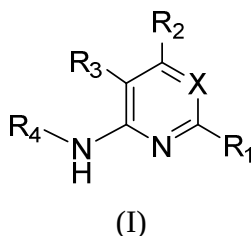


CLAIMS

1. A compound of Formula (I):



or a form thereof, wherein

R₁ is monocyclic or bicyclic heteroaryl substituted on a carbon atom ring member with one, two, three or four R₅ substituents, or on a nitrogen atom ring member with an oxygen atom substituent to form an N-oxide;

X is N or N substituted with an oxygen atom substituent to form an N-oxide;

R₂ is amino;

R₃ is hydrogen, cyano, halo, C₁₋₈alkyl, amino, C₁₋₈alkyl-amino or (C₁₋₈alkyl)₂-amino;

R₄ is phenyl, optionally substituted with one, two, three or four R₆ substituents;

R₅ is independently selected from cyano, halo, hydroxyl, nitro, oxo, C₁₋₈alkyl, cyano-C₁₋₈alkyl, halo-C₁₋₈alkyl, hydroxyl-C₁₋₈alkyl, C₁₋₈alkoxy, C₁₋₈alkoxy-C₁₋₈alkyl, halo-C₁₋₈alkoxy, C₂₋₈alkenyl, C₁₋₈alkoxy-C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy-C₂₋₈alkynyl, carboxyl, amino, C₁₋₈alkyl-amino, (C₁₋₈alkyl)₂-amino, amino-C₁₋₈alkyl, C₁₋₈alkyl-amino-C₁₋₈alkyl, (C₁₋₈alkyl)₂-amino-C₁₋₈alkyl, hydroxyl-C₁₋₈alkyl-amino, hydroxyl-C₁₋₈alkyl-amino-C₁₋₈alkyl, hydroxyl-C₁₋₈alkyl-amino-C₁₋₈alkyl-amino, C₁₋₈alkyl-thio, C₁₋₈alkyl-carbonyl, C₁₋₈alkyl-carbonyl-amino, C₁₋₈alkyl-carbonyl-oxy, C₁₋₈alkyl-carbonyl-oxy-C₁₋₈alkyl, C₁₋₈alkoxy-carbonyl, C₁₋₈alkoxy-carbonyl-C₁₋₈alkyl, C₁₋₈alkoxy-carbonyl-amino, C₁₋₈alkyl-sulfonyl, C₃₋₁₄cycloalkyl, aryl, aryl-C₁₋₈alkyl, aryl-amino, aryl-C₁₋₈alkyl-amino, heteroaryl, heteroaryl-C₁₋₈alkyl or heterocyclyl, wherein C₃₋₁₄cycloalkyl, aryl, heteroaryl or heterocyclyl and the aryl and heteroaryl portions of aryl-C₁₋₈alkyl, aryl-amino, aryl-C₁₋₈alkyl-amino and heteroaryl-C₁₋₈alkyl are each optionally substituted with one, two, three or four halo, C₁₋₈alkyl, halo-C₁₋₈alkyl, hydroxyl-C₁₋₈alkyl, C₁₋₈alkoxy, halo-C₁₋₈alkoxy, hydroxyl-C₁₋₈alkoxy or carboxyl substituents;

R₆ is independently selected from cyano, halo, hydroxyl, nitro, C₁₋₈alkyl, halo-C₁₋₈alkyl, hydroxyl-C₁₋₈alkyl, C₁₋₈alkoxy, halo-C₁₋₈alkoxy, C₂₋₈alkenyl, C₁₋₈alkoxy-C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy-C₂₋₈alkynyl, carboxyl, formyl, formyl-oxy, C₁₋₈alkyl-carbonyl, halo-C₁₋₈alkyl-carbonyl, C₁₋₈alkyl-thio, halo-C₁₋₈alkyl-thio, amino, C₁₋₈alkyl-amino, (C₁₋₈alkyl)₂-amino, C₁₋₈alkyl-carbonyl, C₁₋₈alkyl-carbonyl-oxy, C₁₋₈alkyl-carbonyl-oxy-C₁₋₈alkyl, C₁₋₈alkoxy-carbonyl, halo-C₁₋₈alkoxy-carbonyl, C₁₋₈alkoxy-carbonyl-C₁₋₈alkyl, C₁₋₈alkoxy-carbonyl-amino, C₁₋₈alkoxy-carbonyl-amino-C₁₋₈alkyl, amino-carbonyl, C₁₋₈alkyl-amino-carbonyl, (C₁₋₈alkyl)₂-amino-carbonyl, C₁₋₈alkyl-carbonyl-amino, C₁₋₈alkyl-carbonyl-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, hydroxyl-C₁₋₈alkyl-amino, hydroxyl-C₁₋₈alkyl-amino-C₁₋₈alkyl, hydroxyl-C₁₋₈alkyl-amino-C₁₋₈alkyl-amino, imino-C₁₋₈alkyl, hydroxyl-imino-C₁₋₈alkyl, C₁₋₈alkoxy-imino-C₁₋₈alkyl, C₁₋₈alkyl-sulfonyl, halo-C₁₋₈alkyl-sulfonyl, amino-sulfonyl, C₁₋₈alkyl-amino-sulfonyl, (C₁₋₈alkyl)₂-amino-sulfonyl, B(OR₈)₂, C₃₋₁₄cycloalkyl, heterocyclyl aryl or heteroaryl, wherein C₃₋₁₄cycloalkyl, heterocyclyl, aryl, and heteroaryl are each optionally substituted with one, two, three or four halo or C₁₋₈alkyl substituents;

and,

R₈ is independently hydrogen or C₁₋₈alkyl;

wherein the form of the compound is selected from a free acid, free base, salt, ester, hydrate, solvate, racemate, enantiomer, diastereomer or tautomer thereof.

2. The compound of claim 1, wherein R₁ is substituted monocyclic or bicyclic heteroaryl selected from 1H-pyrazolyl, 1H-imidazolyl, 1,2-oxazolyl, pyridinyl, 1H-indolyl, 2H-indazolyl, 1H-benzimidazolyl, imidazo[2,1-b][1,3]thiazolyl, pyrazolo[1,5-a]pyridinyl, pyrazolo[1,5-c]pyrimidinyl, imidazo[1,2-a]pyridinyl, 1H-imidazo[4,5-b]pyridinyl, 1H-imidazo[4,5-c]pyridinyl, imidazo[1,2-a]pyrazinyl, imidazo[1,2-a]pyrimidinyl, 7H-purinyl or quinolinyl.
3. The compound of claim 2, wherein R₁ is substituted monocyclic or bicyclic heteroaryl selected from 1H-pyrazolyl, 1H-indolyl, 1H-benzimidazolyl, pyrazolo[1,5-a]pyridinyl,

imidazo[1,2-a]pyridinyl, 1H-imidazo[4,5-b]pyridinyl, imidazo[1,2-a]pyrazinyl or imidazo[1,2-a]pyrimidinyl.

4. An effective amount of the compound of any one of claims 1 to 3, for use in the treatment of cancer mediated by Bmi-1, by inhibiting Bmi-1 function and reducing the level of Bmi-1 in a subject.
5. The use of the compound of any one of claims 1 to 3, in the manufacture of a medicament to treat a cancer mediated by Bmi-1 in a subject wherein the medicament is formulated to inhibit Bmi-1 function and reduce the level of Bmi-1 in the subject.
6. A pharmaceutical composition for use in treating a cancer mediated by Bmi-1 comprising an effective amount of the compound of any one of claims 1 to 3, in admixture with a pharmaceutically acceptable excipient.