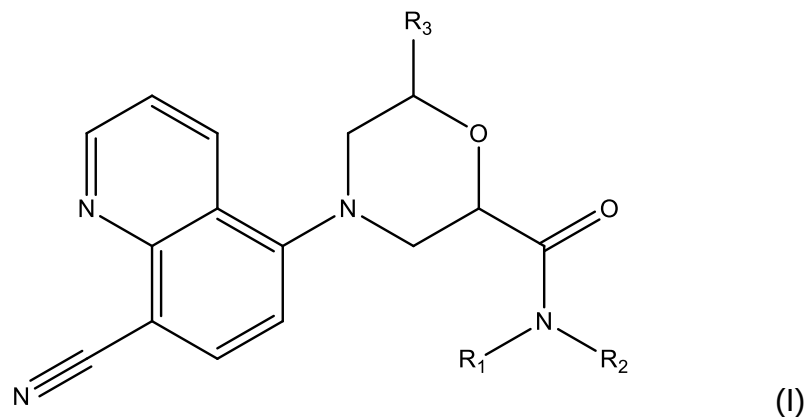


Claims

We claim:

1. A compound of formula (I) or pharmaceutically acceptable salt thereof:



wherein at least one of R_1 and R_2 is $-\text{H}$, methyl, or ethyl, and the other is $-\text{H}$; or the other is

$\text{C}_1\text{-C}_6$ alkyl that is optionally substituted with:

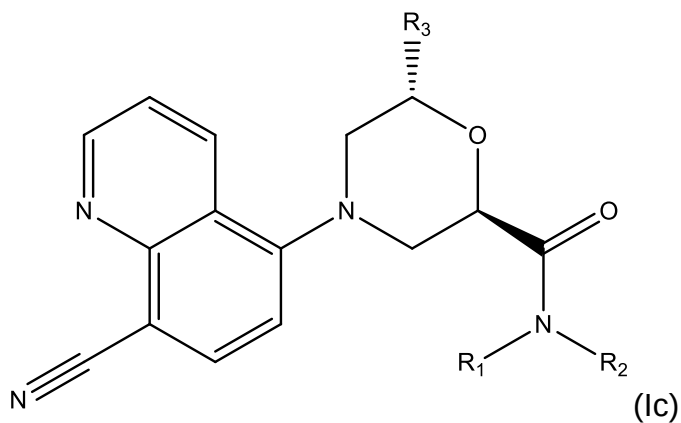
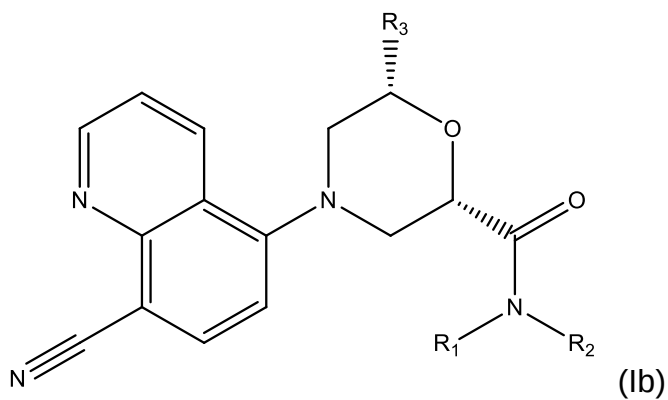
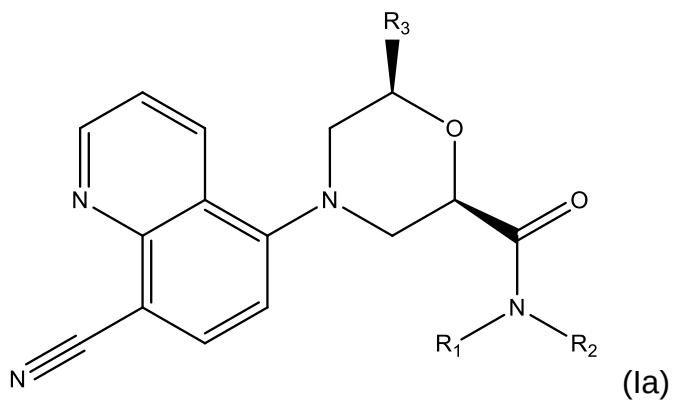
$-\text{OH}$, methoxy, ethoxy, $-\text{OCH}(\text{CH}_3)_2$, $-\text{O}(\text{CH}_2)_2\text{CH}_3$, phenyl, furanyl, $-\text{O}(\text{CH}_2)_2\text{OH}$, phenoxy, methylthio, $-\text{F}$, $-\text{N}(\text{CH}_3)_2$, cyano, pyridinyloxy, fluorophenoxy, isochromanyl, phenol, benzylamino, $-\text{NHCH}_3$, oxo-, amino, carboxyl, 7-member spiroaminy, a three to six member cycloalkyl, saturated or unsaturated and optionally including one or more heteroatoms selected from O and N, and optionally substituted at one or more C or N atoms by methyl, cyano, fluoro, methylamino, or trifluoromethyl; or the other is

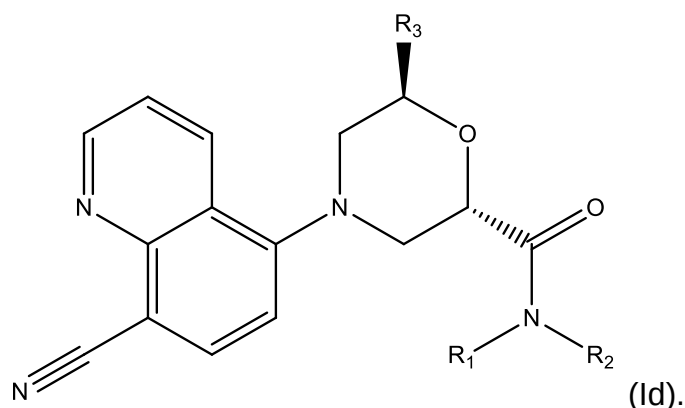
$\text{C}_3\text{-C}_7$ cycloalkane, saturated or unsaturated, optionally bridged, optionally including one or more heteroatoms selected from O, S, and N, and optionally substituted at one or more C or N atoms by methyl, ethyl, pyridinyl, azetidiny, acetamidyl, carboxamidyl, cyano, fluoro, methylamino, or trifluoromethyl; or

R_1 and R_2 , together with the nitrogen atom to which they are attached, form an 8 to 11 member spirodiamine, an 8 member bicyclodiamine, a 7 member spiroxamine, a piperidinyl optionally substituted with ethyl, or a four to six member cycloalkyl, optionally substituted with at least one of carboxamidyl, aminomethyl, methyl, (ethylamino)methyl,

(dimethylamino)methyl, dimethylamino, (methylamino)methyl, and amino; and wherein R_3 is $-H$ or methyl.

2. The compound or pharmaceutically acceptable salt of claim 1, wherein said compound or salt has a stereochemical configuration selected from one of those shown in the group consisting of Formula (Ia), Formula (Ib), Formula (Ic), and Formula (Id):





3. The compound or pharmaceutically acceptable salt of claim 1, wherein said compound or salt is selected from the group consisting of:

- 5-((2*R*,6*R*)-2-formyl-6-methylmorpholino)quinoline-8-carbonitrile;
- (2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-ethyl-6-methylmorpholine-2-carboxamide;
- (2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-isopropyl-6-methylmorpholine-2-carboxamide;
- 5-((2*R*,6*R*)-2-formyl-6-methylmorpholino)quinoline-8-carbonitrile;
- (2*R*,6*R*)-methyl 4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxylate;
- (6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;
- 5-((2*R*,6*R*)-2-((*S*)-3-ethylpiperazine-1-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- (2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(3,4-difluorobenzyl)-6-methylmorpholine-2-carboxamide;
- 5-((2*R*,6*R*)-2-((*S*)-3-aminopyrrolidine-1-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*R*)-2-(azetidine-1-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*R*)-2-([1,4'-bipiperidine]-1'-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- (2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-cyclopropyl-6-methylmorpholine-2-carboxamide;

5-((2*R*,6*R*)-2-(3-aminoazetidine-1-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-hydroxyethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-methoxyethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*R*)-2-hydroxypropyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*S*)-1-hydroxypropan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*R*)-1-hydroxypropan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*S*)-1-hydroxybutan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*S*)-1-hydroxy-3-methylbutan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*S*)-2-hydroxy-1-phenylethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*R*)-2-hydroxy-1-phenylethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-hydroxybutyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-ethoxyethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*R*)-1-hydroxybutan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(1,3-dihydroxypropan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2,3-dihydroxypropyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(((*R*)-tetrahydrofuran-2-yl)methyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((tetrahydrofuran-2-yl)methyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(2-propoxyethyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*R*)-1-hydroxypentan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-isopropoxyethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(1-methoxybutan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-(2-fluorophenyl)-2-hydroxyethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*R*)-1-hydroxy-3-methylbutan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2,2-dimethoxyethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-(2-hydroxyethoxy)ethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((1*S*,2*S*)-2-hydroxycyclohexyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-hydroxycyclohexyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(1-hydroxyhexan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*S*)-1-hydroxy-3,3-dimethylbutan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*S*)-1-hydroxyhexan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((2*S*,3*S*)-1-hydroxy-3-methylpentan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*S*)-1-hydroxy-4-methylpentan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*R*)-1-hydroxy-4-methylpentan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((4-methylmorpholin-2-yl)methyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*S*)-1-hydroxy-4-(methylthio)butan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(2-phenoxyethyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*S*)-1-hydroxy-3-phenylpropan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(2-phenoxypropyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-hydroxy-3-phenylpropyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(2-(pyridin-3-yloxy)propyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-(4-fluorophenoxy)ethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-(3-fluorophenyl)-2-hydroxyethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((*S*)-1-cyclohexyl-3-hydroxypropan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(isochroman-1-ylmethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-hydroxy-3-phenoxypropyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((1*S*,2*R*)-1-hydroxy-1-(4-hydroxyphenyl)propan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((1*S*)-1,3-dihydroxy-1-phenylpropan-2-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-hydroxy-3-(piperazin-1-yl)propyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-*N*-(azetidin-3-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*S*)-pyrrolidin-3-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*R*)-pyrrolidin-3-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*S*)-piperidin-3-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*R*)-piperidin-3-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*S*)-pyrrolidin-2-ylmethyl)morpholine-2-carboxamide;

(2*R*,6*R*)-*N*-(2-(benzylamino)ethyl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(pyridin-2-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(6-methylpyridin-2-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(5-methylisoxazol-3-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(2,2,2-trifluoroethyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2,2-difluoroethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(3,3,3-trifluoropropyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-(dimethylamino)-2-methylpropyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*S*)-morpholin-2-ylmethyl)morpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(((*S*)-4-methylmorpholin-2-yl)methyl)morpholine-2-carboxamide acetic acetate;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-fluoroethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(3-fluoropropyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*S*)-1,1,1-trifluoropropan-2-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*R*)-1,1,1-trifluoropropan-2-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(1,3-dimethyl-1*H*-pyrazol-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(1-methyl-1*H*-pyrazol-5-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-*N*-(cyanomethyl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-*N*-(1-cyanocyclopropyl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(1,2,4-thiadiazol-5-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(3-methyl-1,2,4-thiadiazol-5-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(piperidin-4-yl)morpholine-2-carboxamide;

5-((2*R*,6*R*)-2-methyl-6-(2,6-diazaspiro[3.4]octane-2-carbonyl)morpholino)quinoline-8-carbonitrile hydrochloride;

5-((2*R*,6*R*)-2-methyl-6-(3-((methylamino)methyl)azetidine-1-carbonyl)morpholino)quinoline-8-carbonitrile hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((4-fluoropiperidin-4-yl)methyl)-6-methylmorpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-*N*-(azetidin-3-ylmethyl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

5-((2*R*,6*R*)-2-methyl-6-(2,6-diazaspiro[3.5]nonane-2-carbonyl)morpholino)quinoline-8-carbonitrile hydrochloride;

5-((2*R*,6*R*)-2-methyl-6-(1,6-diazaspiro[3.4]octane-1-carbonyl)morpholino)quinoline-8-carbonitrile hydrochloride;

5-((2*R*,6*R*)-2-methyl-6-(1,7-diazaspiro[4.4]nonane-7-carbonyl)morpholino)quinoline-8-carbonitrile hydrochloride;

(2*R*,6*R*)-*N*-(3-carbamoyl-1-methyl-1*H*-pyrazol-4-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(oxetan-3-ylmethyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(tetrahydro-2*H*-pyran-4-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(tetrahydrofuran-3-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((3-methyloxetan-3-yl)methyl)morpholine-2-carboxamide;

5-((2*R*,6*R*)-2-methyl-6-(2-oxa-6-azaspiro[3.3]heptane-6-carbonyl)morpholino)quinoline-8-carbonitrile;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(oxetan-3-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-*N*-((3-(aminomethyl)oxetan-3-yl)methyl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(oxetan-2-ylmethyl)morpholine-2-carboxamide;

5-((2*R*,6*R*)-2-methyl-6-(piperazine-1-carbonyl)morpholino)quinoline-8-carbonitrile hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(2-(methylamino)ethyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*S*)-3,3,3-trifluoro-2-hydroxypropyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*R*)-3,3,3-trifluoro-2-hydroxypropyl)morpholine-2-carboxamide;

Methyl 2-((2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamido)acetate;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(2-(dimethylamino)ethyl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*S*)-4,4,4-trifluoro-3-hydroxybutyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*R*)-4,4,4-trifluoro-3-hydroxybutyl)morpholine-2-carboxamide;

(2*R*,6*R*)-*N*-(3-amino-4,4,4-trifluorobutyl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

2-((2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamido)acetic acid;

1-((2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carbonyl)azetidine-3-carboxamide;

5-((2*R*,6*R*)-2-methyl-6-(2,7-diazaspiro[4.4]nonane-2-carbonyl)morpholino)quinoline-8-carbonitrile hydrochloride;

5-((2*R*,6*R*)-2-methyl-6-(3,9-diazaspiro[5.5]undecane-3-carbonyl)morpholino)quinoline-8-carbonitrile hydrochloride;

(2*R*,6*R*)-*N*-(3-carbamoylpyridin-4-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*R*)-morpholin-2-ylmethyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(pyridin-4-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(pyridin-3-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(1-(piperidin-4-yl)-1*H*-pyrazol-4-yl)morpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-*N*-(1-(azetidin-3-yl)-1*H*-pyrazol-4-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-*N*-((1*H*-pyrazol-5-yl)methyl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-*N*-((1*H*-pyrazol-4-yl)methyl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((3-(trifluoromethyl)pyridin-2-yl)methyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(1-(pyridin-2-yl)ethyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(pyridin-2-ylmethyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((6-methylpyridin-2-yl)methyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((1-methylpiperidin-2-yl)methyl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((3-methylpyridin-2-yl)methyl)morpholine-2-carboxamide;

(2*R*,6*R*)-*N*-(4-cyano-1*H*-pyrazol-3-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((3*S*,4*R*)-4-fluoro-1-methylpyrrolidin-3-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((3*S*,4*R*)-4-fluoropiperidin-3-yl)-*N*,6-dimethylmorpholine-2-carboxamide hydrochloride;

5-((2*R*,6*R*)-2-(3-(aminomethyl)azetidine-1-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-(3-((ethylamino)methyl)azetidine-1-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-(3-((dimethylamino)methyl)azetidine-1-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(1-methylazepan-4-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(((2*R*,6*R*)-6-methylmorpholin-2-yl)methyl)morpholine-2-carboxamide;

5-((2*R*,6*R*)-2-methyl-6-(octahydropyrrolo[3,4-*c*]pyrrole-2-carbonyl)morpholino)quinoline-8-carbonitrile;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*R*)-pyrrolidin-2-ylmethyl)morpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*S*)-piperidin-2-ylmethyl)morpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-*N*-((1*R*,3*R*,5*S*)-8-azabicyclo[3.2.1]octan-3-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-*N*-(azepan-4-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-*N*-((1*R*,5*S*,6*S*)-3-azabicyclo[3.1.0]hexan-6-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-6-dimethylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(((2*S*,6*R*)-4,6-dimethylmorpholin-2-yl)methyl)-6-methylmorpholine-2-carboxamide;

5-((2*R*,6*R*)-2-(4-(dimethylamino)piperidine-1-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(1-methylpiperidin-4-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((3*S*,4*R*)-4-fluoropyrrolidin-3-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((3*S*,4*R*)-4-fluoropiperidin-3-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(2-azaspiro[3.3]heptan-6-yl)morpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-*N*-(1-(2-amino-2-oxoethyl)piperidin-4-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

5-((2*R*,6*R*)-2-(4-aminopiperidine-1-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-(4-amino-4-methylpiperidine-1-carbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*,6-dimethyl-*N*-((*R*)-piperidin-3-yl)morpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-*N*-(2-carbamoylpyridin-4-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-6-dimethyl-*N*-((*S*)-piperidin-3-yl)morpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(1-ethylpiperidin-4-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(1-ethylpiperidin-3-yl)-6-methylmorpholine-2-carboxamide;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-((*R*)-1-methylpiperidin-3-yl)morpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(4-methylpiperidin-4-yl)morpholine-2-carboxamide;

(2*R*,6*R*)-*N*-(2-amino-2-methylpropyl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide;

rel-(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((3*R*,4*S*)-4-fluoropyrrolidin-3-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

rel-(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((3*S*,4*R*)-4-fluoropyrrolidin-3-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-*N*-(azepan-3-yl)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholine-2-carboxamide hydrochloride;

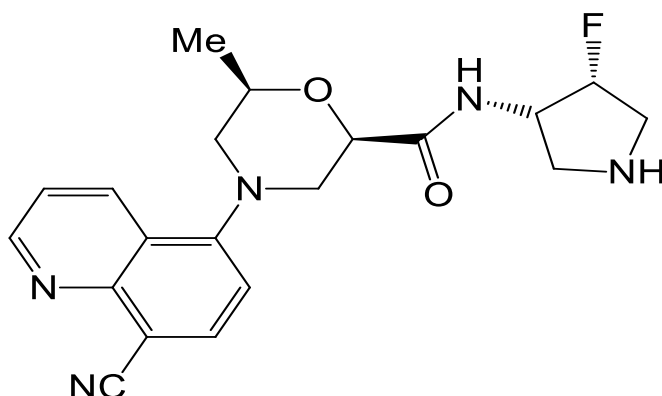
(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(1-methylazepan-3-yl)morpholine-2-carboxamide hydrochloride;

(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(1,4-dimethylpiperidin-4-yl)-6-methylmorpholine-2-carboxamide; and

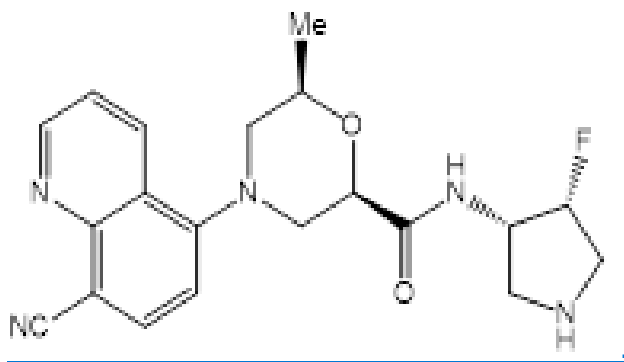
(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-(4-fluoro-1-methylpiperidin-3-yl)-6-methylmorpholine-2-carboxamide hydrochloride.

4. The compound or pharmaceutically acceptable salt of claim 3, wherein said compound or salt is selected from the group consisting of rel-(2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((3*R*,4*S*)-4-fluoropyrrolidin-3-yl)-6-methylmorpholine-2-carboxamide hydrochloride and (2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methyl-*N*-(1-methylpiperidin-4-yl)morpholine-2-carboxamide.

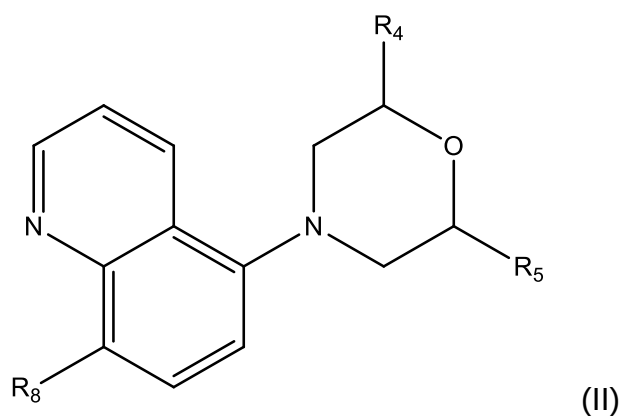
5. A compound as shown below or a pharmaceutically acceptable salt thereof, with relative stereochemistry indicated:



6. [A compound as shown below or a pharmaceutically acceptable salt thereof:](#)



6.7. A compound of Formula (II) or pharmaceutically acceptable salt thereof:



wherein

R₄ is methyl;

R₅ is C₁-C₅ alkyl that is saturated, partially saturated, or unsaturated, and that is optionally substituted with:

-H, -Cl, -F, -OH, -NH₂, oxo-, -N(CH₂CH₃)₂, phenyl, cyclohexyl, phenyltriazolyl, cyclohexyltriazolyl, pyridinyl, pyrrolidinyl,

morpholinyl optionally substituted with methyl or hydroxymethyl,

-O-, substituted with:

C₁-C₆ alkyl, methylphenyl, methylcyclohexyl, pyridinyl, diazinyl, or phenyl optionally substituted with -F or methyl,

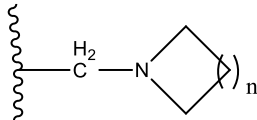
-NH-, substituted with:

C₂-C₇ alkyl that is linear, branched, or cyclic, saturated or unsaturated, and optionally substituted with oxo-, phenyl, methyl, or -OH,

pyridinyl optionally substituted with methyl, methoxy, phenyl, or amino,

diazinyl optionally substituted with ethyl,

benzoimidazolyl, methylphenyl, phenylpyrazolyl, naphthyridyl,
phenyl optionally substituted with -F, methyl, ethyl, or ethoxy,
imidazolidinyl optionally substituted with methyl

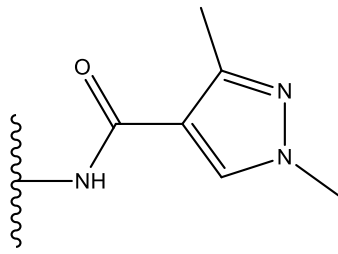
or R₅ is , wherein n is 1-3, and wherein the cyclic amine is optionally substituted with

C₁-C₃ alkyl optionally substituted with

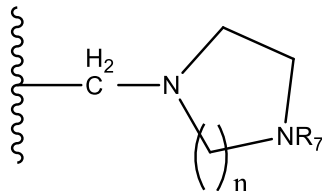
-OH, -F, phenyl, -NH₂, cyclohexyl, -N(CH₃)₂, -C(O)NH₂,
methylsulfonamidyl, benzenesulfonamidyl, methylbenzenesulfonamidyl, or
pyrrolidinyl optionally substituted with methyl or hydroxyl, or
-NHC(O)R₆, wherein R₆ is

C₁-C₅ alkyl, phenyl, pyridinyl, fluorophenyl, methylsulfonyl,
fluorobenzenesulfonyl, dimethyl pyrazole sulfonyl, or
pyrazolyl optionally substituted with methyl;

piperidinyl optionally substituted with -C(O)CH₃, -C(O)CH₂CH₃, methyl, oxo-,

C(O)Ph, -NH₂, -NH-C(O)CH₃, or ,

piperazinyl optionally substituted with -C(O)OC(CH₃)₃, methyl, -C(O)CH₃, -
C(O)Ph, C(O)CH(CH₃)₂, -C(O)CH₃, or methylsulfonyl; or

R₅ is , where n is 1 or 2, and wherein the cyclic diamine is optionally substituted on at least one carbon atom with

methyl, oxo-, -N(CH₃)₂, amino, -CH₂CH₃, or

piperidinyl optionally substituted with methyl, -C(O)CH₃, -C(O)CH(CH₃)₂, -
C(O)Ph, or -C(O)OC(CH₃)₃, and

wherein R₇ is –H, phenyl, –C(O)CH₃, C₁–C₃ alkyl, –C(O)NH₂, or –C(O)Ph; and R₈ is methoxy or cyano.

7.8. The compound or pharmaceutically acceptable salt of claim 67, wherein said compound or salt is selected from the group consisting of:

- 5-((2*R*,6*S*)-2,6-dimethylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*S*)-2,6-dimethylmorpholino)quinoline-8-carbonitrile dihydrochloride;
- 5-((2*R*,6*S*)-2,6-dimethylmorpholino)quinoline-8-carbonitrile methanesulfonate;
- 5-((2*R*,6*S*)-2,6-dimethylmorpholino)quinoline-8-carbonitrile bis(2,2,2-trifluoroacetate);
- 5-((2*R*,6*S*)-2,6-dimethylmorpholino)quinoline-8-carbonitrile bis(sulfonate);
- 5-((2*R*,6*S*)-2,6-dimethylmorpholino)quinoline-8-carbonitrile sulfonate;
- 5-((2*S*,6*R*)-2,6-dimethylmorpholino)quinoline-8-carbonitrile 2,3-dihydroxy-succinate;
- 5-((2*S*,6*R*)-2,6-dimethylmorpholino)quinoline-8-carbonitrile dimethanesulfonate;
- N*-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)acetamide;
- N*-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)pivalamide;
- 5-((2*S*,6*S*)-2,6-dimethylmorpholino)quinoline-8-carbonitrile;
- 5-((2*S*,6*R*)-2-(aminomethyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-(2-(hydroxymethyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- (2*R*,6*S*)-4-(8-methoxyquinolin-5-yl)-2,6-dimethylmorpholine;
- 5-((2*R*,6*S*)-2,6-dimethylmorpholino)quinoline-8-carboxamide;
- 5-((2*R*,6*R*)-2-(hydroxymethyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*R*)-2-(methoxymethyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*R*)-2-((benzyloxy)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*R*)-2-(fluoromethyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*R*)-2-(ethoxymethyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*R*)-2-(isopropoxymethyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*R*)-2-(isobutoxymethyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*R*)-2-(chloromethyl)-6-methylmorpholino)quinoline-8-carbonitrile;
- 5-((2*R*,6*R*)-2-((hexyloxy)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((cyclohexylmethoxy)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-propylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*S*)-2-(fluoromethyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*S*)-2-(chloromethyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((2,6-dimethylphenoxy)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2,6-diethylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-pentylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-formyl-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((benzyloxy)methyl)-6-ethylmorpholino)quinoline-8-carbonitrile;

N-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-2-phenylpropanamide;

5-((2*R*,6*R*)-2-(1-hydroxyallyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((*S*)-1-hydroxyallyl)-6-methylmorpholino)quinoline-8-carbonitrile;

N-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-1-phenylcyclobutanecarboxamide;

5-((2*R*,6*R*)-2-((*R*)-1-hydroxyallyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(pyrrolidin-1-ylmethyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((diethylamino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((benzylamino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-(phenoxymethyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-((*m*-tolylloxy)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-((*p*-tolylloxy)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-(1-hydroxypropyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((*R*)-1-hydroxypentyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((*S*)-1-hydroxypentyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((*R*)-cyclohexyl(hydroxy)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((*S*)-cyclohexyl(hydroxy)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

N-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-2-phenylacetamide;

5-((2*R*,6*R*)-2-((*R*)-1-hydroxy-2-phenylethyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((*S*)-1-hydroxy-2-phenylethyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((*R*)-1-hydroxy-3-phenylpropyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((*S*)-1-hydroxy-3-phenylpropyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((phenylamino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((*m*-tolylamino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((*o*-tolylamino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((*p*-tolylamino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((3,4-difluorophenoxy)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((3-fluorophenoxy)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((1,2-dimethyl-1*H*-benzo[d]imidazol-5-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((1-ethyl-2-methyl-1*H*-benzo[d]imidazol-5-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((cyclohexylamino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((2-fluorophenoxy)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-propionylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-(cyclohexanecarbonyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-(3-phenylpropanoyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-((*o*-tolylloxy)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((pyrimidin-2-ylamino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((pyridin-2-ylamino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((6-methylpyridin-2-yl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((5-methylpyridin-2-yl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*R*)-2-(hydroxymethyl)pyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((2,2-dimethylpyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((2-isopropylpyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*S*)-2-methylpyrrolidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*S*)-3-phenylpyrrolidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*R*)-3-methylpyrrolidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*S*)-3-hydroxypyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((4-methylpyridin-2-yl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((3-phenylpyrrolidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((6-methoxypyridin-3-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*R*)-2-methylpyrrolidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((2,5-dimethylpyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((4-methoxypyridin-2-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((6-methoxypyridin-2-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((1-phenyl-1*H*-pyrazol-5-yl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*S*)-2-(hydroxymethyl)pyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*R*)-3-hydroxypyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*S*)-3-methylpyrrolidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((3,3-dimethylpyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-((*R*)-3-phenyl-1-(pyrrolidin-1-yl)propyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-((*S*)-3-phenyl-1-(pyrrolidin-1-yl)propyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((3-methoxypyridin-2-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*R*)-3-hydroxypiperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((2*R*,6*S*)-2,6-dimethylpiperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*S*)-3-hydroxypiperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-hydroxypiperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((2-(hydroxymethyl)piperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((2-methylpiperidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((2-ethylpiperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((2,3-dimethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((pyridin-3-ylamino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((pyridin-4-ylamino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*S*)-2-(trifluoromethyl)pyrrolidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-methylpiperidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4,4-difluoropiperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-phenylpiperidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-fluoropiperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((cyclopentylamino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((3-methylcyclohexyl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((3-methylpyridin-2-yl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-ethyl-6-(hydroxymethyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((4-phenylpyridin-2-yl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(piperazin-1-ylmethyl)morpholino)quinoline-8-carbonitrile;
5-((2*R*,6*S*)-2-methyl-6-((4-phenylpiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;
5-((2*S*,6*R*)-2-(((6-aminopyridin-2-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;
5-((2*S*,6*R*)-2-((2,5-dimethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;
5-((2*S*,6*R*)-2-((4-acetylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;
5-((2*S*,6*R*)-2-(((2*S*,4*R*)-4-hydroxy-2-(hydroxymethyl)pyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;
5-((2*R*,6*S*)-2-methyl-6-(((*R*)-2-methylpiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;
5-((2*R*,6*S*)-2-methyl-6-(((*R*)-3-methylpiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;
5-((2*R*,6*S*)-2-methyl-6-(((*S*)-3-methylpiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;
5-((2*S*,6*R*)-2-(((2*R*,5*R*)-2,5-dimethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;
5-((2*S*,6*R*)-2-(((2*R*,5*S*)-2,5-dimethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;
5-((2*R*,6*S*)-2-methyl-6-(((4-methylcyclohexyl)amino)methyl)morpholino)quinoline-8-carbonitrile;
5-((2*S*,6*R*)-2-((cyclobutylamino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;
5-((2*S*,6*R*)-2-((cycloheptylamino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;
5-((2*S*,6*R*)-2-(((4-hydroxycyclohexyl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;
5-((2*S*,6*R*)-2-(((2-hydroxycyclopentyl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((2-methylcyclohexyl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((5-phenylpyridin-2-yl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((3-phenylpyridin-2-yl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((((1*S*,3*R*)-3-hydroxycyclopentyl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((3-ethoxypyridin-2-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((2-phenylpyridin-4-yl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((6-phenylpyridin-2-yl)amino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((2-methyl-5-oxopiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-methylpiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-propylpiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-(dimethylamino)piperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-([1,4'-bipiperidin]-1'-ylmethyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-([1,4'-bipiperidin]-1'-ylmethyl)-6-methylmorpholino)quinoline-8-carbonitrile dihydrochloride;

5-((2*S*,6*R*)-2-(((*R*)-3-aminopiperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

(2*R*,6*S*)-4-(8-chloro-1,7-naphthyridin-5-yl)-2,6-dimethylmorpholine;

5-((2*S*,6*R*)-2-((4-aminopiperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((5-fluoropyrimidin-2-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(piperidin-1-ylmethyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(morpholinomethyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((2*S*,6*R*)-2,6-dimethylmorpholino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((2*R*,6*R*)-2-(hydroxymethyl)-6-methylmorpholino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-(pyridin-2-yl)piperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-(pyridin-4-yl)piperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)acetamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)acetamide hydrochloride;

5-((2*S*,6*R*)-2-(((1,8-naphthyridin-2-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidine-4-carboxamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)benzamide;

5-((2*S*,6*R*)-2-((4-isopropylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

4-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperazine-1-carboxamide;

5-((2*S*,6*R*)-2-((4-cyclohexylpiperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-(pyrrolidin-1-yl)piperidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-((4-cyclohexyl-1*H*-1,2,3-triazol-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-((4-phenyl-1*H*-1,2,3-triazol-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((2-oxo-[1,4'-bipiperidin]-1'-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((5-ethylpyrimidin-2-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((6-amino-3,5-dimethylpyridin-2-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((pyrazin-2-ylamino)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((1,3-dimethyl-1*H*-pyrazol-5-yl)amino)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,7*R*)-2-(hydroxymethyl)-7-methyl-1,4-oxazepan-4-yl)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-ethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((3-(pyrrolidin-1-yl)azetidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((3-(piperidin-1-yl)azetidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*R*)-2,4-dimethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-(hydroxymethyl)piperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*R*)-[1,3'-bipyrrolidin]-1'-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*R*)-3-(piperidin-1-yl)pyrrolidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-methyl-1,4-diazepan-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-benzoylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)nicotinamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)isonicotinamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)picolinamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)hexanamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)isobutyramide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)isobutyramide hydrochloride;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)pivalamide;

5-((2*S*,7*R*)-2-([1,4'-bipiperidin]-1'-ylmethyl)-7-methyl-1,4-oxazepan-4-yl)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-morpholinopiperidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

tert-butyl 4-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)piperazine-1-carboxylate;

tert-butyl 4-(4-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperazin-1-yl)piperidine-1-carboxylate;

5-((2*R*,6*S*)-2-methyl-6-((4-(piperazin-1-yl)piperidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-(piperidin-4-yl)piperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((3-(4-methylpiperazin-1-yl)azetidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-([4,4'-bipiperidin]-1-ylmethyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((1'-acetyl-[4,4'-bipiperidin]-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((1'-methyl-[4,4'-bipiperidin]-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((3-(piperazin-1-yl)azetidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((3-(4-acetylpiperazin-1-yl)azetidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((1'-isobutyryl-[4,4'-bipiperidin]-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((3-(4-benzoylpiperazin-1-yl)azetidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((1'-benzoyl-[4,4'-bipiperidin]-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-(1-methylpiperidin-4-yl)piperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-(1-acetylpiperidin-4-yl)piperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-(1-isobutyrylpiperidin-4-yl)piperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-(1-benzoylpiperidin-4-yl)piperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((3-(4-isobutyrylpiperazin-1-yl)azetidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-(4-acetylpiperazin-1-yl)piperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-(4-isobutyrylpiperazin-1-yl)piperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-(4-benzoylpiperazin-1-yl)piperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-(4-methylpiperazin-1-yl)piperidin-1-yl)methyl)-morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((4-(4-methylpiperazin-1-yl)piperidin-1-yl)methyl)-morpholino)quinoline-8-carbonitrile trihydrochloride;

5-((2*S*,6*R*)-2-([1,4'-bipiperidin]-1'-ylmethyl)-6-methylmorpholino)-2-methylquinoline-8-carbonitrile;

N-((*R*)-1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-pyrrolidin-3-yl)acetamide;

N-((*R*)-1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-pyrrolidin-3-yl)isobutyramide;

5-((2*R*,6*S*)-2-methyl-6-(((*S*)-3-(piperidin-1-yl)pyrrolidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*S*)-3-(piperidin-1-yl)pyrrolidin-1-yl)methyl)morpholino)-quinoline-8-carbonitrile dihydrochloride;

5-((2*R*,6*S*)-2-methyl-6-((3-morpholinoazetidin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((3-((*S*)-2-methylpyrrolidin-1-yl)azetidin-1-yl)methyl)-morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*S*)-2-methylpiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*S*)-2,4-dimethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*R*)-3,4-dimethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*S*)-3,4-dimethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*S*)-3-ethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-(((*S*)-3-ethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile dihydrochloride;

5-((2*S*,6*R*)-2-(((*S*)-3-ethyl-4-methylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((4-(azepan-1-yl)piperidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((*S*)-[1,3'-bipyrrolidin]-1'-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((3-(4-aminopiperidin-1-yl)azetidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

N-(1-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-azetidin-3-yl)piperidin-4-yl)acetamide;

5-((2*R*,6*S*)-2-methyl-6-((3-(4-(methylsulfonyl)piperazin-1-yl)azetidin-1-yl)methyl)-morpholino)quinoline-8-carbonitrile;

5-((2*S*,6*R*)-2-((3-((*S*)-3-hydroxypyrrolidin-1-yl)azetidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)-1,3-dimethyl-1*H*-pyrazole-4-carboxamide;

N-(1-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-azetidin-3-yl)piperidin-4-yl)-1,3-dimethyl-1*H*-pyrazole-4-carboxamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)methanesulfonamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)benzenesulfonamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)-4-fluorobenzenesulfonamide;

5-((2*S*,6*R*)-2-((3-aminoazetidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)azetidin-3-yl)acetamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)azetidin-3-yl)-4-fluorobenzamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)azetidin-3-yl)methanesulfonamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)azetidin-3-yl)-4-fluorobenzenesulfonamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)azetidin-3-yl)-1,3-dimethyl-1*H*-pyrazole-4-sulfonamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)piperidin-4-yl)-1,3-dimethyl-1*H*-pyrazole-4-sulfonamide;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)azetidin-3-yl)isobutyramide;

5-((2*R*,6*S*)-2-methyl-6-(((*S*)-2-methyl-5-oxopiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*R*)-2-methyl-5-oxopiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*R*)-2-methyl-3-oxopiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-(((*S*)-2-methyl-3-oxopiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((2,4,5-trimethylpiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*S*)-2-methyl-6-((2,3,4-trimethylpiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile;

N-((*R*)-1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)pyrrolidin-3-yl)benzamide;

5-((2*S*,6*R*)-2-(((*R*)-3-(dimethylamino)pyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

N-((*S*)-1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-pyrrolidin-3-yl)acetamide;

N-((*S*)-1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-pyrrolidin-3-yl)isobutyramide;

N-((*S*)-1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-pyrrolidin-3-yl)benzamide;

5-((2*S*,6*R*)-2-(((*S*)-3-(dimethylamino)pyrrolidin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-((pyrazin-2-yloxy)methyl)morpholino)quinoline-8-carbonitrile;

N-(1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)azetidin-3-yl)benzamide;

5-((2*S*,6*R*)-2-((3,3-dimethylpiperazin-1-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile dihydrochloride;

5-((2*R*,6*S*)-2-methyl-6-((3,3,4-trimethylpiperazin-1-yl)methyl)morpholino)quinoline-8-carbonitrile dihydrochloride;

5-((2*R*,6*R*)-2-((*R*)-1-hydroxyethyl)-6-methylmorpholino)quinoline-8-carbonitrile;

1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-*N*-ethylpiperidine-4-carboxamide;

1-(((2*S*,6*R*)-4-(8-cyanoquinolin-5-yl)-6-methylmorpholin-2-yl)methyl)-*N*-ethylpiperidine-4-carboxamide hydrochloride;

5-((2*R*,6*R*)-2-((*S*)-1-hydroxyethyl)-6-methylmorpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-((pyridin-2-yloxy)methyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-((pyrimidin-2-yloxy)methyl)morpholino)quinoline-8-carbonitrile;

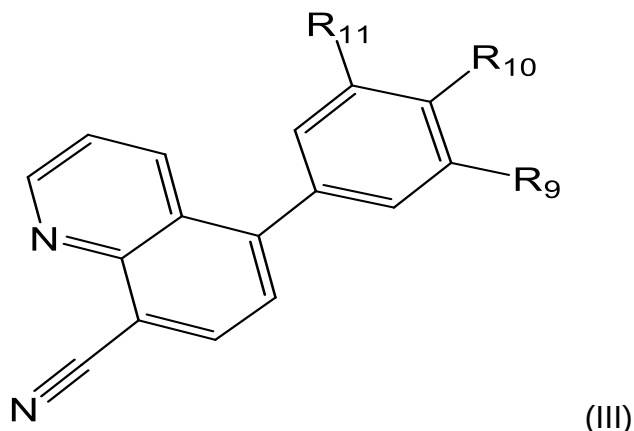
5-((2*R*,6*R*)-2-methyl-6-((*R*)-1-(pyrimidin-2-yloxy)ethyl)morpholino)quinoline-8-carbonitrile;

5-((2*R*,6*R*)-2-methyl-6-((*S*)-1-(pyrimidin-2-yloxy)ethyl)morpholino)quinoline-8-carbonitrile; and

5-((2*R*,6*R*)-2-((*S*)-hydroxy(pyridin-2-yl)methyl)-6-methylmorpholino)quinoline-8-carbonitrile.

[8.9.](#) The compound or pharmaceutically acceptable salt of claim [67](#), wherein said compound or salt is selected from the group consisting of 5-((2*S*,6*R*)-2-([1,4'-bipiperidin]-1'-ylmethyl)-6-methylmorpholino)quinoline-8-carbonitrile and 5-((2*R*,7*R*)-2-(hydroxymethyl)-7-methyl-1,4-oxazepan-4-yl)quinoline-8-carbonitrile.

[9.10.](#) A compound of Formula (III) or pharmaceutically acceptable salt thereof:



wherein

R₁₁ is H or methyl;

R₁₀ is H or, when both R₁₁ and R₉ are H, is methyl-1,4'-bipiperidinyl;

R₉ is -H or is -CH₂- substituted by 1,4'-bipiperidinyl, oxo-, hydroxyl, methylpyridinyl, or piperidinyl optionally substituted with hydroxyl, -N(CH₃)₂, or piperidinyl.

[10.11.](#) The compound or pharmaceutically acceptable salt of claim [910](#), wherein said compound or salt is selected from the group consisting of:

5-(3-([1,4'-bipiperidin]-1'-ylmethyl)phenyl)quinoline-8-carbonitrile;

5-(4-([1,4'-bipiperidin]-1'-ylmethyl)phenyl)quinoline-8-carbonitrile;

5-(3-formyl-5-methylphenyl)quinoline-8-carbonitrile;

5-(3-(hydroxymethyl)-5-methylphenyl)quinoline-8-carbonitrile;

5-(3-(chloromethyl)-5-methylphenyl)quinoline-8-carbonitrile;
5-(3-((4-hydroxypiperidin-1-yl)methyl)-5-methylphenyl)quinoline-8-carbonitrile;
5-(3-((4-(dimethylamino)piperidin-1-yl)methyl)-5-methylphenyl)quinoline-8-carbonitrile;
5-(3-methyl-5-((4-methylpiperazin-1-yl)methyl)phenyl)quinoline-8-carbonitrile;
and
5-(3-([1,4'-bipiperidin]-1'-ylmethyl)-5-methylphenyl)quinoline-8-carbonitrile.

11.12. The compound or pharmaceutically acceptable salt of any of claims 1-~~10~~
11 for use in the treatment of lupus.

12.13. The compound of claim 1112, wherein said compound is to be administered as a pharmaceutically acceptable salt.

13.14. The compound or pharmaceutically acceptable salt of any of claims 1-~~10~~
11 for use in antagonizing TLR7.

14.15. The compound or pharmaceutically acceptable salt of any of claims 1-~~10~~
11 for use in antagonizing TLR8.

15.16. A pharmaceutical composition comprising at least one compound or pharmaceutically acceptable salt of any of claims 1-~~10~~-11 and at least one pharmaceutically acceptable carrier.

16.17. The pharmaceutical composition of claim 1516, wherein said compound or pharmaceutically acceptable salt thereof has an IC₅₀ less than or equal to 100 nM against human TLR7 receptors in a HEK-293 cell line.

17.18. The pharmaceutical composition of claim 1516, wherein said compound or pharmaceutically acceptable salt thereof has an IC₅₀ less than or equal to 20 nM against human TLR7 receptors expressed in a HEK-293 cell line.

18-19. The pharmaceutical composition of claim 1516, wherein said compound or pharmaceutically acceptable salt thereof has an IC₅₀ less than or equal to 5 nM against human TLR7 receptors expressed in a HEK-293 cell line.

19-20. The pharmaceutical composition of any of claims 1617-1819, wherein the IC₅₀ against human TLR7 receptors expressed in a HEK-293 cell line is measured by (1) plating cells of the HEK-293 cell line stably expressing TLR7 in a Dulbecco's modified Eagle's medium containing 10% fetal bovine serum at a density of 2.22X10⁵ cells/ml into a 384-well plate and incubating for 2 days at 37°C, 5% CO₂; (2) adding the compound or pharmaceutically acceptable salt thereof and incubating the cells for 30 minutes; (3) adding CL097 (InvivoGen) at 3ug/ml and incubating the cells for approximately 20 hours; and (4) quantifying NF-kappaB dependent reporter activation by measuring luminescence.

20-21. The compound or pharmaceutically acceptable salt of any of claims 1-1011 for use in the treatment of a systematic lupus erythematosus, cutaneous lupus, or neuropsychiatric lupus.

21-22. The compound of claim 2021, wherein said compound is administered as a pharmaceutically acceptable salt.

22-23. A pharmaceutically acceptable salt of Claim 56, wherein the pharmaceutically acceptable salt, ~~with relative stereochemistry indicated~~, is ~~Rel-~~ (2*R*,6*R*)-4-(8-cyanoquinolin-5-yl)-*N*-((3*S*,4*R*)-4-fluoropyrrolidin-3-yl)-6-methylmorpholine-2-carboxamide hydrochloride.

23-24. The pharmaceutically acceptable salt of claim 22-23 for use in the treatment of lupus.

[24-25.](#) A pharmaceutical composition comprising the pharmaceutically acceptable salt of claim [22-23](#) and at least one pharmaceutically acceptable carrier.

[25-26.](#) The pharmaceutically acceptable salt of claim [22-23](#) for use in the treatment of a systematic lupus erythematosus, cutaneous lupus, or neutropsychiatric lupus.

[26-27.](#) The pharmaceutically acceptable salt of claim [22-23](#) for use in antagonizing TLR7.

[27-28.](#) The pharmaceutically acceptable salt of claim [22-23](#) for use in antagonizing TLR8.

[28-29.](#) The compound or pharmaceutically acceptable salt of claim 5 for use in the treatment of lupus.

[29-30.](#) A pharmaceutical composition comprising the pharmaceutically acceptable salt of claim 5 and at least one pharmaceutically acceptable carrier.

[30-31.](#) The compound or pharmaceutically acceptable salt of claim 5 for use in the treatment of a systematic lupus erythematosus, cutaneous lupus, or neutropsychiatric lupus.

[31-32.](#) The compound or pharmaceutically acceptable salt of claim 5 for use in antagonizing TLR7.

[32-33.](#) The compound or pharmaceutically acceptable salt of claim 5 for use in antagonizing TLR8.