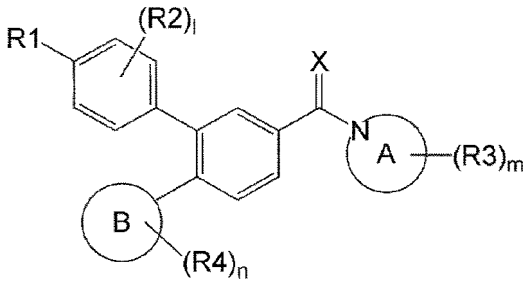
CLAIMS

[Claim 1]

A compound represented by Formula (I) or a salt thereof:



(I)

wherein

ring A represents a monocyclic, bridged cyclic, or spirocyclic 4 to 14-membered nitrogen-containing saturated heterocyclic group having 1 to 3 nitrogen atoms, 0 to 1 sulfur atoms, and 0 to 2 oxygen atoms as heteroatoms,

ring B represents monocyclic or bicyclic 5- to 14-membered unsaturated hydrocarbon or a monocyclic or bicyclic 5- to 14-membered unsaturated heterocyclic group that may be substituted with oxo, that has 0 to 4 nitrogen atoms, 0 to 2 sulfur atoms, and 0 to 3 oxygen atoms as heteroatoms, and that has at least one of nitrogen, sulfur, and oxygen,

X represents O or S,

R1 represents nitro or cyano,

R2 represents halogen,

R3 represents amino, mono- or di(C1-C6 alkyl)amino, (C3-C7 cycloalkyl)amino, or C1-C6 alkyl, and

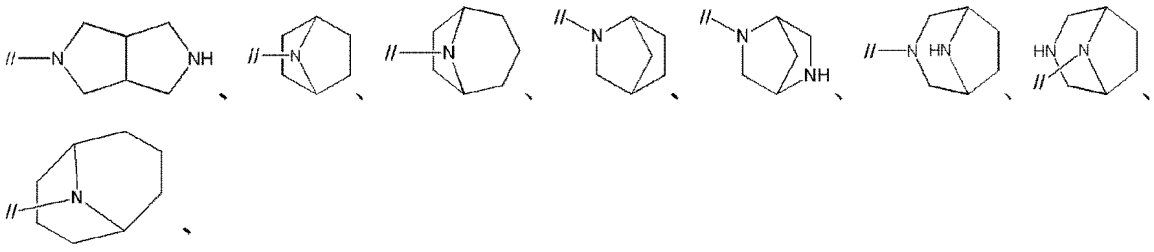
R4 represents halogen, nitro, cyano, carboxy, substituted or unsubstituted C1-C8 alkyl, substituted or unsubstituted C2-C6 alkenyl, substituted or unsubstituted C1-C6 alkoxy, substituted or unsubstituted C3-C7 cycloalkyl, mono- or di(C1-C6 alkyl)amino, or substituted or unsubstituted carbamoyl, wherein when at least one R4 represents substituted C1-C8 alkyl,

substituted C2-C6 alkenyl, substituted C1-C6 alkoxy, substituted C3-C7 cycloalkyl, or substituted carbamoyl, the substituent is halogen, carboxy, C1-C6 alkoxy, hydroxy, C1-C6 alkyl that may be

substituted with hydroxy, monocyclic 5- to 10-membered unsaturated hydrocarbon, carbamoyl that may be substituted with C1-C6 alkyl or monocyclic 5- to 10-membered unsaturated hydrocarbon, (C2-C7 acyl)oxy, amino that may be substituted with C1-C6 alkyl or C2-C7 acyl, C3-C7 cycloalkyl that may be substituted with hydroxy, or (C1-C6 alkoxy)(C1-C6 alkyl), and when two or more of the substituents are present, the substituents may be identical or different,
l is an integer of 0 to 2,
m is an integer of 0 to 2, and
n is an integer of 0 to 5,
wherein when l is 2, two R2s may be identical or different, when m is 2, two R3s may be identical or different, and when n is 2 to 5, two to five R4s may be identical or different.

[Claim 2]

The compound or a salt thereof according to claim 1, which satisfies the following conditions in Formula (I):
ring A represents pyrrolidinyl, piperidinyl, piperazinyl, azepanyl, diazepanyl,



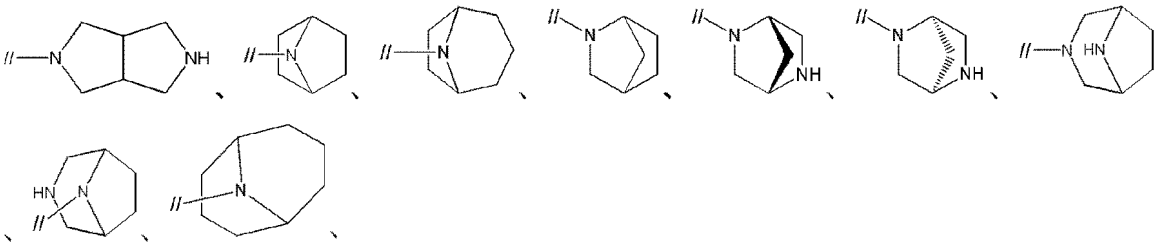
2,7-diazaspiro[3.4]octanyl, 3,7-diazaspiro[3.4]octanyl, 2,7-diazaspiro[3.5]nonanyl, 2,8-diazaspiro[3.5]nonanyl, 3,7-diazaspiro[3.5]nonanyl, 3,8-diazaspiro[4.4]nonanyl, 3,8-diazaspiro[4.5]decanyl, or 9-oxa-diazaspiro[3.5]nonanyl, and
R3 represents amino, methylamino, ethylamino, isopropylamino, dimethylamino, cyclobutylamino, or methyl, wherein when two or more R3s are present, R3s may be identical or different.

[Claim 3]

The compound or a salt thereof according to claim 1 or 2, which satisfies the following conditions in Formula (I):
R4 represents halogen, nitro, cyano, carboxy, C1-C8 alkyl that may be substituted with halogen, amino, hydroxy, carboxy, carbamoyl, (C1-C6 alkyl)carbamoyl, (C1-C6 alkyl)carbonylamino, C1-C6 alkoxy, (C1-C6 alkyl)carbonyl, C3-C7 cycloalkyl, hydroxy(C3-C7 cycloalkyl), or (C1-C6 alkyl)carbonyloxy, C2-C6 alkenyl, C1-C6 alkoxy that may be substituted with hydroxy or monocyclic 5- to 10-membered unsaturated hydrocarbon, C3-C7 cycloalkyl that may be substituted with hydroxy, hydroxy(C1-C4 alkyl), (C1-C4 alkoxy)(C1-C4 alkyl), hydroxy(C3-C7 cycloalkyl), or (C6-C14 aromatic hydrocarbon)-substituted carbamoyl, mono- or di(C1-C6 alkyl)amino, or carbamoyl that may be substituted with C1-C6 alkyl, wherein when two or more R4s are present, R4s may be identical or different.

[Claim 4]

The compound represented by Formula (I) or a salt thereof according to any one of claims 1 to 3,
wherein
ring A represents pyrrolidinyl, piperidinyl, piperazinyl, azepanyl, diazepanyl,



2,7-diazaspiro[3.4]octanyl, 3,7-diazaspiro[3.4]octanyl, 2,7-diazaspiro[3.5]nonanyl, 2,8-diazaspiro[3.5]nonanyl, 3,7-diazaspiro[3.5]nonanyl, 3,8-diazaspiro[4.4]nonanyl, 3,8-diazaspiro[4.5]decanyl, or 9-oxa-diazaspiro[3.5]nonanyl,
ring B represents phenyl, naphthyl, pyridyl, pyrazolopyridyl, pyrazolopyrimidinyl, indolyl, indolinyl, 2-oxo-indolinyl, indazolyl, benzoimidazolyl, benzoisoxazolyl, benzothiazolyl,

benzotriazolyl, imidazopyridinyl, quinolinyl, isoquinolinyl, quinoxalinyl, quinazolinyl, phthalazinyl, 2-oxo-2,3-dihydrobenzo[d]oxazolyl, 1,3-dihydroisobenzofuranyl, dihydrobenzooxazinyl, benzodioxolyl, dihydrobenzodioxynyl, or 2-oxo-2,3-dihydrobenzo[d]thiazolyl,
X represents O or S,
R1 represents nitro or cyano,
R2 represents fluorine, and is present at the ortho position relative to R1 on the phenyl,
R3 represents amino, methylamino, ethylamino, isopropylamino, dimethylamino, cyclobutylamino, or methyl, wherein when two or more R3s are present, R3s may be identical or different, and
R4 represents fluorine, chlorine, bromine, iodine, nitro, cyano, carboxy, methyl, ethyl, n-propyl, isopropyl, tert-butyl, difluoromethyl, trifluoromethyl, fluoroethyl, aminoethyl, hydroxymethyl, hydroxyethyl, hydroxypropyl, hydroxydimethylethyl, hydroxymethylpropyl, hydroxymethylbutyl, hydroxyethylbutyl, carboxymethyl, carbamoylmethyl, methylcarbamoylmethyl, dimethylcarbamoylmethyl, acetylaminoethyl, methoxyethyl, hydroxycyclopropylmethyl, hydroxycyclopropylethyl, hydroxycyclobutylmethyl, methylcarbonyloxyethyl, isobutenyl, methoxy, hydroxypropoxy, cyclopropyl, hydroxymethyl cyclopropyl, methoxymethyl cyclopropyl, hydroxycyclopropyl cyclopropyl, phenylcarbamoyl cyclopropyl, benzyloxy, dimethylamino, carbamoyl, methylcarbamoyl, or dimethylcarbamoyl, wherein when two or more R4s are present, R4s may be identical or different, and
n is an integer of 0 to 3, wherein when n is 2 to 3, two to three R4s may be identical or different.

[Claim 5]

Compounds according to any one of the following (1) to (24) or a salt of the compounds according to any one of the following (1) to (24);

(1) 4-[5-[(3-exo)-3-amino-8-azabicyclo[3.2.1]octane-8-carbonyl]-2-[2-fluoro-4-(2-hydroxy-2-methyl-propyl)phenyl]phenyl]-2-fluoro-

benzonitrile,
(2) 4-[5-[(3S)-3-aminopyrrolidine-1-carbonyl]-2-[2-fluoro-4-(2-hydroxy-2-methyl-propyl)phenyl]phenyl]-2-fluoro-benzonitrile,
(3) 4-[5-[(3-endo)-3-amino-8-azabicyclo[3.2.1]octane-8-carbonyl]-2-[2-fluoro-4-(2-hydroxy-2-methyl-propyl)phenyl]phenyl]-2-fluoro-benzonitrile,
(4) (S)-5'-(3-aminopyrrolidine-1-carbonyl)-3-fluoro-2'-(6-fluoro-1-(2-hydroxy-2-methylpropyl)-1H-indol-5-yl)-[1,1'-biphenyl]-4-carbonitrile,
(5) 5'-((3-endo)-3-amino-8-azabicyclo[3.2.1]octane-8-carbonyl)-2'-(6,7-difluoro-1-(2-hydroxy-2-methylpropyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile,
(6) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'',3-difluoro-4''-(2-hydroxy-2-methylpropyl)-[1,1':2',1''-terphenyl]-4-carbonitrile-isomer-B,
(7) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'-(6,7-difluoro-1-methyl-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile-isomer-B,
(8) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-3-fluoro-2'-(6-fluoro-1-(2-hydroxy-2-methylpropyl)-1H-indazol-5-yl)-[1,1'-biphenyl]-4-carbonitrile-isomer-B,
(9) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'-(6,7-difluoro-1-(2-hydroxy-2-methylpropyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile-isomer-B,
(10) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'-(6,7-difluoro-1-(2-hydroxy-2-methylpropyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile,
(11) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'-(7-chloro-6-fluoro-1-(2-hydroxy-2-methylpropyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile-isomer-X,
(12) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-

carbonyl)-2'-(7-chloro-6-fluoro-1-(2-hydroxy-2-methylpropyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile,

(13) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'-(1-(2-ethyl-2-hydroxybutyl)-6,7-difluoro-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile-isomer-X,

(14) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'-(1-(2-ethyl-2-hydroxybutyl)-6,7-difluoro-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile,

(15) (S)-5'-(3-aminopyrrolidine-1-carbonyl)-3-fluoro-2'-(5-fluoro-3-(2-hydroxy-2-methylpropyl)benzo[d]isoxazol-6-yl)-[1,1'-biphenyl]-4-carbonitrile,

(16) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'-(7-(difluoromethyl)-6-fluoro-1-(2-hydroxy-2-methylpropyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile-isomer-X,

(17) 5-(5-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-4'-cyano-3'-fluoro-[1,1'-biphenyl]-2-yl)-6-fluoro-1-(2-hydroxy-2-methylpropyl)-1H-benzo[d][1,2,3]triazole-7-carbonitrile-isomer-X,

(18) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'-(7-(difluoromethyl)-6-fluoro-1-(2-hydroxy-2-methylpropyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile,

(19) 5'-((3-endo)-3-amino-8-azabicyclo[3.2.1]octane-8-carbonyl)-2'-(7-chloro-6-fluoro-1-((1-hydroxycyclobutyl)methyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile,

(20) 5'-((3-endo)-3-amino-8-azabicyclo[3.2.1]octane-8-carbonyl)-2'-(7-(difluoromethyl)-6-fluoro-1-((1-hydroxycyclobutyl)methyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile,

(21) 5-(5-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-

carbonyl)-4'-cyano-3'-fluoro-[1,1'-biphenyl]-2-yl)-6-fluoro-1-((1-hydroxycyclobutyl)methyl)-1H-benzo[d][1,2,3]triazole-7-carbonitrile-isomer-X,

(22) 5-(5-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-4'-cyano-3'-fluoro-[1,1'-biphenyl]-2-yl)-6-fluoro-1-((1-hydroxycyclobutyl)methyl)-1H-benzo[d][1,2,3]triazole-7-carbonitrile,

(23) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'-(7-bromo-6-fluoro-1-(2-hydroxy-2-methylpropyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile-isomer-X,

(24) 5'-((1S,2S,4R)-rel-2-amino-7-azabicyclo[2.2.1]heptane-7-carbonyl)-2'-(7-bromo-6-fluoro-1-(2-hydroxy-2-methylpropyl)-1H-benzo[d][1,2,3]triazol-5-yl)-3-fluoro-[1,1'-biphenyl]-4-carbonitrile.

[Claim 6]

An LSD1 inhibitor comprising the compound or a salt thereof according to any one of claims 1 to 5, as an active ingredient.

[Claim 7]

A pharmaceutical composition comprising the compound or a salt thereof according to any one of claims 1 to 5.

[Claim 8]

The pharmaceutical composition according to claim 7, which is an orally administered composition.

[Claim 9]

An antitumor agent comprising the compound or a salt thereof according to any one of claims 1 to 5, as an active ingredient.

[Claim 10]

The compound or a salt thereof according to any one of claims 1 to 5, for use in the treatment of a cancer patient.

[Claim 11]

Use of the compound or a salt thereof according to any one of claims 1 to 5 in the manufacture of an agent for treating an LSD1-related cancer.