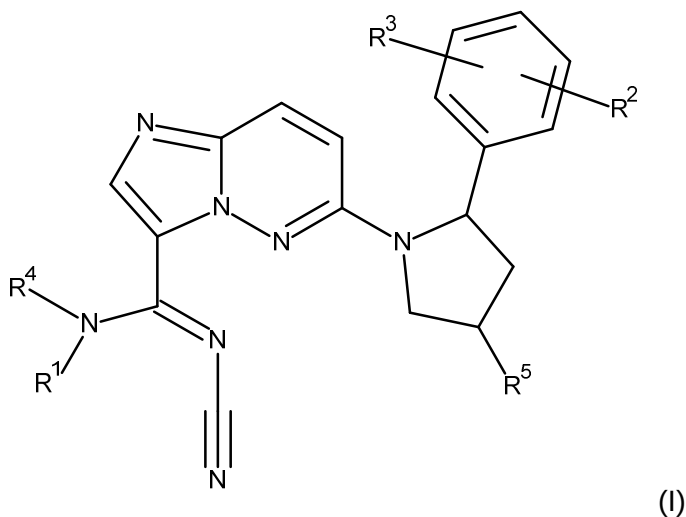


CLAIMS

1. A compound of Formula (I):



or a pharmaceutically acceptable salt or solvate thereof;

wherein:

R¹ is selected from H, -XR⁷, (C₁-C₆)alkyl, (C₃-C₈)cycloalkyl, and a C-linked 4-6 membered heterocycloalkyl containing 1 to 2 heteroatoms selected from N, O and

10 S;

X is selected from -CH₂-;

R² is selected from H and -SR⁶;

R³ is selected from H and halo;

R⁴ is selected from H and (C₁-C₃)alkyl

15 R⁵ is selected from H and halo;

R⁶ is methyl;

R⁷ is phenyl substituted by hydroxy wherein the hydroxyphenyl is optionally further substituted by halo;

provided that if R² is H then R¹ is XR⁷.

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2. A compound according to claim 1 wherein R¹ is selected from -XR⁷, (C₁-C₆)alkyl, (C₃-C₈)cycloalkyl, and a C-linked 4-6 membered heterocycloalkyl containing 1 to 2 heteroatoms selected from N, O and S.

3. A compound according to claim 1 or 2 wherein R¹ is selected from (C₁-C₆)alkyl and (C₃-C₈)cycloalkyl.

4. A compound according to claim 3 wherein R¹ is (C₁-C₆)alkyl.

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5. A compound according to claim 1 or 2 wherein R¹ is selected from -XR⁷ and a C-linked 4-6 membered heterocycloalkyl containing 1 to 2 heteroatoms selected from N, O and S.

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6. A compound according to claim 5 wherein R¹ is selected from -XR⁷ and a C-linked 4-6 membered heterocycloalkyl containing 1 to 2 heteroatoms selected from N and O.

15 7. A compound according to any preceding claim wherein R² is -SR⁶.

8. A compound according to any preceding claim wherein R³ is H or fluoro.

9. A compound according to any preceding claim wherein R⁴ is H.

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10. A compound according to any preceding claim wherein R⁵ is H or fluoro.

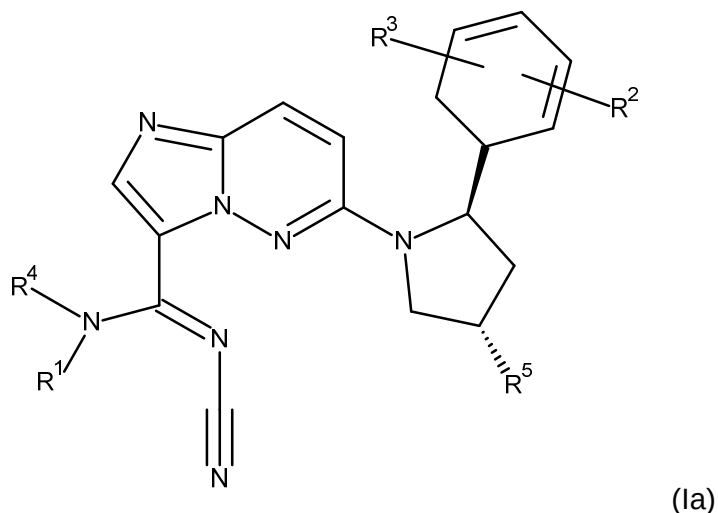
11. A compound according to any preceding claim wherein R⁷ is phenyl substituted by hydroxy wherein the hydroxyphenyl is optionally further substituted by fluoro.

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12. A compound of Formula Ia:



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or a pharmaceutically salt or solvate thereof, wherein R¹, R², R³, R⁴ and R⁵ are as defined in any preceding claim.

13. A compound according to claim wherein said compound is selected from:

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N'-cyano-6-[2-[5-fluoro-2-(methylsulfanyl)phenyl]pyrrolidin-1-yl]-N-[(3R)-oxan-3-yl]imidazo[1,2-b]pyridazine-3-carboximidamide;

N'-cyano-6-[2-[5-fluoro-2-(methylsulfanyl)phenyl]pyrrolidin-1-yl]-N-[(oxan-3-yl]imidazo[1,2-b]pyridazine-3-carboximidamide;

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N'-cyano-N-ethyl-6-[4-fluoro-2-[5-fluoro-3-(methylsulfanyl)phenyl]pyrrolidin-1-yl]imidazo[1,2-b]pyridazine-3-carboximidamide;

N'-cyano-N-ethyl-6-[4-fluoro-2-[5-fluoro-2-(methylsulfanyl)phenyl]pyrrolidin-1-yl]imidazo[1,2-b]pyridazine-3-carboximidamide;

N-butyl-N'-cyano-6-[4-fluoro-2-[5-fluoro-2-(methylsulfanyl)phenyl]pyrrolidin-1-yl]imidazo[1,2-b]pyridazine-3-carboximidamide;

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N'-cyano-N-cyclohexyl-6-[4-fluoro-2-[5-fluoro-2-(methylsulfanyl)phenyl]pyrrolidin-1-yl]imidazo[1,2-b]pyridazine-3-carboximidamide; and

N'-cyano-6-[2-[5-fluoro-2-(methylsulfanyl)phenyl]pyrrolidin-1-yl]-N-[(3-hydroxyphenyl)methyl]imidazo[1,2-b]pyridazine-3-carboximidamide;

or a pharmaceutically acceptable salt or solvate thereof.

14. A compound according to claim wherein said compound is selected from:

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(Z)-N'-cyano-6-[(2R)-2-[5-fluoro-2-(methylsulfanyl)phenyl]pyrrolidin-1-yl]-N-[(3R)-oxan-3-yl]imidazo[1,2-b]pyridazine-3-carboximidamide ;

(Z)-N'-cyano-6-[(2R)-2-[5-fluoro-2-(methylsulfanyl)phenyl]pyrrolidin-1-yl]-N-[(3S)-oxan-3-yl]imidazo[1,2-b]pyridazine-3-carboximidamide;

10

(Z)-N'-cyano-N-ethyl-6-[(2R,4S)-4-fluoro-2-[5-fluoro-3-(methylsulfanyl)phenyl]pyrrolidin-1-

yl]imidazo[1,2-b]pyridazine-3-carboximidamide;

(Z)-N'-cyano-N-ethyl-6-[(2R,4S)-4-fluoro-2-[5-fluoro-2-

(methylsulfanyl)phenyl]pyrrolidin-1-yl]imidazo[1,2-b]pyridazine-3-carboximidamide;

15

(Z)-N-butyl-N'-cyano-6-[(2R,4S)-4-fluoro-2-[5-fluoro-2-

(methylsulfanyl)phenyl]pyrrolidin-1-yl]imidazo[1,2-b]pyridazine-3-carboximidamide;

(Z)-N'-cyano-N-cyclohexyl-6-[(2R,4S)-4-fluoro-2-[5-fluoro-2-

(methylsulfanyl)phenyl]pyrrolidin-1-yl]imidazo[1,2-b]pyridazine-3-carboximidamide;

and

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(Z)-N'-cyano-6-[(2R)-2-[5-fluoro-2-(methylsulfanyl)phenyl]pyrrolidin-1-yl]-N-[(3-hydroxyphenyl)methyl]imidazo[1,2-b]pyridazine-3-carboximidamide;

or a pharmaceutically acceptable salt or solvate thereof.

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15. A pharmaceutical composition comprising a compound according to any of claims 1 to 14 and one or more pharmaceutically acceptable excipients.

16. A pharmaceutical composition according to claim 15 in combination with one or more other therapeutic agents.

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17. A compound according to any of claims 1 to 14 for use as a pharmaceutical.

18. A compound as defined in any of the preceding claims for use in treating or preventing a condition or disorder which is mediated by Trk, wherein the condition or disorder mediated by Trk is selected from: diseases of pruritus and itch; autoimmune diseases of the skin; diseases of skin pain and neuropathy; and
5 diseases of dermatitis.

19. A compound ~~according to claim 18~~ for use according to claim 18 wherein the condition or disorder is mediated by TrkA, TrkB, and TrkC.

10 20. A compound ~~according to claim 18 or 19~~ for use according to claim 18 or 19 wherein the condition or disorder is atopic dermatitis.

15 21. Use of a compound as defined in any of claims 1 to 14 in the manufacture of a medicament for treating or preventing a condition or disorder which is mediated by Trk, wherein the condition or disorder mediated by Trk is selected from: diseases of pruritus and itch; autoimmune diseases of the skin; diseases of skin pain and neuropathy; and diseases of dermatitis.

20 22. Use according to claim 21 wherein the condition or disorder is mediated by TrkA, TrkB, and TrkC.

23. Use according to claim 21 or 22 wherein the condition or disorder is atopic dermatitis.