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(54)	Title	4,6-DIAMINO-PYRIMIDINE DERIVATIVES AS BMI-1 INHIBITORS FOR TREATING CANCER
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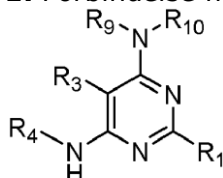
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Enclosed is a translation of the patent claims in Norwegian. Please note that as per the Norwegian Patents Acts, section 66i the patent will receive protection in Norway only as far as there is agreement between the translation and the language of 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/>

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Patentkrav**1. Forbindelse med formel (III):**

(III)

5 eller en form derav, hvori

R₁ er bisyklisk heteroaryl eller bisyklisk heterosyklyl substituert på et karbonatomringelement med én, to, tre eller fire R₅-substituenter eller på et nitrogenatomringelement med en oksygenatoms substituent for å danne et N-oksid; R₃ er halo, hydrogen, cyano, C₁₋₈-alkyl, amino, C₁₋₈-alkyl-amino eller (C₁₋₈-alkyl)₂-amino;

R₄ er fenyl, eventuelt substituert med én, to, tre eller fire R₆-substituenter;

R₅ er uavhengig valgt fra halo, C₁₋₈-alkyl, cyano, hydroksyl, nitro, okso, cyano-C₁₋₈-alkyl, halo-C₁₋₈-alkyl, hydroksyl-C₁₋₈-alkyl, C₁₋₈-alkoksy, C₁₋₈-alkoksy-C₁₋₈-alkyl, halo-C₁₋₈-alkoksy, C₂₋₈-alkenyl, C₁₋₈-alkoksy-C₂₋₈-alkenyl, C₂₋₈-alkynyl, C₁₋₈-alkoksy-C₂₋₈-alkynyl, karboksyl, amino, C₁₋₈-alkyl-amino, (C₁₋₈-alkyl)₂-amino, amino-C₁₋₈-alkyl, C₁₋₈-alkyl-amino-C₁₋₈-alkyl, (C₁₋₈-alkyl)₂-amino-C₁₋₈-alkyl, hydroksyl-C₁₋₈-alkyl-amino, hydroksyl-C₁₋₈-alkyl-amino-C₁₋₈-alkyl, hydroksyl-C₁₋₈-alkyl-amino-C₁₋₈-alkyl-amino, C₁₋₈-alkyl-tio, C₁₋₈-alkyl-karbonyl, C₁₋₈-alkyl-karbonyl-amino, C₁₋₈-alkyl-karbonyl-oksy, C₁₋₈-alkyl-karbonyl-oksy-C₁₋₈-alkyl, C₁₋₈-alkoksy-karbonyl, C₁₋₈-alkoksy-karbonyl-C₁₋₈-alkyl, C₁₋₈-alkoksy-karbonyl-amino, C₁₋₈-alkyl-sulfonyl, C₃₋₁₄-sykloalkyl, aryl, aryl-C₁₋₈-alkyl, aryl-amino, aryl-C₁₋₈-alkyl-amino, heteroaryl, heteroaryl-C₁₋₈-alkyl eller heterosyklyl, hvori C₃₋₁₄-sykloalkyl, aryl, heteroaryl eller heterosyklyl og aryl- og heteroaryldelene av aryl-C₁₋₈-alkyl, aryl-amino, aryl-C₁₋₈-alkyl-amino og heteroaryl-C₁₋₈-alkyl hver eventuelt er substituert med én to, tre eller fire halo, C₁₋₈-alkyl-, halo-C₁₋₈-alkyl-, hydroksyl-C₁₋₈-alkyl-, C₁₋₈-alkoksy-, halo-C₁₋₈-alkoksy-, hydroksyl-C₁₋₈-alkoksy- eller karboksylsubstituenter;

R₆ er uavhengig valgt fra halo-C₁₋₈-alkyl, cyano, halo, nitro, C₁₋₈-alkyl, hydroksyl-C₁₋₈-alkyl, C₁₋₈-alkoksy, halo-C₁₋₈-alkoksy, C₂₋₈-alkenyl, C₁₋₈-alkoksy-C₂₋₈-alkenyl, C₂₋₈-alkynyl, C₁₋₈-alkoksy-C₂₋₈-alkynyl, karboksyl, formel, formyl-oksy, C₁₋₈-alkyl-

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- 5 karbonyl, halo-C₁₋₈alkyl-karbonyl, C₁₋₈alkyl-tio, halo-C₁₋₈alkyl-tio, amino, C₁₋₈alkyl-amino, (C₁₋₈alkyl)₂-amino, C₁₋₈alkyl-karbonyl, C₁₋₈alkyl-karbonyl-oksy, C₁₋₈alkyl-karbonyl-oksy-C₁₋₈alkyl, C₁₋₈alkoksy-karbonyl, halo-C₁₋₈alkoksy-karbonyl, C₁₋₈alkoksy-karbonyl-C₁₋₈alkyl, C₁₋₈alkoksy-karbonyl-amino, C₁₋₈alkoksy-karbonyl-amino-C₁₋₈alkyl, amino-karbonyl, C₁₋₈alkyl-amino-karbonyl, (C₁₋₈alkyl)₂-amino-karbonyl, C₁₋₈alkyl-karbonyl-amino, C₁₋₈alkyl-karbonyl-amino-C₁₋₈alkyl, amino-C₁₋₈alkyl, C₁₋₈alkyl-amino-C₁₋₈alkyl, (C₁₋₈alkyl)₂-amino-C₁₋₈alkyl, amino-C₁₋₈alkyl-amino, C₁₋₈alkyl-amino-C₁₋₈alkyl-amino, (C₁₋₈alkyl)₂-amino-C₁₋₈alkyl-amino, hydroksyl-C₁₋₈alkyl-amino, hydroksyl-C₁₋₈alkyl-amino-C₁₋₈alkyl, hydroksyl-C₁₋₈alkyl-amino-C₁₋₈alkyl-amino, imino-C₁₋₈alkyl, hydroksyl-imino-C₁₋₈alkyl, C₁₋₈alkoksy-imino-C₁₋₈alkyl, C₁₋₈alkyl-sulfonyl, halo-C₁₋₈alkyl-sulfonyl, amino-sulfonyl, C₁₋₈alkyl-amino-sulfonyl, (C₁₋₈alkyl)₂-amino-sulfonyl, B(OR₈)₂, C₃₋₁₄sykloalkyl, heterosyklyl aryl eller heteroaryl, hvori C₃₋₁₄sykloalkyl, heterosyklyl, aryl og heteroaryl hver eventuelt er substituert med én, to, tre eller fire halo eller
- 10 C₁₋₈alkylsubstituent; og,
- 15 R₈ uavhengig er hydrogen eller C₁₋₈alkyl;
- R₉ og R₁₀ er hydrogen; hvori formen av forbindelsen er valgt fra en fri syre, fri base, salt, ester, hydrat, solvat, chelat, clathrat, polymorf, isotopolog, stereoisomer, racemat, enantiomer, diastereomer eller tautomer derav.
- 20
- 2.** Forbindelsen ifølge krav 1 eller en form derav, hvori R₁ er substituert bisyklisk heteroaryl eller bisyklisk heterosyklyl valgt fra 1H-indolyl, 2H-indazolyl, 4,5,6,7-tetrahydro-2H-indazolyl, 1H-benzimidazolyl, imidazo[2,1-b][1,3]tiazolyl, pyrazol[1,5-a]pyridinyl, pyrazol[1,5-c]pyrimidinyl, imidazo[1,2-a]pyridinyl,
- 25 5,6,7,8-tetrahydroimidazo[1,2-a]pyridinyl, 1H-imidazo[4,5-b]pyridinyl, 1H-imidazo[4,5-c]pyridinyl, 4,5,6,7-tetrahydro-3H-imidazo[4,5-c]pyridinyl, imidazo[1,2-a]pyrazinyl, imidazo[1,2-a]pyrimidinyl, 7H-purinyl eller kinolinyl.
- 3.** Forbindelsen ifølge krav 2 eller en form derav, hvori R₁ er substituert bisyklisk heteroaryl valgt fra 1H-indolyl, 1H-benzimidazolyl, pyrazol[1,5-a]pyridinyl, imidazo[1,2-a]pyridinyl, 1H-imidazo[4,5-b]pyridinyl, imidazo[1,2-a]pyrazinyl eller imidazo[1,2-a]pyrimidinyl.
- 30

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4. Forbindelsen ifølge krav 1 eller en form derav, valgt fra gruppen bestående av:

N-[4-(difluormetoksy)fenyl]-2-[2-(trifluormetyl)imidazo[1,2-a]pyridin-3-yl]pyrimidin-4,6-diamin

2-[2-(trifluormetyl)imidazo[1,2-a]pyridin-3-yl]-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

N-[4-(trifluormetoksy)fenyl]-2-[2-(trifluormetyl)imidazo[1,2-a]pyridin-3-yl]pyrimidin-4,6-diamin

2-(6-fluor-2-metylimidazo[1,2-a]pyridin-3-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

N-[4-(difluormetoksy)fenyl]-2-(6-fluor-2-metylimidazo[1,2-a]pyridin-3-yl)pyrimidin-4,6-diamin

2-(2-metyl-1H-imidazo[4,5-b]pyridin-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(5,6-difluor-2-metyl-1H-benzimidazol-1-yl)-N-[3-fluor-4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(2-syklopropyl-6-fluorimidazo[1,2-a]pyridin-3-yl)-N-[4-(difluormetoksy)-3-fluorfenyl]pyrimidin-4,6-diamin

N-[4-(difluormetoksy)-3-fluorfenyl]-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin

N-[4-(difluormetoksy)-3-fluorfenyl]-2-(2-etyl-6-fluor-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin

N-[4-(trifluormetyl)fenyl]-2-(2,5,6-trimetyl-1 H-benzimidazol-1-yl)pyrimidin-4,6-diamin

N-[3-fluor-4-(trifluormetyl)fenyl]-2-(2-metyl-1 H-benzimidazol-1-yl)pyrimidin-4,6-diamin

[3-(4-amino-6-{[4-(trifluormetyl)fenyl]amino}pyrimidin-2-yl)-6-fluorimidazo[1,2-a]pyridin-2-yl]metanol

2-(2,6-dimetyl-1 H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(6-etyl-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(4,6-difluor-2-metyl-1H-benzimidazol-1-yl)-N-[3-fluor-4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

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- 2-(6-etenyl-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 2-(2-syklopropyl-6-fluor-1H-imidazo[4,5-b]pyridin-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 5 2-(6-fluor-2-metyl-1H-imidazo[4,5-b]pyridin-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 2-(6-klor-2-metyl-1H-benzimidazol-1-yl)-N-[3-fluor-4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 2-(5-klor-2-metyl-1H-benzimidazol-1-yl)-N-[3-fluor-4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 10 2-(6-fluor-2-metylimidazo[1,2-a]pyrimidin-3-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- N-[4-(difluormetoksy)-3-fluorfenyl]-2-(5,6-difluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 15 2-(2-syklopropyl-6-fluor-1H-imidazo[4,5-b]pyridin-1-yl)-N-[4-(difluormetoksy)fenyl]pyrimidin-4,6-diamin
- 2-(6-klor-2-metyl-1H-imidazo[4,5-b]pyridin-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 2-(6-klor-2-etyl-1H-imidazo[4,5-b]pyridin-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 20 N-[4-(difluormetoksy)-3-fluorfenyl]-5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- N-[4-(difluormetoksy)-3-fluorfenyl]-5-fluor-2-(5-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 25 2-(6-klor-2-metyl-1H-imidazo[4,5-b]pyridin-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 2-(6-klor-2-metyl-1H-imidazo[4,5-b]pyridin-1-yl)-N-[4-(difluormetoksy)fenyl]-5-fluorpyrimidin-4,6-diamin
- 2-(6-klor-2-metyl-1H-imidazo[4,5-b]pyridin-1-yl)-N-[4-(difluormetoksy)-3-fluorfenyl]-5-fluorpyrimidin-4,6-diamin
- 30 2-(2-syklopropyl-6-fluor-1H-imidazo[4,5-b]pyridin-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

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N-[4-(difluormetoksy)-3-fluorfenyl]-5-fluor-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin

N-[4-(difluormetoksy)-3-fluorfenyl]-2-(2-etyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin

5 N-[4-(difluormetoksy)-3-fluorfenyl]-2-(5,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin

hvor i formen av forbindelsen er valgt fra en fri syre, fri base, salt, ester, hydrat, solvat, chelat, clathrat, polymorf, isotopolog, stereoisomer, racemat, enantiomer, diastereomer eller tautomer derav.

10

5. Forbindelsen eller formen derav ifølge hvilke som helst av de foregående kravene for anvendelse i en fremgangsmåte for å inhibere Bmi-1-funksjon og redusere nivået av Bmi-1 for å behandle kreft mediert av Bmi-1.

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6. Forbindelsen eller formen derav ifølge hvilke som helst av kravene 1 til 4 for anvendelse ifølge krav 5, hvor i fremgangsmåten omfatter å bringe en celle med forhøyede Bmi-1-nivåer fra individet i kontakt med en mengde av forbindelsen eller formen derav, og hvor i cellen er valgt fra en kreftcelle, tumorcelle, kreftstamcelle eller tumorstamcelle, å bestemme en effektiv mengde av forbindelsen eller formen derav, som inhiberer Bmi-1-funksjon i cellen, og deretter å administrere de effektive mengden av forbindelsen eller formen derav til individet.

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7. Forbindelsen eller formen derav ifølge hvilke som helst av kravene 1 til 4 for anvendelse ifølge krav 5, hvor i forbindelsen eller formen derav administreres i en mengde i et område fra ca. 0,001 mg/kg/dag til ca. 500 mg/kg/dag.

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8. Forbindelsen eller formen derav ifølge hvilke som helst av kravene 1 til 4 for anvendelse ifølge krav 5, hvor i forbindelsen eller formen derav administreres i kombinasjon med ett eller flere ytterligere midler valgt fra antikreftmidler, antiproliferative midler, kjemoterapeutiske midler, immunomodulerende midler, anti-angiogene midler, antiinflammatoriske midler, alkyleringsmidler, steroide og ikke-steroid antiinflammatoriske midler, smertestillende midler,

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leukotrienantagonister, P2-agonister, antikolinerge midler, hormonelle midler, biologiske midler, tubulinbindende midler, glukokortikoider, kortikosteroide midler, antibakterielle midler, antihistaminer, antimalarialmidler, antivirale midler og antibiotika; og eventuelt med stråleterapi.

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9. Farmasøytisk sammensetning for anvendelse i behandling av kreft mediert av Bmi-1, omfattende en effektiv mengde av forbindelsen eller formen derav ifølge hvilke som helst av kravene 1 til 4 i blanding med en farmasøytisk akseptabel eksipient.

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10. Forbindelsen eller formen derav ifølge krav 1, valgt fra gruppen bestående av:
2-(5,6-difluor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

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2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(2-etyl-6-fluor-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(6-brom-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

20

2-(4,6-difluor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(6-klor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

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5-fluor-2-(5-fluor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(5,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

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2-(4,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(2-etyl-4,6-difluor-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

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- 5-fluor-2-(4-fluor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
 2-(6-klor-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
 5 2-(2-etyl-5-fluor-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
 2-(2-etyl-6-fluor-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
 10 N-(4-klorfenyl)-5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 N-(3-klorfenyl)-5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 N-(4-klorfenyl)-2-(4,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin
 15 2-(5,7-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
 N-(4-klorfenyl)-5-fluor-2-(5-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 N-(4-klorfenyl)-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 20 **11.** Forbindelsen eller formen derav ifølge krav 1, valgt fra gruppen bestående av:
 N-[4-(difluormetoksy)fenyl]-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 25 2-(6-klor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)fenyl]pyrimidin-4,6-diamin
 2-(5-klor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)fenyl]pyrimidin-4,6-diamin
 N-[4-(difluormetoksy)fenyl]-2-(5,6-difluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 30 N-[4-(difluormetoksy)fenyl]-5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 N-[4-(difluormetoksy)fenyl]-5-fluor-2-(5-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin

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N-[4-(difluormetoksy)fenyl]-2-(4,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin

N-[4-(difluormetoksy)fenyl]-2-(2-etyl-4,6-difluor-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin

5 N-[4-(difluormetoksy)fenyl]-2-(5,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin

N-[4-(difluormetoksy)fenyl]-5-fluor-2-(4-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin

10 N-[4-(difluormetoksy)fenyl]-2-(2-etyl-6-fluor-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin

2-(5-klor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)fenyl]-5-fluorpyrimidin-4,6-diamin

2-(6-klor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)fenyl]-5-fluorpyrimidin-4,6-diamin

15 N-[4-(difluormetoksy)fenyl]-2-(2-etyl-5-fluor-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin

N-[4-(difluormetoksy)fenyl]-2-(2-etyl-6-fluor-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin

20 5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetoksy)fenyl]pyrimidin-4,6-diamin

2-(4,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetoksy)fenyl]pyrimidin-4,6-diamin

5-fluor-2-(5-fluor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetoksy)fenyl]pyrimidin-4,6-diamin

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12. Forbindelsen eller formen derav ifølge krav 1, valgt fra gruppen bestående av:

2-(2-syklopropyl-6-fluor-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)-3-fluorfenyl]pyrimidin-4,6-diamin

30 2-(2-syklopropyl-6-fluor-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(2-syklopropyl-5-fluor-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

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2-(2-syklopropyl-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(2-syklopropyl-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)fenyl]-5-fluorpyrimidin-4,6-diamin

5 2-(2-syklopropyl-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)-3-fluorfenyl]-5-fluorpyrimidin-4,6-diamin

2-(2-syklopropyl-4-fluor-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

10 2-(2-syklopropyl-4-fluor-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)fenyl]-5-fluorpyrimidin-4,6-diamin

2-(2-syklopropyl-4-fluor-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)-3-fluorfenyl]-5-fluorpyrimidin-4,6-diamin

2-(2-syklopropyl-6-fluor-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)fenyl]pyrimidin-4,6-diamin

15 2-(2-syklopropyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

2-(2-syklopropyl-1H-benzimidazol-1-yl)-5-fluor-N-(4-metylfenyl)pyrimidin-4,6-diamin

2-(2-syklopropyl-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetoksy)fenyl]pyrimidin-4,6-diamin

20 N-(4-klorfenyl)-2-(2-syklopropyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin

N-(4-klor-3-fluorfenyl)-2-(2-syklopropyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin

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13. Forbindelsen eller formen derav ifølge krav 1, valgt fra gruppen bestående av:

2-(2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
5-fluor-2-(2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

30 N-[4-(difluormetoksy)fenyl]-5-fluor-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin

2-(2-etyl-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin

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- N-[4-(difluormetoksy)fenyl]-2-(2-etyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin
- 2-(2-etyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 5 N-[4-(difluormetoksy)fenyl]-2-(2-etyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 5-klor-2-(2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 5-klor-2-(2-etyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 10 5-klor-N-[4-(difluormetoksy)fenyl]-2-(2-etyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 5-metyl-2-(2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 15 N-(4-klorfenyl)-5-fluor-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin 5-fluor-2-(2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetoksy)fenyl]pyrimidin-4,6-diamin
- N-(4-klorfenyl)-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin N-(4-bromfenyl)-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 20 N-(4-klorfenyl)-2-(5,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin
- 2-(2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetoksy)fenyl]pyrimidin-4,6-diamin N-[4-(difluormetoksy)fenyl]-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 25 2-(2-etyl-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetoksy)fenyl]pyrimidin-4,6-diamin
- N-(4-klorfenyl)-2-(2-etyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin.
- 14.** Forbindelsen eller formen derav ifølge krav 1, valgt fra gruppen bestående av:
- 30 N-[4-(difluormetoksy)-3-fluorfenyl]-2-(4,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin
- N-[4-(difluormetoksy)-3-fluorfenyl]-2-(2-etyl-4,6-difluor-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin

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- N-[4-(difluormetoksy)-3-fluorfenyl]-5-fluor-2-(4-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 5-fluor-2-(2-metyl-1H-imidazo[4,5-b]pyridin-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 5 N-[4-(difluormetoksy)fenyl]-5-fluor-2-(2-metyl-1H-imidazo[4,5-b]pyridin-1-yl)pyrimidin-4,6-diamin
 N-[4-(difluormetoksy)-3-fluorfenyl]-5-fluor-2-(2-metyl-1H-imidazo[4,5-b]pyridin-1-yl)pyrimidin-4,6-diamin
- 10 N-[4-(difluormetoksy)fenyl]-2-[2-(difluormetyl)-6-fluor-1H-benzimidazol-1-yl]pyrimidin-4,6-diamin
 2-(2-metyl-1H-benzimidazol-1-yl)-N-[4-(metylsulfanyl)fenyl]pyrimidin-4,6-diamin 2-[2-(1-metylsyklopropyl)-1H-benzimidazol-1-yl]-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 15 N-[4-(difluormetoksy)fenyl]-2-[2-(1-metylsyklopropyl)-1H-benzimidazol-1-yl]pyrimidin-4,6-diamin
 2-[2-(metoksymetyl)-1H-benzimidazol-1-yl]-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 2-[2-(propan-2-yl)-1H-benzimidazol-1-yl]-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 20 2-[2-(difluormetyl)-1H-benzimidazol-1-yl]-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
 2-(2-metylpyrazol[1,5-a]pyridin-3-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 2-(2-syklopropyl-6-fluor-1H-imidazo[4,5-b]pyridin-1-yl)-N-[4-(difluormetoksy)fenyl]-5-fluorpyrimidin-4,6-diamin
- 25 N-[4-(difluormetoksy)-3-fluorfenyl]-2-(2-etyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 2-(5-klor-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 30 5-fluor-2-(2-metylimidazo[1,2-a]pyridin-3-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
 N-[4-(difluormetoksy)fenyl]-5-fluor-2-(2-metylimidazo[1,2-a]pyridin-3-yl)pyrimidin-4,6-diamin

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- N-[4-(difluormetoksy)-3-fluorfenyl]-5-fluor-2-(2-metylimidazo[1,2-a]pyridin-3-yl)pyrimidin-4,6-diamin
- N-[4-(difluormetoksy)fenyl]-5-fluor-2-(2-metylpyrazol[1,5-a]pyridin-3-yl)pyrimidin-4,6-diamin
- 5 5-fluor-N-[4-(trifluormetyl)fenyl]-2-(2,6,8-trimetylimidazo[1,2-a]pyrazin-3-yl)pyrimidin-4,6-diamin
- N-[4-(difluormetoksy)fenyl]-5-fluor-2-(2,6,8-trimetylimidazo[1,2-a]pyrazin-3-yl)pyrimidin-4,6-diamin
- 10 N-[4-(difluormetoksy)-3-fluorfenyl]-5-fluor-2-(2,6,8-trimetylimidazo[1,2-a]pyrazin-3-yl)pyrimidin-4,6-diamin
- 5-fluor-2-(2-metylpyrazol[1,5-a]pyridin-3-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 5-fluor-2-[6-fluor-2-(trifluormetyl)imidazo[1,2-a]pyridin-3-yl]-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 15 5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
- 2-(2-syklopropylpyrazol[1,5-a]pyridin-3-yl)-N-[4-(difluormetoksy)fenyl]-5-fluorpyrimidin-4,6-diamin
- [3-(4-amino-6-{[4-(difluormetoksy)fenyl]amino})-5-fluorpyrimidin-2-yl]-5-fluorpyrazol[1,5-a]pyridin-2-yl]metanol
- 20 2-[2-(metylsulfanyl)-1 H-benzimidazol-1-yl]-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)-N-(4-metylfenyl)pyrimidin-4,6-diamin
- 25 N-[4-(difluormetoksy)fenyl]-2-(2-etyl-5-fluorpyrazol[1,5-a]pyridin-3-yl)-5-fluorpyrimidin-4,6-diamin
- 2-(2-etyl-5-fluorpyrazol[1,5-a]pyridin-3-yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- N-[4-(difluormetoksy)-3-fluorfenyl]-2-(2-etyl-5-fluorpyrazol[1,5-a]pyridin-3-yl)-5-fluorpyrimidin-4,6-diamin
- 30 5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)-N-(3-metoksyfenyl)pyrimidin-4,6-diamin

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- 4-{[6-amino-5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4-yl]amino}benzonitril
metyl 4-{[6-amino-5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4-yl]amino}benzoat
- 5 5-fluor-2-(2-metyl-1H-benzimidazol-1-yl)-N-(3-metylfenyl)pyrimidin-4,6-diamin
5-fluor-N-(3-metoksyfenyl)-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
2-(2-metyl-1H-benzimidazol-1-yl)-N-(4-metylfenyl)pyrimidin-4,6-diamin
N-(4-metoksyfenyl)-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 10 N-[4-(dimetylamino)fenyl]-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
5-fluor-2-(2-metyl-1H-benzimidazol-1-yl)-N-(4-metylfenyl)pyrimidin-4,6-diamin
5-fluor-N-(4-metoksyfenyl)-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 15 4-{[6-amino-5-fluor-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4-yl]amino}benzonitril
N-(3-fluor-4-metoksyfenyl)-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 20 4-{[6-amino-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4-yl]amino}benzonitril
2-(2-metyl-1H-benzimidazol-1-yl)-N-(4-nitrofenyl)pyrimidin-4,6-diamin
2-(5,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-(4-metylfenyl)pyrimidin-4,6-diamin
2-(5,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
- 25 N-[4-(difluormetoksy)fenyl]-5-fluor-2-[2-(metoksymetyl)-1H-benzimidazol-1-yl]pyrimidin-4,6-diamin
5-fluor-2-[2-(metoksymetyl)-1 H-benzimidazol-1-yl]-N-(4-metylfenyl)pyrimidin-4,6-diamin
5-fluor-2-[2-(metoksymetyl)-1 H-benzimidazol-1-yl]-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
- 30 N-(4-klorfenyl)-5-fluor-2-[2-(metoksymetyl)-1H-benzimidazol-1-yl]pyrimidin-4,6-diamin

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- 5-fluor-2-[2-(metoksymetyl)-1 H-benzimidazol-1 -yl]-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 1-(4-amino-5-fluor-6-{[4-(trifluormetyl)fenyl]amino}pyrimidin-2-yl)-2-metyl-1H-benzimidazol-6-karbonitril
- 5 1-(4-amino-6-{[4-(difluormetoksy)fenyl]amino}-5-fluorpyrimidin-2-yl)-2-metyl-1H-benzimidazol-6-karbonitril
- 1-{4-amino-5-fluor-6-[(4-metylfenyl)amino]pyrimidin-2-yl}-2-metyl-1H-benzimidazol-6-karbonitril
- 10 1-{4-amino-5-fluor-6-[(4-metoksyfenyl)amino]pyrimidin-2-yl}-2-metyl-1H-benzimidazol-6-karbonitril
- 1-(4-amino-6-{[4-(difluormetoksy)fenyl]amino}-5-fluorpyrimidin-2-yl)-2-metyl-1H-benzimidazol-5-karbonitril
- 1-{4-amino-5-fluor-6-[(4-metylfenyl)amino]pyrimidin-2-yl}-2-metyl-1H-benzimidazol-5-karbonitril
- 15 1-{4-amino-5-fluor-6-[(4-metoksyfenyl)amino]pyrimidin-2-yl}-2-metyl-1H-benzimidazol-5-karbonitril
- 1-(4-amino-5-fluor-6-{[4-(trifluormetyl)fenyl]amino}pyrimidin-2-yl)-2-metyl-1H-benzimidazol-5-karbonitril
- 1-{4-amino-6-[(4-klorfenyl)amino]-5-fluorpyrimidin-2-yl}-2-metyl-1H-benzimidazol-5-karbonitril
- 20 N-[4-(difluormetoksy)fenyl]-5-fluor-2-(2-metyl-6-nitro-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 5-fluor-2-(2-metyl-6-nitro-1H-benzimidazol-1-yl)-N-(4-metylfenyl)pyrimidin-4,6-diamin
- 25 5-fluor-N-(4-metoksyfenyl)-2-(2-metyl-6-nitro-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- N-[4-(difluormetoksy)-3-fluorfenyl]-2-(2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 5-fluor-2-(2-metyl-6-nitro-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- 30 5-fluor-N-(4-metoksyfenyl)-2-(2-metylpyrazol[1,5-a]pyridin-3-yl)pyrimidin-4,6-diamin

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- 2-(4,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-(4-metylfenyl)pyrimidin-4,6-diamin
- 2-(4,6-difluor-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
- 5 5-fluor-2-(6-metoksy-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin
- N-(4-klorfenyl)-5-fluor-2-(6-metoksy-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 5-fluor-2-(6-metoksy-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetoksy)fenyl]pyrimidin-4,6-diamin
- 10 N-[4-(difluormetoksy)fenyl]-5-fluor-2-(6-metoksy-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 5-fluor-2-(6-metoksy-2-metyl-1H-benzimidazol-1-yl)-N-(4-metylfenyl)pyrimidin-4,6-diamin
- 15 5-fluor-2-(6-metoksy-2-metyl-1H-benzimidazol-1-yl)-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
- N-(4-klorfenyl)-5-fluor-2-(6-fluor-2-metylimidazo[1,2-a]pyridin-3-yl)pyrimidin-4,6-diamin
- 5-fluor-2-(6-fluor-2-metylimidazo[1,2-a]pyridin-3-yl)-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
- 20 5-fluor-2-(6-fluor-2-metylimidazo[1,2-a]pyridin-3-yl)-N-(4-metylfenyl)pyrimidin-4,6-diamin
- 5-fluor-2-(2-metyl-5-nitro-1H-benzimidazol-1-yl)-N-(4-metylfenyl)pyrimidin-4,6-diamin
- 25 5-fluor-N-(4-metoksyfenyl)-2-(2-metyl-5-nitro-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
- 2-(6-amino-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
- 2-(6-amino-2-metyl-1H-benzimidazol-1-yl)-N-[4-(difluormetoksy)fenyl]-5-fluorpyrimidin-4,6-diamin
- 30 2-(6-amino-2-metyl-1H-benzimidazol-1-yl)-5-fluor-N-(4-metylfenyl)pyrimidin-4,6-diamin

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- N-(4-klor-3-fluorfenyl)-5-fluor-2-(5-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 N-(4-klor-3-fluorfenyl)-5-fluor-2-(2-metyl-1 H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 5 N-(4-klor-3-fluorfenyl)-5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)-N-(4-metylfenyl)pyrimidin-4,6-diamin
 2-(6-fluor-2-metyl-1 H-benzimidazol-1-yl)-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
 10 N-[4-(dimetylamino)fenyl]-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 N-(4-klor-3-fluorfenyl)-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)pyrimidin-4,6-diamin
 2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)-N-(3-metylfenyl)pyrimidin-4,6-diamin
 15 2-(2-etyl-1H-benzimidazol-1-yl)-5-fluor-N-(4-metylfenyl)pyrimidin-4,6-diamin
 2-(2-etyl-1H-benzimidazol-1-yl)-5-fluor-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
 N-(4-klor-3-fluorfenyl)-2-(2-etyl-1H-benzimidazol-1-yl)-5-fluorpyrimidin-4,6-diamin
 2-(2-syklopropyl-1H-benzimidazol-1-yl)-5-fluor-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
 20 5-fluor-2-(5-fluor-2-metyl-1H-benzimidazol-1-yl)-N-(4-metylfenyl)pyrimidin-4,6-diamin
 5-fluor-2-(5-fluor-2-metyl-1 H-benzimidazol-1-yl)-N-(4-metoksyfenyl)pyrimidin-4,6-diamin
 25 2-(6-amino-2-metyl-1H-benzimidazol-1 -yl)-5-fluor-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin;

- 15.** Forbindelsen eller formen derav ifølge hvilke som helst av kravene 1-10, hvori forbindelsen er 5-fluor-2-(6-fluor-2-metyl-1H-benzimidazol-1-yl)-N-[4-(trifluormetyl)fenyl]pyrimidin-4,6-diamin eller en form derav.
 30