

azabicyclo[3.2.1]octan-8-yl group,
(iii-9) amino group,
(iii-10) methylamino group, ethylamino group, isopropylamino
group, hydroxyethylamino group, dimethylamino group, phenyl
5 methylamino group,
(iii-11) C1-C6 alkoxy group that may have one or more of either
halogen, C3-C7 saturated cycloalkyl group, or monocyclic or
bicyclic unsaturated heterocycloalkyl group having at least one
heteroatom selected from the group consisting of N, S, and O as a
10 substituent,
(iii-12) benzyloxy group that may have one or more carbamoyl
group as a substituent
(iii-13) C1-C4 alkylthio group,
(iii-14) C1-C4 alkylsulfonyl group, or
15 (iii-15) aminosulfonyl group.
[0056]

When R₅ has one or more R₆, R₆ in Formula (A) is more
preferably one of:

(iv-1) fluorine, chlorine
20 (iv-2) hydroxy group,
(iv-3) cyano group,
(iv-4) methyl group,
(iv-7) 3-fluoro pyrrolidinyl group, morpholinyl group,
thiomorpholinyl group, 3-hydroxy azetidiny group, azetidiny
25 group,
(iv-9) amino group,
(iv-10) methylamino group,
(iv-11) C1-C6 alkoxy group that may have one or more of either
halogen or C3-C7 saturated cycloalkyl group as a substituent, or
30 (iv-13) C1-C4 alkylthio group.
[0057]

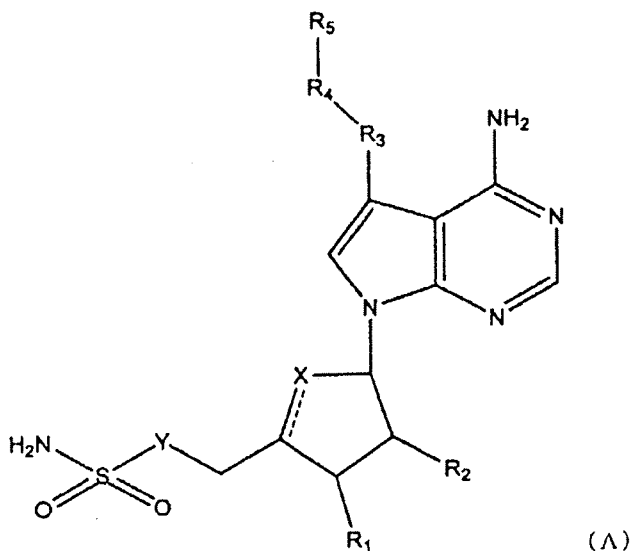
When R₅ has one or more R₆, R₆ in Formula (A) is more
preferably one of:

(v-1) fluorine,
35 (v-4) methyl group,

CLAIMS

[Claim 1]

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



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wherein:

is a single bond or a double bond;

X is -O-, -CH₂-, or -CH=;

10

Y is -NH- or -O-;

R₁ is hydrogen, fluorine, a hydroxy group, a cyano group, or an amino group;

R₂ is hydrogen, fluorine, a hydroxy group, a cyano group, or an amino group;

15

R₃ is a vinylene group, an ethynylene group, a C₆-C₁₄ arylene group, or a monocyclic or bicyclic heteroarylene group having at least one heteroatom selected from the group consisting of N, S, and O;

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R₄ is a bond, a methylene group, or a C₃-C₇ cycloalkylidene group;

R₅ is a C₃-C₇ saturated cycloalkyl group that may have one or more R₆, a C₆-C₁₀ unsaturated cycloalkyl group that may

have one or more R_6 , or a monocyclic or bicyclic unsaturated heterocycloalkyl group that has at least one heteroatom selected from the group consisting of N, S, and O, and that may have one or more R_6 ;

5 R_6 is

halogen,

a hydroxy group,

a cyano group,

10 a C1-C6 alkyl group that may have one or more phenoxy group as a substituent,

a carbamoyl group,

a C1-C6 alkoxy-carbonyl group,

15 a monocyclic or bicyclic unsaturated heterocycloalkyl group having at least one heteroatom selected from the group consisting of N, S, and O,

a monocyclic or bicyclic saturated heterocycloalkyl group that has at least one heteroatom selected from the group consisting of N, S, and O, and that may have one or more of either halogen, hydroxy group, carboxyl group, or C1-C6 alkyl group as a

20 substituent,

an amino group,

a mono- or di-(C1-C4 alkyl) amino group that may have one or more hydroxy group or phenyl group as a substituent,

25 a C1-C6 alkoxy group that may have one or more of either halogen, C3-C7 saturated cycloalkyl group, or monocyclic or bicyclic unsaturated heterocycloalkyl group having at least one heteroatom selected from the group consisting of N, S, and O as a substituent,

30 a benzyloxy group that may have one or more carbamoyl group as a substituent,

a C1-C6 alkylthio group,

a C1-C6 alkylsulfonyl group, or

an aminosulfonyl group,

35 when two or more R_6 are present, the plurality of R_6 may be the same or different.

[Claim 2]

The compound or a salt thereof according to Claim 1, wherein in Formula (A), R₁ is hydrogen, fluorine, or a hydroxy group;

R₂ is hydrogen, fluorine, or a hydroxy group; and

R₃ is an ethynylene group, or a monocyclic or bicyclic heteroarylene group having 1 to 4 of at least one kind of heteroatom selected from the group consisting of N, S, and O.

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[Claim 3]

The compound or a salt thereof according to Claim 1 or 2, wherein in Formula (A), R₁ is a hydroxy group;

R₂ is hydrogen or a hydroxy group;

15

R₃ is an ethynylene group, or a monocyclic heteroarylene group having 2 of at least one kind of heteroatom selected from the group consisting of N, S and O;

R₅ is a C3-C7 saturated cycloalkyl group that may have one or more R₆, a C6-C10 unsaturated cycloalkyl group that may have one or more R₆, or a monocyclic or bicyclic unsaturated heterocycloalkyl group that has at least one heteroatom selected from the group consisting of N, S, and O, and that may have one or more R₆; and

R₆ is halogen; a hydroxy group; a cyano group; a C1-C6 alkyl group that may have one or more phenoxy group as a substituent; a carbamoyl group; a C1-C6 alkoxy carbonyl group; a monocyclic or bicyclic unsaturated heterocycloalkyl group having at least one heteroatom selected from the group consisting of N, S and O; a monocyclic or bicyclic saturated heterocycloalkyl group that has at least one heteroatom selected from the group consisting of N, S, and O, and that may have one or more of either halogen, hydroxy group, carboxyl group, or C1-C6 alkyl group as a substituent; an amino group; a mono- or di-(C1-C4 alkyl) amino group that may have one or more hydroxy group or phenyl group as a substituent; a C1-C6 alkoxy group that may have

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one or more of either halogen, C3-C7 saturated cycloalkyl group, or monocyclic or bicyclic unsaturated heterocycloalkyl group having at least one heteroatom selected from the group consisting of N, S, and O as a substituent; a benzyloxy group that may have
5 one or more carbamoyl group as a substituent; a C1-C4 alkylthio group; a C1-C4 alkylsulfonyl group; or an aminosulfonyl group (when two or more R₆ are present, the plurality of R₆ may be the same or different).

10 [Claim 4]

A compound or a salt thereof according to any one of Claims 1 to 3, wherein, in Formula (A), R₁ is a hydroxy group;

R₂ is hydrogen or a hydroxy group;

R₃ is an ethynylene group, or a monocyclic
15 heteroarylene group having 2 of at least one kind of heteroatom selected from the group consisting of N, S and O;

R₅ is a C3-C7 saturated cycloalkyl group that may have one or more R₆; a C6-C10 unsaturated cycloalkyl group that may have one or more R₆, or a monocyclic or bicyclic unsaturated
20 heterocycloalkyl group that has at least one heteroatom selected from the group consisting of N, S, and O, and that may have one or more R₆; and

R₆ is fluorine; chlorine; a hydroxy group; a cyano group; a C1-C6 alkyl group that may have one or more phenoxy
25 group as a substituent; a carbamoyl group; a C1-C6 alkoxy carbonyl group; a pyridinyl group; an azetidiny group; a hydroxy azetidiny group; a thiomorpholinyl group; a dioxide thiomorpholinyl group; a methyl piperazinyl group; a hydroxy piperidinyl group; an oxopiperidinyl group; a piperidinyl group;
30 a hydroxy pyrrolidinyl group; an oxopyrrolidinyl group; a pyrrolidinyl group; a carboxyl pyrrolidinyl group; a fluoro pyrrolidinyl group; a morpholinyl group; a 9-oxa-3-azabicyclo[3.3.1]nonan-3-yl group; a 3-oxa-8-

azabicyclo[3.2.1]octan-8-yl group; an amino group; a methylamino group; an ethylamino group; an isopropylamino group; a hydroxyethylamino group; a dimethylamino group; a phenyl methylamino group; a C1-C6 alkoxy group that may have one or more
5 of either halogen, C3-C7 saturated cycloalkyl group, or monocyclic or bicyclic unsaturated heterocycloalkyl group having at least one heteroatom selected from the group consisting of N, S, and O as a substituent; a benzyloxy group that may have one or more carbamoyl group as a substituent; a C1-C4 alkylthio group; a
10 C1-C4 alkylsulfonyl group; or an aminosulfonyl group (when two or more R₆ are present, the plurality of R₆ may be the same or different).

[Claim 5]

15 A compound or a salt thereof according to Claims 1 to 4, wherein, in Formula (A), R₁ is a hydroxy group;
R₂ is a hydroxy group;
R₃ is an ethynylene group;
R₄ is a bond;
20 R₅ is a C6-C10 unsaturated cycloalkyl group that may have one or more R₆; or a monocyclic or bicyclic unsaturated heterocycloalkyl group that has at least one heteroatom selected from the group consisting of N, S, and O, and that may have one or more R₆; and
25 R₆ is fluorine; chlorine; a hydroxy group; a cyano group; a methyl group; a 3-fluoro pyrrolidinyl group; a morpholinyl group; a thiomorpholinyl group; a 3-hydroxy azetidiny group; an azetidiny group; an amino group; a N-methylamino group; a C1-C6 alkoxy group that may have one or more
30 of either halogen or C3-C7 saturated cycloalkyl group as a substituent; or a C1-C4 alkylthio group (when two or more R₆ are present, the plurality of R₆ may be the same or different).

[Claim 6]

35 A compound or a salt thereof according to Claims 1 to

5, wherein in Formula (A), Y is -NH-;

R₁ is a hydroxy group;

R₂ is a hydroxy group;

R₃ is an ethynylene group;

5 R₄ is a bond;

R₅ is a phenyl group or a naphthyl group that may have one or more R₆; or a monocyclic or bicyclic unsaturated heterocycloalkyl group that has at least one heteroatom selected from the group consisting of N, S and O, and that may have one or
10 more R₆;

R₆ is fluorine; a methyl group; a 3-fluoro pyrrolidinyl; a 3-hydroxy azetidiny; azetidiny; an amino group; a N-methylamino group; a C1-C6 alkoxy group that may have one or more cyclopropyl group; or a C1-C4 alkylthio group (when two or
15 more R₆ are present, the plurality of R₆ may be the same or different).

[Claim 7]

The compound or a salt thereof according to any one of
20 Claims 1 to 6, wherein the compound or a salt thereof is at least one member selected from the group consisting of: 4-amino-5-[2-(2,6-difluoro phenyl)ethynyl]-7-[(2R,3R,4S,5R)-3,4-dihydroxy-5-[(sulfamoylamino)methyl]tetrahydrofuran-2-yl]pyrrolo[2,3-d]pyrimidine;
25 4-amino-5-[2-(4-amino-2,6-difluoro-phenyl)ethynyl]-7-[(2R,3R,4S,5R)-3,4-dihydroxy-5-[(sulfamoylamino)methyl]tetrahydrofuran-2-yl]pyrrolo[2,3-d]pyrimidine;
4-amino-5-[2-[2,6-difluoro-4-(methylamino)phenyl]ethynyl]-7-
30 [(2R,3R,4S,5R)-3,4-dihydroxy-5-[(sulfamoylamino)methyl]tetrahydrofuran-2-yl]pyrrolo[2,3-d]pyrimidine;
4-amino-5-[2-[2,6-difluoro-4-[(3R)-3-fluoro pyrrolidin-1-yl]phenyl]ethynyl]-7-[(2R,3R,4S,5R)-3,4-dihydroxy-5-
35 [(sulfamoylamino)methyl]tetrahydrofuran-2-yl]pyrrolo[2,3-

- d]pyrimidine;
4-amino-7-[(2R,3R,4S,5R)-3,4-dihydroxy-5-
[(sulfamoylamino)methyl]tetrahydrofuran-2-yl]-5-[2-(2-ethoxy-4,6-
difluoro-phenyl)ethynyl]pyrrolo[2,3-d]pyrimidine;
5 4-amino-5-[2-[2,6-difluoro-4-(3-hydroxy azetidin-1-
yl)phenyl]ethynyl]-7-[(2R,3R,4S,5R)-3,4-dihydroxy-5-
[(sulfamoylamino)methyl]tetrahydrofuran-2-yl]pyrrolo[2,3-
d]pyrimidine;
4-amino-5-[2-[4-(azetidin-1-yl)-2,6-difluoro-phenyl]ethynyl]-7-
10 [(2R,3R,4S,5R)-3,4-dihydroxy-5-[(sulfamoylamino)methyl]-
tetrahydrofuran-2-yl]pyrrolo[2,3-d]pyrimidine;
4-amino-7-[(2R,3R,4S,5R)-3,4-dihydroxy-5-
[(sulfamoylamino)methyl]tetrahydrofuran-2-yl]-5-[2-(2-ethoxy-6-
fluoro-phenyl)ethynyl]pyrrolo[2,3-d]pyrimidine;
15 4-amino-7-[(2R,3R,4S,5R)-3,4-dihydroxy-5-
[(sulfamoylamino)methyl]tetrahydrofuran-2-yl]-5-[2-(2-fluoro-6-
propoxy-phenyl)ethynyl]pyrrolo[2,3-d]pyrimidine;
8-[2-[4-amino-7-[(2R,3R,4S,5R)-3,4-dihydroxy-5-
[(sulfamoylamino)methyl]tetrahydrofuran-2-yl]pyrrolo[2,3-
20 d]pyrimidin-5-yl]ethynyl]-7-fluoro-4-methyl-2,3-dihydro-1,4-
benzoxazine;
4-amino-7-[(2R,3R,4S,5R)-3,4-dihydroxy-5-
[(sulfamoylamino)methyl]tetrahydrofuran-2-yl]-5-[2-(2-
ethylsulfanyl-6-fluoro-phenyl)ethynyl]pyrrolo[2,3-d]pyrimidine;
25 4-amino-5-[2-[2-(cyclopropyl methoxy)-6-fluoro-phenyl]ethynyl]-7-
[(2R,3R,4S,5R)-3,4-dihydroxy-5-
[(sulfamoylamino)methyl]tetrahydrofuran-2-yl]pyrrolo[2,3-
d]pyrimidine;
4-amino-7-[(1R,2S,3R,4R)-2,3-dihydroxy-4-
30 [(sulfamoylamino)methyl]cyclopentyl]-5-[2-(2-fluoro-6-
methylsulfanyl-phenyl)ethynyl]pyrrolo[2,3-d]pyrimidine;
8-[2-[4-amino-7-[(1R,2S,3R,4R)-2,3-dihydroxy-4-
[(sulfamoylamino)methyl]cyclopentyl]pyrrolo[2,3-d]pyrimidin-5-
yl]ethynyl]-7-fluoro-4-methyl-2,3-dihydro-1,4-benzoxazine;
35 4-amino-7-[(1R,4R,5S)-4,5-dihydroxy-3-

[(sulfamoylamino)methyl]cyclopent-2-en-1-yl]-5-[2-(2-ethoxy-6-fluoro-phenyl)ethynyl]pyrrolo[2,3-d]pyrimidine; and
4-amino-7-[(1R,4R,5S)-4,5-dihydroxy-3-
[(sulfamoylamino)methyl]cyclopent-2-en-1-yl]-5-[2-(2-fluoro-6-
5 methylsulfanyl-phenyl)ethynyl]pyrrolo[2,3-d]pyrimidine; and salts
of these compounds.

[Claim 8]

An NAE inhibitor comprising the compound or a salt
10 thereof according to any one of Claims 1 to 7, as an active
ingredient.

[Claim 9]

A pharmaceutical composition comprising the compound
15 or a salt thereof according to any one of Claims 1 to 7.

[Claim 10]

An antitumor agent comprising the compound or a salt
thereof according to any one of Claims 1 to 7.