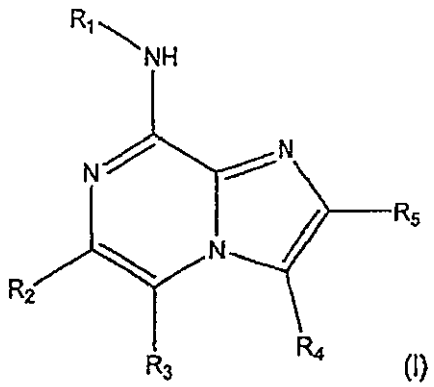
WHAT IS CLAIMED:

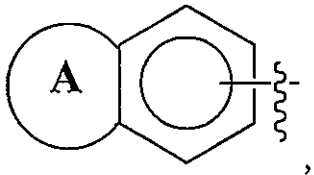
1. At least one chemical entity chosen from compounds of Formula (I):



(I)

or a pharmaceutically acceptable salt thereof, wherein

R_1 is chosen from (2-methyl-2-hydroxypropoxy)pyridin-6-yl, (2-methoxyethoxy)pyridinyl, 2-(dimethylamino)ethoxy-3-pyridinyl, hydroxyethoxy-5-pyridinyl, (3-methyl-3-hydroxyazetidin-1-yl)pyridin-3-yl, (3-methyl-3-hydroxyazetidin-1-yl)pyridin-2-yl, (3-hydroxyazetidin-1-yl)pyridin-2-yl, (hydroxy(dimethylethyl))-5-pyridinyl, (4-methyl-4-hydroxypiperidin-1-yl)pyridin-2-yl, (3-methyl-3-hydroxypiperidin-1-yl)pyridin-2-yl, 5-morpholinopyridin-2-yl, 6-morpholinopyridin-3-yl, ((2-methoxyethyl)(methyl)amino)pyridin-5-yl, ((2-hydroxyethyl)(methyl)amino)pyridin-5-yl, 2-methoxy-4-pyridinyl, 2-hydroxy-5-pyridinyl, (2-hydroxyethyl)-1H-pyrazol-4-yl, (2-hydroxypropyl)-1H-pyrazol-4-yl, (2-methoxyethyl)-1H-pyrazol-4-yl, 1-ethyl-pyrazol-4-yl, 1-isopropyl-pyrazol-4-yl, 3-cyclopropyl-1H-pyrazol-5-yl, and 1-ethyl-5-methyl-pyrazol-3-yl; or

R_1 is 

, wherein A is an optionally substituted heteroaryl group having from 5 to 7 ring atoms including the atoms shared with the 6 membered aromatic ring;

R_2 is chosen from 2,3-dimethyl-2H-indazol-6-yl, 1H-indazolyl-6-yl, 1-methyl-1H-indazol-5-yl, 1-methyl-1H-indazol-6-yl, 3,4-dihydro-2H-1,4-benzoxazin-3-one-6-yl, 1,3-benzoxazol-6-yl, 3-aminoquinolin-6-yl, 2,3-dihydro-1H-indol-6-yl, benzothiazolyl, 2-

aminoquinazolin-6-yl, 3,3-dimethylindolin-2-one, 2,3-dihydro-1H-indol-2-one-6-yl, 4-fluoro-1H-indazol-6-yl, 5-fluoro-1H-indazol-6-yl, and 3-amino-1H-indazol-6-yl;

R_3 is hydrogen;

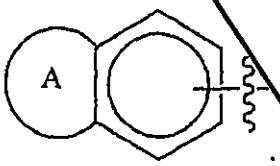
R_4 is hydrogen; and

R_5 is hydrogen.

2. At least one chemical entity of claim 1, wherein R_1 is chosen from (2-methyl-2-hydroxypropoxy)pyridin-6-yl, (2-methoxyethoxy)pyridinyl, 2-(dimethylamino)ethoxy-3-pyridinyl, hydroxyethoxy-5-pyridinyl, (3-methyl-3-hydroxyazetidin-1-yl)pyridin-3-yl, (3-methyl-3-hydroxyazetidin-1-yl)pyridin-2-yl, (3-hydroxyazetidin-1-yl)pyridin-2-yl, (hydroxy(dimethylethyl))-5-pyridinyl, (4-methyl-4-hydroxypiperidin-1-yl)pyridin-2-yl, (3-methyl-3-hydroxypiperidin-1-yl)pyridin-2-yl, 5-morpholinopyridin-2-yl, 6-morpholinopyridin-3-yl, ((2-methoxyethyl)(methyl)amino)pyridin-5-yl, ((2-hydroxyethyl)(methyl)amino)pyridin-5-yl, 2-methoxy-4-pyridinyl, and 2-hydroxy-5-pyridinyl.

3. At least one chemical entity of claim 1, wherein R_1 is chosen from (2-hydroxyethyl)-1H-pyrazol-4-yl, (2-hydroxypropyl)-1H-pyrazol-4-yl, (2-methoxyethyl)-1H-pyrazol-4-yl, 1-ethyl-pyrazol-4-yl, 1-isopropyl-pyrazol-4-yl, 3-cyclopropyl-1H-pyrazol-5-yl, and 1-ethyl-5-methyl-pyrazol-3-yl.

4. At least one chemical entity of claim 1, wherein R_1 is



5. At least one chemical entity of claim 4, wherein A is pyrazolyl, oxazolyl, pyrrolyl, thiazolyl, or imidazolyl, each of which is optionally substituted.

6. At least one chemical entity of claim 4, wherein A is imidazolyl substituted with C_1-C_6 alkyl.

7. At least one chemical entity of claim 4, wherein R_1 is chosen from 1H-benzo[d]imidazol-6-yl, 1H-benzo[d]imidazol-5-yl, 1H-indazol-5-yl, 1H-indazol-6-yl, 1-methyl-1H-benzo[d]imidazol-6-yl, benzoxazol-6-yl, benzoxazol-5-yl, imidazo[1,2-a]pyridine-6-yl, 1H-indole-6-yl, 1H-indole-5-yl, benzothiazol-6-yl, and benzothiazol-5-yl.

8. At least one chemical entity of any one of claims 1 to 7, wherein R_2 is chosen from 1H-indazolyl-6-yl, 1-methyl-1H-indazol-5-yl, 1-methyl-1H-indazol-6-yl, 3,4-dihydro-2H-1,4-benzoxazin-3-one-6-yl, 1,3-benzoxazol-6-yl, 3-aminoquinolin-6-yl, and 2,3-dihydro-1H-indol-2-one-6-yl.

9. At least one chemical entity chosen from compounds having the structure:

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-1,3-benzodiazol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indazol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1-methyl-1H-1,3-benzodiazol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1,3-benzoxazol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1,3-benzoxazol-5-amine;

5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-ol;

N-{imidazo[1,2-a]pyridin-6-yl}-6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indazol-5-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indol-5-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1,3-benzothiazol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1,3-benzothiazol-5-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-2-methoxypyridin-4-amine;

6-[8-(1H-1,3-benzodiazol-5-ylamino)imidazo[1,2-a]pyrazin-6-yl]-3,4-dihydro-2H-1,4-benzoxazin-3-one;

2-(4-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}-1H-pyrazol-1-yl)ethan-1-ol;

3-(4-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}-1H-pyrazol-1-yl)propan-1-ol;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1-(2-methoxyethyl)-1H-pyrazol-4-amine;

1-ethyl-N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-pyrazol-4-amine;

N-[6-(1,3-benzoxazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-1,3-benzodiazol-6-amine;

N-[6-(1H-1,3-benzodiazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indazol-6-amine;

N-[6-(1-methyl-1H-indazol-5-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indazol-5-amine;

N-[6-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indazol-6-amine;

N-[6-(3-aminoquinolin-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1,3-benzothiazol-5-amine;

6-{8-[(2-methoxypyridin-4-yl)amino]imidazo[1,2-a]pyrazin-6-yl}-3,4-dihydro-2H-1,4-benzoxazin-3-one;

N-[6-(2,3-dihydro-1H-indol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-2-methoxypyridin-4-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1-(propan-2-yl)-1H-pyrazol-4-amine;

1-methyl-N-[6-(1-methyl-1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-1,3-benzodiazol-6-amine;

3-cyclopropyl-N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-pyrazol-5-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

N-[6-(2,3-dimethyl-2H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

5-N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-2-N-(2-methoxyethyl)-2-N-methylpyridine-2,5-diamine;

2-[(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)(methyl)amino]ethan-1-ol;

1-[(6-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyridin-8-yl]amino}pyridin-3-yl)oxy]-2-methylpropan-2-ol;

7-(8-{[1-(2-hydroxyethyl)-1H-pyrazol-4-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-1H,2H,3H-pyrido[2,3-b][1,4]oxazin-2-one;

2-(4-{[6-(3,4-dihydro-2H-1,4-benzoxazin-7-yl)imidazo[1,2-a]pyrazin-8-yl]amino}-1H-pyrazol-1-yl)ethan-1-ol;

6-(8-{[1-(2-hydroxyethyl)-1H-pyrazol-4-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-3,4-dihydro-2H-1,4-benzoxazin-3-one;

2-(4-{[6-(1,3-benzothiazol-5-yl)imidazo[1,2-a]pyrazin-8-yl]amino}-1H-pyrazol-1-yl)ethan-1-ol;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(2-methoxyethoxy)pyridin-3-amine;

6-(8-{[6-(morpholin-4-yl)pyridin-3-yl]amino}imidazo[1,2-a]pyrazin-6-yl)quinazolin-2-amine;

2-(4-{[6-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}-1H-pyrazol-1-yl)ethan-1-ol;

6-[2-(dimethylamino)ethoxy]-N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]pyridin-3-amine;

1-(6-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyridin-8-yl]amino}pyridin-3-yl)-3-methylazetidin-3-ol;

2-[(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)oxy]ethan-1-ol;

3,3-dimethyl-6-(8-{[6-(morpholin-4-yl)pyridin-3-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-2,3-dihydro-1H-indol-2-one;

1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-3-methylazetidin-3-ol;

1-(6-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyridin-8-yl]amino}pyridin-3-yl)azetidin-3-ol;

2-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-2-methylpropan-1-ol;

1-[(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)oxy]-2-methylpropan-2-ol;

N-[5-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)pyrazolo[1,5-a]pyrimidin-7-yl]-5-(morpholin-4-yl)pyridin-2-amine;

N-[5-(1-methyl-1H-1,3-benzodiazol-6-yl)pyrazolo[1,5-a]pyrimidin-7-yl]-5-(morpholin-4-yl)pyridin-2-amine;

6-(7-{[5-(morpholin-4-yl)pyridin-2-yl]amino}pyrazolo[1,5-a]pyrimidin-5-yl)-2,3-dihydro-1H-indol-2-one;

N-[6-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

N-[6-(3,4-dihydro-2H-1,4-benzoxazin-7-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

2-[(6-{[5-(1H-indazol-6-yl)pyrazolo[1,5-a]pyrimidin-7-yl]amino}pyridin-3-yl)(methyl)amino]ethan-1-ol;

6-(7-{[5-(morpholin-4-yl)pyridin-2-yl]amino}pyrazolo[1,5-a]pyrimidin-5-yl)-3,4-dihydro-2H-1,4-benzoxazin-3-one;

N-[5-(1H-indol-6-yl)pyrazolo[1,5-a]pyrimidin-7-yl]-5-(morpholin-4-yl)pyridin-2-amine;

6-(8-{[6-(morpholin-4-yl)pyridin-3-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-2,3-dihydro-1H-indol-2-one;

N-[6-(1-methyl-1H-1,3-benzodiazol-5-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

2-[(5-{[6-(3,4-dihydro-2H-1,4-benzoxazin-7-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)(methyl)amino]ethan-1-ol;

N-[6-(1-methyl-1H-1,3-benzodiazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

1-(5-{[6-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-4-methylpiperidin-4-ol;

N-[6-(1H-1,3-benzodiazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

1-(5-{[6-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)azetidin-3-ol;

N-[5-(1H-1,3-benzodiazol-6-yl)pyrazolo[1,5-a]pyrimidin-7-yl]-5-(morpholin-4-yl)pyridin-2-amine;

1-(5-{[6-(1H-indol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-4-methylpiperidin-4-ol;

N-[6-(1H-indol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

6-(8-{[6-(morpholin-4-yl)pyridin-3-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-3,4-dihydro-2H-1,4-benzoxazin-3-one;

1-ethyl-N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyridin-8-yl]-5-methyl-1H-pyrazol-3-amine;

6-[8-({6-[(2-hydroxyethyl)(methyl)amino]pyridin-3-yl}amino)imidazo[1,2-a]pyrazin-6-yl]-3,4-dihydro-2H-1,4-benzoxazin-3-one;

1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-3-methylpiperidin-3-ol;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyridin-8-yl]-6-(morpholin-4-yl)pyridazin-3-amine;

7-(8-{[6-(morpholin-4-yl)pyridin-3-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-1H,2H,3H-pyrido[2,3-b][1,4]oxazin-2-one;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-[4-(propan-2-yl)piperazin-1-yl]pyridin-3-amine;

N-[6-(4-fluoro-1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

2-N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyridin-8-yl]-5-N-(2-methoxyethyl)-5-N-methylpyridine-2,5-diamine;

6-(1H-benzo[d]imidazol-6-yl)-N-(5-morpholinopyridin-2-yl)imidazo[1,2-b]pyridazin-8-amine;

6-(3-amino-1H-indazol-6-yl)-N-(5-morpholinopyridin-2-yl)imidazo[1,2-b]pyridazin-8-amine;

2-[(6-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyridin-8-yl]amino}pyridin-3-yl)(methyl)amino]ethan-1-ol;

1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-4-methylpiperidin-4-ol;

1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)azetidin-3-ol;

6-(1H-indol-6-yl)-N-(5-morpholinopyridin-2-yl)imidazo[1,2-b]pyridazin-8-amine;

N-[6-(5-fluoro-1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine,

(3S)-1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-3-methylpiperidin-3-ol,

(3R)-1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-3-methylpiperidin-3-ol,

1-(5-{[6-(1H-indol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-3-methylazetidin-3-ol,

[(2R)-4-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)morpholin-2-yl]methanol,

[(2S)-4-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)morpholin-2-yl]methanol,

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-2-(morpholin-4-yl)pyrimidin-5-amine,

1-ethyl-N-(6-{1H-pyrrolo[3,2-b]pyridin-6-yl}imidazo[1,2-a]pyrazin-8-yl)-1H-pyrazol-4-amine, and

2-{4-[(6-{1H-pyrrolo[3,2-b]pyridin-6-yl}imidazo[1,2-a]pyrazin-8-yl)amino]-1H-pyrazol-1-yl}ethan-1-ol,

or a pharmaceutically acceptable salt thereof.

10. A pharmaceutical composition comprising at least one chemical entity of any one of claims 1 to 9, and at least one pharmaceutically acceptable vehicle chosen from carriers, adjuvants, and excipients.

11. An effective amount of at least one chemical entity of any one of claims 1-9 for use in the treatment of a human having a disease responsive to the inhibition of Syk activity.

12. The use of an effective amount of at least one chemical entity of any one of claims 1-9 in the manufacture of a medicament for the treatment of a human having a disease responsive to the inhibition of Syk activity.

13. The chemical entity according to claim 11, or the use according to claim 12, wherein the at least one chemical entity is to be administered to the human intravenously, intramuscularly, parenterally, or orally.

14. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease responsive to inhibition of Syk activity is cancer.

15. The chemical entity or the use according to claim 14, wherein the cancer is leukemia, lymphoma, or multiple myeloma.

16. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease responsive to inhibition of Syk activity is B-cell lymphoma or leukemia.

17. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease responsive to inhibition of Syk activity is rheumatoid arthritis, allergic rhinitis, chronic obstructive pulmonary disease (COPD), adult respiratory distress syndrome (ARDs), an allergy-induced inflammatory disease, multiple sclerosis, autoimmune disease, inflammatory disease, acute inflammatory reaction, allergic disorder, or polycystic kidney disease.

18. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease responsive to inhibition of Syk activity is B-cell lymphoma, Hodgkin's lymphoma, non-Hodgkin's lymphoma, hairy cell leukemia, multiple myeloma, chronic myelogenous leukemia, acute myelogenous leukemia, chronic lymphocytic leukemia, or acute lymphocytic leukemia.

19. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease is chronic lymphocytic leukemia.

20. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease is acute lymphocytic leukemia.

21. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease is acute myelogenous leukemia.

22. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease is B-cell lymphoma or B-cell leukemia.

23. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease is systemic lupus erythematosus (SLE), rheumatoid arthritis, multiple vasculitides, idiopathic thrombocytopenic purpura (ITP), myasthenia gravis, allergic

rhinitis, chronic obstructive pulmonary disease (COPD), adult respiratory distress syndrome (ARDs) and asthma.

24. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease is systemic lupus erythematosus (SLE).

25. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease is an allergy-induced inflammatory disease.

26. The chemical entity according to claim 11 or claim 13, or the use according to claim 12 or claim 13, wherein the disease is asthma.

27. A method for determining the presence of Syk in a sample, comprising contacting the sample with at least one chemical entity of any one of claims 1 to 9 under conditions that permit detection of Syk activity, detecting a level of Syk activity in the sample, and therefrom determining the presence or absence of Syk in the sample.

28. A method for inhibiting B-cell activity comprising contacting cells expressing Syk with a compound of at least one chemical entity of any one of claims 1 to 9 in an amount sufficient to detectably decrease B-cell activity *in vitro*.

29. A method for inhibiting ATP hydrolysis, the method comprising contacting cells expressing Syk with at least one chemical entity of any one of claims 1 to 9 in an amount sufficient to detectably decrease B-cell activity *in vitro*.

30. At least one chemical entity of any one of claims 1-9 for use in inhibiting B-cell activity in a human in need thereof.

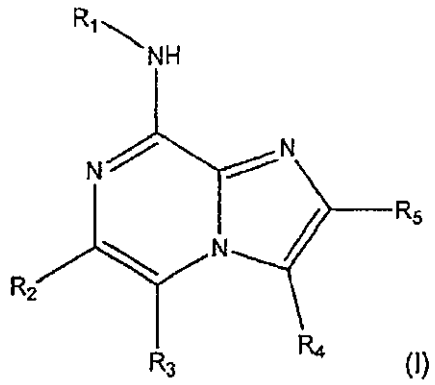
31. The use of at least one chemical entity of any one of claims 1-9 in the manufacture of a medicament for inhibiting B-cell activity in a human in need thereof.

32. At least one chemical entity of any one of claims 1-9 for use in inhibiting ATP hydrolysis in a human in need thereof.

33. The use of at least one chemical entity of any one of claims 1-9 in the manufacture of a medicament for inhibiting ATP hydrolysis in a human in need thereof.

CLAIMS

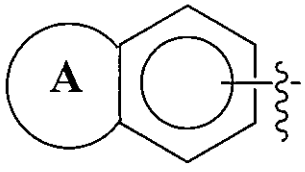
1. At least one chemical entity chosen from compounds of Formula (I):



(I)

or a pharmaceutically acceptable salt thereof, wherein

R_1 is chosen from (2-methyl-2-hydroxypropoxy)pyridin-6-yl, (2-methoxyethoxy)pyridinyl, 2-(dimethylamino)ethoxy-3-pyridinyl, hydroxyethoxy-5-pyridinyl, (3-methyl-3-hydroxyazetidin-1-yl)pyridin-3-yl, (3-methyl-3-hydroxyazetidin-1-yl)pyridin-2-yl, (3-hydroxyazetidin-1-yl)pyridin-2-yl, (hydroxy(dimethylethyl))-5-pyridinyl, (4-methyl-4-hydroxypiperidin-1-yl)pyridin-2-yl, (3-methyl-3-hydroxypiperidin-1-yl)pyridin-2-yl, 5-morpholinopyridin-2-yl, 6-morpholinopyridin-3-yl, ((2-methoxyethyl)(methyl)amino)pyridin-5-yl, ((2-hydroxyethyl)(methyl)amino)pyridin-5-yl, 2-methoxy-4-pyridinyl, 2-hydroxy-5-pyridinyl, (2-hydroxyethyl)-1H-pyrazol-4-yl, (2-hydroxypropyl)-1H-pyrazol-4-yl, (2-methoxyethyl)-1H-pyrazol-4-yl, 1-ethyl-pyrazol-4-yl, 1-isopropyl-pyrazol-4-yl, 3-cyclopropyl-1H-pyrazol-5-yl, and 1-ethyl-5-methyl-pyrazol-3-yl; or

R_1 is 

, wherein A is an optionally substituted heteroaryl group having from 5 to 7 ring atoms including the atoms shared with the 6 membered aromatic ring;

R_2 is chosen from 2,3-dimethyl-2H-indazol-6-yl, 1H-indazolyl-6-yl, 1-methyl-1H-indazol-5-yl, 1-methyl-1H-indazol-6-yl, 3,4-dihydro-2H-1,4-benzoxazin-3-one-6-yl, 1,3-benzoxazol-6-yl, 3-aminoquinolin-6-yl, 2,3-dihydro-1H-indol-6-yl, benzothiazolyl, 2-

aminoquinazolin-6-yl, 3,3-dimethylindolin-2-one, 2,3-dihydro-1H-indol-2-one-6-yl, 4-fluoro-1H-indazol-6-yl, 5-fluoro-1H-indazol-6-yl, and 3-amino-1H-indazol-6-yl;

R_3 is hydrogen;

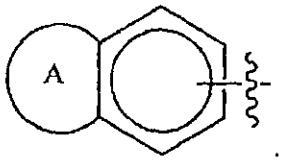
R_4 is hydrogen; and

R_5 is hydrogen.

2. At least one chemical entity of claim 1, wherein R_1 is chosen from (2-methyl-2-hydroxypropoxy)pyridin-6-yl, (2-methoxyethoxy)pyridinyl, 2-(dimethylamino)ethoxy-3-pyridinyl, hydroxyethoxy-5-pyridinyl, (3-methyl-3-hydroxyazetidin-1-yl)pyridin-3-yl, (3-methyl-3-hydroxyazetidin-1-yl)pyridin-2-yl, (3-hydroxyazetidin-1-yl)pyridin-2-yl, (hydroxy(dimethylethyl))-5-pyridinyl, (4-methyl-4-hydroxypiperidin-1-yl)pyridin-2-yl, (3-methyl-3-hydroxypiperidin-1-yl)pyridin-2-yl, 5-morpholinopyridin-2-yl, 6-morpholinopyridin-3-yl, ((2-methoxyethyl)(methyl)amino)pyridin-5-yl, ((2-hydroxyethyl)(methyl)amino)pyridin-5-yl, 2-methoxy-4-pyridinyl, and 2-hydroxy-5-pyridinyl.

3. At least one chemical entity of claim 1, wherein R_1 is chosen from (2-hydroxyethyl)-1H-pyrazol-4-yl, (2-hydroxypropyl)-1H-pyrazol-4-yl, (2-methoxyethyl)-1H-pyrazol-4-yl, 1-ethyl-pyrazol-4-yl, 1-isopropyl-pyrazol-4-yl, 3-cyclopropyl-1H-pyrazol-5-yl, and 1-ethyl-5-methyl-pyrazol-3-yl.

4. At least one chemical entity of claim 1, wherein R_1 is



5. At least one chemical entity of claim 4, wherein A is pyrazolyl, oxazolyl, pyrrolyl, thiazolyl, or imidazolyl, each of which is optionally substituted.

6. At least one chemical entity of claim 4, wherein A is imidazolyl substituted with C_1-C_6 alkyl.

7. At least one chemical entity of claim 4, wherein R_1 is chosen from 1H-benzo[d]imidazol-6-yl, 1H-benzo[d]imidazol-5-yl, 1H-indazol-5-yl, 1H-indazol-6-yl, 1-methyl-1H-benzo[d]imidazol-6-yl, benzoxazol-6-yl, benzoxazol-5-yl, imidazo[1,2-a]pyridine-6-yl, 1H-indole-6-yl, 1H-indole-5-yl, benzothiazol-6-yl, and benzothiazol-5-yl.

8. At least one chemical entity of any one of claims 1 to 7, wherein R_2 is chosen from 1H-indazolyl-6-yl, 1-methyl-1H-indazol-5-yl, 1-methyl-1H-indazol-6-yl, 3,4-dihydro-2H-1,4-benzoxazin-3-one-6-yl, 1,3-benzoxazol-6-yl, 3-aminoquinolin-6-yl, and 2,3-dihydro-1H-indol-2-one-6-yl.

9. At least one chemical entity chosen from compounds having the structure:

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-1,3-benzodiazol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indazol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1-methyl-1H-1,3-benzodiazol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1,3-benzoxazol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1,3-benzoxazol-5-amine;

5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-ol;

N-{imidazo[1,2-a]pyridin-6-yl}-6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indazol-5-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indol-5-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1,3-benzothiazol-6-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1,3-benzothiazol-5-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-2-methoxypyridin-4-amine;

6-[8-(1H-1,3-benzodiazol-5-ylamino)imidazo[1,2-a]pyrazin-6-yl]-3,4-dihydro-2H-1,4-benzoxazin-3-one;

2-(4-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}-1H-pyrazol-1-yl)ethan-1-ol;

3-(4-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}-1H-pyrazol-1-yl)propan-1-ol;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1-(2-methoxyethyl)-1H-pyrazol-4-amine;

1-ethyl-N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-pyrazol-4-amine;

N-[6-(1,3-benzoxazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-1,3-benzodiazol-6-amine;

N-[6-(1H-1,3-benzodiazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indazol-6-amine;

N-[6-(1-methyl-1H-indazol-5-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indazol-5-amine;

N-[6-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-indazol-6-amine;

N-[6-(3-aminoquinolin-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1,3-benzothiazol-5-amine;

6-{8-[(2-methoxypyridin-4-yl)amino]imidazo[1,2-a]pyrazin-6-yl}-3,4-dihydro-2H-1,4-benzoxazin-3-one;

N-[6-(2,3-dihydro-1H-indol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-2-methoxypyridin-4-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1-(propan-2-yl)-1H-pyrazol-4-amine;

1-methyl-N-[6-(1-methyl-1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-1,3-benzodiazol-6-amine;

3-cyclopropyl-N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-1H-pyrazol-5-amine;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

N-[6-(2,3-dimethyl-2H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

5-N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-2-N-(2-methoxyethyl)-2-N-methylpyridine-2,5-diamine;

2-[(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)(methyl)amino]ethan-1-ol;

1-[(6-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyridin-8-yl]amino}pyridin-3-yl)oxy]-2-methylpropan-2-ol;

7-(8-{[1-(2-hydroxyethyl)-1H-pyrazol-4-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-1H,2H,3H-pyrido[2,3-b][1,4]oxazin-2-one;

2-(4-{[6-(3,4-dihydro-2H-1,4-benzoxazin-7-yl)imidazo[1,2-a]pyrazin-8-yl]amino}-1H-pyrazol-1-yl)ethan-1-ol;

6-(8-{[1-(2-hydroxyethyl)-1H-pyrazol-4-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-3,4-dihydro-2H-1,4-benzoxazin-3-one;

2-(4-{[6-(1,3-benzothiazol-5-yl)imidazo[1,2-a]pyrazin-8-yl]amino}-1H-pyrazol-1-yl)ethan-1-ol;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(2-methoxyethoxy)pyridin-3-amine;

6-(8-{[6-(morpholin-4-yl)pyridin-3-yl]amino}imidazo[1,2-a]pyrazin-6-yl)quinazolin-2-amine;

2-(4-{[6-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}-1H-pyrazol-1-yl)ethan-1-ol;

6-[2-(dimethylamino)ethoxy]-N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]pyridin-3-amine;

2-[(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)oxy]ethan-1-ol;

3,3-dimethyl-6-(8-{[6-(morpholin-4-yl)pyridin-3-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-2,3-dihydro-1H-indol-2-one;

1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-3-methylazetidin-3-ol;

1-(6-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyridin-8-yl]amino}pyridin-3-yl)azetidin-3-ol;

2-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-2-methylpropan-1-ol;

1-[(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)oxy]-2-methylpropan-2-ol;

N-[6-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

N-[6-(3,4-dihydro-2H-1,4-benzoxazin-7-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

6-(8-{[6-(morpholin-4-yl)pyridin-3-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-2,3-dihydro-1H-indol-2-one;

N-[6-(1-methyl-1H-1,3-benzodiazol-5-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

2-[(5-{[6-(3,4-dihydro-2H-1,4-benzoxazin-7-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)(methyl)amino]ethan-1-ol;

N-[6-(1-methyl-1H-1,3-benzodiazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

1-(5-{[6-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-4-methylpiperidin-4-ol;

N-[6-(1H-1,3-benzodiazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

1-(5-{[6-(3,4-dihydro-2H-1,4-benzoxazin-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)azetidin-3-ol;

1-(5-{[6-(1H-indol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-4-methylpiperidin-4-ol;

N-[6-(1H-indol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

6-(8-{[6-(morpholin-4-yl)pyridin-3-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-3,4-dihydro-2H-1,4-benzoxazin-3-one;

6-[8-({6-[(2-hydroxyethyl)(methyl)amino]pyridin-3-yl}amino)imidazo[1,2-a]pyrazin-6-yl]-3,4-dihydro-2H-1,4-benzoxazin-3-one;

1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-3-methylpiperidin-3-ol;

7-(8-{[6-(morpholin-4-yl)pyridin-3-yl]amino}imidazo[1,2-a]pyrazin-6-yl)-1H,2H,3H-pyrido[2,3-b][1,4]oxazin-2-one;

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-[4-(propan-2-yl)piperazin-1-yl]pyridin-3-amine;

N-[6-(4-fluoro-1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine;

1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-4-methylpiperidin-4-ol;

1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)azetidin-3-ol;

N-[6-(5-fluoro-1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-6-(morpholin-4-yl)pyridin-3-amine,

(3S)-1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-3-methylpiperidin-3-ol,

(3R)-1-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-3-methylpiperidin-3-ol,

1-(5-{[6-(1H-indol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)-3-methylazetidin-3-ol,

[(2R)-4-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)morpholin-2-yl]methanol,

[(2S)-4-(5-{[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]amino}pyridin-2-yl)morpholin-2-yl]methanol,

N-[6-(1H-indazol-6-yl)imidazo[1,2-a]pyrazin-8-yl]-2-(morpholin-4-yl)pyrimidin-5-amine,

1-ethyl-N-(6-{1H-pyrrolo[3,2-b]pyridin-6-yl}imidazo[1,2-a]pyrazin-8-yl)-1H-pyrazol-4-amine, and

2-{4-[(6-{1H-pyrrolo[3,2-b]pyridin-6-yl}imidazo[1,2-a]pyrazin-8-yl)amino]-1H-pyrazol-1-yl}ethan-1-ol,

or a pharmaceutically acceptable salt thereof.

10. A pharmaceutical composition comprising at least one chemical entity of any one of claims 1 to 9, and at least one pharmaceutically acceptable vehicle chosen from carriers, adjuvants, and excipients.

11. The use of an effective amount of at least one chemical entity of any one of claims 1 to 9 in the manufacture of a medicament for the treatment of a human having a disease responsive to the inhibition of Syk activity, wherein the disease responsive to inhibition of Syk activity is cancer.

12. The use according to claim 11, wherein the at least one chemical entity is to be administered to the human intravenously, intramuscularly, parenterally, or orally.

13. The use according to claim 11 or 12, wherein the cancer is leukemia, lymphoma, or multiple myeloma.

14. The use according to claim 11 or 12, wherein the cancer is B-cell lymphoma or leukemia.

15. The use according to claim 11 or 12, wherein the cancer is Hodgkin's lymphoma, non-Hodgkin's lymphoma, hairy cell leukemia, chronic myelogenous leukemia, acute myelogenous leukemia, chronic lymphocytic leukemia, or acute lymphocytic leukemia.

16. The use according to claim 11 or 12, wherein the cancer is chronic lymphocytic leukemia.

17. The use according to claim 11 or 12, wherein the cancer is acute lymphocytic leukemia.

18. The use according to claim 11 or 12, wherein the cancer is acute myelogenous leukemia.

19. The use according to claim 11 or 12, wherein the cancer is B-cell lymphoma or B-cell leukemia.

20. The use of an effective amount of at least one chemical entity of any one of claims 1 to 9 in the manufacture of a medicament for the treatment of a human having a disease responsive to the inhibition of Syk activity, wherein the disease responsive to inhibition of Syk activity is rheumatoid arthritis, allergic rhinitis, chronic obstructive pulmonary disease (COPD), adult respiratory distress syndrome (ARDs), an allergy-induced inflammatory disease, multiple sclerosis, autoimmune disease, inflammatory disease, acute inflammatory reaction, allergic disorder, polycystic kidney disease, systemic lupus erythematosus (SLE), multiple vasculitides, idiopathic thrombocytopenic purpura (ITP), myasthenia gravis, or asthma.

21. The use according to claim 20, wherein the disease is systemic lupus erythematosus (SLE).

22. The use according to claim 20, wherein the disease is an allergy-induced inflammatory disease.

23. The use according to claim 20, wherein the disease is asthma.

24. A method for determining the presence of Syk in a sample, comprising contacting the sample with at least one chemical entity of any one of claims 1 to 9 under conditions that permit

detection of Syk activity, detecting a level of Syk activity in the sample, and therefrom determining the presence or absence of Syk in the sample.

25. A method for inhibiting B-cell activity comprising contacting cells expressing Syk with a compound of at least one chemical entity of any one of claims 1 to 9 in an amount sufficient to detectably decrease B-cell activity *in vitro*.

26. A method for inhibiting ATP hydrolysis, the method comprising contacting cells expressing Syk with at least one chemical entity of any one of claims 1 to 9 in an amount sufficient to detectably decrease B-cell activity *in vitro*.