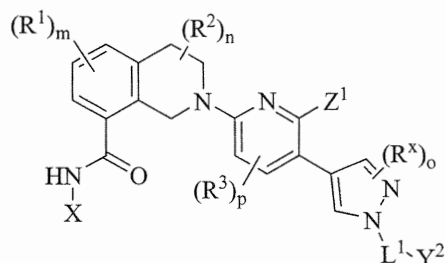


Claims

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;

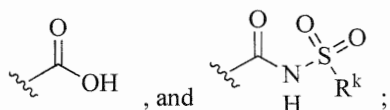
L¹ is selected from the group consisting of (CR⁶R⁷)_q, (CR⁶R⁷)_s-C(O)-(CR⁶R⁷)_r, and (CR⁶R⁷)_s-S(O)₂-(CR⁶R⁷)_r; and

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.

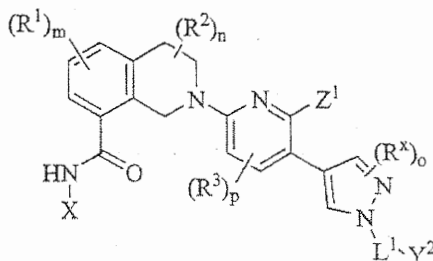
5. A pharmaceutical composition comprising an excipient and a therapeutically effective amount of the compound or therapeutically acceptable salt of any one of claims 1-4.

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, 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, 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.

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;

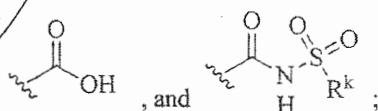
L¹ is selected from the group consisting of (CR⁶R⁷)_q, (CR⁶R⁷)_s-C(O)-(CR⁶R⁷)_r, and (CR⁶R⁷)_s-S(O)₂-(CR⁶R⁷)_r; and

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-[(3,6-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.