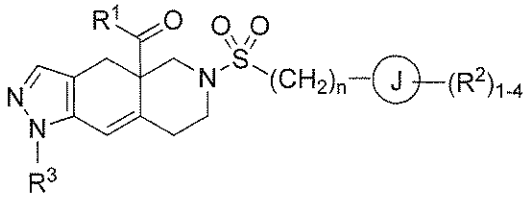
What is needed in the art are new compositions and methods for modulating GR receptors. Surprisingly, the present invention meets these and other needs.

~~BRIEF~~ SUMMARY ~~OF THE INVENTION~~

[0004] The present ~~invention~~disclosure provides many fused azadecalin compounds. In ~~some embodiments~~an aspect, the present ~~invention~~disclosure provides compounds having the structure of formula I:



(I)

wherein R^1 of formula I is a heteroaryl ring having from 5 to 6 ring members and from 1 to 4 heteroatoms which can each independently be N, O or S, optionally substituted with 1-4 groups which can each independently be R^{1a}. Each R^{1a} of formula I can independently be hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, N-oxide, C_{3-8} cycloalkyl, or C_{3-8} heterocycloalkyl. Ring J of formula I can be a cycloalkyl ring, a heterocycloalkyl ring, an aryl ring or a heteroaryl ring, wherein the heterocycloalkyl and heteroaryl rings have from 5 to 6 ring members and from 1 to 4 heteroatoms which can each independently be N, O or S. Each R^2 of formula I can independently be hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, C_{1-6} alkyl-C_{1-6} alkoxy, -CN, -OH, $-NR^{2a}R^{2b}$, $-C(O)R^{2a}$, $-C(O)OR^{2a}$, $-C(O)NR^{2a}R^{2b}$, $-SR^{2a}$, $-S(O)R^{2a}$, $-S(O)_2R^{2a}$, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl, wherein the heterocycloalkyl groups are optionally substituted with 1-4 R^{2c} groups. Alternatively, two R^2 groups linked to the same carbon can be combined to form an oxo group (=O). Alternatively, two R^2 groups can be combined to form a heterocycloalkyl ring having from 5 to 6 ring members and from 1 to 3 heteroatoms wherein each can independently be N, O or S, wherein the heterocycloalkyl ring is optionally substituted with from 1 to 3 R^{2d} groups. R^{2a} and R^{2b} of formula I can each independently be hydrogen or C_{1-6} alkyl. Each R^{2c} can independently be hydrogen, halogen, hydroxy, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, or $-NR^{2a}R^{2b}$. Each R^{2d} can independently be hydrogen or C_{1-6} alkyl, or two R^{2d} groups attached to the same ring atom can be combined to form (=O). R^3 of formula I can be phenyl or pyridyl, each optionally substituted with 1-4 R^{3a} groups. Each R^{3a} of formula I can independently be hydrogen, halogen, or

C_{1-6} haloalkyl. Subscript n of formula I can be an integer from 0 to 3. The compounds of formula I can also be the salts and isomers thereof.

[0005] In ~~some embodiments~~another aspect, the present ~~invention~~disclosure provides a pharmaceutical composition including a pharmaceutically acceptable excipient and the compound of formula I as defined above.

[0006] In some embodiments, the present ~~invention~~disclosure provides a method of modulating a glucocorticoid receptor, the method including contacting a glucocorticoid receptor with a compound of formula I, thereby modulating the glucocorticoid receptor.

[0007] In some embodiments, the present ~~invention~~disclosure provides a method of treating a disorder through modulating a glucocorticoid receptor, the method including administering to a subject in need of such treatment, a therapeutically effective amount of a compound of formula I, thereby treating the disorder.

[0008] In some embodiments, the present ~~invention~~disclosure provides a method of treating a disorder through antagonizing a glucocorticoid receptor, the method including administering to a subject in need of such treatment, an effective amount of the compound of formula I, thereby treating the disorder.

BRIEF DESCRIPTION OF THE DRAWINGS

[0009] Figures 1 and 2 show various synthetic schemes for making the compounds of the present invention.

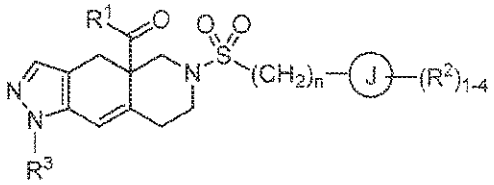
DETAILED DESCRIPTION OF THE INVENTION

I. General

[0010] The present invention provides compounds capable of modulating a glucocorticoid receptor (GR) and thereby providing beneficial therapeutic effects. The compounds include heteroaryl ketone fused azadecalins. The present invention also provides methods of treating diseases and disorders by modulating a GR receptor with the compounds of the present invention.

WHAT IS CLAIMED IS:

1. A compound having the formula:



(I)

wherein

R^1 is a heteroaryl ring having from 5 to 6 ring members and from 1 to 4 heteroatoms each independently selected from the group consisting of N, O and S, optionally substituted with 1-4 groups each independently selected from R^{1a};

each R^{1a} is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, N-oxide, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl;

ring J is selected from the group consisting of a cycloalkyl ring, a heterocycloalkyl ring, an aryl ring and a heteroaryl ring, wherein the heterocycloalkyl and heteroaryl rings have from 5 to 6 ring members and from 1 to 4 heteroatoms each independently selected from the group consisting of N, O and S;

each R^2 is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, C_{1-6} alkyl-C_{1-6} alkoxy, -CN, -OH, $-NR^{2a}R^{2b}$, $-C(O)R^{2a}$, $-C(O)OR^{2a}$, $-C(O)NR^{2a}R^{2b}$, $-SR^{2a}$, $-S(O)R^{2a}$, $-S(O)_2R^{2a}$, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl, wherein the heterocycloalkyl groups are optionally substituted with 1-4 R^{2c} groups;

alternatively, two R^2 groups linked to the same carbon are combined to form an oxo group (=O);

alternatively, two R^2 groups are combined to form a heterocycloalkyl ring having from 5 to 6 ring members and from 1 to 3 heteroatoms each independently selected from the group consisting of N, O and S, wherein the heterocycloalkyl ring is optionally substituted with from 1 to 3 R^{2d} groups;

R^{2a} and R^{2b} are each independently selected from the group consisting of hydrogen and C_{1-6} alkyl;

each R^{2c} is independently selected from the group consisting of hydrogen, halogen, hydroxy, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, and $-NR^{2a}R^{2b}$;
each R^{2d} is independently selected from the group consisting of hydrogen and C_{1-6} alkyl, or two R^{2d} groups attached to the same ring atom are combined to form (=O);
R^3 is selected from the group consisting of phenyl and pyridyl, each optionally substituted with 1-4 R^{3a} groups;
each R^{3a} is independently selected from the group consisting of hydrogen, halogen, and C_{1-6} haloalkyl;
subscript n is an integer from 0 to 3;
or salts and isomers thereof.

2. The compound of claim 1, wherein
R^1 is a heteroaryl ring having from 5 to 6 ring members and from 1 to 4 heteroatoms each independently selected from the group consisting of N, O and S, optionally substituted with 1-4 groups each independently selected from R^{1a};
each R^{1a} is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl;
ring J is selected from the group consisting of a cycloalkyl ring, a heterocycloalkyl ring, an aryl ring and a heteroaryl ring, wherein the heterocycloalkyl and heteroaryl rings have from 5 to 6 ring members and from 1 to 3 heteroatoms each independently selected from the group consisting of N, O and S;
each R^2 is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, $-NR^{2a}R^{2b}$, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl, wherein the heterocycloalkyl groups are optionally substituted with 1-4 R^{2c} groups;
R^{2a} and R^{2b} are each independently selected from the group consisting of hydrogen and C_{1-6} alkyl;
each R^{2c} is independently selected from the group consisting of hydrogen, halogen, hydroxy, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, and $-NR^{2a}R^{2b}$;
R^3 is selected from the group consisting of phenyl and pyridyl, each optionally substituted with 1-4 R^{3a} groups;

each R^{3a} is independently selected from the group consisting of hydrogen, halogen, and C_{1-6} haloalkyl; and

subscript n is an integer from 0 to 3.

3. The compound of any of claims 1 to 2, wherein

R^1 is a heteroaryl ring having from 5 to 6 ring members and from 1 to 3 heteroatoms each independently selected from the group consisting of N, O and S, optionally substituted with 1-4 groups each independently selected from R^{1a};

each R^{1a} is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, C_{1-6} haloalkyl, and C_{1-6} alkoxy;

ring J is selected from the group consisting of tetrahydrofuran, phenyl and pyridyl;

each R^2 is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, $-NR^{2a}R^{2b}$, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl;

R^{2a} and R^{2b} are each independently selected from the group consisting of hydrogen and C_{1-6} alkyl;

R^3 is selected from the group consisting of phenyl and pyridyl;

R^{3a} is F; and

subscript n is 0 or 1.

4. The compound of any of claims 1 to 3, wherein

R^1 is a heteroaryl ring having from 5 to 6 ring members and from 1 to 3 heteroatoms each independently selected from the group consisting of N, O and S, optionally substituted with 1-4 groups each independently selected from R^{1a};

each R^{1a} is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, C_{1-6} haloalkyl, and C_{1-6} alkoxy;

ring J is selected from the group consisting of phenyl and pyridyl;

each R^2 is independently selected from the group consisting of hydrogen, halogen, C_{1-6} haloalkyl, -CN and C_{5-6} heterocycloalkyl;

R^3 is selected from the group consisting of phenyl and pyridyl; and

R^{3a} is F.

5. The compound of any of claims 1 to 4, wherein R^1 is selected from the group consisting of pyrrole, pyrazole, imidazole, triazole, tetrazole, furan, oxazole, isoxazole, oxadiazole, thiophene, thiazole, isothiazole, thiadiazole, pyridine, pyrazine, pyrimidine, and pyridazine.

6. The compound of any of claims 1 to 5, wherein R^1 is selected from the group consisting of 2-pyrrole, 3-pyrrole, 3-pyrazole, 4-pyrazole, 5-pyrazole, 2-imidazole, 4-imidazole, 5-imidazole, 1,2,3-triazol-4-yl, 1,2,3,-triazol-5-yl, 1,2,4-triazol-3-yl, 1,2,4-triazol-5-yl, 1,2,3,4-tetrazol-1-yl, 1,2,3,4,tetrazol-5-yl, 2-furan, 3-furan, 2-oxazole, 4-oxazole, 5-oxazole, 3-isoxazole, 4-isooxazole, 5-isooxazole, 1,2,4-oxadiazol-3-yl, 1,2,4-oxadiazol-5-yl, 1,2,5-oxadiazol-3-yl, 1,3,4-oxadiazol-2-yl, 2-thiophene, 3-thiophene, 2-thiazole, 4-thiazole, 5-thiazole, 3-isothiazole, 4-isothiazole, 5-isothiazole, 1,2,3-thiadiazol-4-yl, 1,2,3-thiadiazol-5-yl, 1,2,4-thiadiazol-3-yl, 1,2,4-thiadiazol-5-yl, 1,2,5-thiadiazol-3-yl, 1,3,4-thiadiazol-2-yl, 2-pyridine, 3-pyridine, 4-pyridine, pyrazine, 2-pyrimidine, 4-pyrimidine, 5-pyrimidine, 6-pyrimidine, 3-pyridazine, 4-pyridazine, 5-pyridazine, and 6-pyrdazine.

7. The compound of any of claims 1 to 5, wherein R^1 is selected from the group consisting of pyrazole, imidazole, triazole, furan, oxazole, oxadiazole, thiophene, thiazole, pyridine, pyrazine and pyrimidine.

8. The compound of any of claims 1 to 7, wherein R^1 is selected from the group consisting of 1-pyrazole, 3-pyrazole, 4-pyrazole, 5-pyrazole, 2-imidazole, 4-imidazole, 5-imidazole, 1,2,3-triazol-4-yl, 1,2,3,-triazol-5-yl, 1,2,4-triazol-3-yl, 1,2,4-triazol-5-yl, 2-furan, 3-furan, 2-oxazole, 4-oxazole, 5-oxazole, 1,2,4-oxadiazol-3-yl, 1,2,4-oxadiazol-5-yl, 1,2,5-oxadiazol-3-yl, 1,3,4-oxadiazol-2-yl, 2-thiophene, 3-thiophene, 2-thiazole, 4-thiazole, 5-thiazole, 2-pyridine, 3-pyridine, 4-pyridine, pyrazine, 2-pyrimidine, 4-pyrimidine, 5-pyrimidine, and 6-pyrimidine.

9. The compound of any of claims 1, 2 and 5 to 8, wherein each R^{1a} is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, C_{1-6} haloalkyl, C_{1-6} alkoxy and C_{3-8} heterocycloalkyl.

10. The compound of any of claims 1 to 9, wherein each R^{1a} is independently selected from the group consisting of hydrogen, methyl, ethyl, trifluoromethyl, and methoxy.

11. The compound of any of claims 1, 2 and 5 to 10, wherein ring J is selected from the group consisting of heterocycloalkyl, aryl and heteroaryl.

12. The compound of any of claims 1, 2 and 5 to 11, wherein ring J is selected from the group consisting of aryl and heteroaryl.

13. The compound of any of claims 1, 2 and 5 to 12, wherein ring J is selected from the group consisting of phenyl, pyridine, imidazole, pyrazole, triazole, tetrazole, thiadiazole, isothiazole, isoxazole, cyclohexyl, tetrahydrofuran and tetrahydro-2H-pyran.

14. The compound of any of claims 1 to 13, wherein ring J is phenyl.

15. The compound of any of claims 1 to 13, wherein ring J is pyridyl.

16. The compound of any of claims 1 and 5 to 15, wherein each R^2 is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, C_{1-6} alkyl-C_{1-6} alkoxy, -CN, $-NR^{2a}R^{2b}$, $-C(O)OR^{2a}$, $-S(O)_2R^{2a}$, C_{3-8} cycloalkyl, or C_{3-8} heterocycloalkyl, wherein the heterocycloalkyl group has 5-6 ring members and 1 to 2 heteroatoms.

17. The compound of any of claims 1 and 5 to 16, wherein each R^2 is independently selected from the group consisting of hydrogen, methyl, ethyl, propyl, isopropyl, F, Cl, $-CF_3$, CH_2OMe, OMe, $OCHF_2$, -CN, $-NMe_2$, -C(O)OH, $-C(O)NMe_2$, $-S(O)_2Me$, pyrrolidine, piperidine or morpholine.

18. The compound of any of claims 1 to 17, wherein R^3 is 4-F-phenyl.

19. The compound of claim 1, wherein the compound is selected from the group consisting of:

(R)-(6-((6-chloropyridin-3-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((6-chloropyridin-3-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(1-methyl-1H-imidazol-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-3-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methyl-1,3,4-oxadiazol-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(oxazol-4-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(oxazol-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(furan-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiophen-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(oxazol-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyrimidin-2-yl)methanone,
(R)-(6-((3,5-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methoxypyridin-2-yl)methanone,
(R)-(4-ethylpyridin-2-yl)(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methoxypyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-fluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-fluorobenzyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

((4aR)-1-(4-fluorophenyl)-6-((((R/S)-tetrahydrofuran-2-yl)methyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(o-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((4-ethylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-chlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-methoxyphenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((4-chloro-3-fluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-methoxyphenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-fluoro-4-methylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(phenylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2-fluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-(trifluoromethyl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-tosyl-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((4-fluoro-3-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-4-((1-(4-fluorophenyl)-4a-picolinoyl-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzonitrile,

(R)-(1-(4-fluorophenyl)-6-((6-methoxypyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((tetrahydro-2H-pyran-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-(cyclohexylsulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((3,5-dimethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1H-imidazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-morpholinopyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-(pyrrolidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-fluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-fluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-4-(((1-(4-fluorophenyl)-4a-(thiazole-2-carbonyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)methyl)benzonitrile,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(1-methyl-1H-1,2,4-triazol-5-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyrazin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-fluoropyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-fluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methoxypyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-5-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-fluoropyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(pyrrolidin-1-yl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(pyrrolidin-1-yl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-(piperidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-(pyrrolidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-(pyrrolidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

((R)-1-(4-fluorophenyl)-6-((6-((R)-3-fluoropyrrolidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(pyrrolidin-1-yl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-(piperidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-(pyrrolidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(6-((4-chlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-methoxy-3-methylphenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-chloro-4-methoxyphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-fluoro-4-methoxyphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((2-fluoro-4-methylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-3-((1-(4-fluorophenyl)-4a-picolinoyl-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzonitrile,

(R)-(6-((4-(difluoromethoxy)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethoxy)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,5-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-tosyl-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(6-((3-(difluoromethoxy)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-dimethylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,5-dimethylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-methylpyridin-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-chloro-4-fluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-3-((1-(4-fluorophenyl)-4a-(4-methylpicolinoyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzonitrile,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-imidazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3,5-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(methylsulfonyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-3-((1-(4-fluorophenyl)-4a-picolinoyl-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzoic acid,

(R)-(1-(4-fluorophenyl)-6-((3-(methoxymethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-methyl-3,4-dihydro-2H-pyrido[3,2-b][1,4]oxazin-7-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2,3,4-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-(trifluoromethyl)pyridin-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-(trifluoromethyl)pyridin-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((1,5-dimethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((4-fluoro-3-methylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-methyl-3,4-dihydro-2H-pyrido[3,2-b][1,4]oxazin-7-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((2,3-dihydrobenzofuran-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-5-((1-(4-fluorophenyl)-4a-(4-methylpicolinoyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)-1-methylindolin-2-one,

(R)-(1-(4-fluorophenyl)-6-((3-(methylsulfonyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((1,3-dimethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((2,3-dihydrobenzo[b][1,4]dioxin-6-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3-fluoro-4-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3-fluoro-4-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-3-((4a-(4-ethylpicolinoyl)-1-(4-fluorophenyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzonitrile,

(R)-(4-ethylpyridin-2-yl)(1-(4-fluorophenyl)-6-((6-(trifluoromethyl)pyridin-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)methanone,

(R)-3-((1-(4-fluorophenyl)-4a-(4-methylpicolinoyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzoic acid,

(R)-(6-((3,5-dimethylisoxazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-phenyl-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((1,3-dimethyl-1H-pyrazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2-(trifluoromethyl)pyridin-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)pyridin-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2-(trifluoromethyl)pyridin-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((4-chloro-3-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-chloro-4-methylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(2-(pyrrolidin-1-yl)pyridin-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-3-((1-(4-fluorophenyl)-4a-(4-(trifluoromethyl)picolinoyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzonitrile,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1,5-dimethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-3-(trifluoromethyl)-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-chlorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-1-(4-(trifluoromethyl)phenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(6-((1,2-dimethyl-1H-imidazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1,2-dimethyl-1H-imidazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-imidazol-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-imidazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-propyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-(2-methoxyethyl)-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(1-methyl-1H-pyrazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-isopropyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2-methyl-2H-1,2,3-triazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-1,2,3-triazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-1,2,3-triazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((2-ethyl-2H-1,2,3-triazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-1,2,3-triazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-1,2,3-triazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((2-ethyl-2H-1,2,3-triazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-1,2,3-triazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-1,2,3-triazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2-propyl-2H-1,2,3-triazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-propyl-1H-1,2,3-triazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-propyl-1H-1,2,3-triazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2-propyl-2H-1,2,3-triazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-propyl-1H-1,2,3-triazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-propyl-1H-1,2,3-triazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2-isopropyl-2H-1,2,3-triazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-isopropyl-1H-1,2,3-triazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-isopropyl-1H-1,2,3-triazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-propyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-propyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-methoxyphenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3-fluoro-4-methylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(phenylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3-chlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(6-methylpyridin-2-yl)methanone,

(R)-(6-((4-fluoro-3-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3-fluoro-4-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylthiazol-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(6-((4-chloro-3-fluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(6-((1,3-dimethyl-1H-pyrazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-propyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-tosyl-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(1-methyl-1H-imidazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(1-methyl-1H-imidazol-2-yl)methanone ,

(R)-3-((1-(4-fluorophenyl)-4a-picolinoyl-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)-N,N-dimethylbenzamide,

(R)-(6-((1-ethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,
(R)-pyridin-2-yl(1-(pyridin-3-yl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)methanone,
(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-phenyl-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,
(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(3,4-difluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,
(R)-(6-((3,5-difluoro-4-methoxyphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,
(R)-(6-((6-(dimethylamino)pyridin-3-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,
(R)-5-((1-(4-fluorophenyl)-4a-picolinoyl-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)-1-methylpyridin-2(1H)-one,
(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone, and
(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,
or salts and isomers thereof.

20. The compound of claim 1, wherein the compound is selected from the group consisting of:
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,
(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,
(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,
(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,
(R)-(6-((3-fluoro-4-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone ,
(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,
(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,
(R)-(6-((1-ethyl-1H-pyrazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone, or
(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone.

21. A pharmaceutical composition comprising a compound of any of claims 1 to 20 and a pharmaceutically acceptable excipient.

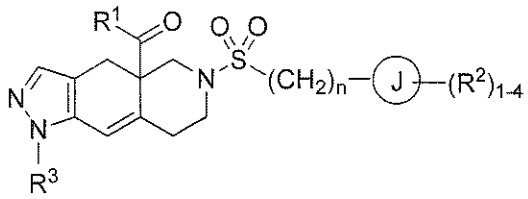
22. The composition of claim 21, further comprising an anti-inflammatory glucocorticosteroid.

23. A method of modulating a glucocorticoid receptor, comprising contacting a glucocorticoid receptor with a compound of any of claims 1 to 20, thereby modulating the glucocorticoid receptor.

24. A method of treating a disorder through antagonizing a glucocorticoid receptor, the method comprising administering to a subject in need thereof, a therapeutically effective amount of a compound of any of claims 1 to 20, thereby treating the disorder.

WHAT IS CLAIMED IS:

1. A compound having the formula:



(I)

wherein

R^1 is a heteroaryl ring having from 5 to 6 ring members and from 1 to 4 heteroatoms each independently selected from the group consisting of N, O and S, optionally substituted with 1-4 groups each independently selected from R^{1a};

each R^{1a} is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, N-oxide, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl;

ring J is selected from the group consisting of a cycloalkyl ring, a heterocycloalkyl ring, an aryl ring and a heteroaryl ring, wherein the heterocycloalkyl and heteroaryl rings have from 5 to 6 ring members and from 1 to 4 heteroatoms each independently selected from the group consisting of N, O and S;

each R^2 is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, C_{1-6} alkyl-C_{1-6} alkoxy, -CN, -OH, $-NR^{2a}R^{2b}$, $-C(O)R^{2a}$, $-C(O)OR^{2a}$, $-C(O)NR^{2a}R^{2b}$, $-SR^{2a}$, $-S(O)R^{2a}$, $-S(O)_2R^{2a}$, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl, wherein the heterocycloalkyl groups are optionally substituted with 1-4 R^{2c} groups;

alternatively, two R^2 groups linked to the same carbon are combined to form an oxo group (=O);

alternatively, two R^2 groups are combined to form a heterocycloalkyl ring having from 5 to 6 ring members and from 1 to 3 heteroatoms each independently selected from the group consisting of N, O and S, wherein the heterocycloalkyl ring is optionally substituted with from 1 to 3 R^{2d} groups;

R^{2a} and R^{2b} are each independently selected from the group consisting of hydrogen and C_{1-6} alkyl;

each R^{2c} is independently selected from the group consisting of hydrogen, halogen, hydroxy, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, and $-NR^{2a}R^{2b}$;

each R^{2d} is independently selected from the group consisting of hydrogen and C_{1-6} alkyl, or two R^{2d} groups attached to the same ring atom are combined to form (=O);

R^3 is selected from the group consisting of phenyl and pyridyl, each optionally substituted with 1-4 R^{3a} groups;

each R^{3a} is independently selected from the group consisting of hydrogen, halogen, and C_{1-6} haloalkyl;

subscript n is an integer from 0 to 3;

or salts and isomers thereof.

2. The compound of claim 1, wherein

R^1 is a heteroaryl ring having from 5 to 6 ring members and from 1 to 4 heteroatoms each independently selected from the group consisting of N, O and S, optionally substituted with 1-4 groups each independently selected from R^{1a};

each R^{1a} is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl;

ring J is selected from the group consisting of a cycloalkyl ring, a heterocycloalkyl ring, an aryl ring and a heteroaryl ring, wherein the heterocycloalkyl and heteroaryl rings have from 5 to 6 ring members and from 1 to 3 heteroatoms each independently selected from the group consisting of N, O and S;

each R^2 is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, $-NR^{2a}R^{2b}$, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl, wherein the heterocycloalkyl groups are optionally substituted with 1-4 R^{2c} groups;

R^{2a} and R^{2b} are each independently selected from the group consisting of hydrogen and C_{1-6} alkyl;

each R^{2c} is independently selected from the group consisting of hydrogen, halogen, hydroxy, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, and $-NR^{2a}R^{2b}$;

R^3 is selected from the group consisting of phenyl and pyridyl, each optionally substituted with 1-4 R^{3a} groups;

each R^{3a} is independently selected from the group consisting of hydrogen, halogen, and C_{1-6} haloalkyl; and

subscript n is an integer from 0 to 3.

3. The compound of claim 1, wherein

R^1 is a heteroaryl ring having from 5 to 6 ring members and from 1 to 3 heteroatoms each independently selected from the group consisting of N, O and S, optionally substituted with 1-4 groups each independently selected from R^{1a};

each R^{1a} is independently selected from the group consisting of hydrogen and C_{1-6} alkyl;

ring J is selected from the group consisting of tetrahydrofuran, phenyl and pyridyl;

each R^2 is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, -CN, $-NR^{2a}R^{2b}$, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl;

R^{2a} and R^{2b} are each independently selected from the group consisting of hydrogen and C_{1-6} alkyl;

R^3 is selected from the group consisting of phenyl and pyridyl;

R^{3a} is F; and

subscript n is 0 or 1.

4. The compound of claim 1, wherein

R^1 is a heteroaryl ring having from 5 to 6 ring members and from 1 to 3 heteroatoms each independently selected from the group consisting of N, O and S, optionally substituted with 1-4 groups each independently selected from R^{1a};

each R^{1a} is independently selected from the group consisting of hydrogen and C_{1-6} alkyl;

ring J is selected from the group consisting of phenyl and pyridyl;

each R^2 is independently selected from the group consisting of hydrogen, halogen, C_{1-6} haloalkyl, -CN and C_{5-6} heterocycloalkyl;

R^3 is selected from the group consisting of phenyl and pyridyl; and

R^{3a} is F.

5. The compound of claim 1, wherein R^1 is selected from the group consisting of pyrrole, pyrazole, imidazole, triazole, tetrazole, furan, oxazole, isoxazole, oxadiazole, thiophene, thiazole, isothiazole, thiadiazole, pyridine, pyrazine, pyrimidine, and pyridazine.

6. The compound of claim 1, wherein R^1 is selected from the group consisting of 2-pyrrole, 3-pyrrole, 3-pyrazole, 4-pyrazole, 5-pyrazole, 2-imidazole, 4-imidazole, 5-imidazole, 1,2,3-triazol-4-yl, 1,2,3,-triazol-5-yl, 1,2,4-triazol-3-yl, 1,2,4-triazol-5-yl, 1,2,3,4-tetrazol-1-yl, 1,2,3,4,tetrazol-5-yl, 2-furan, 3-furan, 2-oxazole, 4-oxazole, 5-oxazole, 3-isoxazole, 4-isooxazole, 5-isooxazole, 1,2,4-oxadiazol-3-yl, 1,2,4-oxadiazol-5-yl, 1,2,5-oxadiazol-3-yl, 1,3,4-oxadiazol-2-yl, 2-thiophene, 3-thiophene, 2-thiazole, 4-thiazole, 5-thiazole, 3-isothiazole, 4-isothiazole, 5-isothiazole, 1,2,3-thiadiazol-4-yl, 1,2,3-thiadiazol-5-yl, 1,2,4-thiadiazol-3-yl, 1,2,4-thiadiazol-5-yl, 1,2,5-thiadiazol-3-yl, 1,3,4-thiadiazol-2-yl, 2-pyridine, 3-pyridine, 4-pyridine, pyrazine, 2-pyrimidine, 4-pyrimidine, 5-pyrimidine, 6-pyrimidine, 3-pyridazine, 4-pyridazine, 5-pyridazine, and 6-pyridazine.

7. The compound of claim 1, wherein R^1 is selected from the group consisting of pyrazole, imidazole, triazole, furan, oxazole, oxadiazole, thiophene, thiazole, pyridine, pyrazine and pyrimidine.

8. The compound of claim 1, wherein R^1 is selected from the group consisting of 1-pyrazole, 3-pyrazole, 4-pyrazole, 5-pyrazole, 2-imidazole, 4-imidazole, 5-imidazole, 1,2,3-triazol-4-yl, 1,2,3,-triazol-5-yl, 1,2,4-triazol-3-yl, 1,2,4-triazol-5-yl, 2-furan, 3-furan, 2-oxazole, 4-oxazole, 5-oxazole, 1,2,4-oxadiazol-3-yl, 1,2,4-oxadiazol-5-yl, 1,2,5-oxadiazol-3-yl, 1,3,4-oxadiazol-2-yl, 2-thiophene, 3-thiophene, 2-thiazole, 4-thiazole, 5-thiazole, 2-pyridine, 3-pyridine, 4-pyridine, pyrazine, 2-pyrimidine, 4-pyrimidine, 5-pyrimidine, and 6-pyrimidine.

9. The compound of claim 1, wherein each R^{1a} is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, C_{1-6} haloalkyl, C_{1-6} alkoxy and C_{3-8} heterocycloalkyl.

10. The compound of claim 1, wherein each R^{1a} is independently selected from the group consisting of hydrogen, methyl, ethyl, trifluoromethyl, and methoxy.

11. The compound of claim 1, wherein ring J is selected from the group consisting of heterocycloalkyl, aryl and heteroaryl.

12. The compound of claim 1, wherein ring J is selected from the group consisting of aryl and heteroaryl.

13. The compound of claim 1, wherein ring J is selected from the group consisting of phenyl, pyridine, imidazole, pyrazole, triazole, tetrazole, thiadiazole, isothiazole, cyclohexyl, tetrahydrofuran and tetrahydro-2H-pyran.

14. The compound of claim 1, wherein ring J is phenyl.

15. The compound of claim 1, wherein ring J is pyridyl.

16. The compound of claim 1, wherein each R^2 is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, halogen, C_{1-6} haloalkyl, C_{1-6} alkoxy, C_{1-6} haloalkoxy, C_{1-6} alkyl-C_{1-6} alkoxy, -CN, $-NR^{2a}R^{2b}$, $-C(O)OR^{2a}$, $-S(O)_2R^{2a}$, C_{3-8} cycloalkyl, and C_{3-8} heterocycloalkyl, wherein the heterocycloalkyl group has 5-6 ring members and 1 to 2 heteroatoms.

17. The compound of claim 1, wherein each R^2 is independently selected from the group consisting of hydrogen, methyl, ethyl, F, Cl, $-CF_3$, OMe, $OCHF_2$, -CN, $-NMe_2$, $-S(O)_2Me$, pyrrolidine, piperidine and morpholine.

18. The compound of claim 1, wherein R^3 is 4-F-phenyl.

19. The compound of claim 1, wherein the compound is selected from the group consisting of:

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(1-methyl-1H-imidazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-fluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((6-chloropyridin-3-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-morpholinopyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-3-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methyl-1,3,4-oxadiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(oxazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(oxazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(furan-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiophen-2-yl)methanone,

(R)-(6-((6-chloropyridin-3-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-(pyrrolidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3-fluorobenzyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

((4aR)-1-(4-fluorophenyl)-6-((((R/S)-tetrahydro furan-2-yl)methyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-fluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(pyrrolidin-1-yl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-fluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(pyrrolidin-1-l)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-4-(((1-(4-fluorophenyl)-4a-(thiazole-2-carbonyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)methyl)benzonitrile,

(R)-(1-(4-fluorophenyl)-6-((5-(piperidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-(pyrrolidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-(pyrrolidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

((R)-1-(4-fluorophenyl)-6-((6-((R)-3-fluoropyrrolidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(pyrrolidin-1-yl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(1-methyl-1H-1,2,4-triazol-5-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyrazin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-fluoropyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-fluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-(piperidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(o-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((4-ethylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-(pyrrolidin-1-yl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-chlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-methoxyphenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((4-chloro-3-fluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-methoxyphenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-fluoro-4-methylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(phenylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2-fluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-(trifluoromethyl)pyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-tosyl-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((4-fluoro-3-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methoxypyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-4-((1-(4-fluorophenyl)-4a-picolinoyl-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzonitrile,

(R)-(1-(4-fluorophenyl)-6-((6-methoxypyridin-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((tetrahydro-2H-pyran-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-5-yl)methanone,

(R)-(6-(cyclohexylsulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-methoxyphenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((4-chlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-methoxy-3-methylphenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-chloro-4-methoxyphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-fluoro-4-methylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(phenylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3-chlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3-fluoro-4-methoxyphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((2-fluoro-4-methylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-tosyl-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(1-methyl-1H-imidazol-2-yl)methanone,

(R)-3-((1-(4-fluorophenyl)-4a-picolinoyl-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzonitrile,

(R)-(6-((4-(difluoromethoxy)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethoxy)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(1-methyl-1H-imidazol-2-yl)methanone,

(R)-(6-((3,5-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-tosyl-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3-(difluoromethoxy)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-dimethylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,5-dimethylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(oxazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-methylpyridin-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(6-methylpyridin-2-yl)methanone,

(R)-(6-((3-chloro-4-fluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-3-((1-(4-fluorophenyl)-4a-(4-methylpicolinoyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4*H*)-yl)sulfonyl)benzonitrile,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-imidazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-(m-tolylsulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyrimidin-2-yl)methanone,

(R)-(6-((4-fluoro-3-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3,5-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(methylsulfonyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-3-((1-(4-fluorophenyl)-4a-picolinoyl-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzoic acid,

(R)-3-((1-(4-fluorophenyl)-4a-picolinoyl-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)-N,N-dimethylbenzamide,

(R)-(1-(4-fluorophenyl)-6-((3-(methoxymethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-methyl-3,4-dihydro-2H-pyrido[3,2-b][1,4]oxazin-7-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2,3,4-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-(trifluoromethyl)pyridin-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((6-(trifluoromethyl)pyridin-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((1,5-dimethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((4-fluoro-3-methylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-methyl-3,4-dihydro-2H-pyrido[3,2-b][1,4]oxazin-7-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((2,3-dihydrobenzofuran-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-5-((1-(4-fluorophenyl)-4a-(4-methylpicolinoyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)-1-methylindolin-2-one,

(R)-(1-(4-fluorophenyl)-6-((3-(methylsulfonyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3,5-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methoxypyridin-2-yl)methanone,

(R)-(6-((1,3-dimethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((2,3-dihydrobenzo[b][1,4]dioxin-6-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3-fluoro-4-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((3-fluoro-4-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-fluoro-4-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-3-((4a-(4-ethylpicolinoyl)-1-(4-fluorophenyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzonitrile,

(R)-(4-ethylpyridin-2-yl)(1-(4-fluorophenyl)-6-((6-(trifluoromethyl)pyridin-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)methanone,

(R)-3-((1-(4-fluorophenyl)-4a-(4-methylpicolinoyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzoic acid,

(R)-(6-((3,5-dimethylisoxazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-phenyl-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((1,3-dimethyl-1H-pyrazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2-(trifluoromethyl)pyridin-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(4-ethylpyridin-2-yl)(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methoxypyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylthiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)pyridin-2-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-methylpyridin-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((2-(trifluoromethyl)pyridin-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-pyridin-2-yl(1-(pyridin-3-yl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(6-((4-chloro-3-(trifluoromethyl)phenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3-chloro-4-methylphenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-phenyl-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(2-(pyrrolidin-1-yl)pyridin-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((5-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-3-((1-(4-fluorophenyl)-4a-(4-(trifluoromethyl)picolinoyl)-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)benzonitrile,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(3,4-difluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-4-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1,5-dimethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-3-(trifluoromethyl)-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(6-((4-chloro-3-fluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(6-((6-(dimethylamino)pyridin-3-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((3,4,5-trifluorophenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-chlorophenyl)-6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(6-((1-ethyl-1H-pyrazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((3,5-dimethyl-1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1H-imidazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((1,3-dimethyl-1H-pyrazol-5-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-5-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone,

(R)-(6-((1H-pyrazol-4-yl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-5-((1-(4-fluorophenyl)-4a-picolinoyl-4a,5,7,8-tetrahydro-1H-pyrazolo[3,4-g]isoquinolin-6(4H)-yl)sulfonyl)-1-methylpyridin-2(1H)-one,

(R)-(6-((1-methyl-1H-pyrazol-3-yl)sulfonyl)-1-(4-(trifluoromethyl)phenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

or salts and isomers thereof.

20. The compound of claim 1, wherein the compound is selected from the group consisting of:

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone, and

(R)-(1-(4-fluorophenyl)-6-((3-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone.

21. The compound of claim 1, wherein the compound is selected from the group consisting of:

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((4-(trifluoromethyl)phenyl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone,

(R)-(6-((3,4-difluorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(pyridin-2-yl)methanone,

(R)-(1-(4-fluorophenyl)-6-((1-methyl-1H-pyrazol-4-yl)sulfonyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(4-(trifluoromethyl)pyridin-2-yl)methanone,

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(thiazol-2-yl)methanone, and

(R)-(6-((3,4-dichlorophenyl)sulfonyl)-1-(4-fluorophenyl)-4,4a,5,6,7,8-hexahydro-1H-pyrazolo[3,4-g]isoquinolin-4a-yl)(5-methylthiazol-2-yl)methanone.

22. A pharmaceutical composition comprising a compound of claim 1 and a pharmaceutically acceptable excipient.

23. The composition of claim 22, further comprising an anti-inflammatory glucocorticosteroid.