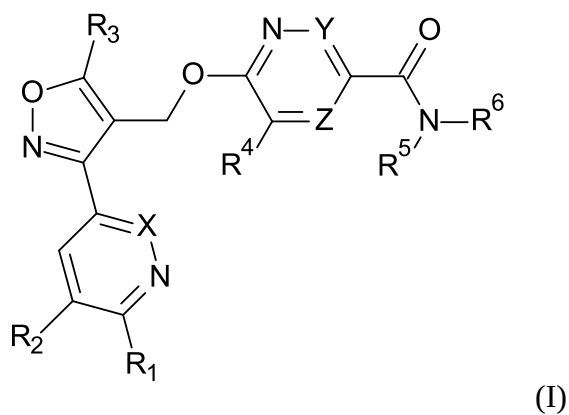


distribution, selective GABA_A $\alpha 5$ PAMs will restore GABAergic signaling in key brain regions (*e.g.* hippocampus, amygdala, nucleus accumbens and prefrontal cortex) without the side-effects of non-selective GABA_A modulators (*e.g.* benzodiazepines).

Summary

5 In one aspect, the present disclosure provides for compounds of formula (I)



wherein

X is selected from

i) N, and

10 ii) CH;

Y is selected from

i) N, and

ii) CR¹⁰;

Z is selected from

- i) N, and
- ii) CR¹¹;

R¹ is selected from

- 5 i) C₁₋₆-alkyl,
- ii) halo-C₁₋₆-alkyl,
- iii) C₁₋₆-alkoxy,
- iv) halo-C₁₋₆-alkoxy,
- v) hydroxy-C₁₋₆-alkyl,
- 10 vi) C₃₋₈-cycloalkyl,
- vii) halogen, and
- viii) amino substituted on the nitrogen atom by one or two substituents independently selected from
 - a. H,
 - 15 b. C₁₋₆-alkyl, and
 - c. C₃₋₈-cycloalkyl;

R² is selected from

- i) H, and
- ii) halogen;

20 R³ is selected from

- i) H,
- ii) C₁₋₆-alkyl,

- iii) C₃₋₈-cycloalkyl,
- iv) hydroxy-C₁₋₆-alkyl, and
- v) halo-C₁₋₆-alkyl;

R⁴ is selected from

- 5 i) H,
- ii) C₁₋₆-alkyl,
- iii) C₁₋₆-alkoxy,
- iv) C₃₋₈-cycloalkyl, and
- v) halogen;

10 R⁵ is H;

R⁶ is selected from

- i) H,
- ii) C₁₋₆-alkyl,
- iii) C₃₋₈-cycloalkyl substituted with R⁷, R⁸ and R⁹,
- 15 iv) C₃₋₈-cycloalkyl-C₁₋₆-alkyl substituted with R⁷, R⁸ and R⁹,
- v) C₁₋₆-alkylsulfonyl-C₁₋₆-alkyl,
- vi) cyano-C₁₋₆-alkyl,
- vii) hydroxy-C₁₋₆-alkyl,
- viii) dihydroxy-C₁₋₆-alkyl,
- 20 ix) halo-C₁₋₆-alkyl,
- x) heterocycloalkyl substituted with R⁷, R⁸ and R⁹, and
- xi) heterocycloalkyl-C₁₋₆-alkyl substituted with R⁷, R⁸ and R⁹;

R⁷, R⁸ and R⁹ are independently selected from

- i) H,
- ii) C₁₋₆-alkyl,
- iii) C₁₋₆-alkoxy,
- 5 iv) C₁₋₆-alkoxyalkyl,
- v) C₁₋₆-alkoxycarbonyl,
- vi) cyano,
- vii) C₃₋₈-cycloalkoxy,
- viii) C₃₋₈-cycloalkyl,
- 10 ix) halo-C₁₋₆-alkoxy,
- x) halo-C₁₋₆-alkyl,
- xi) halogen,
- xii) hydroxy,
- xiii) hydroxy-C₁₋₆-alkyl, and
- 15 xiv) oxo;

R¹⁰ is selected from

- i) H,
- ii) C₁₋₆-alkyl,
- iii) C₁₋₆-alkoxy,
- 20 iv) C₃₋₈-cycloalkyl, and
- v) halogen;

R¹¹ is selected from

- i) H,
- ii) C₁₋₆-alkyl,
- iii) C₁₋₆-alkoxy,
- iv) C₃₋₈-cycloalkyl, and
- 5 v) halogen;

or R⁵ and R¹⁰ together form -(CH₂)_n;

or R⁵ and R¹¹ together form -(CH₂)_n;

or R⁵ and R⁶ together with the nitrogen atom to which they are attached form a heterocycloalkyl substituted with R⁷, R⁸ and R⁹;

10 n is selected from 1 and 2;

or pharmaceutically acceptable salts.

In another aspect, the present disclosure provides for a compound as described herein for use as therapeutically active substance.

15 In another aspect, the present disclosure provides for a pharmaceutical composition comprising a compound as described herein and a therapeutically inert carrier.

In another aspect, the present disclosure provides for a compound as described herein for use in the treatment or prophylaxis of Alzheimer's disease, mild cognitive impairment, age-related cognitive decline, negative and/or cognitive symptoms associated with schizophrenia, bipolar disorders, autism spectrum disorder Angelman syndrome, Rett syndrome, Prader-Willi syndrome, epilepsy, post-traumatic stress disorder, amyotrophic lateral sclerosis, fragile-X disorder.

25 In another aspect, the present disclosure provides for the use of a compound as described herein for in the preparation of a medicament for the treatment or prophylaxis of Alzheimer's disease, mild cognitive impairment, age-related cognitive decline, negative and/or cognitive symptoms associated with schizophrenia, bipolar disorders, autism spectrum disorder Angelman syndrome, Rett syndrome, Prader-Willi syndrome, epilepsy, post-traumatic stress disorder, amyotrophic lateral sclerosis, fragile-X disorder.

The term “C₁₋₆-alkoxy” denotes a group of the formula -O-R', wherein R' is an C₁₋₆-alkyl group. Examples of C₁₋₆-alkoxy groups include methoxy, ethoxy, n-propoxy, isopropoxy, n-butoxy, isobutoxy and tert-butoxy. Particular examples are methoxy and ethoxy. In the case of R¹, particular example is methoxy.

- 5 The term “C₁₋₆-alkoxy-C₁₋₆-alkyl” denotes an C₁₋₆-alkyl group wherein one of the hydrogen atoms of the C₁₋₆-alkyl group has been replaced by an C₁₋₆-alkoxy group. Examples of C₁₋₆-alkoxy-C₁₋₆-alkyl groups include include methoxymethyl, ethoxymethyl, methoxymethyl, ethoxyethyl, methoxypropyl and ethoxypropyl. Particular alkoxyalkyl group is methoxyethyl.

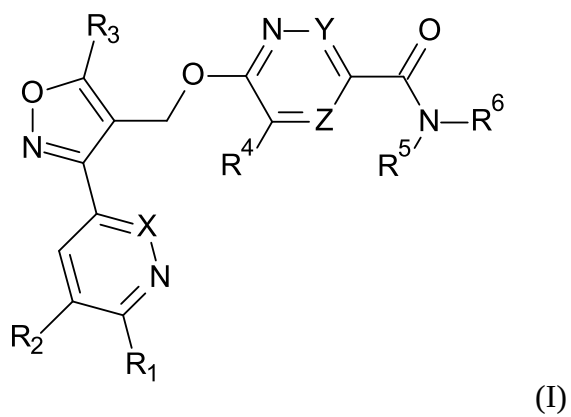
- 10 The term “C₁₋₆-alkoxycarbonyl” denotes a group of the formula -C(O)-R', wherein R' is a C₁₋₆-alkoxy group. Examples of C₁₋₆-alkoxycarbonyl groups include groups wherein R' is methoxy, ethoxy, n-propoxy, isopropoxy, n-butoxy, isobutoxy or tert-butoxy Particular example of C₁₋₆-alkoxycarbonyl is a group wherein R' is ethoxy.

- 15 The term “C₁₋₆-alkyl” denotes a monovalent linear or branched saturated hydrocarbon group of 1 to 6 carbon atoms. Examples of C₁₋₆-alkyl include methyl, ethyl, propyl, isopropyl, n-butyl, iso-butyl, sec-butyl, tert-butyl and pentyl. Particular C₁₋₆-alkyl groups are methyl, ethyl, isopropyl and tert-butyl. In the case of R¹, more particular example is methyl. In the case of R³, more particular examples are methyl and ethyl. In the case of R³, further more particular example is methyl.

- 20 The term “C₁₋₆-alkylsulfonyl” denotes a group of the formula -S(O)₂-R', wherein R' is a C₁₋₆-alkyl group. Examples of C₁₋₆-alkylsulfonyl include groups wherein R' is methyl, ethyl, propyl, isopropyl, n-butyl, iso-butyl, sec-butyl, tert-butyl or pentyl.

Claims

1. Compounds of formula (I)



wherein

5 X is selected from

- i) N, and
- ii) CH;

Y is selected from

- i) N, and
- 10 ii) CR^{10} ;

Z is selected from

- i) N, and
- ii) CR^{11} ;

R^1 is selected from

- 15 i) C_{1-6} -alkyl,
- ii) halo- C_{1-6} -alkyl,
- iii) C_{1-6} -alkoxy,
- iv) halo- C_{1-6} -alkoxy,

- v) hydroxy-C₁₋₆-alkyl,
- vi) C₃₋₈-cycloalkyl,
- vii) halogen, and
- viii) amino substituted on the nitrogen atom by one or two substituents independently selected from

5

- a. H,
- b. C₁₋₆-alkyl, and
- c. C₃₋₈-cycloalkyl;

R² is selected from

10

- i) H, and
- ii) halogen;

R³ is selected from

15

- i) H,
- ii) C₁₋₆-alkyl,
- iii) C₃₋₈-cycloalkyl,
- iv) hydroxy-C₁₋₆-alkyl, and
- v) halo-C₁₋₆-alkyl;

R⁴ is selected from

20

- i) H,
- ii) C₁₋₆-alkyl,
- iii) C₁₋₆-alkoxy,
- iv) C₃₋₈-cycloalkyl, and

v) halogen;

R⁵ is H;

R⁶ is selected from

i) H,

5 ii) C₁₋₆-alkyl,

iii) C₃₋₈-cycloalkyl substituted with R⁷, R⁸ and R⁹,

iv) C₃₋₈-cycloalkyl-C₁₋₆-alkyl substituted with R⁷, R⁸ and R⁹,

v) C₁₋₆-alkylsulfonyl-C₁₋₆-alkyl,

vi) cyano-C₁₋₆-alkyl,

10 vii) hydroxy-C₁₋₆-alkyl,

viii) dihydroxy-C₁₋₆-alkyl,

ix) halo-C₁₋₆-alkyl,

x) heterocycloalkyl substituted with R⁷, R⁸ and R⁹, and

xi) heterocycloalkyl-C₁₋₆-alkyl substituted with R⁷, R⁸ and R⁹;

15 R⁷, R⁸ and R⁹ are independently selected from

i) H,

ii) C₁₋₆-alkyl,

iii) C₁₋₆-alkoxy,

iv) C₁₋₆-alkoxyalkyl,

20 v) C₁₋₆-alkoxycarbonyl,

vi) cyano,

vii) C₃₋₈-cycloalkoxy,

- viii) C₃₋₈-cycloalkyl,
- ix) halo-C₁₋₆-alkoxy,
- x) halo-C₁₋₆-alkyl,
- xi) halogen,
- 5 xii) hydroxy,
- xiii) hydroxy-C₁₋₆-alkyl, and
- xiv) oxo;

R¹⁰ is selected from

- i) H,
- 10 ii) C₁₋₆-alkyl,
- iii) C₁₋₆-alkoxy,
- iv) C₃₋₈-cycloalkyl, and
- v) halogen;

R¹¹ is selected from

- 15 i) H,
- ii) C₁₋₆-alkyl,
- iii) C₁₋₆-alkoxy,
- iv) C₃₋₈-cycloalkyl, and
- v) halogen;

20 or R⁵ and R¹⁰ together form -(CH₂)_n;

or R⁵ and R¹¹ together form -(CH₂)_n;

or R⁵ and R⁶ together with the nitrogen atom to which they are attached form a heterocycloalkyl substituted with R⁷, R⁸ and R⁹;

n is selected from 1 and 2;

or pharmaceutically acceptable salts.

5 2. A compound according to claim 1, wherein

X is selected from

i) N, and

ii) CH;

Y is selected from

10 i) N, and

ii) CR¹⁰;

Z is selected from

i) N, and

ii) CR¹¹;

15 R¹ is selected from

i) C₁₋₆-alkyl,

ii) halo-C₁₋₆-alkyl,

iii) C₁₋₆-alkoxy,

iv) C₃₋₈-cycloalkyl,

20 v) halogen, and

vi) amino substituted on the nitrogen atom by two independently selected C₁₋₆-alkyl;

R² is selected from

- i) H,
- ii) halogen;

R³ is selected from

- i) H,
- 5 ii) C₁₋₆-alkyl,
- iii) C₃₋₈-cycloalkyl, and
- iv) halo-C₁₋₆-alkyl;

R⁴ is selected from

- i) H, and
- 10 ii) C₁₋₆-alkyl;

R⁵ is H;

R⁶ is selected from

- i) H,
- ii) C₁₋₆-alkyl,
- 15 iii) C₃₋₈-cycloalkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from
 - a. H,
 - b. C₁₋₆-alkyl,
 - c. C₁₋₆-alkoxy,
 - 20 d. C₁₋₆-alkoxyalkyl,
 - e. C₁₋₆-alkoxycarbonyl,
 - f. cyano,

g. C₃₋₈-cycloalkoxy,

h. halo-C₁₋₆-alkoxy,

i. halo-C₁₋₆-alkyl,

j. halogen,

5 k. hydroxy, and

l. hydroxy-C₁₋₆-alkyl;

iv) C₃₋₈-cycloalkyl-C₁₋₆-alkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from

a. H,

10 b. C₁₋₆-alkyl,

c. C₁₋₆-alkoxy,

d. C₁₋₆-alkoxyalkyl,

e. C₁₋₆-alkoxycarbonyl,

f. cyano,

15 g. C₃₋₈-cycloalkoxy,

h. halo-C₁₋₆-alkoxy,

i. halo-C₁₋₆-alkyl,

j. halogen,

k. hydroxy, and

20 l. hydroxy-C₁₋₆-alkyl;

v) C₁₋₆-alkylsulfonyl-C₁₋₆-alkyl,

vi) cyano-C₁₋₆-alkyl,

- vii) dihydroxy-C₁₋₆-alkyl,
- viii) halo-C₁₋₆-alkyl,
- ix) heterocycloalkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from

- 5
- a. H,
 - b. C₁₋₆-alkyl,
 - c. hydroxy, and
 - d. oxo;

and wherein the heterocycloalkyl is selected from

- 10
- a. oxetanyl,
 - b. tetrahydrofuranyl,
 - c. tetrahydropyranyl,
 - d. oxepanyl,
 - e. oxabicyclo[2.2.1]heptanyl,
- 15
- f. oxaspiro[3.3]heptanyl,
 - g. azetidiny,
 - h. tetrahydrothiophenyl, and
 - i. tetrahydrothiopyranyl; and

- 20
- x) oxetanyl-C₁₋₆-alkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from
- a. H,
 - b. hydroxy;

R¹⁰ is selected from

- i) H, and
- ii) halogen;

R¹¹ is selected from

- i) H,
- ii) C₁₋₆-alkyl, and
- iii) C₁₋₆-alkoxy;

or R⁵ and R¹⁰ together form -(CH₂)_n-;

or R⁵ and R¹¹ together form -(CH₂)_n-;

or R⁵ and R⁶ together with the nitrogen atom to which they are attached form a heterocycloalkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from

- a. H,
- b. C₁₋₆-alkyl,
- c. C₁₋₆-alkoxy,
- d. cyano,
- e. halogen,
- f. hydroxy, and
- g. oxo;

and wherein the heterocycloalkyl is selected from

- a. azetidinyI,
- b. pyrrolidinyl,
- c. piperidinyl,
- d. morpholinyl,

- e. thiomorpholinyl,
- f. oxaazabicyclo[3.1.1]heptanyl,
- g. oxaazabicyclo[2.2.1]heptanyl,
- h. azaspiro[3.3]heptanyl,
- 5 i. oxaazaspiro[3.3]heptanyl,
- j. thiaazaspiro[3.3]heptanyl;

n is 1;

or pharmaceutically acceptable salts.

3. A compound according to claim 1, wherein

10 X is selected from

- i) N, and
- ii) CH;

Y is selected from

- i) N, and
- 15 ii) CR¹⁰;

Z is selected from

- i) N, and
- ii) CR¹¹;

R¹ is selected from

- 20 i) C₁₋₆-alkyl,
- ii) halo-C₁₋₆-alkyl,
- iii) C₁₋₆-alkoxy,

- iv) C₃₋₈-cycloalkyl,
- v) halogen, and
- vi) amino substituted on the nitrogen atom by two independently selected C₁₋₆-alkyl;

5 R² is selected from

- i) H,
- ii) halogen;

R³ is selected from

- i) H,
- 10 ii) C₁₋₆-alkyl,
- iii) C₃₋₈-cycloalkyl, and
- iv) halo-C₁₋₆-alkyl;

R⁴ is selected from

- i) H, and
- 15 ii) C₁₋₆-alkyl;

R⁵ is H;

R⁶ is selected from

- i) H,
- ii) C₁₋₆-alkyl,
- 20 iii) C₃₋₈-cycloalkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from
 - a. H,
 - b. C₁₋₆-alkyl,

- c. C₁₋₆-alkoxy,
 - d. C₁₋₆-alkoxyalkyl,
 - e. C₁₋₆-alkoxycarbonyl,
 - f. cyano,
 - 5 g. C₃₋₈-cycloalkoxy,
 - h. halo-C₁₋₆-alkoxy,
 - i. halo-C₁₋₆-alkyl,
 - j. halogen,
 - k. hydroxy, and
 - 10 l. hydroxy-C₁₋₆-alkyl;
- iv) C₃₋₈-cycloalkyl-C₁₋₆-alkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from
- a. H,
 - b. C₁₋₆-alkyl,
 - 15 c. C₁₋₆-alkoxy,
 - d. C₁₋₆-alkoxyalkyl,
 - e. C₁₋₆-alkoxycarbonyl,
 - f. cyano,
 - g. C₃₋₈-cycloalkoxy,
 - 20 h. halo-C₁₋₆-alkoxy,
 - i. halo-C₁₋₆-alkyl,
 - j. halogen,

k. hydroxy, and

l. hydroxy-C₁₋₆-alkyl;

v) C₁₋₆-alkylsulfonyl-C₁₋₆-alkyl,

vi) cyano-C₁₋₆-alkyl,

5 vii) dihydroxy-C₁₋₆-alkyl,

viii) halo-C₁₋₆-alkyl,

ix) heterocycloalkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from

a. H,

10 b. C₁₋₆-alkyl,

c. hydroxy, and

d. oxo;

and wherein the heterocycloalkyl is selected from

a. oxetanyl,

15 b. tetrahydrofuranyl,

c. tetrahydropyranyl,

d. oxepanyl,

e. oxabicyclo[2.2.1]heptanyl,

f. oxaspiro[3.3]heptanyl,

20 g. azetidiny,

h. tetrahydrothiophenyl, and

i. tetrahydrothiopyranyl; and

x) oxetanyl-C₁₋₆-alkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from

a. H,

b. hydroxy;

5 R¹⁰ is selected from

i) H, and

ii) halogen;

R¹¹ is selected from

i) H,

10 ii) C₁₋₆-alkyl, and

iii) C₁₋₆-alkoxy;

or pharmaceutically acceptable salts.

4. A compound according to claim 1, wherein

X is selected from

15 i) N, and

ii) CH;

Y is selected from

i) N, and

ii) CR¹⁰;

20 Z is CR¹¹;

R¹ is C₁₋₆-alkyl;

R² is selected from

- i) H,
- ii) halogen;

R³ is selected from

- i) C₁₋₆-alkyl,
- 5 ii) C₃₋₈-cycloalkyl, and
- iii) halo-C₁₋₆-alkyl;

R⁴ is selected from

- i) H, and
- ii) C₁₋₆-alkyl;

10 R⁵ is H;

R⁶ is selected from

- i) C₁₋₆-alkyl,
- ii) C₃₋₈-cycloalkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from

- 15 a. H,
- b. C₁₋₆-alkyl,
- c. C₁₋₆-alkoxyalkyl, and
- d. halo-C₁₋₆-alkyl;

iii) hydroxy-C₁₋₆-alkyl,

20 iv) halo-C₁₋₆-alkyl,

v) heterocycloalkyl substituted with R⁷, R⁸ and R⁹, wherein R⁷, R⁸ and R⁹ are independently selected from

- a. H, and

b. C₁₋₆-alkyl;

and wherein the heterocycloalkyl is selected from

a. oxetanyl,

b. tetrahydrofuranyl,

5 c. tetrahydropyranyl,

d. oxepanyl, and

e. oxaspiro[3.3]heptanyl;

R¹⁰ is H;

R¹¹ is H;

10 or pharmaceutically acceptable salts.

5. A compound according to claim 1, wherein

X is CH;

Y is N;

Z is CR¹¹;

15 R¹ is C₁₋₆-alkyl;

R² is H;

R³ is C₁₋₆-alkyl;

R⁴ is H;

R⁵ is H;

20 R⁶ is heterocycloalkyl substituted with R⁷, R⁸ and R⁹;

R⁷, R⁸ and R⁹ are H;

R¹¹ is H;

or pharmaceutically acceptable salts.

6. A compound according to claim 1, wherein

X is CH;

Y is N;

5 Z is CR¹¹;

R¹ is C₁₋₆-alkyl;

R² is H;

R³ is C₁₋₆-alkyl;

R⁴ is H;

10 R⁵ is H;

R⁶ is tetrahydropyranyl substituted with R⁷, R⁸ and R⁹;

R⁷, R⁸ and R⁹ are H;

R¹¹ is H;

or pharmaceutically acceptable salts.

15 7. A compound according to any one of claims 1 to 6, selected from

6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-N-tetrahydropyran-4-yl-pyridine-3-carboxamide;

N-(cyclopropylmethyl)-6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyridine-3-carboxamide;

20 N-ethyl-6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyridine-3-carboxamide;

N-((1S)-1-(hydroxymethyl)butyl)-6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyridine-3-carboxamide;

6-((5-cyclopropyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-N-((1S)-1-(hydroxymethyl)butyl)pyridine-3-carboxamide;

6-((5-cyclopropyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-N-tetrahydropyran-4-yl-pyridine-3-carboxamide;

5 (S)-N-(1-hydroxypentan-2-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

10 2-isobutyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-1H-pyrrolo[3,4-c]pyridin-3(2H)-one;

2-methyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-1H-pyrrolo[3,4-c]pyridin-3(2H)-one;

N-isobutyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)nicotinamide;

15 2-(2-hydroxyethyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-1H-pyrrolo[3,4-c]pyridin-3(2H)-one;

(S)-2-(1-hydroxypentan-2-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-1H-pyrrolo[3,4-c]pyridin-3(2H)-one;

(S)-N-(1-hydroxypentan-2-yl)-5-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyrazine-2-carboxamide;

20 5-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyrazine-2-carboxamide;

N-(1,1-dioxothiolan-3-yl)-6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyridine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)nicotinamide;

25 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-((3S)-1,1-dioxothiolan-3-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)-1,2-oxazol-4-yl)methoxy)pyridine-3-carboxamide;

N-((3R)-1,1-dioxothiolan-3-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)-1,2-oxazol-4-yl)methoxy)pyridine-3-carboxamide;

5 N-(1,1-dioxothian-4-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)-1,2-oxazol-4-yl)methoxy)pyridine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-2-(tetrahydro-2H-pyran-4-yl)-1H-pyrrolo[3,4-c]pyridin-3(2H)-one;

10 N-((1S,2R)-2-Hydroxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-((1S,2S)-2-Hydroxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-((1R,2R)-2-Hydroxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

15 N-cyclopropyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-((1R,2S)-2-hydroxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

20 N-((3S,4R)-3-hydroxytetrahydropyran-4-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-(2-hydroxyethyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-(1,1-dioxothian-4-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)-1,2-oxazol-4-yl)methoxy)pyridazine-3-carboxamide;

25 N-(cyclopropylmethyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-(cyclopropylmethyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-(2-cyanoethyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

5 (RS)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1,1,1-trifluoropropan-2-yl)pyridazine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(oxetan-3-yl)pyridazine-3-carboxamide;

10 (RS)-N-(1,1-dioxothiolan-3-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)-1,2-oxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-ethyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-isopropyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

15 N-isobutyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(2,2,2-trifluoroethyl)pyridazine-3-carboxamide;

20 N-tert-butyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-(3,3-difluorocyclobutyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-(4,4-difluorocyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

25 6-((3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-N-tetrahydropyran-4-yl-pyridine-3-carboxamide;

(RS)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydrofuran-3-yl)pyridazine-3-carboxamide;

N-methyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

5 (3,3-difluoroazetidin-1-yl)(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

(3,3-difluoropyrrolidin-1-yl)(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

10 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-((3-methyloxetan-3-yl)methyl)pyridazine-3-carboxamide;

6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-N-(oxetan-3-ylmethyl)pyridazine-3-carboxamide;

N-((3-hydroxyoxetan-3-yl)methyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

15 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-((3R,4R)-3-methyltetrahydropyran-4-yl)pyridazine-3-carboxamide;

(4,4-difluoropiperidin-1-yl)(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

20 N-(1-(methoxymethyl)cyclopropyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

(3-methoxyazetidin-1-yl)(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

(3-hydroxy-3-methylazetidin-1-yl)(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

25 azetidin-1-yl(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

(RS)-N-(2,2-dimethyltetrahydropyran-4-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1-(trifluoromethyl)cyclopropyl)pyridazine-3-carboxamide;

5 (6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)(morpholino)methanone;

(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)(2-oxa-6-azaspiro[3.3]heptan-6-yl)methanone;

10 4-methyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-4-yl)pyridazine-3-carboxamide;

(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)(6-oxa-1-azaspiro[3.3]heptan-1-yl)methanone;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1,1,1-trifluoro-2-methylpropan-2-yl)pyridazine-3-carboxamide;

15 (3-fluoroazetidin-1-yl)(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

(3-hydroxyazetidin-1-yl)(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

20 (3-fluoro-3-methylazetidin-1-yl)(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

ethyl 1-(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamido)cyclopropanecarboxylate;

N-(1-cyanocyclopropyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

25 5-methyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

5-methyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-4-yl)pyridazine-3-carboxamide;

N-(1,1-dioxothian-4-yl)-5-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyrazine-2-carboxamide;

5 N-(2-hydroxy-1,1-dimethyl-ethyl)-5-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyrazine-2-carboxamide;

N-cyclopropyl-5-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyrazine-2-carboxamide;

10 (RS)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(3-methyltetrahydrofuran-3-yl)pyridazine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1-methylcyclopropyl)pyridazine-3-carboxamide;

5-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-N-(oxetan-3-yl)pyrazine-2-carboxamide;

15 5-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-N-(2,2,2-trifluoroethyl)pyrazine-2-carboxamide;

N-(4-hydroxy-2-methylbutan-2-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

20 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(2-methyl-4-(methylsulfonyl)butan-2-yl)pyridazine-3-carboxamide;

(S)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(3-methyltetrahydrofuran-3-yl)pyridazine-3-carboxamide;

(R)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(3-methyltetrahydrofuran-3-yl)pyridazine-3-carboxamide;

25 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(3-methyloxetan-3-yl)pyridazine-3-carboxamide;

1-(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carbonyl)azetidine-3-carbonitrile;

N-(1-(hydroxymethyl)cyclopropyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

5 N-(4,4-difluorocyclohexyl)-5-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyrazine-2-carboxamide;

(S)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydrofuran-3-yl)pyridazine-3-carboxamide;

10 (S)-N-(1-cyanobutan-2-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

(R)-5-((5-Methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydrofuran-3-yl)pyrazine-2-carboxamide;

N-(2-Hydroxyethyl)-5-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyrazine-2-carboxamide;

15 2-(1,1-Dioxothian-4-yl)-6-[[5-methyl-3-(6-methylpyridin-3-yl)-1,2-oxazol-4-yl]methoxy]-1H-pyrrolo[3,4-c]pyridin-3-one;

(S)-5-((5-Methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydrofuran-3-yl)pyrazine-2-carboxamide;

20 2-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-6-tetrahydropyran-4-yl-7H-pyrrolo[3,4-b]pyridin-5-one;

N-(1,1-Dioxothiolo-3-yl)-5-((5-methyl-3-(6-methylpyridin-3-yl)-1,2-oxazol-4-yl)methoxy)pyrazine-2-carboxamide;

N-(Cyclopropylmethyl)-5-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyrazine-2-carboxamide;

25 2-(4,4-Difluorocyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-1H-pyrrolo[3,4-c]pyridin-3(2H)-one;

6-((5-Methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(4-methyltetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

(R)-5-((5-Methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-3-yl)pyrazine-2-carboxamide;

5 (R)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-3-yl)pyridazine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(2-oxaspiro[3.3]heptan-6-yl)pyridazine-3-carboxamide;

10 5-((5-Methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(2-oxaspiro[3.3]heptan-6-yl)pyrazine-2-carboxamide;

5-((5-Methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-((cis)-4-(trifluoromethyl)cyclohexyl)pyrazine-2-carboxamide;

(S)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-3-yl)pyridazine-3-carboxamide;

15 N-((cis)-4-Hydroxy-4-methylcyclohexyl)-5-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyrazine-2-carboxamide;

N-((trans)-4-Hydroxy-4-methylcyclohexyl)-5-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyrazine-2-carboxamide;

20 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(4-methyltetrahydro-2H-pyran-4-yl)nicotinamide;

6-((5-Methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-2-(tetrahydro-2H-thiopyran-4-yl)-1H-pyrrolo[3,4-c]pyridin-3(2H)-one;

6-((5-methyl-3-(6-methylpyridazin-3-yl)isoxazol-4-yl)methoxy)-N-tetrahydropyran-4-ylpyridine-3-carboxamide;

25 N-((1S)-1-(hydroxymethyl)butyl)-6-((5-methyl-3-(6-methylpyridazin-3-yl)isoxazol-4-yl)methoxy)pyridine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(4-methyltetrahydrothiopyran-4-yl)pyridazine-3-carboxamide;

N-(4-methyl-1,1-dioxidotetrahydro-2H-thiopyran-4-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

5 (2,2-dioxido-2-thia-6-azaspiro[3.3]heptan-6-yl)(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

(2,2-Dioxido-2-thia-6-azaspiro[3.3]heptan-6-yl)(5-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyrazin-2-yl)methanone;

10 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1-methylcyclopentyl)pyridazine-3-carboxamide;

5-((5-Methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(4,4,4-trifluorobutyl)pyrazine-2-carboxamide;

N-(1-isopropylazetidin-3-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

15 6-((5-Methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-2-(2-oxaspiro[3.3]heptan-6-yl)-1H-pyrrolo[3,4-c]pyridin-3(2H)-one;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1-methylcyclobutyl)pyridazine-3-carboxamide;

20 6-((5-Ethyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)nicotinamide;

N-(1,1-Dioxidotetrahydro-2H-thiopyran-4-yl)-6-((5-ethyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)nicotinamide;

6-((5-ethyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-4-yl)pyridazine-3-carboxamide;

25 6-((5-ethyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(3-methyloxetan-3-yl)pyridazine-3-carboxamide;

N-cyclopropyl-6-((5-ethyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

(R)-N-(1-hydroxypentan-2-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)nicotinamide;

5 6-((5-(difluoromethyl)-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-N-((1S)-1-(hydroxymethyl)butyl)pyridine-3-carboxamide;

6-((5-(difluoromethyl)-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-N-tetrahydropyran-4-yl-pyridine-3-carboxamide;

10 N-((3R,4S)-3-hydroxytetrahydropyran-4-yl)-6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyridine-3-carboxamide;

6-((3-(6-cyclopropyl-3-pyridyl)-5-methyl-isoxazol-4-yl)methoxy)-N-tetrahydropyran-4-yl-pyridine-3-carboxamide;

6-((3-(6-cyclopropyl-3-pyridyl)-5-methyl-isoxazol-4-yl)methoxy)-N-((1S)-1-(hydroxymethyl)butyl)pyridine-3-carboxamide;

15 6-((3-(6-cyclopropyl-3-pyridyl)-5-methyl-isoxazol-4-yl)methoxy)-N-(1,1-dioxothian-4-yl)pyridine-3-carboxamide;

N-((1R,2S)-3,3-difluoro-2-hydroxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)nicotinamide;

20 N-((1R,2R)-2-hydroxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)nicotinamide;

N-((1S,2R)-3,3-difluoro-2-hydroxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)nicotinamide;

N-((3S,4R)-3-hydroxytetrahydro-2H-pyran-4-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)nicotinamide;

25 N-((1S,2S)-2-hydroxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)nicotinamide;

2-fluoro-N-methyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)nicotinamide;

6-((3-(6-methoxypyridin-3-yl)-5-methylisoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)nicotinamide;

5 N-isopropyl-6-((3-(6-methoxypyridin-3-yl)-5-methylisoxazol-4-yl)methoxy)nicotinamide;

(S)-N-(1-hydroxypentan-2-yl)-6-((3-(6-methoxypyridin-3-yl)-5-methylisoxazol-4-yl)methoxy)nicotinamide;

(1,1-dioxidothiomorpholino)(6-((3-(6-methoxypyridin-3-yl)-5-methylisoxazol-4-yl)methoxy)pyridin-3-yl)methanone;

10 (S)-6-((3-(6-(dimethylamino)pyridin-3-yl)-5-methylisoxazol-4-yl)methoxy)-N-(1-hydroxypentan-2-yl)nicotinamide;

4-methoxy-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)nicotinamide;

15 6-((3-(5-fluoro-6-methyl-3-pyridyl)-5-methyl-isoxazol-4-yl)methoxy)-N-tetrahydropyran-4-yl-pyridine-3-carboxamide;

6-((3-(5-fluoro-6-methyl-3-pyridyl)-5-methyl-isoxazol-4-yl)methoxy)-N-isopropyl-pyridine-3-carboxamide;

6-((3-(5-fluoro-6-methyl-3-pyridyl)-5-methyl-isoxazol-4-yl)methoxy)-N-((1S)-1-(hydroxymethyl)butyl)pyridine-3-carboxamide;

20 N-((1r,4r)-4-hydroxy-4-methylcyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-(1,3-dihydroxy-2-methylpropan-2-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

25 N-((1s,4s)-4-hydroxy-4-methylcyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-((1R,3S)-3-hydroxy-3-(trifluoromethyl)cyclopentyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-((1S,3R)-3-hydroxy-3-(trifluoromethyl)cyclopentyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

(1S,4S)-2-oxa-5-azabicyclo[2.2.1]heptan-5-yl(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

5 N-((1RS,3RS)-3-hydroxycyclopentyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

trans-N-(4-hydroxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

10 trans-N-(4-methoxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

(6,6-difluoro-2-azaspiro[3.3]heptan-2-yl)(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

N-(3-hydroxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

15 3-oxa-6-azabicyclo[3.1.1]heptan-6-yl(6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazin-3-yl)methanone;

cis-N-(4-methoxycyclohexyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

20 6-((3-(6-chloropyridin-3-yl)-5-methylisoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

(S)-6-((3-(6-chloropyridin-3-yl)-5-methylisoxazol-4-yl)methoxy)-N-(tetrahydrofuran-3-yl)pyridazine-3-carboxamide;

6-((3-(6-chloropyridin-3-yl)-5-methylisoxazol-4-yl)methoxy)-N-(3-methyloxetan-3-yl)pyridazine-3-carboxamide;

25 6-((3-(6-chloropyridin-3-yl)-5-methylisoxazol-4-yl)methoxy)-N-(2-oxaspiro[3.3]heptan-6-yl)pyridazine-3-carboxamide;

trans-N-(3-methoxycyclobutyl)-6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

cis-N-(3-methoxycyclobutyl)-6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

5 6-((5-cyclopropyl-3-(6-methylpyridazin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

trans-N-((1RS,3RS)-3-methoxycyclopentyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

10 N-(6,6-difluorospiro[3.3]heptan-2-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

6-((5-(fluoromethyl)-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-4-yl)pyridazine-3-carboxamide;

cis-N-3-((2,2-difluoroethoxy)cyclobutyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

15 trans-N-3-((2,2-difluoroethoxy)cyclobutyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

cis-N-(3-ethoxycyclobutyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

20 trans-N-(3-ethoxycyclobutyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

cis-N-(3-(difluoromethoxy)cyclobutyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

trans-N-(3-(difluoromethoxy)cyclobutyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

25 trans-N-(3-cyclopropoxycyclobutyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

cis-N-(3-cyclopropoxycyclobutyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

6-((3-(6-chloropyridin-3-yl)-5-cyclopropylisoxazol-4-yl)methoxy)-N-(3-methyloxetan-3-yl)pyridazine-3-carboxamide;

5 6-((3-(6-chloropyridin-3-yl)-5-cyclopropylisoxazol-4-yl)methoxy)-N-(2-oxaspiro[3.3]heptan-6-yl)pyridazine-3-carboxamide;

(S)-6-((3-(6-chloropyridin-3-yl)-5-cyclopropylisoxazol-4-yl)methoxy)-N-(tetrahydrofuran-3-yl)pyridazine-3-carboxamide;

10 6-((5-cyclopropyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(3-methyloxetan-3-yl)pyridazine-3-carboxamide;

6-((5-cyclopropyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(2-oxaspiro[3.3]heptan-6-yl)pyridazine-3-carboxamide;

(S)-6-((5-cyclopropyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydrofuran-3-yl)pyridazine-3-carboxamide;

15 6-((5-cyclopropyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-4-yl)pyridazine-3-carboxamide;

6-((5-(fluoromethyl)-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1-(trifluoromethyl)cyclopropyl)pyridazine-3-carboxamide;

20 6-((5-methyl-3-(6-(trifluoromethyl)pyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-4-yl)pyridazine-3-carboxamide;

N-((2S)-7-oxabicyclo[2.2.1]heptan-2-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

(RS)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(oxepan-4-yl)pyridazine-3-carboxamide;

25 6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)-N-((1R,3S,4R)-7-oxabicyclo[2.2.1]heptan-3-yl)pyridazine-3-carboxamide;

N-((2R)-7-oxabicyclo[2.2.1]heptan-2-yl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-((1R,3R)-3-ethoxycyclopentyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

5 N-((1R,3R)-3-(2,2-difluoroethoxy)cyclopentyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

6-((5-(fluoromethyl)-3-(6-methylpyridazin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

10 6-((5-methyl-3-(6-methylpyridazin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

N-[4-(2,2-difluoroethoxy)cyclohexyl]-6-[[5-methyl-3-(6-methylpyridin-3-yl)-1,2-oxazol-4-yl]methoxy]pyridazine-3-carboxamide;

N-(4-ethoxycyclohexyl)-6-[[5-methyl-3-(6-methylpyridin-3-yl)-1,2-oxazol-4-yl]methoxy]pyridazine-3-carboxamide;

15 or pharmaceutically acceptable salts.

8. A compound according to any one of claims 1 to 7, selected from

N-((1S)-1-(hydroxymethyl)butyl)-6-((5-methyl-3-(6-methyl-3-pyridyl)isoxazol-4-yl)methoxy)pyridine-3-carboxamide;

20 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

N-cyclopropyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

(RS)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1,1,1-trifluoropropan-2-yl)pyridazine-3-carboxamide;

25 N-isopropyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-tert-butyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

N-(1-(methoxymethyl)cyclopropyl)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)pyridazine-3-carboxamide;

5 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1-(trifluoromethyl)cyclopropyl)pyridazine-3-carboxamide;

5-methyl-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-4-yl)pyridazine-3-carboxamide;

10 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1-methylcyclopropyl)pyridazine-3-carboxamide;

(S)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(3-methyltetrahydrofuran-3-yl)pyridazine-3-carboxamide;

(R)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(3-methyltetrahydrofuran-3-yl)pyridazine-3-carboxamide;

15 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(3-methyloxetan-3-yl)pyridazine-3-carboxamide;

(S)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydrofuran-3-yl)pyridazine-3-carboxamide;

20 6-((5-Methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(4-methyltetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

(R)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-3-yl)pyridazine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(2-oxaspiro[3.3]heptan-6-yl)pyridazine-3-carboxamide;

25 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(1-methylcyclobutyl)pyridazine-3-carboxamide;

6-((3-(5-fluoro-6-methyl-3-pyridyl)-5-methyl-isoxazol-4-yl)methoxy)-N-((1S)-1-(hydroxymethyl)butyl)pyridine-3-carboxamide;

6-((5-cyclopropyl-3-(6-methylpyridazin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

5 6-((5-(fluoromethyl)-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydropyran-4-yl)pyridazine-3-carboxamide;

6-((5-cyclopropyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(3-methyloxetan-3-yl)pyridazine-3-carboxamide;

10 (RS)-6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(oxepan-4-yl)pyridazine-3-carboxamide;

6-((5-(fluoromethyl)-3-(6-methylpyridazin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

6-((5-methyl-3-(6-methylpyridazin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide;

15 or pharmaceutically acceptable salts.

9. A compound according to any one of claims 1 to 8, wherein the compound is 6-((5-methyl-3-(6-methylpyridin-3-yl)isoxazol-4-yl)methoxy)-N-(tetrahydro-2H-pyran-4-yl)pyridazine-3-carboxamide or pharmaceutically acceptable salts.

20 10. A compound according to any one of claims 1 to 9 for use as therapeutically active substance.

11. A pharmaceutical composition comprising a compound according to any one of claims 1 to 9 and a therapeutically inert carrier.

25 12. A compound according to any one of claims 1 to 9 for use in the treatment or prophylaxis of Alzheimer's disease, mild cognitive impairment, age-related cognitive decline, negative and/or cognitive symptoms associated with schizophrenia, bipolar disorders, autism spectrum disorder, Angelman syndrome, Rett syndrome, Prader-Willi syndrome, epilepsy, post-traumatic stress disorder, amyotrophic lateral sclerosis, fragile-X disorder.

13. The use of a compound according to any one of claims 1 to 9 in the preparation of a medicament for the treatment or prophylaxis of Alzheimer's disease, mild cognitive impairment, age-related cognitive decline, negative and/or cognitive symptoms associated with schizophrenia, bipolar disorders, autism spectrum disorder Angelman syndrome, Rett syndrome, Prader-Willi syndrome, epilepsy, post-traumatic stress disorder, amyotrophic lateral sclerosis, fragile-X disorder.
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