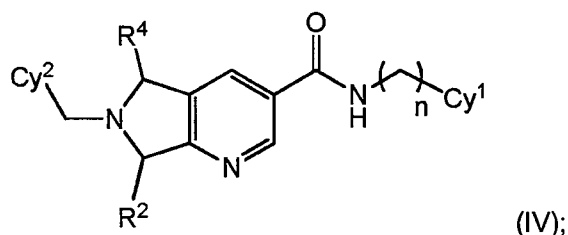


Claims

1. A compound of Formula (IV):



or a pharmaceutically acceptable salt thereof, wherein

R^2 is (C₁-C₃)alkyl;

R^4 is hydrogen, (C₁-C₃)alkyl, or =O;

n is 1 or 2;

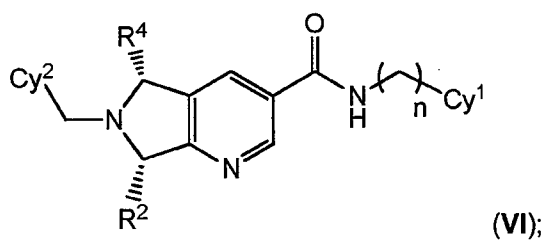
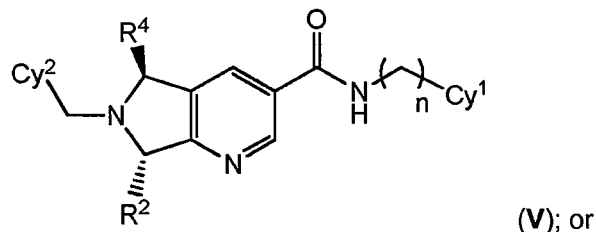
Cy^1 is phenyl, piperidiny, tetrahydro-2H-thiopyranyl 1,1-dioxide, pyridiny, piperaziny, azetidiny, imidazolyl, tetrahydropyranyl, 1,4-dioxanyl, pyridaziny, pyrazolyl, pyrrolidiny, cyclohexyl, morpholiny, 6,7-dihydro-5H-pyrrolo[2,1-c][1,2,4]triazolyl, 1,2,3,4-tetrahydro-1,8-naphthyridiny, 2,3-dihydro-1H-indenyl, or imidazo[1,2-a]pyrimidiny, each of which is optionally substituted with 1 to 2 groups independently selected from R^5 ;

Cy^2 is phenyl, pyrimidiny, cyclohexyl, or pyridiny, each of which are optionally substituted with 1 to 2 groups independently selected from R^6 ; and

R^5 and R^6 are each independently selected from halo, cyano, nitro, amino, hydroxy, carboxy, (C₁-C₆)alkyl, heterocyclyl, hydroxy(C₁-C₆)alkyl, CO₂H, (CH₂)₁₋₃COOH, (C₁-C₃)alkylcarbonyloxy, (C₃-C₆)cycloalkyl, hydroxy(C₃-C₆)cycloalkyl, (C₄-C₇)cycloalkylalkyl, (C₂-C₆)alkenyl, halo(C₂-C₆)alkenyl, hydroxy(C₂-C₆)alkenyl, (C₂-C₆)alkynyl, (C₃-C₆)cycloalkyl(C₂-C₄)alkynyl, halo(C₁-C₆)alkyl, halo(C₃-C₆)cycloalkyl, halo(C₄-C₇)cycloalkylalkyl, (C₁-C₆)alkoxy, (C₃-C₆)cycloalkoxy, (C₄-C₇)cycloalkylalkoxy, halo(C₁-C₆)alkoxy, halo(C₃-C₆)cycloalkoxy, halo(C₄-C₇)cycloalkylalkoxy, (C₁-C₆)alkylthio, (C₃-C₆)cycloalkylthio, (C₄-C₇)cycloalkylalkylthio, halo(C₁-C₆)alkylthio, halo(C₃-C₆)cycloalkylthio, halo(C₄-C₇)cycloalkylalkylthio, (C₁-C₆)alkylsulfinyl, (C₃-C₆)cycloalkylsulfinyl, (C₄-C₇)cycloalkylalkylsulfinyl, halo(C₁-C₆)alkylsulfonyl, halo(C₃-C₆)cycloalkylsulfonyl, halo(C₄-C₇)cycloalkylalkylsulfonyl, (C₁-C₆)alkylsulfonyl, (C₃-C₆)cycloalkylsulfonyl, halo(C₄-C₇)cycloalkylalkylsulfonyl, (C₁-C₆)alkylamino, di(C₁-C₆)alkylamino, (C₁-C₆)alkoxy(C₁-C₆)alkoxy, halo(C₁-C₆)alkoxy(C₁-C₆)alkoxy, (C₁-C₆)alkoxycarbonyl, H₂NCO, H₂NSO₂, (C₁-C₆)alkylaminocarbonyl, di(C₁-C₆)alkylaminocarbonyl, (C₁-C₃)alkoxy(C₁-

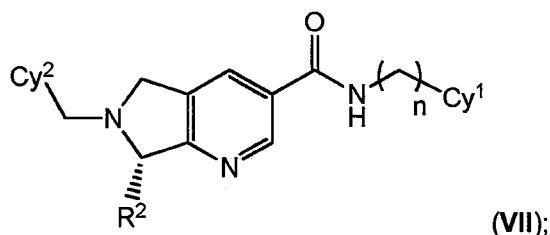
C₃)alkylaminocarbonyl, heterocyclcarbonyl, (C₁-C₆)alkylaminosulfonyl, di(C₁-C₆)alkylaminosulfonyl, heterocyclsulfonyl, (C₁-C₆)alkylcarbonylamino, (C₁-C₆)alkylcarbonylamino(C₁-C₆)alkyl, (C₁-C₆)alkylsulfonylamino, (C₁-C₆)alkylsulfonylamino(C₁-C₆)alkyl, (C₁-C₆)alkoxycarbonyl(C₁-C₆)alkoxy, (C₁-C₆)alkoxy(C₁-C₆)alkyl, halo(C₁-C₆)alkoxy(C₁-C₆)alkyl, hydroxy(C₁-C₆)alkoxy, aryl, heteroaryl, oxo, amino(C₁-C₆)alkyl, (C₁-C₆)alkylamino(C₁-C₆)alkyl, di(C₁-C₆)alkylamino(C₁-C₆)alkyl amino(C₂-C₆)alkoxy, (C₁-C₆)alkylamino(C₂-C₆)alkoxy, di(C₁-C₆)alkylamino(C₂-C₆)alkoxy, (C₁-C₆)alkylcarbonyl, hydroxy(C₁-C₆)alkylcarbonyl, (C₁-C₆)alkylhydroxycarbonyl, (C₁-C₆)alkylhydroxy(C₁-C₆)alkyl, (C₃-C₆)cycloalkylcarbonyl, (C₃-C₆)cycloalkylaminocarbonyl, {(C₃-C₆)cycloalkyl}{(C₁-C₆)alkyl}aminocarbonyl, di(C₃-C₆)cycloalkylaminocarbonyl, (C₃-C₆)cycloalkylaminosulfonyl, {(C₃-C₆)cycloalkyl}{(C₁-C₆)alkyl}aminosulfonyl, di(C₃-C₆)cycloalkylaminosulfonyl, cyano(C₁-C₆)alkyl, aminocarbonyl(C₁-C₆)alkyl, (C₁-C₆)alkylaminocarbonyl(C₁-C₆)alkyl, di(C₁-C₆)alkylaminocarbonyl(C₁-C₆)alkyl, (C₃-C₆)cycloalkylaminocarbonyl(C₁-C₆)alkyl, {(C₃-C₆)cycloalkyl}{(C₁-C₆)alkyl}aminocarbonyl(C₁-C₆)alkyl, [(C₁-C₆)alkyl(C₄-C₆)heterocycl](C₁-C₆)alkyl, and di(C₃-C₆)cycloalkylaminocarbonyl(C₁-C₆)alkyl.

2. The compound of Claim 1, wherein the compound is of Formula (V) or (VI):



or a pharmaceutically acceptable salt thereof.

3. The compound of Claim 1 or 2, wherein the compound is of Formula (VII):



or a pharmaceutically acceptable salt thereof.

4. The compound of any one of Claims 1 to 3, wherein Cy¹ is phenyl, piperidinyl, tetrahydro-2H-thiopyranyl 1,1-dioxide, pyridinyl, piperazinyl, azetidiny, imidazolyl, tetrahydropyranyl, 1,4-dioxanyl, pyridazinyl, pyrazolyl, pyrrolidinyl, cyclohexyl, morpholinyl, 6,7-dihydro-5H-pyrrolo[2,1-c][1,2,4]triazolyl, 1,2,3,4-tetrahydro-1,8-naphthyridinyl, 2,3-dihydro-1H-indenyl, or imidazo[1,2-a]pyrimidinyl, each of which is optionally substituted with 1 to 2 groups independently selected from R⁵, wherein at least one R⁵ is (C₁-C₃)alkylsulfonyl or (C₁-C₃)alkylaminosulfonyl.
5. The compound of any one of Claims 1 to 4, wherein R² is (C₁-C₃)alkyl; and Cy¹ is phenyl, pyridinyl, or piperidinyl, each of which is optionally substituted with 1 to 2 groups independently selected from R⁵, wherein at least one R⁵ is (C₁-C₃)alkylsulfonyl or (C₁-C₃)alkylaminosulfonyl.
6. The compound of any one of Claims 1 to 5, wherein Cy² is cyclohexyl optionally substituted with 1 to 2 groups independently selected from R⁵.
7. The compound of any one of Claims 1 to 6, wherein
R⁵ is selected from halo, (C₁-C₃)alkyl, halo(C₁-C₃)alkyl, cyano, hydroxy(C₁-C₃)alkyl, (C₁-C₃)alkoxycarbonyl, (C₁-C₃)alkylsulfonyl, (C₁-C₃)alkoxy, halo(C₁-C₃)alkoxy, oxo, hydroxy, (C₁-C₃)alkylcarbonyl, hydroxy(C₁-C₃)alkylcarbonyl, (C₁-C₃)alkylhydroxycarbonyl, (C₁-C₃)alkylaminosulfonyl, (C₁-C₃)alkylaminocarbonyl, di(C₁-C₃)alkylamino(C₂-C₆)alkoxy, (C₁-C₆)alkoxycarbonyl, [(C₁-C₃)alkyl(C₄-C₆)heterocyclyl](C₁-C₃)alkyl, and (C₁-C₃)alkylhydroxy(C₁-C₃)alkyl; and
R⁶ is selected from halo, (C₁-C₃)alkyl, halo(C₁-C₃)alkyl, cyano, hydroxy(C₁-C₃)alkyl, (C₁-C₃)alkoxycarbonyl, (C₁-C₃)alkylsulfonyl, (C₁-C₃)alkoxy, halo(C₁-C₃)alkoxy, oxo, hydroxy, aryl(C₁-C₃)alkoxycarbonyl, (C₁-C₃)alkylhydroxy(C₁-C₃)alkyl, heteroaryl, and (C₁-C₃)alkoxycarbonyl.

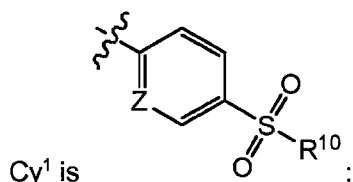
8. The compound of any one of Claims 1 to 7, wherein

R⁵ is selected from halo, (C₁-C₃)alkoxy, hydroxy, (C₁-C₃)alkyl, hydroxy(C₁-C₃)alkyl, halo(C₁-C₃)alkyl, (C₁-C₆)alkoxycarbonyl, di(C₁-C₃)alkylamino(C₂-C₆)alkoxy, [(C₁-C₃)alkyl(C₄-C₆)heterocyclyl](C₁-C₃)alkyl, oxo, (C₁-C₃)alkylcarbonyl, (C₁-C₃)alkylaminosulfonyl, (C₁-C₃)alkylsulfonyl, and cyano; and

R⁶ is selected from halo(C₁-C₃)alkyl, (C₁-C₃)alkoxy, halo, cyano, (C₁-C₃)alkoxycarbonyl, (C₁-C₃)alkylhydroxy(C₁-C₃)alkyl, hydroxy(C₁-C₃)alkyl, and halo(C₁-C₃)alkoxy.

9. The compound of any one of Claims 1 to 8, wherein R⁵ is selected from halo, cyano, (C₁-C₃)alkyl, (C₁-C₃)alkylaminosulfonyl, and (C₁-C₃)alkylsulfonyl; and R⁶ is selected from halo(C₁-C₃)alkyl, (C₁-C₃)alkoxy, halo, cyano, (C₁-C₃)alkoxycarbonyl, and halo(C₁-C₃)alkoxy.

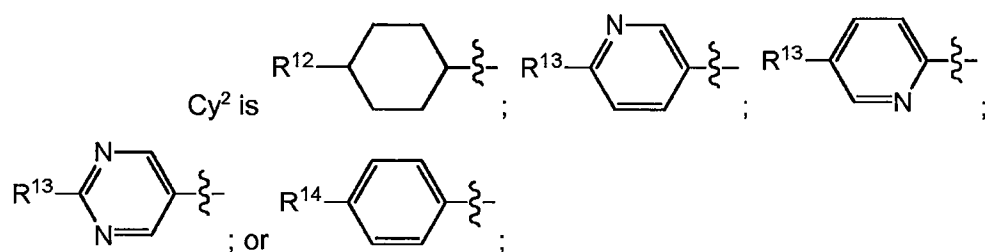
10. The compound of any one Claims 1 to 9, wherein



R¹⁰ is (C₁-C₃)alkyl or (C₁-C₃)alkylamino; and

Z is CH or N.

11. The compound of any one of Claims 1 to 5 and 7 to 10, wherein



R¹² is (C₁-C₃)alkoxycarbonyl, halo, dihalo, (C₁-C₃)alkoxy, or halo(C₁-C₃)alkyl;

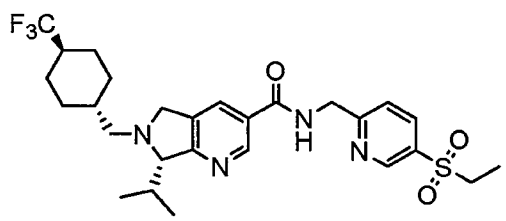
R¹³ is halo or halo(C₁-C₃)alkyl; and

R¹⁴ is halo, cyano, halo(C₁-C₃)alkyl, or halo(C₁-C₃)alkoxy.

12. The compound of Claim 11, wherein R¹² to R¹⁴ are each CF₃.

13. The compound of any one of Claims 1 to 12, wherein R² is isopropyl.

14. The compound of Claim 1, wherein the compound is of the Formula:



or a pharmaceutically acceptable salt thereof.

15. A pharmaceutical composition comprising a compound of any one of Claims 1 to 14, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier.

16. A compound of any one of Claims 1 to 14, or a pharmaceutically acceptable salt thereof, or a composition of Claim 15, for use in treating a disease or disorder selected from the group consisting of asthma, chronic obstructive pulmonary disease (COPD), bronchitis, allergic rhinitis, atopic dermatitis, contact dermatitis, acne, cystic fibrosis, allograft rejection, multiple sclerosis, scleroderma, arthritis, rheumatoid arthritis, juvenile rheumatoid arthritis, osteoarthritis, ankylosing spondylitis, systemic lupus erythematosus (SLE), psoriasis, Hashimoto's disease, pancreatitis, autoimmune diabetes, type I diabetes, autoimmune ocular disease, ulcerative colitis, Crohn's disease, regional enteritis, inflammatory bowel disease (IBD), inflammatory bowel syndrome (IBS), Sjögren's syndrome, optic neuritis, obesity, hepatosteatorrhea, adipose tissue-associated inflammation, insulin resistance, type II diabetes, neuromyelitis optica, myasthenia gravis, age related macular degeneration, dry eye, uveitis, Guillain-Barré syndrome, psoriasis, psoriatic arthritis (PsA), steroid resistant asthma, Graves' disease, scleritis, major depression, seasonal affective disorder, PTSD, bipolar disorder, autism, epilepsy, Alzheimer's, CNS disorders associated with altered sleep and/or circadian rhythms, endometriosis, obstructive sleep apnea syndrome (OSAS), Behçet's disease, dermatomyositis, polymyositis, graft versus host disease, primary biliary cirrhosis, liver fibrosis, non-alcoholic fatty liver disease (NAFLD), sarcoidosis, primary sclerosing cholangitis, autoimmune thyroid disease, autoimmune polyendocrine syndrome type I, autoimmune polyendocrine syndrome type II, celiac disease, neuromyelitis, juvenile idiopathic arthritis, systemic sclerosis, myocardial infarction, pulmonary hypertension, osteoarthritis, cutaneous leishmaniasis, sinonasal polyposis, and cancer.

17. A compound of any one of Claims 1 to 14, or a pharmaceutically acceptable salt thereof, or a composition of Claim 15, for use in treating a disease or disorder selected from the group consisting of asthma, atopic dermatitis, acne, Crohn's disease, regional enteritis, ulcerative colitis, Sjögren's syndrome, uveitis, Behçet's disease, dermatomyositis, multiple sclerosis, ankylosing spondylitis, systemic lupus erythematosus (SLE), scleroderma, psoriasis, psoriatic arthritis (PsA), steroid resistant asthma, and rheumatoid arthritis.

18. Use of a compound of any one of Claims 1 to 14, or a pharmaceutically acceptable salt thereof, or a composition of Claim 15, for the manufacture of a medicament for treating a disease or disorder selected from the group consisting of asthma, chronic obstructive pulmonary disease (COPD), bronchitis, allergic rhinitis, atopic dermatitis, contact dermatitis, acne, cystic fibrosis, allograft rejection, multiple sclerosis, scleroderma, arthritis, rheumatoid arthritis, juvenile rheumatoid arthritis, osteoarthritis, ankylosing spondylitis, systemic lupus erythematosus (SLE), psoriasis, Hashimoto's disease, pancreatitis, autoimmune diabetes, type I diabetes, autoimmune ocular disease, ulcerative colitis, Crohn's disease, regional enteritis, inflammatory bowel disease (IBD), inflammatory bowel syndrome (IBS), Sjögren's syndrome, optic neuritis, obesity, hepatosteatorrhea, adipose tissue-associated inflammation, insulin resistance, type II diabetes, neuromyelitis optica, myasthenia gravis, age related macular degeneration, dry eye, uveitis, Guillain-Barré syndrome, psoriasis, psoriatic arthritis (PsA), steroid resistant asthma, Graves' disease, scleritis, major depression, seasonal affective disorder, PTSD, bipolar disorder, autism, epilepsy, Alzheimer's, CNS disorders associated with altered sleep and/or circadian rhythms, endometriosis, obstructive sleep apnea syndrome (OSAS), Behçet's disease, dermatomyositis, polymyositis, graft versus host disease, primary biliary cirrhosis, liver fibrosis, non-alcoholic fatty liver disease (NAFLD), sarcoidosis, primary sclerosing cholangitis, autoimmune thyroid disease, autoimmune polyendocrine syndrome type I, autoimmune polyendocrine syndrome type II, celiac disease, neuromyelitis, juvenile idiopathic arthritis, systemic sclerosis, myocardial infarction, pulmonary hypertension, osteoarthritis, cutaneous leishmaniasis, sinonasal polyposis, and cancer.

19. The use of Claim 18, wherein the disease or disorder is selected from the group consisting of asthma, atopic dermatitis, acne, Crohn's disease, regional enteritis, ulcerative colitis, Sjögren's syndrome, uveitis, Behçet's disease, dermatomyositis, multiple sclerosis, ankylosing spondylitis, systemic lupus erythematosus (SLE), scleroderma, psoriasis, psoriatic arthritis (PsA), steroid resistant asthma, and rheumatoid arthritis.