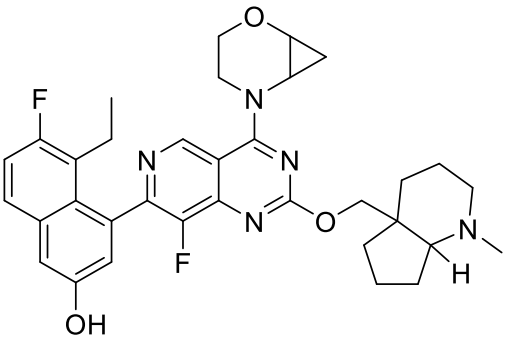
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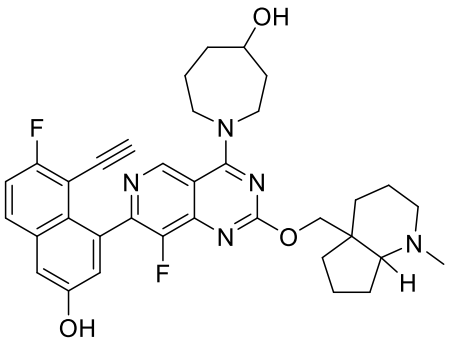
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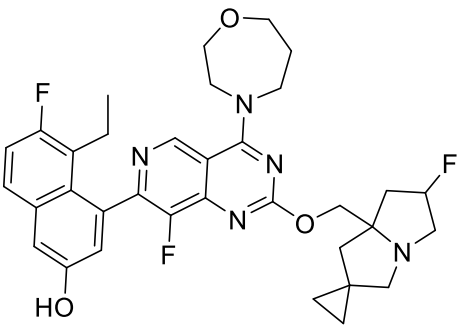
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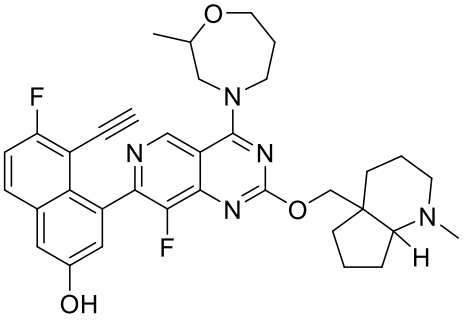
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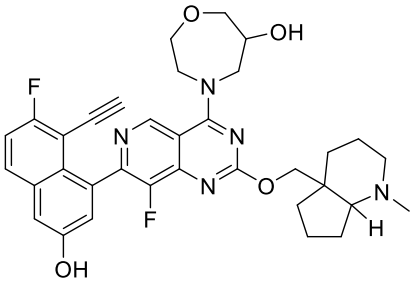
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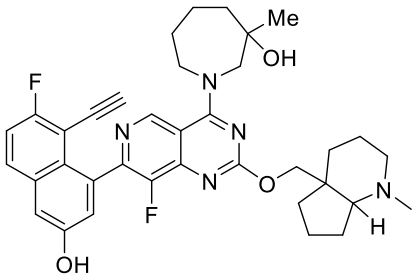
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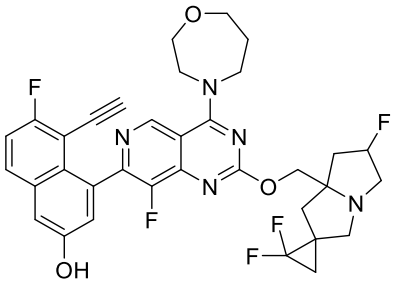
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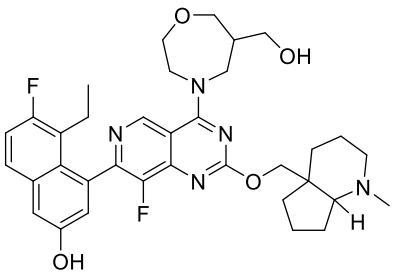
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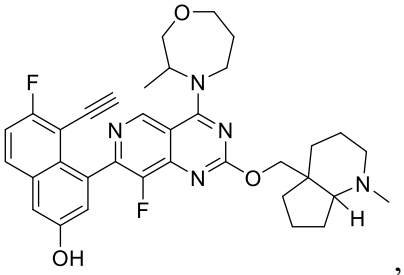
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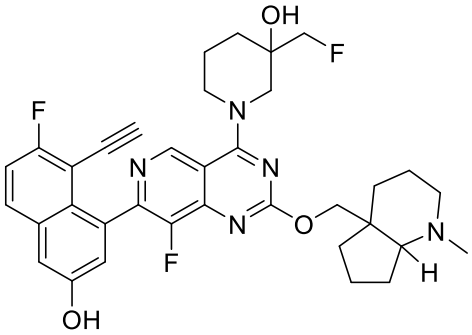
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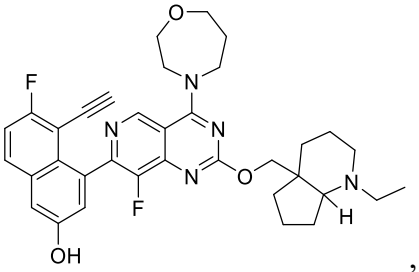
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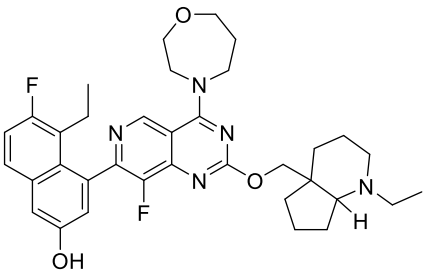
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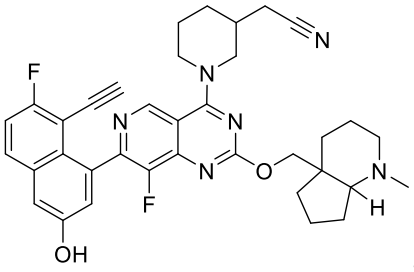
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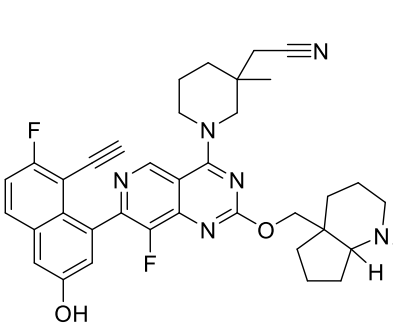
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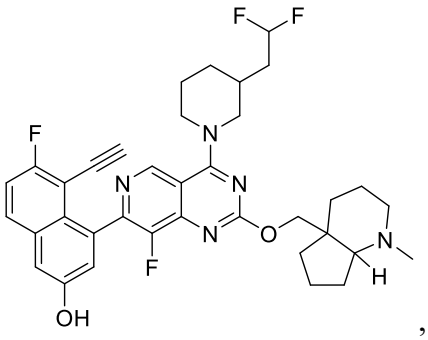
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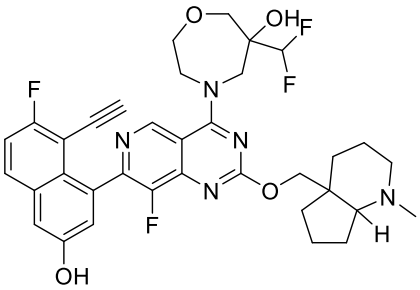
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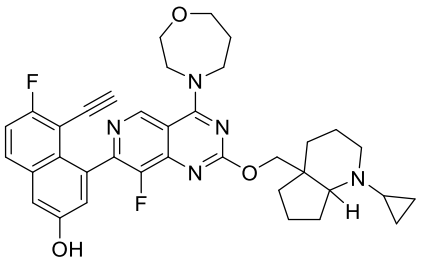
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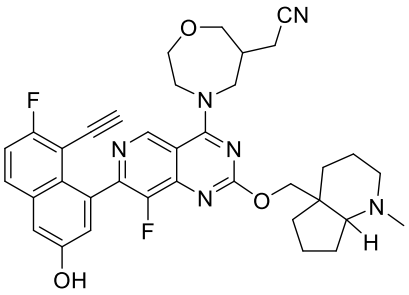
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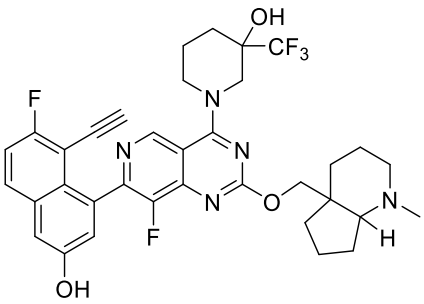
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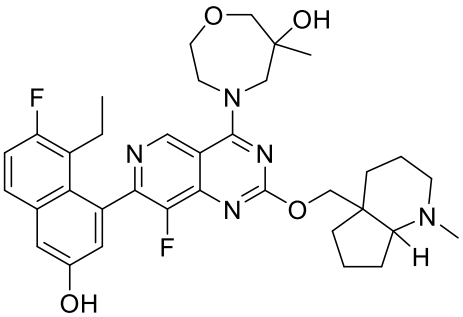
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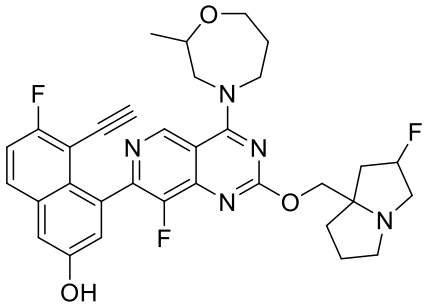
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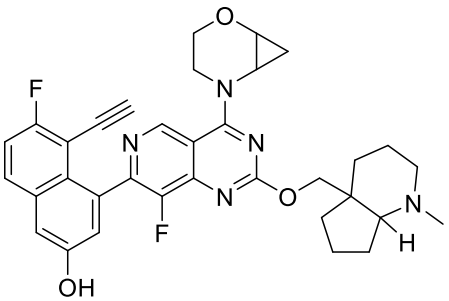
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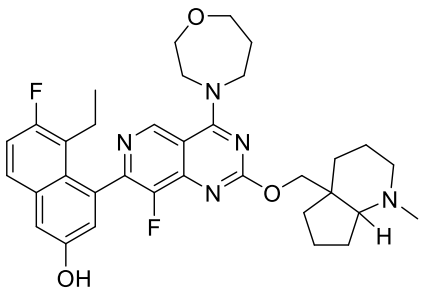
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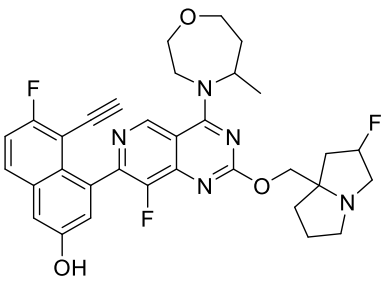
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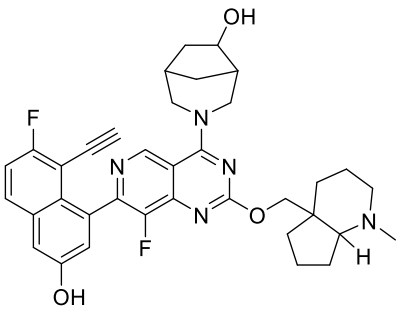
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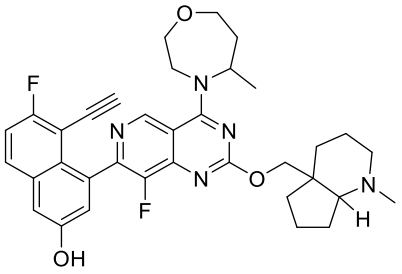
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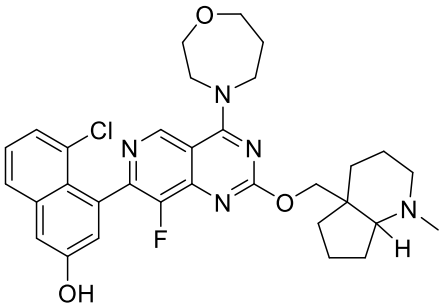
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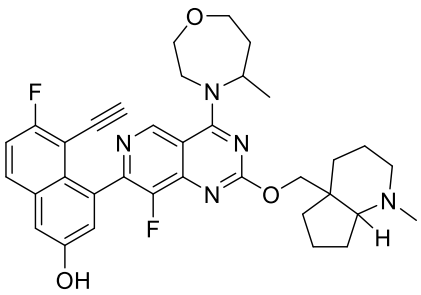
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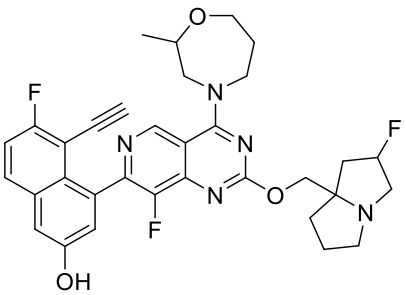
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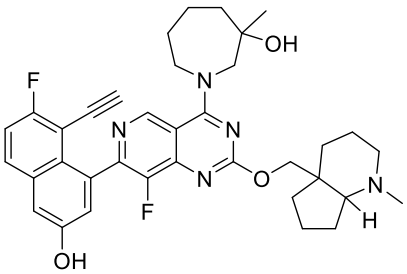
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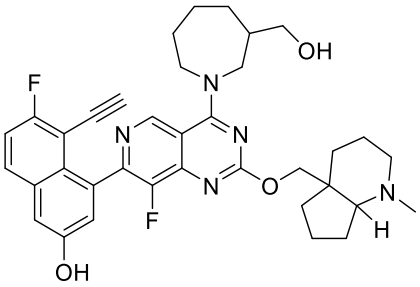
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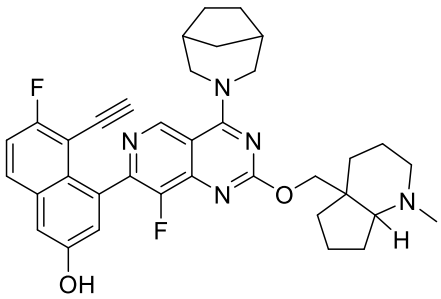
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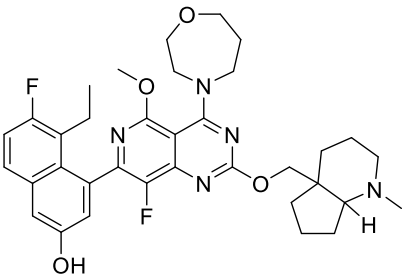
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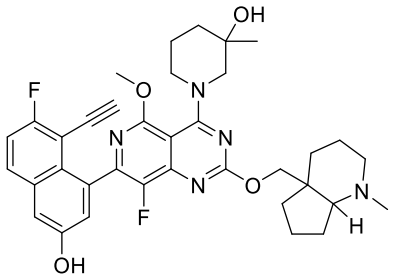
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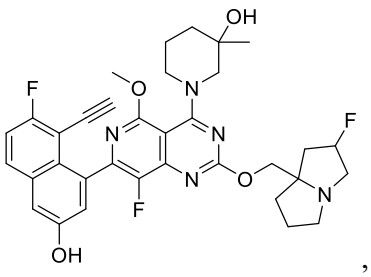
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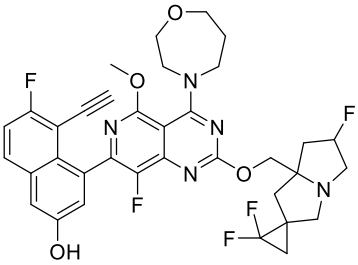
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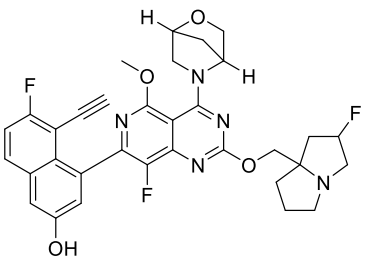
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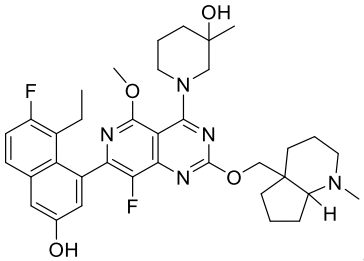
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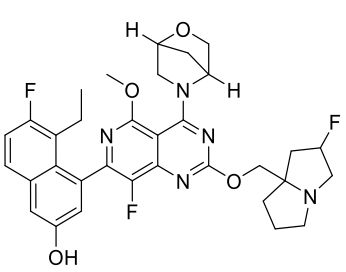
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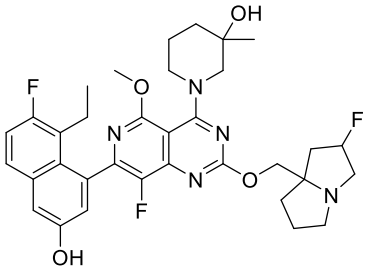
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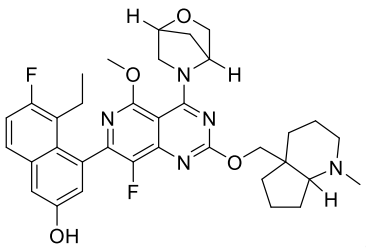
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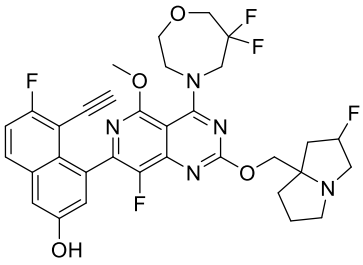
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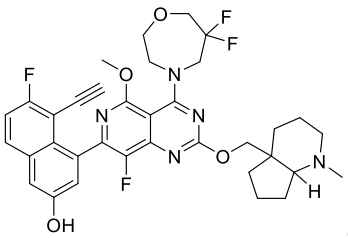
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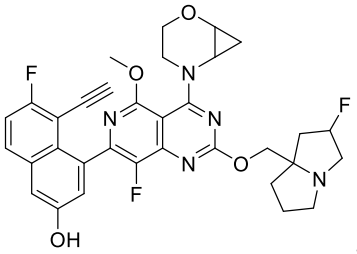
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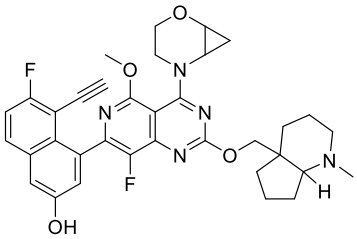
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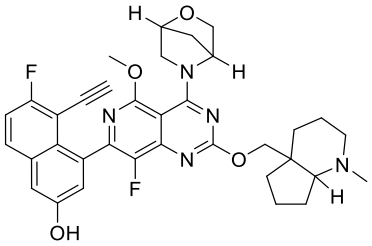
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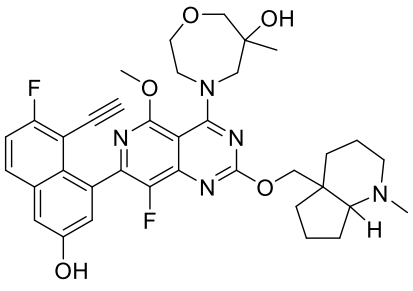
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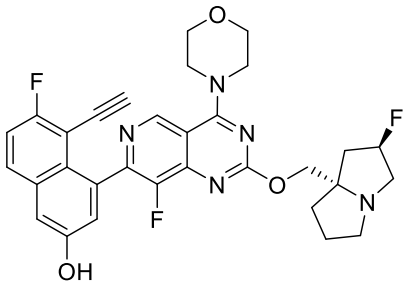
or a pharmaceutically acceptable salt thereof.

43. The compound of claim 40 that is selected from the group consisting of: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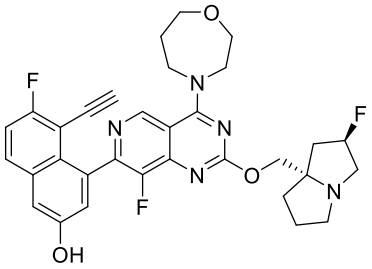
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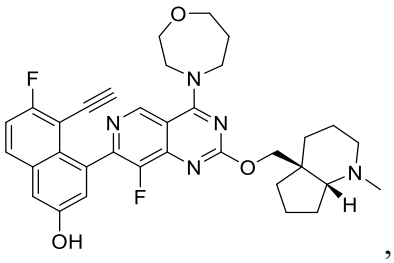
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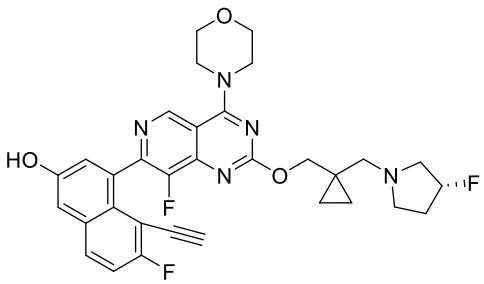
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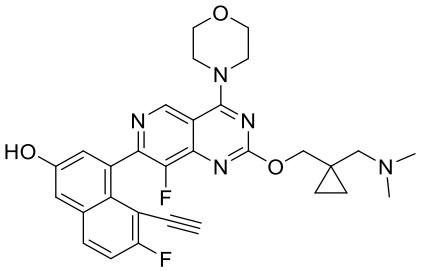
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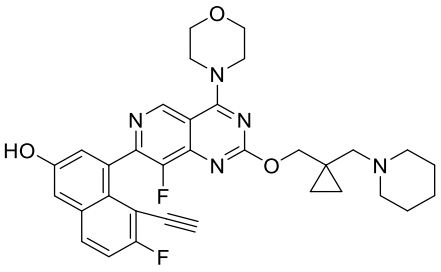
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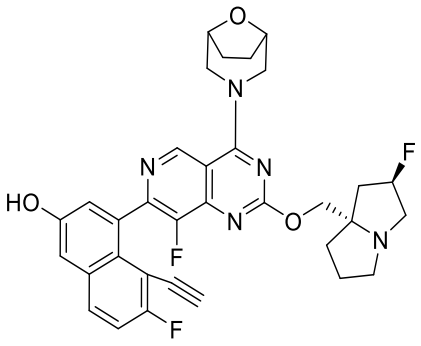
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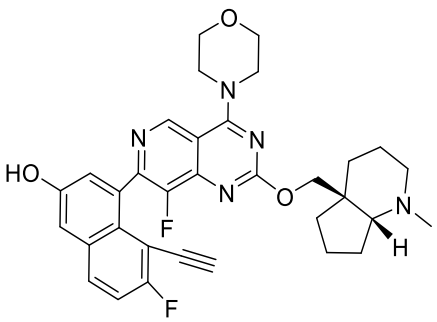
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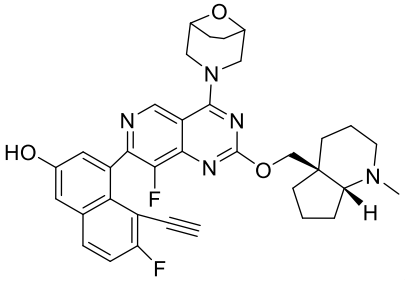
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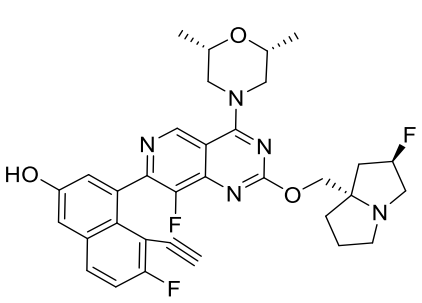
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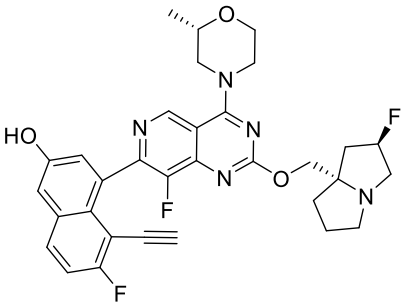
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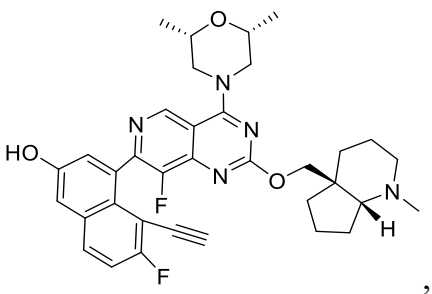
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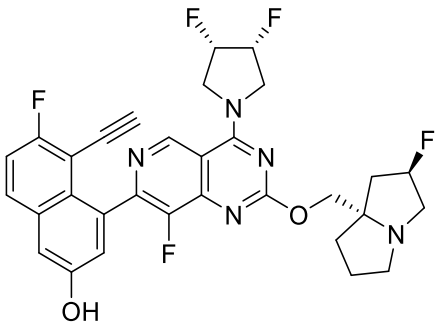
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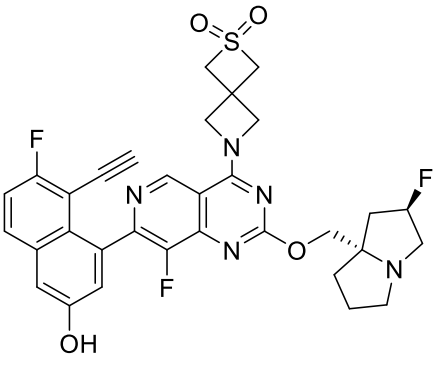
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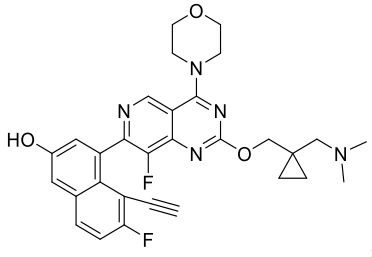
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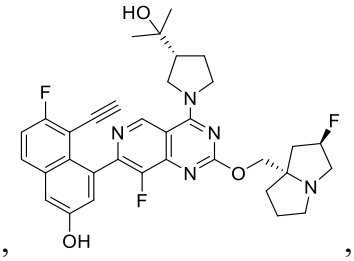
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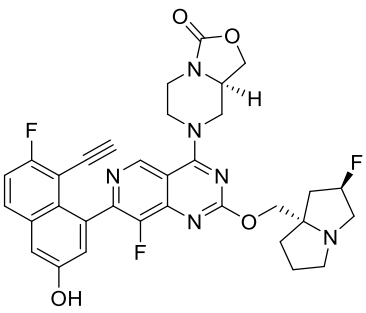
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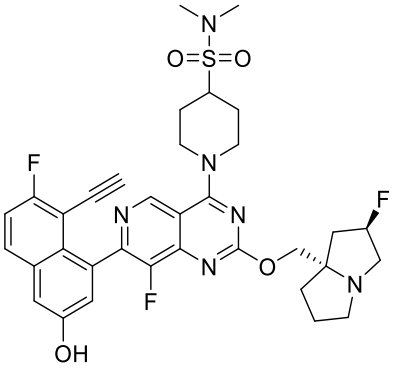
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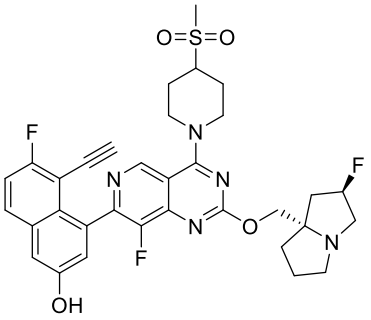
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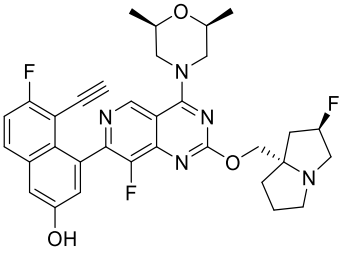
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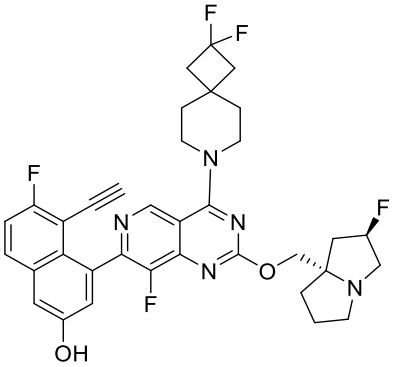
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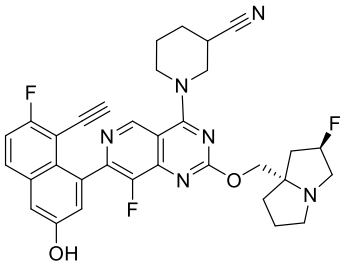
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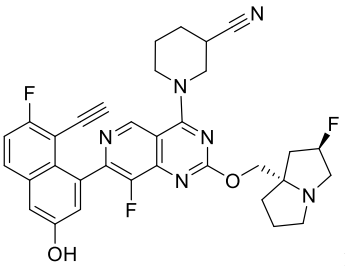
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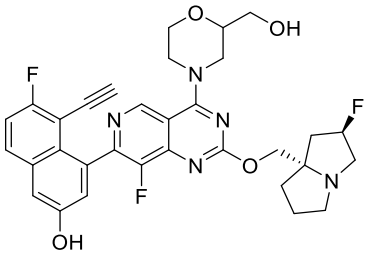
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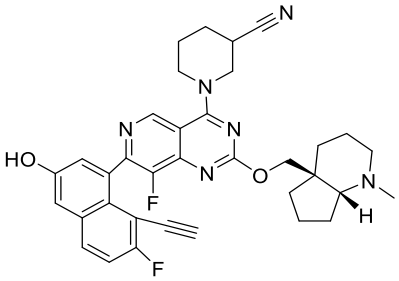
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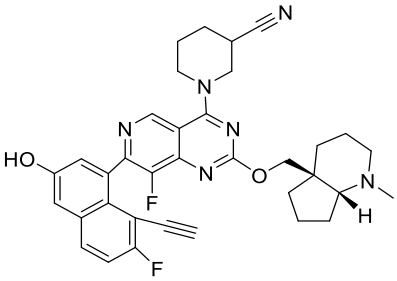
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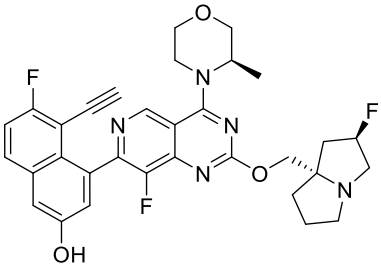
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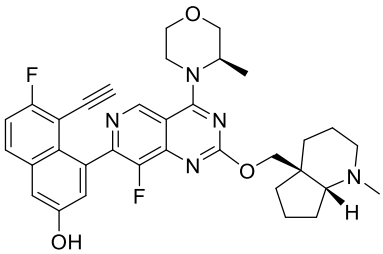
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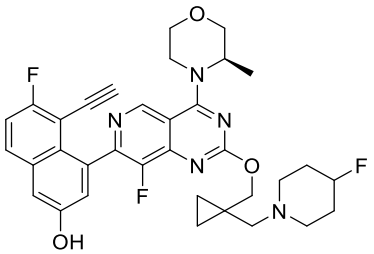
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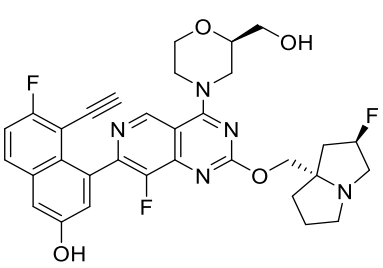
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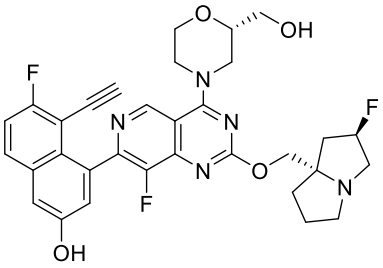
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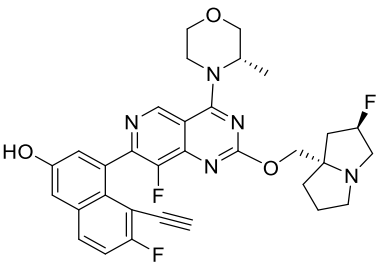
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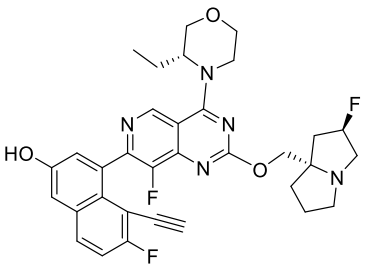
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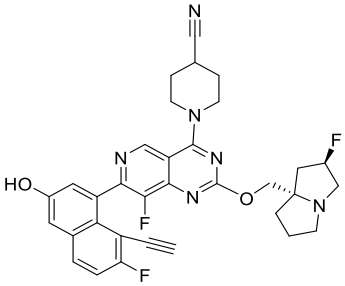
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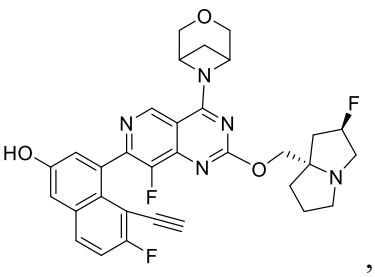
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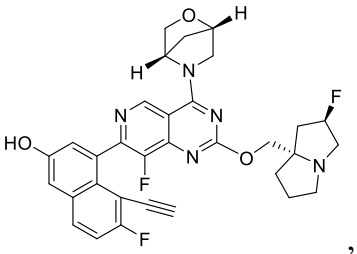
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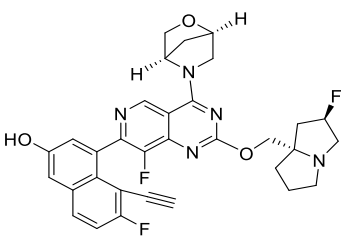
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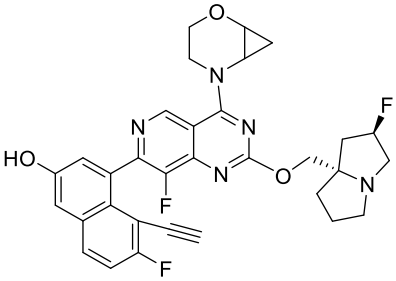
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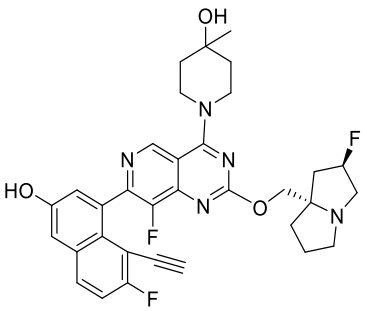
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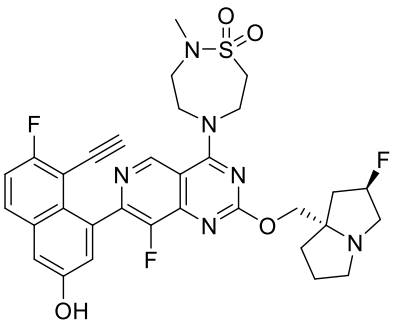
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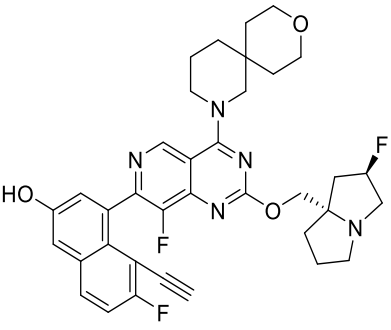
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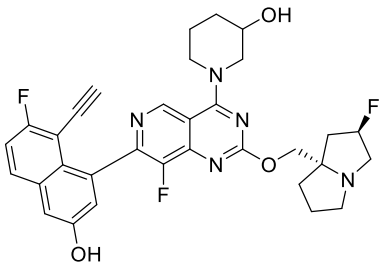
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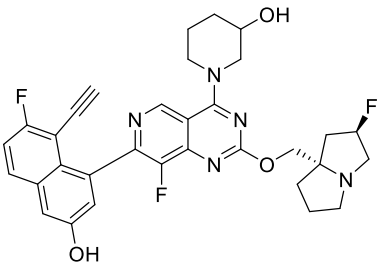
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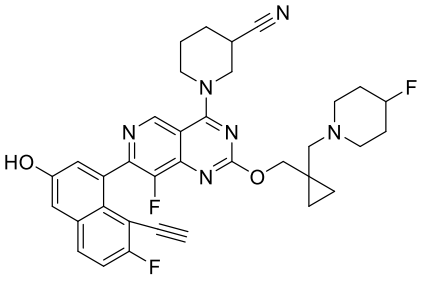

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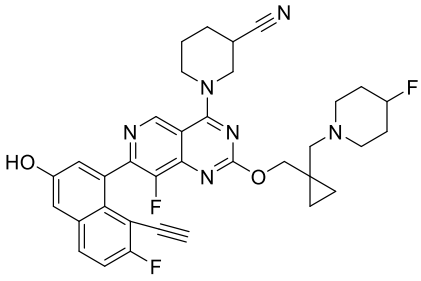

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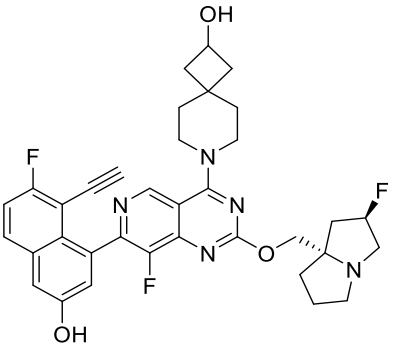

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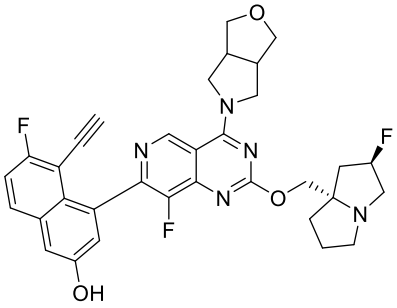

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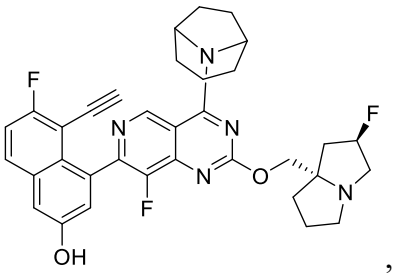

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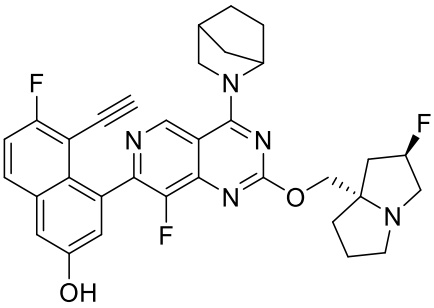

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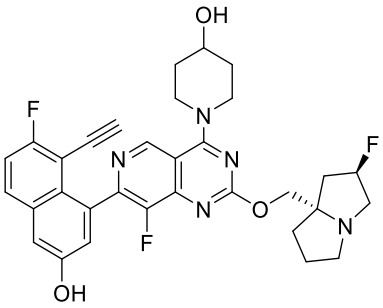

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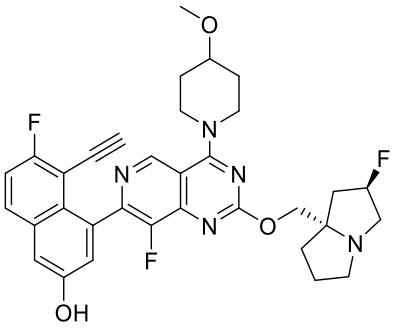

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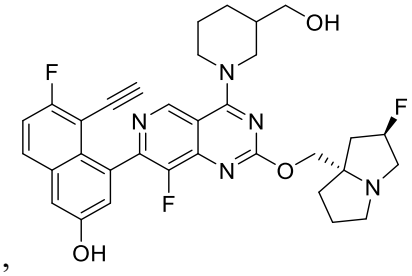

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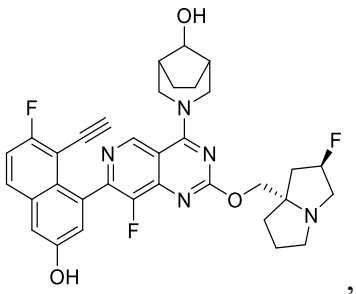

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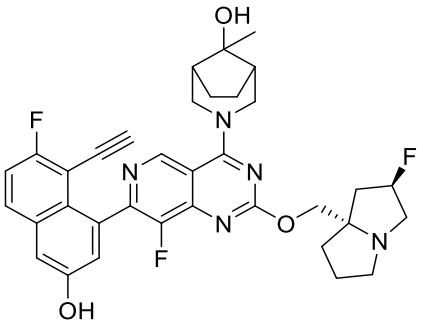

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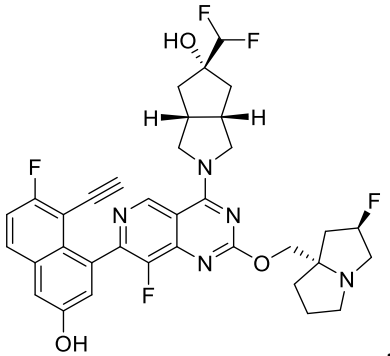

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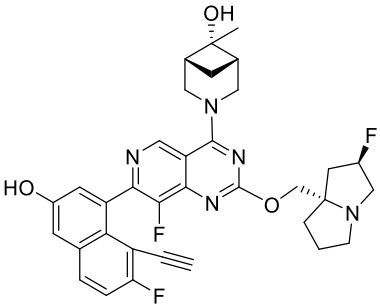

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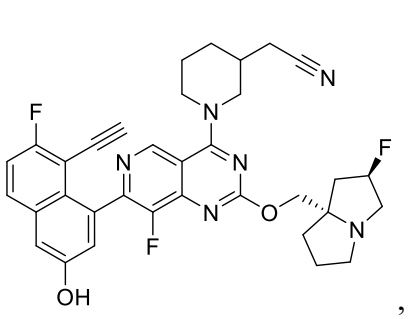

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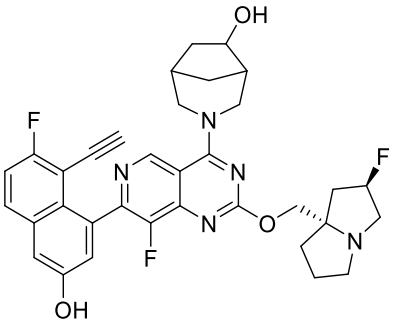
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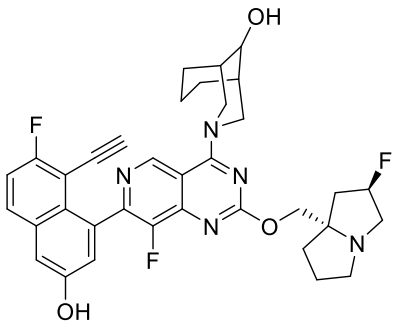
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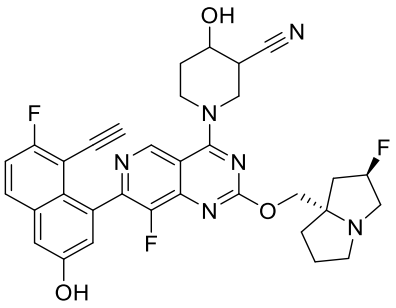

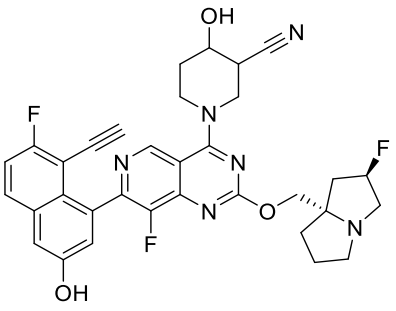
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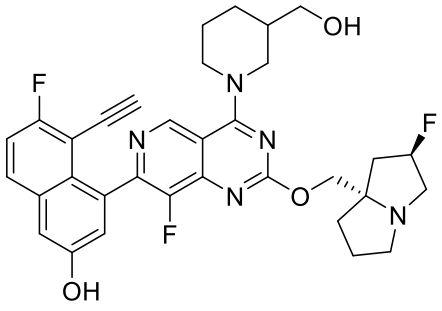
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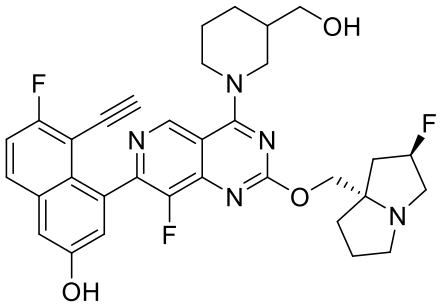
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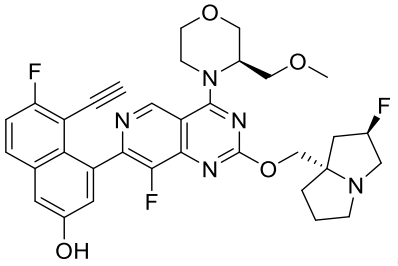
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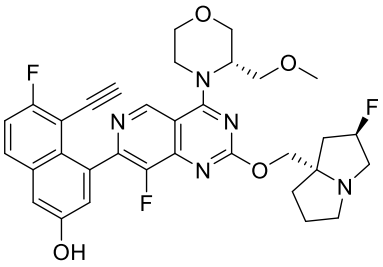
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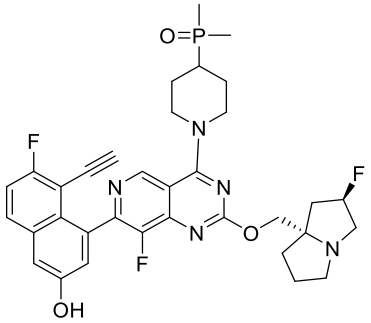
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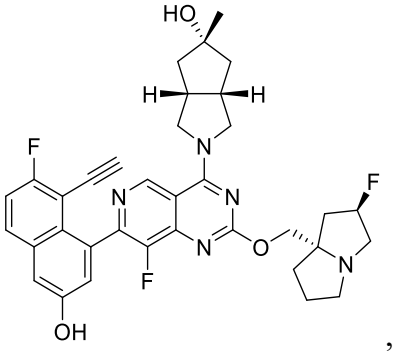
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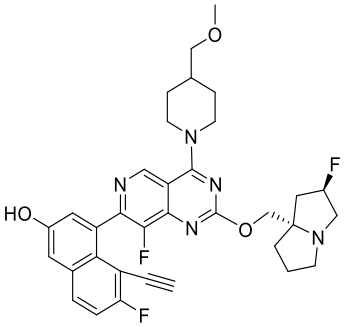
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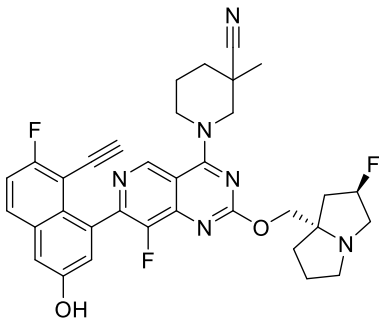
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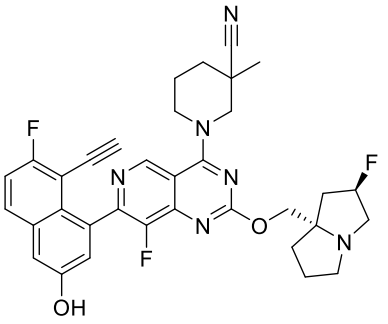
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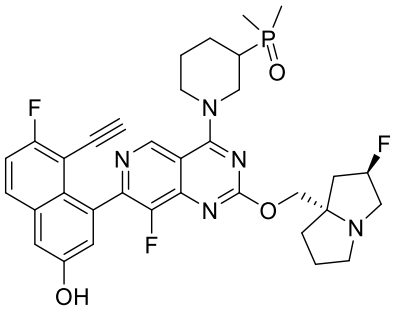
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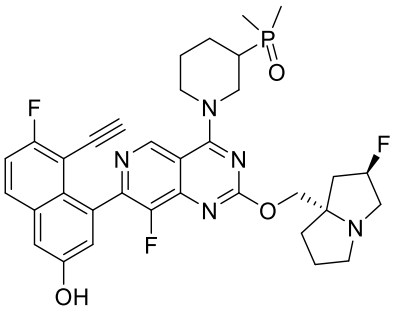
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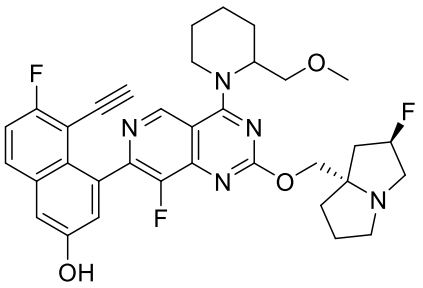
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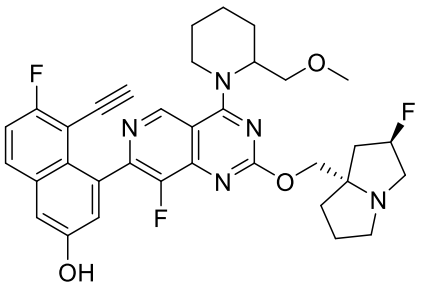
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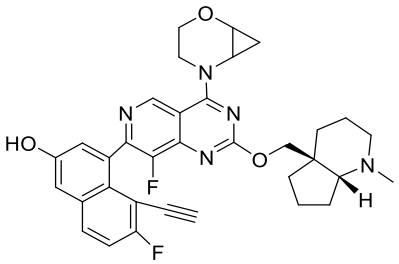
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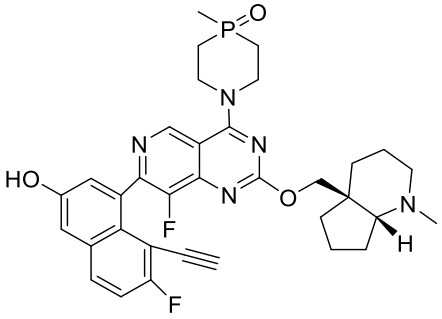
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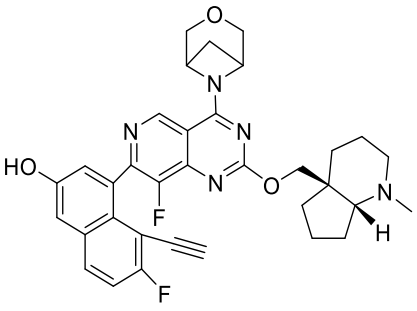
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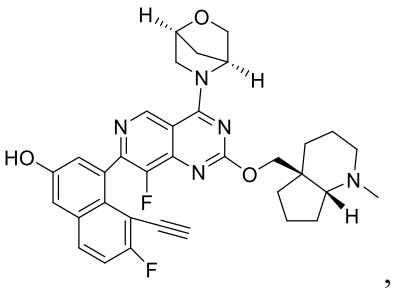
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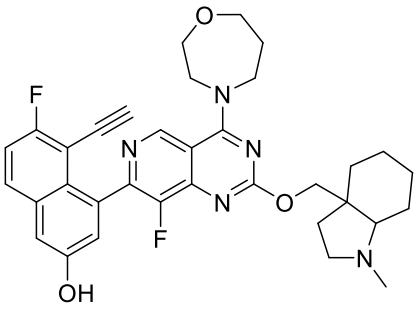
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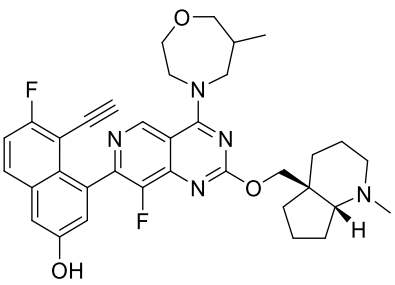
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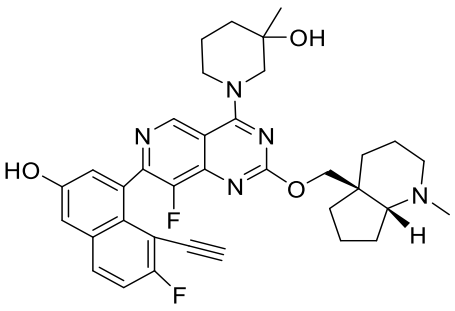
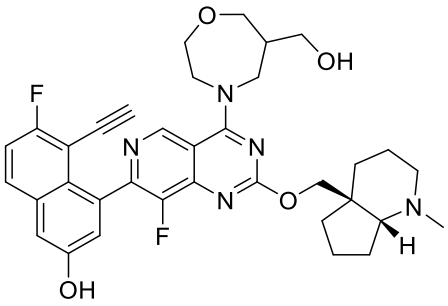
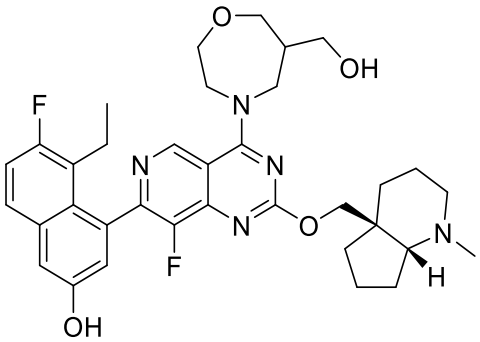
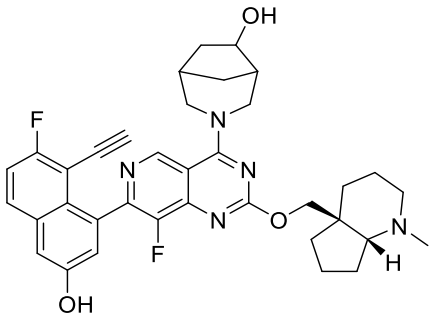
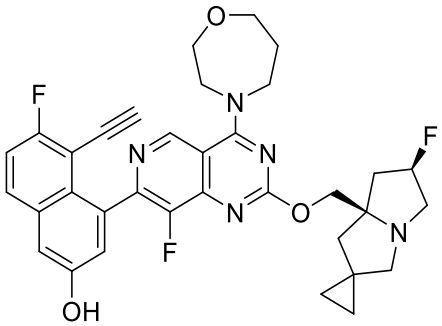
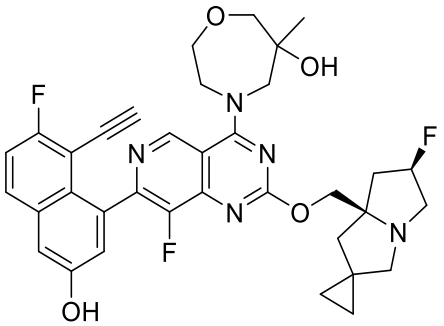
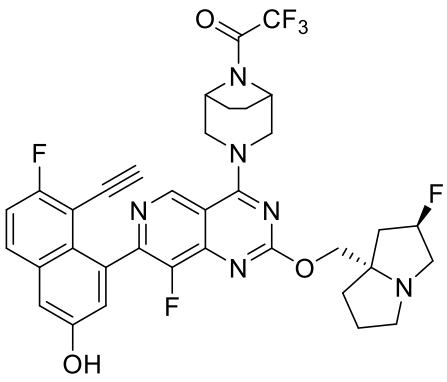
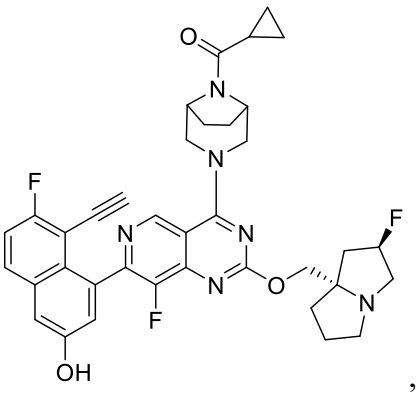

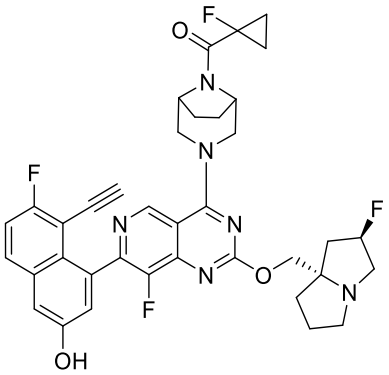
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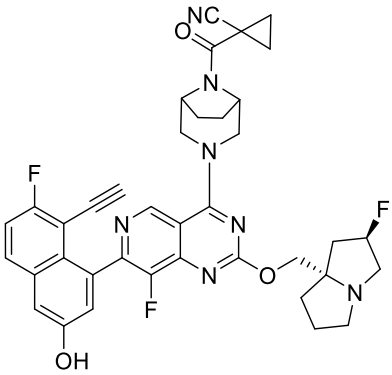
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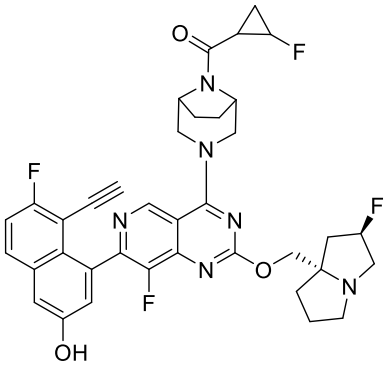
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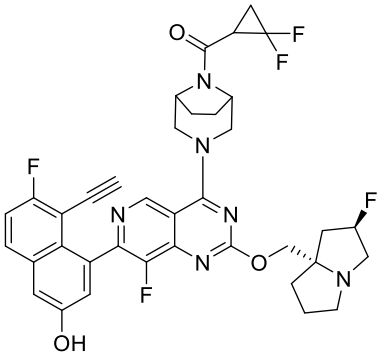
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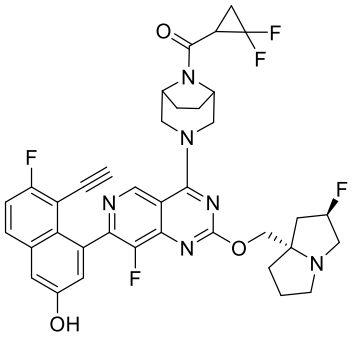
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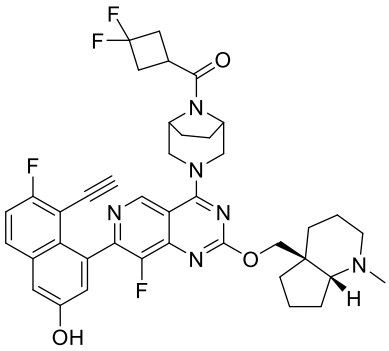
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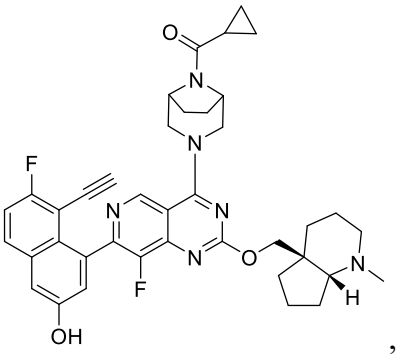
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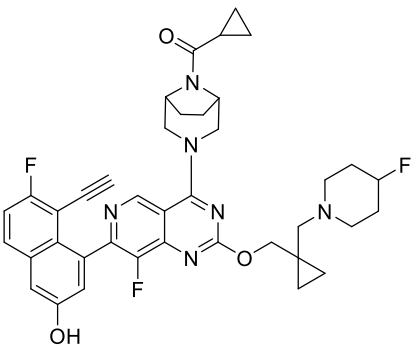
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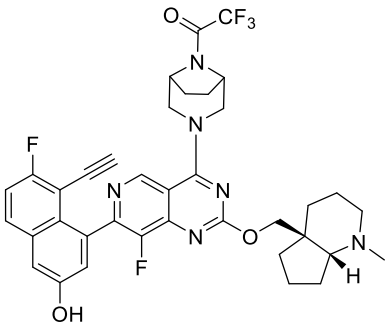
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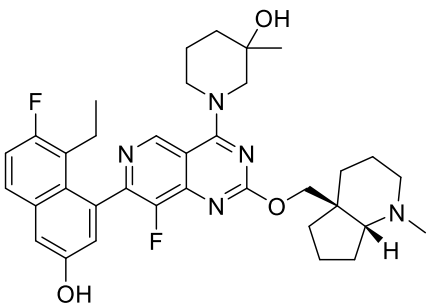
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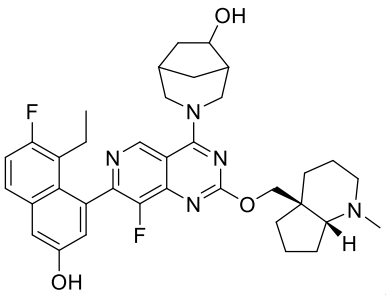
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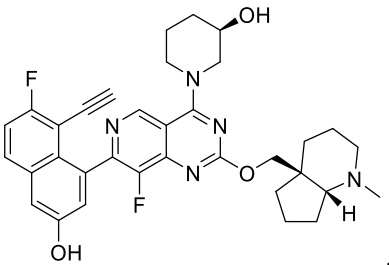
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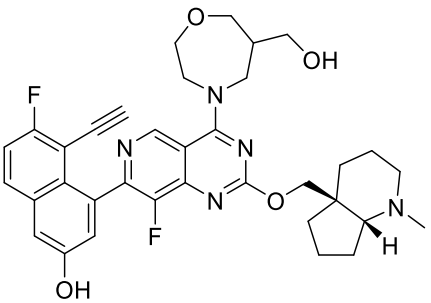
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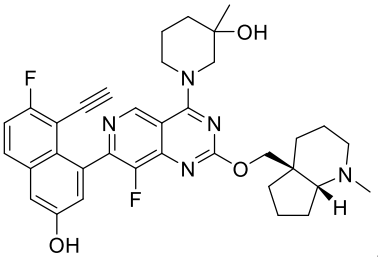
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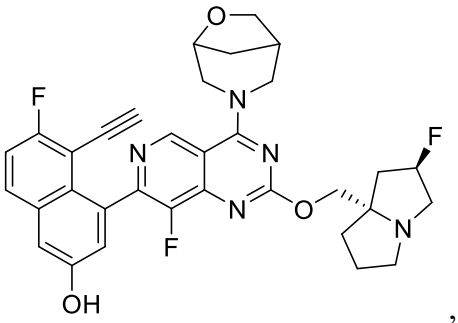
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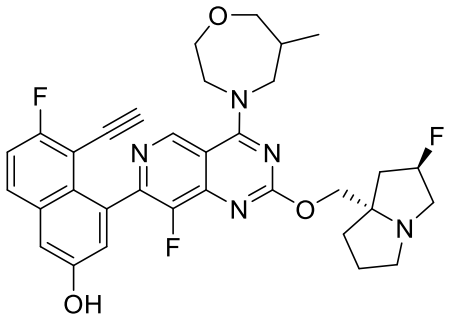
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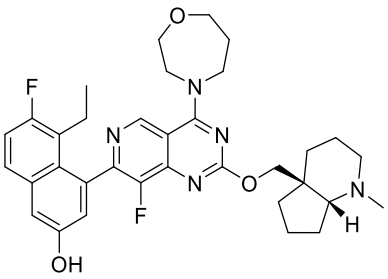
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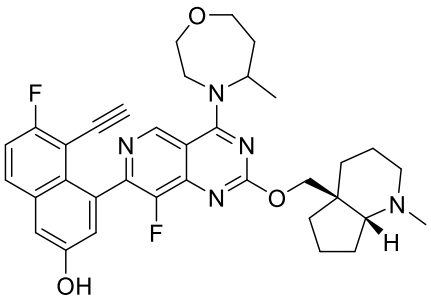
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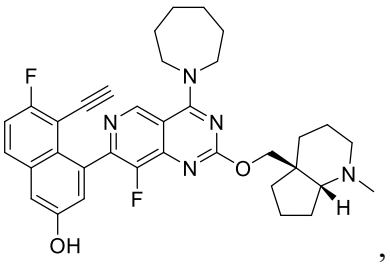
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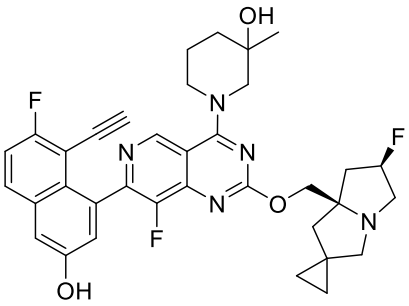
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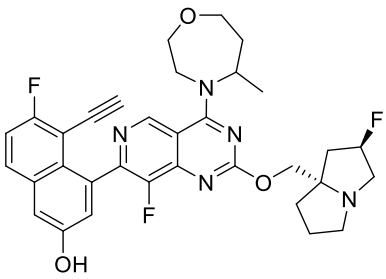
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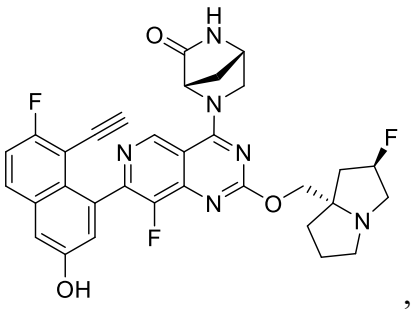
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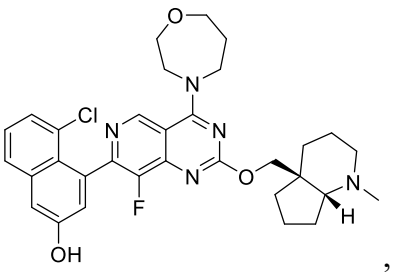
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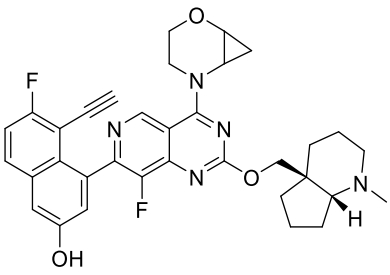
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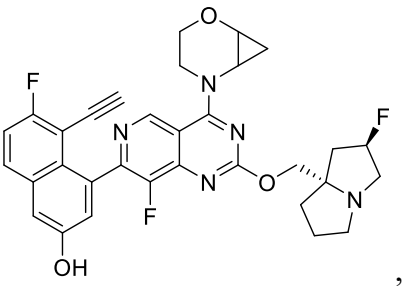
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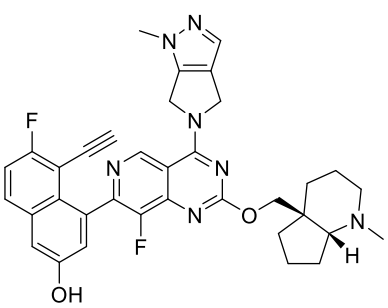
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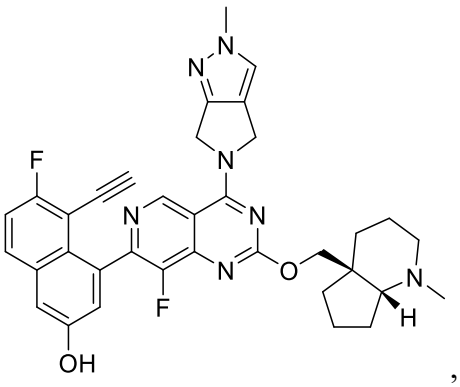
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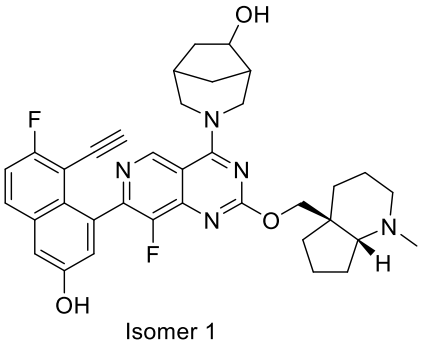


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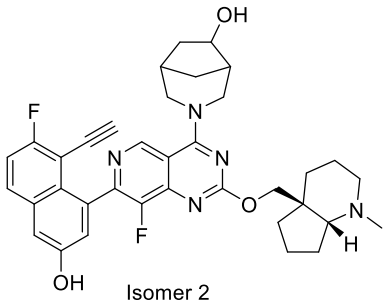


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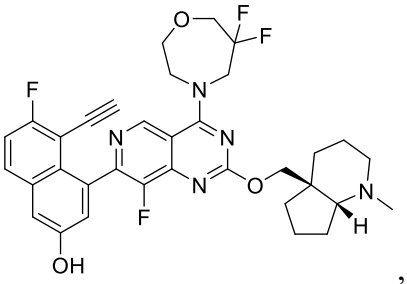


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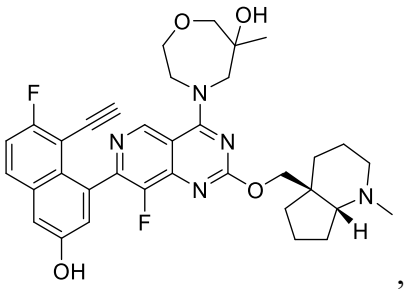


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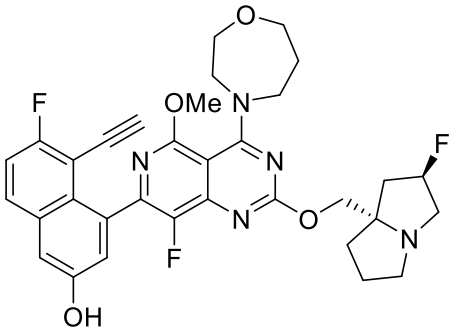


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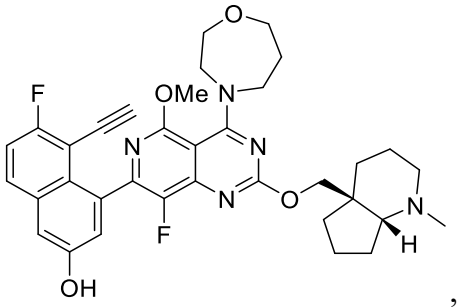


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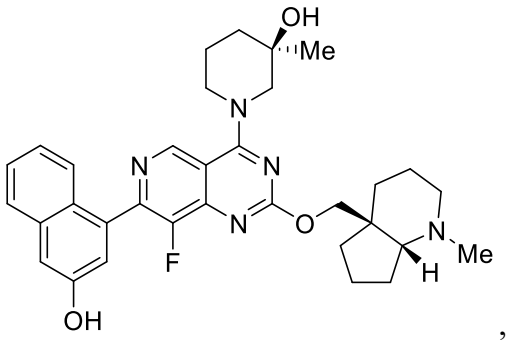


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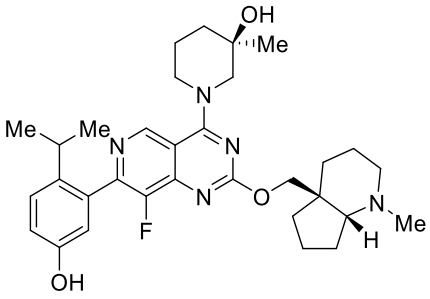


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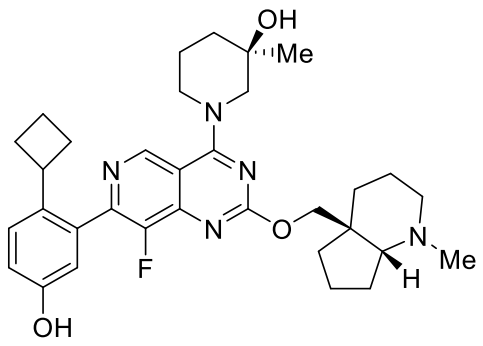


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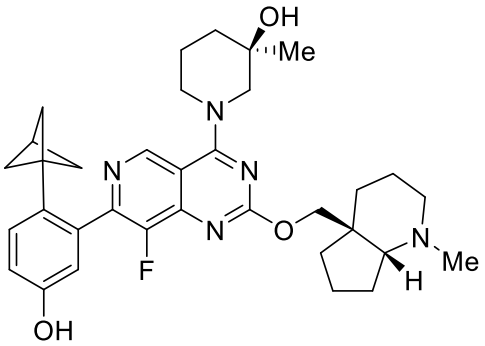
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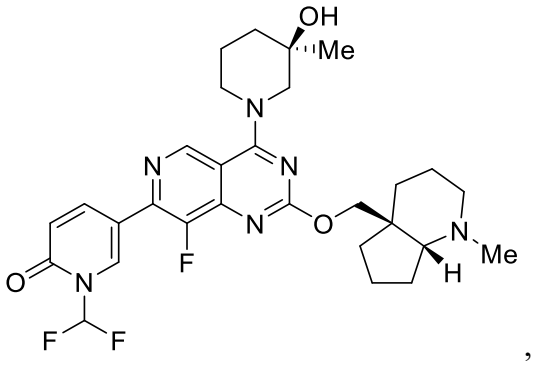
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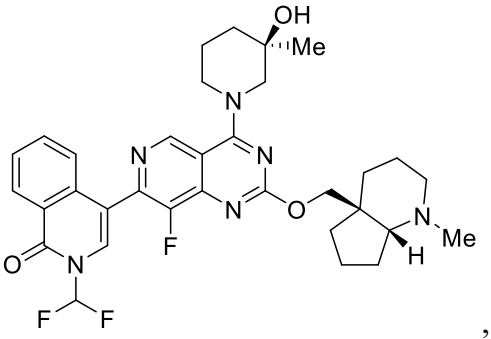
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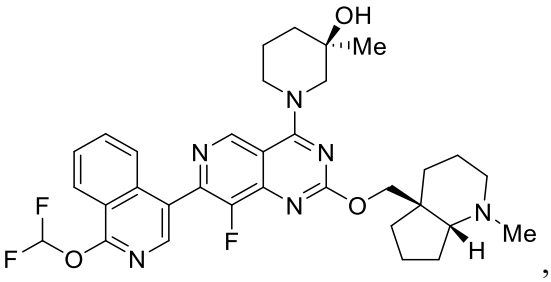
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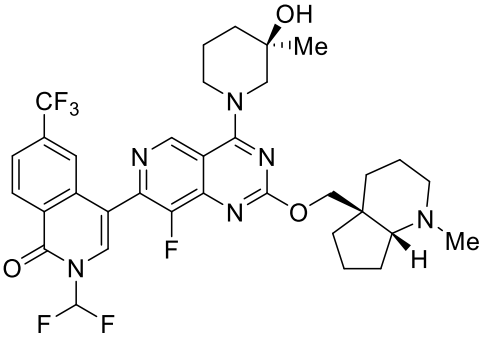
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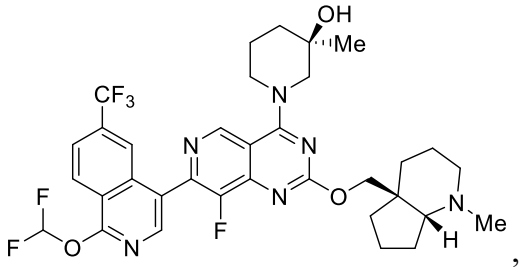
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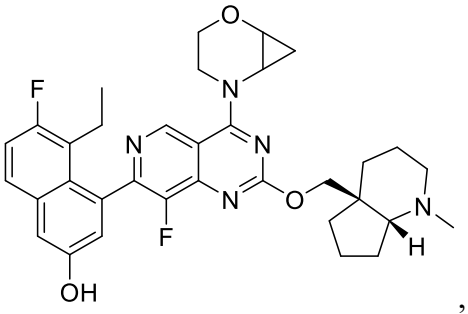
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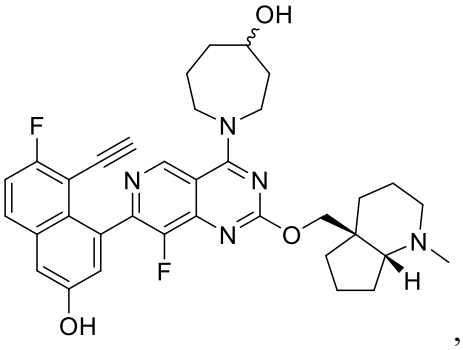
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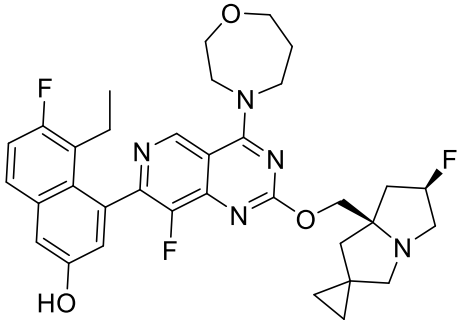

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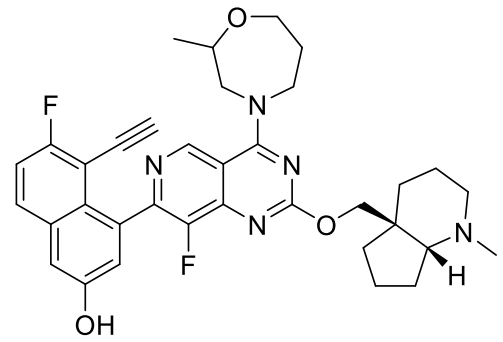

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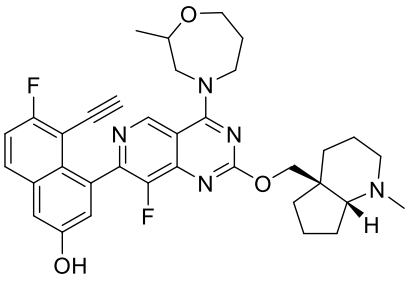

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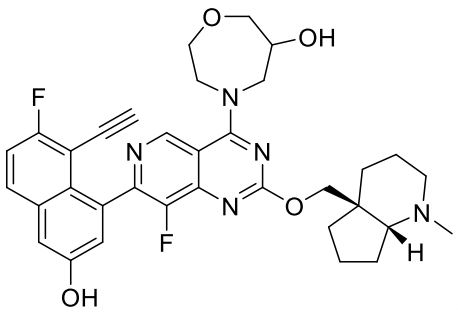

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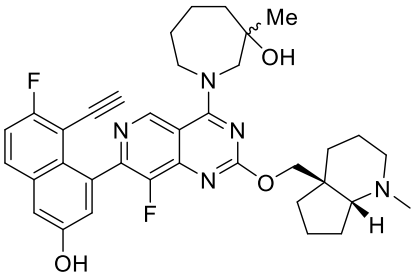

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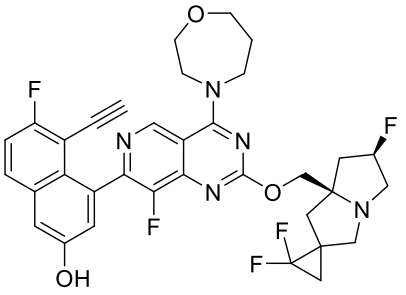

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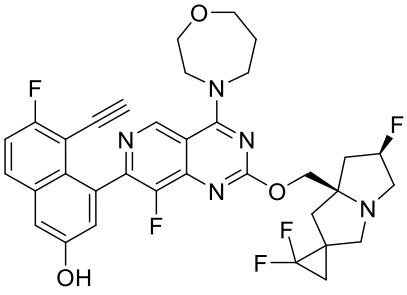

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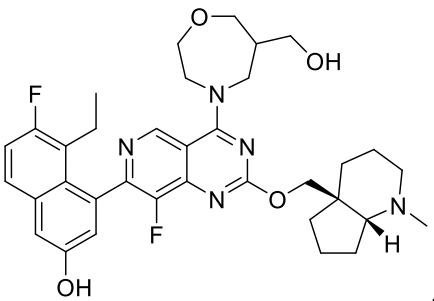

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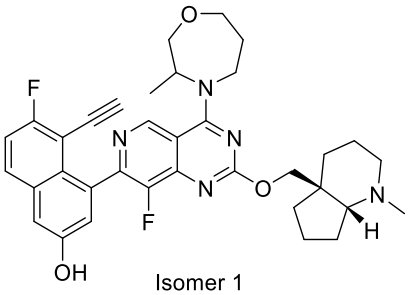
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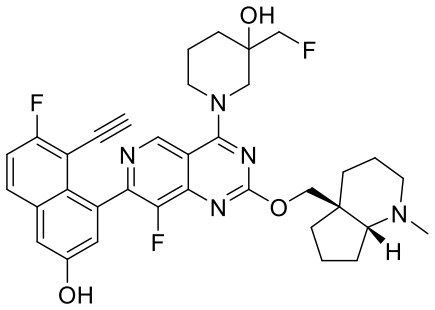
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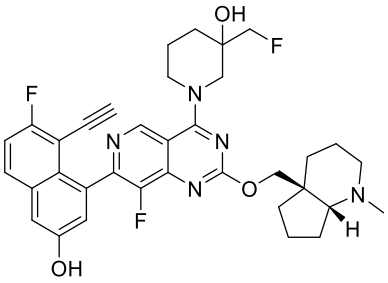
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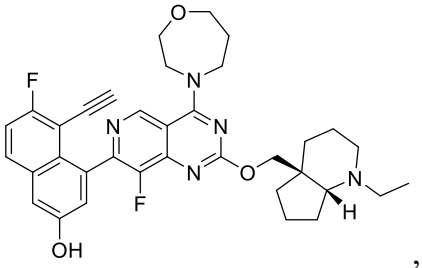

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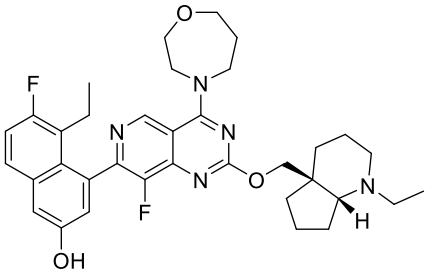

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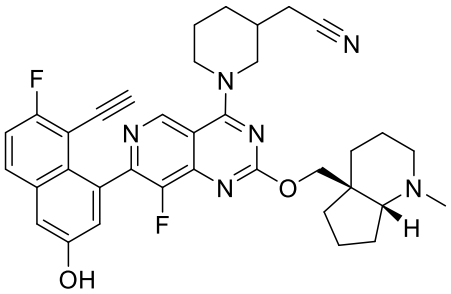

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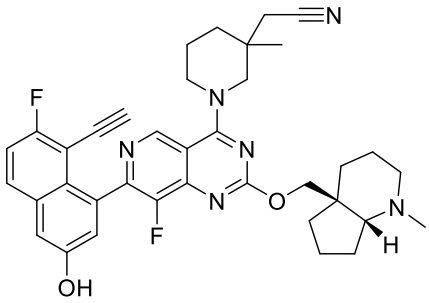

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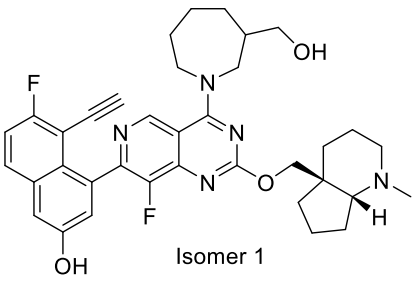
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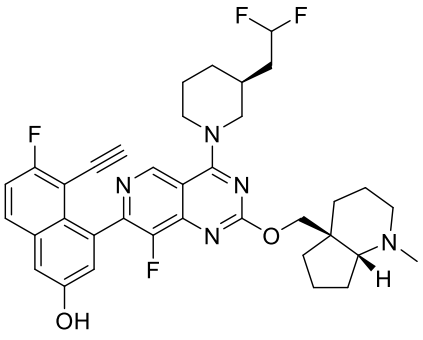

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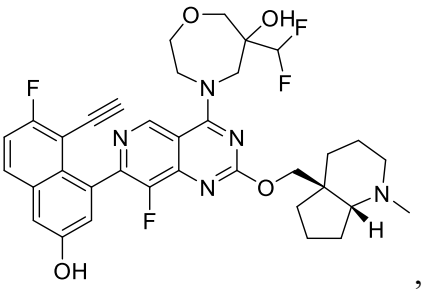

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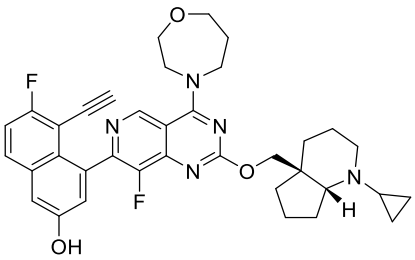
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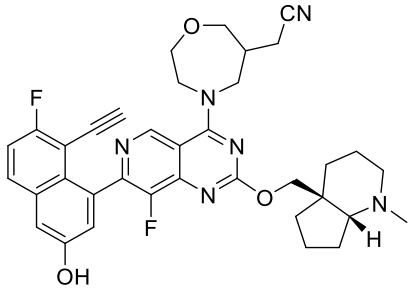
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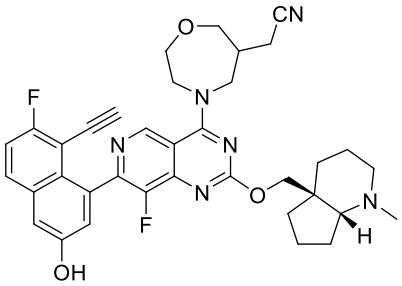
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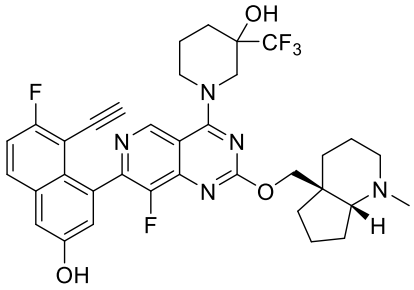
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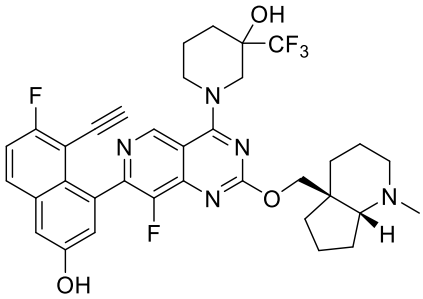
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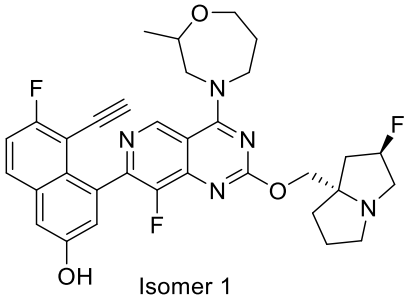
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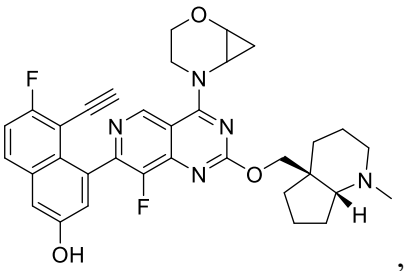
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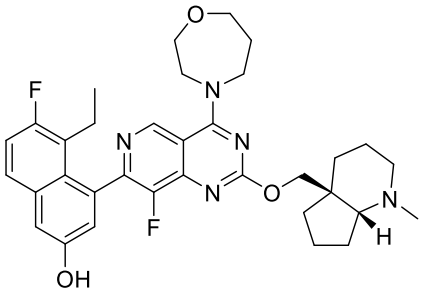
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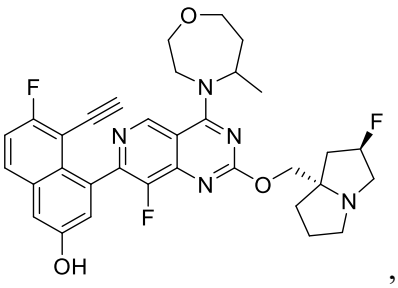
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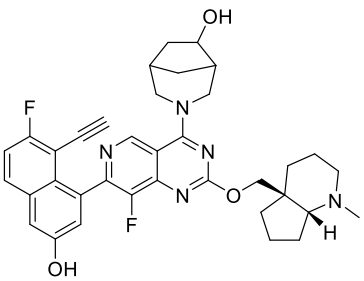
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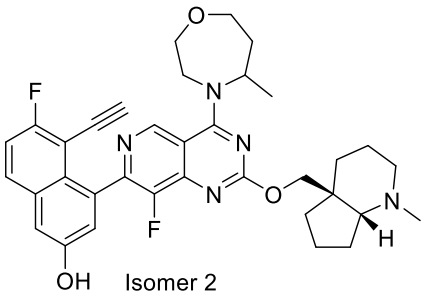
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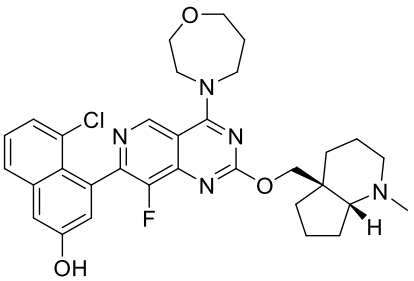
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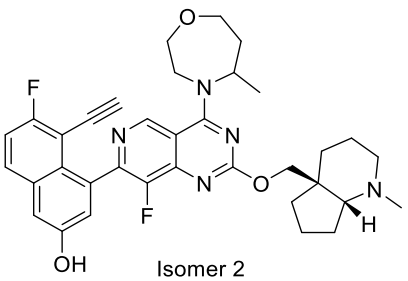
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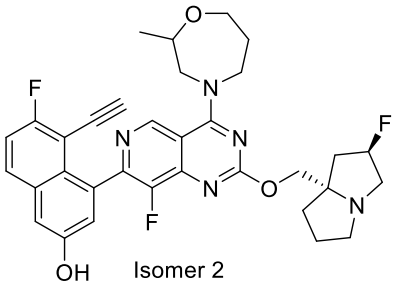
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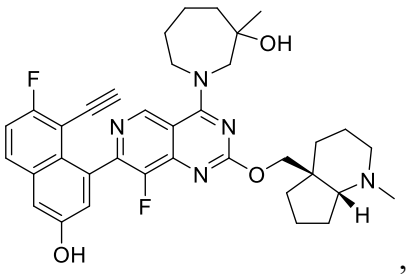
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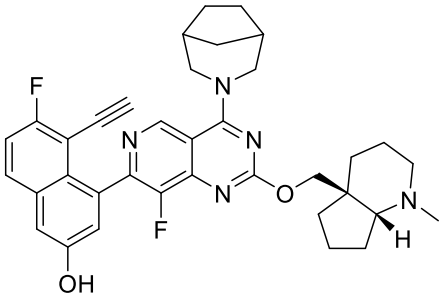
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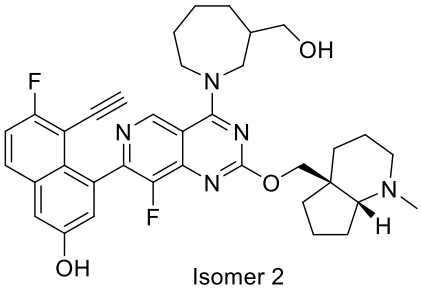
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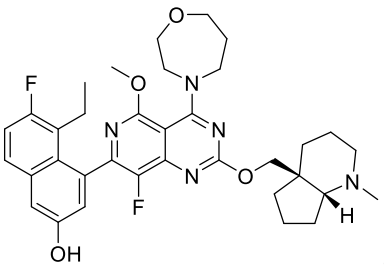
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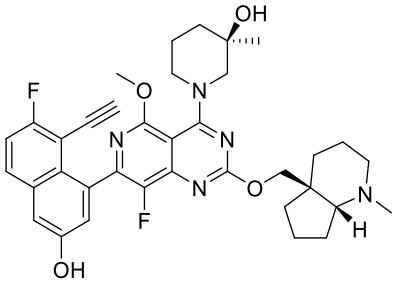
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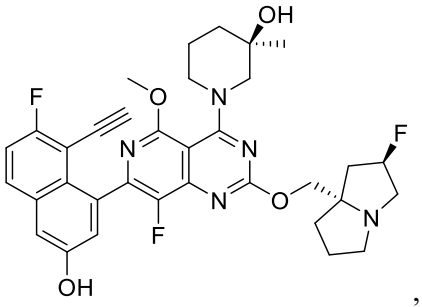
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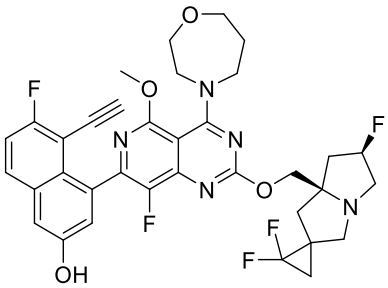
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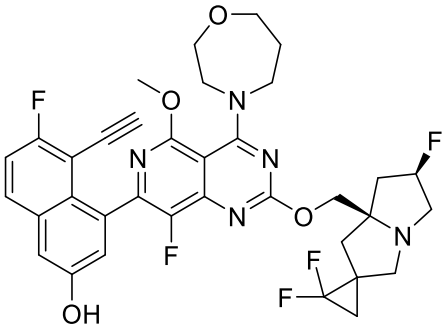
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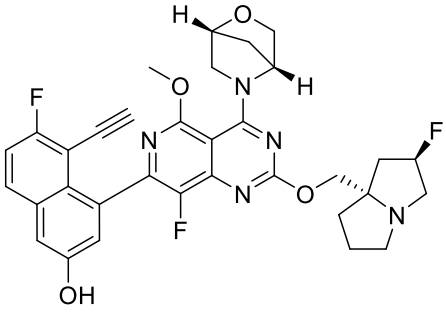
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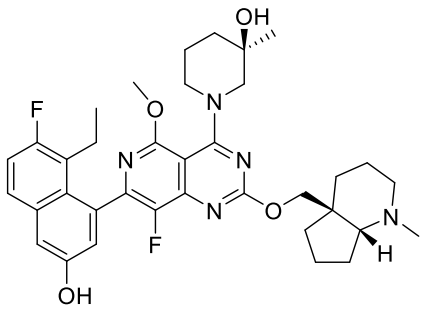
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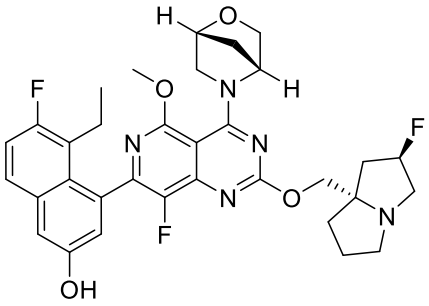
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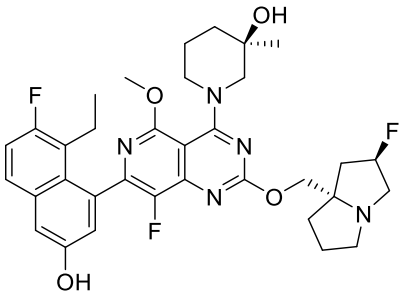
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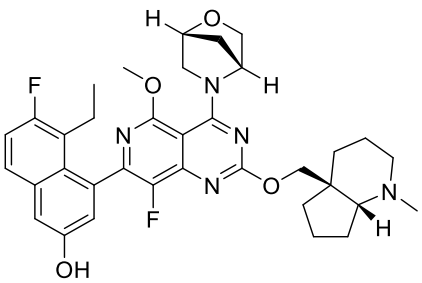
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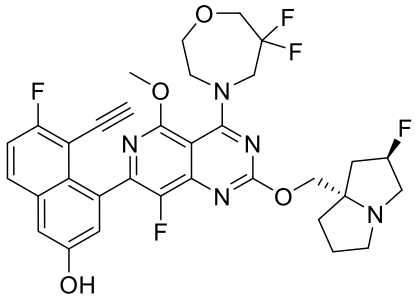
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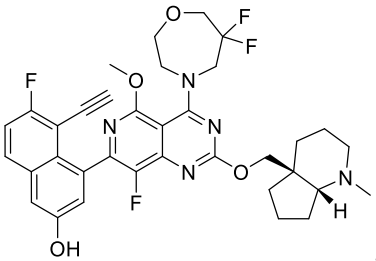
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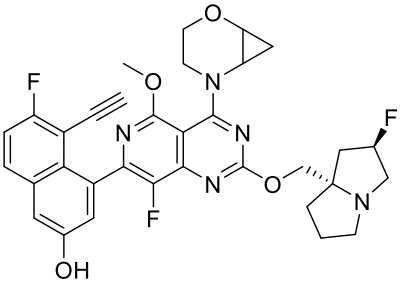
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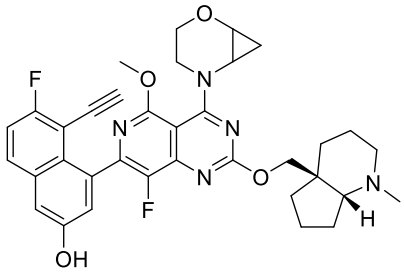
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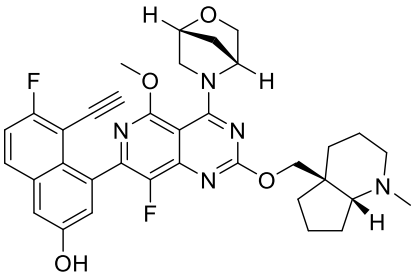
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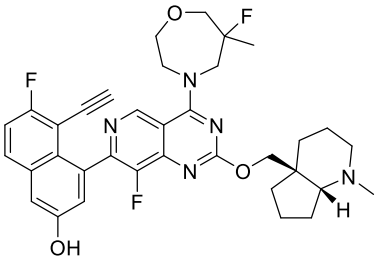
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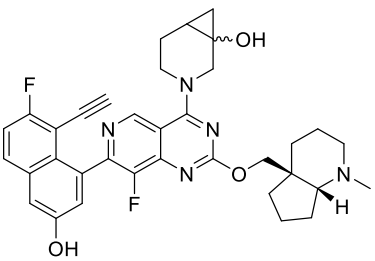
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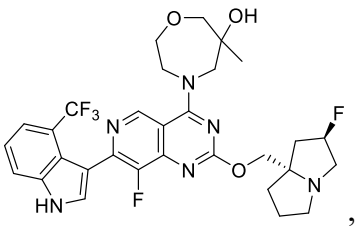
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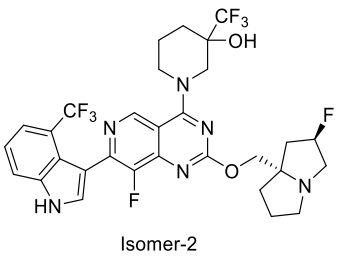
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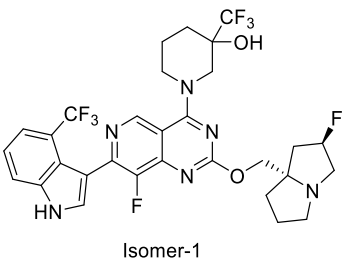
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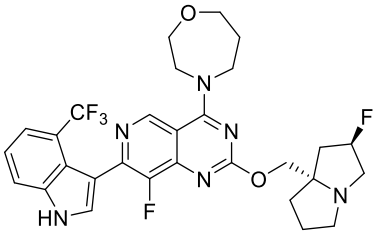
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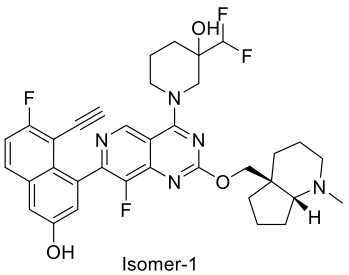
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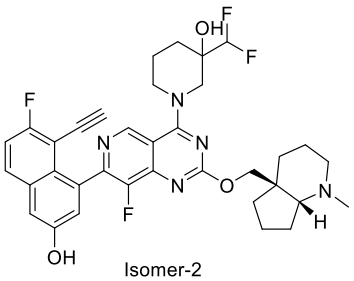
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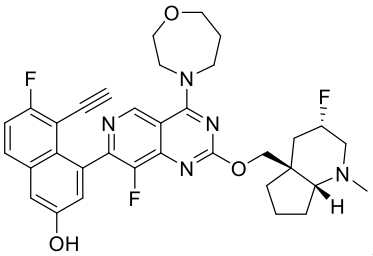
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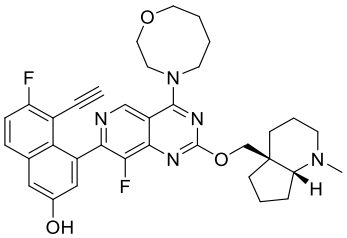
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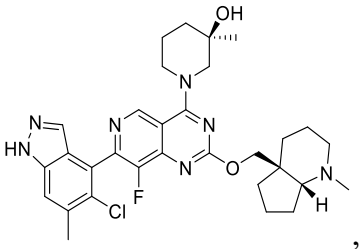
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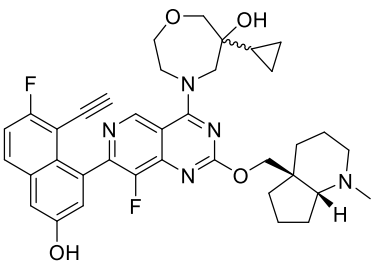
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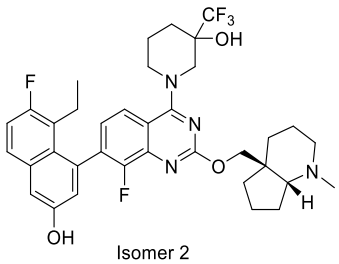
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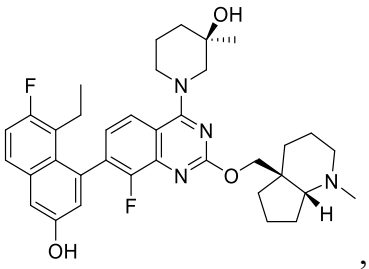
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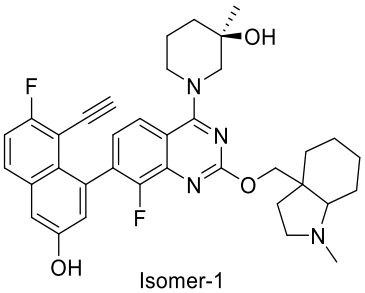
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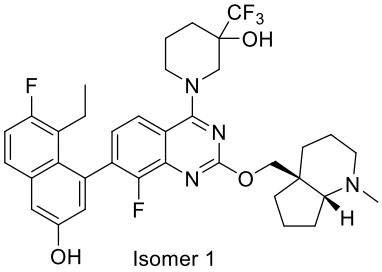
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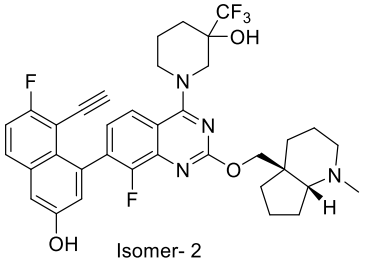
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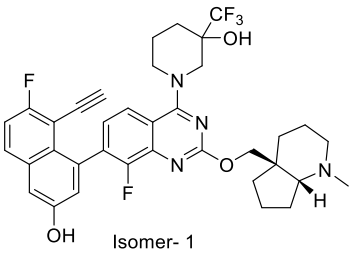
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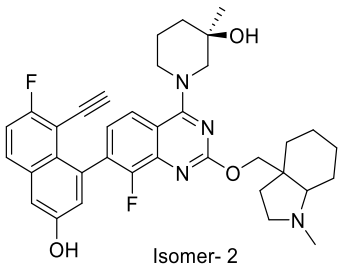
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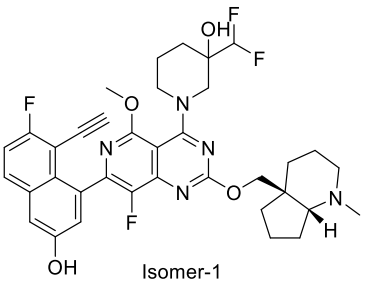
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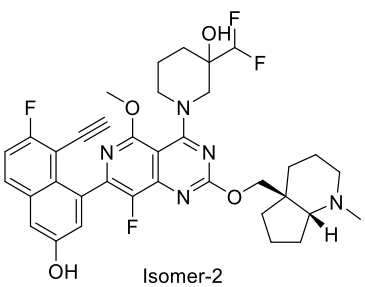
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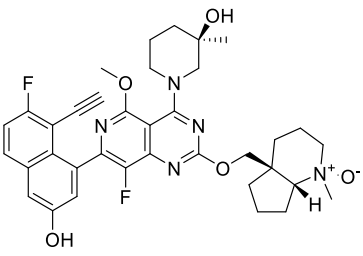
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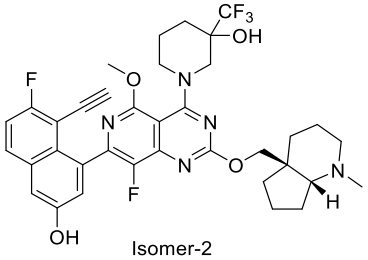
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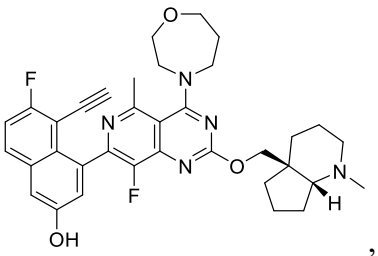
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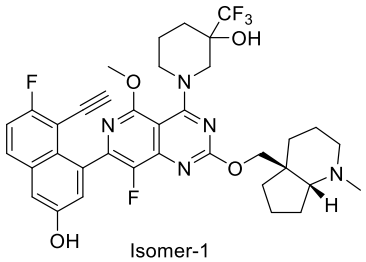
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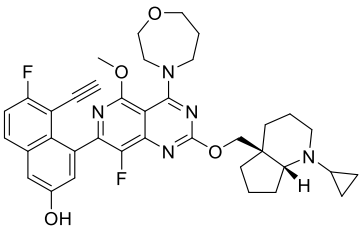
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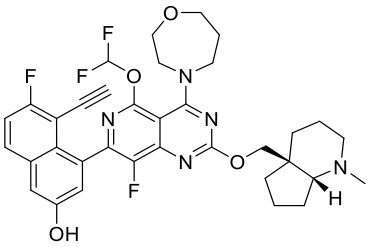
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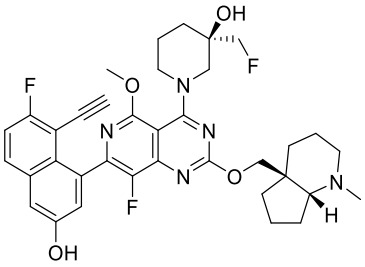
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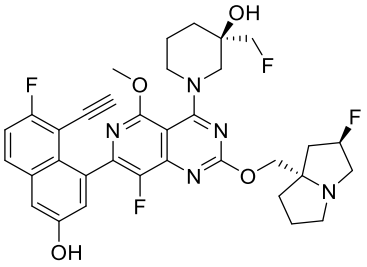
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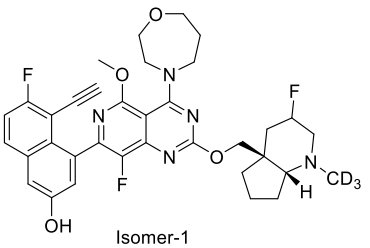
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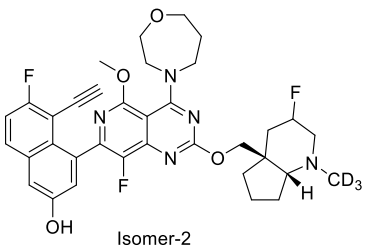
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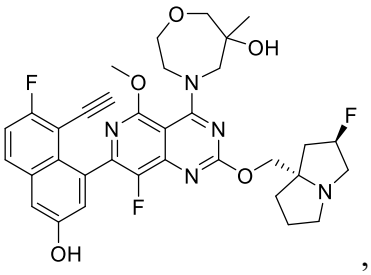
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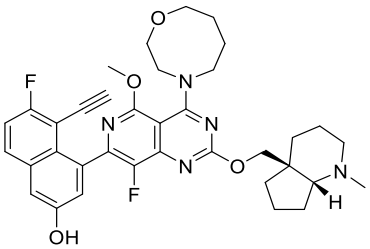
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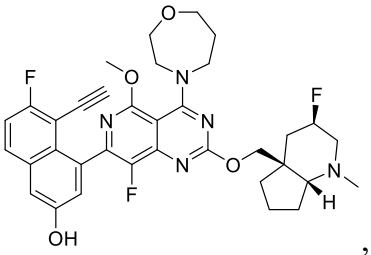
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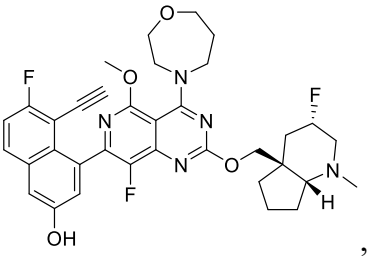
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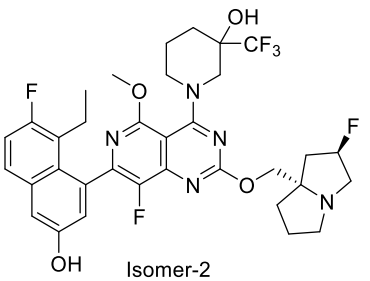
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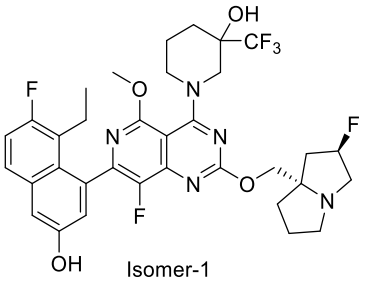
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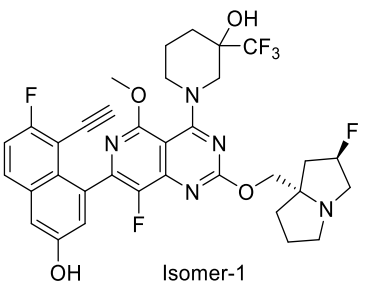
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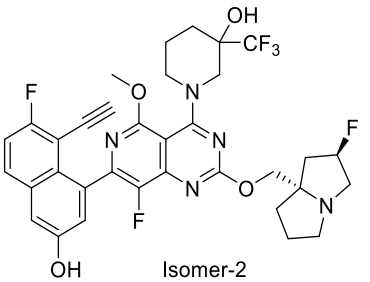
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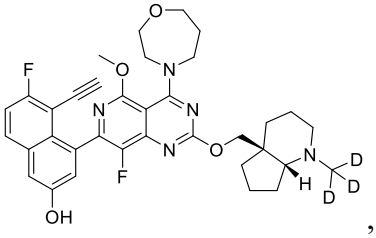
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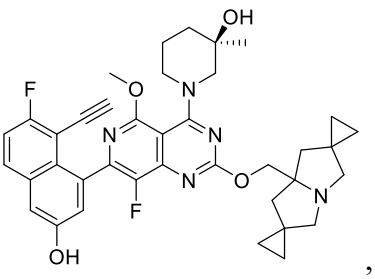
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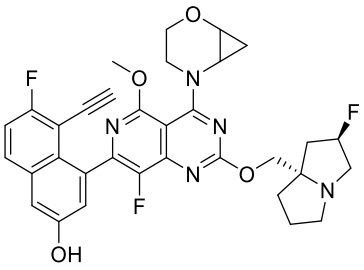
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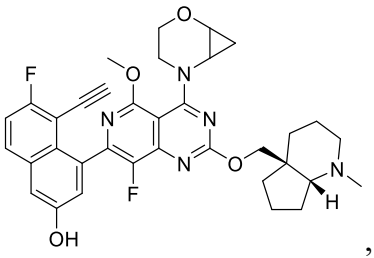
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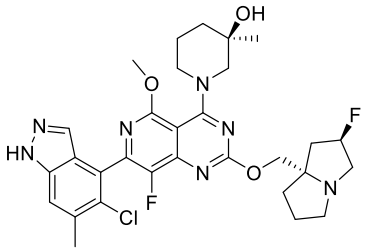
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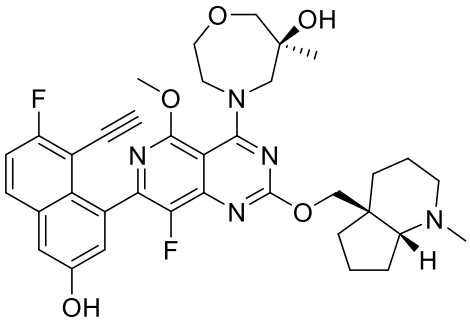
; or a pharmaceutically acceptable salt thereof.

44. A compound selected from the group consisting of:

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(morpholin-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin- 7a-yl]methoxy}-8-fluoro-4-(1,4-oxazepan-4- yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6- fluoronaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4- (1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

5-ethynyl-6-fluoro-4-{8-fluoro-2-[(1-{[(3R)-3-fluoropyrrolidin-1-yl]methyl}cyclopropyl)methoxy]-4-(morpholin-4-yl)pyrido[4,3-d]pyrimidin-7-yl}naphthalen-2-ol;

4-[2-({1-[(dimethylamino)methyl]cyclopropyl}methoxy)-8-fluoro-4-(morpholin-4-yl)pyrido[4,3-d]pyrimidin-7-yl]-5-ethynyl-6-fluoronaphthalen-2-ol;

5-ethynyl-6-fluoro-4-[8-fluoro-4-(morpholin-4-yl)-2-({1-[(piperidin-1-yl)methyl]cyclopropyl}methoxy)pyrido[4,3-d]pyrimidin-7-yl]naphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{8-oxa-3-azabicyclo[3.2.1]octan-3-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-(morpholin-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-{8-oxa-3-azabicyclo[3.2.1]octan-3-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[(2R,6S)-2,6-dimethylmorpholin-4-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(2S)-2-methylmorpholin-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-4-[(2R,6S)-2,6-dimethylmorpholin-4-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[(3R,4S)-3,4-difluoropyrrolidin-1-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

6-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-2¿6-thia-6-azaspiro[3.3]heptane-2,2-dione;

4-[2-({1-[(dimethylamino)methyl]cyclopropyl}methoxy)-8-fluoro-4-(morpholin-4-yl)pyrido[4,3-d]pyrimidin-7-yl]-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(3R)-3-(2-hydroxypropan-2-yl)pyrrolidin-1-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(8aR)-7-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-hexahydro-1H-[1,3]oxazolo[3,4-a]piperazin-3-one;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-N,N-dimethylpiperidine-4-sulfonamide;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(4-methanesulfonylpiperidin-1-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[(2R,6S)-2,6-dimethylmorpholin-4-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-{2,2-difluoro-7-azaspiro[3.5]nonan-7-yl}-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)piperidine-3-carbonitrile;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)piperidine-3-carbonitrile;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[2-(hydroxymethyl)morpholin-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)piperidine-3-carbonitrile isomer 1;

1-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)piperidine-3-carbonitrile isomer 2;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(3R)-3-methylmorpholin-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-[(3R)-3-methylmorpholin-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

5-ethynyl-6-fluoro-4-[8-fluoro-2-({1-[(4-fluoropiperidin-1-yl)methyl]cyclopropyl}methoxy)-4-[(3R)-3-methylmorpholin-4-yl]pyrido[4,3-d]pyrimidin-7-yl]naphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(2R)-2-(hydroxymethyl)morpholin-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(2S)-2-(hydroxymethyl)morpholin-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(3S)-3-methylmorpholin-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[(3R)-3-ethylmorpholin-4-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)piperidine-4-carbonitrile;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{3-oxa-6-azabicyclo[3.1.1]heptan-6-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(1S,4S)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(1R,4R)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{2-oxa-5-azabicyclo[4.1.0]heptan-5-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-4-methylpiperidin-4-ol;

5-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-2-methyl-1¿6,2,5-thiadiazepane-1,1-dione;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{9-oxa-2-azaspiro[5.5]undecan-2-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)piperidin-3-ol isomer 1;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)piperidin-3-ol isomer 2;

1-[7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-({1-[(4-fluoropiperidin-1-yl)methyl]cyclopropyl}methoxy)pyrido[4,3-d]pyrimidin-4-yl]piperidine-3-carbonitrile isomer 1;

1-[7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-({1-[(4-fluoropiperidin-1-yl)methyl]cyclopropyl}methoxy)pyrido[4,3-d]pyrimidin-4-yl]piperidine-3-carbonitrile isomer 2;

7-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-7-azaspiro[3.5]nonan-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{hexahydro-1H-furo[3,4-c]pyrrol-5-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-{8-azabicyclo[3.2.1]octan-8-yl}-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-{2-azabicyclo[2.2.1]heptan-2-yl}-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)piperidin-4-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(4-methoxypiperidin-1-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[3-(hydroxymethyl)piperidin-1-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

3-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-azabicyclo[3.2.1]octan-8-ol;

3-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-8-methyl-3-azabicyclo[3.2.1]octan-8-ol;

(3aR,5R,6aS)-2-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-5-(difluoromethyl)-octahydrocyclopenta[c]pyrrol-5-ol;

(1R,5S,6R)-3-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-6-methyl-3-azabicyclo[3.1.1]heptan-6-ol;

2-[1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)piperidin-3-yl]acetonitrile;

3-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-azabicyclo[3.2.1]octan-6-ol;

3-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-azabicyclo[3.3.1]nonan-9-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-4-hydroxypiperidine-3-carbonitrile isomer 1;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-4-hydroxypiperidine-3-carbonitrile isomer 2;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[3-(hydroxymethyl)piperidin-1-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol isomer 1;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[3-(hydroxymethyl)piperidin-1-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol isomer 2;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(3R)-3-(methoxymethyl)morpholin-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(3S)-3-(methoxymethyl)morpholin-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[4-(dimethylphosphoryl)piperidin-1-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(3aR,5S,6aS)-2-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-5-methyl-octahydrocyclopenta[c]pyrrol-5-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[4-(methoxymethyl)piperidin-1-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidine-3-carbonitrile isomer 1;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidine-3-carbonitrile isomer 2;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[3-(dimethylphosphoryl)piperidin-1-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol isomer 1;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[3-(dimethylphosphoryl)piperidin-1-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol isomer 2;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[2-(methoxymethyl)piperidin-1-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol isomer 1;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[2-(methoxymethyl)piperidin-1-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol isomer 2;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-{2-oxa-5-azabicyclo[4.1.0]heptan-5-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4aS,7aR)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-4-methyl-1,4-azaphosphinane 4-oxide;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-{3-oxa-6-azabicyclo[3.1.1]heptan-6-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-[(1S,4S)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-(6-methyl-1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-[6-(hydroxymethyl)-1,4-oxazepan-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-[6-(hydroxymethyl)-1,4-oxazepan-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethyl-6-fluoronaphthalen-2-ol;

3-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-azabicyclo[3.2.1]octan-6-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(morpholin-4-yl)quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin- 7a-yl]methoxy}-8-fluoro-4-(1,4-oxazepan-4-yl)quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2ol ;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-(1,4-oxazepan-4-yl)quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{8-oxa-3-azabicyclo[3.2.1]octan-3-yl}quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(2S)-2-methylmorpholin-4-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(3R)-3-methylmorpholin-4-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[(2R,6S)-2,6-dimethylmorpholin-4-yl]-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[2-(hydroxymethyl)morpholin-4-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(3S)-3-methylmorpholin-4-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(pyrrolidin-1-yl)quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-{6,6-dimethyl-3-azabicyclo[3.1.0]hexan-3-yl}-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[(3R,4S)-3,4-difluoropyrrolidin-1-yl]-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)piperidine-3-carbonitrile;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(3R)-3-(2-hydroxypropan-2-yl)pyrrolidin-1-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-1¿6-thiomorpholine-1,1-dione;

(9aR)-8-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-octahydropiperazino[2,1-c]morpholin-4-one;

(8aR)-7-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-hexahydro-1H-[1,3]oxazolo[3,4-a]piperazin-3-one;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(4-methanesulfonylpiperidin-1-yl)quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-N,N-dimethylpiperidine-4-sulfonamide;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{2-oxa-7-azaspiro[3.5]nonan-7-yl}quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-{2,2-difluoro-7-azaspiro[3.5]nonan-7-yl}-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(piperidin-1-yl)quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-{2-azabicyclo[2.2.1]heptan-2-yl}-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-{8-azabicyclo[3.2.1]octan-8-yl}-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{hexahydro-1H-furo[3,4-c]pyrrol-5-yl}quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(1S,4S)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{3-oxa-6-azabicyclo[3.1.1]heptan-6-yl}quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(1R,4R)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(2R)-2-(hydroxymethyl)morpholin-4-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(2S)-2-(hydroxymethyl)morpholin-4-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(8aS)-7-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-hexahydro-1H-[1,3]oxazolo[3,4-a]piperazin-3-one;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)morpholine-2-carbonitrile;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[(3R)-3-ethylmorpholin-4-yl]-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{2-oxa-5-azabicyclo[4.1.0]heptan-5-yl}quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-[(3R)-3-methylmorpholin-4-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)piperidine-4-carbonitrile;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)piperidin-3-ol;

7-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-7-azaspiro[3.5]nonan-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-4-methylpiperidin-4-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)piperidine-3-carbonitrile isomer 1;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)piperidine-3-carbonitrile isomer 2;

5-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-2-methyl-1¿6,2,5-thiadiazepane-1,1-dione;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[(3S)-3-ethylmorpholin-4-yl]-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)piperidin-4-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(4-methoxypiperidin-1-yl)quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

3-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-3-azabicyclo[3.2.1]octan-8-ol;

3-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-8-methyl-3-azabicyclo[3.2.1]octan-8-ol;

(8aS)-7-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-hexahydro-1H-[1,3]oxazolo[3,4-a]piperazin-3-one;

3-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-3-azabicyclo[3.3.1]nonan-9-ol;

(3aR,5R,6aS)-2-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-5-(difluoromethyl)-octahydrocyclopenta[c]pyrrol-5-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(3R)-3-(methoxymethyl)morpholin-4-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[(3S)-3-(methoxymethyl)morpholin-4-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[4-(dimethylphosphoryl)piperidin-1-yl]-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(3aR,5S,6aS)-2-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-5-methyl-octahydrocyclopenta[c]pyrrol-5-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[4-(methoxymethyl)piperidin-1-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

3-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-3-azabicyclo[3.1.1]heptan-6-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{9-oxa-3-azabicyclo[4.2.1]nonan-3-yl}quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(3R)-1-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)piperidin-3-ol;

(3R)-1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)piperidin-3-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(6-methyl-1,4-oxazepan-4-yl)quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(3S)-1-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)piperidin-3-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-1,4-oxazepan-6-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-1,4-oxazepan-6-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(6-methoxy-1,4-oxazepan-4-yl)quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-(6-methoxy-1,4-oxazepan-4-yl)quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

6-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-6-chloro-8-fluoro-4-{8-oxa-3-azabicyclo[3.2.1]octan-3-yl}quinazolin-7-yl)-4-methyl-5-(trifluoromethyl)pyridin-2-amine;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-(morpholin-4-yl)quinazolin-7-yl)-5-ethylnaphthalen-2-ol;

4-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-(morpholin-4-yl)quinazolin-7-yl)-5-ethylnaphthalen-2-ol;

(1S,4S)-5-(2-{[(4aS,7aR)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-2,5-diazabicyclo[2.2.1]heptan-3-one;

1-[3-(2-{[(*2R,7aS*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3,8-diazabicyclo[3.2.1]octan-8-yl]-2,2,2-trifluoroethan-1-one;

4-(2-{[(*2R,7aS*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-{8-cyclopropanecarbonyl-3,8-diazabicyclo[3.2.1]octan-3-yl}-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(*2R,7aS*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[8-(1-fluorocyclopropanecarbonyl)-3,8-diazabicyclo[3.2.1]octan-3-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-[3-(2-{[(*2R,7aS*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3,8-diazabicyclo[3.2.1]octane-8-carbonyl]cyclopropane-1-carbonitrile;

4-(2-{[(*2R,7aS*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[8-(2-fluorocyclopropanecarbonyl)-3,8-diazabicyclo[3.2.1]octan-3-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(*2R,7aS*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[8-(2,2-difluorocyclopropanecarbonyl)-3,8-diazabicyclo[3.2.1]octan-3-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol isomer 1;

4-(2-{[(*2R,7aS*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[8-(2,2-difluorocyclopropanecarbonyl)-3,8-diazabicyclo[3.2.1]octan-3-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol isomer 2;

4-(2-{[(*4aS,7aR*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-4-[8-(3,3-difluorocyclobutanecarbonyl)-3,8-diazabicyclo[3.2.1]octan-3-yl]-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(*4aS,7aR*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-4-{8-cyclopropanecarbonyl-3,8-diazabicyclo[3.2.1]octan-3-yl}-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(4-{8-cyclopropanecarbonyl-3,8-diazabicyclo[3.2.1]octan-3-yl}-8-fluoro-2-({1-[(4-fluoropiperidin-1-yl)methyl]cyclopropyl}methoxy)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-[3-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3,8-diazabicyclo[3.2.1]octan-8-yl]-2,2,2-trifluoroethan-1-one;

4-(2-{[(*2R,7aS*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[8-(4,4-difluorocyclohexanecarbonyl)-3,8-diazabicyclo[3.2.1]octan-3-yl]-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-[8-(oxane-4-carbonyl)-3,8-diazabicyclo[3.2.1]octan-3-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-[3-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-3,8-diazabicyclo[3.2.1]octan-8-yl]-2,2,2-trifluoroethan-1-one;

4-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-[8-(3,3-difluorocyclobutanecarbonyl)-3,8-diazabicyclo[3.2.1]octan-3-yl]-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-4-{8-cyclopropanecarbonyl-3,8-diazabicyclo[3.2.1]octan-3-yl}-8-fluoroquinazolin-7-yl)-5-ethynyl-6-7-6fluoronaphthalen-2-ol;

4-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{8-[(1s,4s)-4-hydroxycyclohexanecarbonyl]-3,8-diazabicyclo[3.2.1]octan-3-yl}quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-[3-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-3,8-diazabicyclo[3.2.1]octan-8-yl]-2,2,2-trifluoroethan-1-one;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-4-{8-cyclopropanecarbonyl-3,8-diazabicyclo[3.2.1]octan-3-yl}-8-fluoroquinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{8-methanesulfonyl-3,8-diazabicyclo[3.2.1]octan-3-yl}quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

6-(2-{[(*4aS,7aR*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-6-chloro-4-[8-(3,3-difluorocyclobutanecarbonyl)-3,8-diazabicyclo[3.2.1]octan-3-yl]-8-fluoroquinazolin-7-yl)-4-methyl-5-(trifluoromethyl)pyridin-2-amine;

1-[3-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-[6-amino-4-methyl-3-(trifluoromethyl)pyridin-2-yl]-6-chloro-8-fluoroquinazolin-4-yl)-3,8-diazabicyclo[3.2.1]octan-8-yl]-2,2,2-trifluoroethan-1-one;

4-(2-{[(6'*R*,7'a*R*)-6'-fluoro-hexahydrospiro[cyclopropane-1,2'-pyrrolizine]-7'a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

3-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-azabicyclo[3.2.1]octan-6-ol;

(3*R*)-1-(2-{[(4aS,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)piperidin-3-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-[6-(hydroxymethyl)-1,4-oxazepan-4-yl]pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

4-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{6-oxa-3-azabicyclo[3.2.1]octan-3-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(6-methyl-1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-(5-methyl-1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-4-(azepan-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(6'*R*,7'a*R*)-6'-fluoro-hexahydrospiro[cyclopropane-1,2'-pyrrolizine]-7'a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

4-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-(5-methyl-1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(1*S*,4*S*)-5-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-2,5-diazabicyclo[2.2.1]heptan-3-one;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-chloronaphthalen-2-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-{2-oxa-5-azabicyclo[4.1.0]heptan-5-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-4-{2-oxa-5-azabicyclo[4.1.0]heptan-5-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-{1-methyl-1H,4H,5H,6H-pyrrolo[3,4-c]pyrazol-5-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

5-ethynyl-6-fluoro-4-{8-fluoro-2-[(1-methyl-octahydro-1H-indol-3a-yl)methoxy]-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl}naphthalen-2-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-{2-methyl-2H,4H,5H,6H-pyrrolo[3,4-c]pyrazol-5-yl}pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

3-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-azabicyclo[3.2.1]octan-6-ol isomer 1;

3-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-azabicyclo[3.2.1]octan-6-ol isomer 2;

4-(2-{[(6'*R*,7'a*R*)-6'-fluoro-hexahydrospiro[cyclopropane-1,2'-pyrrolizine]-7'a-yl]methoxy}-8-fluoro-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-4-(6,6-difluoro-1,4-oxazepan-4-yl)-8-fluoropyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

(4*R*)-4-[(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)amino]pyrrolidin-2-one

4-[2-({1H,2H,3H,5H,9bH-benzo[a]pyrrolizin-9b-yl}methoxy)-8-fluoro-4-(1,4-oxazepan-4-yl)quinazolin-7-yl]-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-[(1R,4R)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl]quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-3-methylpiperidin-3-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-{2-methyl-2H,4H,5H,6H-pyrrolo[3,4-c]pyrazol-5-yl}quinazolin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoroquinazolin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

4-(2-{[(2R,7aS)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-8-fluoro-5-methoxy-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-5-methoxy-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(3*R*)-1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-7-(3-hydroxynaphthalen-1-yl)pyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

(3*R*)-1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-7-[5-hydroxy-2-(propan-2-yl)phenyl]pyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

(3*R*)-1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(2-cyclobutyl-5-hydroxyphenyl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

(3*R*)-1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(2-{bicyclo[1.1.1]pentan-1-yl}-5-hydroxyphenyl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

5-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-[(3R)-3-hydroxy-3-methylpiperidin-1-yl]pyrido[4,3-d]pyrimidin-7-yl)-1-(difluoromethyl)-1,2-dihydropyridin-2-one;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-[(3R)-3-hydroxy-3-methylpiperidin-1-yl]pyrido[4,3-d]pyrimidin-7-yl)-2-(difluoromethyl)-1,2-dihydroisoquinolin-1-one;

(3*R*)-1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-[1-(difluoromethoxy)isoquinolin-4-yl]-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-4-[(3R)-3-hydroxy-3-methylpiperidin-1-yl]pyrido[4,3-d]pyrimidin-7-yl)-2-(difluoromethyl)-6-(trifluoromethyl)-1,2-dihydroisoquinolin-1-one;

(3*R*)-1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-[1-(difluoromethoxy)-6-(trifluoromethyl)isoquinolin-4-yl]-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

(3*R*)-1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-7-(3-hydroxynaphthalen-1-yl)quinazolin-4-yl)-3-methylpiperidin-3-ol;

(3*R*)-1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-7-[5-hydroxy-2-(propan-2-yl)phenyl]quinazolin-4-yl)-3-methylpiperidin-3-ol;

(3*R*)-1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(2-cyclobutyl-5-hydroxyphenyl)-8-fluoroquinazolin-4-yl)-3-methylpiperidin-3-ol;

4-(4-(2-oxa-5-azabicyclo[4.1.0]heptan-5-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)-5-ethyl-6-fluoronaphthalen-2-ol;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)azepan-4-ol;

5-ethyl-6-fluoro-4-(8-fluoro-2-(((6'*R*,7a'*R*)-6'-fluorodihydro-1'H,3'H-spiro[cyclopropane-1,2'-pyrrolizin]-7a'(5'H)-yl)methoxy)-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-4-(2-methyl-1,4-oxazepan-4-yl)-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 1;

5-ethynyl-6-fluoro-4-(8-fluoro-4-(2-methyl-1,4-oxazepan-4-yl)-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 2;

4-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-1,4-oxazepan-6-ol;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-methylazepan-3-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-4-(1,4-oxazepan-4-yl)-2-(((6'*R*,7a'*R*)-2,2,6'-trifluorodihydro-1'H,3'H-spiro[cyclopropane-1,2'-pyrrolizin]-7a'(5'H)-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 1;

5-ethynyl-6-fluoro-4-(8-fluoro-4-(1,4-oxazepan-4-yl)-2-(((6'*R*,7a'*R*)-2,2,6'-trifluorodihydro-1'H,3'H-spiro[cyclopropane-1,2'-pyrrolizin]-7a'(5'H)-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 2;

5-ethyl-6-fluoro-4-(8-fluoro-4-(6-(hydroxymethyl)-1,4-oxazepan-4-yl)-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-4-(3-methyl-1,4-oxazepan-4-yl)-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 1;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-(fluoromethyl)piperidin-3-ol isomer 1;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-(fluoromethyl)piperidin-3-ol isomer 2;

4-(2-(((4a*S*,7a*R*)-1-ethyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-8-fluoro-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

5-ethyl-4-(2-(((4a*S*,7a*R*)-1-ethyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-8-fluoro-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-6-fluoronaphthalen-2-ol;

2-(1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)piperidin-3-yl)acetonitrile;

2-(1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-yl)acetonitrile;

5-ethynyl-6-fluoro-4-(8-fluoro-4-(3-(hydroxymethyl)azepan-1-yl)-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 1;

4-(4-((*S*)-3-(2,2-difluoroethyl)piperidin-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

6-(difluoromethyl)-4-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-1,4-oxazepan-6-ol;

4-(2-(((4a*S*,7a*R*)-1-cyclopropyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-8-fluoro-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

2-(4-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-1,4-oxazepan-6-yl)acetonitrile isomer 1;

2-(4-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-1,4-oxazepan-6-yl)acetonitrile isomer 2;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-(trifluoromethyl)piperidin-3-ol isomer 1;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-(trifluoromethyl)piperidin-3-ol isomer 2;

4-(7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-4-(2-methyl-1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 1;

4-(4-(2-oxa-5-azabicyclo[4.1.0]heptan-5-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

5-ethyl-6-fluoro-4-(8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-4-(5-methyl-1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

3-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-azabicyclo[3.2.1]octan-6-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-4-(5-methyl-1,4-oxazepan-4-yl)-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

5-chloro-4-(8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-4-(5-methyl-1,4-oxazepan-4-yl)-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-4-(2-methyl-1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 2;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-methylazepan-3-ol;

4-(4-(3-azabicyclo[3.2.1]octan-3-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-4-(3-(hydroxymethyl)azepan-1-yl)-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 2;

4-(7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)quinazolin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

4-(7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)quinazolin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-8-fluoro-5-methoxy-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethyl-6-fluoronaphthalen-2-ol;

(3*R*)-1-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

(3*R*)-1-(2-{[(2*R*,7a*S*)-2-fluoro-hexahydro-1H-pyrrolizin-7a-yl]methoxy}-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-5-methoxy-4-(1,4-oxazepan-4-yl)-2-(((6'*R*,7a'*R*)-2,2,6'-trifluorodihydro-1'H,3'H-spiro[cyclopropane-1,2'-pyrrolizin]-7a'(5'H)-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 1;

5-ethynyl-6-fluoro-4-(8-fluoro-5-methoxy-4-(1,4-oxazepan-4-yl)-2-(((6'*R*,7a'*R*)-2,2,6'-trifluorodihydro-1'H,3'H-spiro[cyclopropane-1,2'-pyrrolizin]-7a'(5'H)-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol isomer 2;

4-(4-((1*R*,4*R*)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(*R*)-1-(7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

4-(4-((1*R*,4*R*)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-7-yl)-5-ethyl-6-fluoronaphthalen-2-ol;

(*R*)-1-(7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

4-(4-((1*R*,4*R*)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)-5-ethyl-6-fluoronaphthalen-2-ol;

4-(4-(6,6-difluoro-1,4-oxazepan-4-yl)-8-fluoro-2-(((2*R*,7aS)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(4-(6,6-difluoro-1,4-oxazepan-4-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(4-(2-oxa-5-azabicyclo[4.1.0]heptan-5-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(4-(2-oxa-5-azabicyclo[4.1.0]heptan-5-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(4-((1*R*,4*R*)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-4-(6-fluoro-6-methyl-1,4-oxazepan-4-yl)-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

3-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-azabicyclo[4.1.0]heptan-1-ol;

4-(8-fluoro-2-(((2R,7aS)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-7-(4-(trifluoromethyl)-1H-indol-3-yl)pyrido[4,3-d]pyrimidin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

1-(8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-7-(4-(trifluoromethyl)-1H-indol-3-yl)pyrido[4,3-d]pyrimidin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

1-(8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-7-(4-(trifluoromethyl)-1H-indol-3-yl)pyrido[4,3-d]pyrimidin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

4-(8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-7-(4-(trifluoromethyl)-1H-indol-3-yl)pyrido[4,3-d]pyrimidin-4-yl)-1,4-oxazepane;

3-(difluoromethyl)-1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)piperidin-3-ol;

3-(difluoromethyl)-1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)piperidin-3-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-2-(((3*S*,4a*S*,7a*R*)-3-fluoro-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-4-(1,4-oxazocan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

(3*R*)-1-(7-(5-chloro-6-methyl-1H-indazol-4-yl)-8-fluoro-2-(((4a*S*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

6-cyclopropyl-4-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-1,4-oxazepan-6-ol;

1-(7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)quinazolin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

(*R*)-1-(7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)quinazolin-4-yl)-3-methylpiperidin-3-ol;

(3*R*)-1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-((1-methyloctahydro-3aH-indol-3a-yl)methoxy)quinazolin-4-yl)-3-methylpiperidin-3-ol;

1-(7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4aS,7aR)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)quinazolin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)quinazolin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)quinazolin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

(3*R*)-1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-((1-methyloctahydro-3aH-indol-3a-yl)methoxy)quinazolin-4-yl)-3-methylpiperidin-3-ol;

3-(difluoromethyl)-1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)piperidin-3-ol;

3-(difluoromethyl)-1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)piperidin-3-ol;

(4a*S*,7a*R*)-4a-(((7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-4-((*R*)-3-hydroxy-3-methylpiperidin-1-yl)-5-methoxypyrido[4,3-d]pyrimidin-2-yl)oxy)methyl)-1-methyloctahydro-1H-cyclopenta[b]pyridine 1-oxide;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-5-methyl-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

4-(2-(((4a*S*,7a*R*)-1-cyclopropyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-8-fluoro-5-methoxy-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(5-(difluoromethoxy)-8-fluoro-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(*R*)-1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-4-yl)-3-(fluoromethyl)piperidin-3-ol;

(*R*)-1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-3-(fluoromethyl)piperidin-3-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-2-(((4a*S*,7a*R*)-3-fluoro-1-(methyl-d3)octahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-5-methoxy-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-2-(((4a*S*,7a*R*)-3-fluoro-1-(methyl-d3)octahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-5-methoxy-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

4-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-4-(1,4-oxazocan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-2-(((3*R*,4aS,7a*R*)-3-fluoro-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-5-methoxy-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-2-(((3*S*,4a*S*,7a*R*)-3-fluoro-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-5-methoxy-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

1-(7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

1-(7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

1-(7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-3-(trifluoromethyl)piperidin-3-ol;

5-ethynyl-6-fluoro-4-(8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-(methyl-d3)octahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)-4-(1,4-oxazepan-4-yl)pyrido[4,3-d]pyrimidin-7-yl)naphthalen-2-ol;

(*R*)-1-(2-((1'H,3'H,5'H-dispiro[cyclopropane-1,2'-pyrrolizine-6',1''-cyclopropan]-7a'(7'H)-yl)methoxy)-7-(8-ethynyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoro-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol;

4-(4-(2-oxa-5-azabicyclo[4.1.0]heptan-5-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

4-(4-(2-oxa-5-azabicyclo[4.1.0]heptan-5-yl)-8-fluoro-5-methoxy-2-(((4a*S*,7a*R*)-1-methyloctahydro-4aH-cyclopenta[b]pyridin-4a-yl)methoxy)pyrido[4,3-d]pyrimidin-7-yl)-5-ethynyl-6-fluoronaphthalen-2-ol;

(3*R*)-1-(7-(5-chloro-6-methyl-1H-indazol-4-yl)-8-fluoro-2-(((2*R*,7a*S*)-2-fluorotetrahydro-1H-pyrrolizin-7a(5H)-yl)methoxy)-5-methoxypyrido[4,3-d]pyrimidin-4-yl)-3-methylpiperidin-3-ol; and

4-(2-{[(4a*S*,7a*R*)-1-methyl-octahydro-1H-cyclopenta[b]pyridin-4a-yl]methoxy}-7-(8-ethyl-7-fluoro-3-hydroxynaphthalen-1-yl)-8-fluoropyrido[4,3-d]pyrimidin-4-yl)-6-methyl-1,4-oxazepan-6-ol;

or a pharmaceutically acceptable salt thereof.

45. A pharmaceutical composition comprising a compound of any one of claims 1 to 44, or a pharmaceutically acceptable salt thereof, and one or more pharmaceutically acceptable excipients.

46. An oral dosage form comprising a compound of any one of claims 1 to 44, or a pharmaceutically acceptable salt thereof, and one or more pharmaceutically acceptable excipients.

47. A compound, composition, or dosage form of any one of claims 1 to 44, or a pharmaceutically acceptable salt thereof for use in inhibiting KRAS G12A, KRAS G12C, KRAS G12D, KRAS G12R, KRAS G12S, KRAS G12V, KRAS G13D and/or KRAS Q61H in a subject in need thereof, wherein the compound, composition, or dosage form, or a pharmaceutically acceptable salt thereof is to be administered to the subject.

48. A compound, composition, or dosage form of any one of claims 1 to 44, or a pharmaceutically acceptable salt thereof for use in treating a cancer susceptible to KRAS G12A,

KRAS G12C, KRAS G12D, KRAS G12R, KRAS G12S, KRAS G12V, KRAS G13D and/or KRAS Q61H inhibition in a subject in need thereof.

49. Use of a compound, composition, or dosage form of any one of claims 1 to 44, or a pharmaceutically acceptable salt thereof in the manufacture of a medicament for treating a cancer susceptible to KRAS G12A, KRAS G12C, KRAS G12D, KRAS G12R, KRAS G12S, KRAS G12V, KRAS G13D and/or KRAS Q61H inhibition in a subject in need thereof.

50. A compound, composition, or dosage form of any one of claims 1 to 44, or a pharmaceutically acceptable salt thereof for use in treating a cancer expressing KRAS G12A, KRAS G12C, KRAS G12D, KRAS G12R, KRAS G12S, KRAS G12V, KRAS G13D and/or KRAS Q61H mutation in a subject in need thereof.

51. Use of a compound, composition, or dosage form of any one of claims 1 to 44, or a pharmaceutically acceptable salt thereof in the manufacture of a medicament for treating a cancer expressing KRAS G12A, KRAS G12C, KRAS G12D, KRAS G12R, KRAS G12S, KRAS G12V, KRAS G13D and/or KRAS Q61H mutation in a subject in need thereof.

52. A compound, composition, or dosage form of any one of claims 1 to 44, or a pharmaceutically acceptable salt thereof for use in treating a cancer in a subject in need thereof.

53. Use of a compound, composition, or dosage form of any one of claims 1 to 44, or a pharmaceutically acceptable salt thereof in the manufacture of a medicament for treating a cancer in a subject in need thereof.

54. The compound, composition, or dosage form of any one of claims, or the pharmaceutically acceptable salt thereof for use of claim 52; or the use of claim 53, wherein the cancer is pancreatic cancer, colorectal cancer, lung cancer, gastric cancer, breast cancer, bladder cancer, cervical cancer, ovarian cancer, cancer of the uterus, or a combination thereof.

55. The compound, composition, or dosage form of any one of claims, or the pharmaceutically acceptable salt thereof for use; or the use of claim 54, wherein the cancer is non-small cell lung cancer.