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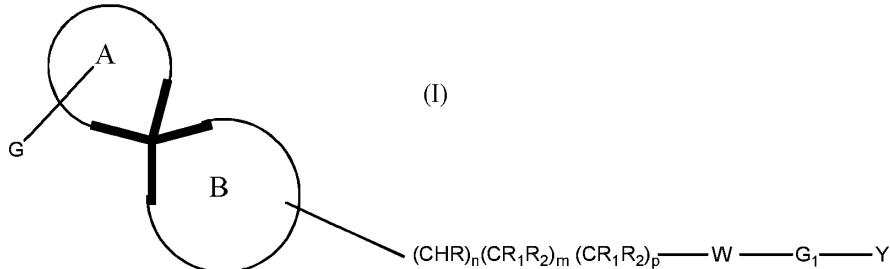
(54) Title **DOPAMINE D3 RECEPTOR ANTAGONISTS COMPOUNDS**

(56) References
Cited: WO-A2-2007/133561
WO-A1-2013/092893
WO-A1-03/091220
US-A- 3 654 305

Enclosed is a translation of the patent claims in Norwegian. Please note that as per the Norwegian Patents Acts, section 66i the patent will receive protection in Norway only as far as there is agreement between the translation and the language of the application/patent granted at the EPO. In matters concerning the validity of the patent, language of the application/patent granted at the EPO will be used as the basis for the decision. The patent documents published by the EPO are available through Espacenet (<http://worldwide.espacenet.com>) or via the search engine on our website here: <https://search.patentstyret.no/>

Patentkrav

1. Forbindelse av formel (I) eller et farmasøytisk akseptabelt salt derav,



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hvor

A er en mettet 3–6-leddet karbosyklig ring og en slik ring kan være substituert med én eller flere C_{1–4}-alkylgrupper;

B er en mettet 4–6-leddet heterosyklig ring, hvor ett eller to karbonatomer kan erstattes av et heteroatom valgt fra minst ett nitrogen eller et oksygen og det bindende atomet 10 er alltid et nitrogenatom; en slik ring kan også være substituert ved karbonatomene, eller muligens ved et annet nitrogenatom, med én eller flere C_{1–4}-alkylgrupper;

G er fenyl, bifenyl, naftyl eller pyridyl, som eventuelt kan være substituert med 1, 2, 3, 4 eller 5 substituenter valgt fra gruppen som består av: halogen, cyano, hydroksyl, amino, C_{1–4}-alkylamino, C_{1–4}-alkyl, C_{1–4}-alkoksy, halogenC_{1–4}-alkyl, halogenC_{1–4}-alkoksy, SF₅, 15 C(=O)NH₂ og C(=O)(O)_zR₃;

W er S, SO₂, O, CHR₂ eller NR₃;

n er 0 eller 1;

m er 1 eller 2;

p er 1 eller 2;

20 z er hver uavhengig 0 eller 1;

R er hydrogen eller C_{1–4}-alkyl; C_{1–4}-alkoksy;

R₁ er hver uavhengig hydrogen eller F, C_{1–4}-alkyl; OH, C_{1–4}-alkoksy;

R₂ er hver uavhengig hydrogen eller F, C_{1–4}-alkyl; OH, C_{1–4}-alkoksy;

R₃ er hver uavhengig hydrogen eller C_{1–4}-alkyl;

25 R₄ er hver uavhengig hydrogen eller C_{1–4}-alkyl; eller -C(=O)C_{1–4}-alkyl; -C(=O)C_{1–4}-alkoksyC_{1–4}-alkyl; -C(=O)C_{3–6}-sykloalkyl;

R₅ er hver uavhengig hydrogen eller C_{1–4}-alkyl;

R₆ er hver uavhengig hydrogen eller C_{1–4}-alkyl;

R₇ er hver uavhengig halogen, C_{1–4}-alkyl; OH, C_{1–4}-alkoksy;

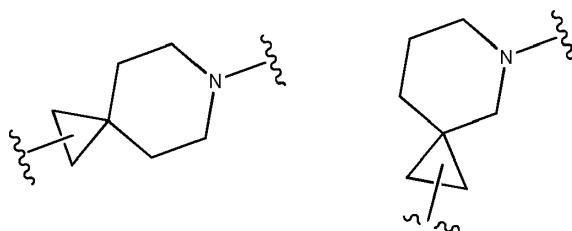
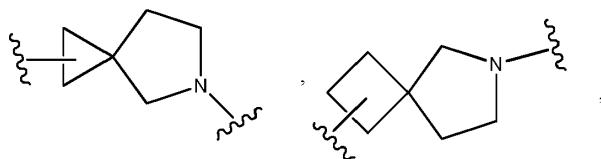
G_1 er en 5–6-leddet heteroaromatisk gruppe som inneholder minst ett ringnitrogen, som eventuelt kan være substituert med 1, 2 eller 3 substituenter uavhengig valgt fra gruppen som består av: halogen, cyano, hydroksyl, amino, C_{1-4} -alkylamino, C_{1-4} -alkyl, halogen C_{1-4} -alkyl, halogen C_{1-4} -alkoksy, C_{1-4} -alkoksy, $C(=O)NH_2$ og $C(=O)(O)_zR_3$;

- 5 Y er fenyl eller en del valgt fra gruppen som består av: 5–6-leddet heteroaromatisk gruppe, en 8–11-leddet heteroaromatisk gruppe, en mettet mono-3–7-leddet karbosyklisk gruppe og en 8–11-leddet bisyklistisk karbosyklisk gruppe, og for en hvilken som helst av slike grupper kan ett eller flere ringkarboner erstattes av $N(R_4)_z$, O, S; en hvilken som helst av gruppene kan eventuelt substitueres med 1, 2 eller 3 substituenter valgt fra: halogen, cyano, 10 hydroksyl, C_{1-4} -alkylamino, C_{1-4} -alkyl, C_{1-4} -alkoksy, halogen C_{1-4} -alkyl, halogen C_{1-4} -alkoksy, okso, $-NHC(=O)C_{1-4}$ -alkyl, $-NR_5R_6$, SF_5 , $-(CH_2)_zC(=O)NR_5R_6$, $-C(=O)(O)_zR_3$, $-C_{1-4}$ -alkylCN, $-SO_2NR_5R_6$, Y' eller OY';

Y' er fenyl, eller en 5–6-leddet heteroaromatisk gruppe eventuelt substituert med 1 eller 2 R_7 -grupper.

15

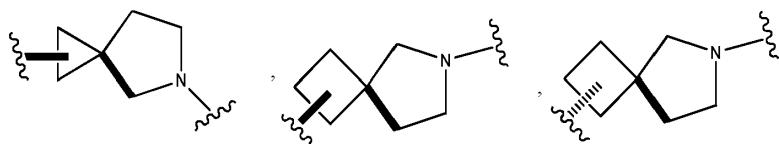
2. Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge krav 1, av formel (IA) der A og B av forbindelser av formel (I) er valgt fra følgende:



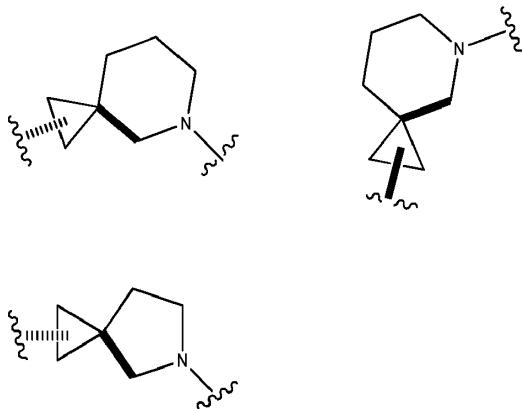
20

og hvori G , G_1 , W, Y, n, m, p, z, R, R_1 og R_2 er som definert i krav 1.

3. Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge krav 1, av formel (IB) der A og B av forbindelser av formel (I) er valgt fra følgende:



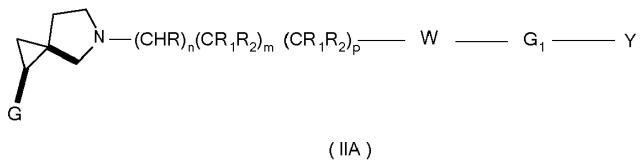
25



og hvori G, G₁, W, Y, n, m, p, z, R, R₁ og R₂ er som definert i krav 1.

5

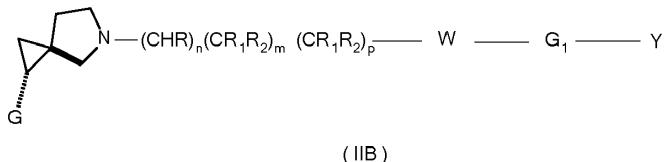
- 4.** Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge krav 1, av formel (IIA)



hvor G, G₁, W, Y, n, m, p, z, R, R₁ og R₂ er som definert i krav 1.

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- 5.** Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge krav 1, av formel (IIB)



hvor G, G₁, W, Y, n, m, p, z, R, R₁ og R₂ er som definert i krav 1.

15

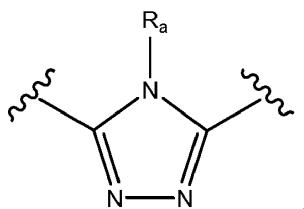
- 6.** Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge et hvilket som helst av kravene 1 til 5, hvor W er S og R, R₁ og R₂ er hydrogen.

- 7.** Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge et hvilket som helst av kravene 1 til 6, hvor G er fenyl eller pyridyl, eventuelt substituert med 1, 2, 3, 4 eller 5, for eksempel 1 eller 2, substituenter uavhengig valgt fra gruppen som består av: halogen, cyano, hydroksyl, amino, C₁₋₄-alkylamino, C₁₋₄-alkyl, halogenC₁₋₄-alkyl, halogenC₁₋₄-alkoksy, C₁₋₄-alkoksy, -C(=O)NH₂ og -C(=O)(O)_zR₃.

8. Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge et hvilket som helst av kravene 1 til 6, hvor G er fenyl eller pyridyl, eventuelt substituert med 1, 2 eller 3 grupper uavhengig valgt fra halogen, C₁₋₄-alkyl og halogenC₁₋₄-alkyl.

5 9. Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge et hvilket som helst av kravene 1 til 8, hvor G er valgt fra: fenyl, 4-trifluormetyl-fenyl, 2-fluor-4-trifluormetyl-fenyl, 2,4-difluorfenyl, 4-fluorfenyl, 2-trifluormetyl-fenyl, 2-trifluormetyl-4-fluorfenyl eller 3,5-diklorfenyl.

10 10. Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge et hvilket som helst av kravene 1 til 9, hvor G₁ er



hvor R_a er H eller C₁₋₄-alkyl; eventuelt hvor R_a er C₁₋₄-alkyl, for eksempel methyl.

15 11. Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge et hvilket som helst av kravene 1 til 10, hvor Y er valgt fra gruppen som består av:

- fenyl eventuelt substituert med én eller to substituenter valgt fra: cyano, -C(=O)NH₂, sulfonamid, acetyl, -CH₂CN, -CH₂C(=O)NH₂ og Y';
- en mettet mono-3–7-leddet karbosyklig gruppe der 0 eller 1 eller 2 karbonatomer er erstattet av et heteroatom uavhengig valgt fra O eller NR₃;
- en 8–11-leddet bisyklig karbosyklig gruppe, eventuelt substituert med ett eller flere C₁₋₄-alkyl;
- en 5–6-leddet heteroaromatisk gruppe eventuelt substituert med én eller to substituenter valgt fra: halogen, cyano, hydroksyl, C₁₋₄-alkyl, halogenC₁₋₄-alkyl, C₁₋₄-alkoksy, -(CH₂)_zC(=O)N(R₄R₅), Y' og OY';
- en 8–11-leddet heteroaromatisk gruppe der 1 eller 2 eller 3 atomkarboner kan erstattes av N, eventuelt substitueres med ett eller flere C₁₋₄-alkyl.

30 12. Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge et hvilket som helst av kravene 1 til 11, hvor Y er fenyl substituert med 1 eller 2 substituenter valgt fra

cyno,

-CH₂CN, -C(=O)R₃, -(CH₂)_zC(=O)NR₅R₆, -SO₂NH₂ og Y', hvori Y' er valgt fra oksadiazolyl, tetrazolyl, triazolyl og oksazolyl, der Y' eventuelt er substituert med C₁₋₄-alkyl; eventuelt er Y fenyl substituert med 1 substituent valgt fra cyano, CH₂CN, acetyl, -CH₂C(=O)NH₂,

- 5 -C(=O)NH₂, -SO₂NH₂ og Y', hvori Y' er valgt fra oksadiazolyl, tetrazolyl, triazolyl og oksazolyl, der Y' eventuelt er substituert med methyl;

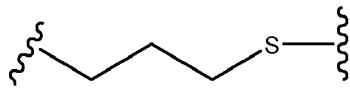
og hvori R₃, R₅, R₆ og z er som definert i krav 1.

- 13.** Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge et hvilket som helst av kravene 1 til 12, hvori Y er valgt fra pyridyl, pyrimidinyl og pyrazinyl, hvilken som helst av disse gruppene kan eventuelt substitueres med 1 eller 2 substituenter valgt fra:

fluor, cyno, C₁₋₄-alkyl, C₁₋₄-alkoksy, halogenC₁₋₄-alkyl, og -C(=O)NR₅R₆; eventuelt hvori Y er pyridyl, eventuelt substituert med 1 eller 2 substituenter valgt fra: C₁₋₄-alkyl og -C(=O)NH₂;

- 15 hvori, R₅ og R₆ er som definert i krav 1.

- 14.** Forbindelse eller et farmasøytisk akseptabelt salt derav, ifølge et hvilket som helst av kravene 1 til 13, hvori gruppe -(CHR)_n(CR₁R₂)_m(CR₁R₂)_pW- er



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15. Forbindelse, ifølge krav 1, valgt fra:

(1R,3R eller 1S,3S)-5-(3-{[4-metyl-5-(4-metyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (Enantiomer 1);

- 25 (1R,3S/1S,3R)-5-(3-{[4-metyl-5-(4-metyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S)-5-(3-{[4-metyl-5-(4-metyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}-propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (Enantiomer 1);

- 30 (1R,3S)-5-(3-{[4-metyl-5-(4-metyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}-propyl)-1-fenyl-5-azaspiro[2.4]heptan (Enantiomer 2);
 (1R,3S)-5-(3-{[4-metyl-5-(4-metyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}-propyl)-1-fenyl-5-azaspiro[2.4]heptan (Enantiomer 2);

(1R,3S/1S,3R)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (TRANS);
 (1S,3R eller 1R,3S)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (Enantiomer 1);
 5 (1R,3S eller 1S,3R)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (Enantiomer 2);
 (1S,3S/1R,3R)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (CIS);
 10 (1S,3S)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (Enantiomer 2);
 (1S,3S/1R,3R)-1-(2,4-difluorfenyl)-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (CIS);
 (1S,3S)-1-(2,4-difluorfenyl)-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);
 15 (1R,3S/1S,3R)-1-(4-fluorfenyl)-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (CIS);
 (1R,3S)-1-(4-fluorfenyl)-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);
 (1S,3S/1R,3R)-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[2-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (TRANS);
 20 (1S,3S eller 1R,3R)-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[2-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (TRANS, Enantiomer 2);
 (1S,3R eller 1R,3S)-1-[4-fluor-2-(trifluormetyl)fenyl]-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);
 25 (1S,3S eller 1R,3R)-1-[4-fluor-2-(trifluormetyl)fenyl]-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (TRANS, Enantiomer 1);
 30 (1S,3S/1R,3R)-1-(3,5-diklorfenyl)-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (TRANS);
 (1R,3S/1S,3R)-1-(3,5-diklorfenyl)-5-(3-{{[4-methyl-5-(4-methyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl}-5-azaspiro[2.4]heptan (CIS);

- (1S,3S/1R,3R)-5-(3-{[4-metyl-5-(oksan-4-yl)-4H-1,2,4-triazol-3-yl]-sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (TRANS);
- (1R,3R eller 1S,3S)-5-(3-{[4-metyl-5-(oksan-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (Enantiomer 1);
- 5 (1S,3S eller 1R,3R)-5-(3-{[4-metyl-5-(oksan-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (Enantiomer 2);
- (1R,3S/1S,3R)-5-(3-{[4-metyl-5-(oksan-4-yl)-4H-1,2,4-triazol-3-yl]-sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
- (1R,3S)-5-(3-{[4-metyl-5-(oksan-4-yl)-4H-1,2,4-triazol-3-yl]-sulfanyl}-propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (Enantiomer 1);
- 10 (1R,3S/1S,3R)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{[4-metyl-5-(oksan-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (TRANS);
- (1S,3R eller 1R,3S)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{[4-metyl-5-(oksan-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (Enantiomer 1);
- 15 (1S,3S/1R,3R)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{[4-metyl-5-(oksan-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (CIS);
- (1S,3S)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{[4-metyl-5-(oksan-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (Enantiomer 2);
- (1R,3S/1S,3R)-5-{3-[(4-metyl-5-{8-oksabisyklo[3.2.1]oktan-3-yl}-4H-1,2,4-triazol-3-yl)sulfanyl]propyl}-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
- 20 (1R,3S)-5-{3-[(4-metyl-5-{8-oksabisyklo[3.2.1]oktan-3-yl}-4H-1,2,4-triazol-3-yl)sulfanyl]propyl}-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (Enantiomer 1);
- (1S,3S/1R,3R)-5-{3-[(5-sykloheksyl-4-metyl-4H-1,2,4-triazol-3-yl)sulfanyl]propyl}-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (TRANS);
- 25 (1R,3R eller 1S,3S)-5-{3-[(5-sykloheksyl-4-metyl-4H-1,2,4-triazol-3-yl)sulfanyl]propyl}-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (Enantiomer 1);
- (1S,3S eller 1R,3R)-5-{3-[(5-sykloheksyl-4-metyl-4H-1,2,4-triazol-3-yl)sulfanyl]propyl}-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (Enantiomer 2);
- (1R,3S/1S,3R)-5-{3-[(5-sykloheksyl-4-metyl-4H-1,2,4-triazol-3-yl)-sulfanyl]propyl}-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
- 30 (1R,3S)-5-{3-[(5-sykloheksyl-4-metyl-4H-1,2,4-triazol-3-yl)-sulfanyl]propyl}-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (Enantiomer 1);

1-{4-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]piperidin-1-yl}etan-1-on (CIS);

5 1-{4-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]piperidin-1-yl}etan-1-on (CIS, Enantiomer 1);

1-{4-[5-(3-[(1S,3S)-1-[2-fluor-4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4-methyl-4H-1,2,4-triazol-3-yl]piperidin-1-yl}etan-1-on (CIS);

10 1-{4-[5-(3-[(1S,3S)-1-[2-fluor-4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4-methyl-4H-1,2,4-triazol-3-yl]piperidin-1-yl}etan-1-on (CIS);

15 3-metoksy-1-{4-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]piperidin-1-yl}propan-1-on (CIS, Enantiomer 1);

(1R,3S)-5-(3-[5-(1-syklopropankarbonylpiperidin-4-yl)-4-methyl-4H-1,2,4-triazol-3-yl]-sulfanyl)propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);

20 15 N-{4-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]sykloheksyl}acetamid (CIS);

N-{4-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]sykloheksyl}acetamid (CIS, Enantiomer 1);

25 20 (1R,3S/1S,3R)-5-(3-[(4-methyl-5-(morpholin-4-yl)-4H-1,2,4-triazol-3-yl)sulfanyl]propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S)-5-(3-[(4-methyl-5-(morpholin-4-yl)-4H-1,2,4-triazol-3-yl)sulfanyl]propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);

30 25 4-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]piperidin-2-on (CIS, Enantiomer 1);

25 30 5-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]piperidin-2-on (CIS, Enantiomer 1);

6-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (CIS);

5-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (CIS);

5-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (CIS, Enantiomer 1);

5-[5-(3-[(1S,3S)-1-[2-fluor-4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4-methyl-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (CIS);

5-[5-(3-[(1S,3S/1R,3R)-1-(2,4-difluorfenyl)-5-azaspiro[2.4]heptan-5-yl]propyl]-sulfanyl)-4-methyl-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (CIS);

5-[5-(3-[(1R,3S/1S,3R)-1-(4-fluorfenyl)-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4-methyl-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (CIS);

5 5-[4-methyl-5-(3-[(1S,3S/1R,3R)-1-[2-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (TRANS);

1-methyl-5-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (CIS, Enantiomer 1);

10 4-[4-methyl-5-(3-[(1R,3S/1S,3R)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (CIS);

4-[4-methyl-5-(3-[(1S,3S/1R,3R)-1-[2-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (TRANS);

15 1-methyl-4-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]-1,2-dihydropyridin-2-on (CIS, Enantiomer 1);

4-[5-(3-[(1S,3S)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)-4-methyl-4H-1,2,4-triazol-3-yl]-1-methyl-1,2-dihydropyridin-2-on (CIS, Enantiomer 1);

(1S,3S/1R,3S)-5-(3-{[4-methyl-5-(pyridin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (TRANS);

20 (1R,3S/1S,3R)-5-(3-{[4-methyl-5-(pyridin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S/1S,3R)-5-(3-{[4-methyl-5-(pyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

25 (1R,3S)-5-(3-{[4-methyl-5-(pyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);

(1S,3S/1R,3R)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{[4-methyl-5-(pyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (CIS);

(1S,3S)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{[4-methyl-5-(pyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);

30 (1S,3S/1R,3R)-1-(2,4-difluorfenyl)-5-(3-{[4-methyl-5-(pyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (CIS);

(1S,3S)-1-(2,4-difluorfenyl)-5-(3-{[4-methyl-5-(pyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);

- (1R,3S/1S,3R)-1-(4-fluorfenyl)-5-(3-{[4-methyl-5-(pyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (CIS);
- (1R,3S)-1-(4-fluorfenyl)-5-(3-{[4-methyl-5-(pyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);
- 5 (1S,3S/1R,3R)-5-(3-{[4-methyl-5-(pyridin-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (TRANS);
- (1R,3S/1S,3R)-5-(3-{[4-methyl-5-(pyridin-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
- (1R,3S/1S,3R)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-(3-{[4-methyl-5-(pyridin-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (TRANS);
- 10 (1R,3S/1S,3R)-5-(3-{[4-methyl-5-(2-metylpyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
- (1R,3S)-5-(3-{[4-methyl-5-(2-metylpyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
- 15 (1S,3S/1R,3R)-5-(3-{[4-methyl-5-(2-metylpyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[2-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (TRANS);
- (1R,3S/1S,3R)-5-(3-{[4-methyl-5-(6-metylpyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
- (1R,3S)-5-(3-{[4-methyl-5-(3-metylpyridin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
- 20 (1R,3S)-5-(3-{[5-(2,6-dimetylpyridin-3-yl)-4-methyl-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
- (1R,3S)-5-(3-{[5-(2-fluorpyridin-3-yl)-4-methyl-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
- 25 (1R,3S)-5-[3-(4-methyl-5-[2-(trifluormetyl)pyridin-3-yl]-4H-1,2,4-triazol-3-yl]sulfanyl)propyl]-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
- (1R,3S)-5-(3-{[5-(2-metoksypyridin-3-yl)-4-methyl-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
- (1R,3S)-5-(3-{[5-(2-metoksypyridin-3-yl)-4-methyl-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
- 30 (1R,3S)-5-[3-(4-methyl-5-[2-(trifluormetyl)pyridin-3-yl]-4H-1,2,4-triazol-3-yl]sulfanyl)propyl]-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
- 5-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl]-4H-1,2,4-triazol-3-yl]pyridin-2-karbonitril (CIS, Enantiomer 1);
- 4-[4-methyl-5-(3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl]-4H-1,2,4-triazol-3-yl]pyridin-2-karbonitril (CIS, Enantiomer 1);

5-[4-metyl-5-({3-[(1S,3S/1R,3R)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-2-karboksamid (TRANS);

5-[4-metyl-5-({3-[(1S,3S eller 1R,3R)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-2-karboksamid

5 (TRANS, Enantiomer 1);

5-[4-metyl-5-({3-[(1R,3R eller 1S,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-2-karboksamid
(TRANS, Enantiomer 2);

5-[4-metyl-5-({3-[(1R,3S/1S,3R)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-2-karboksamid (CIS);

5-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-2-karboksamid (CIS, Enantiomer 1);

5-[5-({3-[(1S,3S)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4-metyl-4H-1,2,4-triazol-3-yl]pyridin-2-karboksamid (CIS, Enantiomer 1);

15 6-metyl-5-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-2-karboksamid (CIS, Enantiomer 1);

6-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-3-karboksamid (CIS, Enantiomer 1);

20 6-[5-({3-[(1S,3S)-1-[2-fluor-4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4-metyl-4H-1,2,4-triazol-3-yl]pyridin-3-karboksamid (CIS, Enantiomer 1);

4-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-2-karboksamid (CIS, Enantiomer 1);

25 5-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-3-karboksamid (CIS, Enantiomer 1);

6-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-2-karboksamid (CIS, Enantiomer 1);

30 N-metyl-6-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]pyridin-2-karboksamid (CIS, Enantiomer 1);

(1R,3S/1S,3R)-5-(3-{[4-metyl-5-(pyridazin-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S)-5-(3-{[4-metyl-5-(pyridazin-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);

(1S,3S/1R,3R)-5-(3-{[4-methyl-5-(pyrimidin-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (TRANS);
 (1S,3S eller 1R,3R)-5-(3-{[4-methyl-5-(pyrimidin-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (TRANS, Enantiomer
 5 1);
 (1R,3R eller 1S,R3)-5-(3-{[4-methyl-5-(pyrimidin-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (TRANS, Enantiomer
 10 2);
 (1R,3S/1S,3R)-5-(3-{[4-methyl-5-(pyrimidin-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
 (1S,3S/1R,3R)-1-[2-fluor-4-(trifluormethyl)fenyl]-5-(3-{[4-methyl-5-(pyrimidin-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (CIS);
 (1S,3S eller 1R,3R)-5-(3-{[4-methyl-5-(pyrazin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (TRANS, Enantiomer
 15 1);
 (1R,3R eller 1S,R3)-5-(3-{[4-methyl-5-(pyrazin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (TRANS, Enantiomer
 2);
 (1R,3S/1S,3R)-5-(3-{[4-methyl-5-(pyrazin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
 (1R,3S)-5-(3-{[4-methyl-5-(pyrazin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);
 (1S,3S/1R,3R)-1-[2-fluor-4-(trifluormethyl)fenyl]-5-(3-{[4-methyl-5-(pyrazin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-5-azaspiro[2.4]heptan (CIS);
 25 (1R,3S/1S,3R)-5-(3-{[4-methyl-5-(6-metylpyrazin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
 (1R,3S/1S,3R)-5-(3-{[4-methyl-5-(5-metylpyrazin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
 (1R,3S/1S,3R)-5-(3-{[4-methyl-5-(3-metylpyrazin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
 30 (1R,3S/1S,3R)-5-(3-{[4-methyl-5-(3-metylpyrazin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS);
 (1R,3S)-5-(3-{[4-methyl-5-(3-metylpyrazin-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);
 5-[4-metyl-5-((3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]propyl)sulfanyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-karboksamid (CIS);

(1R,3S/1S,3R)-5-(3-{[4-metyl-5-(3-metyl-1,2-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S)-5-(3-{[4-metyl-5-(3-metyl-1,2-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}-propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);

5 (1R,3S/1S,3R)-5-(3-{[4-metyl-5-(4-metyl-1,3-tiazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S)-5-(3-{[4-metyl-5-(4-metyl-1,3-tiazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}-propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);

(1R,3S/1S,3R)-5-(3-{[4-metyl-5-(1,3-tiazol-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

10 (1R,3S)-5-(3-{[4-metyl-5-(1,3-tiazol-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);

(1R,3S/1S,3R)-5-(3-{[4-metyl-5-(1,3-tiazol-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);

(1R,3S/1S,3R)-5-(3-{[4-metyl-5-(1-metyl-1H-pyrazol-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

15 (1R,3S)-5-(3-{[4-metyl-5-(1-metyl-1H-pyrazol-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);

(1R,3S/1S,3R)-5-(3-{[4-metyl-5-(1-metyl-1H-pyrazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S)-5-(3-{[4-metyl-5-(1-metyl-1H-pyrazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 2);

20 (1R,3S/1S,3R)-5-(3-{[5-(furan-2-yl)-4-metyl-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S/1S,3R)-5-(3-{[5-(furan-3-yl)-4-metyl-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

25 (1R,3S/1S,3R)-5-(3-{[4-metyl-5-(tiofen-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S/1S,3R)-5-(3-{[4-metyl-5-(tiofen-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S/1S,3R)-5-(3-{[4-metyl-5-(1-metyl-1H-pyrrol-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

30 (1R,3S/1S,3R)-5-(3-{[4-metyl-5-(1-metyl-1H-pyrrol-2-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1S,3S/1R,3R)-5-(3-{[4-metyl-5-(1,2,3-tiadiazol-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (TRANS);

(1R,3S/1S,3R)-5-(3-{[4-metyl-5-(1,2,3-tiadiazol-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspiro[2.4]heptan (CIS);

(1R,3S)-5-(3-{[4-methyl-5-(1,2,3-tiadiazol-4-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 (1R,3S)-5-(3-{[4-methyl-5-(4-methyl-1,2,3-tiadiazol-5-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}-propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 5 (1R,3S)-5-[3-({4-methyl-5-[2-(pyridin-3-yl)-1,3-oksazol-5-yl]-4H-1,2,4-triazol-3-yl}-sulfanyl)propyl]-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 (1R,3S)-5-(3-{[4-methyl-5-(6-fenoksypyridin-3-yl)-4H-1,2,4-triazol-3-yl]sulfanyl}propyl)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 (1R,3S)-5-{3-[(4-methyl-5-{{1,2,4}triazolo[4,3-a]pyridin-6-yl}-4H-1,2,4-triazol-3-yl)-sulfanyl]propyl}-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 10);
 (1R,3S)-5-{3-[(4-methyl-5-{{1,2,4}triazolo[4,3-a]pyridin-7-yl}-4H-1,2,4-triazol-3-yl)-sulfanyl]propyl}-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 (1S,3S)-1-[2-fluor-4-(trifluormethyl)fenyl]-5-{3-[(4-methyl-5-{{1,2,4}triazolo[4,3-a]pyridin-7-yl}-4H-1,2,4-triazol-3-yl)sulfanyl]propyl}-5-azaspiro[2.4]heptan (CIS, Enantiomer 15);
 (1R,3S)-5-{3-[(4-methyl-5-{3-methyl-[1,2,4]triazolo[4,3-a]pyridin-6-yl}-4H-1,2,4-triazol-3-yl)sulfanyl]propyl}-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 (1R,3S)-5-{3-[(5-{1H-imidazo[4,5-b]pyridin-5-yl}-4-methyl-4H-1,2,4-triazol-3-yl)-sulfanyl]propyl}-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 20);
 (1R,3S)-5-[3-({4-methyl-5-[4-(1H-1,2,3,4-tetrazol-5-yl)fenyl]-4H-1,2,4-triazol-3-yl}-sulfanyl)propyl]-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 (1R,3S)-5-[3-({4-methyl-5-[4-(1,3,4-oksadiazol-2-yl)fenyl]-4H-1,2,4-triazol-3-yl}-sulfanyl)propyl]-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 25 (1R,3S)-5-[3-({4-methyl-5-[4-(5-methyl-1,2,4-oksadiazol-3-yl)fenyl]-4H-1,2,4-triazol-3-yl}sulfanyl)propyl]-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 (1R,3S)-5-[3-({4-methyl-5-[4-(4H-1,2,4-triazol-4-yl)fenyl]-4H-1,2,4-triazol-3-yl}sulfanyl)propyl]-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 (1R,3S)-5-[3-({4-methyl-5-[4-(1,3-oksazol-2-yl)fenyl]-4H-1,2,4-triazol-3-yl}sulfanyl)-propyl]-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 30);
 (1R,3S)-5-[3-({4-methyl-5-[3-(1,3-oksazol-2-yl)fenyl]-4H-1,2,4-triazol-3-yl}-sulfanyl)propyl]-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 (1R,3S)-5-[3-({4-methyl-5-[3-(1,3-oksazol-2-yl)fenyl]-4H-1,2,4-triazol-3-yl}-sulfanyl)propyl]-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan (CIS, Enantiomer 1);
 4-[4-methyl-5-({3-[(1R,3S)-1-[4-(trifluormethyl)fenyl]-5-azaspiro[2.4]heptan-5-yl]-propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]benzamid (CIS, Enantiomer 1);

4-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspido[2.4]heptan-5-yl]-propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]benzonitril (CIS, Enantiomer 1);

1-{4-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspido[2.4]heptan-5-yl]-propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]fenyl}etan-1-on (CIS, Enantiomer 1);

5 4-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspido[2.4]heptan-5-yl]-propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]benzen-1-sulfonamid (CIS, Enantiomer 1);

10 2-{4-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspido[2.4]heptan-5-yl]-propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]fenyl}acetonitril (CIS, Enantiomer 1);

2-{4-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspido[2.4]heptan-5-yl]-

10 propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]fenyl}acetamid (CIS, Enantiomer 1);

3-[4-metyl-5-({3-[(1R,3S)-1-[4-(trifluormetyl)fenyl]-5-azaspido[2.4]heptan-5-yl]-propyl}sulfanyl)-4H-1,2,4-triazol-3-yl]benzamid (CIS, Enantiomer 1);

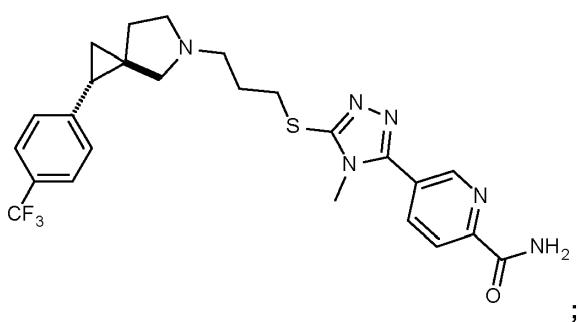
(1S,3S)-1-[2-fluor-4-(trifluormetyl)-fenyl]-5-{4-[4-metyl-5-(oksazan-4-yl)-4H-1,2,4-triazol-3-yl]butyl}-5-azaspido[2.4]heptan (Enantiomer 2);

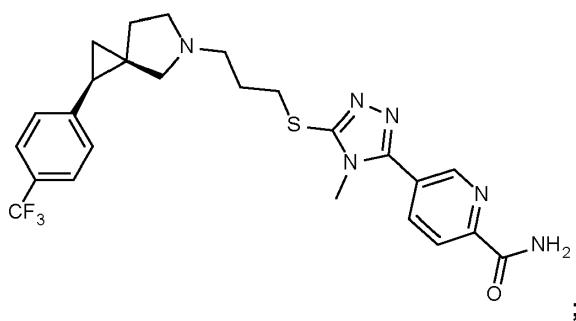
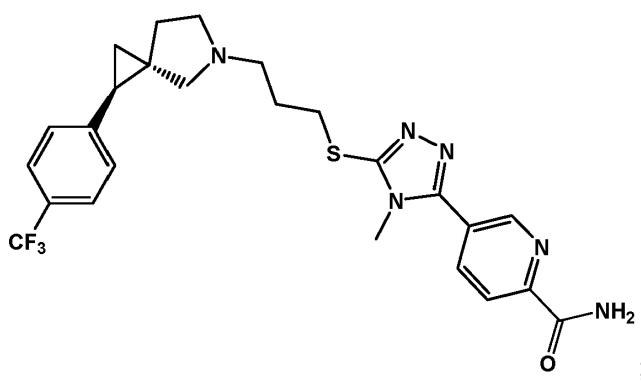
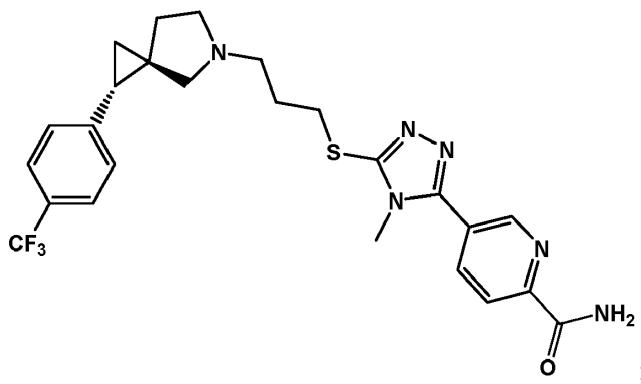
15 (1R,3S)-5-{4-[4-metyl-5-(4-metyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl]butyl}-1-[4-(trifluormetyl)fenyl]-5-azaspido[2.4]heptan (CIS, Enantiomer 1);

(1R,3S/1S,3R)-5-(3-{{4-metyl-5-(4-metyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl}sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspido[2.5]oktan (TRANS);

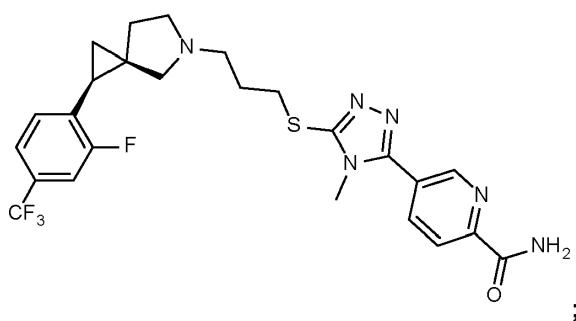
20 (1S,3R eller 1R,3S)-5-(3-{{4-metyl-5-(4-metyl-1,3-oksazol-5-yl)-4H-1,2,4-triazol-3-yl}sulfanyl}propyl)-1-[4-(trifluormetyl)fenyl]-5-azaspido[2.5]oktan (TRANS, Enantiomer 2);
eller et farmasøytisk akseptabelt salt derav.

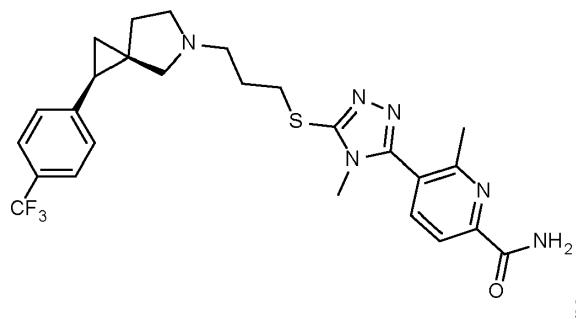
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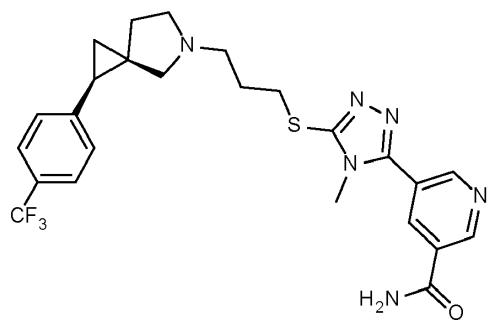


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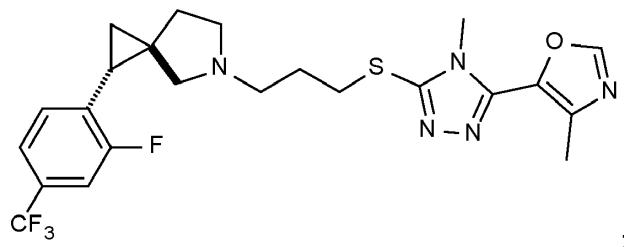
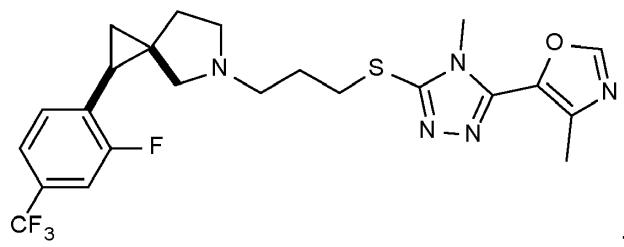
eller



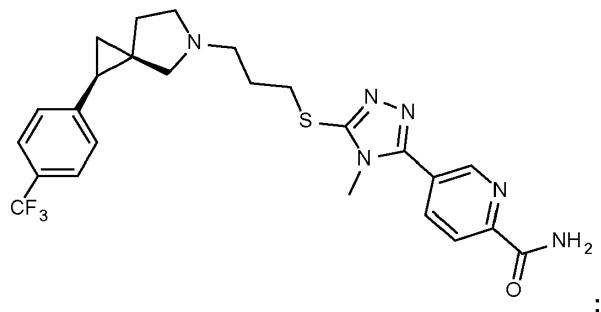
eller et farmasøytisk akseptabelt salt derav.

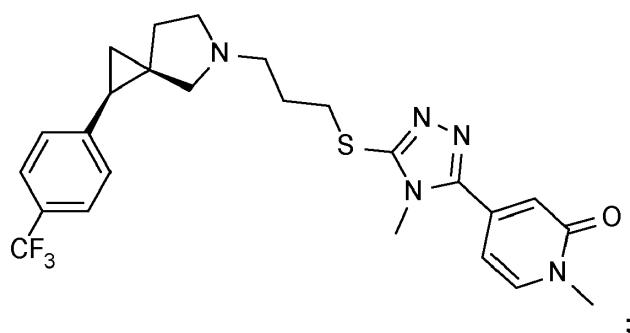
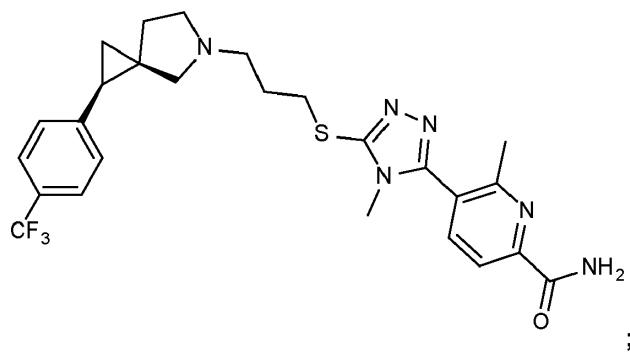
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17. Forbindelse ifølge krav 1 som har formelen:

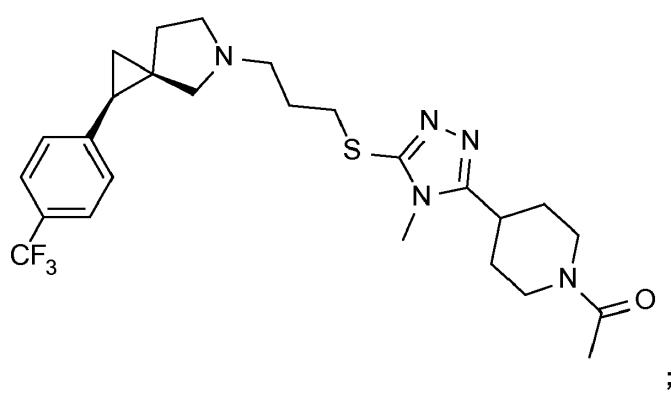
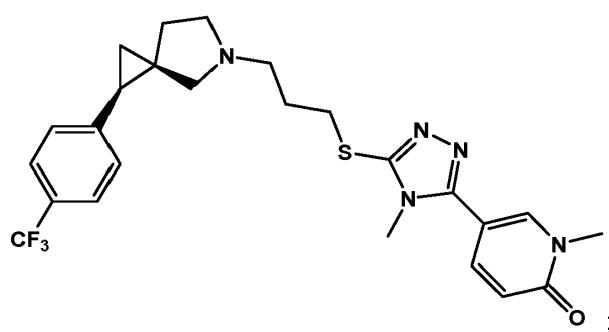


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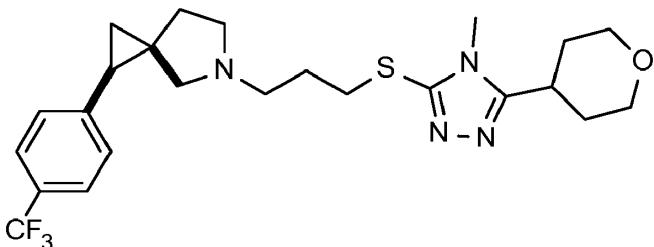




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eller



eller et farmasøytisk akseptabelt salt derav.

18. Farmasøytisk sammensetning omfattende en forbindelse ifølge et hvilket som

5 helst av kravene 1–17 og en farmasøytisk akseptabel bærer.

19. Forbindelse ifølge et hvilket som helst av kravene 1 til 17 for anvendelse som

et medikament.

10 **20.** Forbindelse ifølge et hvilket som helst av kravene 1–17 for anvendelse ved behandling av en tilstand, hvori tilstanden er valgt fra:

(i) stoffmisbruk, eventuelt hvori stoffmisbruket er alkohol-, kokain-, heroin- eller nikotinmisbruk; eller

(ii) en dyskinetisk lidelse, eventuelt hvori den dyskinetiske lidelsen er Parkinsons

15 sykdom, nevroleptisk indusert parkinsonisme eller tardive dyskinesier; eller

(iii) kognitiv svikt, eventuelt hvori den kognitive svikten er en hukommelsesforstyrrelse eller Alzheimers sykdom; eller

(iv) depresjon, angst, en spiseforstyrrelse, seksuell dysfunksjon, for tidlig utløsnings, en søvnforstyrrelse, oppkast, en bevegelsesforstyrrelse, en obsessiv-kompulsiv lidelse,

20 amnesi, aggressjon, autisme, svimmelhet, demens, en døgnrytmeforstyrrelse, eller en gastrisk motilitetsforstyrrelse, eventuelt hvori den gastriske motilitetsforstyrrelsen er IBS; eller

(v) en psykotisk tilstand, eventuelt hvori den psykotiske tilstanden er schizofreni, en schizo-affektiv lidelse, psykotisk depresjon, mani, paranoid forstyrrelse eller en paranoid personlighetsforstyrrelse; eller

25 (vi) medikamentavhengighet, abstinenssymptomer fra legemidler med misbruk eller inhibering av toleranse indusert av opioider, eventuelt hvori legemidlet er alkohol, kokain, et opiat, nikotin eller et benzodiazepin; eller

(vii) ikke-substansrelaterte lidelser og vanedannende lidelser, eventuelt hvori lidelsen er en spillesykdom.