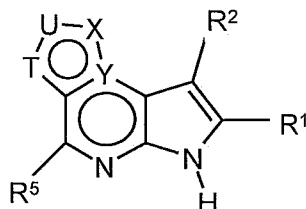


SUMMARY OF THE INVENTION

In one aspect, the present invention provides a compound of formula (I)



Formula (I)

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pharmaceutically acceptable salts thereof wherein

T is N, U is N, X is CR³ and Y is N; or

T is CR⁶, U is N, X is CR³ and Y is N; or

T is N, U is CR⁴, X is CR³ and Y is N; or

10 T is CR⁶, U is CR⁴, X is CR³ and Y is N;

R¹ and R² are each independently hydrogen, deuterium, -N(R^a)(R^b), halogen, -OR^a, -SR^a, -S(O)R^a, -S(O)₂R^a, -NO₂, -C(O)OR^a, -CN, -C(O)N(R^a)(R^b), -N(R^a)C(O)(R^b), -C(O)R^a, -C(OH)R^aR^b, -N(R^a)S(O)₂-R^b, -S(O)₂N(R^a)(R^b), -CF₃, -OCF₃, optionally substituted (C₁-C₆)alkyl, optionally substituted (C₂-C₆)alkenyl, optionally substituted (C₂-C₆)alkynyl, optionally substituted (C₃-C₁₀)cycloalkyl, optionally substituted (C₁-C₁₀)heteroaryl, optionally substituted (C₁-C₁₀) heterocyclyl, or optionally substituted (C₆-C₁₀)aryl;

R⁵ is hydrogen, deuterium, -N(R^a)(R^b), halogen, -SR^a, -S(O)R^a, -S(O)₂R^a, -NO₂, -C(O)OR^a, -CN, -C(O)N(R^a)(R^b), -N(R^a)C(O)(R^b), -C(O)R^a, -C(OH)R^aR^b, -N(R^a)S(O)₂-R^b, -S(O)₂N(R^a)(R^b), -CF₃, -OCF₃, optionally substituted (C₁-C₆)alkyl, optionally substituted (C₂-C₆)alkenyl, optionally substituted (C₂-C₆)alkynyl, optionally substituted (C₃-C₁₀)cycloalkyl, optionally substituted (C₁-C₁₀)heteroaryl, optionally substituted (C₁-C₁₀) heterocyclyl, or optionally substituted (C₆-C₁₀)aryl;

20

wherein in a moiety containing $-N(R^a)(R^b)$, the nitrogen, R^a and R^b may form a ring such that $-N(R^a)(R^b)$ represents an optionally substituted (C_2-C_{10}) heterocyclyl or optionally substituted (C_1-C_{10}) heteroaryl linked through a nitrogen;

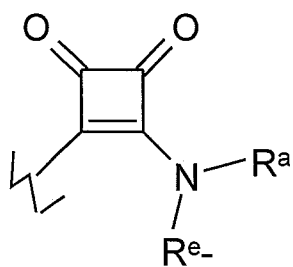
R^3 is optionally substituted cyclopropyl, optionally substituted cyclobutyl, optionally substituted cyclopentyl, optionally substituted cyclohexyl, optionally substituted phenyl, optionally substituted adamantanyl, optionally substituted azetidiny, optionally substituted bicyclo[2.1.1]hexyl, optionally substituted bicyclo[2.2.1]heptyl, optionally substituted bicyclo[2.2.2]octyl, optionally substituted bicyclo[3.2.1]octyl, optionally substituted bicyclo[3.1.1]heptyl, optionally substituted azabicyclo[3.2.1]octanyl, optionally substituted azabicyclo[2.2.1]heptanyl, optionally substituted 2-azabicyclo[3.2.1]octanyl, optionally substituted azabicyclo[3.2.2]nonanyl, optionally substituted bicyclo[2.2.1]hept-2-enyl, optionally substituted piperidinyl, optionally substituted pyrrolidinyl or optionally substituted tetrahydrofuranyl; or

R^3 is -A-D-E-G, wherein:

A is a bond or (C_1-C_6) alkylene;

D is an optionally substituted azetidiny, optionally substituted bicyclo[2.2.2]octanylene, optionally substituted bicyclo[2.2.1]heptylene, optionally substituted bicyclo[2.1.1]hexylene, optionally substituted cyclobutylene, optionally substituted cyclopentylene, optionally substituted cyclohexylene, optionally substituted bicyclo[2.2.1]hept-2-enylene, optionally substituted piperidinyl, or optionally substituted pyrrolidinyl;

E is a bond, $-R^e-$, $-R^e-C(O)-R^e-$, $-R^e-C(O)C(O)-R^e-$, $-R^e-C(O)O-R^e-$, $-R^e-C(O)C(O)N(R^a)-R^e-$, $-R^e-N(R^a)-C(O)C(O)-R^e-$, $-R^e-O-R^e-$, $-R^e-S(O)_2-R^e-$, $-R^e-S(O)-R^e-$, $-R^e-S-R^e-$, $-R^e-N(R^a)-R^e-$, $-R^e-N(R^a)C(O)-R^e-$, $-R^eC(O)N(R^a)R^e-$, $-R^e-OC(O)N(R^a)-R^e-$, $-R^e-N(R^a)C(O)OR^e-$, $-R^e-OC(O)-R^e-$, $-R^e-N(R^a)C(O)N(R^b)-R^e-$, $-R^e-N(R^a)S(O)_2-R^e-$, or $-R^e-S(O)_2N(R^a)-R^e-$; or



E is ;

where in all cases, E is linked to either a carbon or a nitrogen atom in D;

G is hydrogen, deuterium, $-N(R^a)(R^b)$, halogen, $-OR^a$, $-SR^a$, $-S(O)R^a$, $-S(O)_2R^a$, $-NO_2$, $-C(O)OR^a$, $-CN$, $-C(O)N(R^a)(R^b)$, $-N(R^a)C(O)R^b$, $-N(R^a)C(O)OR^b$, $-OC(O)N(R^a)$, $-N(R^a)C(O)N(R^b)_2$, $-C(O-R^a)(R^b)_2$, $-C(O)R^a$, $-CF_3$, $-OCF_3$, $-N(R^a)S(O)_2R^b$, $-S(O)_2N(R^a)(R^b)$, $-S(O)_2N(R^a)C(O)R^b$, an optionally substituted $-(C_1-C_6)$ alkyl, an optionally substituted $-(C_2-C_6)$ alkenyl, an optionally substituted $-(C_2-C_6)$ alkynyl, an optionally substituted $-(C_3-C_{10})$ cycloalkyl, an optionally substituted $-(C_1-C_{10})$ heteroaryl, an optionally substituted $-(C_1-C_{10})$ heterocyclyl, an optionally substituted $-(C_6-C_{10})$ aryl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_3-C_{10}) cycloalkyl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_6-C_{10}) aryl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_1-C_{10}) heteroaryl, or an optionally substituted $-(C_1-C_6)$ alkyl- (C_1-C_{10}) heterocyclyl;

wherein in a moiety containing $-N(R^a)(R^b)$, the nitrogen, R^a and R^b may form a ring such that $-N(R^a)(R^b)$ represents an optionally substituted (C_2-C_{10}) heterocyclyl or an optionally substituted (C_1-C_{10}) heteroaryl linked through a nitrogen;

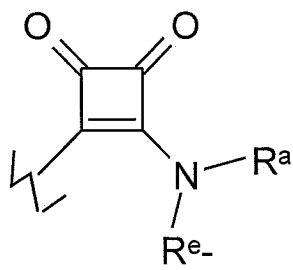
R^4 and R^6 are each independently a hydrogen, halogen, deuterium, an optionally substituted bridged (C_5-C_{12}) cycloalkyl group, optionally substituted bridged (C_2-C_{10}) heterocyclyl group, optionally substituted (C_1-C_8) alkyl, optionally substituted (C_3-C_{10}) cycloalkyl, optionally substituted (C_3-C_8) cycloalkenyl, optionally substituted (C_6-C_{10}) aryl, optionally substituted (C_1-C_{10}) heteroaryl, optionally substituted (C_2-C_{10}) heterocyclyl or $-J-L-M-Q$;

wherein:

J is a bond, $-C(O)-$, optionally substituted (C_1-C_6) alkylene, optionally substituted (C_2-C_6) alkenylene, optionally substituted (C_2-C_6) alkynylene, optionally substituted (C_3-C_{12}) cycloalkylene, optionally substituted (C_2-C_6) heterocyclylene, $-C(O)N(R^a)-R^e-$, $-N(R^a)C(O)-R^e-$, $-O-R^e-$, $-N(R^a)-R^e-$, $-S-R^e-$, $-S(O)_2-R^e-$, $-S(O)R^e-$, $-C(O-R^a)(R^b)-R^e-$, $-S(O)_2N(R^a)-R^e-$, $-N(R^a)S(O)_2-R^e-$ or $-N(R^a)C(O)N(R^b)-R^e-$;

L is a bond, an optionally substituted (C_1-C_8) alkylene, optionally substituted bridged (C_5-C_{12}) cycloalkylene, optionally substituted (C_3-C_{10}) cycloalkylene, optionally substituted bridged (C_5-C_{10}) cycloalkenylene, optionally substituted (C_3-C_{10}) cycloalkenylene, optionally substituted (C_6-C_{10}) arylene, optionally substituted (C_1-C_{10}) heteroarylene, optionally substituted bridged (C_2-C_{10}) heterocyclylene or an optionally substituted (C_2-C_{10}) heterocyclylene;

M is a bond, $-R^e$ -, $-R^e-C(O)-R^e$ -, $-R^e-C(O)C(O)-R^e$ -, $-R^e-C(O)O-R^e$ -, $-R^e-OC(O)-R^e$ -, $-R^e-C(O)C(O)N(R^a)-R^e$ -, $-R^e-N(R^a)-C(O)C(O)-R^e$ -, $-R^e-O-R^e$ -, $-R^e-S(O)_2-R^e$ -, $-R^e-S(O)-R^e$ -, $-R^e-S-R^e$ -, $-R^e-N(R^a)-R^e$ -, $-R^e-N(R^a)C(O)-R^e$ -, $-R^e-C(O)N(R^a)R^e$ -, $-R^e-OC(O)N(R^a)-R^e$ -, $-R^e-N(R^a)C(O)OR^e$ -, $-R^e-N(R^a)C(O)N(R^b)-R^e$ -, $-R^e-N(R^a)S(O)_2-R^e$ -, or $-R^e-S(O)_2N(R^a)-R^e$ -; or



5 M is ;

where in all cases, M is linked to either a carbon or a nitrogen atom in L;

Q is hydrogen, deuterium, $-N(R^a)(R^b)$, halogen, $-OR^a$, $-SR^a$, $-S(O)R^a$, $-S(O)_2R^a$, $-NO_2$, $-C(O)OR^a$, $-CN$, $-C(O)N(R^a)(R^b)$, $-N(R^a)C(O)R^b$, $-N(R^a)C(O)OR^b$, $-N(R^a)C(O)N(R^b)_2$, $-C(O-R^a)(R^b)_2$, $-C(O)R^a$, $-CF_3$, $-OCF_3$, $-N(R^a)S(O)_2R^b$, $-S(O)_2N(R^a)(R^b)$, $-S(O)_2N(R^a)C(O)R^b$, an
 10 optionally substituted (C_1-C_6) alkyl, an optionally substituted (C_2-C_6) alkenyl, an optionally substituted (C_2-C_6) alkynyl, an optionally substituted (C_3-C_{10}) cycloalkyl, an optionally substituted (C_1-C_{10}) heteroaryl, an optionally substituted (C_1-C_{10}) heterocyclyl, an optionally substituted (C_6-C_{10}) aryl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_3-C_{10}) cycloalkyl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_6-C_{10}) aryl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_1-C_{10}) heteroaryl, or an
 15 optionally substituted $-(C_1-C_6)$ alkyl- (C_1-C_{10}) heterocyclyl;

wherein in a moiety containing $-N(R^a)(R^b)$, the nitrogen, R^a and R^b may form a ring such that $-N(R^a)(R^b)$ represents an optionally substituted (C_2-C_{10}) heterocyclyl or an optionally substituted (C_1-C_{10}) heteroaryl linked through a nitrogen;

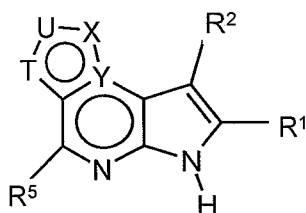
R^a and R^b are each independently hydrogen, deuterium, an optionally substituted (C_1-C_{10}) alkyl, an optionally substituted (C_2-C_{10}) alkenyl, an optionally substituted (C_2-C_{10}) alkynyl, an optionally substituted (C_1-C_{10}) alkyl-O- (C_1-C_{10}) alkyl, an optionally substituted (C_3-C_{10}) cycloalkyl, an optionally substituted (C_6-C_{10}) aryl, an optionally substituted (C_1-C_{10}) heteroaryl, an optionally substituted (C_1-C_{10}) heterocyclyl, an optionally substituted $-(C_1-C_6)$ alkylene- (C_3-C_{10}) cycloalkyl, an optionally substituted $-(C_1-C_6)$ alkylene- (C_6-C_{10}) aryl, an
 20

optionally substituted $-(C_1-C_6)\text{alkylene}-(C_1-C_{10})\text{heteroaryl}$, or an optionally substituted $-(C_1-C_6)\text{alkylene}-(C_1-C_{10})\text{heterocyclyl}$; and

R^e for each occurrence is independently a bond, an optionally substituted $(C_1-C_{10})\text{alkylene}$, an optionally substituted $(C_2-C_{10})\text{alkenylene}$, an optionally substituted $(C_2-C_{10})\text{alkynylene}$, an optionally substituted $-(C_1-C_{10})\text{alkylene-O}-(C_1-C_{10})\text{alkylene}$ group, an optionally substituted $(C_3-C_{10})\text{cycloalkylene}$, an optionally substituted $(C_6-C_{10})\text{arylene}$, an optionally substituted $(C_1-C_{10})\text{heteroarylene}$, or an optionally substituted $(C_1-C_{10})\text{heterocyclylene}$; and

wherein the optional substituents are independently selected from $(C_1-C_8)\text{alkyl}$ groups, $(C_2-C_8)\text{alkenyl}$ groups, $(C_2-C_8)\text{alkynyl}$ groups, $(C_3-C_{10})\text{cycloalkyl}$ groups, halogen, halogenated $(C_1-C_8)\text{alkyl}$ groups, $-\text{O}-(C_1-C_8)\text{alkyl}$ groups, $-\text{OH}$, $-\text{S}-(C_1-C_8)\text{alkyl}$ groups, $-\text{SH}$, $-\text{NH}(C_1-C_8)\text{alkyl}$ groups, $-\text{N}((C_1-C_8)\text{alkyl})_2$ groups, $-\text{NH}_2$, $-\text{C}(\text{O})\text{NH}_2$, $-\text{C}(\text{O})\text{NH}(C_1-C_8)\text{alkyl}$ groups, $-\text{C}(\text{O})\text{N}((C_1-C_8)\text{alkyl})_2$, $-\text{NHC}(\text{O})\text{H}$, $-\text{NHC}(\text{O})(C_1-C_8)\text{alkyl}$ groups, $-\text{NHC}(\text{O})(C_3-C_8)\text{cycloalkyl}$ groups, $-\text{N}((C_1-C_8)\text{alkyl})\text{C}(\text{O})\text{H}$, $-\text{N}((C_1-C_8)\text{alkyl})\text{C}(\text{O})(C_1-C_8)\text{alkyl}$ groups, $-\text{NHC}(\text{O})\text{NH}_2$, $-\text{NHC}(\text{O})\text{NH}(C_1-C_8)\text{alkyl}$ groups, $-\text{N}((C_1-C_8)\text{alkyl})\text{C}(\text{O})\text{NH}_2$ groups, $-\text{NHC}(\text{O})\text{N}((C_1-C_8)\text{alkyl})_2$ groups, $-\text{N}((C_1-C_8)\text{alkyl})\text{C}(\text{O})\text{N}((C_1-C_8)\text{alkyl})_2$ groups, $-\text{N}((C_1-C_8)\text{alkyl})\text{C}(\text{O})\text{NH}((C_1-C_8)\text{alkyl})$, $-\text{C}(\text{O})\text{H}$, $-\text{C}(\text{O})(C_1-C_8)\text{alkyl}$ groups, $-\text{CN}$, $-\text{NO}_2$, $-\text{S}(\text{O})(C_1-C_8)\text{alkyl}$ groups, $-\text{S}(\text{O})_2(C_1-C_8)\text{alkyl}$ groups, $-\text{S}(\text{O})_2\text{N}((C_1-C_8)\text{alkyl})_2$ groups, $-\text{S}(\text{O})_2\text{NH}(C_1-C_8)\text{alkyl}$ groups, $-\text{S}(\text{O})_2\text{NH}(C_3-C_8)\text{cycloalkyl}$ groups, $-\text{S}(\text{O})_2\text{NH}_2$ groups, $-\text{NHS}(\text{O})_2(C_1-C_8)\text{alkyl}$ groups, $-\text{N}((C_1-C_8)\text{alkyl})\text{S}(\text{O})_2(C_1-C_8)\text{alkyl}$ groups, $-(C_1-C_8)\text{alkyl-O}-(C_1-C_8)\text{alkyl}$ groups, $-\text{O}-(C_1-C_8)\text{alkyl-O}-(C_1-C_8)\text{alkyl}$ groups, $-\text{C}(\text{O})\text{OH}$, $-\text{C}(\text{O})\text{O}(C_1-C_8)\text{alkyl}$ groups, NHOH , $\text{NHO}(C_1-C_8)\text{alkyl}$ groups, $-\text{O-halogenated}(C_1-C_8)\text{alkyl}$ groups, $-\text{S}(\text{O})_2\text{-halogenated}(C_1-C_8)\text{alkyl}$ groups, $-\text{S-halogenated}(C_1-C_8)\text{alkyl}$ groups, $-(C_1-C_6)\text{heterocycle}$, $-(C_1-C_6)\text{heteroaryl}$, $-\text{phenyl}$, $-\text{NHC}(\text{O})\text{O}-(C_1-C_6)\text{alkyl}$ groups, $-\text{N}((C_1-C_6)\text{alkyl})\text{C}(\text{O})\text{O}-(C_1-C_6)\text{alkyl}$ groups, $-\text{C}(\text{=NH})-(C_1-C_6)\text{alkyl}$ groups, $-\text{C}(\text{=NOH})-(C_1-C_6)\text{alkyl}$ groups, or $-\text{C}(\text{=N-O}-(C_1-C_6)\text{alkyl})-(C_1-C_6)\text{alkyl}$ groups.

In another aspect, the present invention provides a pharmaceutical composition comprising a compound of Formula (I) as defined herein



Formula (I)

a pharmaceutically acceptable carrier and excipient and a second therapeutic agent selected from the group consisting of cytokine suppressive anti-inflammatory drugs, antibodies to or antagonists of other human cytokines or growth factors, IL-1, IL-2, IL-3, IL-4, IL-5, IL-6, IL-7, IL-8, IL-12, IL-15, IL-16, IL-21, IL-23, interferons, EMAP-II, GM-CSF, FGF, PDGF, CTLA or
 5 IL-8, IL-12, IL-15, IL-16, IL-21, IL-23, interferons, EMAP-II, GM-CSF, FGF, PDGF, CTLA or their ligands including CD154, adalimumab golimumab, certolizumab pegol (CDP870), tocilizumab, CDP 571, soluble p55 or p75 TNF receptors, Lenercept, TNF α converting enzyme inhibitors, IL-1 inhibitors, Interleukin 11, IL-18 antagonists, IL-12 antagonists, IL-12 antibodies, soluble IL-12 receptors, IL-12 binding proteins, non-depleting anti-CD4 inhibitors FK506,
 10 rapamycin, mycophenolate mofetil, leflunomide, NSAIDs, ibuprofen, corticosteroids, phosphodiesterase inhibitors, adenosine agonists, antithrombotic agents, complement inhibitors, adrenergic agents, IL-1 β converting enzyme inhibitors, T-cell signalling kinase inhibitors, metalloproteinase inhibitors, sulfasalazine, 6-mercaptapurines, derivatives p75TNFR1gG, sIL-1RI, sIL-1RII, sIL-6R, celecoxib, hydroxychloroquine sulfate, rofecoxib, infliximab, naproxen,
 15 valdecoxib, meloxicam, acetate, gold sodium thiomalate, aspirin, triamcinolone acetonide, propoxyphene napsylate/apap, folate, nabumetone, diclofenac, piroxicam, etodolac, diclofenac sodium, oxaprozin, oxycodone HCl, hydrocodone bitartrate/apap, diclofenac sodium/misoprostol, fentanyl, anakinra, tramadol HCl, salsalate, sulindac, cyanocobalamin/fa/pyridoxine, acetaminophen, alendronate sodium, morphine sulfate, lidocaine
 20 hydrochloride, indomethacin, glucosamine sulf/chondroitin, amitriptyline HCl, sulfadiazine, oxycodone HCl/acetaminophen, olopatadine HCl misoprostol, naproxen sodium, omeprazole, cyclophosphamide, rituximab, IL-1 TRAP, MRA, CTLA4-IG, IL-18 BP, anti-IL-12, anti-IL15, VX-740, Roflumilast, IC-485, CDC-801, S1P1 agonists, FTY720, PKC family inhibitors,

Ruboxistaurin, AEB-071, Mesopram, methotrexate, budenoside, dexamethasone, 5-aminosalicylic acid, olsalazine, IL-1ra, T cell signaling inhibitors, tyrosine kinase inhibitors, IL-11, mesalamine, prednisone, azathioprine, mercaptopurine, methylprednisolone sodium succinate, diphenoxylate/atrop sulfate, loperamide hydrochloride, ciprofloxacin/dextrose-water,

5 tetracycline hydrochloride, fluocinonide, metronidazole, thimerosal/boric acid, cholestyramine/sucrose, ciprofloxacin hydrochloride, hyoscyamine sulfate, meperidine hydrochloride, midazolam hydrochloride, promethazine hydrochloride, sodium phosphate, sulfamethoxazole/trimethoprim, polycarbophil, propoxyphene napsylate, hydrocortisone, multivitamins, balsalazide disodium, codeine phosphate/apap, colesevelam HCl,

10 cyanocobalamin, folic acid, levofloxacin, natalizumab, interferon-gamma, methylprednisolone, cyclosporine, 4-aminopyridine, tizanidine, interferon- β 1a, interferon- β 1b, interferon α -n3, interferon- α , interferon β 1A-IF, Peginterferon α 2b, hyperbaric oxygen, intravenous immunoglobulin, cladribine, FK506, prednisolone, antiinflammatory cytokines, interferon- β , caspase inhibitors, inhibitors of caspase-1, antibodies to CD40 ligand and CD80, alemtuzumab,

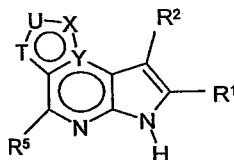
15 dronabinol, daclizumab, mitoxantrone, xaliproden hydrochloride, fampridine, glatiramer acetate, sinnabidol, α -immunokine NNSO3, ABR-215062, AnergiX.MS, chemokine receptor antagonists, BBR-2778, calagualine, CPI-1189, liposome encapsulated mitoxantrone, THC.CBD, cannabinoid agonists, MBP-8298, MNA-715, anti-IL-6 receptor antibody, neurovax, pirfenidone allotrap 1258 (RDP-1258), sTNF-R1, talampanel, teriflunomide, TGF-beta2, tiplimotide, VLA-4

20 antagonists, interferon gamma antagonists, IL-4 agonists, misoprostol, minocyclin, etanercept, betamethasone diprop augmented, dimethylsulfoxide, ketoprofen, tolmetin sodium, calcipotriene, glucosamine sulfate, risedronate sodium, thioguanine, alefacept, efalizumab, COX2 inhibitors, hydroxychloroquine, steroids, cytotoxics, inhibitors of PDE4, purine synthesis inhibitor, CTLA-4-IgG, anti-B7 family antibodies, anti-PD-1 family antibodies, anti-cytokine antibodies,

25 fonotolizumab, anti-IFN γ antibody, anti-receptor receptor antibodies, antibodies to B-cell surface molecules, LJP 394, anti-CD20 antibody and lymphostat.

LIST OF EMBODIMENTS

In a first embodiment the invention provides a compound of Formula (I)



Formula (I)

pharmaceutically acceptable salts, pro-drugs, biologically active metabolites, stereoisomers and isomers thereof wherein

T is N, U is N, X is CR³ and Y is N; or

10 T is CR⁶, U is N, X is CR³ and Y is N; or

T is N, U is CR⁴, X is CR³ and Y is N; or

T is CR⁶, U is CR⁴, X is CR³ and Y is N; or

T is CR⁶, U is N, X is NR³ and Y is C; or

T is O, U is N, X is CR³ and Y is C; or

15 T is NR⁶, U is N, X is CR³ and Y is C; or

T is CR⁶, U is CR⁴, X is NR³ and Y is C; or

T is S, U is N, X is CR³ and Y is C;

R¹, R² and R⁵ are each independently hydrogen, deuterium, -N(R^a)(R^b), halogen, -OR^a, -SR^a, -S(O)R^a, -S(O)₂R^a, -NO₂, -C(O)OR^a, -CN, -C(O)N(R^a)(R^b), -N(R^a)C(O)(R^b), -C(O)R^a, -

20 C(OH)R^aR^b, -N(R^a)S(O)₂R^b, -S(O)₂N(R^a)(R^b), -CF₃, -OCF₃, optionally substituted (C₁-C₆)alkyl, optionally substituted (C₂-C₆)alkenyl, optionally substituted (C₂-C₆)alkynyl, optionally substituted (C₃-C₁₀)cycloalkyl, optionally substituted (C₁-C₁₀)heteroaryl, optionally substituted (C₁-C₁₀)heterocyclyl, or optionally substituted (C₆-C₁₀)aryl;

wherein in a moiety containing -N(R^a)(R^b), the nitrogen, R^a and R^b may form a ring

25 such that -N(R^a)(R^b) represents an optionally substituted (C₂-C₁₀)heterocyclyl or

optionally substituted (C₁-C₁₀)heteroaryl linked through a nitrogen;

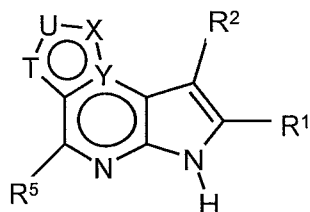
R³ is hydrogen, an optionally substituted bridged (C₅-C₁₂)cycloalkyl, optionally substituted bridged (C₂-C₁₀)heterocyclyl, optionally substituted (C₁-C₈)alkyl, optionally substituted (C₃-C₁₀)cycloalkyl, optionally substituted (C₃-C₈)cycloalkenyl, optionally substituted (C₆-C₁₀)aryl, optionally substituted (C₁-C₁₀)heteroaryl, optionally substituted (C₂-C₁₀)heterocyclyl; or

R³ is -A-D-E-G, wherein A is attached to X and:

A is a bond, -C(O)-, optionally substituted (C₁-C₆)alkylene, optionally substituted (C₂-C₆)alkenylene, optionally substituted (C₂-C₆)alkynylene, optionally substituted (C₃-

CLAIMS

1. A compound of formula (I)



Formula (I)

pharmaceutically acceptable salts thereof wherein

T is N, U is N, X is CR³ and Y is N; or

T is CR⁶, U is N, X is CR³ and Y is N; or

T is N, U is CR⁴, X is CR³ and Y is N; or

T is CR⁶, U is CR⁴, X is CR³ and Y is N;

R¹ and R² are each independently hydrogen, deuterium, -N(R^a)(R^b), halogen, -OR^a, -SR^a, -S(O)R^a, -S(O)₂R^a, -NO₂, -C(O)OR^a, -CN, -C(O)N(R^a)(R^b), -N(R^a)C(O)(R^b), -C(O)R^a, -C(OH)R^aR^b, -N(R^a)S(O)₂-R^b, -S(O)₂N(R^a)(R^b), -CF₃, -OCF₃, optionally substituted (C₁-C₆)alkyl, optionally substituted (C₂-C₆)alkenyl, optionally substituted (C₂-C₆)alkynyl, optionally substituted (C₃-C₁₀)cycloalkyl, optionally substituted (C₁-C₁₀)heteroaryl, optionally substituted (C₁-C₁₀) heterocyclyl, or optionally substituted (C₆-C₁₀)aryl;

R⁵ is hydrogen, deuterium, -N(R^a)(R^b), halogen, -SR^a, -S(O)R^a, -S(O)₂R^a, -NO₂, -C(O)OR^a, -CN, -C(O)N(R^a)(R^b), -N(R^a)C(O)(R^b), -C(O)R^a, -C(OH)R^aR^b, -N(R^a)S(O)₂-R^b, -S(O)₂N(R^a)(R^b), -CF₃, -OCF₃, optionally substituted (C₁-C₆)alkyl, optionally substituted (C₂-C₆)alkenyl, optionally substituted (C₂-C₆)alkynyl, optionally substituted (C₃-C₁₀)cycloalkyl, optionally substituted (C₁-C₁₀)heteroaryl, optionally substituted (C₁-C₁₀) heterocyclyl, or optionally substituted (C₆-C₁₀)aryl;

wherein in a moiety containing -N(R^a)(R^b), the nitrogen, R^a and R^b may form a ring such that -N(R^a)(R^b) represents an optionally substituted (C₂-C₁₀)heterocyclyl or optionally substituted (C₁-C₁₀)heteroaryl linked through a nitrogen;

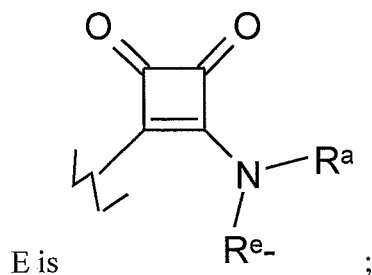
R^3 is optionally substituted cyclopropyl, optionally substituted cyclobutyl, optionally substituted cyclopentyl, optionally substituted cyclohexyl, optionally substituted phenyl, optionally substituted adamantanyl, optionally substituted azetidiny, optionally substituted bicyclo[2.1.1]hexyl, optionally substituted bicyclo[2.2.1]heptyl, optionally substituted bicyclo[2.2.2]octyl, optionally substituted bicyclo[3.2.1]octyl, optionally substituted bicyclo[3.1.1]heptyl, optionally substituted azabicyclo[3.2.1]octanyl, optionally substituted azabicyclo[2.2.1]heptanyl, optionally substituted 2-azabicyclo[3.2.1]octanyl, optionally substituted azabicyclo[3.2.2]nonanyl, optionally substituted bicyclo[2.2.1]hept-2-enyl, optionally substituted piperidiny, optionally substituted pyrrolidiny or optionally substituted tetrahydrofuranyl; or

R^3 is -A-D-E-G, wherein:

A is a bond or (C_1-C_6) alkylene;

D is an optionally substituted azetidiny, optionally substituted bicyclo[2.2.2]octanylene, optionally substituted bicyclo[2.2.1]heptylene, optionally substituted bicyclo[2.1.1]hexylene, optionally substituted cyclobutylene, optionally substituted cyclopentylene, optionally substituted cyclohexylene, optionally substituted bicyclo[2.2.1]hept-2-enylene, optionally substituted piperidiny, or optionally substituted pyrrolidiny;

E is a bond, $-R^e$ -, $-R^e-C(O)-R^e$ -, $-R^e-C(O)C(O)-R^e$ -, $-R^e-C(O)O-R^e$ -, $-R^e-C(O)C(O)N(R^a)-R^e$ -, $-R^e-N(R^a)-C(O)C(O)-R^e$ -, $-R^e-O-R^e$ -, $-R^e-S(O)_2-R^e$ -, $-R^e-S(O)-R^e$ -, $-R^e-S-R^e$ -, $-R^e-N(R^a)-R^e$ -, $-R^e-N(R^a)C(O)-R^e$ -, $-R^eC(O)N(R^a)R^e$ -, $-R^e-OC(O)N(R^a)-R^e$ -, $-R^e-N(R^a)C(O)OR^e$ -, $-R^e-OC(O)-R^e$ -, $-R^e-N(R^a)C(O)N(R^b)-R^e$ -, $-R^e-N(R^a)S(O)_2-R^e$ -, or $-R^e-S(O)_2N(R^a)-R^e$;- or



where in all cases, E is linked to either a carbon or a nitrogen atom in D;

G is hydrogen, deuterium, $-N(R^a)(R^b)$, halogen, $-OR^a$, $-SR^a$, $-S(O)R^a$, $-S(O)_2R^a$, $-NO_2$, $-C(O)OR^a$, $-CN$, $-C(O)N(R^a)(R^b)$, $-N(R^a)C(O)R^b$, $-N(R^a)C(O)OR^b$, $-OC(O)N(R^a)$, -

$N(R^a)C(O)N(R^b)_2$, $-C(O-R^a)(R^b)_2$, $-C(O)R^a$, $-CF_3$, $-OCF_3$, $-N(R^a)S(O)_2R^b$, $-S(O)_2N(R^a)(R^b)$, $-S(O)_2N(R^a)C(O)R^b$, an optionally substituted $-(C_1-C_6)$ alkyl, an optionally substituted $-(C_2-C_6)$ alkenyl, an optionally substituted $-(C_2-C_6)$ alkynyl, an optionally substituted $-(C_3-C_{10})$ cycloalkyl, an optionally substituted $-(C_1-C_{10})$ heteroaryl, an optionally substituted $-(C_1-C_{10})$ heterocyclyl, an optionally substituted $-(C_6-C_{10})$ aryl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_3-C_{10}) cycloalkyl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_6-C_{10}) aryl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_1-C_{10}) heteroaryl, or an optionally substituted $-(C_1-C_6)$ alkyl- (C_1-C_{10}) heterocyclyl;

wherein in a moiety containing $-N(R^a)(R^b)$, the nitrogen, R^a and R^b may form a ring such that $-N(R^a)(R^b)$ represents an optionally substituted (C_2-C_{10}) heterocyclyl or an optionally substituted (C_1-C_{10}) heteroaryl linked through a nitrogen;

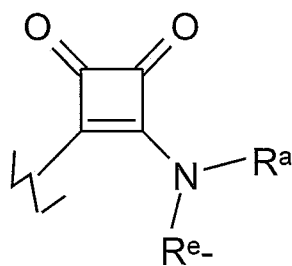
R^4 and R^6 are each independently a hydrogen, halogen, deuterium, an optionally substituted bridged (C_5-C_{12}) cycloalkyl group, optionally substituted bridged (C_2-C_{10}) heterocyclyl group, optionally substituted (C_1-C_8) alkyl, optionally substituted (C_3-C_{10}) cycloalkyl, optionally substituted (C_3-C_8) cycloalkenyl, optionally substituted (C_6-C_{10}) aryl, optionally substituted (C_1-C_{10}) heteroaryl, optionally substituted (C_2-C_{10}) heterocyclyl or $-J-L-M-Q$;

wherein:

J is a bond, $-C(O)-$, optionally substituted (C_1-C_6) alkylene, optionally substituted (C_2-C_6) alkenylene, optionally substituted (C_2-C_6) alkynylene, optionally substituted (C_3-C_{12}) cycloalkylene, optionally substituted (C_2-C_6) heterocyclylene, $-C(O)N(R^a)-R^e-$, $-N(R^a)C(O)-R^e-$, $-O-R^e-$, $-N(R^a)-R^e-$, $-S-R^e-$, $-S(O)_2-R^e-$, $-S(O)R^e-$, $-C(O-R^a)(R^b)-R^e-$, $-S(O)_2N(R^a)-R^e-$, $-N(R^a)S(O)_2-R^e-$ or $-N(R^a)C(O)N(R^b)-R^e-$;

L is a bond, an optionally substituted (C_1-C_8) alkylene, optionally substituted bridged (C_5-C_{12}) cycloalkylene, optionally substituted (C_3-C_{10}) cycloalkylene, optionally substituted bridged (C_5-C_{10}) cycloalkenylene, optionally substituted (C_3-C_{10}) cycloalkenylene, optionally substituted (C_6-C_{10}) arylene, optionally substituted (C_1-C_{10}) heteroarylene, optionally substituted bridged (C_2-C_{10}) heterocyclylene or an optionally substituted (C_2-C_{10}) heterocyclylene;

M is a bond, $-R^e-$, $-R^e-C(O)-R^e-$, $-R^e-C(O)C(O)-R^e-$, $-R^e-C(O)O-R^e-$, $-R^e-OC(O)-R^e-$, $-R^e-C(O)C(O)N(R^a)-R^e-$, $-R^e-N(R^a)-C(O)C(O)-R^e-$, $-R^e-O-R^e-$, $-R^e-S(O)_2-R^e-$, $-R^e-S(O)-R^e-$, $-R^e-S-R^e-$, $-R^e-N(R^a)-R^e-$, $-R^e-N(R^a)C(O)-R^e-$, $-R^e-C(O)N(R^a)-R^e-$, $-R^e-OC(O)N(R^a)-R^e-$, $-R^e-N(R^a)C(O)OR^e-$, $-R^e-N(R^a)C(O)N(R^b)-R^e-$, $-R^e-N(R^a)S(O)_2-R^e-$, or $-R^e-S(O)_2N(R^a)-R^e-$; or



M is ;

where in all cases, M is linked to either a carbon or a nitrogen atom in L;

Q is hydrogen, deuterium, $-N(R^a)(R^b)$, halogen, $-OR^a$, $-SR^a$, $-S(O)R^a$, $-S(O)_2R^a$, $-NO_2$, $-C(O)OR^a$, $-CN$, $-C(O)N(R^a)(R^b)$, $-N(R^a)C(O)R^b$, $-N(R^a)C(O)OR^b$, $-N(R^a)C(O)N(R^b)_2$, $-C(O)R^a(R^b)_2$, $-C(O)R^a$, $-CF_3$, $-OCF_3$, $-N(R^a)S(O)_2R^b$, $-S(O)_2N(R^a)(R^b)$, $-S(O)_2N(R^a)C(O)R^b$, an optionally substituted (C_1-C_6) alkyl, an optionally substituted (C_2-C_6) alkenyl, an optionally substituted (C_2-C_6) alkynyl, an optionally substituted (C_3-C_{10}) cycloalkyl, an optionally substituted (C_1-C_{10}) heteroaryl, an optionally substituted (C_1-C_{10}) heterocyclyl, an optionally substituted (C_6-C_{10}) aryl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_3-C_{10}) cycloalkyl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_6-C_{10}) aryl, an optionally substituted $-(C_1-C_6)$ alkyl- (C_1-C_{10}) heteroaryl, or an optionally substituted $-(C_1-C_6)$ alkyl- (C_1-C_{10}) heterocyclyl;

wherein in a moiety containing $-N(R^a)(R^b)$, the nitrogen, R^a and R^b may form a ring such that $-N(R^a)(R^b)$ represents an optionally substituted (C_2-C_{10}) heterocyclyl or an optionally substituted (C_1-C_{10}) heteroaryl linked through a nitrogen;

R^a and R^b are each independently hydrogen, deuterium, an optionally substituted (C_1-C_{10}) alkyl, an optionally substituted (C_2-C_{10}) alkenyl, an optionally substituted (C_2-C_{10}) alkynyl, an optionally substituted (C_1-C_{10}) alkyl-O- (C_1-C_{10}) alkyl, an optionally substituted (C_3-C_{10}) cycloalkyl, an optionally substituted (C_6-C_{10}) aryl, an optionally substituted (C_1-C_{10}) heteroaryl, an optionally substituted (C_1-C_{10}) heterocyclyl, an optionally substituted $-(C_1-C_6)$ alkylene- (C_3-C_{10}) cycloalkyl, an optionally substituted $-(C_1-C_6)$ alkylene- (C_6-C_{10}) aryl, an optionally substituted $-(C_1-C_6)$ alkylene- (C_1-C_{10}) heteroaryl, or an optionally substituted $-(C_1-C_6)$ alkylene- (C_1-C_{10}) heterocyclyl; and

R^e for each occurrence is independently a bond, an optionally substituted (C_1-C_{10}) alkylene, an optionally substituted (C_2-C_{10}) alkenylene, an optionally substituted (C_2-C_{10}) alkynylene, an optionally substituted $-(C_1-C_{10})$ alkylene-O- (C_1-C_{10}) alkylene group, an

optionally substituted (C₃-C₁₀)cycloalkylene, an optionally substituted (C₆-C₁₀)arylene, an optionally substituted (C₁-C₁₀)heteroarylene, or an optionally substituted (C₁-C₁₀)heterocyclylene; and

wherein the optional substituents are independently selected from (C₁-C₈)alkyl groups, (C₂-C₈)alkenyl groups, (C₂-C₈)alkynyl groups, (C₃-C₁₀)cycloalkyl groups, halogen, halogenated (C₁-C₈)alkyl groups, -O-(C₁-C₈)alkyl groups, -OH, -S-(C₁-C₈)alkyl groups, -SH, -NH(C₁-C₈)alkyl groups, -N((C₁-C₈)alkyl)₂ groups, -NH₂, -C(O)NH₂, -C(O)NH(C₁-C₈)alkyl groups, -C(O)N((C₁-C₈)alkyl)₂, -NHC(O)H, -NHC(O)(C₁-C₈)alkyl groups, -NHC(O)(C₃-C₈)cycloalkyl groups, -N((C₁-C₈)alkyl)C(O)H, -N((C₁-C₈)alkyl)C(O)(C₁-C₈)alkyl groups, -NHC(O)NH₂, -NHC(O)NH(C₁-C₈)alkyl groups, -N((C₁-C₈)alkyl)C(O)NH₂ groups, -NHC(O)N((C₁-C₈)alkyl)₂ groups, -N((C₁-C₈)alkyl)C(O)N((C₁-C₈)alkyl)₂ groups, -N((C₁-C₈)alkyl)C(O)NH((C₁-C₈)alkyl), -C(O)H, -C(O)(C₁-C₈)alkyl groups, -CN, -NO₂, -S(O)(C₁-C₈)alkyl groups, -S(O)₂(C₁-C₈)alkyl groups, -S(O)₂N((C₁-C₈)alkyl)₂ groups, -S(O)₂NH(C₁-C₈)alkyl groups, -S(O)₂NH(C₃-C₈)cycloalkyl groups, -S(O)₂NH₂ groups, -NHS(O)₂(C₁-C₈)alkyl groups, -N((C₁-C₈)alkyl)S(O)₂(C₁-C₈)alkyl groups, -(C₁-C₈)alkyl-O-(C₁-C₈)alkyl groups, -O-(C₁-C₈)alkyl-O-(C₁-C₈)alkyl groups, -C(O)OH, -C(O)O(C₁-C₈)alkyl groups, NHOH, NHO(C₁-C₈)alkyl groups, -O-halogenated (C₁-C₈)alkyl groups, -S(O)₂-halogenated (C₁-C₈)alkyl groups, -S-halogenated (C₁-C₈)alkyl groups, -(C₁-C₆) heterocycle, -(C₁-C₆) heteroaryl, -phenyl, -NHC(O)O-(C₁-C₆)alkyl groups, -N((C₁-C₆)alkyl)C(O)O-(C₁-C₆)alkyl groups, -C(=NH)-(C₁-C₆)alkyl groups, -C(=NOH)-(C₁-C₆)alkyl groups, or -C(=N-O-(C₁-C₆)alkyl)-(C₁-C₆)alkyl groups.

2. The compound of claim 1 wherein E is -R^e-C(O)-R^e-, R^e-O-R^e-, -R^e-S(O)₂-R^e-, -R^e-N(R^a)-R^e-, -R^e-N(R^a)C(O)-R^e-, -R^e-C(O)N(R^a)R^e-, -R^e-N(R^a)S(O)₂-R^e-, or -R^e-S(O)₂N(R^a)R^e-.

3. The compound of claim 2 wherein G is -OR^a, -CN, -N(R^a)S(O)₂R^b, -S(O)₂N(R^a)(R^b), optionally substituted (C₁-C₆)alkyl, optionally substituted cyclopropyl, optionally substituted cyclobutyl, optionally substituted cyclopentyl, optionally substituted phenyl, optionally substituted pyridazine, optionally substituted pyrazine, optionally substituted pyrimidine, optionally substituted pyrazole, optionally substituted pyrrolidine, optionally substituted

quinazoline, optionally substituted pyridine, optionally substituted thiazolidine or optionally substituted triazole.

4. The compound of claim 3, wherein R^1 , R^2 , R^4 , R^5 and R^6 , where present, are each independently hydrogen or an optionally substituted $-(C_1-C_4)$ alkyl.

5. The compound of claim 3 wherein

A is a bond;

D is optionally substituted cyclopentylene, optionally substituted bicyclo[2.2.2]octanylene, optionally substituted azetidiny, or optionally substituted piperidiny;

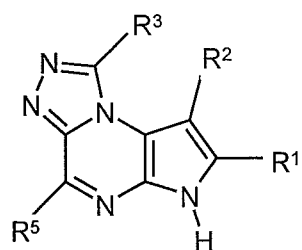
E is $-R^e-C(O)-R^e-$, $-R^e-N(R^a)-R^e-$, $-R^e-S(O)_2N(R^a)-R^e$, $-R^e-S(O)_2-R^e-$, or $-R^e-N(R^a)S(O)_2-R^e-$;

wherein R^e for each occurrence is independently a bond or an optionally substituted (C_1-C_6) alkylene; and

G is $-CN$, optionally substituted cyclopropyl, optionally substituted cyclobutyl, optionally substituted cyclopentyl, optionally substituted phenyl, optionally substituted pyrazine, optionally substituted pyridazine, optionally substituted pyrazole, or optionally substituted pyridine.

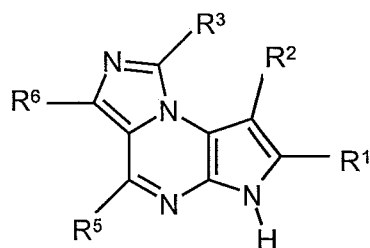
6. The compound of claim 5 wherein R^1 , R^2 , R^4 , R^5 and R^6 where present are each independently hydrogen or an optionally substituted $-(C_1-C_4)$ alkyl.

7. The compound of claim 6 wherein the compound is selected from the group consisting of a compound of Formula (Ia)



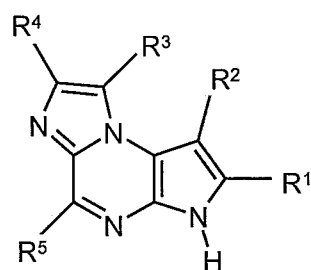
Formula (Ia) ,

a compound of Formula (Ib)



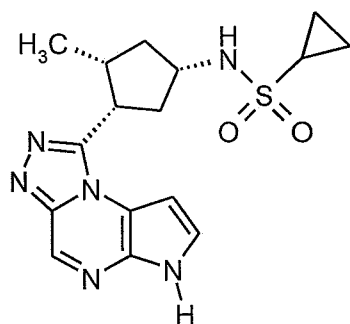
Formula (Ib) , and

a compound of Formula (Ic)

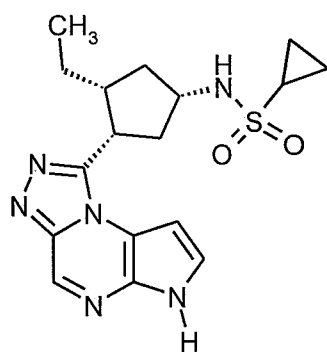


Formula (Ic) .

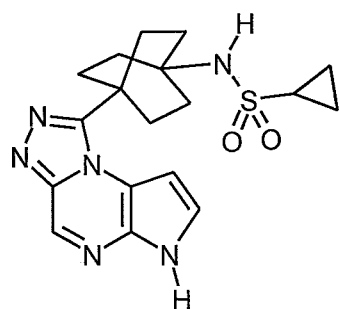
8. The compound of claim 7 wherein the compound is selected from the group consisting of



,

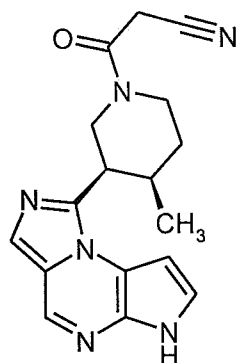


, and



9. The compound of claim 7 wherein the compound is of formula (Ic), wherein
- A is a bond;
 - D is optionally substituted cyclopentylene or optionally substituted piperidine;
 - E is $-R^e-N(R^a)-R^e$, $-R^e-S(O)_2-N(R^a)-R^e$, $-R^e-C(O)-R^e$, $-R^e-S(O)_2-R^e$, or $-R^e-N(R^a)S(O)_2-R^e$; and
 - G is $-CN$, optionally substituted phenyl, optionally substituted pyrazine, optionally substituted pyridazine, optionally substituted pyrazole, or optionally substituted pyridine.

10. The compound of claim 7 wherein the compound is of formula (Ib), wherein the compound is



11. The compound of claim 6 wherein G is optionally substituted phenyl, optionally substituted pyrazine, optionally substituted pyrazole, optionally substituted pyridazine or optionally substituted pyridine.

12. The compound of claim 1 wherein R³ is optionally substituted cyclopropyl, optionally substituted cyclobutyl, optionally substituted cyclopentyl, optionally substituted cyclohexyl, optionally substituted phenyl, optionally substituted adamantanyl, optionally substituted azetidiny, optionally substituted bicyclo[2.1.1]hexyl, optionally substituted bicyclo[2.2.1]heptyl, optionally substituted bicyclo[2.2.2]octyl, optionally substituted bicyclo[3.2.1]octyl, optionally substituted bicyclo[3.1.1]heptyl, optionally substituted azabicyclo[3.2.1]octanyl, optionally substituted azabicyclo[2.2.1]heptanyl, optionally substituted 2-azabicyclo[3.2.1]octanyl, optionally substituted azabicyclo[3.2.2]nonanyl, optionally substituted bicyclo[2.2.1]hept-2-enyl, optionally substituted piperidinyl, optionally substituted pyrrolidinyl or optionally substituted tetrahydrofuranyl;

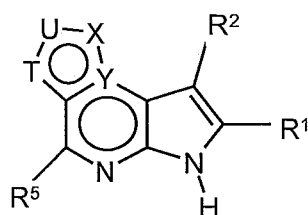
R² and R⁵ are each independently hydrogen, deuterium, -N(R^a)(R^b), halogen, -OR^a, -SR^a, -S(O)R^a, -S(O)₂R^a, -NO₂, -C(O)OR^a, -CN, -C(O)N(R^a)(R^b), -N(R^a)C(O)(R^b), -C(O)R^a, -C(OH)R^aR^b, -N(R^a)S(O)₂-R^b, -S(O)₂N(R^a)(R^b), -CF₃, -OCF₃, optionally substituted (C₁-C₆)alkyl, optionally substituted (C₃-C₁₂)cycloalkyl, optionally substituted (C₆-C₁₀)aryl, optionally substituted (C₁-C₁₀)heteroaryl or optionally substituted (C₁-C₁₀)heterocyclyl.

13. The compound of claim 12 wherein R¹ is optionally substituted (C₆-C₁₀)aryl or optionally substituted (C₁-C₁₀)heteroaryl.

14. The compound of claim 13 wherein R² is hydrogen, halogen, -CN, -C(O)NR^aR^b, -CF₃, optionally substituted -(C₁-C₆)alkyl, optionally substituted -(C₃-C₆)cycloalkyl, optionally substituted benzo(b)thienyl, optionally substituted benzimidazolyl, optionally substituted benzofuranyl, optionally substituted benzoxazolyl, optionally substituted benzothiazolyl, optionally substituted benzothiadiazolyl, optionally substituted furanyl, optionally substituted imidazolyl, optionally substituted indolyl, optionally substituted indolyl, optionally substituted indazolyl, optionally substituted isoxazolyl, optionally substituted isoindolyl, optionally substituted morpholyl, optionally substituted oxadiazolyl, optionally substituted phenyl, optionally substituted piperazyl, optionally substituted piperidyl, optionally substituted pyranyl, optionally substituted pyrazolyl, optionally substituted pyrazolo[3,4-*d*]pyrimidyl, optionally substituted pyridyl, optionally substituted pyrimidyl, optionally substituted pyrrolidyl, optionally substituted pyrrolyl, optionally substituted optionally pyrrolo[2,3-*d*]pyrimidyl, substituted quinolyl, optionally substituted thiomorpholyl, optionally substituted tetrahydropyranyl, optionally substituted tetrahydrofuranyl, optionally substituted tetrahydroindolyl, optionally substituted thiazolyl, or optionally substituted thienyl.

15. The compound of claim 14 wherein R¹ is optionally substituted azaindole, optionally substituted benzofuran, optionally substituted benzothiazole, optionally substituted benzoxazole, optionally substituted dihydropyrroloimidazole, optionally substituted furan, optionally substituted imidazole, optionally substituted imidazoxazole, optionally substituted imidazopyrazine, optionally substituted imidazopyridine, optionally substituted indazole, optionally substituted indole, optionally substituted isoquinoline, optionally substituted isothiazole, optionally substituted isoxazole, optionally substituted oxadiazole, optionally substituted oxazole, optionally substituted pyrazole, optionally substituted pyridine, optionally substituted pyrimidine, optionally substituted pyrazolopyridine, optionally substituted pyrrole, optionally substituted quinoline, optionally substituted quinazoline, optionally substituted thiazole, or optionally substituted thiophene.

16. The compound of claim 15 wherein R^5 is hydrogen, halogen, $-NH_2$ or $-N(R^a)(R^b)$.
17. A pharmaceutical composition comprising a compound of Formula (I) as defined in claim 1



Formula (I)

a pharmaceutically acceptable carrier and excipient and a second therapeutic agent selected from the group consisting of cytokine suppressive anti-inflammatory drugs, antibodies to or antagonists of other human cytokines or growth factors, IL-1, IL-2, IL-3, IL-4, IL-5, IL-6, IL-7, IL-8, IL-12, IL-15, IL-16, IL-21, IL-23, interferons, EMAP-II, GM-CSF, FGF, PDGF, CTLA or their ligands including CD154, adalimumab golimumab, certolizumab pegol (CDP870), tocilizumab, CDP 571, soluble p55 or p75 TNF receptors, Lenercept, TNF α converting enzyme inhibitors, IL-1 inhibitors, Interleukin 11, IL-18 antagonists, IL-12 antagonists, IL-12 antibodies, soluble IL-12 receptors, IL-12 binding proteins, non-depleting anti-CD4 inhibitors FK506, rapamycin, mycophenolate mofetil, leflunomide, NSAIDs, ibuprofen, corticosteroids, phosphodiesterase inhibitors, adenosine agonists, antithrombotic agents, complement inhibitors, adrenergic agents, IL-1 β converting enzyme inhibitors, T-cell signalling kinase inhibitors, metalloproteinase inhibitors, sulfasalazine, 6-mercaptapurines, derivatives p75TNFR IgG, sIL-1RI, sIL-1RII, sIL-6R, celecoxib, hydroxychloroquine sulfate, rofecoxib, infliximab, naproxen, valdecoxib, meloxicam, acetate, gold sodium thiomalate, aspirin, triamcinolone acetonide, propoxyphene napsylate/apap, folate, nabumetone, diclofenac, piroxicam, etodolac, diclofenac sodium, oxaprozin, oxycodone HCl, hydrocodone bitartrate/apap, diclofenac sodium/misoprostol, fentanyl, anakinra, tramadol HCl, salsalate, sulindac, cyanocobalamin/fa/pyridoxine, acetaminophen, alendronate sodium, morphine sulfate, lidocaine hydrochloride, indomethacin, glucosamine sulf/chondroitin, amitriptyline HCl, sulfadiazine,

oxycodone HCl/acetaminophen, olopatadine HCl misoprostol, naproxen sodium, omeprazole, cyclophosphamide, rituximab, IL-1 TRAP, MRA, CTLA4-IG, IL-18 BP, anti-IL-12, anti-IL15, VX-740, Roflumilast, IC-485, CDC-801, S1P1 agonists, FTY720, PKC family inhibitors, Ruboxistaurin, AEB-071, Mesopram, methotrexate, budenoside, dexamethasone, 5-aminosalicylic acid, olsalazine, IL-1ra, T cell signaling inhibitors, tyrosine kinase inhibitors, IL-11, mesalamine, prednisone, azathioprine, mercaptopurine, methylprednisolone sodium succinate, diphenoxylate/atrop sulfate, loperamide hydrochloride, ciprofloxacin/dextrose-water, tetracycline hydrochloride, fluocinonide, metronidazole, thimerosal/boric acid, cholestyramine/sucrose, ciprofloxacin hydrochloride, hyoscyamine sulfate, meperidine hydrochloride, midazolam hydrochloride, promethazine hydrochloride, sodium phosphate, sulfamethoxazole/trimethoprim, polycarbophil, propoxyphene napsylate, hydrocortisone, multivitamins, balsalazide disodium, codeine phosphate/apap, colesevelam HCl, cyanocobalamin, folic acid, levofloxacin, natalizumab, interferon-gamma, methylprednisolone, cyclosporine, 4-aminopyridine, tizanidine, interferon- β 1a, interferon- β 1b, interferon α -n3, interferon- α , interferon β 1A-IF, Peginterferon α 2b, hyperbaric oxygen, intravenous immunoglobulin, cladribine, FK506, prednisolone, antiinflammatory cytokines, interferon- β , caspase inhibitors, inhibitors of caspase-1, antibodies to CD40 ligand and CD80, alemtuzumab, dronabinol, daclizumab, mitoxantrone, xaliproden hydrochloride, fampridine, glatiramer acetate, sinnabidol, α -immunokine NNSO3, ABR-215062, Anergix.MS, chemokine receptor antagonists, BBR-2778, calagualine, CPI-1189, liposome encapsulated mitoxantrone, THC.CBD, cannabinoid agonists, MBP-8298, MNA-715, anti-IL-6 receptor antibody, neurovax, pirfenidone allotrap 1258 (RDP-1258), sTNF-R1, talampanel, teriflunomide, TGF-beta2, tiplimotide, VLA-4 antagonists, interferon gamma antagonists, IL-4 agonists, misoprostol, minocyclin, etanercept, betamethasone diprop augmented, dimethylsulfoxide, ketoprofen, tolmetin sodium, calcipotriene, glucosamine sulfate, risedronate sodium, thioguanine, alefacept, efalizumab, COX2 inhibitors, hydroxychloroquine, steroids, cytotoxics, inhibitors of PDE4, purine synthesis inhibitor, CTLA-4-IgG, anti-B7 family antibodies, anti-PD-1 family antibodies, anti-cytokine antibodies, fonotolizumab, anti-IFN γ antibody, anti-receptor antibodies, antibodies to B-cell surface molecules, LJP 394, anti-CD20 antibody and lymphostat-B.