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A61K 31/4725 (2006.01)

A61K 31/502 (2006.01)

Norwegian Industrial Property Office

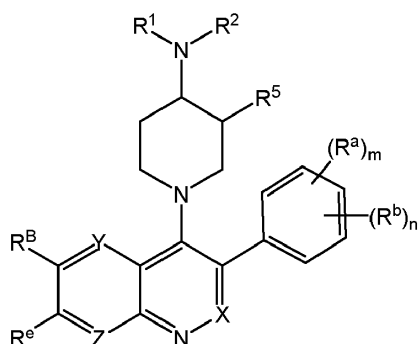
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(54)	Title	SOMATOSTATIN MODULATORS AND USES THEREOF
(56)	References Cited:	KR-A- 20160 062 023 EP-A1- 2 871 179 US-A1- 2015 232 478 US-A1- 2003 153 553 WO-A2-2008/051272 WO-A1-2010/041054 Scott E. Wolkenberg ET AL: "Design, Synthesis, and Evaluation of Novel 3,6-Diaryl-4-aminoalkoxyquinolines as Selective Agonists of Somatostatin Receptor Subtype 2", Journal of Medicinal Chemistry, vol. 54, no. 7, 14 April 2011 (2011-04-14) , pages 2351-2358, XP055569581, US ISSN: 0022-2623, DOI: 10.1021/jm101501b

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 som har følgende struktur, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav:

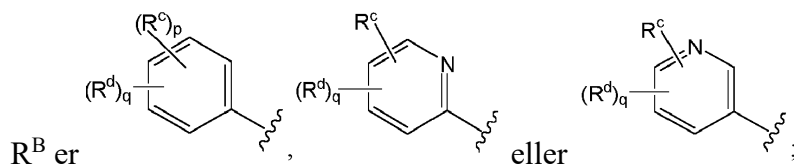


5 hvori:

hver R^a er uavhengig hydrogen, F, Cl, Br, $-\text{CH}_3$, $-\text{CH}_2\text{F}$, $-\text{CHF}_2$, $-\text{CF}_3$, $-\text{CN}$, $-\text{OH}$, $-\text{OCH}_3$, $-\text{OCF}_3$, $-\text{CH}_2\text{OH}$ eller $-\text{CH}_2\text{CH}_2\text{OH}$;

hver R^b er uavhengig hydrogen, F, Cl, Br, $-\text{CH}_3$, $-\text{CH}_2\text{F}$, $-\text{CHF}_2$, $-\text{CF}_3$, $-\text{CN}$, $-\text{OH}$, $-\text{OCH}_3$ eller $-\text{OCF}_3$;

10 m er 1 eller 2; n er 0, 1 eller 2;



hver R^c er uavhengig hydrogen, F, Cl, Br, $-\text{CH}_3$, $-\text{CH}_2\text{CH}_3$, $-\text{CH}_2\text{CH}_2\text{CH}_3$, $-\text{CH}(\text{CH}_3)_2$, $-\text{CH}_2\text{CH}_2\text{CH}_2\text{CH}_3$, $-\text{CH}_2\text{CH}(\text{CH}_3)_2$, $-\text{CH}(\text{CH}_3)(\text{CH}_2\text{CH}_3)$, $-\text{C}(\text{CH}_3)_3$, $-\text{CH}_2\text{OH}$, $-\text{CH}_2\text{CN}$, $-\text{CH}_2\text{F}$, $-\text{CHF}_2$, $-\text{CF}_3$, $-\text{CN}$, $-\text{OH}$, $-\text{OCH}_3$, $-\text{OCH}_2\text{CH}_3$, $-\text{OCF}_3$, $-\text{CO}_2\text{H}$, $-\text{CO}_2\text{CH}_3$, $-\text{CO}_2\text{CH}_2\text{CH}_3$, $-\text{CH}_2\text{CO}_2\text{H}$, $-\text{CH}_2\text{CO}_2\text{CH}_3$, $-\text{CH}_2\text{CO}_2\text{CH}_2\text{CH}_3$, $-\text{C}(=\text{O})\text{NH}_2$, $-\text{C}(=\text{O})\text{NHCH}_3$, $-\text{C}(=\text{O})\text{NHOCH}_3$, $-\text{C}(=\text{O})\text{N}(\text{CH}_3)_2$, $-\text{SO}_2\text{N}(\text{CH}_3)_2$, $-\text{C}(=\text{NOCH}_3)\text{H}$, $-\text{CH}_2\text{C}(=\text{O})\text{NH}_2$, $-\text{CH}_2\text{C}(=\text{O})\text{NHCH}_3$, $-\text{CH}_2\text{C}(=\text{O})\text{N}(\text{CH}_3)_2$, $-\text{NH}_2$, $-\text{NHCH}_3$, $-\text{N}(\text{CH}_3)_2$, $-\text{NHCO}_2\text{CH}_3$, $-\text{NHSO}_2\text{CH}_3$, $-\text{CH}_2\text{NH}_2$, $-\text{CH}_2\text{NHCH}_3$, $-\text{CH}_2\text{N}(\text{CH}_3)_2$, $-\text{CH}(\text{CF}_3)\text{NH}_2$, azetidinyll eller pyrrolidinyll;

20 hver R^d er uavhengig hydrogen, F, Cl, Br, $-\text{CH}_3$, $-\text{CH}_2\text{CH}_3$, $-\text{CH}_2\text{CH}_2\text{CH}_3$, $-\text{CH}(\text{CH}_3)_2$, $-\text{CH}_2\text{CH}_2\text{CH}_2\text{CH}_3$, $-\text{CH}_2\text{CH}(\text{CH}_3)_2$, $-\text{CH}(\text{CH}_3)(\text{CH}_2\text{CH}_3)$, $-\text{C}(\text{CH}_3)_3$, $-\text{CH}_2\text{OH}$, $-\text{CH}_2\text{CN}$, $-\text{CH}_2\text{F}$, $-\text{CHF}_2$, $-\text{CF}_3$, pyrrolyll, imidazolyl, pyrazolyl, triazolyl, tetrazolyl, $-\text{CN}$, $-\text{OH}$, $-\text{OCH}_3$, $-\text{OCH}_2\text{CH}_3$, $-\text{OCF}_3$, $-\text{OCH}_2\text{OCH}_3$, $-\text{OCH}_2\text{OCH}_2\text{CH}_3$, $-\text{OCH}_2\text{CH}_2\text{OH}$, $-\text{C}(=\text{O})\text{NHOCH}_3$, $-\text{C}(=\text{NOH})\text{H}$, $-\text{C}(=\text{NOCH}_3)\text{H}$, $-\text{CH}_2\text{C}(=\text{O})\text{NH}_2$, $-\text{NH}_2$, $-\text{NHCO}_2\text{CH}_3$, $-\text{NHSO}_2\text{CH}_3$, $-\text{NH}(\text{C}=\text{O})\text{NHCH}_3$, $-\text{NH}(\text{C}=\text{O})\text{NHOCH}_3$ eller $-\text{CH}(\text{CF}_3)\text{NH}_2$;

eller dersom en R^c og en R^d er på tilstøtende atomer av R^B , blir de tilstøtende R^c - og R^d -gruppene tatt sammen med de mellomliggende atomene som de er festet til for å danne en 5- eller 6-leddet monosyklisk heterosyklisk ring;

p er 1 eller 2; q er 0, 1 eller 2;

5 X er CR^f eller N;

Y er CR^f eller N;

Z er CR^f eller N;

R^e er hydrogen, F, Cl, Br, $-CH_3$, $-CH_2CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$, $-OCH_2CH_3$ eller $-OCF_3$;

10 hver R^f er uavhengig hydrogen, F, Cl, Br, $-CH_3$, $-CH_2CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$, $-OCH_2CH_3$ eller $-OCF_3$;

R^1 er hydrogen;

R^2 er hydrogen, metyl, etyl, 2-fluoretyl, 2-hydroksyetyl, 2-metoksyetyl, n-propyl, i-propyl, syklopropyl, 3-fluorpropyl, 3-metoksypropyl, n-butyl, i-butyl, sek-butyl, syklobutyl, tert-butyl eller oksetanyl;

15 R^5 er hydrogen, F, Cl, Br, $-OH$, $-OCH_3$, $-NH_2$, $-NHCH_3$, $-N(CH_3)_2$, $-OCH_2CH_3$, $-OCF_3$, $-CH_3$, $-CH_2CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-C(=O)OCH_3$, $-C(=O)NH_2$, $-C(=O)NHCH_3$ eller $-C(=O)N(CH_3)_2$;

20 eller R^2 og R^5 blir tatt sammen med de mellomliggende atomene som de er festet til for å danne et azetidinyll, pyrrolidinyll, piperidinyll, morfolinyll, tiomorfolinyll, piperazinyll eller azepanyll.

2. Forbindelsen ifølge krav 1, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav, hvori:

25 hver R^a er uavhengig hydrogen, F, Cl, Br, $-CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$, $-OCF_3$, $-CH_2OH$ eller $-CH_2CH_2OH$;

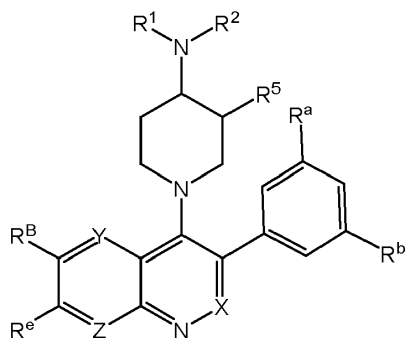
hver R^b er uavhengig hydrogen, F, Cl, Br, $-CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$ eller $-OCF_3$;

30 hver R^c er uavhengig hydrogen, F, Cl, Br, $-CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$, $-OCF_3$, $-NH_2$, $-C(=O)NH_2$, $-C(=NOCH_3)H$, $-SO_2N(CH_3)_2$, azetidinyll eller pyrrolidinyll;

hver R^d er uavhengig hydrogen, F, Cl, Br, $-CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$, $-OCF_3$, $-OCH_2OCH_3$, $-CH_2OH$, $-OCH_2CH_2OH$, $-C(=O)NHOCH_3$, $-NH_2$, $-NHCO_2CH_3$, $-NH(C=O)NHOCH_3$ eller $-CH_2C(=O)NH_2$;

eller hvis en R^c og en R^d er på tilstøtende atomer av R^B , blir de tilstøtende R^c - og R^d -gruppene tatt sammen med de mellomliggende atomene som de er festet til for å danne 5-leddet monosyklisk heterosyklisk ring.

- 5 3. Forbindelsen ifølge krav 1 eller krav 2, hvori forbindelsen har følgende struktur, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav:



- 10 4. Forbindelsen ifølge et hvilket som helst av kravene 1 til 3, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav, hvori:

hver R^a er uavhengig hydrogen, F, Cl, Br, $-CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$, $-OCF_3$, $-CH_2OH$ eller $-CH_2CH_2OH$;

15 hver R^b er uavhengig hydrogen, F, Cl, Br, $-CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$ eller $-OCF_3$;

hver R^c er uavhengig hydrogen, F, Cl, Br, $-CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$, $-OCF_3$, $-NH_2$, $-C(=O)NH_2$, $-C(=NOCH_3)H$, $-SO_2N(CH_3)_2$, azetidinyll eller pyrrolidinyll;

hver R^d er uavhengig hydrogen, F, Cl, Br, $-CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$, $-OCF_3$, $-OCH_2OCH_3$, $-CH_2OH$, $-OCH_2CH_2OH$, $-C(=O)NHOCH_3$, $-NH_2$, $-NHCO_2CH_3$, -

- 20 $NH(C=O)NHOCH_3$ eller $-CH_2C(=O)NH_2$;

R^e er hydrogen, F, Cl, Br, $-CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$ eller $-OCF_3$;

hver R^f er hydrogen;

R^1 er hydrogen;

R^2 er hydrogen;

- 25 R^5 er hydrogen, F, Cl, Br, $-OH$, $-OCH_3$, $-NH_2$, $-NHCH_3$, $-N(CH_3)_2$, $-OCH_2CH_3$, $-OCF_3$, $-CH_3$, $-CH_2CH_3$, $-CH_2F$, $-CHF_2$, $-CF_3$, $-CN$, $-C(=O)OCH_3$, $-C(=O)NH_2$, $-C(=O)NHCH_3$ eller $-C(=O)N(CH_3)_2$;

eller R^2 og R^5 blir tatt sammen med de mellomliggende atomene som de er festet til for å danne et piperidinyll, morfolinyll, tiomorfolinyll eller piperazinyll.

5. Forbindelsen ifølge et hvilket som helst av kravene 1 til 4, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav, hvori:

hver R^a er uavhengig hydrogen, F, Cl, $-\text{CH}_3$, $-\text{CF}_3$, $-\text{CN}$, $-\text{OH}$, $-\text{OCH}_3$, $-\text{OCF}_3$, $-\text{CH}_2\text{OH}$ eller $-\text{CH}_2\text{CH}_2\text{OH}$

hver R^b er uavhengig hydrogen, F, Cl, $-\text{CH}_3$, $-\text{CF}_3$, $-\text{CN}$, $-\text{OH}$, $-\text{OCH}_3$ eller $-\text{OCF}_3$;

hver R^c er uavhengig hydrogen, F, Cl, $-\text{CH}_3$, $-\text{CF}_3$, $-\text{CN}$, $-\text{OH}$, $-\text{OCH}_3$ eller $-\text{OCF}_3$;

hver R^d er uavhengig hydrogen, F, Cl, $-\text{CH}_3$, $-\text{CF}_3$, $-\text{CN}$, $-\text{OH}$, $-\text{OCH}_3$, $-\text{OCF}_3$ eller $-\text{NH}_2$;

X er CR^f ; Y er CR^f ; og Z er CR^f ;

eller X er N; Y er CR^f ; og Z er CR^f ;

eller X er CR^f ; Y er N; og Z er CR^f ;

eller X er CR^f ; Y er CR^f ; og Z er N;

R^e er hydrogen, F eller Cl;

hver R^f er hydrogen;

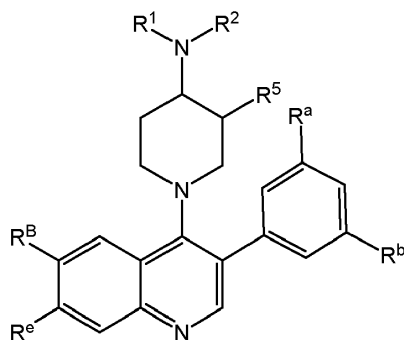
R^1 er hydrogen;

R^2 er hydrogen;

R^5 er hydrogen;

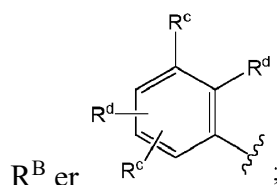
eller R^2 og R^5 blir tatt sammen med de mellomliggende atomene som de er festet til for å danne et morfolinyll.

6. Forbindelsen ifølge et hvilket som helst av kravene 1 til 3, hvori forbindelsen har følgende struktur av formel (AVb), eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav:



Formel (AVb)

hvori:



R^1 er hydrogen;

R^2 er hydrogen, metyl, etyl, 2-fluoretyl, 2-hydroksyetyl, 2-metoksyetyl, n-propyl, i-propyl, syklopropyl, 3-fluorpropyl, 3-metoksypropyl, n-butyl, i-butyl, sek-butyl, syklobutyl, tert-butyl eller oksetanyl;

R^5 er hydrogen, F, Cl, Br, -OH, -OCH₃, -NH₂, -NHCH₃, -N(CH₃)₂, -OCH₂CH₃, -OCF₃, -CH₃, -CH₂CH₃, -CH₂F, -CHF₂, -CF₃, -CN, -C(=O)OCH₃, -C(=O)NH₂, -C(=O)NHCH₃ eller -C(=O)N(CH₃)₂;

eller R^2 og R^5 blir tatt sammen med de mellomliggende atomene som de er festet til for å danne et azetidinyll, pyrrolidinyl, piperidinyl, morfolinyl, tiomorfolinyl, piperazinyl eller azepanyl.

7. Forbindelsen ifølge krav 6, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav, hvori:

R^a er hydrogen, F, Cl, Br, -CH₃, -CH₂F, -CHF₂, -CF₃, -CN, -OH, -OCH₃, -OCF₃, -CH₂OH eller -CH₂CH₂OH;

R^b er hydrogen, F, Cl, Br, -CH₃, -CH₂F, -CHF₂, -CF₃, -CN, -OH, -OCH₃ eller -OCF₃;

hver R^c er uavhengig hydrogen, F, Cl, Br, -CH₃, -CH₂F, -CHF₂, -CF₃, -CN, -OH, -OCH₃, -OCF₃, -NH₂, -C(=O)NH₂, -C(=NOCH₃)H, -SO₂N(CH₃)₂, azetidinyll eller pyrrolidinyl;

hver R^d er uavhengig hydrogen, F, Cl, Br, -CH₃, -CH₂F, -CHF₂, -CF₃, -CN, -OH, -OCH₃, -OCF₃, -OCH₂OCH₃, -CH₂OH, -OCH₂CH₂OH, -C(=O)NHOCH₃, -NH₂, -NHCO₂CH₃, -NH(C=O)NHOCH₃ eller -CH₂C(=O)NH₂;

R^e er hydrogen, F, Cl, Br, -CH₃, -CH₂F, -CHF₂, -CF₃, -CN, -OH, -OCH₃ eller -OCF₃;

R^1 er hydrogen;

R^2 er hydrogen;

R^5 er hydrogen, F, Cl, Br, -OH, -OCH₃, -NH₂, -NHCH₃, -N(CH₃)₂, -OCH₂CH₃, -OCF₃, -CH₃, -CH₂CH₃, -CH₂F, -CHF₂, -CF₃, -CN, -C(=O)OCH₃, -C(=O)NH₂, -C(=O)NHCH₃ eller -C(=O)N(CH₃)₂;

eller R^2 og R^5 blir tatt sammen med de mellomliggende atomene som de er festet til for å danne et piperidinyl, morfolinyl, tiomorfolinyl eller piperazinyl.

8. Forbindelsen ifølge krav 6 eller krav 7, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav, hvori:

R^a er hydrogen, F, Cl, $-CH_3$, $-CF_3$, $-CN$, $-OH$, $-CH_2OH$, $-CH_2CH_2OH$, $-OCH_3$ eller $-OCF_3$;

5 R^b er hydrogen, F, Cl, $-CH_3$, $-CF_3$, $-CN$, $-OH$, $-OCH_3$ eller $-OCF_3$;

hver R^c er uavhengig hydrogen, F, Cl, $-CH_3$, $-CF_3$, $-CN$, $-OH$, $-NH_2$, $-OCH_3$, $-OCF_3$, $-CONH_2$ eller $-C(=NOCH_3)H$;

hver R^d er uavhengig hydrogen, F, Cl, $-CH_3$, $-CF_3$, $-CN$, $-OH$, $-NH_2$, $-OCH_3$ eller $-OCF_3$;

R^1 er hydrogen;

10 R^2 er hydrogen;

R^5 er hydrogen;

eller R^2 og R^5 blir tatt sammen med de mellomliggende atomene som de er festet til for å danne et morfolinyl.

15 9. Forbindelsen ifølge krav 1, hvori forbindelsen er:

1-1: 3-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3,5-dimetylphenyl)cinnolin-6-yl]-5-fluorbenzamid;

1-2: 3-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3,5-dimetylphenyl)cinnolin-6-yl]-2-hydroksybenzonitril;

20 **1-4:** 3-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3-fluor-5-metylphenyl)cinnolin-6-yl]-5-fluorbenzamid;

1-5: 3-{4-[trans-4-amino-3-fluorpiperidin-1-yl]-7-klor-3-(3-fluor-5-metylphenyl)cinnolin-6-yl}-5-fluorbenzamid;

1-6: 3-{4-[cis-4-amino-3-fluorpiperidin-1-yl]-7-klor-3-(3-fluor-5-metylphenyl)cinnolin-6-yl}-5-fluorbenzamid;

1-7: 3-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3-fluor-5-metylphenyl)cinnolin-6-yl]-2-hydroksybenzonitril;

1-8: 5-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3-fluor-5-metylphenyl)cinnolin-6-yl]-2,3-dihydro-1H-1,3-benzodiazol-2-on;

30 **1-9:** 4-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3-fluor-5-metylphenyl)cinnolin-6-yl]-2,3-dihydro-1H-1,3-benzodiazol-2-on;

1-10: 3-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-7-klor-3-(3-fluor-5-metylphenyl)cinnolin-6-yl}-5-fluorbenzamid;

- 1-11:** 3-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3,5-diklorfenyl)cinnolin-6-yl]-2-hydroksybenzonitril;
- 1-12:** 3-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3-klor-5-metylfenyl)cinnolin-6-yl]-5-fluor-2-hydroksybenzonitril;
- 5 **1-13:** 3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)cinnolin-6-yl]-5-fluorbenzamid;
- 1-14:** 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)cinnolin-6-yl]-5-fluorbenzamid;
- 1-16:** 5-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)cinnolin-6-yl]-2,3-dihydro-1H-1,3-benzodiazol-2-on;
- 10 **1-17:** 3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)cinnolin-6-yl]-2-hydroksybenzonitril;
- 1-18:** 3-[4-(4-aminopiperidin-1-yl)-3-(3-klor-5-metylfenyl)cinnolin-6-yl]-2-hydroksybenzonitril;
- 1-19:** 3-[4-(4-aminopiperidin-1-yl)-3-(3-klor-5-fluorfenyl)cinnolin-6-yl]-2-
- 15 hydroksybenzonitril;
- 1-20:** 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)cinnolin-6-yl]-2-hydroksybenzonitril;
- 1-21:** 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-diklorfenyl)cinnolin-6-yl]-2-hydroksybenzonitril;
- 20 **1-22:** 3-[4-(4-aminopiperidin-1-yl)-3-(3-klor-5-metylfenyl)cinnolin-6-yl]-5-fluor-2-hydroksybenzonitril;
- 1-23:** 6-[4-(4-aminopiperidin-1-yl)-3-(3-klor-5-metylfenyl)cinnolin-6-yl]-4-fluor-2,3-dihydro-1H-1,3-benzodiazol-2-on;
- 1-24:** 4-[4-(4-aminopiperidin-1-yl)-3-(3-klor-5-metylfenyl)cinnolin-6-yl]-6-fluor-2,3-
- 25 dihydro-1H-1,3-benzodiazol-2-on;
- 1-25:** 3-[7-klor-3-(3-fluor-5-metylfenyl)-4-{4-[(2-fluoretyl)amino]piperidin-1-yl}cinnolin-6-yl]-5-fluorbenzamid;
- 1-26:** 3-{4-[trans-4-amino-3-hidroksypiperidin-1-yl]-7-klor-3-(3-fluor-5-metylfenyl)cinnolin-6-yl}-5-fluorbenzamid;
- 30 **1-27:** 3-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3,5-diklorfenyl)cinnolin-6-yl]-5-fluor-2-hydroksybenzonitril;
- 1-28:** 6-[4-(4-aminopiperidin-1-yl)-3-(3,5-diklorfenyl)cinnolin-6-yl]-4-fluor-2,3-dihydro-1H-1,3-benzodiazol-2-on;
- 1-29:** 2-[4-(4-aminopiperidin-1-yl)-3-(3-klor-5-metylfenyl)cinnolin-6-yl]-6-fluorfenol;

1-30: 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-diklorfenyl)cinnolin-6-yl]-5-fluorbenzamid;

1-31: 3-[3-(3,5-diklorfenyl)-4-{4-[(oksetan-3-yl)amino]piperidin-1-yl}cinnolin-6-yl]-5-fluorbenzamid;

1-33: 3-[3-(3,5-diklorfenyl)-4-{4-[(2-fluoretyl)amino]piperidin-1-yl}cinnolin-6-yl]-5-fluorbenzamid;

1-35: 3-[7-klor-3-(3-klor-5-metylfenyl)-4-{4-[(2-fluoretyl)amino]piperidin-1-yl}cinnolin-6-yl]-5-fluorbenzamid;

1-36: 3-[7-klor-3-(3-klor-5-metylfenyl)-4-{4-[(2-fluoretyl)amino]piperidin-1-yl}cinnolin-6-yl]-5-fluor-2-hidroksybenzonitril;

1-38: 3-[7-klor-3-(3-klor-5-metylfenyl)-4-{4-[(oksetan-3-yl)amino]piperidin-1-yl}cinnolin-6-yl]-5-fluorbenzamid;

1-39: 3-[7-klor-3-(3-klor-5-metylfenyl)-4-{4-[(oksetan-3-yl)amino]piperidin-1-yl}cinnolin-6-yl]-5-fluor-2-hidroksybenzonitril;

1-40: 3-[3-(3-klor-5-metylfenyl)-4-{4-[(oksetan-3-yl)amino]piperidin-1-yl}cinnolin-6-yl]-5-fluor-2-hidroksybenzonitril;

1-41: 3-{4-[trans-4-amino-3-fluoropiperidin-1-yl]-7-klor-3-(3-klor-5-metylfenyl)cinnolin-6-yl}-5-fluorbenzamid;

1-42: 6-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3-klor-5-metylfenyl)cinnolin-6-yl]pyridin-2-karboksamid;

1-43: 6-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3-klor-5-metylfenyl)cinnolin-6-yl]-5-klorpyridin-2-karboksamid;

1-44: 2-[4-(4-aminopiperidin-1-yl)-7-klor-3-(3-klor-5-metylfenyl)cinnolin-6-yl]pyridin-4-karboksamid;

1-45: 2-[4-(4-aminopiperidin-1-yl)-3-(3-klor-5-metylfenyl)cinnolin-6-yl]-6-fluor-3-metylfenol;

1-48: 3-{4-[cis-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-klor-5-metylfenyl)cinnolin-6-yl}-5-fluorbenzamid;

1-49: 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-klor-5-metylfenyl)cinnolin-6-yl}-5-fluorbenzamid;

1-51: 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-diklorfenyl)cinnolin-6-yl}-2-hidroksybenzonitril;

1-52: 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-klor-5-metylfenyl)cinnolin-6-yl}-2-hidroksybenzonitril;

1-53: 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metylfenyl)cinnolin-6-yl}-2-hydroksybenzonitril;

1-54: 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)cinnolin-6-yl}-2-hydroksybenzonitril;

5 **1-55:** 5-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)cinnolin-6-yl}-4-aminopyridin-3-karbonitril;

1-56: 5-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metylfenyl)cinnolin-6-yl}-4-aminopyridin-3-karbonitril;

10 **1-57:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)cinnolin-6-yl}-3-aminopyridin-4-karbonitril;

1-58: 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metylfenyl)cinnolin-6-yl}-3-aminopyridin-4-karbonitril;

eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav.

15

10. Forbindelsen ifølge krav 1, hvori forbindelsen er:

2-1: 3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-2-hydroksybenzonitril;

20 **2-2:** 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-2-hydroksybenzonitril;

2-3: 3-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-hydroksybenzonitril;

2-4: 3-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}benzonitril;

25 **2-5:** 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]benzonitril;

2-6: 2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]fenol;

2-7: 6-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]pyridin-2-karboksamid;

30 **2-8:** 4-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-2,3-dihydro-1H-1,3-benzodiazol-2-on;

2-9: 5-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-2,3-dihydro-1H-1,3-benzodiazol-2-on;

2-10: 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-5-fluor-2-hydroksybenzonitril;

2-16: 3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-fluor-2-hydroksybenzonitril;

2-19: 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-hydroksybenzonitril;

5 **2-20:** 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-2-hydroksybenzonitril;

2-58: 3-{4-[cis-4-amino-3-hidroksypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-(metoksymetoksy)benzonitril;

10 **2-59:** 3-{4-[(3R,4R)-4-amino-3-fluorpiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-hydroksybenzonitril;

2-60: 3-{4-[cis-4-amino-3-hidroksypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-hydroksybenzonitril;

2-61: 3-{4-[(3S,4R)-4-amino-3-metoksyypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-hydroksybenzonitril;

15 **2-62:** 3-{4-[(3R,4S)-4-amino-3-metoksyypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-(metoksymetoksy)benzonitril;

2-63: 3-{4-[cis-4-amino-3-hidroksypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-5-fluor-2-hydroksybenzonitril;

20 **2-65:** 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-2-(2-hidroksyethoksy)benzonitril;

2-66: 3-{4-[(3R,4S)-4-amino-3-metoksyypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-hydroksybenzonitril;

2-75: 5-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-6-(azetidin-1-yl)pyridin-3-karbonitril;

25 **2-76:** 5-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-6-(azetidin-1-yl)pyridin-3-karboksamid;

2-77: 2-amino-3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]benzonitril;

30 **2-78:** 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-2-(metoksymetoksy)benzonitril;

2-80: 6-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-fluor-2,3-dihydro-1H-1,3-benzodiazol-2-on;

2-81: 2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-6-[(metoksyimino)metyl]fenol;

2-83: metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4,6-difluorfenyl}karbamat;

2-84: 3-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-(metoksymetoksy)benzonitril;

5 **2-85:** 6-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}pyridin-2-karboksamid;

2-86: N-{2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4,6-difluorfenyl}metansulfonamid;

10 **2-92:** trans-4-amino-1-[6-(3-cyano-2-hidroksyfenyl)-3-(3,5-difluorfenyl)kinolin-4-yl]piperidin-3-karbonitril;

2-94: cis-4-amino-1-[6-(3-cyano-2-hidroksyfenyl)-3-(3,5-difluorfenyl)kinolin-4-yl]piperidin-3-karbonitril;

2-99: 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4,5-difluor-2-hidroksybenzonitril;

15 **2-100:** 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4,5-difluor-2-hidroksybenzonitril;

2-102: 3-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4,5-difluor-2-hidroksybenzonitril;

20 **2-103:** 3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-4,5-difluor-2-hidroksybenzonitril;

2-104: 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-4,5-difluor-2-hidroksybenzonitril;

2-106: 3-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-4,5-difluor-2-hidroksybenzonitril;

25 **2-107:** 3-{4-[cis-4-amino-3-hidroksypiperidin-1-yl]-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl}-4,5-difluor-2-hidroksybenzonitril;

2-108: 2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4,6-difluor-3-metylfenol;

30 **2-109:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4,6-difluor-3-metylfenol;

2-111: 2-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4,6-difluor-3-metylfenol;

2-112: 2-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-4,6-difluor-3-metylfenol;

2-113: cis-4-amino-1-[6-(3,5-difluor-2-hidroksy-6-metylfenyl)-3-(3-fluor-5-metoksyfenyl)kinolin-4-yl]piperidin-3-ol;

2-114: 2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4-fluor-6-[(metoksyimino)metyl]fenol;

5 **2-115:** 2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-6-[(metoksyimino)metyl]fenol;

2-116: 2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-4-fluor-6-[(metoksyimino)metyl]fenol;

10 **2-117:** 2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-3,4-difluor-6-[(metoksyimino)metyl]fenol;

2-118: 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-4-fluor-6-[(1E)-(metoksyimino)metyl]fenol;

2-119: trans-4-amino-1-[6-(3-cyano-5,6-difluor-2-hidroksyfenyl)-3-(3,5-difluorfenyl)kinolin-4-yl]piperidin-3-karbonitril;

15 **2-120:** cis-4-amino-1-[6-(3-cyano-5,6-difluor-2-hidroksyfenyl)-3-(3,5-difluorfenyl)kinolin-4-yl]piperidin-3-karbonitril;

2-121: trans-4-amino-1-[6-(3-cyano-5,6-difluor-2-hidroksyfenyl)-3-(3-fluor-5-metylfenyl)kinolin-4-yl]piperidin-3-karbonitril;

20 **2-122:** cis-4-amino-1-[6-(3-cyano-5,6-difluor-2-hidroksyfenyl)-3-(3-fluor-5-metylfenyl)kinolin-4-yl]piperidin-3-karbonitril;

2-123: metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-3,4,6-trifluorfenyl}karbamat;

2-124: metyl-N-{3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-5-fluorpyridin-4-yl}karbamat;

25 **2-125:** metyl-N-{3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-5-cyanopyridin-4-yl}karbamat;

2-126: 1-[3-(3,5-difluorfenyl)-6-{3-fluor-2-[(hidroksyimino)metyl]fenyl}kinolin-4-yl]piperidin-4-amin;

30 **2-127:** 1-(6-{3-klor-2-[(hidroksyimino)metyl]fenyl}-3-(3,5-difluorfenyl)kinolin-4-yl)piperidin-4-amin;

2-128: 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-2-[(hidroksyimino)metyl]benzonitril;

2-129: 1-[3-(3,5-difluorfenyl)-6-{3-fluor-2-[(metoksyimino)metyl]fenyl}kinolin-4-yl]piperidin-4-amin;

2-130: 1-(6-{3-klor-2-[(metoksyimino)metyl]fenyl}-3-(3,5-difluorfenyl)kinolin-4-yl)piperidin-4-amin;

2-131: 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-2-[(1E)-(metoksyimino)metyl]benzonitril;

5 **2-132:** 1-{6-[3-(1-amino-2,2,2-trifluoretyl)fenyl]-3-(3,5-difluorfenyl)kinolin-4-yl}piperidin-4-amin;

2-133: 1-{6-[2-(1-amino-2,2,2-trifluoretyl)-3-fluorfenyl]-3-(3,5-difluorfenyl)kinolin-4-yl}piperidin-4-amin;

10 **2-134:** 2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-6-klor-3,4-difluorfenol;

2-135: 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-6-klor-3,4-difluorfenol;

2-136: 2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-6-klor-4-fluorfenol;

15 **2-137:** 2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-6-klorfenol;

2-138: 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-6-klorfenol;

2-139: 1-{2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4,6-difluorfenyl}-3-metoksyurea;

20 **2-140:** 1-(2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4,6-difluorfenyl)-3-metoksyurea;

2-141: 6-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4-klor-2,3-dihydro-1H-1,3-benzodiazol-2-on;

25 **2-142:** 5-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-2,3-dihydro-1,3-benzoksazol-2-on;

2-143: metyl-trans-4-amino-1-[6-(3-cyano-2-hidroksyfenyl)-3-(3,5-difluorfenyl)kinolin-4-yl]piperidin-3-karboksylat;

2-144: 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-6-fluorfenol;

30 **2-146:** trans-4-amino-1-[6-(3-cyano-2-hidroksyfenyl)-3-(3,5-difluorfenyl)kinolin-4-yl]piperidin-3-karboksamid;

2-147: 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-5-klor-4-fluor-2-hidroksybenzonitril;

2-148: metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-6-cyanofenyl}karbamat;

2-149: trans-4-amino-1-[6-(3-cyano-2-hydroksyfenyl)-3-(3,5-difluorfenyl)kinolin-4-yl]-N-metylpiperidin-3-karboksamid;

5 **2-150:** 2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-3,4,6-trifluorfenol;

2-151: trans-4-amino-1-[6-(3-cyano-2-hydroksyfenyl)-3-(3,5-difluorfenyl)kinolin-4-yl]-N,N-dimetylpiperidin-3-karboksamid;

10 **2-152:** 1-[3-(3,5-difluorfenyl)-6-{5-fluor-4-[(hydroksyimino)metyl]pyridin-3-yl}kinolin-4-yl]piperidin-4-amin;

2-153: N-[(2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-6-fluorfenyl)metyliden]hydroksylamin;

2-154: N-[(2-{4-[(trans)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4,6-difluorfenyl)metyliden]hydroksylamin;

15 **2-155:** N-[(2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl}-6-fluorfenyl)metyliden]hydroksylamin;

2-156: 4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-[3-fluor-2-(1H-imidazol-2-yl)fenyl]-3-(3-fluor-5-metoksyfenyl)kinolin;

20 **2-157:** 4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-[3-fluor-2-(1H-imidazol-5-yl)fenyl]-3-(3-fluor-5-metoksyfenyl)kinolin;

2-158: 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl}-4,5-difluor-2-hydroksybenzonitril;

2-159: 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl}-4,5-difluor-2-hydroksybenzonitril;

25 **2-160:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl}-3,4,6-trifluorfenol;

2-161: 1-[3-(3-fluor-5-metoksyfenyl)-6-(2,3,5-trifluor-6-hydroksyfenyl)kinolin-4-yl]piperidin-4-ol;

30 **2-162:** 1-{3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-5-hydroksypyridin-4-yl}-3-metoksyurea;

2-163: 1-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl]-6-klorfenyl}-3-metoksyurea;

2-164: 1-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl]-4,6-difluorfenyl}-3-metoksyurea;

2-165: metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl]-4,6-difluorfenyl} karbamat;

2-166: 1-(6-{3-fluor-2-[(hydroksyimino)metyl]fenyl}-3-(3-fluor-5-metoksyfenyl)kinolin-4-yl)piperidin-4-amin;

5 **2-167:** metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl]-6-cyanofenyl} karbamat;

2-168: 1-(6-{5-fluor-4-[(hydroksyimino)metyl]pyridin-3-yl}-3-(3-fluor-5-metylfenyl)kinolin-4-yl)piperidin-4-amin;

10 **2-169:** 3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-fluorpyridin-4-amin;

2-170: metyl-(trans)-4-amino-1-[6-(3-cyano-2-hidroksyfenyl)-3-(3,5-difluorfenyl)kinolin-4-yl]piperidin-3-karboksylat;

2-171: metyl-N-{3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-fluorpyridin-4-yl} karbamat;

15 **2-172:** 1-{3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-fluorpyridin-4-yl}-3-metoksyurea;

2-173: metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl]-6-fluorfenyl} karbamat;

20 **2-174:** 1-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metoksyfenyl)kinolin-6-yl]-6-fluorfenyl}-3-metoksyurea;

2-176: 1-(6-{5-fluor-4-[(hydroksyimino)metyl]pyridin-3-yl}-3-[3-fluor-5-(trifluormetyl)fenyl]kinolin-4-yl)piperidin-4-amin;

2-177: 4-amino-5-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]pyridin-3-karbonitril;

25 **2-178:** 3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-fluorpyridin-2-amin;

2-179: 2-amino-3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4,5-difluorbenzonitril;

30 **2-180:** metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-6-cyano-3,4-difluorfenyl} karbamat;

2-181: 3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-klorpyridin-4-amin;

2-182: metyl-N-{3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-fluorpyridin-2-yl} karbamat;

2-183: 1-{3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-fluorpyridin-2-yl}-3-metoksyurea;

2-184: metyl-N-{3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-klorpyridin-4-yl}karbamat;

5 **2-185:** metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4-cyanofenyl}karbamat;

2-186: 1-{3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-klorpyridin-4-yl}-3-metoksyurea;

10 **2-187:** 4-amino-5-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]pyridin-3-karbonitril;

2-189: 1-{2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4-cyanofenyl}-3-metoksyurea;

2-190: 3-{4-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-hidroksybenzonitril;

15 **2-192:** 1-(6-{5-klor-4-[(hidroksyimino)metyl]pyridin-3-yl}-3-(3,5-difluorfenyl)kinolin-4-yl)piperidin-4-amin;

2-193: 3-{4-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-2-hidroksybenzonitril;

20 **2-194:** 3-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-5-fluor-2-(metoksymetoksy)benzonitril;

2-195: 1-[3-(3,5-difluorfenyl)-6-{3-[(hidroksyimino)metyl]pyridin-2-yl}kinolin-4-yl]piperidin-4-amin;

2-196: 1-[3-(3,5-difluorfenyl)-6-{2-[(hidroksyimino)metyl]pyridin-3-yl}kinolin-4-yl]piperidin-4-amin;

25 **2-197:** metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-4,6-difluorfenyl}karbamat;

2-200: 1-[3-(3,5-difluorfenyl)-6-{6-[(hidroksyimino)metyl]pyridin-2-yl}kinolin-4-yl]piperidin-4-amin;

30 **2-202:** 1-[3-(3,5-difluorfenyl)-6-{3-fluor-2-[(hidroksyimino)metyl]-6-metylfenyl}kinolin-4-yl]piperidin-4-amin;

2-205: metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-6-cyano-3,4-difluorfenyl}karbamat;

2-207: 1-(2-{4-[cis-4-amino-3-metoksypiperidin-1-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-4,6-difluorfenyl)-3-metoksyurea;

2-208: 1-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-6-cyano-3,4-difluorfenyl}-3-metylurea;

2-209: metyl-N-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-6-fluorfenyl}karbamat;

5 **2-213:** 4-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-2-[(metoksyimino)metyl]fenol;

2-215: 1-{3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-4-metylpiperidin-2-yl}-3-metoksyurea;

10 **2-217:** trans-1-(6-{3-fluor-2-[(hydroksyimino)metyl]fenyl}-3-(3-fluor-5-metylfenyl)kinolin-4-yl)-3-metoksypiperidin-4-amin;

2-218: metyl-N-(2-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-6-cyanofenyl)karbamat;

2-219: cis-3-{4-[4-amino-3-hydroksypiperidin-1-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-5-fluor-2-hydroksybenzonitril;

15 **2-221:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-6-[(metoksyimino)metyl]fenol;

2-222: 1-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-4-cyanofenyl}-3-metoksyurea;

20 **2-223:** 1-{3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-5-fluorfenyl}-3-metoksyurea;

2-226: 1-{3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]pyridin-2-yl}-3-metoksyurea;

2-227: cis-4-amino-1-(6-{3-fluor-2-[(hydroksyimino)metyl]fenyl}-3-(3-fluor-5-metylfenyl)kinolin-4-yl)piperidin-3-ol;

25 **2-235:** metyl-N-(2-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-4,6-difluorfenyl)karbamat;

2-236: 1-(2-{4-[trans-4-amino-3-metoksypiperidin-1-yl]-3-(3-fluor-5-metylfenyl)kinolin-6-yl}-4,6-difluorfenyl)-3-metoksyurea;

30 **2-237:** 1-{2-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-4-cyano-6-fluorfenyl}-3-metoksyurea;

2-239: 1-{3-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]-6-metylpiperidin-2-yl}-3-metoksyurea;

2-240: 1-{4-[4-(4-aminopiperidin-1-yl)-3-(3-fluor-5-metylfenyl)kinolin-6-yl]fenyl}-3-metoksyurea;

- 2-241:** 2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-6-brom-3,4-difluorfenol;
- 2-243:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-6-[(1E)-(metoksyimino)metyl]fenol;
- 5 **2-244:** 5-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-aminopyridin-3-karbonitril;
- 2-246:** 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}pyridin-2-amin;
- 2-247:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-metylpyridin-3-amin;
- 10 **2-248:** 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}pyridin-2-ol;
- 2-250:** 2-[4-(4-aminopiperidin-1-yl)-3-(3,5-difluorfenyl)kinolin-6-yl]-4-metylpyridin-3-amin;
- 15 **2-251:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-3-hidroksypyridin-4-karbonitril;
- 2-254:** 5-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-hidroksypyridin-3-karbonitril;
- 2-256:** 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-5-fluorpyridin-4-ol;
- 20 **2-258:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-metylpyridin-3-ol;
- 2-259:** 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-5-fluorpyridin-2-ol;
- 25 **2-261:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-3-fluorpyridin-4-karbonitril;
- 2-262:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-3-aminopyridin-4-karbonitril;
- 2-263:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}pyridin-3-ol;
- 30 **2-264:** N-[(3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}pyridin-2-yl)metyliden]hidroksylamin;
- 2-265:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-5-fluorpyridin-3-amin;

- 2-266:** 3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-5-[(metoksyimino)metyl]pyridin-4-amin;
- 2-269:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-[(metoksyimino)metyl]pyridin-3-amin;
- 5 **2-273:** 1-(3-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}pyridin-2-yl)-3-metoksyurea;
- 2-275:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-klorpyridin-3-amin;
- 2-283:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}pyridin-3-amin;
- 10 **2-284:** 2-{4-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-metoksypyridin-3-amin;
- 2-285:** 2-{4-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-6-[(metoksyimino)metyl]fenol;
- 15 **2-286:** 2-{4-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-6-[(metoksyimino)metyl]fenol;
- 2-287:** 2-{4-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-[(metoksyimino)metyl]pyridin-3-amin;
- 2-288:** 2-{4-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-[(metoksyimino)metyl]pyridin-3-amin;
- 20 **2-289:** 2-{4-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-3-aminopyridin-4-karbonitril;
- 2-290:** 2-{4-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-3-aminopyridin-4-karbonitril;
- 25 **2-291:** (5-{4-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-aminopyridin-3-karbonitril;
- 2-292:** (5-{4-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-aminopyridin-3-karbonitril;
- 2-293:** 3-{4-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}pyridin-2-amin;
- 30 **2-294:** 3-{4-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}pyridin-2-amin;
- 2-295:** 2-{4-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-metylpyridin-3-amin;

2-296: 2-{4-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-metylpyridin-3-amin;

2-297: 2-{4-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-metylpyridin-3-ol;

5 **2-298:** 2-{4-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-4-metylpyridin-3-ol;

2-299: 3-{4-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-5-fluor-2-hydroksybenzonitril;

10 **2-300:** 3-{4-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-3-(3,5-difluorfenyl)kinolin-6-yl}-5-fluor-2-hydroksybenzonitril;

eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav.

11. Forbindelsen ifølge krav 1, hvori forbindelsen er:

15 **3-1:** 3-[8-(4-aminopiperidin-1-yl)-7-(3-fluor-5-metylfenyl)-1,5-naftyridin-2-yl]-5-fluorbenzamid;

3-2: 3-[8-(4-aminopiperidin-1-yl)-7-(3-fluor-5-metylfenyl)-1,5-naftyridin-2-yl]-5-fluor-2-hydroksybenzonitril;

20 **3-3:** 4-[8-(4-aminopiperidin-1-yl)-7-(3-fluor-5-metylfenyl)-1,5-naftyridin-2-yl]-6-fluor-2,3-dihydro-1H-1,3-benzodiazol-2-on;

3-4: 5-[8-(4-aminopiperidin-1-yl)-7-(3-fluor-5-metylfenyl)-1,5-naftyridin-2-yl]-2,3-dihydro-1H-1,3-benzodiazol-2-on;

3-5: 6-[8-(4-aminopiperidin-1-yl)-7-(3-fluor-5-metylfenyl)-1,5-naftyridin-2-yl]-4-fluor-2,3-dihydro-1H-1,3-benzodiazol-2-on;

25 **3-6:** 3-[8-(4-aminopiperidin-1-yl)-7-(3-klor-5-metylfenyl)-1,5-naftyridin-2-yl]-5-fluor-2-hydroksybenzonitril;

3-7: 3-[8-(4-aminopiperidin-1-yl)-7-(3-fluor-5-metylfenyl)-1,5-naftyridin-2-yl]-2-hydroksybenzonitril;

30 **3-8:** 2-[8-(4-aminopiperidin-1-yl)-7-(3-fluor-5-metylfenyl)-1,5-naftyridin-2-yl]-6-fluorfenol;

3-9: 6-[8-(4-aminopiperidin-1-yl)-7-(3,5-difluorfenyl)-1,5-naftyridin-2-yl]-4-fluor-2,3-dihydro-1H-1,3-benzodiazol-2-on;

3-10: 6-{8-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-7-(3,5-difluorfenyl)-1,5-naftyridin-2-yl}-4-fluor-2,3-dihydro-1H-1,3-benzodiazol-2-on;

3-11: 4-{8-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-7-(3-fluor-5-metylfenyl)-1,5-naftyridin-2-yl}-6-fluor-2,3-dihydro-1H-1,3-benzodiazol-2-on;

3-12: 4-{8-[trans-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-7-(3,5-difluorfenyl)-1,5-naftyridin-2-yl}-1H,2H,3H-imidazo[4,5-c]pyridin-2-on;

5 eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav.

12. Forbindelsen ifølge krav 1, hvori forbindelsen er:

4-1: 3-[5-(4-aminopiperidin-1-yl)-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl]-2-
10 hydroksybenzonitril;

4-2: 3-[5-(4-aminopiperidin-1-yl)-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl]-2-hydroksybenzonitril;

4-3: 3-[5-(4-aminopiperidin-1-yl)-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl]-5-fluor-2-hydroksybenzonitril;

15 **4-4:** 3-[5-(4-aminopiperidin-1-yl)-6-(3-klor-5-metylfenyl)-1,8-naftyridin-3-yl]-2-hydroksybenzonitril;

4-5: 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl}-2-hydroksybenzonitril;

20 **4-6:** 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl}-2-hydroksybenzonitril;

4-7: 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl}-5-fluor-2-hydroksybenzonitril;

4-8: 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl}-5-fluor-2-hydroksybenzonitril;

25 **4-9:** 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl}-5-fluor-2-hydroksybenzonitril;

4-10: 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl}-5-fluor-2-hydroksybenzonitril;

30 **4-11:** 2-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl}-3-aminopyridin-4-karbonitril;

4-12: 2-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl}-3-aminopyridin-4-karbonitril;

4-13: 5-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl}-4-aminopyridin-3-karbonitril;

4-14: 5-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b]morfolin-6-yl]-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl}-4-aminopyridin-3-karbonitril;

4-15: 2-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl}-3-aminopyridin-4-karbonitril;

5 **4-16:** 2-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl}-3-aminopyridin-4-karbonitril;

4-17: 2-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-fluorfenyl)-1,8-naftyridin-3-yl}-3-aminopyridin-4-karbonitril;

10 **4-18:** 2-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-fluorfenyl)-1,8-naftyridin-3-yl}-3-aminopyridin-4-karbonitril;

4-19: 2-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-metylfenyl)-1,8-naftyridin-3-yl}-3-aminopyridin-4-karbonitril;

4-20: 2-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-metylfenyl)-1,8-naftyridin-3-yl}-3-aminopyridin-4-karbonitril;

15 **4-21:** 2-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl}-4-[(metoksyimino)metyl]pyridin-3-amin;

4-22: 2-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl}-4-[(metoksyimino)metyl]pyridin-3-amin;

20 **4-23:** 2-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-metylfenyl)-1,8-naftyridin-3-yl}-4-[(metoksyimino)metyl]pyridin-3-amin;

4-24: 2-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-metylfenyl)-1,8-naftyridin-3-yl}-4-[(metoksyimino)metyl]pyridin-3-amin;

4-25: 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl}-2-hidroksybenzonitril;

25 **4-26:** 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl}-2-hidroksybenzonitril;

4-27: 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-fluorfenyl)-1,8-naftyridin-3-yl}-2-hidroksybenzonitril;

30 **4-28:** 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-fluorfenyl)-1,8-naftyridin-3-yl}-2-hidroksybenzonitril;

4-29: 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-fluor-5-metoksyfenyl)-1,8-naftyridin-3-yl}-2-hidroksybenzonitril;

4-30: 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-fluor-5-metoksyfenyl)-1,8-naftyridin-3-yl}-2-hidroksybenzonitril;

4-31: 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-metylfenyl)-1,8-naftyridin-3-yl}-2-hydroksybenzonitril;

4-32: 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-metylfenyl)-1,8-naftyridin-3-yl}-2-hydroksybenzonitril;

5 **4-33:** 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3,5-diklorfenyl)-1,8-naftyridin-3-yl}-2-hydroksybenzonitril;

4-34: 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3,5-diklorfenyl)-1,8-naftyridin-3-yl}-2-hydroksybenzonitril;

10 **4-35:** 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

4-36: 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3,5-difluorfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

4-37: 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

15 **4-38:** 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-fluor-5-metylfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

4-39: 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-fluor-5-metoksyfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

20 **4-40:** 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-fluor-5-metoksyfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

4-41: 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-fluorfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

4-42: 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-fluorfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

25 **4-43:** 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-metylfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

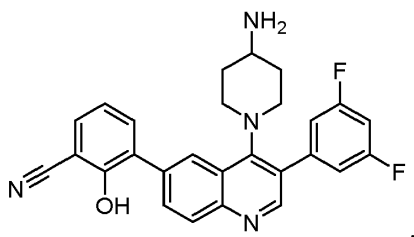
4-44: 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3-klor-5-metylfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

30 **4-45:** 3-{5-[(4 α R,8 α R)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3,5-diklorfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

4-46: 3-{5-[(4 α S,8 α S)-oktahydro-1H-pyrido[3,4-b][1,4]oksazin-6-yl]-6-(3,5-diklorfenyl)-1,8-naftyridin-3-yl}pyridin-2-amin;

eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav.

- 13.** Forbindelsen ifølge krav 1, hvori forbindelsen har følgende struktur:



eller et farmasøytisk akseptabelt salt eller solvat derav.

5

- 14.** Farmasøytisk sammensetning omfattende en forbindelse ifølge et hvilket som helst av kravene 1 til 12, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav, og minst én farmasøytisk akseptabel eksipiens.

- 10 **15.** Farmasøytisk sammensetning omfattende en forbindelse ifølge krav 13, eller et farmasøytisk akseptabelt salt, eller solvat derav, og minst én farmasøytisk akseptabel eksipiens.

- 16.** Den farmasøytiske sammensetningen ifølge krav 14 eller krav 15, hvori den farmasøytiske sammensetningen er i form av en tablett, en pille eller en kapsel.

15

- 17.** Forbindelse ifølge et hvilket som helst av kravene 1 til 12, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav, for anvendelse som en medisin.

- 20 **18.** Forbindelse ifølge krav 13, eller et farmasøytisk akseptabelt salt, eller solvat derav, for anvendelse som en medisin.

- 19.** Forbindelse ifølge et hvilket som helst av kravene 1 til 12, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav, for anvendelse i behandlingen av akromegali, en nevroendokrin svulst, en oftalmisk sykdom eller tilstand, nevropati, nefropati, en respiratorisk sykdom eller tilstand, kreft, smerte, en nevrodegenerativ sykdom eller tilstand, en inflammatorisk sykdom eller tilstand, en psykiatrisk sykdom eller tilstand, eller kombinasjoner derav.

25

20. Forbindelse ifølge krav 13, eller et farmasøytisk akseptabelt salt, eller solvat derav, for anvendelse i behandlingen av akromegali, en nevroendokrin svulst, en oftalmisk sykdom eller tilstand, nevropati, nefropati, en respiratorisk sykdom eller tilstand, kreft, smerte, en nevrodegenerativ sykdom eller tilstand, en inflammatorisk sykdom eller tilstand, en psykiatrisk sykdom eller tilstand, eller kombinasjoner derav.
21. Farmasøytisk sammensetning ifølge et hvilket som helst av kravene 14 til 16 for anvendelse i behandlingen av akromegali, en nevroendokrin svulst, en oftalmisk sykdom eller tilstand, nevropati, nefropati, en respiratorisk sykdom eller tilstand, kreft, smerte, en nevrodegenerativ sykdom eller tilstand, en inflammatorisk sykdom eller tilstand, en psykiatrisk sykdom eller tilstand, eller kombinasjoner derav.
22. Forbindelse ifølge et hvilket som helst av kravene 1 til 12, eller et farmasøytisk akseptabelt salt, solvat, diastereomerblanding eller individuelle enantiomerer derav, for anvendelse i behandlingen av akromegali, en nevroendokrin svulst eller kombinasjoner derav.
23. Forbindelse ifølge krav 13, eller et farmasøytisk akseptabelt salt, eller solvat derav, for anvendelse i behandlingen av akromegali, en nevroendokrin svulst eller kombinasjoner derav.
24. Farmasøytisk sammensetning ifølge et hvilket som helst av kravene 14 til 16 for anvendelse i behandlingen av akromegali, en nevroendokrin svulst eller kombinasjoner derav.