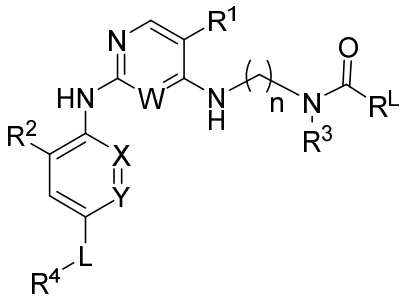
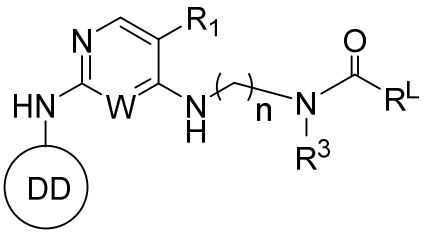
CLAIMS

What is claimed is:

1. A compound represented by Formula IA or Formula ID:



Formula IA



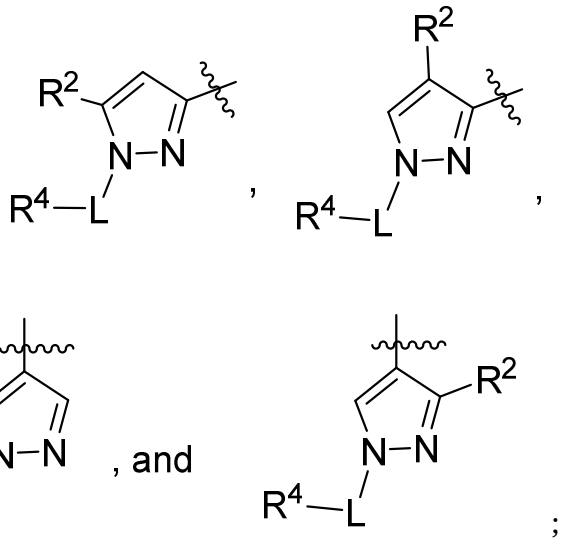
Formula ID

or a pharmaceutically acceptable salt, enantiomer, stereoisomer, or tautomer thereof, wherein:

X is CH or N;

Y is C(R^{33}) or N;

ring DD is selected from the group consisting of



W is CH or N;

R^1 is selected from the group consisting of halogen, cyano, C_1-C_5alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_5alkyl and C_3-C_5cycloalkyl may be optionally substituted by one, two or three independent occurrences of fluorine;

R^2 is selected from the group consisting of halogen, cyano, C_1-C_5alkyl, C_3-C_6cycloalkyl, C_2-C_5alkenyl, C_2-C_5alkynyl, C_1-C_5alkoxy, and C_1-C_5alkoxy-C_2-C_5alkyl, wherein each C_1-C_5alkyl, C_3-C_6cycloalkyl, C_2-C_5alkenyl, C_2-C_5alkynyl, and C_1-C_5alkoxy

may be optionally substituted by one, two, or three independent occurrences of fluorine or cyano;

R^3 is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^4 is selected from the group consisting of B, D, NR^6R^9, NR^6-$(C(R^{10})_2)_p$-NR^6R^9, and C(O)-D;

B is selected from an N-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen and heteroaryl, wherein B may be optionally substituted on one or more available carbons by R^7 and may be optionally substituted on an available nitrogen by R^9;

D is selected from a C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen and heteroaryl, wherein D may be optionally substituted on one or more available carbons by R^7 and may be optionally substituted on an available nitrogen by R^9;

each occurrence of R^7 is independently selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, cyano, and $(C(R^{10})_2)_h$-NR^6R^9, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^7 are joined together with the atom to which they are attached to form oxo;

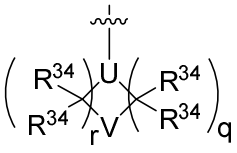
R^6 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, C_1-C_5alkoxy-C_2-C_5alkyl, $C(=O)R^5$, SO_2R^5, C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen, and heteroaryl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^9 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, C_1-C_5alkoxy-C_2-C_5alkyl, $C(=O)R^5$, SO_2R^5, C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen, and heteroaryl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^5 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, and heterocyclyl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

each occurrence of R^{10} is independently selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally

substituted by one or more independent occurrences of fluorine, or two R^{10} are joined together with the carbon to which they are attached to form a C_3-C_5cycloalkyl;

R^L is selected from the group consisting of C_1-C_6alkyl and C_3-C_6cycloalkyl, $NR^{11}R^{12}$, and , wherein each C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by 1, 2, or 3 independent occurrences of fluorine;

U is N or CR^{13};

V is selected from the group consisting of oxygen, $C(R^{34})_2$, and NR^6;

r is 0, 1, or 2;

q is 1, 2, or 3;

R^{11} is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{12} is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{13} is selected from H and C_1-C_3alkyl;

R^{33} is selected from the group consisting of H, halogen, C_1-C_6alkyl, and C_1-C_6alkoxy, wherein C_1-C_6alkyl and C_1-C_6alkoxy may be optionally substituted by one or more independent occurrences of fluorine;

each occurrence of R^{34} is independently selected from H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^{34} are joined together with the carbon to which they are attached to form a C_3-C_6cycloalkyl;

L is $-(C(R^{10})_2)_m-$;

h is 1, 2, or 3;

m is 0, 1, 2, or 3;

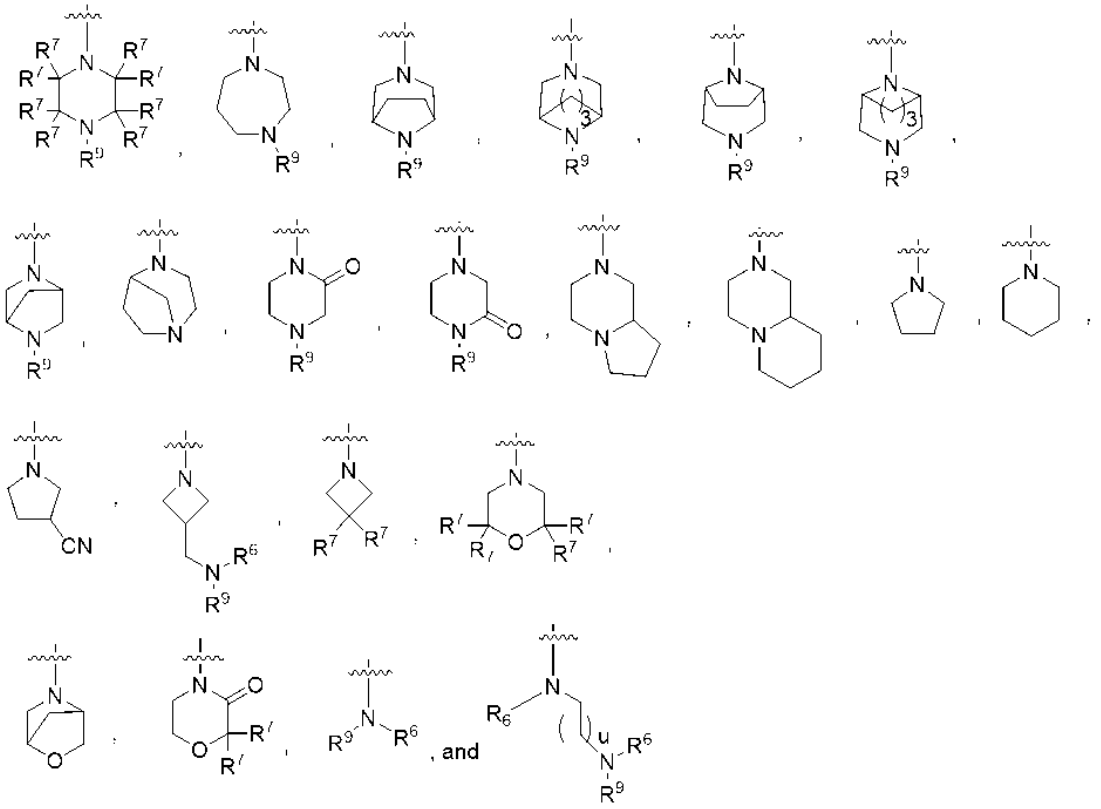
n is 2, 3, or 4; and

p is 2 or 3;

provided that (i) when r is 0 and q is 1, then U is not CR^{13} and V is not O, and when r or q is 1, then U is not N and V is not O or NR^6; (ii) when the compound is of Formula IA:

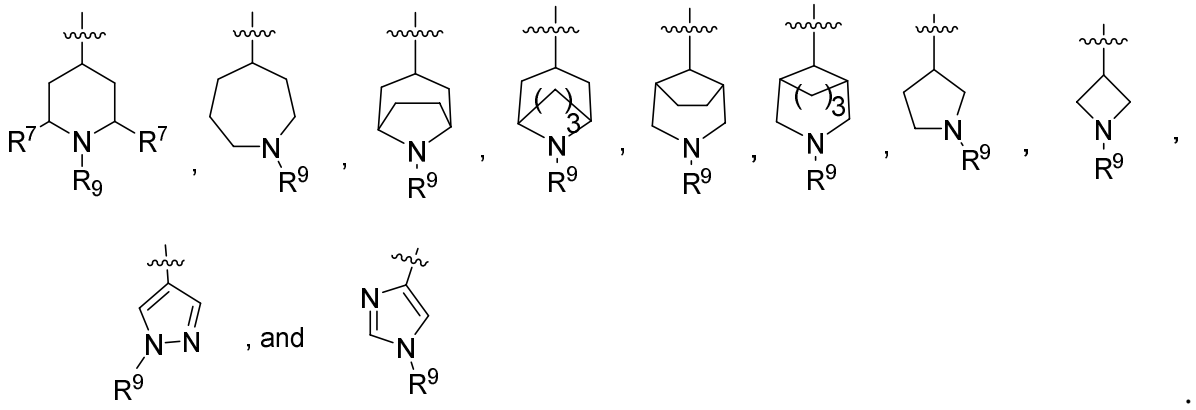
both of X and Y are not N; and (iii) when the compound is of Formula ID: when m is 0, R^4 is C-linked to the pyrazolyl ring, and when m is 1, R^4 is C-linked to L.

2. The compound of claim 1, wherein R^4 is selected from the group consisting of:



wherein u is 1 or 2.

3. The compound of claim 1, wherein R^4 is selected from the group consisting of:



.

4. The compound of any one of claims 1-3, wherein R^1 is selected from the group consisting of halogen, C_1-C_5alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_5alkyl and C_3-

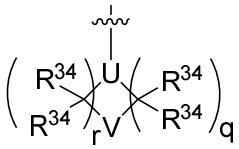
C_5cycloalkyl may be optionally substituted with one, two, or three independent occurrences of fluorine.

5. The compound of any one of claims 1-4, wherein R^2 is selected from the group consisting of C_3-C_5cycloalkyl, C_1-C_5alkyl, halogen, CN, C_2-C_5alkenyl, and C_2-C_5alknyl, wherein C_3-C_5cycloalkyl, C_1-C_5alkyl, C_2-C_5alkenyl, and C_2-C_5alknyl may be optionally substituted with one, two, or three independent occurrences of fluorine.

6. The compound of any one of claims 1-5, wherein R^3 is selected from the group consisting of H and C_1-C_3alkyl, wherein C_1-C_3alkyl may be optionally substituted by one or more independent occurrences of fluorine.

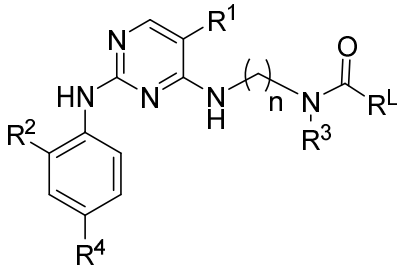
7. The compound of any one of claims 1-6, wherein R^L is selected from the group consisting of C_1-C_6alkyl and C_3-C_6cycloalkyl, each of which may be optionally substituted by 1, 2, or 3 independent occurrences of fluorine.

8. The compound of any one of claims 1-6, wherein R^L is $NR^{11}R^{12}$.

9. The compound of any one of claims 1-6, wherein R^L is 

, wherein U is N or CR^{13}; V is selected from the group consisting of oxygen, $C(R^{34})_2$, and NR^6; r is 0, 1, or 2; q is 1, 2, or 3; R^{13} is selected from H and C_1-C_3alkyl; and each occurrence of R^{34} is independently selected from H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^{34} are joined together with the carbon to which they are attached to form a C_3-C_6cycloalkyl; provided that when r is 0 and q is 1, then U is not CR^{13} and V is not O, and when r or q is 1, then U is not N and V is not O or NR^6.

10. The compound of claim 1 represented by:



Formula IB

or a pharmaceutically acceptable salt thereof, wherein:

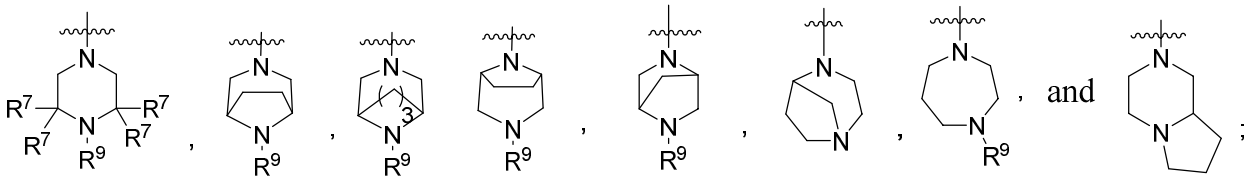
n is 2, 3, or 4;

R^1 is selected from the group consisting of halogen, cyano, C_1-C_5alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_5alkyl and C_3-C_5cycloalkyl may be optionally substituted by one, two or three independent occurrences of fluorine;

R^2 is selected from the group consisting of C_1-C_2alkyl and C_3-C_4cycloalkyl, and halogen, wherein C_1-C_2alkyl and C_3-C_4cycloalkyl may be optionally substituted by one, two or three independent occurrences of fluorine;

R^3 is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

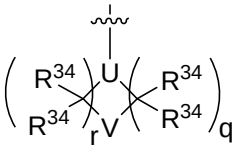
R^4 is selected from the group consisting of:



R^9 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, $C(=O)R^5$, SO_2R^5, C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen and heteroaryl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^5 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, and heterocyclyl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

each occurrence of R^7 is independently selected from the group consisting of H, C_1-C_6 alkyl, and C_3-C_6cycloalkyl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^7 are joined together with the atom to which they are attached to form oxo;

R^L is selected from the group consisting of C_1-C_6alkyl and C_3-C_6cycloalkyl, $NR^{11}R^{12}$, and  , wherein each C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by 1, 2, or 3 independent occurrences of fluorine;

U is N or CR^{13};

V is selected from the group consisting of oxygen, $C(R^{34})_2$, and NR^6;

r is 0, 1, or 2;

q is 1, 2, or 3;

R^6 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, $C(=O)R^5$, SO_2R^5, C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen and heteroaryl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{11} is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

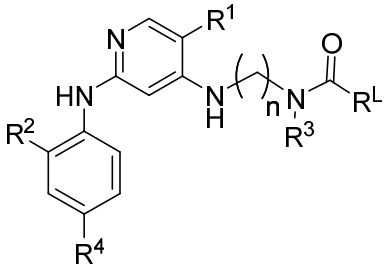
R^{12} is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{13} is selected from H and C_1-C_3alkyl; and

each occurrence of R^{34} is independently selected from H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^{34} are joined together with the carbon to which they are attached to form a C_3-C_6cycloalkyl;

provided that when r is 0 and q is 1, then U is not CR^{13} and V is not O, and when r or q is 1, then U is not N and V is not O or NR^6.

11. The compound of claim 1 represented by:



or a pharmaceutically acceptable salt thereof, wherein:

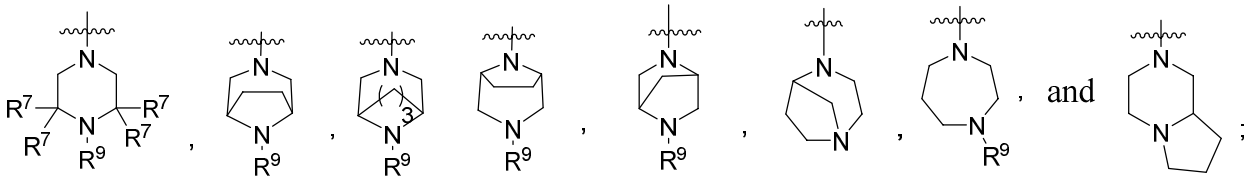
n is 2, 3, or 4;

R^1 is selected from the group consisting of halogen, cyano, C_1-C_5alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_5alkyl and C_3-C_5cycloalkyl may be optionally substituted by one, two or three independent occurrences of fluorine;

R^2 is selected from the group consisting of C_1-C_2alkyl and C_3-C_4cycloalkyl, and halogen, wherein C_1-C_2alkyl and C_3-C_4cycloalkyl may be optionally substituted by one, two or three independent occurrences of fluorine;

R^3 is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

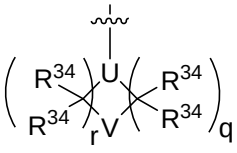
R^4 is selected from the group consisting of:



R^9 is independently selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, $C(=O)R^5$, SO_2R^5, C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen and heteroaryl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^5 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, and heterocyclyl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

each occurrence of R^7 is independently selected from the group consisting of H, C_1-C_6 alkyl, and C_3-C_6cycloalkyl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^7 are joined together with the atom to which they are attached to form oxo;

R^L is selected from the group consisting of C_1-C_6alkyl and C_3-C_6cycloalkyl, $NR^{11}R^{12}$, and  , wherein each C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by 1, 2, or 3 independent occurrences of fluorine;

U is N or CR^{13};

V is selected from the group consisting of oxygen, $C(R^{34})_2$, and NR^6;

r is 0, 1, or 2;

q is 1, 2, or 3;

R^6 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, $C(=O)R^5$, SO_2R^5, C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen and heteroaryl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{11} is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

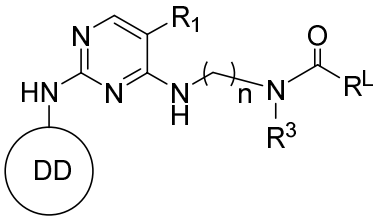
R^{12} is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{13} is selected from H and C_1-C_3alkyl; and

each occurrence of R^{34} is independently selected from H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^{34} are joined together with the carbon to which they are attached to form a C_3-C_6cycloalkyl;

provided that when r is 0 and q is 1, then U is not CR^{13} and V is not O, and when r or q is 1, then U is not N and V is not O or NR^6.

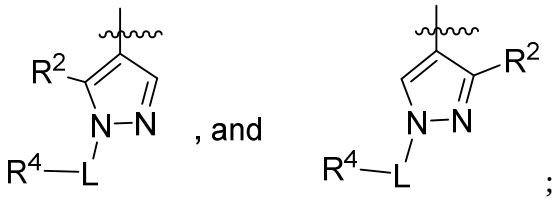
12. The compound of claim 1 represented by:



Formula IE

or a pharmaceutically acceptable salt thereof, wherein:

ring DD is selected from the group consisting of:



;

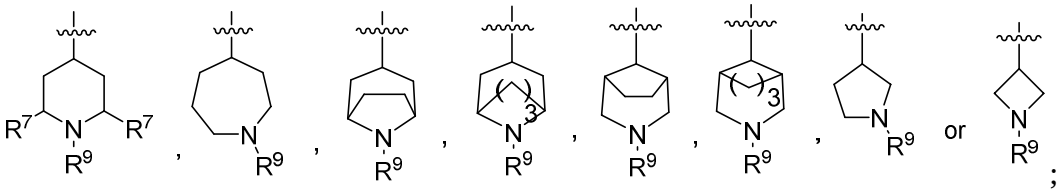
n is 2, 3, or 4;

R^1 is selected from the group consisting of halogen, cyano, C_1-C_5alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_5alkyl and C_3-C_5cycloalkyl may be optionally substituted by one, two or three independent occurrences of fluorine;

R^2 is selected from the group consisting of C_1-C_2alkyl and C_3-C_4cycloalkyl, and halogen, wherein C_1-C_2alkyl and C_3-C_4cycloalkyl may be optionally substituted by one, two or three independent occurrences of fluorine;

R^3 is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^4 is selected from the group consisting of:

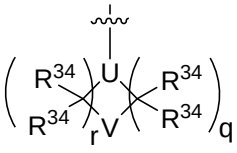


;

R^9 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, $C(=O)R^5$, SO_2R^5, C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen and heteroaryl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^5 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, and heterocyclyl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

each occurrence of R^7 is independently selected from the group consisting of H, C_1-C_6 alkyl, and C_3-C_6cycloalkyl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^7 are joined together with the atom to which they are attached to form oxo;

R^L is selected from the group consisting of C_1-C_6alkyl and C_3-C_6cycloalkyl, $NR^{11}R^{12}$, and  , wherein each C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by 1, 2, or 3 independent occurrences of fluorine;

U is N or CR^{13};

V is selected from the group consisting of oxygen, $C(R^{34})_2$, and NR^6;

r is 0, 1, or 2;

q is 1, 2, or 3;

R^6 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, $C(=O)R^5$, SO_2R^5, C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen and heteroaryl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{11} is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{12} is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{13} is selected from H and C_1-C_3alkyl;

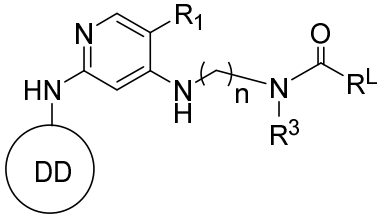
each occurrence of R^{34} is independently selected from H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^{34} are joined together with the carbon to which they are attached to form a C_3-C_6cycloalkyl; and

L is $-(C(R^{10})_2)_m-$;

m is 0, 1, 2, or 3;

provided that when r is 0 and q is 1, then U is not CR^{13} and V is not O, and when r or q is 1, then U is not N and V is not O or NR^6.

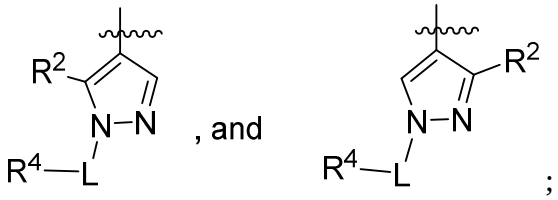
13. The compound of claim 1 represented by:



Formula IF

or a pharmaceutically acceptable salt thereof, wherein:

ring DD is selected from the group consisting of:



;

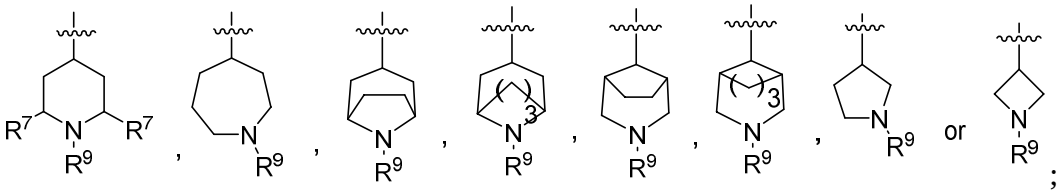
n is 2, 3, or 4;

R^1 is selected from the group consisting of halogen, cyano, C_1-C_5alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_5alkyl and C_3-C_5cycloalkyl may be optionally substituted by one, two or three independent occurrences of fluorine;

R^2 is selected from the group consisting of C_1-C_2alkyl and C_3-C_4cycloalkyl, and halogen, wherein C_1-C_2alkyl and C_3-C_4cycloalkyl may be optionally substituted by one, two or three independent occurrences of fluorine;

R^3 is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^4 is selected from the group consisting of:

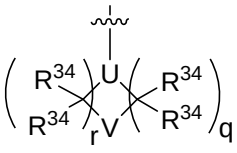


;

R^9 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, $C(=O)R^5$, SO_2R^5, C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen and heteroaryl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^5 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, and heterocyclyl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

each occurrence of R^7 is independently selected from the group consisting of H, C_1-C_6 alkyl, and C_3-C_6cycloalkyl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^7 are joined together with the atom to which they are attached to form oxo;

R^L is selected from the group consisting of C_1-C_6alkyl and C_3-C_6cycloalkyl, $NR^{11}R^{12}$, and  , wherein each C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by 1, 2, or 3 independent occurrences of fluorine;

U is N or CR^{13};

V is selected from the group consisting of oxygen, $C(R^{34})_2$, and NR^6;

r is 0, 1, or 2;

q is 1, 2, or 3;

R^6 is selected from the group consisting of H, C_1-C_6alkyl, C_3-C_6cycloalkyl, $C(=O)R^5$, SO_2R^5, C-linked heterocyclyl having at least one nitrogen and optionally having an additional ring nitrogen or oxygen and heteroaryl, wherein C_1-C_6alkyl and C_3-C_6cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{11} is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{12} is selected from the group consisting of H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine;

R^{13} is selected from H and C_1-C_3alkyl;

each occurrence of R^{34} is independently selected from H, C_1-C_3alkyl, and C_3-C_5cycloalkyl, wherein C_1-C_3alkyl and C_3-C_5cycloalkyl may be optionally substituted by one or more independent occurrences of fluorine, or two R^{34} are joined together with the carbon to which they are attached to form a C_3-C_6cycloalkyl; and

L is $-(C(R^{10})_2)_m-$;

m is 0, 1, 2, or 3;

provided when r is 0 and q is 1, then U is not CR^{13} and V is not O, and when r or q is 1, then U is not N and V is not O or NR^6.

14. The compound of claim 1 selected from the group consisting of: N-(3-((5-cyclopropyl-2-((3-methyl-1-(piperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-cyclopropyl-2-((2-methoxy-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-cyclopropyl-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-methyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)pivalamide, N-(3-((5-bromo-2-((2-isopropyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-cyclopropyl-2-((2-isopropyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-cyclopropyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((1-(1-amino-2-methyl-1-oxopropan-2-yl)-3-methyl-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-methoxy-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((1-(2-cyanopropan-2-yl)-3-methyl-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-methyl-6-(4-methylpiperazin-1-yl)pyridin-3-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-chloro-2-((2-isopropoxy-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-chloro-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((5-chloro-2-((2-methyl-4-(piperidin-4-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-methyl-N-(3-((2-((2-methyl-4-(piperidin-4-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-chloro-2-((2-methyl-4-(1-methylpiperidin-4-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-methyl-N-(3-((2-((2-methyl-4-(1-methylpiperidin-4-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((1-(1-cyclopropylpiperidin-4-yl)-3-methyl-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((2-((3-methyl-1-(piperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-

yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((1-(1-isopropylpiperidin-4-yl)-3-methyl-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((1-(1-isobutylpiperidin-4-yl)-3-methyl-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-bromo-2-((3-methyl-1-(1-d3-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((3-methyl-1-(1-d3-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-methyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-methyl-6-(4-methylpiperazin-1-yl)pyridin-3-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-methyl-6-morpholinopyridin-3-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-isopropyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-cyclopropyl-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-bromo-2-((2-cyclopropyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-((1R,5S)-8-methyl-3,8-diazabicyclo[3.2.1]octan-3-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-(5-methyl-2,5-diazabicyclo[2.2.1]heptan-2-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-(1,4-diazabicyclo[3.2.1]octan-4-yl)-2-ethylphenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-(hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-(difluoromethyl)-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-3,3-difluorocyclobutane-1-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)isobutyramide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)pivalamide, N-(3-((2-((2-

ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclopropanecarboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)tetrahydrofuran-3-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)tetrahydro-2H-pyran-4-carboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)tetrahydrofuran-3-carboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)tetrahydro-2H-pyran-4-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-((1R,5S)-8-methyl-3,8-diazabicyclo[3.2.1]octan-3-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-(5-methyl-2,5-diazabicyclo[2.2.1]heptan-2-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((2-((4-(1,4-diazabicyclo[3.2.1]octan-4-yl)-2-ethylphenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-(hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((5-(difluoromethyl)-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-3,3-difluoro-N-methylcyclobutane-1-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methylisobutyramide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methylpivalamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methyloxetane-3-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methyltetrahydrofuran-3-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-

yl)amino)propyl)-N-methyltetrahydro-2H-pyran-4-carboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-N-methyloxetane-3-carboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-N-methyltetrahydrofuran-3-carboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-N-methyltetrahydro-2H-pyran-4-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((2-((2-ethyl-4-((1R,5S)-8-methyl-3,8-diazabicyclo[3.2.1]octan-3-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((2-((2-ethyl-4-(5-methyl-2,5-diazabicyclo[2.2.1]heptan-2-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((2-((4-(1,4-diazabicyclo[3.2.1]octan-4-yl)-2-ethylphenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((2-((2-ethyl-4-(hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N,1-dimethylazetidine-3-carboxamide, N-(3-((5-chloro-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((5-chloro-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-N,1-dimethylazetidine-3-carboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)-N,1-dimethylazetidine-3-carboxamide, N-(3-((2-((2-bromo-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-bromo-4-((1R,5S)-8-methyl-3,8-diazabicyclo[3.2.1]octan-3-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-bromo-4-(5-methyl-2,5-diazabicyclo[2.2.1]heptan-2-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-(1,4-diazabicyclo[3.2.1]octan-4-yl)-2-bromophenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-bromo-4-(hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-bromo-4-(4-methylpiperazin-1-

yl)phenyl)amino)-5-(difluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-chloro-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-chloro-4-((1R,5S)-8-methyl-3,8-diazabicyclo[3.2.1]octan-3-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-chloro-4-(5-methyl-2,5-diazabicyclo[2.2.1]heptan-2-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-(1,4-diazabicyclo[3.2.1]octan-4-yl)-2-chlorophenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-chloro-4-(hexahydropyrrolo[1,2-a]pyrazin-2(1H)-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-chloro-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(difluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-cyano-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-cyano-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(difluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-(4-methylpiperazin-1-yl)-2-(trifluoromethyl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-(difluoromethyl)-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-cyclopropyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(difluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((5-chloro-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-(difluoromethyl)-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-1-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methylazetidine-1-carboxamide, 3-(3-((2-((2-ethyl-4-((1R,5S)-8-methyl-3,8-diazabicyclo[3.2.1]octan-3-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-1,1-dimethylurea, 1-(3-((2-((2-ethyl-4-((1R,5S)-8-methyl-3,8-diazabicyclo[3.2.1]octan-3-yl)phenyl)amino)-5-

(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-1,3,3-trimethylurea, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)azetidine-1-carboxamide, N-(3-((5-bromo-2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)pyrimidin-4-yl)amino)propyl)pyrrolidine-1-carboxamide, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-3,3-dimethylazetidine-1-carboxamide, 3-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-1,1-dimethylurea, N-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-2,2-dimethylazetidine-1-carboxamide, 3-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)-1,1-dimethylurea, 1-(3-((2-((2-ethyl-4-(4-methylpiperazin-1-yl)phenyl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)-1,3,3-trimethylurea, N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((1-(2-(dimethylamino)ethyl)-3-methyl-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((3-methyl-1-(1-methylpyrrolidin-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-bromo-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-chloro-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)tetrahydrofuran-3-carboxamide, N-methyl-N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-bromo-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((5-chloro-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)tetrahydro-2H-pyran-4-carboxamide, N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((1-(2-(dimethylamino)ethyl)-3-methyl-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((3-methyl-1-(1-methylpyrrolidin-3-yl)-

1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-methyl-N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((1-(2-(dimethylamino)ethyl)-3-methyl-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N-methyloxetane-3-carboxamide, N-methyl-N-(3-((2-((3-methyl-1-(1-methylpyrrolidin-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((5-bromo-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, 1-methyl-N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, N-(3-((2-((1-(2-(dimethylamino)ethyl)-3-methyl-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, 1-methyl-N-(3-((2-((3-methyl-1-(1-methylpyrrolidin-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, N,1-dimethyl-N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, N-(3-((2-((1-(2-(dimethylamino)ethyl)-3-methyl-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)-N,1-dimethylazetidine-3-carboxamide, N,1-dimethyl-N-(3-((2-((3-methyl-1-(1-methylpyrrolidin-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, N-(3-((5-bromo-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((5-(difluoromethyl)-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-(difluoromethyl)-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-N-methylcyclobutanecarboxamide, N-(3-((5-(difluoromethyl)-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((5-(difluoromethyl)-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-methyl-N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)-5-

(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-methyl-N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, 1-methyl-N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, N,1-dimethyl-N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, N-(3-((5-(difluoromethyl)-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-(difluoromethyl)-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((5-(difluoromethyl)-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((5-bromo-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-bromo-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((5-bromo-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((5-chloro-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((5-chloro-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((5-chloro-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-1-methylazetidine-3-carboxamide, N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)isobutyramide, N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)pivalamide, N-methyl-N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)pivalamide, N-(3-((5-bromo-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)pivalamide, N-(3-((5-bromo-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)pivalamide, 3,3-difluoro-N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutane-1-carboxamide, 3,3-difluoro-N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-

1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutane-1-carboxamide, 3,3-difluoro-N-methyl-N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutane-1-carboxamide, 3,3-difluoro-N-methyl-N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutane-1-carboxamide, N-(3-((5-(difluoromethyl)-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-3,3-difluorocyclobutane-1-carboxamide, N-(3-((5-bromo-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-3,3-difluorocyclobutane-1-carboxamide, N-(3-((5-bromo-2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-3,3-difluorocyclobutane-1-carboxamide, N-(3-((5-chloro-2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)pyrimidin-4-yl)amino)propyl)-3,3-difluorocyclobutane-1-carboxamide, N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-methyl-N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)oxetane-3-carboxamide, 1-methyl-N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)azetidine-3-carboxamide, N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)oxetane-3-carboxamide, 1-methyl-N-(3-((2-((3-methyl-1-(8-methyl-8-azabicyclo[3.2.1]octan-3-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)azetidine-3-carboxamide, N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-1-carboxamide, 1,1-dimethyl-3-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)urea, N-methyl-N-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-1-carboxamide, 1,1,3-trimethyl-3-(3-((2-((3-methyl-1-(1-methylpiperidin-4-yl)-1H-pyrazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)urea, and pharmaceutically acceptable salts, enantiomers, stereoisomers, and tautomers thereof.

15. A compound selected from the group consisting of: N-(3-((2-((5-methyl-1-(1-methylpiperidin-4-yl)-1H-1,2,3-triazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)-2H-1,2,3-triazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)thiazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)oxazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-methyl-2-(1-methylpiperidin-4-yl)thiazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-methyl-2-(1-methylpiperidin-4-yl)oxazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((5-methyl-2-(4-methylpiperazin-1-yl)thiazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((5-methyl-2-(4-methylpiperazin-1-yl)oxazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-methyl-2-(4-methylpiperazin-1-yl)thiazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-methyl-2-(4-methylpiperazin-1-yl)oxazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((5-methyl-1-(1-methylpiperidin-4-yl)-1H-1,2,3-triazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)-2H-1,2,3-triazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)thiazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)oxazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((4-methyl-2-(1-methylpiperidin-4-yl)thiazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((4-methyl-2-(1-methylpiperidin-4-yl)oxazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((5-methyl-2-(4-methylpiperazin-1-yl)thiazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((5-methyl-2-(4-methylpiperazin-1-yl)oxazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((4-methyl-

2-(4-methylpiperazin-1-yl)thiazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, N-(3-((2-((4-methyl-2-(4-methylpiperazin-1-yl)oxazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)oxetane-3-carboxamide, 3,3-difluoro-N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)-2H-1,2,3-triazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)cyclobutane-1-carboxamide, 1-methyl-N-(3-((2-((5-methyl-1-(1-methylpiperidin-4-yl)-1H-1,2,3-triazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, 1-methyl-N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)-2H-1,2,3-triazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, 1-methyl-N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)thiazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, 1-methyl-N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)oxazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, 1-methyl-N-(3-((2-((4-methyl-2-(1-methylpiperidin-4-yl)thiazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, 1-methyl-N-(3-((2-((4-methyl-2-(1-methylpiperidin-4-yl)oxazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, 1-methyl-N-(3-((2-((5-methyl-2-(4-methylpiperazin-1-yl)thiazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, 1-methyl-N-(3-((2-((5-methyl-2-(4-methylpiperazin-1-yl)oxazol-4-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, 1-methyl-N-(3-((2-((4-methyl-2-(4-methylpiperazin-1-yl)thiazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, 1-methyl-N-(3-((2-((4-methyl-2-(4-methylpiperazin-1-yl)oxazol-5-yl)amino)-5-(trifluoromethyl)pyrimidin-4-yl)amino)propyl)azetidine-3-carboxamide, N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)-2H-1,2,3-triazol-4-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)thiazol-4-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((5-methyl-2-(1-methylpiperidin-4-yl)oxazol-4-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-methyl-2-(1-methylpiperidin-4-yl)thiazol-5-yl)amino)-5-(trifluoromethyl)pyridin-4-yl)amino)propyl)cyclobutanecarboxamide, N-(3-((2-((4-methyl-2-(1-methylpiperidin-4-yl)oxazol-5-yl)amino)-5-(trifluoromethyl)pyridin-4-

yl)amino)propyl)cyclobutanecarboxamide, and pharmaceutically acceptable salts, enantiomers, stereoisomers, and tautomers thereof.

16. A pharmaceutical composition comprising a compound of any one of claims 1-15, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable excipient.

17. A pharmaceutical composition comprising a compound of any one of claims 1-15, or a pharmaceutically acceptable salt thereof, one or more additional therapeutic agents, and a pharmaceutically acceptable excipient.

18. The pharmaceutical composition of claim 17, wherein the composition comprises one additional therapeutic agent.

19. The pharmaceutical composition of claim 17 or claim 18, wherein the additional therapeutic agent is a MAPKAP pathway inhibitor.

20. The pharmaceutical composition of claim 19, wherein the MAPKAP pathway inhibitor is selected from the group consisting of a MEK inhibitor, an ERK inhibitor, a RAF inhibitor, and a Ras inhibitor.

21. The pharmaceutical composition of claim 20, wherein the MEK inhibitor is selected from the group consisting of trametinib, selumetinib, cobimetinib, binimetinib, and pharmaceutically acceptable salts thereof; the ERK inhibitor is selected from the group consisting of ulixertinib, SCH772984, LY3214996, ravoxertinib, VX-11e, and pharmaceutically acceptable salts thereof; the RAF inhibitor is selected from the group consisting of LY3009120, LXH254, RAF709, dabrafenib, vemurafenib, and pharmaceutically acceptable salts thereof; or
the Ras inhibitor is selected from the group consisting of AMG-510, MRTX849, and pharmaceutically acceptable salts thereof.

22. The pharmaceutical composition of claim 17 or claim 18, wherein the additional therapeutic agent is a chemotherapeutic agent.

23. The pharmaceutical composition of claim 22, wherein the chemotherapeutic agent is a selected from the group consisting of anti-tubulin agents, vinorelbine, DNA-alkylating agents, DNA intercalating agents, 5-fluorouracil, capecitabine, cytarabine, decitabine, 5-aza cytadine, gemcitabine, and methotrexate.

24. Use of a compound of any one of claims 1-15, or a pharmaceutically acceptable salt thereof, in the manufacture of a medicament for treating a tumor in a patient in need thereof.

25. Use of a compound of any one of claims 1-15, or a pharmaceutically acceptable salt thereof, in the manufacture of a medicament for treating a cancer in a patient in need thereof.

26. The use of claim 25, wherein the cancer is selected from the group consisting of gastrointestinal stromal tumors, esophageal cancer, gastric cancer, melanomas, gliomas, glioblastomas, ovarian cancer, bladder cancer, pancreatic cancer, prostate cancer, lung cancers, breast cancers, renal cancers, hepatic cancers, osteosarcomas, multiple myelomas, cervical carcinomas, cancers that are metastatic to bone, papillary thyroid carcinoma, non-small cell lung cancer, and colorectal cancers.

27. The use of claim 25 or 26, wherein one or more additional therapeutic agents is to be administered to the patient.

28. The use of claim 27, wherein the additional therapeutic agent is a MAPKAP pathway inhibitor.

29. The use of claim 28, wherein the MAPKAP pathway inhibitor is selected from the group consisting of a MEK inhibitor, an ERK inhibitor, a RAF inhibitor, and a Ras inhibitor.

30. The use of claim 29, wherein the MEK inhibitor is selected from the group consisting of trametinib, selumetinib, cobimetinib, binimetinib, and pharmaceutically acceptable salts thereof; the ERK inhibitor is selected from the group consisting of ulixertinib, SCH772984, LY3214996, ravoxertinib, VX-11e, and pharmaceutically acceptable salts thereof; he RAF inhibitor is selected from the group consisting of LY3009120, LXH254, RAF709, dabrafenib, vemurafenib, and pharmaceutically acceptable salts thereof; or the Ras inhibitor is selected

from the group consisting of AMG-510, MRTX849, and pharmaceutically acceptable salts thereof.

31. The use of claim 27, wherein the additional therapeutic agent is a chemotherapeutic agent.

32. The use of claim 31, wherein the chemotherapeutic agent is selected from the group consisting of anti-tubulin agents, vinorelbine, DNA-alkylating agents, DNA intercalating agents, 5-fluorouracil, capecitabine, cytarabine, decitabine, 5-aza cytadine, gemcitabine, and methotrexate.

33. A compound of any one of claims 1-15, or a pharmaceutically acceptable salt thereof, for use in therapy.