(12) **Translation of European patent specification**

(11) **NO/EP 3468966 B1**

NORWAY

(19) NO

(51) Int Cl.

C07D 405/14 (2006.01)
A61K 31/407 (2006.01)
A61P 3/10 (2006.01)
A61P 35/00 (2006.01)
C07D 403/04 (2006.01)
C07D 403/14 (2006.01)
C07D 471/10 (2006.01)
C07D 487/10 (2006.01)
C07D 491/10 (2006.01)

Norwegian Industrial Property Office

(45)	Translation Published	2021.01.18
(80)	Date of The European Patent Office Publication of the Granted Patent	2020.08.05
(86)	European Application Nr.	17739383.2
(86)	European Filing Date	2017.06.08
(87)	The European Application's Publication Date	2019.04.17
(30)	Priority	2016.06.10, US, 201662348496 P
(84)	Designated Contracting States:	AL ; AT ; BE ; BG ; CH ; CY ; CZ ; DE ; DK ; EE ; ES ; FI ; FR ; GB ; GR ; HR ; HU ; IE ; IS ; IT ; LI ; LT ; LU ; LV ; MC ; MK ; MT ; NL ; NO ; PL ; PT ; RO ; RS ; SE ; SI ; SK ; SM ; TR
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(54) Title **INHIBITORS OF THE MENIN-MLL INTERACTION**

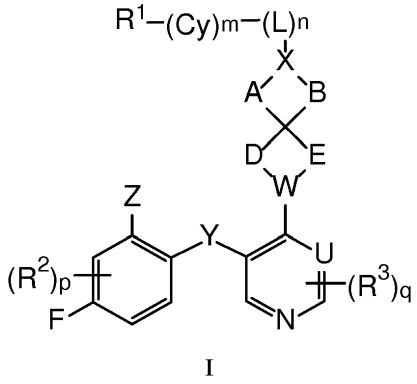
(56) References Cited:

WO-A1-2014/164543
WO-A1-2015/191701
A. SHI ET AL: "Structural insights into inhibition of the bivalent menin-MLL interaction by small molecules in leukemia", BLOOD, vol. 120, no. 23, 29 November 2012 (2012-11-29), pages 4461-4469, XP055245187, US ISSN: 0006-4971, DOI: 10.1182/blood-2012-05-429274

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. En forbindelse med formel I:



I

eller et farmasøytisk akseptabelt salt derav, hvor:

A, B, D og E er hver uavhengig valgt fra -C(R^{A1})(R^{A2})-, -C(R^{A1})(R^{A2})-C(R^{A1})(R^{A2})-, -C(R^{A1})(R^{A2})-O-, -C(R^{A1})(R^{A2})-NR^{A3}-, -C(=O)-, -C(R^{A1})(R^{A2})-C(=O)-, og -N=C(NH_2)- hvor ikke mer enn en av A, B, D og E er -C(R^{A1})(R^{A2})-O-, -C(R^{A1})(R^{A2})-NR^{A3}-, -C(R^{A1})(R^{A2})-C(=O)-, -C(=O)-, eller -N=C(NH_2)-;

U er N eller CR^U, hvor R^U er H, halo, CN, OH, C_{1-4} alkyl, C_{1-4} alkoksy, amino, C_{1-4} alkylamino, eller C_{2-8} dialkylamino;

W er N eller CR^W, hvor R^W er H, halo, CN, OH, C_{1-4} alkyl, C_{1-4} alkoksy, amino, C_{1-4} alkylamino, eller C_{2-8} dialkylamino;

X er N eller CR^X, hvor R^X er H, halo, CN, OH, C_{1-4} alkyl, C_{1-4} alkoksy, amino, C_{1-4} alkylamino, eller C_{2-8} dialkylamino, hvor hvis X er N, atomet av L som er direkte bundet til X er et annet enn N, O eller S;

L er valgt fra -C_{1-6} alkylen- og -(C_{1-4} alkylen)$_a$-Q-(C_{1-4} alkylen)b-, hvor C_{1-6} alkylengruppe og hvilken som helst C_{1-4} alkylengruppe av -(C_{1-4} alkylen)$_a$-Q-(C_{1-4} alkylen)b- gruppen er eventuelt substituert med 1, 2 eller 3 substituenter som er uavhengig valgt fra halo, CN, OH, C_{1-3} alkyl, C_{1-3} alkoksy, C_{1-3} haloalkyl, C_{1-3} haloalkoksy, amino, C_{1-3} alkylamino, og di(C_{1-3} alkyl)amino;

Q er -O-, -S-, -S(=O)-, -$S(=O)_2$-, -C(=O)-, -C(=O)NR^{q1}-, -C(=O)O-, -OC(=O)NR^{q1}-,

-NR^{q1}-, -NR^{q1}C(=O)O-, -NR^{q1}C(=O)NR^{q1}-, -$S(=O)_2NR^{q1}$-, -C(=NR^{q2})-, eller -C(=NR^{q2})-NR^{q1}-, hvor hver R^{q1} er uavhengig valgt fra H eller C_{1-6} alkyl, og hvor hver R^{q2} er uavhengig valgt fra H, C_{1-6} alkyl, og CN;

Cy er en koblings-C_{6-14} aryl, C_{3-18} sykloalkyl-, 5-16-leddet heteroaryl- eller 4-18-leddet heterosykloalkylgruppe, som hver er eventuelt substituert med 1, 2, 3 eller 4 substituenter som er uavhengig valgt fra R^{Cy};

hver R^{Cy} er uavhengig valgt fra halo, C_{1-6} alkyl, C_{1-4} haloalkyl, C_{1-4} cyanoalkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-10} aryl, C_{3-10} sykloalkyl, 5-10-leddet heteroaryl, 4-10-leddet heterosykloalkyl, CN,NO_2, OR^{a1} SR^{a1}, $C(O)R^{b1}$, $C(O)NR^{c1}R^{d1}$, $C(O)OR^{a1}$, $OC(O)R^{b1}$, $OC(O)NR^{c1}R^{d1}$, $C(=NR^{e1})NR^{c1}R^{d1}$, $NR^{c1}C(=NR^{e1})NR^{c1}R^{d1}$, $NR^{c1}R^{d1}$, $NR^{c1}C(O)R^{b1}$, $NR^{c1}C(O)OR^{a1}$, $NR^{c1}C(O)NR^{c1}R^{d1}$, $NR^{c1}S(O)R^{b1}$, $NR^{c1}S(O)_2R^{b1}$, $NR^{c1}S(O)_2NR^{c1}R^{d1}$, $S(O)R^{b1}$, $S(O)NR^{c1}R^{d1}$, $S(O)_2R^{b1}$ og $S(O)_2NR^{c1}R^{d1}$, hvor nevnte C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-10} aryl, C_{3-10} sykloalkyl, 5-10-leddet heteroaryl og 4-10-leddet heterosykloalkyl er hver eventuelt substituert med 1, 2, 3 eller 4 substituenter som er uavhengig valgt fra CN,NO_2, OR^{a1}, SR^{a1}, $C(O)R^{b1}$, $C(O)NR^{c1}R^{d1}$, $C(O)OR^{a1}$, $OC(O)R^{b1}$, $OC(O)NR^{c1}R^{d1}$, $C(=NR^{e1})NR^{c1}R^{d1}$, $NR^{c1}C(=NR^{e1})NR^{c1}R^{d1}$, $NR^{c1}R^{d1}$, $NR^{c1}C(O)R^{b1}$, $NR^{c1}C(O)OR^{a1}$, $NR^{c1}C(O)NR^{c1}R^{d1}$, $NR^{c1}S(O)R^{b1}$, $NR^{c1}S(O)_2R^{b1}$, $NR^{c1}S(O)_2NR^{c1}R^{d1}$, $S(O)R^{b1}$, $S(O)NR^{c1}R^{d1}$, $S(O)_2R^{b1}$, og $S(O)_2NR^{c1}R^{d1}$;

R^1 er H, Cy^1, halo, C_{1-6} alkyl, C_{1-4} haloalkyl, C_{1-4} cyanoalkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, CN,NO_2, OR^{a2}, SR^{a2}, $C(O)R^{b2}$, $C(O)NR^{c2}R^{d2}$, $C(O)OR^{a2}$, $OC(O)R^{b2}$, $OC(O)NR^{c2}R^{d2}$, $C(=NR^{e2})NR^{c2}R^{d2}$, $NR^{c2}C(=NR^{e2})NR^{c2}R^{d2}$, $NR^{c2}R^{d2}$, $NR^{c2}C(O)R^{b2}$, $NR^{c2}C(O)OR^{a2}$, $NR^{c2}C(O)NR^{c2}R^{d2}$, $NR^{c2}S(O)R^{b2}$, $NR^{c2}S(O)_2R^{b2}$, $NR^{c2}S(O)_2NR^{c2}R^{d2}$, $S(O)R^{b2}$, $S(O)NR^{c2}R^{d2}$, $S(O)_2R^{b2}$ og $S(O)_2NR^{c2}R^{d2}$, hvor nevnte C_{1-6} alkyl, C_{2-6} alkenyl og C_{2-6} alkynyl er hver eventuelt substituert med 1, 2, 3 eller 4 substituenter som er uavhengig valgt fra halo, CN,NO_2, OR^{a2}, SR^{a2}, $C(O)R^{b2}$, $C(O)NR^{c2}R^{d2}$, $C(O)OR^{a2}$, $OC(O)R^{b2}$, $OC(O)NR^{c2}R^{d2}$, $C(=NR^{e2})NR^{c2}R^{d2}$, $NR^{c2}C(=NR^{e2})NR^{c2}R^{d2}$, $NR^{c2}R^{d2}$, $NR^{c2}C(O)R^{b2}$, $NR^{c2}C(O)OR^{a2}$, $NR^{c2}C(O)NR^{c2}R^{d2}$, $NR^{c2}S(O)R^{b2}$, $NR^{c2}S(O)_2R^{b2}$, $NR^{c2}S(O)_2NR^{c2}R^{d2}$, $S(O)R^{b2}$, $S(O)NR^{c2}R^{d2}$, $S(O)_2R^{b2}$, og $S(O)_2NR^{c2}R^{d2}$;

Y er O, S,$CR^{Y1}R^{Y2}$ eller NR^{Y3}, hvor R^{Y1}, R^{Y2}og R^{Y3} er hver uavhengig valgt fra H og C_{1-4} alkyl;

Z er Cy^2, halo, C_{1-6} alkyl, C_{1-4} haloalkyl, C_{1-4} cyanoalkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, CN,NO_2, OR^{a3}, SR^{a3}, $C(O)R^{b3}$, $C(O)NR^{c3}R^{d3}$, $C(O)OR^{a3}$, $OC(O)R^{b3}$, $OC(O)NR^{c3}R^{d3}$, $C(=NR^{e3})NR^{c3}R^{d3}$, $NR^{c3}C(=NR^{e3})NR^{c3}R^{d3}$, $NR^{c3}R^{d3}$, $NR^{c3}C(O)R^{b3}$, $NR^{c3}C(O)OR^{a3}$, $NR^{c3}C(O)NR^{c3}R^{d3}$, $NR^{c3}S(O)R^{b3}$, $NR^{c3}S(O)_2R^{b3}$, $NR^{c3}S(O)_2NR^{c3}R^{d3}$, $S(O)R^{b3}$, $S(O)NR^{c3}R^{d3}$, $S(O)_2R^{b3}$, $S(O)_2NR^{c3}R^{d3}$og $P(O)R^{c3}R^{d3}$ hvor C_{1-6} alkyl, C_{2-6} alkenyl og C_{2-6} alkynyl er hver eventuelt substituert med 1, 2, 3 eller 4substituenter som er uavhengig valgt fra Cy^2, halo, CN,NO_2, CN,NO_2, OR^{a3}, SR^{a3}, $C(O)R^{b3}$, $C(O)NR^{c3}R^{d3}$, $C(O)OR^{a3}$, $OC(O)R^{b3}$, $OC(O)NR^{c3}R^{d3}$, $C(=NR^{e3})NR^{c3}R^{d3}$, $NR^{c3}C(=NR^{e3})NR^{c3}R^{d3}$,

$NR^{c3}R^{d3}$, $NR^{c3}C(O)R^{b3}$, $NR^{c3}C(O)OR^{a3}$, $NR^{c3}C(O)NR^{c3}R^{d3}$, $NR^{c3}S(O)R^{b3}$, $NR^{c3}S(O)R^{b3}$, $NR^{c3}S(O)_2NR^{c3}R^{d3}$, $S(O)R^{b3}$, $S(O)NR^{c3}R^{d3}$, $S(O)_2R^{b3}$, og $S(O)_2NR^{c3}R^{d3}$;

hver R^2 og R^3 er uavhengig valgt fra H, halo, C_{1-6} alkyl, C_{1-4} haloalkyl, C_{1-4} cyanoalkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, CN,NO_2, OR^{a4}, SR^{a4}, $C(O)R^{b4}$, $C(O)NR^{c4}R^{d4}$, $C(O)OR^{a4}$, $OC(O)R^{b4}$, $OC(O)NR^{c4}R^{d4}$, $C(=NR^{e4})NR^{c4}R^{d4}$, $NR^{c4}C(=NR^{e4})NR^{c4}R^{d4}$, $NR^{c4}R^{d4}$, $NR^{c4}C(O)R^{b4}$, $NR^{c4}C(O)OR^{a4}$, $NR^{c4}C(O)NR^{c4}R^{d4}$, $NR^{c4}S(O)R^{b4}$, $NR^{c4}S(O)_2R^{b4}$, $NR^{c4}S(O)_2NR^{c4}R^{d4}$, $S(O)R^{b4}$, $S(O)NR^{c4}R^{d4}$, $S(O)_2R^{b4}$, og $S(O)_2NR^{c4}R^{d4}$, hvor nevnte C_{1-6} alkyl, C_{2-6} alkenyl og C_{2-6} alkynyl er hver eventuelt substituert med 1, 2, 3 eller 4 substituenter som er uavhengig valgt fra halogen, CN,NO_2, OR^{a4}, SR^{a4}, $C(O)R^{b4}$, $C(O)NR^{c4}R^{d4}$, $C(O)OR^{a4}$, $OC(O)R^{b4}$, $OC(O)NR^{c4}R^{d4}$, $C(=NR^{e4})NR^{c4}R^{d4}$, $NR^{c4}C(=NR^{e4})NR^{c4}R^{d4}$, $NR^{c4}R^{d4}$, $NR^{c4}C(O)R^{b4}$, $NR^{c4}C(O)OR^{a4}$, $NR^{c4}C(O)NR^{c4}R^{d4}$, $NR^{c4}S(O)R^{b4}$, $NR^{c4}S(O)_2R^{b4}$, $NR^{c4}S(O)_2NR^{c4}R^{d4}$, $S(O)R^{b4}$, $S(O)NR^{c4}R^{d4}$, $S(O)_2R^{b4}$, og $S(O)_2NR^{c4}R^{d4}$;

hver R^{A1} er uavhengig valgt fra H, halo, C_{1-4} alkyl, C_{1-4} alkoksy, C_{1-4} haloalkyl, C_{1-4} haloalkoksy, amino, C_{1-4} alkylamino, C_{2-8} dialkylamino, CN,NO_2og OH;

hver R^{A2} er uavhengig valgt fra H, halo, C_{1-4} alkyl, C_{1-4} alkoksy, C_{1-4} haloalkyl, C_{1-4} haloalkoksy, amino, C_{1-4} alkylamino, C_{2-8} dialkylamino, CN,NO_2og OH;

hver R^{A3} er uavhengig valgt fra H, C_{1-4} alkyl, C_{1-4} alkoksy, C_{1-4} haloalkyl, $C(O)R^z$, og $C(O)OR^z$, hvor nevnte C_{1-4} alkyl er eventuelt substituert med fenyl, C_{1-4} alkoksy, C_{1-4} haloalkoksy, CN,NO_2, eller OH;

R^z er H, C_{1-4} alkyl, eller fenyl;

hver Cy^1 er uavhengig valgt fra C_{6-14} aryl, C_{3-18} sykloalkyl, 5-16 leddet heteroaryl og 4-18 leddet heterosykloalkyl, som hver er eventuelt substituert med 1, 2, 3 eller 4 substituenter som er uavhengig valgt fra R^{Cy1};

hver Cy^2 er uavhengig valgt fra C_{6-14} aryl, C_{3-18} sykloalkyl, 5-16 leddet heteroaryl og 4-18 leddet heterosykloalkyl, som hver er eventuelt substituert med 1, 2, 3 eller 4 substituenter uavhengig valgt fra R^{Cy2};

hver R^{Cy1} og R^{Cy2} er uavhengig valgt fra halo, C_{1-6} alkyl, C_{1-4} haloalkyl, C_{1-4} cyanoalkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, fenyl, C_{3-7} sykloalkyl, 5-6-leddet heteroaryl og 4-7-leddet heterosykloalkyl, CN,NO_2, OR^{a5}, SR^{a5}, $C(O)R^{b5}$, $C(O)NR^{c5}R^{d5}$, $C(O)OR^{a5}$, $OC(O)R^{b5}$, $OC(O)NR^{c5}R^{d5}$, $C(=NR^{e5})NR^{c5}R^{d5}$, $NR^{c5}C(=NR^{e5})NR^{c5}R^{d5}$, $NR^{c5}R^{d5}$, $NR^{c5}C(O)R^{b5}$, $NR^{c5}C(O)OR^{a5}$, $NR^{c5}C(O)NR^{c5}R^{d5}$, $NR^{c5}S(O)R^{b5}$, $NR^{c5}S(O)_2R^{b5}$, $NR^{c5}S(O)_2NR^{c5}R^{d5}$, $S(O)R^{b5}$, $S(O)NR^{c5}R^{d5}$, $S(O)_2R^{b5}$, og $S(O)_2NR^{c5}R^{d5}$, hvor nevnte

C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, fenyl, C_{3-7} sykloalkyl, 5-6-leddet heteroaryl og 4-7-leddet heterosykloalkyl er hver eventuelt substituert med 1, 2, 3 eller 4 substituenter som er uavhengig valgt fra CN,NO_2, OR^{a5}, SR^{a5}, $C(O)R^{b5}$, $C(O)NR^{c5}R^{d5}$, $C(O)OR^{a5}$, $OC(O)R^{b5}$, $OC(O)NR^{c5}R^{d5}$, $C(=NR^{e5})NR^{c5}R^{d5}$, $NR^{c5}C(=NR^{e5})NR^{c5}R^{d5}$, $NR^{c5}R^{d5}$, $NR^{c5}C(O)R^{b5}$, $NR^{c5}C(O)OR^{a5}$, $NR^{c5}C(O)NR^{c5}R^{d5}$, $NR^{c5}S(O)R^{b5}$, $NR^{c5}S(O)_2R^{b5}$, $NR^{c5}S(O)_2NR^{c5}R^{d5}$, $S(O)R^{b5}$, $S(O)NR^{c5}R^{d5}$, $S(O)_2R^{b5}$, og $S(O)_2NR^{c5}R^{d5}$;

hver R^{a1}, R^{b1}, R^{c1}, R^{d1}, R^{a2}, R^{b2}, R^{c2}, R^{d2}, R^{a3}, R^{b3}, R^{c3}, R^{d3}, R^{a4}, R^{b4}, R^{c4}, R^{d4}, R^{a5}, R^{b5}, R^{c5}og R^{d5} er uavhengig valgt fra H, C_{1-6} alkyl, C_{1-4} haloalkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-10} aryl, C_{3-10} sykloalkyl, 5-10-leddet heteroaryl, 4-10-leddet heterosykloalkyl, C_{6-10} aryl-C_{1-6} alkyl, C_{3-10} sykloalkyl-C_{1-6} alkyl, (5-10-leddet heteroaryl)-C_{1-6} alkyl, og (4-10-leddet heterosykloalkyl)-C_{1-6} alkyl, hvor nevnte C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{6-10} aryl, C_{3-10} sykloalkyl, 5-10-leddet heteroaryl, 4-10-leddet heterosykloalkyl, C_{6-10} aryl-C_{1-6} alkyl, C_{3-10} sykloalky-C_{1-6} alkyl, (5-10-leddet heteroaryl)-C_{1-6} alkyl, og (4-10-leddet heterosykloalkyl)-C_{1-6} alkyl er hver eventuelt substituert med 1, 2, 3, 4 eller 5 substituenter som er uavhengig valgt fra R^g;

hver R^{e1}, R^{e2}, R^{e3}, R^{e4} og R^{e5} er uavhengig valgt fra H, C_{1-4} alkyl, og CN;

hver R^g er uavhengig valgt fra gruppen bestående av OH, NO_2, CN, halo, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{1-4} haloalkyl, C_{1-6} alkoksy, C_{1-6} haloalkoksy, cyano-C_{1-3} alkyl, HO-C_{1-3} alkyl, amino, C_{1-6} alkylamino, di(C_{1-6} alkyl)amino, tiol, C_{1-6} alkyltio, C_{1-6} alkylsulfinyl, C_{1-6} alkylsulfonyl, karboksy, aminokarbonyl, C_{1-6} alkylkarbonyl og C_{1-6} alkoksykarbonyl;

n er 0 eller 1;

m er 0 eller 1;

p er 0, 1, 2 eller 3;

q er 0, 1 eller 2;

a er 0 eller 1; og

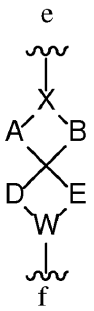
b er 0 eller 1,

hvor en hvilken som helst sykloalkyl- eller heterosykloalkylgruppe eventuelt er ytterligere substituert med 1 eller 2 okso-grupper.

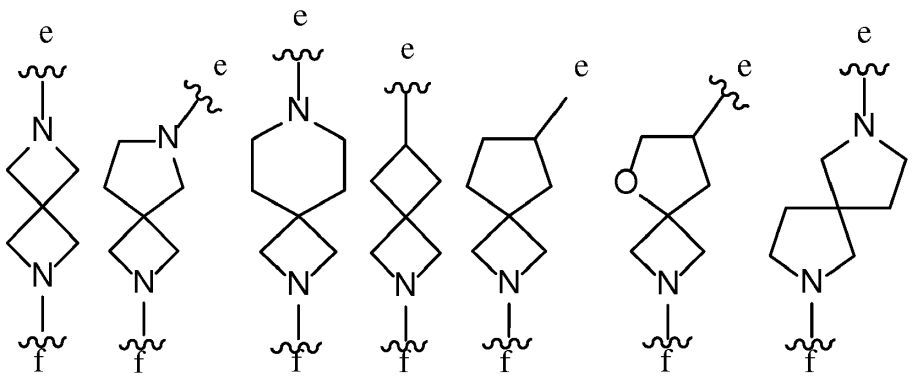
2. Forbindelsen ifølge krav 1, eller et farmasøytisk akseptabelt salt derav, hvor
i) Y er O, eller ii) Y er NR^{Y3},
og/eller i) U er N, eller ii) U er CR^{U},
og/eller i) W er N, eller ii) W er CR^{W}
og/eller i) X er N, eller ii) X er CR^{X}
og/eller i) A, B, D og E er hver uavhengig valgt fra $-C(R^{A1})(R^{A2})-$, og $-C(R^{A1})(R^{A2})-C(R^{A1})(R^{A2})-$, eller ii) A, B, D og E er hver uavhengig valgt fra $-CH_2-$ og $-CH_2-CH_2-$.

3. Forbindelsen ifølge krav 1 eller krav 2, eller et farmasøytisk akseptabelt salt derav, hvor:

i) spirodelen er representert ved formelen nedenfor:



hvor e og f indikerer festepunkter til resten av molekylet, og er valgt fra:



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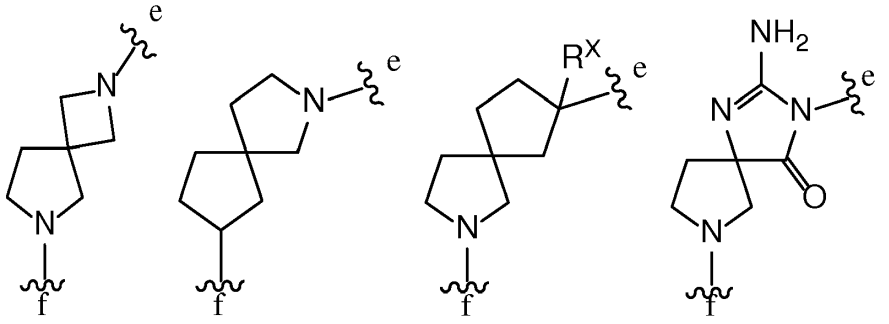

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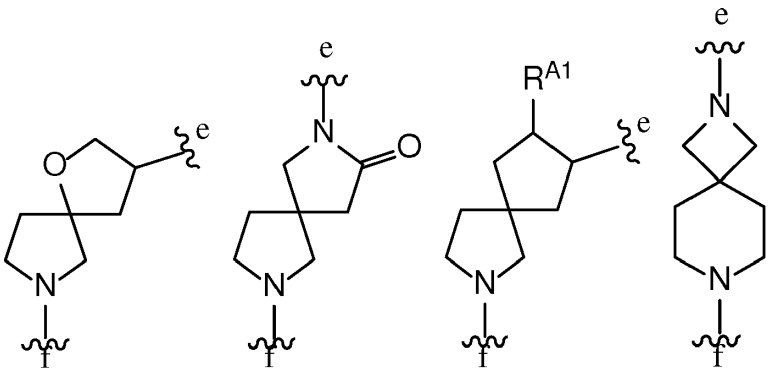

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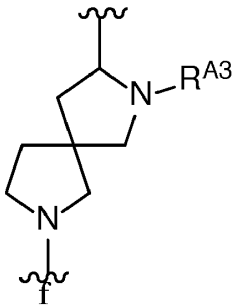
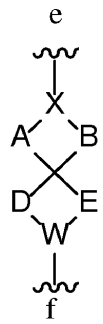


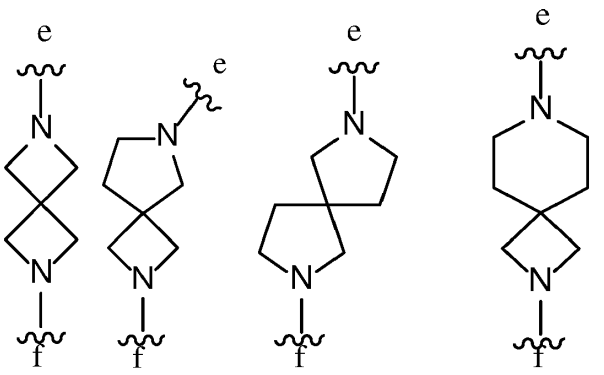
,

eller

ii) spirodelen er representert ved formelen nedenfor:



hvor e og f indikerer festepunkter til resten av molekylet, og er valgt fra:



, , og .

4. Forbindelsen ifølge et hvilket som helst av kravene 1 til 3, eller et farmasøytisk akseptabelt salt derav, hvor

i) L er valgt fra -C_{1-6} alkylen- som er eventuelt substituert med 1, 2 eller 3 substituenter som er uavhengig valgt fra halo, CN, OH, C_{1-3} alkyl, C_{1-3} alkoksy, C_{1-3} haloalkyl, C_{1-3} haloalkoksy, amino, C_{1-3} alkylamino, og di(C_{1-3} alkyl)amino, eller

ii) L er valgt fra metylen, etylen og -CH_2-CH(OH)-, eller

iii) L er metylen, eller

iv) L er valgt fra -(C_{1-4} alkylen)a-Q-(C_{1-4} alkylen)b-, hvor hvilken som helst C_{1-4} alkylengruppe av de N-(C_{1-4} alkylen)a-Q-(C_{1-4} alkylen)b- gruppen er eventuelt substituert med 1, 2 eller 3 substituenter som er uavhengig valgt fra halo, CN, OH, C_{1-3} alkyl, C_{1-3} alkoksy, C_{1-3} haloalkyl, C_{1-3} haloalkoksy, amino, C_{1-3} alkylamino, og di(C_{1-3} alkyl)amino, eller

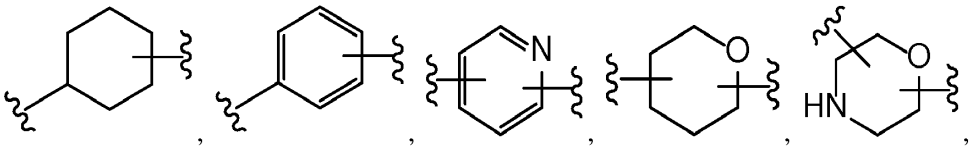
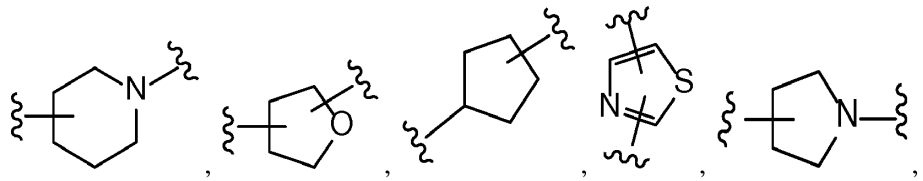
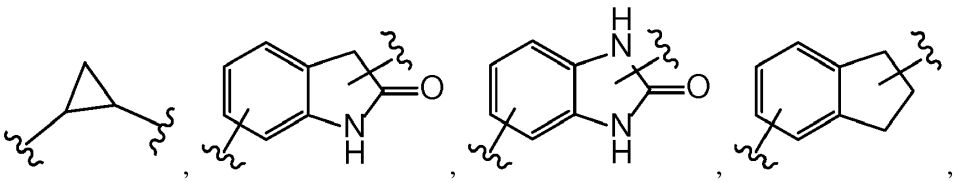
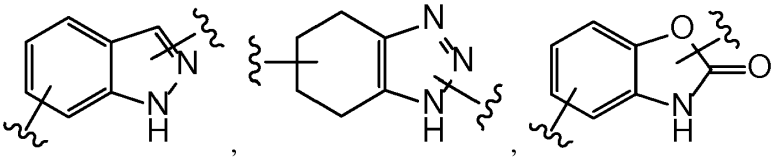
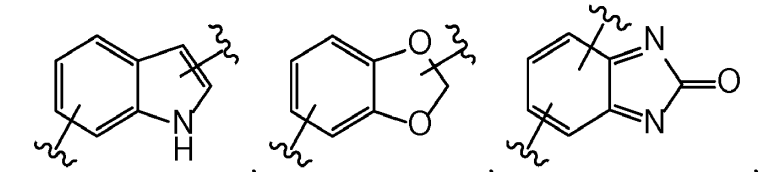
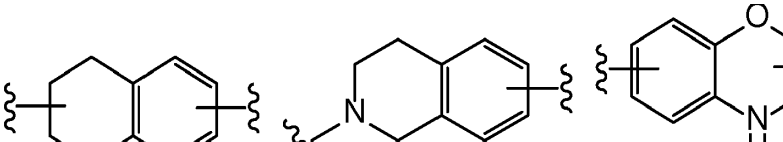
v) L er valgt fra -C(O)-CH_2-, -C(O)-CH_2-CH_2-, C(O), -NH-CH_2-, NH, -C(O)-CH(NH_2)-, -NH-CH(CH_3)-, -N(CH_3)-C(O)-, N(CH_3)-CH_2-, -CH_2-CH_2-O-, og -C(O)-NH-.

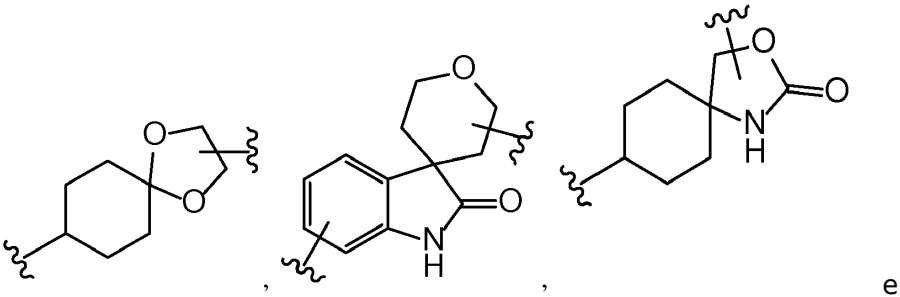
5. Forbindelsen ifølge et hvilket som helst av kravene 1 til 4, eller et farmasøytisk akseptabelt salt derav, hvor

i) Cy er et koblingsfenyl, C_{3-18} sykloalkyl-, 5-10-leddet heteroaryl- eller 4-9-leddet heterosykloalkylgruppe, som hver er eventuelt substituert med 1, 2, 3 eller 4 substituenter som er uavhengig valgt fra R^{Cy}, eller

ii) Cy er et koblingsfenyl, C_{3-18} sykloalkyl-, 5-10-leddet heteroaryl- eller 4-9-leddet heterosykloalkylgruppe, som hver er eventuelt substituert med 1, 2, 3 eller 4 substituenter som er uavhengig valgt fra R^{Cy}, eller

iii) Cy er en koblingsgruppe som har formelen:

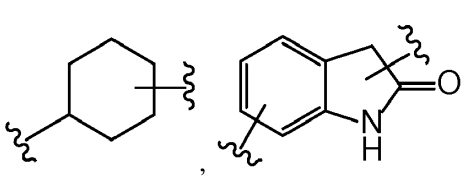


eller

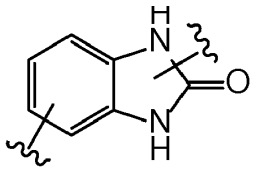
,

hver av disse er eventuelt substituert med 1, 2, 3 eller 4 substituenter som er uavhengig valgt fra R^{Cy}, eller

iv) Cy er en koblingsgruppe som har formelen:



, eller



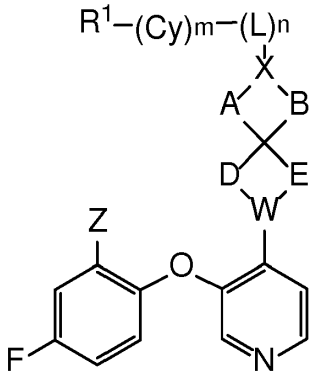
eller

v) Z er Cy^2 eller $C(O)NR^{c3}R^{d3}$.

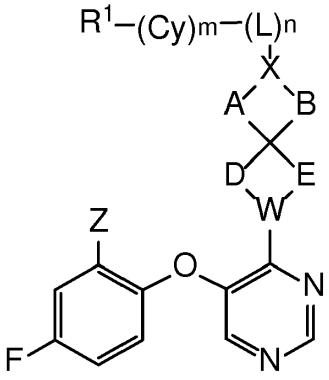
6. Forbindelsen ifølge et hvilket som helst av kravene 1 til 5, eller et farmasøytisk akseptabelt salt derav, hvor
i) n er 0, eller ii) n er 1,
og/eller i) m er 0 eller ii) m er 1,
og/eller i) p er 0 eller ii) p er 1,
og/eller i) q er 0 eller ii) q er 1.

7. Forbindelsen ifølge et hvilket som helst av kravene 1 og 3 til 6, eller et farmasøytisk akseptabelt salt derav, som har

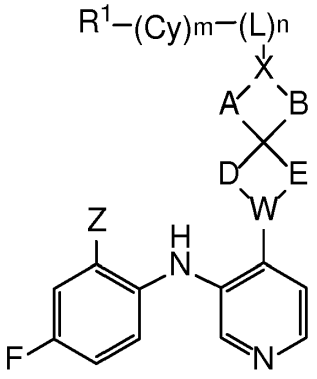
i) formel IIa, IIb, IIIa eller IIIb:



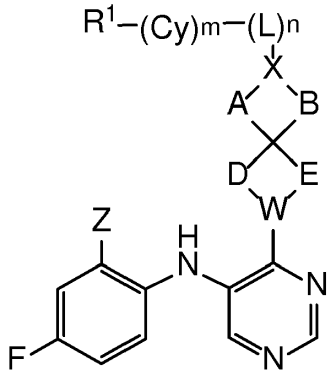
IIa



IIb



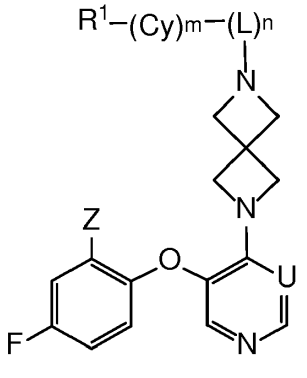
IIIa



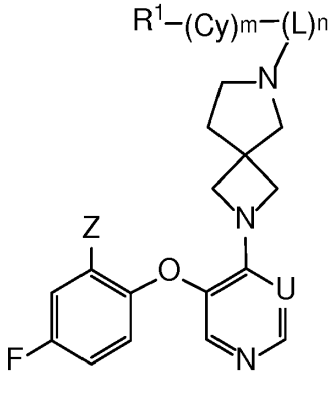
IIIb ,

eller

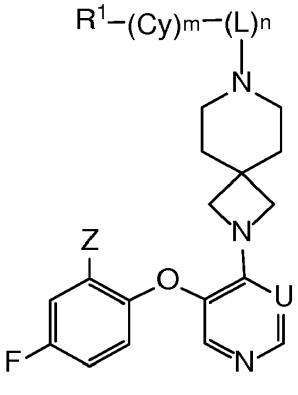
ii) formel IVa, IVb, IVc, IVd, IVe eller IVf:



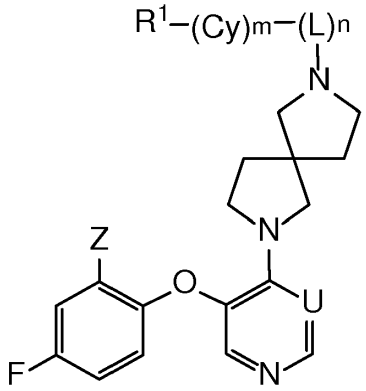
IVa



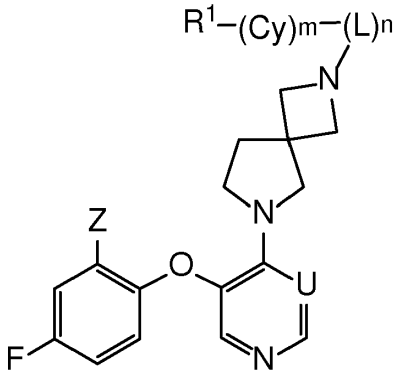
IVb



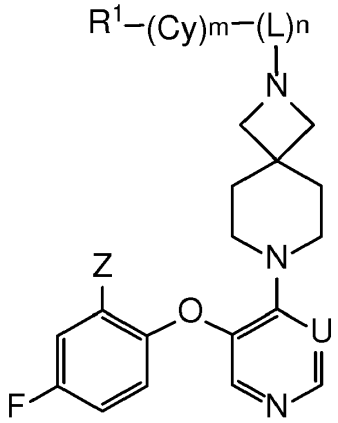
IVc



IVd



IVe



IVf.

8. Forbindelsen ifølge krav 1, hvor forbindelsen er valgt fra:

5-fluor-N,N-diisopropyl-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

N-etyl-5-fluor-N-isopropyl-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-2-((4-(7-((1-(2-hydroksyetyl)-2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-N,N-diisopropylbenzamid;

N-etyl-5-fluor-2-((4-(7-((1-(2-hydroksyetyl)-2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-N-isopropylbenzamid;

5-fluor-N-(2-hydroksyetyl)-N-isopropyl-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-N,N-diisopropyl-2-((4-(7-(((1r,4r)-4-(metylsulfonamido)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-N,N-diisopropyl-2-((4-(7-(((1s,4s)-4-(metylsulfonamido)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-((7-(5-(2-(amino(syklopentyl)metyl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-(2-(syklopentyl(dimetylamino)metyl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

N-(syklopentyl(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)metyl) acetamid;

6-((7-(5-(4-fluor-2-(1-hydroksy-2-metylpropyl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-3,3-dimetylindolin-2-on;

6-((7-(5-(4-fluor-2-isobutyrylfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-3-metyl-2-oksoindolin-3-karbonitril;

5-fluor-2-((4-(6-(3-(4-fluorfenyl)propanoyl)-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)-N,N-diisopropylbenzamid;

5-((7-(5-(4-fluor-2-(1-isopropyl-1H-pyrazol-5-yl)fenoksy)pyrimidin-4-yl)-3-okso-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

N-(4-fluor-2-(5-isopropyl-3-metylisoksazol-4-yl)fenyl)-4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-amin;

4-(5-fluor-2-((4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)fenyl)-5-isopropyl-3-metylisoksazol;

N-(5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)-4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-amin;

5-fluor-2-((4-(2-(2-hydroksy-2-metylpropyl)-2,7-diazaspiro[3.5]nonan-7-yl)pyrimidin-5-yl)amino)-N,N-diisopropylbenzamid;

5-((7-(5-(2-(dimetylfosforyl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-N-(4-fluorbenzyl)-5-oksa-2-azaspiro[3.4]oktan-7-amin;

4-(((2-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-5-oksa-2-azaspiro[3.4]oktan-7-yl)amino)metyl)benzonitril;

7-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-N-(4-fluorbenzyl)-1-oksa-7-azaspiro[4.4]nonan-3-amin;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-(((1r,4r)-4-(metylkarbamoyl)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

2-((4-(7-amino-7-(4-cyanobensyl)-2-azaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N,N-diisopropylbenzamid;

5-fluor-2-((4-(7-hydroksy-8-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2-azaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-N,N-diisopropylbenzamid;

2-((4-(7-amino-8-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2-azaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N,N-diisopropylbenzamid;

5-fluor-2-((4-(8-(4-fluorbenzyl)-7-(2-hydroksyetyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-N,N-diisopropylbenzamid;

6-((7-(5-(2-klor-4-fluorfenoksy)pyrimidin-4-yl)-2-azaspiro[4.4]nonan-2-yl)metyl)-1-metyl-1H-benzo[d]imidazol-2(3H)-on;

5-((7-(3-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyridin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-(2-(3-syklopropyl-1-metyl-6-okso-1,6-dihydropyridin-2-yl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

N-(5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)-4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyridin-3-amin;

2-(5-((4',5-difluor-2'-(2-fluorpropan-2-yl)-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan;

5-fluor-N-isopropyl-2-((4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)benzensulfonamid;

5-((7-(5-(4-fluor-2-(2-metoksybutan-2-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-(4-fluor-2-(3-hydroksypentan-3-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

2-(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)-N-metylsyklopropankarboksamid;

5-((7-(5-(4-fluor-2-(3-hydroksy-3-metylbutyl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

metyl 2-(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)syklopropankarboksylat;

2-(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)-N-metylsyklopropankarboksamid;

6-((2-(5-(2-klor-4-fluorfenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.4]oktan-6-yl)metyl)-3,3-dimetylindolin-2-on;

2-(6-(5-(2-klor-4-fluorfenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-karbonyl)-2,3-dihydro-1H-inden-5-sulfonamid;

5-((7-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

5-((7-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pynmidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

2-syklopropyl-5'-fluor-2’-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

2-syklopropyl-5'-fluor-2’-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karboksamid;

2-syklopropyl-5'-fluor-2’-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karboksylsyre;

2-syklopropyl-5'-fluor-N,N-dimetyl-2’-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karboksamid;

5-((7-(2-klor-5-(4-fluor-2-(1-isopropyl-1H-pyrazol-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-((4,5-difluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

5'-fluor-2-metyl-2’-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

5-((7-(5-((2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

5-((7-(5-((5-fluor-2’-(trifluormetyl)-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

5'-fluor-2,6-dimetyl-2’-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

2-syklopropyl-3’, 5'-difluor-2’-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

5-((7-(5-(4-fluor-2-(2-isopropyl-1H-imidazol-1-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

5-((7-(5-(2-(syklopropylmetoksy)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H -benzo[d]imidazol-2-on;

etyl 2-(7-(5-((4'-cyano-2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)tiazol-4-karboksylat;

2-(7-(5-((4'-cyano-2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)tiazol-4-karboksylsyre;

2-(7-(5-((4'-cyano-2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)-N-metyltiazol-4-karboksamid;

2-(7-(5-((4'-cyano-2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)-N,N-dimetyltiazol-4-karboksamid;

7-benzyl-2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2-azaspiro[4.4]nonan;

5-((7-(5-((5-fluor-2'-(1-hydroksyetyl)-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-(3-(2-oksooxazolidin-3-yl)benzyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-(4-(2-oksooxazolidin-3-yl)benzyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-((7-(5-((5-fluor-2'-(2-hydroksypropan-2-yl)-[1,1'-bifenyl]-2-yl)oksy)pyrimidiN-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

2-(1,4-dioksaspiro[4.5]dekan-8-ylmetyl)-6-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan;

4-((6-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)metyl)sykloheksanol;

2-syklopropyl-5'-fluor-2'-((4-(6-((4-hydroksysykloheksyl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

2-(5-((5-fluor-2'-(1-metoksyetyl)-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan;

5-(5-fluor-2-((4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)fenyl)-2,3-dihydro-1H-inden-2-amin;

5-((7-(5-((5-fluor-2'-(1-hydroksypropan-2-yl)-[1,1'-bifenyl]-2-yl)oksy)pyrimidiN-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

5-((7-(5-(4-fluor-2-(morfolinometyl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d]imidazol-2(3H)-on;

1-(7-(5-((2'-etyl-5-fluor-[1,1'-bifenyl]-2-yl)amino)pyrimidin-4-yl)-2,7-diazaspiro[3.5]nonan-2-yl)-2-metylpropan-2-ol;

1-((6-(5-(4-fluor-2-(1-isopropyl-3-(trifluormetyl)-1H-pyrazol-5-yl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)metyl)sykloheksan-1-ol;

N-(2-amino-2-oksoetyl)-N-(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)isobutyramid;

N-(5-fluor-2-((4-(7-(((1r,4r)-4-(metylsulfonamido)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl) propan-2-sulfonamid;

tert-butyl 7-(5-(4-fluor-2-(N-metylisobutyramido)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-karboksylat;

N-(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4,4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)-N-metylisobutyramid;

5-((7-(5-(4-fluor-2-isobutylfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazo1-2-on;

2-(3-((2'-etyl-5-fluor-[1,1'-bifenyl]-2-yl)metyl)pyridin-4-yl)-6-((tetrahydro-2H-pyran-4-yl))metyl)-2,6-diazaspiro[3.3]heptan;

N-((1r,4r)-4-((2-(5-(2-(3-syklopropyl-1-metyl-6-okso-1,6-dihydropyridin-2-yl)-4-fluorfenoksy)pyrimidiN-4-yl)-2,7-diazaspiro[3.5]nonan-7-yl)metyl)sykloheksyl)-2,2,2-trifluoracetamid;

N-(4-((2-(5-(2-(3-syklopropyl-1-metyl-6-okso-1,6-dihydropyridin-2-yl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[3.5]nonan-7-yl)metyl)sykloheksyl)metansulfonamid;

5-((7-(5-(2-(3-syklopropyl-1-metyl-6-okso-1,6-dihydropyridin-2-yl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1-(2-hydroksyetyl)-1H-benzo[d]imidazol-2(3H)-on;

(1r,4r)-4-(2-(6-(5-(2-(2-syklopropylpyridin-3-yl)-4-fluorfenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)etyl)sykloheksan-1-amin;

tert-butyl((1r,4r)-4-(((2-(5-(4-fluor-2-(isopropyl(metyl) karbamoyl)fenoksy)pyrimidin-4-yl)-2-azaspiro[3.3]heptaN-6-yl)amino)metyl)sykloheksyl)karbamat;

tert-butyl((1r,4r)-4-((2-(5-(2-(N-etylisobutyramido)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[3.5]nonan-7-yl)metyl)sykloheksyl)karbamat;

metyl((1r,4r)-4-((2-(5-(2-(N-etylisobutyramido)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[3.5]nonan-7-yl)metyl)sykloheksyl)karbamat;

N-etyl-N-(5-fluor-2-((4-(7-(((1r,4r)-4-(metylsulfonamido)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)isobutyramid;

2-((4-(6-(2-((1r,4r)-4-(3,3-dimetylbutanamido)sykloheksyl)etyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N,N-diisopropylbenzamid;

tert-butyl((1r,4r)-4-(2-(6-(5-(2-(2-syklopropylpyridin-3-yl)-4-fluorfenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)etyl)sykloheksyl)karbamat;

5-fluor-2-((4-(7-(2-hydroksy-2-metylpropyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)-N,N-diisopropylbenzamid;

2-((4-(7-((3-cyano-3-metyl-2-oksoindolin-6-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N,N-diisopropylbenzamid;

metyletyl(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4,4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)karbamat;

5-fluor-2-((4-(7-((1-(2-hydroksyetyl)-2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-N-isopropyl-N-metylbenzamid;

5-fluor-N-isopropyl-N-metyl-2-((4-(6-((tetrahydro-2H-pyran-4-yl)amino)-2-azaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-(((1r,4r)-4-(metylsulfonamido)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

tert-butyl((1r,4r)-4-((2-(5-(4-fluor-2-(isopropyl(metyl)karbamoyl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[3.5]nonan-7-yl)metyl)sykloheksyl)karbamat;

metyl((1r,4r)-4-((2-(5-(4-fluor-2-(isopropyl(metyl)karbamoyl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[3.5]nonan-7-yl)metyl)sykloheksyl)karbamat;

N-(tert-butyl)-2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2-azaspiro[3.4]oktan-6-amin;

2-((4-(7-(((1r,4r)-4-(3,3-dimetylureido)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N,N-diisopropylbenzamid;

5-fluor-2-((4-(7-((4-hydroksysykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)-N,N-diisopropylbenzamid;

5-fluor-2-((4-(6-((4-hydroksysykloheksyl)metyl)-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)-N,N-diisopropylbenzamid;

2-((4-(7-((1,4-dioksaspiro[4,5]dekan-8-yl)metyl)-2,7-diazaspiro[3,5]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N,N-diisopropylbenzamid;

5-fluor-N,N-diisopropyl-2-((4-(7-((tetrahydro-2H-pyran-4-yl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-N,N-diisopropyl-2-((4-(6-neopentyl-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)benzamid;

2-((4-(6-(syklopropylmetyl)-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N,N-diisopropylbenzamid;

2-((4-(6-(6-cyano-1,2,3,4-tetrahydronaftalen-2-yl)-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N,N-diisopropylbenzamid;

5-fluor-N,N-diisopropyl-2-((4-(6-(2-((1r,4r)-4-pivalamidosykloheksyl)etyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)benzamid;

N-(2-((4-(6-(sykloheksylmetyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)-5-fluorfenyl)-N-etylisobutyramid;

N-etyl-5-fluor-N-isopropyl-2-((4-(7-((1-metyl-2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

2-(5-(2-(syklopentyloksy)-4-fluorfenoksy)pyrimidin-4-yl)-7-((tetrahydro-2H-pyran-4-yl)metyl)-2,7-diazaspiro[4.4]nonan;

2-(5-(2-syklopropoksy-4-fluorfenoksy)pyrimidin-4-yl)-7-((tetrahydro-2H-pyran-4-yl)metyl)-2,7-diazaspiro[4.4]nonan;

N-etyl-N-(5-fluor-2-((4-(7-((tetrahydro-2H-pyran-4-yl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)isobutyramid;

5-fluor-N,N-diisopropyl-2-((4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-N,N-diisopropyl-2-((4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)benzamid;

2-((4-(6-(2-(4-cyanofenyl) acetyl)-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N,N-diisopropylbenzamid;

5-fluor-2-((4-(6-(6-fluor-1,2,3,4-tetrahydronaftalen-2-yl)-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)-N,N-diisopropylbenzamid;

tert-butyl((1r,4r)-4-(2-(6-(5-(2-(diisopropylcarbamoyl)-4-fluorfenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)etyl)sykloheksyl)karbamat;

2-((4-(6-(2-(4-cyanofenyl) acetyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N,N-diisopropylbenzamid;

N-etyl-N-(5-fluor-2-((4-(6-(5-(metylsulfonyl)-2,3-dihydro-1H-inden-2-karbonyl)-2,6diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)fenyl)isobutyramid;

3-(((2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2-azaspiro[3.3]heptan-6-yl)amino)metyl)bisyklo[1.1.1]pentan-1-karbonitril;

N-etyl-N-(5-fluor-2-((4-(6-(2-(4-(metylsulfonyl)fenyl) acetyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)fenyl)isobutyramid;

N-(2-((4-(6-(5-brom-2,3-dihydro-1H-inden-2-karbonyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)-5-fluorfenyl)-N-etylisobutyramid;

N-etyl-N-(5-fluor-2-((4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)fenyl)isobutyramid;

N-syklopropyl-5-fluor-N-isopropyl-2-((4-(7-((2-oxo-2,3-dihydro-1H-benzo[d]imidazol-5-yl)methyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

2-((4-(7-((1-(2-acetamidoetyl)-2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4,4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid;

2-((4-(7-((1-(2-(dimetylamino)etyl)-2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid;

2-((4-(7-((3-cyano-3-metyl-2-oksoindolin-6-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid;

5-((7-(5-(4-fluor-2-(1-isopropyl-1H-pyrazol-5-yl)fenoksy)-2-metylpyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

2-((4-(2-(2-(4-cyanofenyl) acetyl)-2,6-diazaspiro[3.4]oktan-6-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid;

2-((4-(7-((1-etyl-2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid;

5-fluor-N-isopropyl-2-((4-(7-((1-(2-metoksyetyl)-2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-N-metylbenzamid;

4-(2-(6-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.4]oktan-2-yl)-2-oksoetyl)benzonitril;

5-((7-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1-(2-metoksyetyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

1-(6-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)-2-(6-metoksypyridin-3-yl)ethan-1-on;

6-(2-(6-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)-2-oksoetyl)-3,3-dimetylindolin-2-on;

tert-butyl((1r,4r)-4-(2-(6-(5-(4-fluor-2-(isopropyl(metyl) karbamoyl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)etyl)sykloheksyl)karbamat;

5-((7-(5-(4-fluor-2-((isopropyl(metyl)amino)metyl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

N-etyl-N-(5-fluor-2-((4-(6-isobutyl-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)fenyl)isobutyramid;

N-(2-((4-(6-((4,4-difluorsykloheksyl)metyl)-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)-5-fluorfenyl)-N-etylisobutyramid;

tert-butyl((1r,4r)-4-(2-(6-(5-(4-fluor-2-(N-metylisobutyramido)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)etyl)sykloheksyl)karbamat;

2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-7-(6-fluor-3,4-dihydroisokinolin-2(1H)-yl)-5-oxa-2-azaspiro[3.4]oktan;

4-(((2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2-azaspiro[3.3]heptan-6-yl)amino)metyl)-1-metylsykloheksan-1-karbonitril;

4-(1-((2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2-azaspiro[3.3]heptan-6-yl)amino)etyl)benzonitril;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-(4-(2-oksooxazolidin-3-yl)benzyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

N-((1r,4r)-4-(2-(6-(5-((4'-cyano-2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)etyl)sykloheksyl) acetamid;

metyl(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)(isopropyl)karbamat;

2-((4-(7-((1H-indazol-6-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid;

2-((4-(7-((3-cyano-1H-indazol-6-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid;

tert-butyl((1r,4r)-4-((7-(5-(2-(syklopentyloksy)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)sykloheksyl)karbamat;

4-((2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2-azaspiro[3.3]heptan-6-yl)amino)-1-metylsykloheksankarbonitril;

4-(2-(2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.4]oktan-6-yl)-2-oksoetyl)benzonitril;

5-((7-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1-metyl-1,3-dihydro-2H-benzo[d]imidazol-2-on;

2-syklopropyl-5'-fluor-2’-((4-(6-((4-hydroksysykloheksyl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

4-(((2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2-azaspiro[3.3]heptan-6-yl)amino)metyl)benzonitril;

5-((7-(5-(2-(2,5-dimetylpyrrolidin-1-karbonyl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-(4-fluor-2-(pyrrolidin-1-karbonyl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-(4-fluor-2-(morfolin-4-karbonyl)fenoksy)pyrimidin-4-yl)-2,7-diaza spiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

N-etyl-N-(5-fluor-2-((4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)fenyl)isobutyramid;

7-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-N-(4-fluorbenzyl)-1-oksa-7-azaspiro[4.4]nonan-3-amin;

N-(2-((4-(6-(sykloheksylmetyl)-2,6-diazaspiro[3.4]oktan-2-yl)pyrimidin-5-yl)oksy)-5-fluorfenyl)-N-etylisobutyramid;

N-benzyl-2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-5-oxa-2-azaspiro[3.4]oktan-7-amin;

5-((7-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-((5-fluor-2'-(prop-1-en-2-yl)-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

2-(5-(4-fluor-2-(2-isopropoksypyridin-3-yl)fenoksy)pyrimidin-4-yl)-6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-((1-metyl-2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

etyl(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)(metyl)karbamat;

N-syklopropyl-5-fluor-N-metyl-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-N-metyl-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4,4]nonan-2-yl)pyrimidin-5-yl)oksy)-N-fenylbenzamid;

2-((4-(6-(sykloheksylmetyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid;

2-(5-(2-(syklopentyloksy)-4-fluorfenoksy)pyrimidin-4-yl)-6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan;

2-syklopropyl-5'-fluor-2'-((4-(7-((2-okso-2,3-dihydrobenso[d]oksazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

metyl(3-((7-(5-((4'-cyano-2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)fenyl)karbamat;

2'-((4-(7-((1H-benzo[d][1,2,3]triazol-6-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-2-syklopropyl-5'-fluor-[1,1'-bifenyl]-4-karbonitril;

N-(2-klor-4-((7-(5-((4'-cyano-2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)fenyl) acetamid;

N,N-dietyl-5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-((2-okso-2,3-dihydrobenso[d]oksazol-5-yl)metyl)-2,7-diazaspiro[4,4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

N-(tert-butyl)-5-fluor-N-metyl-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

1-(7-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)-2-metylpropaN-2-ol;

2-(5-(2-(2-syklopropylpyridin-3-yl)-4-fluorfenoksy)pyrimidin-4-yl)-6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan;

6-((7-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-3,3-dimetylindolin-2-on;

6-((7-(5-(2-(2-syklopropylpyridin-3-yl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-3,3-dimetylindolin-2-on;

5-((7-(5-(2-(2-syklopropylpyridin-3-yl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

4-(((2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-5-oxa-2-azaspiro[3.4]oktan-7-yl)(metyl)amino)metyl)benzonitril;

6-((7-(5-(4-fluor-2-(2,2,2-trifluoroetoksy)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-3,3-dimetylindolin-2-on;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

N-(sykloheksylmetyl)-2-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-5-oxa-2-azaspiro[3.4]oktan-7-amin;

N-(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4,4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)-N-(2-hydroksyetyl)isobutyramid;

N-etyl-N-(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)isobutyramid;

N-(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4,4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)-N-(2,2,2-trifluoretyl)isobutyramid;

N-((1r,4r)-4-((7-(5-((4'-cyano-2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidiN-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)sykloheksyl) acetamid;

tert-butyl((1r,4r)-4-(2-(6-(5-((4'-cyano-2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)etyl)sykloheksyl)karbamat;

5-((7-(5-(4-fluor-2-(5-isopropyltiazol-4-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

N-((1s,4s)-4-((7-(5-((4'-cyano-2'-syklopropyl-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidiN-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)sykloheksyl) acetamid;

2-syklopropyl-2’-((4-(7-((1-etyl-2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5'-fluor-[1,1'-bifenyl]-4-karbonitril;

3-((7-(5-(2-(syklopentyloksy)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-indol-6-karbonitril;

6-((7-(5-(2-(syklopentyloksy)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-3,3-dimetylindolin-2-en;

2-((4-(7-((6-cyano-1H-indol-3-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid;

2-syklopropyl-5'-fluor-2’-((4-(7-(4-(2-oksopyrrolidin-1-yl)benzyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

2-syklopropyl-5'-fluor-2’-((4-(7-((2-oksoindolin-6-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

6-((7-(5-(4-fluor-2-(1-isopropyl-1H-pyrazol-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-benzo[d][1,2,3]triazol;

2-syklopropyl-3’,5'-difluor-2’-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

3-((7-(5-(2-(syklopropylmetoksy)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-indol-6-karboksamid;

3-((7-(5-(2-(syklopropylmetoksy)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1H-indol-6-karbonitril;

2-((4-(7-((3,3-dimetyl-2-oksoindolin-6-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid;

2'-((4-(6-(4-cyanofenetyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)-2-syklopropyl-5'-fluor-[1,1’-bifenyl]-4-karbonitril;

2-syklopropyl-5'-fluor-2’-((4-(7-((2-oksoindolin-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

2-syklopropyl-2’-((4-(7-((3,3-dimetyl-2-oksoindolin-6-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-5'-fluor-[1,1'-bifenyl]-4-karbonitril;

2-amino-2-sykloheksyl-l-(7-(5-(4-fluor-2-(1-isopropyl-1H-pyrazol-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)etanon;

metyl(5-fluor-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)fenyl)(metyl)karbamat;

5-((7-(5-(2-(benzyloksy)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H -benzo[d]imidazol-2-on;

5-((7-(5-(4-fluor-2-metoksyfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-((3-okso-3,4-dihydro-2H-benzo[b][1,4]oksazin-6-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-((7-(5-(4-fluor-2-(2-metylpyrrolidin-1-karbonyl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-(2-((1s,4s)-7-azabicyclo[2.2.1]heptan-7-karbonyl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-((2’-(1,1-difluoretyl)-5-fluor-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

2-syklopropyl-5'-fluor-2’-((4-(6-((4-hydroksytetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

2-syklopropyl-5'-fluor-2’-((4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-((7-(5-(4-fluor-2-(1-isopropyl-3-(trifluormetyl)-1H-pyrazol-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-(4-fluor-2-(2-isopropyl-5-oksopyrrolidin-1-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

(1r,4r)-4-((7-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)sykloheksan-1-amin;

tert-butyl((1r,4r)-4-((7-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)sykloheksyl)karbamat;

N-(4-((7-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)fenyl)acetamid;

5-fluor-N-isopropyl-N-metyl-2-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)benzensulfonamid;

etyl-5'-fluor-2'-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4,4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-2-karboksylat;

5-((7-(5-(4-fluor-2-(4-isopropyltiazol-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-fluor-N-isopropyl-N-metyl-2-((4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5'-fluor-2-metyl-2'-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-4-karbonitril;

4-(2-(6-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)-2-oksoetyl)benzonitril;

4-(2-(6-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)-2-oksoetyl)benzonitril;

1-(6-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan-2-yl)-2-(4-(metylsulfonyl)fenyl)ethan-1-on;

5'-fluor-2-metyl-2'-((4-(7-((2-okso-2,3-dihydro-1H-benzo[d]imidazol-5-yl)metyl)-2,7-diazaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-[1,1'-bifenyl]-3-karbonitril;

2-((3,3-difluorsykloheksyl)metyl)-6-(5-(4-fluor-2-(1-isopropyl-1H-pyrazol-5-yl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan;

2-((3,3-difluorsykloheksyl)metyl)-6-(5-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenoksy)pyrimidin-4-yl)-2,6-diazaspiro[3.3]heptan;

4-(((2-(5-(4-fluor-2-(1-isopropyl-1H-pyrazol-5-yl)fenoksy)pyrimidin-4-yl)-5-oxa-2-azaspiro[3.4]oktan-7-yl)amino)metyl)benzonitril;

5-((7-(5-(2-(2-etylpyridin-3-yl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-(4-fluor-2-isopentylfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-(4-fluor-2-isobutylfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

5-((7-(5-(4-fluor-2-(1-isopropyl-1H-pyrazol-5-yl)fenoksy)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

2-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pyrimidin-4-yl)-6-((tetrahydro-2H-pyran-4-yl))metyl)-2,6-diazaspiro[3.3]heptan;

N-etyl-5-fluor-N-isopropyl-2-((4-(7-(((1r,4r)-4-(metylsulfonamido)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-N,N-diisopropyl-2-((4-(6-((tetrahydro-2H-pyran-4-yl)amino)-2-azaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)benzamid;

5-fluor-N,N-diisopropyl-2-((4-(6-(metyl(tetrahydro-2H-pyran-4-yl)amino)-2-azaspiro[3.3]heptan-2-yl)pyrimidin-5-yl)oksy)benzamid;

tert-butyl((1r,4r)-4-((7-(5-((2-(diisopropylcarbamoyl)-4-fluorfenyl)amino)pyrimidin-4-yl)-2,7-diazaspiro[3.5]nonan-2-yl)metyl)sykloheksyl)karbamat;

1-((6-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)amino)pyrimidin-4-yl)-2,6-diaza spiro[3.3]heptan-2-yl)metyl)sykloheksan-1-ol;

5-((7-(5-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)amino)pyrimidin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

N-(4-fluor-2-(4-isopropylpyrimidin-5-yl)fenyl)-4-(6-((tetrahydro-2H-pyran-4-yl)metyl)-2,6-diazaspiro[3.3]heptaN-2-yl)pyrimidin-5-amin;

N-(5-fluor-2'-isopropoksy-[1,1'-bifenyl]-2-yl)-4-(2-isobutyl-2,7-diazaspiro[3.5]nonan-7-yl)pyrimidin-5-amin;

N-(5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)-4-(2-isobutyl-2,7-diazaspiro[3.5]nonan-7-yl)pyrimidin-5-amin;

N-(2'-etyl-5-fluor-[1,1'-bifenyl]-2-yl)-4-(2-isobutyl-2,7-diazaspiro[3.5]nonan-7-yl)pyrimidin-5-amin;

5-fluor-N,N-diisopropyl-2-((4-(2-(4-(metylsulfonamido)sykloheksyl)-2,7-diazaspiro[3.5]nonan-7-yl)pyrimidin-5-yl)amino)benzamid;

5-((7-(3-((5-fluor-2'-isopropyl-[1,1'-bifenyl]-2-yl)oksy)pyridin-4-yl)-2,7-diazaspiro[4.4]nonan-2-yl)metyl)-1,3-dihydro-2H-benzo[d]imidazol-2-on;

2'-((4-(7-amino-7-benzyl-2-azaspiro[4.4]nonan-2-yl)pyrimidin-5-yl)oksy)-2-syklopropyl-5'-fluor-[1,1'-bifenyl]-4-karbonitril;

tert-butyl((1r,4r)-4-((2-(5-(2-(diisopropylcarbamoyl)-4-fluorfenoksy)pyrimidin-4-yl)-2,7-diazaspiro[3.5]nonan-7-yl)metyl)sykloheksyl)karbamat;

2-((4-(3-(4-acetamidobensyl)-2-amino-4-okso-1,3,7-triazaspiro[4.4]ikke-1-en-7-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropyl-N-metylbenzamid; og

N-etyl-2-((4-(7-(((1r,4r)-4-(etylsulfonamido)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropylbenzamid;

eller et farmasøytisk akseptabelt salt derav.

9. Forbindelsen ifølge krav 1, som er

i) 5-fluor-N,N-diisopropyl-2-((4-(7-(((1r,4r)-4-(metylsulfonamido)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)benzamid, eller et farmasøytisk akseptabelt salt derav, eller

ii)N-etyl-2-((4-(7-(((1r,4r)-4-(etylsulfonamido)sykloheksyl)metyl)-2,7-diazaspiro[3.5]nonan-2-yl)pyrimidin-5-yl)oksy)-5-fluor-N-isopropylbenzamid, eller et farmasøytisk akseptabelt salt derav.

10. Et farmasøytisk akseptabelt salt av forbindelsen ifølge krav 9, som er

i) et bis-metansulfonsyresalt,

ii) et bis-saltsyresalt, eller

iii) et sesquifumarsyresalt.

11. En krystallinsk form av saltet ifølge krav 10, som er

i) i det vesentlige vannfri, eller

ii) hydratisert eller solvatisert, eventuelt hvor den krystallinske formen er et monohydrat.

12. En farmasøytisk sammensetning som omfatter en forbindelse, et farmasøytisk akseptabelt salt eller en krystallinsk form ifølge et hvilket som helst av de foregående kravene og i det minste en farmasøytisk akseptabel bærer.

13. Forbindelsen, det farmasøytisk akseptable saltet, saltet eller den krystallinske formen ifølge et hvilket som helst av kravene 1-11 eller sammensetningen ifølge krav 12 for bruk ved behandling av kreft. for eksempel hvor kreften er: i) en hematologisk kreft, ii) leukemi, iii) lymfom eller iv) der kreften er blandet avstammingsleukemi (MLL), MLL-relatert leukemi, MLL-assosiert leukemi, MLL-positiv leukemi, MLL-indusert leukemi, omorganisert blandet avstammingsleukemi (MLL-r), leukemi som er assosiert med en MLL-omorganisering eller en omorganisering av *MLL* gen, akutt leukemi, kronisk leukemi, indolent leukemi, lymfoblastisk leukemi, lymfocytisk leukemi, myeloid leukemi, myelogen leukemi, barndomsleukemi, akutt lymfocytisk leukemi (ALL), akutt myeloid leukemi (AML), akutt myelogen leukemi, akutt ikke-lymfocytisk leukemi, kronisk lymfocytisk leukemi (CLL), kronisk myelogen leukemi (CML), terapirelatert leukemi, myelodysplastisk syndrom (MDS), myeloproliferativ sykdom (MPD), myeloproliferativ neoplasi (MPN), plasmacelle neoplasma, multippelt myelom, myelodysplasi, kutan T-celle lymfom, lymfoid neoplasm, AIDS-relatert lymfom, thymom, thymuskarsinom, mykos fungus, Alibert-Bazin syndrom, granulom fungus, Sézary syndrom, hårcelleleukemi, T-celle prolymfocytisk leukemi (T-PLL), stor granulær lymfocytisk leukemi, meningeal leukemi, leukemi leptomeningitt, leukemisk meningitt, multippel myelom, Hodgkins lymfom, ikke Hodgkins lymfom (ondartet lymfom) eller Waldenstroms makroglobulinemi.

14. Forbindelsen, det farmasøytisk akseptable saltet eller den krystallinske formen ifølge et hvilket som helst av kravene 1-11 eller sammensetningen ifølge krav 12 for bruk i behandlingen av insulinresistens, pre-diabetes, diabetes eller risiko for diabetes.

15. Forbindelsen, det farmasøytisk akseptable saltet eller den krystallinske formen ifølge et hvilket som helst av kravene 1-11 eller sammensetningen ifølge krav 12 for bruk i behandlingen av hyperglykemi.