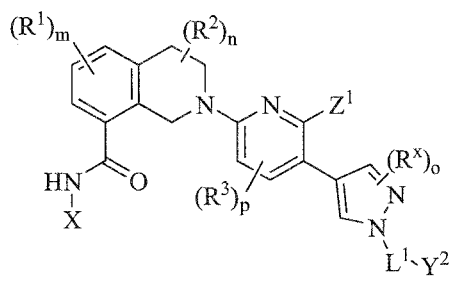


certain cancer type. Such a selective compound also may confer certain advantages in the clinical setting, by providing, for example, the flexibility to select a dosing regimen, a reduced on-target toxic effect in normal cells, among others (e.g., lymphopenia has been observed in Bcl-2 deficient mice). See, Nakayama, K. *et al. PNAS* (1994) 91, 3700-3704.

5 In view of the above, there is a need in the art for small molecules therapeutics that can selectively inhibit the activity of one type or a subset of anti-apoptotic Bcl-2 proteins, for example, of a Bcl-xL anti-apoptotic protein. The present invention fulfills at least this need.

SUMMARY OF THE INVENTION

10 In one aspect, there is provided a compound having Formula (III)



Formula (III),

or a therapeutically acceptable salt thereof, wherein

X is benzo[d]thiazolyl optionally substituted with one or two R⁴;

15 R^x, at each occurrence, is independently selected from the group consisting of R⁵, CO(O)R⁵, CO(O)H, CN, F, Cl, Br and I;

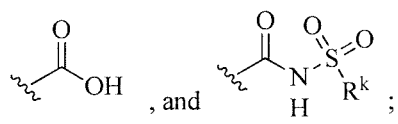
L¹ is selected from the group consisting of (CR⁶R⁷)_q, (CR⁶R⁷)₂-C(O)-(CR⁶R⁷)_p, and (CR⁶R⁷)₂-S(O)₂-(CR⁶R⁷)_p; and

20 Y² is selected from the group consisting of C₃₋₁₁ branched chain alkyl, C₃₋₇ cycloalkyl, C₄₋₇ cycloalkenyl, phenyl, and C₃₋₇ heterocyclyl; wherein the C₃₋₇ cycloalkyl, C₄₋₇ cycloalkenyl, phenyl, and C₃₋₇ heterocyclyl are optionally fused to one benzene ring; wherein Y² is optionally substituted with one, two, or three substituents independently selected from the group consisting of R⁸, OR⁸, SO₂R⁸, CO(O)R⁸, NHR⁸, N(R⁸)₂, C(O)H, OH, CN, NO₂, F, Cl, Br and I; or

25 L¹ is a bond; and

30 Y² is selected from the group consisting of C₃₋₇ cycloalkyl, phenyl, and C₃₋₇ heterocyclyl; wherein the C₃₋₇ cycloalkyl, phenyl, and C₃₋₇ heterocyclyl represented by Y² are optionally fused to one benzene ring; wherein each Y² and each ring fused to Y² are optionally substituted with one substituent independently selected from the group consisting of R⁸ and C(O)NHR⁸;

Z¹ is selected from the group consisting of



R¹ is absent;

R², at each occurrence, is independently C₁₋₆ alkyl;

5 R³ is absent;

R⁴, at each occurrence, is independently selected from the group consisting of OR¹²

and halogen;

R⁵, at each occurrence, is independently selected from the group consisting of C₁₋₆ alkyl, C₁₋₆ haloalkyl, C₁₋₆ hydroxyalkyl, aryl, and cycloalkyl;

10 R⁶ and R⁷, at each occurrence, are each independently selected from the group consisting of hydrogen, R¹⁵, and CO(O)R¹⁵;

R⁸, at each occurrence, is independently selected from the group consisting of C₁₋₆ alkyl, C₂₋₆ alkynyl, aryl, heterocyclyl, and cycloalkyl; wherein the R⁸ C₁₋₆ alkyl, and C₂₋₆ alkynyl are optionally substituted with one, two, or three substituents independently selected
15 from the group consisting of R¹⁶, OR¹⁶, SO₂R¹⁶, C(O)R¹⁶, N(R¹⁶)₂, OH, F, Cl, Br and I;
wherein the R⁸ aryl and heterocyclyl are optionally substituted with one substituent
independently selected from the group consisting of C₁₋₆ alkyl, F, Cl, Br and I;

R^k, at each occurrence, is independently C₁₋₆ alkyl;

R¹² is C₁₋₄ alkyl;

20 R¹⁵, at each occurrence, is independently selected from the group consisting of C₁₋₄ alkyl, and aryl; wherein the R¹⁵ C₁₋₄ alkyl is optionally substituted with one substituent independently selected from the group consisting C₁₋₄ alkoxy, and heterocycloalkyl;

R¹⁶, at each occurrence, is independently selected from the group consisting of C₁₋₄ alkyl, aryl, heterocycloalkyl, and heteroaryl;

25 q is 1, 2, or 3;

s is 0;

r is 0, or 1;

m is 0;

n is 0, or 1;

30 o is 0, 1, or 2; and

p is 0.

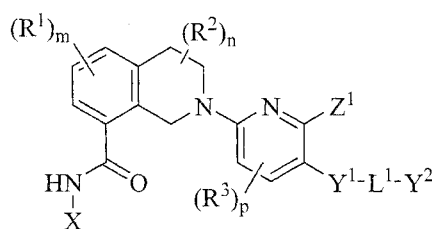
In one aspect, there is provided 6-[8-(1,3-Benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(1-methoxy-3,3-dimethylcyclohexyl)methyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid or a therapeutically acceptable salt thereof.

In one aspect, there is provided a pharmaceutical composition comprising an excipient and a therapeutically effective amount of the compound or therapeutically acceptable salt as described herein.

In one aspect, there is provided use of a therapeutic effective amount of a compound as described herein, or a pharmaceutically acceptable salt thereof, in the manufacture of a medicament for treating bladder cancer, brain cancer, breast cancer, bone marrow cancer, cervical cancer, chronic lymphocytic leukemia, colorectal cancer, esophageal cancer, hepatocellular cancer, lymphoblastic leukemia, follicular lymphoma, a lymphoid malignancy of T-cell or B-cell origin, melanoma, myelogenous leukemia, myeloma, oral cancer, ovarian cancer, non-small cell lung cancer, prostate cancer, small cell lung cancer or spleen cancer in a patient.

In one aspect, there is provided use of a therapeutic effective amount of a compound as described herein, or a pharmaceutically acceptable salt thereof, in the manufacture of a medicament for treating bladder cancer, brain cancer, breast cancer, bone marrow cancer, cervical cancer, chronic lymphocytic leukemia, colorectal cancer, esophageal cancer, hepatocellular cancer, lymphoblastic leukemia, follicular lymphoma, a lymphoid malignancy of T-cell or B-cell origin, melanoma, myelogenous leukemia, myeloma, oral cancer, ovarian cancer, non-small cell lung cancer, prostate cancer, small cell lung cancer or spleen cancer in a patient.

One embodiment of this invention disclosure, therefore, pertains to compounds or therapeutically acceptable salts thereof, which are useful as inhibitors of anti-apoptotic Bcl-xL proteins, the compounds having Formula (I)



Formula (I)

or a therapeutically acceptable salt thereof, wherein

X is heteroaryl; wherein the heteroaryl represented by X is optionally substituted with one, two, three, or four R⁴;

Y¹ is phenylene, or C₅₋₆ heteroarylene; optionally fused to one or two rings selected from the group consisting of C₃₋₈ cycloalkane, C₃₋₈ cycloalkene, benzene, C₅₋₆ heteroarene, C₃.

₈ heterocycloalkane, and C₃₋₈ heterocycloalkene; wherein Y¹ is optionally substituted with one, two, three, or four substituents independently selected from the group consisting of R⁵, OR⁵, SR⁵, S(O)R⁵, SO₂R⁵, C(O)R⁵, CO(O)R⁵, OC(O)R⁵, OC(O)OR⁵, NH₂, NHR⁵, N(R⁵)₂, NHC(O)R⁵, NR⁵C(O)R⁵, NHS(O)₂R⁵, NR⁵S(O)₂R⁵, NHC(O)OR⁵, NR⁵C(O)OR⁵,

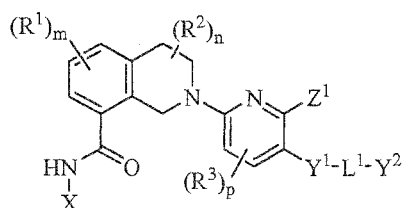
- 5 NHC(O)NH₂, NHC(O)NHR⁵, NHC(O)N(R⁵)₂, NR⁵C(O)NHR⁵, NR⁵C(O)N(R⁵)₂, C(O)NH₂, C(O)NHR⁵, C(O)N(R⁵)₂, C(O)NHOH, C(O)NHOR⁵, C(O)NHSO₂R⁵, C(O)NR⁵SO₂R⁵, SO₂NH₂, SO₂NHR⁵, SO₂N(R⁵)₂, CO(O)H, C(O)H, OH, CN, N₃, NO₂, F, Cl, Br and I;

- L¹ is selected from the group consisting of (CR⁶R⁷)_q, (CR⁶R⁷)_s-O-(CR⁶R⁷)_r, (CR⁶R⁷)_s-C(O)-(CR⁶R⁷)_r, (CR⁶R⁷)_s-S-(CR⁶R⁷)_r, (CR⁶R⁷)_s-S(O)₂-(CR⁶R⁷)_r, (CR⁶R⁷)_s-NR^{6A}C(O)-(CR⁶R⁷)_r, (CR⁶R⁷)_s-C(O)NR^{6A}-(CR⁶R⁷)_r, (CR⁶R⁷)_s-NR^{6A}-(CR⁶R⁷)_r, (CR⁶R⁷)_s-S(O)₂NR^{6A}-(CR⁶R⁷)_r, and (CR⁶R⁷)_s-NR^{6A}S(O)₂-(CR⁶R⁷)_r; and
- 10

Y² is selected from the group consisting of C₃₋₁₁ branched chain alkyl, C₃₋₇ cycloalkyl, C₄₋₇ cycloalkenyl, phenyl, and C₃₋₇ heterocyclyl; wherein the C₃₋₇ cycloalkyl, C₄₋₇ cycloalkenyl, phenyl, and C₃₋₇ heterocyclyl are optionally fused to one or two rings selected from the group

WHAT IS CLAIMED IS:

1. A compound having Formula (I)



Formula (I),

- 5 or a therapeutically acceptable salt thereof, or a therapeutically acceptable salt thereof, wherein

X is heteroaryl; wherein the heteroaryl represented by X is optionally substituted with one, two, three, or four R⁴;

- Y¹ is phenylene, or C₅₋₆ heteroarylene; optionally fused to one or two rings selected
 10 from the group consisting of C₃₋₈ cycloalkane, C₃₋₈ cycloalkene, benzene, C₅₋₆ heteroarene, C₃₋₈ heterocycloalkane, and C₃₋₈ heterocycloalkene; wherein Y¹ is optionally substituted with one, two, three, or four substituents independently selected from the group consisting of R⁵, OR⁵, SR⁵, S(O)R⁵, SO₂R⁵, C(O)R⁵, CO(O)R⁵, OC(O)R⁵, OC(O)OR⁵, NH₂, NHR⁵, N(R⁵)₂, NHC(O)R⁵, NR⁵C(O)R⁵, NHS(O)₂R⁵, NR⁵S(O)₂R⁵, NHC(O)OR⁵, NR⁵C(O)OR⁵,
 15 NHC(O)NH₂, NHC(O)NHR⁵, NHC(O)N(R⁵)₂, NR⁵C(O)NHR⁵, NR⁵C(O)N(R⁵)₂, C(O)NH₂, C(O)NHR⁵, C(O)N(R⁵)₂, C(O)NHOH, C(O)NHOR⁵, C(O)NHSO₂R⁵, C(O)NR⁵SO₂R⁵, SO₂NH₂, SO₂NHR⁵, SO₂N(R⁵)₂, CO(O)H, C(O)H, OH, CN, N₃, NO₂, F, Cl, Br and I;

- L¹ is selected from the group consisting of (CR⁶R⁷)_q, (CR⁶R⁷)_s-O-(CR⁶R⁷)_r, (CR⁶R⁷)_s-C(O)-(CR⁶R⁷)_r, (CR⁶R⁷)_s-S-(CR⁶R⁷)_r, (CR⁶R⁷)_s-S(O)₂-(CR⁶R⁷)_r, (CR⁶R⁷)_s-NR^{6A}C(O)-
 20 (CR⁶R⁷)_r, (CR⁶R⁷)_s-C(O)NR^{6A}-(CR⁶R⁷)_r, (CR⁶R⁷)_s-NR^{6A}-(CR⁶R⁷)_r, (CR⁶R⁷)_s-S(O)₂NR^{6A}-(CR⁶R⁷)_r, and (CR⁶R⁷)_s-NR^{6A}S(O)₂-(CR⁶R⁷)_r; and

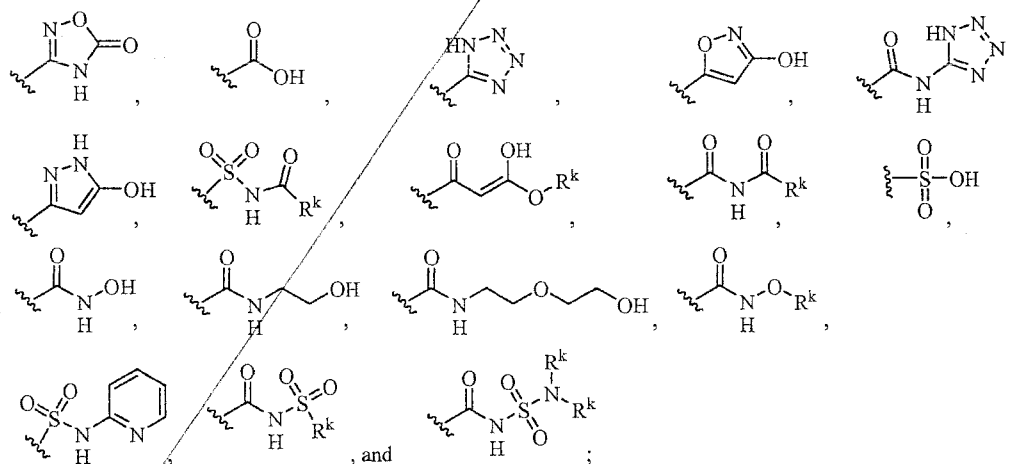
- Y² is selected from the group consisting of C₃₋₁₁ branched chain alkyl, C₃₋₇ cycloalkyl, C₄₋₇ cycloalkenyl, phenyl, and C₃₋₇ heterocyclyl; wherein the C₃₋₇ cycloalkyl, C₄₋₇ cycloalkenyl, phenyl, and C₃₋₇ heterocyclyl are optionally fused to one or two rings selected from the group
 25 consisting of C₃₋₈ cycloalkane, C₃₋₈ cycloalkene, benzene, C₅₋₆ heteroarene, C₃₋₈ heterocycloalkane, and C₃₋₈ heterocycloalkene; wherein Y² is optionally substituted with one, two, three, four, or five substituents independently selected from the group consisting of R⁸, OR⁸, SR⁸, S(O)R⁸, SO₂R⁸, C(O)R⁸, CO(O)R⁸, OC(O)R⁸, OC(O)OR⁸, NH₂, NHR⁸, N(R⁸)₂, NHC(O)R⁸, NR⁸C(O)R⁸, NHS(O)₂R⁸, NR⁸S(O)₂R⁸, NHC(O)OR⁸, NR⁸C(O)OR⁸,
 30 NHC(O)NH₂, NHC(O)NHR⁸, NHC(O)N(R⁸)₂, NR⁸C(O)NHR⁸, NR⁸C(O)N(R⁸)₂, C(O)NH₂, C(O)NHR⁸, C(O)N(R⁸)₂, C(O)NHOH, C(O)NHOR⁸, C(O)NHSO₂R⁸, C(O)NR⁸SO₂R⁸, SO₂NH₂, SO₂NHR⁸, SO₂N(R⁸)₂, CO(O)H, C(O)H, OH, CN, N₃, NO₂, F, Cl, Br and I; or

L¹ is a bond; and

- Y^2 is selected from the group consisting of C_{3-7} cycloalkyl, C_{4-7} cycloalkenyl, phenyl, and C_{3-7} heterocyclyl; wherein the C_{3-7} cycloalkyl, C_{4-7} cycloalkenyl, phenyl, and C_{3-7} heterocyclyl represented by Y^2 are optionally fused to one or two rings selected from the group consisting of C_{3-8} cycloalkane, C_{3-8} cycloalkene, benzene, C_{5-6} heteroarene, C_{3-8} heterocycloalkane, and C_{3-8} heterocycloalkene; wherein each Y^2 and each ring fused to Y^2 are optionally substituted with one, two, three, four, or five substituents independently selected from the group consisting of R^8 , OR^8 , SR^8 , $S(O)R^8$, SO_2R^8 , $C(O)R^8$, $CO(O)R^8$, $OC(O)R^8$, $OC(O)OR^8$, NH_2 , NHR^8 , $N(R^8)_2$, $NHC(O)R^8$, $NR^8C(O)R^8$, $NHS(O)_2R^8$, $NR^8S(O)_2R^8$, $NHC(O)OR^8$, $NR^8C(O)OR^8$, $NHC(O)NH_2$, $NHC(O)NHR^8$, $NHC(O)N(R^8)_2$, $NR^8C(O)NHR^8$, $NR^8C(O)N(R^8)_2$, $C(O)NH_2$, $C(O)NHR^8$, $C(O)N(R^8)_2$, $C(O)NHOH$, $C(O)NHOR^8$, $C(O)NHSO_2R^8$, $C(O)NR^8SO_2R^8$, SO_2NH_2 , SO_2NHR^8 , $SO_2N(R^8)_2$, $CO(O)H$, $C(O)H$, OH , CN , N_3 , NO_2 , F , Cl , Br and I ;

- Z^1 is selected from the group consisting of $C(O)OR^9$, $C(O)NR^{10}R^{11}$, $C(O)R^{11}$, $NR^{10}C(O)R^{11}$, $NR^{10}C(O)NR^{10}R^{11}$, $OC(O)NR^{10}R^{11}$, $NR^{10}C(O)OR^9$, $C(=NOR^{10})NR^{10}R^{11}$, $NR^{10}C(=NCN)NR^{10}R^{11}$, $NR^{10}S(O)_2NR^{10}R^{11}$, $S(O)_2R^9$, $S(O)_2NR^{10}R^{11}$, $N(R^{10})S(O)_2R^{11}$, $NR^{10}C(=NR^{11})NR^{10}R^{11}$, $C(=S)NR^{10}R^{11}$, $C(=NR^{10})NR^{10}R^{11}$, halogen, NO_2 , and CN ; or

Z^1 is selected from the group consisting of



- R^1 , at each occurrence, is independently selected from the group consisting of halo, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, and C_{1-6} haloalkyl;
- R^2 , at each occurrence, is independently selected from the group consisting of deuterium, halo, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, and C_{1-6} haloalkyl;
- two R^2 that are attached to the same carbon atom, together with said carbon atom, optionally form a ring selected from the group consisting of heterocycloalkyl, heterocycloalkenyl, cycloalkyl, and cycloalkenyl;
- R^3 , at each occurrence, is independently selected from the group consisting of halo, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, and C_{1-6} haloalkyl;

R^4 , at each occurrence, is independently selected from the group consisting of $NR^{12}R^{13}$, OR^{12} , CN, NO_2 , halogen, $C(O)OR^{12}$, $C(O)NR^{12}R^{13}$, $NR^{12}C(O)R^{13}$, $NR^{12}S(O)_2R^{14}$, $NR^{12}S(O)R^{14}$, $S(O)_2R^{14}$, $S(O)R^{14}$ and R^{14} ;

R^5 , at each occurrence, is independently selected from the group consisting of C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{1-6} haloalkyl, C_{1-6} hydroxyalkyl, aryl, heterocyclyl, cycloalkyl, and cycloalkenyl;

R^{6A} is independently selected from the group consisting of hydrogen, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, and C_{1-6} haloalkyl;

R^6 and R^7 , at each occurrence, are each independently selected from the group consisting of hydrogen, R^{15} , OR^{15} , SR^{15} , $S(O)R^{15}$, SO_2R^{15} , $C(O)R^{15}$, $CO(O)R^{15}$, $OC(O)R^{15}$, $OC(O)OR^{15}$, NH_2 , NHR^{15} , $N(R^{15})_2$, $NHC(O)R^{15}$, $NR^{15}C(O)R^{15}$, $NHS(O)_2R^{15}$, $NR^{15}S(O)_2R^{15}$, $NHC(O)OR^{15}$, $NR^{15}C(O)OR^{15}$, $NHC(O)NH_2$, $NHC(O)NHR^{15}$, $NHC(O)N(R^{15})_2$, $NR^{15}C(O)NHR^{15}$, $NR^{15}C(O)N(R^{15})_2$, $C(O)NH_2$, $C(O)NHR^{15}$, $C(O)N(R^{15})_2$, $C(O)NHOH$, $C(O)NHOR^{15}$, $C(O)NHSO_2R^{15}$, $C(O)NR^{15}SO_2R^{15}$, SO_2NH_2 , SO_2NHR^{15} , $SO_2N(R^{15})_2$, $CO(O)H$, $C(O)H$, OH , CN, N_3 , NO_2 , F, Cl, Br and I; or

R^8 , at each occurrence, is independently selected from the group consisting of C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{1-6} haloalkyl, aryl, heterocyclyl, cycloalkyl, and cycloalkenyl; wherein the R^8 C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, and C_{1-6} haloalkyl are optionally substituted with one, two, three, four, five, or six substituents independently selected from the group consisting of R^{16} , OR^{16} , SR^{16} , $S(O)R^{16}$, SO_2R^{16} , $C(O)R^{16}$, $CO(O)R^{16}$, $OC(O)R^{16}$, $OC(O)OR^{16}$, NH_2 , NHR^{16} , $N(R^{16})_2$, $NHC(O)R^{16}$, $NR^{16}C(O)R^{16}$, $NHS(O)_2R^{16}$, $NR^{16}S(O)_2R^{16}$, $NHC(O)OR^{16}$, $NR^{16}C(O)OR^{16}$, $NHC(O)NH_2$, $NHC(O)NHR^{16}$, $NHC(O)N(R^{16})_2$, $NR^{16}C(O)NHR^{16}$, $NR^{16}C(O)N(R^{16})_2$, $C(O)NH_2$, $C(O)NHR^{16}$, $C(O)N(R^{16})_2$, $C(O)NHOH$, $C(O)NHOR^{16}$, $C(O)NHSO_2R^{16}$, $C(O)NR^{16}SO_2R^{16}$, SO_2NH_2 , SO_2NHR^{16} , $SO_2N(R^{16})_2$, $CO(O)H$, $C(O)H$, OH , CN, N_3 , NO_2 , F, Cl, Br and I; wherein the R^8 aryl, heterocyclyl, cycloalkyl, and cycloalkenyl are optionally substituted with one, two, or three substituents independently selected from the group consisting of C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{1-6} haloalkyl, NH_2 , $C(O)NH_2$, SO_2NH_2 , $C(O)H$, (O), OH, CN, NO_2 , OCF_3 , OCF_2CF_3 , F, Cl, Br and I;

R^9 is selected from the group consisting of C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{1-6} haloalkyl, cycloalkyl, phenyl and $(CH_2)_{1-4}$ phenyl; and

R^{10} and R^{11} , at each occurrence, are each independently selected from the group consisting of hydrogen, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{3-6} cycloalkyl, C_{1-6} haloalkyl, phenyl and $(CH_2)_{1-4}$ -phenyl; or

R^{10} and R^{11} , or R^{10} and R^9 , together with the atom to which each is attached are combined to form a heterocyclyl;

R^k , at each occurrence, is independently selected from the group consisting of C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{3-7} heterocycloalkyl, C_{3-7} cycloalkyl and C_{1-6} haloalkyl; wherein the R^k C_{1-6} alkyl, C_{2-6} alkenyl, and C_{2-6} alkynyl are optionally substituted with aryl, heterocyclyl, cycloalkyl, or cycloalkenyl;

5 R^{12} and R^{13} , at each occurrence, are each independently selected from the group consisting of hydrogen, C_{1-4} alkyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{1-4} haloalkyl and $(CH_2)_4$ phenyl;

R^{14} , at each occurrence, is independently selected from the group consisting of C_{1-4} alkyl, C_{2-4} alkenyl, C_{2-4} alkynyl and C_{1-4} haloalkyl;

10 R^{12} and R^{13} , or R^{12} and R^{14} , at each occurrence, together with the atom to which each is attached, are optionally combined to form a heterocyclyl;

R^{15} , at each occurrence, is independently selected from the group consisting of C_{1-4} alkyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{1-4} haloalkyl, C_{1-4} hydroxyalkyl, aryl, heterocyclyl, cycloalkyl, and cycloalkenyl; wherein the R^{15} C_{1-4} alkyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{1-4} haloalkyl, and C_{1-4} hydroxyalkyl are optionally substituted with one, two, or three substituents independently selected from the group consisting of C_{1-4} alkyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{1-4} haloalkyl, C_{1-4} hydroxyalkyl, C_{1-4} alkoxy, aryl, heterocycloalkyl, heterocycloalkenyl, heteroaryl, cycloalkyl, and cycloalkenyl, NH_2 , $C(O)NH_2$, SO_2NH_2 , $C(O)H$, $C(O)OH$, (O), OH, CN, NO_2 , OCF_3 , OCF_2CF_3 , F, Cl, Br and I;

20 R^{16} , at each occurrence, is independently selected from the group consisting of C_{1-4} alkyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{1-4} haloalkyl, C_{1-4} hydroxyalkyl, aryl, heterocycloalkyl, heterocycloalkenyl, heteroaryl, cycloalkyl, and cycloalkenyl; wherein the R^{16} C_{1-4} alkyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{1-4} haloalkyl, and C_{1-4} hydroxyalkyl are optionally substituted with one substituent independently selected from the group consisting of OCH_3 , $OCH_2CH_2OCH_3$, and

25 $OCH_2CH_2NHCH_3$;

q is 1, 2, or 3;

s is 0, 1, 2, or 3;

r is 0, 1, 2, or 3;

wherein the sum of s and r is 0, 1, or 2;

30 m is 0, 1, 2, or 3;

n is 0, 1, 2, 3, 4, 5, or 6; and

p is 0, 1, or 2.

35 2. The compound of claim 1, or a therapeutically acceptable salt thereof, wherein

Y^1 is pyrrolyl, pyrazolyl, or triazolyl.

3. The compound of claim 1, or a therapeutically acceptable salt thereof,
wherein

Y^1 is pyridinyl or phenyl.

5 4. The compound of claim 1, or a therapeutically acceptable salt thereof,
wherein

X is benzo[d]thiazolyl; which is optionally substituted with one, two, three or
four R^4 .

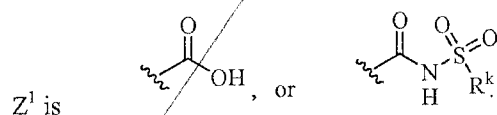
10 5. The compound of claim 2, or a therapeutically acceptable salt thereof,
wherein

X is benzo[d]thiazolyl; which is optionally substituted with one, two, three or
four R^4 .

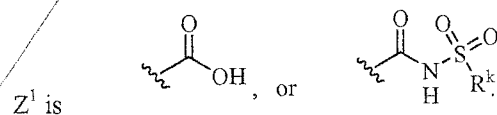
15 6. The compound of claim 3, or a therapeutically acceptable salt thereof,
wherein

wherein
X is benzo[d]thiazolyl; which is optionally substituted with one, two, three or
four R^4 .

20 7. The compound of claim 2, or a therapeutically acceptable salt thereof,
wherein



25 8. The compound of claim 3, or a therapeutically acceptable salt thereof,
wherein



30 9. The compound of claim 2, or a therapeutically acceptable salt thereof,
wherein

L^1 is $(CR^6R^7)_q$; and

Y^2 is selected from the group consisting of C_{3-7} cycloalkyl, C_{4-7} cycloalkenyl,
phenyl, C_{5-6} heteroaryl, and C_{3-7} heterocycloalkyl; wherein

R^6 and R^7 , at each occurrence, are R^{15} or hydrogen; and
 q is 1, 2, or 3.

10. The compound of claim 3, or a therapeutically acceptable salt thereof,
 5 wherein

L^1 is $(CR^6R^7)_q$; and

Y^2 is selected from the group consisting of C_{3-7} cycloalkyl, C_{4-7} cycloalkenyl,
 phenyl, C_{5-6} heteroaryl, and C_{3-7} heterocycloalkyl; wherein

R^6 and R^7 , at each occurrence, are R^{15} or hydrogen; and
 10 q is 1, 2, or 3.

11. The compound of claim 2, or a therapeutically acceptable salt thereof,
 wherein

L^1 is selected from the group consisting of $(CR^6R^7)_s-C(O)NR^{6A}-(CR^6R^7)_t$, and
 15 $(CR^6R^7)_s-S(O)_2NR^{6A}-(CR^6R^7)_t$;

s is 0;

t is 0 or 1;

R^{6A} is independently selected from the group consisting of hydrogen, and C_{1-6}
 alkyl; and

20 R^6 and R^7 , at each occurrence, are hydrogen.

12. The compound of claim 3, or a therapeutically acceptable salt thereof,
 wherein

L^1 is selected from the group consisting of $(CR^6R^7)_s-NR^{6A}C(O)-(CR^6R^7)_t$, $(CR^6R^7)_s-$
 25 $C(O)NR^{6A}-(CR^6R^7)_t$, $(CR^6R^7)_s-NR^{6A}-(CR^6R^7)_t$, $(CR^6R^7)_s-S(O)_2NR^{6A}-(CR^6R^7)_t$, and $(CR^6R^7)_s-$
 $NR^{6A}S(O)_2-(CR^6R^7)_t$;

s is 0;

t is 0 or 1;

R^{6A} is independently selected from the group consisting of hydrogen, and C_{1-6}
 30 alkyl; and

R^6 and R^7 , at each occurrence, are hydrogen.

13. The compound of claim 1, or a pharmaceutically acceptable salt thereof,
 selected from the group consisting of

35 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-
 benzyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(pyridin-4-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(pyridin-3-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 5 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(4-hydroxybenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(1-phenylethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 10 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{4-[2-(dimethylamino)ethoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 3-(1-benzyl-1H-pyrazol-4-yl)-6-{8-[(5,6-difluoro-1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(4-fluorophenyl)ethyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 15 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-chlorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 3-(1-benzyl-1H-pyrazol-4-yl)-6-{8-[(6-fluoro-1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl}pyridine-2-carboxylic acid;
- 3-(1-benzyl-1H-pyrazol-4-yl)-6-{8-[(6-methoxy-1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl}pyridine-2-carboxylic acid;
- 20 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-3,5-dimethyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-methylbenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 25 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-methoxybenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(4-chlorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-(benzyloxy)benzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 30 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-hydroxybenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{3-[2-(dimethylamino)ethoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 35 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-3-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 3-(1-benzyl-1H-pyrazol-4-yl)-6-{8-[(7-chloro-1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl}pyridine-2-carboxylic acid;
- 5 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{6-(pyrrolidin-1-yl)pyridin-2-yl}methyl)-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-phenyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-cyanobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 10 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-chlorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(pyridin-2-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 15 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-3-cyano-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(naphthalen-2-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[3-(3-methyl-1,2,4-oxadiazol-5-yl)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 20 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-benzyl-3-(hydroxymethyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(benzyloxy)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 25 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-methyl-1-{6-(pyrrolidin-1-yl)pyridin-2-yl}methyl)-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-benzyl-3-(ethoxycarbonyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 30 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[3-(dimethylamino)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-3-carboxy-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-hydroxybenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 35 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2,3-dihydro-1H-inden-1-yl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-phenylethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3,4-dihydro-2H-chromen-4-yl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 5 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-(dimethylamino)ethoxy)benzyl]-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-fluorobenzyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 10 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(cyclohexylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(cyclohexylmethyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(4-{[3-(dimethylamino)propyl]amino}-3-nitrobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 15 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(4-fluoro-3-nitrobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-[2-(morpholin-4-yl)ethoxy]benzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-[3-(dimethylamino)propoxy]benzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 20 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-(pyridin-4-ylmethoxy)benzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-(dimethylamino)ethoxy)benzyl]-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 25 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(tetrahydro-2H-pyran-4-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-[3-(dimethylamino)prop-1-yn-1-yl]benzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2,3-difluorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 30 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3,5-difluorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2,5-difluorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 35 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2,6-difluorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(tetrahydro-2H-pyran-3-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-nitrobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 5 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(biphenyl-3-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2,2-dimethylpropyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 10 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-cyclohexylethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-[3-(trifluoromethyl)benzyl]-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(biphenyl-2-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 15 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(cyclopentylmethyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-formylbenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-5-phenyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 20 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[5-methyl-1-(tetrahydro-2H-pyran-3-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(1-phenylcyclohexyl)methyl]-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 25 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{3-[(dimethylamino)methyl]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-[3-(methylsulfonyl)benzyl]-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(cyclohexylmethyl)-5-cyclopropyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 30 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3,5-di-tert-butylbenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-[2-(morpholin-4-ylsulfonyl)benzyl]-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 35 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-[(4,4-difluorocyclohexyl)methyl]-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{{1-(trifluoromethyl)cyclohexyl}methyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(diphenylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 5 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(morpholin-4-yl)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[3-(morpholin-4-yl)-1-phenylpropyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 10 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{{1-(tert-butoxycarbonyl)piperidin-4-yl}methyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-methyl-1-{2-[2-(morpholin-4-yl)ethoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(dimethylamino)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 15 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-methyl-1-{2-[3-(morpholin-4-yl)propoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{5-methyl-1-[(1-methylcyclohexyl)methyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 20 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{{2-(4-chlorophenyl)-4,4-dimethylcyclohex-1-en-1-yl}methyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(cyclohexylmethyl)-5-ethyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 25 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{{[(1R,2R,5R)-6,6-dimethylbicyclo[3.1.1]hept-2-yl]methyl}-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{{[(1S,2R,5S)-6,6-dimethylbicyclo[3.1.1]hept-2-yl]methyl}-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 30 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{{1-(2-methylpropyl)cyclohexyl}methyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{{1-(2-methoxyethyl)cyclohexyl}methyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 35 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(2-methoxyethoxy)benzyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethyl)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(1R,2R,4R)-bicyclo[2.2.1]hept-5-en-2-ylmethyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-{(3,3-dimethylcyclohexyl)methyl}-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-methoxy-1-phenylpropyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(4-methoxy-1-phenylbutyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-methoxy-2-oxo-1-phenylethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-cyclohexyl-1-phenylethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(3-methoxypropyl)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{2-[3-hydroxy-2-(hydroxymethyl)-2-methylpropoxy]benzyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{5-methyl-1-[2-(tetrahydro-2H-pyran-4-ylmethoxy)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(1,4-dioxan-2-ylmethoxy)benzyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethoxy)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(4-phenoxyphenyl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(3-phenoxyphenyl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[3-(4-nitrophenoxy)phenyl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[3-(4-chlorophenoxy)phenyl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(3-benzylphenyl)pyridine-2-carboxylic acid;

- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[3-(cyclohexylmethyl)-2-methylphenyl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(4-methyl-3-phenoxyphenyl)pyridine-2-carboxylic acid;
- 5 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(2-methyl-5-phenoxyphenyl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(2-methyl-3-phenoxyphenyl)pyridine-2-carboxylic acid;
- 10 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(2-methyl-4-phenoxyphenyl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[3-(cyclohexylmethoxy)-2-methylphenyl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[4-(cyclohexyloxy)-2-methylphenyl]pyridine-2-carboxylic acid;
- 15 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[3-(cyclohexyloxy)-2-methylphenyl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[4-(cyclohexylmethoxy)-2-methylphenyl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[2-cyano-3-(cyclohexyloxy)phenyl]pyridine-2-carboxylic acid;
- 20 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[2-chloro-3-(cyclohexyloxy)phenyl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[3-(cyclohexylamino)-2-methylphenyl]pyridine-2-carboxylic acid;
- 25 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[3-(cyclohexyloxy)-2-fluorophenyl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[3-(cyclohexyloxy)-2-(trifluoromethyl)phenyl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{3-[(3,3-dimethylcyclohexyl)oxy]-2-methylphenyl}pyridine-2-carboxylic acid;
- 30 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-1H-1,2,3-triazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-benzyl-5-(ethoxycarbonyl)-2-methyl-1H-pyrrol-3-yl]pyridine-2-carboxylic acid;
- 35 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-5-carboxy-2-methyl-1H-pyrrol-3-yl)pyridine-2-carboxylic acid;

- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-1H-pyrrol-3-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(4-methylphenyl)sulfonyl]-1H-pyrrol-3-yl}pyridine-2-carboxylic acid;
- 5 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-1H-pyrrol-3-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(phenylsulfonyl)-1H-pyrrol-3-yl]pyridine-2-carboxylic acid;
- 10 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-5-cyano-2-methyl-1H-pyrrol-3-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[5-cyano-1-(cyclohexylmethyl)-2-methyl-1H-pyrrol-3-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-cyano-2-methyl-1-{[1-(piperidin-1-yl)cyclohexyl]methyl}-1H-pyrrol-3-yl)pyridine-2-carboxylic acid;
- 15 acid;
- 6,6'-bis[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3,3'-bipyridine-2,2'-dicarboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(2-benzyl-1,2,3,4-tetrahydroisoquinolin-6-yl)pyridine-2-carboxylic acid;
- 20 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethyl)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(piperidin-4-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[2-(pyridin-2-ylmethyl)-1,2,3,4-tetrahydroisoquinolin-6-yl]pyridine-2-carboxylic acid;
- 25 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[2-(cyclohexylmethyl)-1,2,3,4-tetrahydroisoquinolin-6-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-2'-(cyclohexyloxy)-3'-methyl-3,4'-bipyridine-2-carboxylic acid;
- 30 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-2'-(cyclohexyloxy)-3,4'-bipyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-2'-phenoxy-3,4'-bipyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-1H-pyrrolo[2,3-b]pyridin-3-yl)pyridine-2-carboxylic acid;
- 35 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3'-methyl-2'-phenoxy-3,4'-bipyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3'-methyl-2'-[methyl(phenyl)amino]-3,4'-bipyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(methoxymethyl)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

5 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[3,3-dimethyl-1-(morpholin-4-yl)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethyl)-3,3-dimethylcyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-

10 carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(3-methoxypropyl)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

N-(1,3-benzothiazol-2-yl)-2-{5-(1-{[1-(2-methoxyethyl)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)-6-[(methylsulfonyl)carbamoyl]pyridin-2-yl}-1,2,3,4-

15 tetrahydroisoquinoline-8-carboxamide;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(tetrahydro-2H-pyran-2-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[(1R,2R,3R,5S)-2-(2-methoxyethyl)-6,6-dimethylbicyclo[3.1.1]hept-3-yl]methyl}-1H-

20 pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethoxy)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[1-cyclohexyl-3-(morpholin-4-yl)propyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

25 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-1H-indazol-5-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[5-methyl-1-(tetrahydro-2H-pyran-2-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[5-methyl-1-({1-[3-(morpholin-4-yl)propoxy]cycloheptyl}methyl)-1H-pyrazol-4-yl]pyridine-2-

30 carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{2-methyl-3-[methyl(phenyl)amino]phenyl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(1-methoxy-3,3-dimethylcyclohexyl)methyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic

35 acid;

6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-
{[(1S,2R,5S)-6,6-dimethylbicyclo[3.1.1]hept-2-yl]methyl}-5-methyl-1H-1,2,3-triazol-4-
yl)pyridine-2-carboxylic acid;

5 6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-3-{3-[(3,3-
dimethylcyclohexyl)(methyl)amino]-2-methylphenyl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-
methyl-1-{[1-(morpholin-4-yl)cyclohexyl]methyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic
acid;

10 6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-2'-
[cyclohexyl(methyl)amino]-3'-methyl-3,4'-bipyridine-2-carboxylic acid;

N-(1,3-benzothiazol-2-yl)-2-(5-{1-[(1-methoxy-3,3-dimethylcyclohexyl)methyl]-5-
methyl-1H-pyrazol-4-yl}-6-[(methylsulfonyl)carbamoyl]pyridin-2-yl)-1,2,3,4-
tetrahydroisoquinoline-8-carboxamide;

15 6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-1-methyl-3,4-dihydroisoquinolin-2(1H)-yl]-3-
{5-methyl-1-[(1-methylcyclohexyl)methyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-3'-cyano-2'-
[cyclohexyl(methyl)amino]-3,4'-bipyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(1-
methoxycyclohexyl)methyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

20 6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(1-
methoxy-3,3-dimethylcyclohexyl)methyl]-5-methyl-1H-1,2,3-triazol-4-yl}pyridine-2-
carboxylic acid;

25 6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-({1-[2-
(1,1-dioxidothiomorpholin-4-yl)ethoxy]cyclohexyl}methyl)-5-methyl-1H-pyrazol-4-
yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-cyano-
2-methyl-1-{[1-(morpholin-4-yl)cyclohexyl]methyl}-1H-pyrrol-3-yl)pyridine-2-carboxylic
acid;

30 6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-
hydroxyethoxy)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-
(2,3-dimethoxypropoxy)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-
carboxylic acid;

35 N-(1,3-benzothiazol-2-yl)-2-{5-[5-cyano-1-(cyclohexylmethyl)-2-methyl-1H-pyrrol-
3-yl]-6-[(methylsulfonyl)carbamoyl]pyridin-2-yl}-1,2,3,4-tetrahydroisoquinoline-8-
carboxamide;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-cyano-1-[[1-(2-methoxyethoxy)cycloheptyl]methyl]-2-methyl-1H-pyrrol-3-yl)pyridine-2-carboxylic acid;

5 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-[[1-(1,4-dioxan-2-ylmethoxy)cycloheptyl]methyl]-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[5-methyl-1-({1-[2-(morpholin-4-yl)-2-oxoethoxy]cyclohexyl}methyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

10 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-[[1-(2,3-dihydroxypropoxy)cycloheptyl]methyl]-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

15 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-cyano-1-[[1-(dimethylamino)cyclohexyl]methyl]-2-methyl-1H-pyrrol-3-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{3-[(cyclohexylcarbonyl)(methyl)amino]-2-methylphenyl}pyridine-2-carboxylic acid;

20 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-({3,3-dimethyl-1-[2-(methylsulfonyl)ethoxy]cyclohexyl}methyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(2-methyl-3-{methyl[(1-methylcyclohexyl)carbonyl]amino}phenyl)pyridine-2-carboxylic acid;

25 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-[[1-(2-methoxyethoxy)-3,3-dimethylcyclohexyl]methyl]-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{2-cyano-3-[(cyclohexylsulfonyl)(methyl)amino]phenyl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{2-cyano-3-[2-(tricyclo[3.3.1.1^{3,7}]dec-1-yl)pyrrolidin-1-yl]phenyl}pyridine-2-carboxylic acid;

30 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3'-methyl-2'-(piperidin-1-yl)-3,4'-bipyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(1-cyclohexyl-3-methoxypropyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

35 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-({3,3-dimethyl-1-[2-(methylamino)ethoxy]cyclohexyl}methyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{5-methyl-1-[(2,6,6-trimethyltetrahydro-2H-pyran-2-yl)methyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-1H-indazol-4-yl)pyridine-2-carboxylic acid; and

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-1H-pyrrolo[2,3-c]pyridin-3-yl)pyridine-2-carboxylic acid.

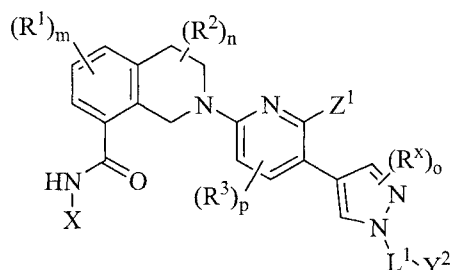
14. A composition for treating bladder cancer, brain cancer, breast cancer, bone marrow cancer, cervical cancer, chronic lymphocytic leukemia, colorectal cancer, esophageal cancer, hepatocellular cancer, lymphoblastic leukemia, follicular lymphoma, lymphoid malignancies of T-cell or B-cell origin, melanoma, myelogenous leukemia, myeloma, oral cancer, ovarian cancer, non-small cell lung cancer, chronic lymphocytic leukemia, myeloma, prostate cancer, small cell lung cancer or spleen cancer, said composition comprising an excipient and a therapeutically effective amount of a compound of claim 1 or a pharmaceutically acceptable salt thereof.

15. Use of a therapeutically effective amount of a compound of claim 1, or a pharmaceutically acceptable salt thereof, in the manufacture of a medicament for treating bladder cancer, brain cancer, breast cancer, bone marrow cancer, cervical cancer, chronic lymphocytic leukemia, colorectal cancer, esophageal cancer, hepatocellular cancer, lymphoblastic leukemia, follicular lymphoma, lymphoid malignancies of T-cell or B-cell origin, melanoma, myelogenous leukemia, myeloma, oral cancer, ovarian cancer, non-small cell lung cancer, chronic lymphocytic leukemia, myeloma, prostate cancer, small cell lung cancer or spleen cancer in a patient.

16. Use of a therapeutically effective amount of the compound of claim 1, or a pharmaceutically acceptable salt thereof, and a therapeutically effective amount of one additional therapeutic agent or more than one additional therapeutic agent, in the manufacture of a medicament for treating bladder cancer, brain cancer, breast cancer, bone marrow cancer, cervical cancer, chronic lymphocytic leukemia, colorectal cancer, esophageal cancer, hepatocellular cancer, lymphoblastic leukemia, follicular lymphoma, lymphoid malignancies of T-cell or B-cell origin, melanoma, myelogenous leukemia, myeloma, oral cancer, ovarian cancer, non-small cell lung cancer, chronic lymphocytic leukemia, myeloma, prostate cancer, small cell lung cancer or spleen cancer in a patient.

Claims

1. A compound having Formula (III)



Formula (III),

or a therapeutically acceptable salt thereof, wherein

X is benzo[d]thiazolyl optionally substituted with one or two R⁴;

R^x, at each occurrence, is independently selected from the group consisting of R⁵, CO(O)R⁵, CO(O)H, CN, F, Cl, Br and I;

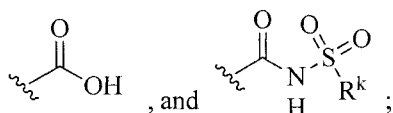
L¹ is selected from the group consisting of (CR⁶R⁷)_q, (CR⁶R⁷)_s-C(O)-(CR⁶R⁷)_r, and (CR⁶R⁷)_s-S(O)₂-(CR⁶R⁷)_r; and

Y² is selected from the group consisting of C₃₋₁₁ branched chain alkyl, C₃₋₇ cycloalkyl, C₄₋₇ cycloalkenyl, phenyl, and C₃₋₇ heterocyclyl; wherein the C₃₋₇ cycloalkyl, C₄₋₇ cycloalkenyl, phenyl, and C₃₋₇ heterocyclyl are optionally fused to one benzene ring; wherein Y² is optionally substituted with one, two, or three substituents independently selected from the group consisting of R⁸, OR⁸, SO₂R⁸, CO(O)R⁸, NHR⁸, N(R⁸)₂, C(O)H, OH, CN, NO₂, F, Cl, Br and I; or

L¹ is a bond; and

Y² is selected from the group consisting of C₃₋₇ cycloalkyl, phenyl, and C₃₋₇ heterocyclyl; wherein the C₃₋₇ cycloalkyl, phenyl, and C₃₋₇ heterocyclyl represented by Y² are optionally fused to one benzene ring; wherein each Y² and each ring fused to Y² are optionally substituted with one substituent independently selected from the group consisting of R⁸ and C(O)NHR⁸;

Z¹ is selected from the group consisting of



R¹ is absent;

R^2 , at each occurrence, is independently C_{1-6} alkyl;
 R^3 is absent;
 R^4 , at each occurrence, is independently selected from the group consisting of OR^{12} and halogen;
 R^5 , at each occurrence, is independently selected from the group consisting of C_{1-6} alkyl, C_{1-6} haloalkyl, C_{1-6} hydroxyalkyl, aryl, and cycloalkyl;
 R^6 and R^7 , at each occurrence, are each independently selected from the group consisting of hydrogen, R^{15} , and $CO(O)R^{15}$;
 R^8 , at each occurrence, is independently selected from the group consisting of C_{1-6} alkyl, C_{2-6} alkynyl, aryl, heterocyclyl, and cycloalkyl; wherein the R^8 C_{1-6} alkyl, and C_{2-6} alkynyl are optionally substituted with one, two, or three substituents independently selected from the group consisting of R^{16} , OR^{16} , SO_2R^{16} , $C(O)R^{16}$, $N(R^{16})_2$, OH, F, Cl, Br and I; wherein the R^8 aryl and heterocyclyl are optionally substituted with one substituent independently selected from the group consisting of C_{1-6} alkyl, F, Cl, Br and I;
 R^k , at each occurrence, is independently C_{1-6} alkyl;
 R^{12} is C_{1-4} alkyl;
 R^{15} , at each occurrence, is independently selected from the group consisting of C_{1-4} alkyl, and aryl; wherein the R^{15} C_{1-4} alkyl is optionally substituted with one substituent independently selected from the group consisting C_{1-4} alkoxy, and heterocycloalkyl;
 R^{16} , at each occurrence, is independently selected from the group consisting of C_{1-4} alkyl, aryl, heterocycloalkyl, and heteroaryl;
 q is 1, 2, or 3;
 s is 0;
 r is 0, or 1;
 m is 0;
 n is 0, or 1;
 o is 0, 1, or 2; and
 p is 0.

2. The compound or therapeutically acceptable salt of claim 1, wherein L^1 is $(CR^6R^7)_q$; and

Y² is selected from the group consisting of C₃₋₇ cycloalkyl, C₄₋₇ cycloalkenyl, phenyl, and C₃₋₇ heterocycloalkyl; wherein

R⁶ and R⁷, at each occurrence, are R¹⁵ or hydrogen; and

q is 1, 2, or 3.

3. The compound or therapeutically acceptable salt of claim 1, selected from the group consisting of

- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(pyridin-4-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(pyridin-3-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(4-hydroxybenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(1-phenylethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{4-[2-(dimethylamino)ethoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 3-(1-benzyl-1H-pyrazol-4-yl)-6-{8-[(5,6-difluoro-1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(4-fluorophenyl)ethyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-chlorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 3-(1-benzyl-1H-pyrazol-4-yl)-6-{8-[(6-fluoro-1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl}pyridine-2-carboxylic acid;
- 3-(1-benzyl-1H-pyrazol-4-yl)-6-{8-[(6-methoxy-1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-3,5-dimethyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-methylbenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-methoxybenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(4-chlorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[3-(benzyloxy)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-hydroxybenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{3-[2-(dimethylamino)ethoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-3-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

3-(1-benzyl-1H-pyrazol-4-yl)-6-{8-[(7-chloro-1,3-benzothiazol-2-yl)carbamoyl]-3,4-dihydroisoquinolin-2(1H)-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[6-(pyrrolidin-1-yl)pyridin-2-yl]methyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-phenyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-cyanobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-chlorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(pyridin-2-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-3-cyano-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(naphthalen-2-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[3-(3-methyl-1,2,4-oxadiazol-5-yl)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-benzyl-3-(hydroxymethyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(benzyloxy)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-methyl-1-{[6-(pyrrolidin-1-yl)pyridin-2-yl]methyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-benzyl-3-(ethoxycarbonyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[3-(dimethylamino)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-3-carboxy-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-hydroxybenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2,3-dihydro-1H-inden-1-yl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-phenylethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3,4-dihydro-2H-chromen-4-yl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{2-[2-(dimethylamino)ethoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-fluorobenzyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(cyclohexylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(cyclohexylmethyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(4-{[3-(dimethylamino)propyl]amino}-3-nitrobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(4-fluoro-3-nitrobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{2-[2-(morpholin-4-yl)ethoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{2-[3-(dimethylamino)propoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(pyridin-4-ylmethoxy)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{2-[2-(dimethylamino)ethoxy]benzyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(tetrahydro-2H-pyran-4-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{2-[3-(dimethylamino)prop-1-yn-1-yl]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2,3-difluorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3,5-difluorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2,5-difluorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2,6-difluorobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(tetrahydro-2H-pyran-3-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-nitrobenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(biphenyl-3-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2,2-dimethylpropyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-cyclohexylethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[3-(trifluoromethyl)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(biphenyl-2-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(cyclopentylmethyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-formylbenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-benzyl-5-phenyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[5-methyl-1-(tetrahydro-2H-pyran-3-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(1-phenylcyclohexyl)methyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{3-[(dimethylamino)methyl]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[3-(methylsulfonyl)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(cyclohexylmethyl)-5-cyclopropyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3,5-di-tert-butylbenzyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(morpholin-4-ylsulfonyl)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(4,4-difluorocyclohexyl)methyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{1-(trifluoromethyl)cyclohexyl}methyl)-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(diphenylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(morpholin-4-yl)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[3-(morpholin-4-yl)-1-phenylpropyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{1-(tert-butoxycarbonyl)piperidin-4-yl}methyl)-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-methyl-1-{2-[2-(morpholin-4-yl)ethoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(dimethylamino)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-methyl-1-{2-[3-(morpholin-4-yl)propoxy]benzyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{5-methyl-1-[(1-methylcyclohexyl)methyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{2-(4-chlorophenyl)-4,4-dimethylcyclohex-1-en-1-yl}methyl)-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(cyclohexylmethyl)-5-ethyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[(1R,2R,5R)-6,6-dimethylbicyclo[3.1.1]hept-2-yl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[(1S,2R,5S)-6,6-dimethylbicyclo[3.1.1]hept-2-yl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{1-(2-methylpropyl)cyclohexyl}methyl)-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{1-(2-methoxyethyl)cyclohexyl}methyl)-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(2-methoxyethoxy)benzyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethyl)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(1R,2R,4R)-bicyclo[2.2.1]hept-5-en-2-ylmethyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(3,3-dimethylcyclohexyl)methyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(3-methoxy-1-phenylpropyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(4-methoxy-1-phenylbutyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-methoxy-2-oxo-1-phenylethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(2-cyclohexyl-1-phenylethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(3-methoxypropyl)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{2-[3-hydroxy-2-(hydroxymethyl)-2-methylpropoxy]benzyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{5-methyl-1-[2-(tetrahydro-2H-pyran-4-ylmethoxy)benzyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[2-(1,4-dioxan-2-ylmethoxy)benzyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethoxy)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethyl)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(piperidin-4-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(methoxymethyl)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[3,3-dimethyl-1-(morpholin-4-yl)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethyl)-3,3-dimethylcyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(3-methoxypropyl)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- N-(1,3-benzothiazol-2-yl)-2-{5-(1-{[1-(2-methoxyethyl)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)-6-[(methylsulfonyl)carbamoyl]pyridin-2-yl}-1,2,3,4-tetrahydroisoquinoline-8-carboxamide;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(tetrahydro-2H-pyran-2-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[(1R,2R,3R,5S)-2-(2-methoxyethyl)-6,6-dimethylbicyclo[3.1.1]hept-3-yl]methyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethoxy)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[1-cyclohexyl-3-(morpholin-4-yl)propyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;
- 6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[5-methyl-1-(tetrahydro-2H-pyran-2-ylmethyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[5-methyl-1-({1-[3-(morpholin-4-yl)propoxy]cycloheptyl}methyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(1-methoxy-3,3-dimethylcyclohexyl)methyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(5-methyl-1-{{1-(morpholin-4-yl)cyclohexyl}methyl}-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

N-(1,3-benzothiazol-2-yl)-2-(5-{1-[(1-methoxy-3,3-dimethylcyclohexyl)methyl]-5-methyl-1H-pyrazol-4-yl}-6-[(methylsulfonyl)carbamoyl]pyridin-2-yl)-1,2,3,4-tetrahydroisoquinoline-8-carboxamide;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-1-methyl-3,4-dihydroisoquinolin-2(1H)-yl]-3-{5-methyl-1-[(1-methylcyclohexyl)methyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(1-methoxycyclohexyl)methyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-({1-[2-(1,1-dioxidothiomorpholin-4-yl)ethoxy]cyclohexyl}methyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-hydroxyethoxy)cyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2,3-dimethoxypropoxy)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(1,4-dioxan-2-ylmethoxy)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[5-methyl-1-({1-[2-(morpholin-4-yl)-2-oxoethoxy]cyclohexyl}methyl)-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2,3-dihydroxypropoxy)cycloheptyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-({3,3-dimethyl-1-[2-(methylsulfonyl)ethoxy]cyclohexyl}methyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-(1-{[1-(2-methoxyethoxy)-3,3-dimethylcyclohexyl]methyl}-5-methyl-1H-pyrazol-4-yl)pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-(1-cyclohexyl-3-methoxypropyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid;

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-[1-({3,3-dimethyl-1-[2-(methylamino)ethoxy]cyclohexyl}methyl)-5-methyl-1H-pyrazol-4-yl]pyridine-2-carboxylic acid; and

6-[8-(1,3-benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{5-methyl-1-[(2,6,6-trimethyltetrahydro-2H-pyran-2-yl)methyl]-1H-pyrazol-4-yl}pyridine-2-carboxylic acid.

4. 6-[8-(1,3-Benzothiazol-2-ylcarbamoyl)-3,4-dihydroisoquinolin-2(1H)-yl]-3-{1-[(1-methoxy-3,3-dimethylcyclohexyl)methyl]-5-methyl-1H-pyrazol-4-yl}pyridine-2-carboxylic acid or a therapeutically acceptable salt thereof.

5. A pharmaceutical composition comprising an excipient and a therapeutically effective amount of the compound or therapeutically acceptable salt of any one of claims 1-4.

6. Use of a therapeutic effective amount of a compound of any one of claims 1-4, or a pharmaceutically acceptable salt thereof, in the manufacture of a medicament for treating bladder cancer, brain cancer, breast cancer, bone marrow cancer, cervical cancer, chronic lymphocytic leukemia, colorectal cancer, esophageal cancer, hepatocellular cancer, lymphoblastic leukemia, follicular lymphoma, a lymphoid malignancy of T-cell or B-cell origin, melanoma, myelogenous leukemia, myeloma, oral cancer, ovarian cancer, non-small cell lung cancer, prostate cancer, small cell lung cancer or spleen cancer in a patient.

7. Use of a therapeutic effective amount of a compound of any one of claims 1-4, or a pharmaceutically acceptable salt thereof, in the manufacture of a medicament for

treating bladder cancer, brain cancer, breast cancer, bone marrow cancer, cervical cancer, chronic lymphocytic leukemia, colorectal cancer, esophageal cancer, hepatocellular cancer, lymphoblastic leukemia, follicular lymphoma, a lymphoid malignancy of T-cell or B-cell origin, melanoma, myelogenous leukemia, myeloma, oral cancer, ovarian cancer, non-small cell lung cancer, prostate cancer, small cell lung cancer or spleen cancer in a patient.