STING can be activated by other types of cyclic-di-nucleotides (CDNs), such as c-di-AMP, c-di-GMP, and 3',3'-cGAMP from bacteria. Following the signal transduction, STING is rapidly degraded to prevent it from constitutive signaling of the inflammatory responses.

While eliciting robust host defense responses, aberrant STING signaling fuels dysregulated immune responses associated with many pathologies. Gain-of-function (GoF) human STING mutations are the root cause of STING-associated vasculopathy with onset in infancy (SAVI), a monogenic disease characterized by the onset of auto-inflammation conditions called type I interferonopathies. Mechanistically, the disease-causing substitutions trigger ligand-independent, constitutive STING activation. Besides, STING is implicated in DNA-driven inflammations, such as Aicardi-Goutières Syndrome (AGS) and genetic forms of lupus. Unlike SAVI, the STING mediated continuous innate immune activation in AGS is caused by deficiencies in self-DNA clearance and metabolisms due to mutations in endonuclease gene TREX1 and/or DNASE2. Consistently, genetic and pharmacological inhibition of STING ameliorates systemic inflammation and morbidity in the Trex1-/- mouse model. Apart from genetic disorders, robust preclinical and clinical evidence supports a general pathogenic role of STING in a range of inflammation-associated disorders such as SLE and RA. A direct link between the cGAS-STING pathway and SLE was established by observing that PBMC from a subset of SLE patients has elevated cytosolic cGAMP than healthy controls. In addition, membrane vesicles from apoptotic cells in SLE sera have high ISGs-stimulating activities dependent on cGAS-STING. And that disrupting STING signaling ameliorated the development of lupus-like phenotypes in FcγrIIb-/- mice. Furthermore, multiple recent studies associate STING with distinct types of neurodegeneration. Taking Parkinson's disease as an example, missense mutations in PARKIN and PINK resulted in the accumulation of mitochondrial DNA that triggers neuronal inflammation in a cGAS-STING dependent manner. The absence of STING rescued the motor deficit and neuronal cell loss in the mouse disease model. Finally, STING also mediates tumorigenic DNA responses caused by chromosomal instability during cancer metastasis, and that STING-deficiency confers protection against colorectal and skin cancer in the mouse.

SUMMARY

In one aspect, the present disclosure relates to a compound of formula (Ia),

$_{7}$cycloalkyl)$_{2}$aminocarbonyl, C_{1-6}alkylC_{3-7}cycloalkylaminocarbonyl, haloC_{1-6}alkylaminocarbonyl, halo(C_{1-6}alkyl)$_{2}$aminocarbonyl, haloC_{3-7}cycloalkylaminocarbonyl, halo(C_{3-7}cycloalkyl)$_{2}$aminocarbonyl, haloC_{1-6}alkylC_{3-7}cycloalkylaminocarbonyl, C_{1-6}alkylhaloC_{3-7}cycloalkylaminocarbonyl, (C_{1-6}alkyl)$_{3}$ammonio, C_{1-6}alkylsulfanyl, C_{1-6}alkylsulfinyl, C_{1-6}alkylsulfonyl, C_{3-7}cycloalkylsulfanyl, C_{3-7}cycloalkylsulfinyl, C_{3-7}cycloalkylsulfonyl, haloC_{1-6}alkylsulfanyl, haloC_{1-6}alkylsulfinyl, haloC_{1-6}alkylsulfonyl, haloC_{3-7}cycloalkylsulfanyl, haloC_{3-7}cycloalkylsulfinyl, haloC_{3-7}cycloalkylsulfonyl, haloC_{1-6}alkoxysulfonyl, C_{1-6}alkylaminosulfonyl, (haloC_{1-6}alkylamino)sulfonyl, (C_{1-6}alkyl)$_{2}$aminosulfonyl, (haloC_{1-6}alkyl)$_{2}$aminosulfonyl, (C_{1-6}alkyl)$_{2}$phosphoryl, (C_{1-6}alkoxy)$_{2}$phosphoryl, hydroxy(C_{1-6}alkoxy)phosphoryl, C_{1-6}alkoxyphosphoryl, hydroxy, hydroxyC_{1-6}alkyl, hydroxyhaloC_{1-6}alkyl, hydroxyC_{1-6}alkoxy, hydroxylhaloC_{1-6}alkoxy, hydroxyC_{3-7}cycloalkyl, hydroxyhaloC_{3-7}cycloalkyl and sulfonimidoyl optionally substituted by H, C_{1-6}alkyl, C_{1-6}alkoxy, aryl, heteroaryl and heterocyclyl;

M^2 is arylene, heterocyclylene or heteroarylene, said arylene, heterocyclylene or heteroarylene being optionally substituted by R^1;

M^3 is heterocyclylene which is unsubstituted or optionally substituted by R^3;

R^2 is deuterio, R, ROC_{1-6}alkyl, RSC_{1-6}alkyl, RS$O_2$$C_{1-6}$alkyl, RNH$C_{1-6}$alkyl or (R)$_2NC_{1-6}$alkyl;

R^3 is heterocyclyl, heteroaryl, aryl, heterocyclylcarbonyl, heteroarylcarbonyl or arylcarbonyl, said heterocyclyl, heteroaryl, aryl, heterocyclylcarbonyl, heteroarylcarbonyl and arylcarbonyl being unsubstituted or optionally substituted by R^1;

Q^1 is -M-, -M-O-M-, -M-S-M-, -M-SO_2-M-, -M-SO(NH)-M-, -M-SO(NR)-M-, -M-SiR_2-M- or -M-N(R)-M-; wherein M is C_{1-6}alkylene unsubstituted or optionally substituted by the substituents independently selected from deuterio, halogen, cyano, carboxy, R, RO, RS, RNH, R_2N, RSO_2, ROC_{1-6}alkyl, RSC_{1-6}alkyl, RNHC_{1-6}alkyl, (R)$_2$NC_{1-6}alkyl and RS$O_2$$C_{1-6}$alkyl;

Q^2 is NH, O, S, SO, SO_2, P(O)R, PO(NR), Se, N(R), SiR_2, heterocyclyl, heteroaryl or aryl;

or a pharmaceutically acceptable salt thereof.

Another object of the present disclosure is related to novel compounds of formula (I) or (Ia). Their manufacture, medicaments based on a compound in accordance with the disclosure

and their production as well as the use of compounds of formula (I) or (Ia) as STING antagonist, and for the treatment or prophylaxis of autoimmune diseases, inflammatory diseases, neurological disorders diseases, metabolic diseases, cardiovascular diseases, or selective types of cancers where overexpression or activation of STING is implicated. The compounds of formula (I) or (Ia) show superior STING antagonism activity. In addition, the compounds of formula (I) or (Ia) also show good cytotoxicity, phototoxicity, solubility, hPBMC, human microsome stability and SDPK profiles, as well as low CYP inhibition.

In another aspect, the present disclosure relates to a compound selected from:

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[2-(2,4-difluorophenyl)-3-methyl-imidazole-4-carbonyl]-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7-oxa-10,13,16,18-tetrazapentacyclo[14.6.1.1^{2,6}.1^{8,11}.0^{19,23}]pentacosa-1(22),2,4,6(25),17,19(23),20-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(2-hydroxyethyl)-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-13-benzyl-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(2-methoxyethyl)-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18, 20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(3-morpholinopropyl)-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

3-[(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24), 21-heptaen-13-yl]propyl-trimethyl-ammonium;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyridazin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-13-methyl-10-[3-(3-pyridyl)imidazo[1,5-a]pyrazin-8-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[7-(2,4-difluorophenyl)imidazo[5,1-f][1,2,4]triazin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl] -15-(2-methoxyethoxy)-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl] -15-(2-methoxyethoxy)-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15- (2-methoxyethoxy)-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-4-(oxetan-3-yl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-22-fluoro-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(2-fluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(2-fluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-[4-fluoro-2-(trideuteriomethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluoro-6-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,6-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(1-bicyclo[1.1.1]pentanyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[3-(2,4-difluorophenyl)-[1,2,4]triazolo[4,3-a]pyrazin-8-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(3,4-difluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-15-methoxy-13,18-dimethyl-10-[1-(3-methyl-1-bicyclo[1.1.1]pentanyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(3-fluoro-1-bicyclo[1.1.1]pentanyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

5-fluoro-2-[4-[(8S,11S,15R)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-1-yl]benzonitrile;

[(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaene-4-carbonitrile;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaene-4-carbonitrile;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl) pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(cyclopropoxy) -4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18, 20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-4,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-4,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-4,10,13,17,19,26-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-4,10,13,17,19,26-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-4,5,10,13,17,19-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-4,5,10,13,17,19-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18, 20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-4,5,10,13,17,19-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-4,5,10,13,17,19-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-(hydroxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-(hydroxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-(4-fluoro-2-methoxy-phenyl)-4-[(8S,11S)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidine-6-carboxamide;

1-(4-fluoro-2-methoxy-phenyl)-N,N-dimethyl-4-[(8S,11S)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidine-6-carboxamide;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(methylamino)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-methoxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-hydroxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

8S,11S)-10-[1-(2,4-difluorophenyl)-6-ethoxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(2-methoxyethylamino)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(2-morpholinoethoxy)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-10-[1-(4-fluoro-2-methylsulfanyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(difluoromethoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(difluoromethoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(difluoromethoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(difluoromethoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26 hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(6-fluoro-2-hydroxy-3-pyridyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,6-difluoro-3-pyridyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-13-methyl-10-[3-(2-thienyl)imidazo[1,5-a]pyrazin-8-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(5-fluoro-3-methoxy-2-pyridyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(3-fluoro-5-methoxy-2-pyridyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(3,5-difluoro-2-pyridyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[2-(2-chloro-4-fluoro-phenyl)pyrimidin-5-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(methylamino)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,22,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-13-(cyclopropylmethyl)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,21,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(2-morpholinoethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(2-morpholinoethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(2-methoxyethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[2-fluoro-4-(2-methoxyethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(oxetan-3-ylmethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[2-fluoro-4-(oxetan-3-ylmethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(oxetan-3-yloxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[2-fluoro-4-(oxetan-3-yloxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-[(4-methylmorpholin-2-yl)methoxy]phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-18-(2-methoxyethyl)-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-18-(2-morpholinoethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-18-(2-hydroxyethyl)-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-18-[2-[(4aR,7aS)-2,3,4a,5,7,7a-hexahydro-[1,4]dioxino[2,3-c]pyrrol-6-yl]ethyl]-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-18-[2-(4-methylsulfonylpiperazin-1-yl)ethyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-18-[2-(1,2,4-triazol-1-yl)ethyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-18-(2-pyrazol-1-ylethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-23-fluoro-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,19,21,28-hexazapentacyclo[17.6.1.1^{2,6}.1^{8,11}.0^{22,26}]octacosa-1(25),2(28),3,5,20,22(26),23-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-3-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(trifluoromethyl)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

5-fluoro-2-[4-[(8S,11S)-13-methyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-1-yl]benzonitrile;

5-fluoro-2-[4-[(8S,11S)-13-methyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-1-yl]benzoic acid;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-ethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-13-ethyl-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(2-methoxyethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-13-(2-morpholinoethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(3-hydroxypropyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-[3-(tetrazol-2-yl)propyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-[3-(tetrazol-1-yl)propyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-[3-(4-methylsulfonylpiperazin-1-yl)propyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-(2-hydroxyethoxy)-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-(2-hydroxyethoxy)-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-15-(2-morpholinoethoxy)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-15-(2-morpholinoethoxy)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-18,22-dideuterio-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-15-hydroxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-hydroxy-13,15,18-trimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-hydroxy-13,15,18-trimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-hydroxy-13,15,18-trimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-hydroxy-13,15,18-trimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

[(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-dimethylphosphoryl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

1-(2,4-difluorophenyl)-4-[(8S,11S)-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidine-6-carboxylic acid;

1-(2,4-difluorophenyl)-4-[(8S,11S)-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidine-6-carbonitrile;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-c]pyridazin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-c]pyridazin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-b]pyridin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-b]pyridin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-b]pyridin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-b]pyridin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-23-chloro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(hydroxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-morpholino-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

Ethyl 1-(2,4-difluorophenyl)-4-[(8S,11S)-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidine-6-carboxylate;

(8S,11S)-10-[1-(4-fluoro-2-methylsulfanyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methylsulfinyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methylsulfonyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-dimethylphosphoryl-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-4-[(3,3-difluoroazetidin-1-yl)methyl]-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-4-(morpholinomethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-4,15-dimethoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-4,15-dimethoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-4,13-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-4,13-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[6-amino-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[6-amino-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-13-(cyclopropylmethyl)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-hydroxy-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-vinyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2-cyclopropyl-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2-bromo-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-18-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,22-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-4-(difluoromethoxy)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-4-(cyclopropoxy)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-13-cyclopropyl-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-(trideuteriomethoxy)-13-(trideuteriomethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[2-(difluoromethyl)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)-1H-pyrazolo[4,3-d]pyrimidin-7-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-18-ethyl-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-18-ethyl-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-(hydroxymethyl)-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[6-cyclobutyl-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4,5-difluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-[2-(difluoromethoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-[2-fluoro-4-(trifluoromethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-15-methoxy-10-[1-(2-methoxyphenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-chloro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(5-chloro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-4-(methoxymethyl)-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaene-4-carbonitrile;

(8S,11S)-10-[1-[4-fluoro-2-(trifluoromethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-18-(trifluoromethyl)-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-4,13-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13-ethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13-ethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-4,13-dimethyl-3,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-3,5,7,10,13,17,19-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-thia-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(trideuteriomethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)triazolo[4,5-c]pyridin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(methoxymethyl)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluoro-6-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(5-fluoropyrimidin-2-yl)pyrazolo[4,3-c]pyridin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-3,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-3,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-3,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

8S,11S,15S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-4,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-4,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-4,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(1-bicyclo[1.1.1]pentanyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(5-fluoropyrimidin-2-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-fluoro-4-phenyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(4,4,4-trifluorobutyl)-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-chloro-4-fluoro-phenyl)pyrazole-4-carbonyl]-22-fluoro-13-(3-pyrimidin-2-yloxypropyl)-4,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-22-chloro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-21-chloro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-21-fluoro-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S,15R)-10-[1-(3,5-difluoro-2-pyridyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(3,5-difluoro-2-pyridyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-4,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(3-methoxypropyl)-4,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-4,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-4,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-16-oxa-7,10,13,19,21,28-hexazapentacyclo[17.6.1.1^{2,6}.1^{8,11}.0^{22,26}]octacosa-1(25),2(28),3,5,20,22(26),23-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluoro-6-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-15-methoxy-13,18-dimethyl-10-(1-pyrimidin-5-ylpyrazolo[3,4-d]pyrimidin-4-yl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(11S,14S,18R)-13-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-25-fluoro-18-methoxy-16,21-dimethyl-5-oxa-10,13,16,20,22,29-hexazahexacyclo[18.6.1.1^{2,9}.1^{11,14}.0^{3,7}.0^{23,27}]nonacosa-1(26),2(29),3(7),8,21,23(27),24-heptaen-15-one;

(8S,11S,15R)-22-fluoro-15-methoxy-13,18-dimethyl-10-[1-methyl-6-(3-thienyl)pyrazolo[4,3-c]pyridin-3-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[6-(2,4-difluorophenyl)-1-methyl-pyrazolo[4,3-c]pyridin-3-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-21,23-difluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluoro-6-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[2-(2,4-difluorophenoxy)-4-pyridyl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-15-ethoxy-10-[1-[4-fluoro-2-(2-oxopyrrolidin-1-yl)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,15,18-trimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,15,18-trimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,15,18-trimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,15,18-trimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,21,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,21,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-21-fluoro-18-hydroxy-13-methyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaene-14-carboxylic acid;

(8S,11S,15R)-10-[6-ethoxy-1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-methoxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-ethoxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(morpholinomethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-18-isopropyl-13-methyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

methyl (8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-12-oxo-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaene-18-carboxylate;

(8S,11S)-18-(chloromethyl)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-18-hydroxy-13-methyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-18-amino-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-N,13-dimethyl-12-oxo-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaene-18-carboxamide;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

N-[(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-12-oxo-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-18-yl]acetamide;

(8S,11S,15R)-10-[1-(2,4-difluoro-6-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-18-cyclopropyl-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-fluoro-4-tetrahydrofuran-3-yl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22,23-difluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-[2-fluoro-4-(2-oxopyrrolidin-1-yl)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-18-methyl-15-(trideuteriomethoxy)-13-(trideuteriomethyl)-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(oxetan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-tetrahydrofuran-3-yl-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-tetrahydropyran-4-yl-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(oxetan-3-ylmethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(oxetan-2-ylmethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1-methylsulfonylazetidin-3-yl)methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(1-acetylazetidin-3-yl)methyl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(oxetan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[2-hydroxy-1-(hydroxymethyl)ethyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[1-(hydroxymethyl)-2-methoxy-ethyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(1-methylsulfonylazetidin-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(1-acetylazetidin-3-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(1-methylsulfonylpyrrolidin-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[1-(2,2,2-trifluoroacetyl)pyrrolidin-3-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(1-acetylpyrrolidin-3-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1-methylsulfonylpyrrolidin-3-yl)methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[[1-(2,2,2-trifluoroacetyl)pyrrolidin-3-yl]methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(1-acetylpyrrolidin-3-yl)methyl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(1-methyl-4-piperidyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-18-methyl-13-(3-morpholinopropyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-18-methyl-13-[3-(2-oxa-5-azabicyclo[2.2.1]heptan-5-yl)propyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-18-methyl-13-[3-(2-oxa-6-azaspiro[3.3]heptan-6-yl)propyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-13-[3-[(4a*R*,7a*S*)-2,3,4a,5,7,7a-hexahydro-[1,4]dioxino[2,3-c]pyrrol-6-yl]propyl]-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-18-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1-ethylsulfonylazetidin-3-yl)methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(1-cyclopropylsulfonylazetidin-3-yl)methyl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[[1-(2,2,2-trifluoroethyl)azetidin-3-yl]methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)-6-(pyrrolidine-1-carbonyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-(2,4-difluorophenyl)-4-[(8S,11S)-22-fluoro-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]-N-(2-hydroxyethyl)pyrazolo[3,4-d]pyrimidine-6-carboxamide;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)-6-(morpholine-4-carbonyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-7-(methoxymethyl)pyrazolo[4,3-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-5,13,18-trimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(1,3-dihydroisobenzofuran-4-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-7-methyl-6-oxo-pyrazolo[3,4-b]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2,2,2-trifluoro-1-hydroxy-ethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[7-cyclopropyl-1-(2,4-difluorophenyl)-6-oxo-pyrazolo[3,4-b]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-7-(2-methoxyethyl)-6-oxo-pyrazolo[3,4-b]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-7-methoxy-pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26- hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-7-methoxy-pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(1-methoxycyclopropyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(1-hydroxycyclopropyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-[2-(azetidin-1-yl)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-morpholino-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(2-fluoro-4-morpholino-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-[4-fluor-2-(2-oxa-6-azaspiro[3.3]heptan-6-yl)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluoro-6-methoxy-phenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-[4-fluoro-2-(3-methoxycyclobutoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-[4-fluoro-2-[1-(methoxymethyl)cyclopropoxy]phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(hydroxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-4,22-difluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2,2-dioxo-2λ⁶-thia-6-azaspiro[3.3]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(hydroxymethyl)azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-oxa-6-azaspiro[3.3]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(1-hydroxy-1-methyl-ethyl)azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(3*R*,4*R*)-3-amino-4-fluoro-pyrrolidin-1-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-[(dimethylamino)methyl]azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-methylsulfonylazetidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(tetrahydrofuran-3-ylmethoxy)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(1-cyclopropylazetidin-3-yl)oxy-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[2-(4-methylpiperazin-1-yl)ethoxy]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[[1-(2,2-difluoroacetyl)azetidin-3-yl]methoxy]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(4-methylpiperazin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1-methylazetidin-3-yl)methoxy]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1-methylpyrrolidin-3-yl)methoxy]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[1-(2-fluoroethyl)azetidin-3-yl]oxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(oxetan-3-ylmethoxy)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(3*S*,4*S*)-3-amino-4-methoxy-pyrrolidin-1-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(dimethylamino)azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(3-amino-3-methyl-azetidin-1-yl)-1-(2,4-difluorophenyl)pyrazolo [3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[1-(2,2-difluoroethyl)azetidin-3-yl]oxy-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-morpholinoazetidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[[1-(2,2-difluoroethyl)azetidin-3-yl]methoxy]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[1-(2,2-difluoroethyl)azetidin-3-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]ethyl acetate;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(5-fluoro-3-methylsulfanyl-2-pyridyl)-6-methylsulfanyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*)-1-hydroxyethyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*S*)-1-hydroxyethyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-oxa-6-azaspiro[3.3]heptan-6-ylmethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(morpholinomethyl)azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

2-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]acetonitrile;

(8*S*,11*S*,15*R*)-10-[6-(4,7-diazaspiro[2.5]octan-7-ylmethyl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[4-(oxetan-3-yl)piperazin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(4,7-diazaspiro[2.5]octan-7-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[4-(2H-tetrazol-5-yl)-1-piperidyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(4-methyl-4,7-diazaspiro[2.5]octan-7-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(3,5-difluoro-2-pyridyl)-6-(3-morpholinoazetidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-methyl-3,6-diazabicyclo[3.1.1]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(3,4,6,7,8,8a-hexahydro-1H-pyrrolo[1,2-a]pyrazin-2-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[[(2*R*,8*S*)-2-fluoro-1,2,3,5,6,7-hexahydropyrrolizin-8-yl]methoxy]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-

dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(3a*R*,6a*S*)-2-methyl-1,3,3a,4,6,6a-hexahydropyrrolo[3,4-c]pyrrol-5-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-cyclopentyl-1-(3,5-difluoro-2-pyridyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-oxa-6-azabicyclo[3.1.1]heptan-6-ylmethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(4a*R*,7a*R*)-4-methyl-2,3,4a,5,7,7a-hexahydropyrrolo[3,4-b][1,4]oxazin-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-benzyl-4-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]piperazine-2-carboxylic acid;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(9-methyl-3-oxa-7,9-diazabicyclo[3.3.1]nonan-7-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(4-oxazol-5-yl-1-piperidyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-methyl-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-methylsulfonyl-pyrazolo[3,4-*d*]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-10-(3-prop-1-ynylimidazo[1,5-*a*]pyrazin-8-yl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-methylsulfanyl-pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluoro-6-methoxy-phenyl)-6-[[1-(2-methoxyacetyl)azetidin-3-yl]methyl]pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(3-methoxy-3-methyl-but-1-ynyl)phenyl]pyrazolo[3,4-*d*]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(3-methoxypyrrolidin-1-yl)phenyl]pyrazolo[3,4-*d*]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluoro-6-tetrahydrofuran-3-yloxy-phenyl)pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

2-[2-[4-[(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-*d*]pyrimidin-1-yl]-3,5-difluoro-phenoxy]acetonitrile;

(8*S*,11*S*,15*S*)-10-[1-[2,4-difluoro-6-[(1-methylsulfonylazetidin-3-yl)methoxy]phenyl]pyrazolo[3,4-*d*]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-4-(cyclopropylamino)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(6-fluoro-1H-indazol-5-yl)pyrazolo[3,4-*d*]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)indazol-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(trideuteriomethoxy)phenyl]-6-(oxetan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(trideuteriomethoxy)phenyl]-6-[3-oxa-6-azabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-methyl-7-oxo-pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-methoxyethyl)-7-oxo-pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[7-benzyloxy-1-(2,4-difluorophenyl)pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[6-(cyclopropylmethyl)-1-(2,4-difluorophenyl)-7-oxo-pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[(1,1-dioxothietan-3-yl)methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

2-[3-[1-(2,4-difluorophenyl)-4-[(8S,11S,15S)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]azetidin-1-yl]acetonitrile;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-(1,1-dioxothian-4-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(methylsulfonylmethyl)azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[6-(1-cyclopropylazetidin-3-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[(1,1-dioxothietan-3-yl)methylsulfanyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-[4-fluoro-2-(trideuteriomethoxy)phenyl]-6-(3-oxa-6-azabicyclo[3.1.1]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(azetidin-1-yl)-1-[4-fluoro-2-(trideuteriomethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(azetidin-1-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-oxa-6-azabicyclo[3.1.1]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-oxa-6-azabicyclo[3.1.1]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(dimethylamino)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-oxa-5-azabicyclo[2.2.1]heptan-5-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(8-oxa-3-azabicyclo[3.2.1]octan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-oxa-8-azabicyclo[3.2.1]octan-8-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-imidazol-1-yl-pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2,2,2-trifluoroethoxy)pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-(oxetan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(3*S*)-1-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]pyrrolidine-3-carbonitrile;

8-[1-(2,4-difluorophenyl)-4-[(8S,11S,15R)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]-6,7,9,9a-tetrahydro-1*H*-pyrazino[2,1-c][1,4]oxazin-4-one;

(8*S*,11*S*,15*R*)-10-[6-(3,4,6,7,9,9a-hexahydro-1*H*-pyrazino[2,1-c][1,4]oxazin-8-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-

5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]azetidine-3-carbonitrile;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-oxa-6-azabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-oxa-6-azabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-methyl-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]azetidine-3-carbonitrile;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-

dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[8-(2-fluoroethyl)-5-oxa-2,8-diazaspiro[3.5]nonan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[8-(2-methoxyethyl)-5-oxa-2,8-diazaspiro[3.5]nonan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[5-(2-fluoroethyl)-8-oxa-2,5-diazaspiro[3.5]nonan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

4-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]morpholin-3-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-methyl-2-oxo-pyrrolidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,8-diazabicyclo[3.2.1]octan-8-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-5-(3-phenylpropylamino)pyrazole-4-carbonyl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(1*S*,5*S*)-3-cyclopropyl-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-oxopyrrolidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(6-oxa-3-azabicyclo[3.1.1]heptan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[6-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-3-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-[(3-fluorooxetan-3-yl)methyl]-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-[(3-fluorooxetan-3-yl)methyl]-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[6-hydroxy-6-(trifluoromethyl)-2-azaspiro[3.3]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(6-oxa-1-azaspiro[3.3]heptan-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[6-(2-fluoroethyl)-1,6-diazaspiro[3.3]heptan-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-(3-oxa-6-azabicyclo[3.1.1]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(2-fluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(2-fluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(2-fluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-

dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2,2,2-trifluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-hydroxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

2-[3-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]azetidin-1-yl]acetonitrile;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

3-[6-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]-3,6-diazabicyclo[3.1.1]heptan-3-yl]propanenitrile;

(8*S*,11*S*,15*R*)-10-[6-[(1*R*,4*R*)-5-(2,2-difluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(1*R*,4*R*)-5-(2,2-difluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(2,2,2-trifluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[6-[(1*R*,4*R*)-5-(2,2-difluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-5-(2-morpholinoethylamino)pyrazole-4-carbonyl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[11-benzyl-3-(2,4-difluorophenyl)-3,4,8,11-tetrazatricyclo[7.3.0.0^{2,6}]dodeca-1(9),2(6),4,7-tetraen-7-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[3-(2,4-difluorophenyl)-11-methyl-3,4,8,11-tetrazatricyclo[7.3.0.0^{2,6}]dodeca-1(9),2(6),4,7-tetraen-7-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[5-amino-1-(2,4-difluorophenyl)pyrazole-4-carbonyl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(2*S*)-2-(methoxymethyl)pyrrolidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

2-[6-[1-(2,4-difluorophenyl)-4-[(8*S*,11S,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]-3,6-diazabicyclo[3.1.1]heptan-3-yl]acetonitrile;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]azetidine-3-carbonitrile;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-methyl-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-

7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[7-(2-fluoroethyl)-1,7-diazaspiro[3.5]nonan-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[7-(oxetan-3-yl)-1,7-diazaspiro[3.5]nonan-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[6-(oxetan-3-yl)-1,6-diazaspiro[3.3]heptan-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(6,6-dioxo-6λ^{6}-thia-1-azaspiro[3.3]heptan-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,7-diazabicyclo[4.2.0]octan-7-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[3-(2,4-difluorophenyl)-11-(oxetan-3-yl)-3,4,8,11-tetrazatricyclo[7.3.0.0^{2,6}]dodeca-1(9),2(6),4,7-tetraen-7-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,5*S*)-3-(oxetan-3-yl)-3,6-diazabicyclo[3.2.0]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-

heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-hydroxy-2-oxo-pyrrolidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-oxopyrrolidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

3-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]oxazolidin-2-one;

3-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-

heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]oxazolidin-2-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(4-methyl-2-oxo-piperazin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-22-fluoro-10-[6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(2,2,2-trifluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[6-[3-(2,2-difluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2,2,2-trifluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-hydroxy-2-methyl-propanoyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(1-hydroxycyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(1-hydroxycyclobutanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

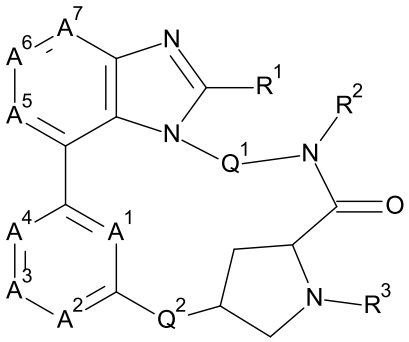
(8*S*,11*S*,15*R*)-10-[6-[3-(cyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one; and

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(3-fluorocyclobutyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

or a pharmaceutically acceptable salt thereof.

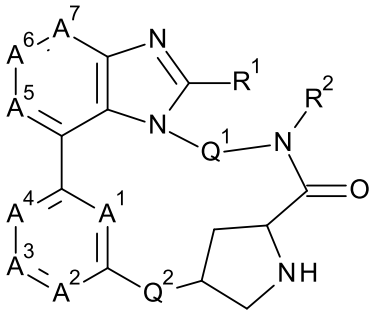
In another aspect, the present disclosure relates to a process for the preparation of a compound as described herein comprising any of the following steps:

a) the formation of compound of formula (I),



(I),

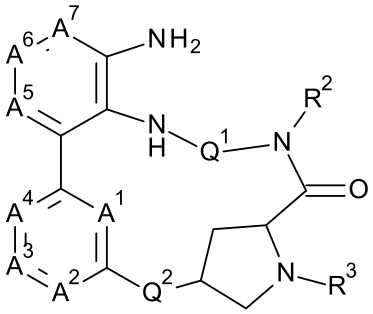
via nucleophilic substitution between compound of formula (VIII),

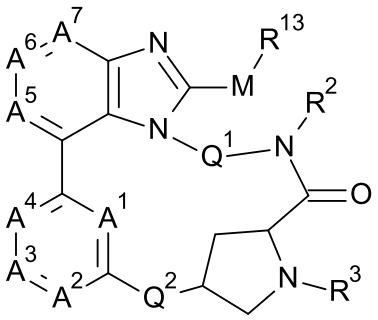
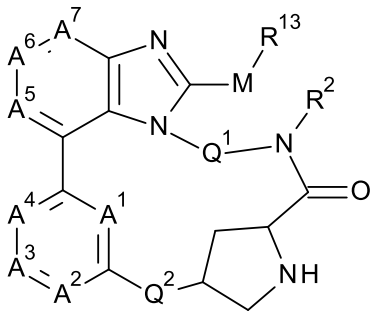


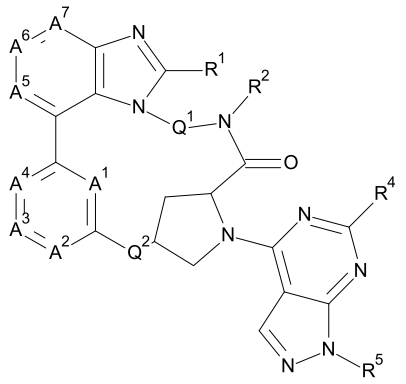
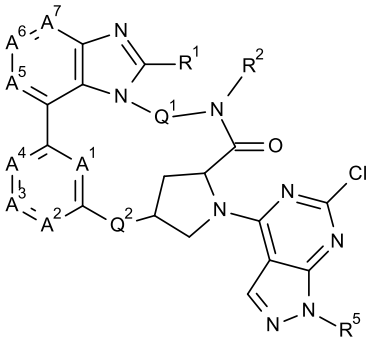
(VIII), and R^3X, in the presence of a base;

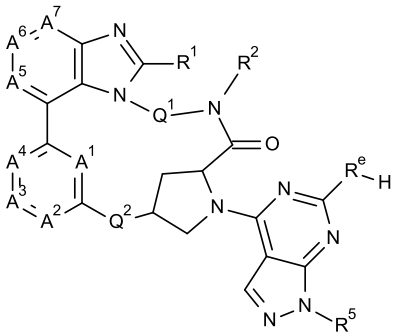
b) the formation of compound of formula (I) via Buchwald-Hartwig reaction between compound of formula (VIII) and R^3X, in the presence of a catalyst;

c) the formation of compound of formula (I) via condensation reaction between compound of formula (VIII) and R^3OH, in the presence of coupling reagent;

d) the formation of compound of formula (I) via cyclization reaction between compound of formula (XVIII) ,  (XVIII), and regents;

e) the formation of compound of formula (XXI),  (XXI), via nucleophilic substitution between compound of formula (XX),  (XX), and R^3X in the presence of a base;

f) the formation of compound of formula (XXII),  (XXII), via reaction between compound of formula (Ic),  (Ic), and alcohols or amines (R^4-X) in the presence of a base;

g) the amino group of compound of formula (XXIV),  (XXIV), reacts with halide via nucleophilic substitution or with ketone via reductive amination to provide final compound of formula (XXII);

wherein

in step a) and e) , the base is DIEA;

in step b), the catalyst is $Pd_2(dba)_3$;

in step c), the coupling reagent is HATU;

in step d), the regents is CDI, BrCN or $R^1(OCH_3)_3$;

in step e), the base is DIEA;

in step f), the base is DIPEA, K_2CO_3 or NaH;

X is halogen; R^{13} is R^a, R^b or R^c; M is $C_{1\text{-}6}$alkylene; R^e is a substituent bearing primary or secondary amino groups; R^1 to R^7, Q^1, Q^2, A^1 to A^7 are as described herein.

In another aspect, the present disclosure relates to a compound or pharmaceutically acceptable salt as described herein for use as therapeutically active substance.

In another aspect, the present disclosure relates to a pharmaceutical composition comprising a compound as described herein and a therapeutically inert carrier.

In another aspect, the present disclosure relates to the use of a compound as described herein in the manufacture of a medicament for the treatment or prophylaxis of autoimmune diseases, inflammatory diseases, neurological disorders diseases, metabolic diseases, cardiovascular diseases, ocular diseases, or selective types of cancers where overexpression or activation of STING is implicated.

In another aspect, the present disclosure relates to the use of a compound as described herein in the manufacture of a medicament for the treatment to subjects suffered from an interferonopathy or auto-inflammatory diseases in which the STING activation are the root-cause of disease pathologies.

In another aspect, the present disclosure relates to a compound as described herein for use in the inhibition of STING.

In another aspect, the present disclosure relates to the use of a compound as described herein in the preparation of a medicament for the inhibition of STING.

In another aspect, the present disclosure relates to a compound or pharmaceutically acceptable salt as described herein, when manufactured according to a process as described herein.

In another aspect, the present disclosure relates to use of a compound as described herein in the manufacture of a medicament for the treatment or prophylaxis of autoimmune diseases.

DETAILED DESCRIPTION OF THE INVENTION

DEFINITIONS

The term “deuterio” or “deuterium” denotes the isotope of hydrogen that has a single neutron as well as a proton in the nucleus.

The term "C_{1-6}alkyl" denotes a saturated, linear or branched chain alkyl group containing 1 to 6, particularly 1 to 4 carbon atoms, for example methyl, ethyl, *n*-propyl, isopropyl, *n*-butyl, isobutyl, *tert*-butyl and the like. Particular "C_{1-6}alkyl" groups are methyl, ethyl and *n*-propyl.

The term "C_{1-6}alkylene" denotes a linear or branched saturated divalent hydrocarbon group of 1 to 6 carbon atoms or a divalent branched saturated divalent hydrocarbon group of 3 to 6 carbon atoms. Examples of C_{1-6}alkylene groups include methylene, ethylene, propylene, 2-methylpropylene, butylene, 2-ethylbutylene, pentylene, hexylene.

The term "C_{1-6}alkoxy" denotes C_{1-6}alkyl-O-.

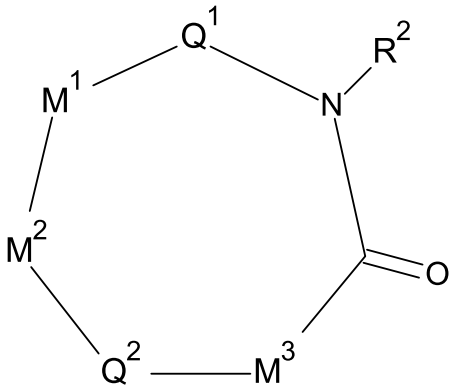
The term "trideuterioC_{1-6}alkyl" denotes a C_{1-6}alkyl group wherein three of the hydrogen atoms of the C_{1-6}alkyl group has been replaced by deuterium. Example of trideuterioC_{1-6}alkyl includes trideuteriomethyl.

The term "C_{2-6}alkenyl" denotes a monovalent linear or branched hydrocarbon group of 2 to 6 carbon atoms with at least one double bond. In particular embodiments, alkenyl has 2 to 4 carbon atoms with at least one double bond. Examples of C_{2-6}alkenyl include ethenyl (or vinyl), propenyl, prop-2-enyl, isopropenyl, n-butenyl, and iso-butenyl.

The term "C_{2-6}alkynyl" denotes a monovalent linear or branched hydrocarbon group of 2 to 6 carbon atoms with at least one triple bond. In particular embodiments, alkynyl has 2 to 4 carbon atoms with at least one triple bond. Examples of C_{2-6}alkynyl include ethynyl (–C≡CH), prop-1-ynyl (–C≡CCH_3), prop-2-ynyl (propargyl, –CH_2C≡CH), but-1-ynyl, but-2-ynyl, and but-3-ynyl.The term "halogen" and "halo" are used interchangeably herein and denote fluoro, chloro, bromo, or iodo.

CLAIMS

1. A compound of formula (Ia),



(Ia),

wherein

M^1 is heteroarylene optionally substituted by R^1; wherein

R^1 is deuterio, halogen, cyano, carboxy, R, RO, RS, RNH, $(R)_2N$, RCO, RSO_2, $RNHSO_2$, R_2NSO_2, RSO(NR), R_3Si, $ROC_{1-6}alkyl$, $RSC_{1-6}alkyl$, $RNHC_{1-6}alkyl$, $(R)_2NC_{1-6}alkyl$, $RSO_2C_{1-6}alkyl$, $RNHSO_2C_{1-6}alkyl$, $R_2NSO_2C_{1-6}alkyl$, $RSO(NR)C_{1-6}alkyl$ or $R_3SiC_{1-6}alkyl$; wherein R is H, R^a, R^b or R^c;

R^a is $C_{1-6}alkyl$ optionally substituted by R^b or R^c;

R^b is $C_{3-7}cycloalkyl$, heterocyclyl, heteroaryl or aryl, said $C_{3-7}cycloalkyl$, heterocyclyl, heteroaryl and aryl being unsubstituted or optionally substituted by deuterio, halogen or R^c;

R^c is selected from deuterio, $C_{1-6}alkyl$, $(C_{1-6}alkyl)_3Si$, $haloC_{1-6}alkyl$, $deuterioC_{1-6}alkyl$, $C_{1-6}alkoxy$, $haloC_{1-6}alkoxy$, $deuterioC_{1-6}alkoxy$, $C_{2-6}alkenyl$, $haloC_{2-6}alkenyl$, $deuterioC_{2-6}alkenyl$, $C_{2-6}alkynyl$, $haloC_{2-6}alkynyl$, $deuterioC_{2-6}alkynyl$, $C_{3-7}cycloalkyl$, $haloC_{3-7}cycloalkyl$, $deuterioC_{3-7}cycloalkyl$, amino, $C_{1-6}alkylamino$, $(C_{1-6}alkyl)_2amino$, $C_{1-6}alkylcarbonylamino$, $haloC_{1-6}alkylamino$, $haloC_{1-6}alkylcarbonylamino$, $(haloC_{1-6}alkyl)_2amino$, $C_{3-7}cycloalkylamino$, $C_{3-7}cycloalkylcarbonylamino$, $(C_{3-7}cycloalkyl)_2amino$, $haloC_{3-7}cycloalkylamino$, $(haloC_{3-7}cycloalkyl)_2amino$, $C_{3-7}cycloalkyl(C_{1-6}alkyl)amino$, $haloC_{3-7}cycloalkyl(C_{1-6}alkyl)amino$, $C_{1-6}alkylsulfonylamino$, $haloC_{1-6}alkylsulfonylamino$, $C_{1-6}alkoxyC_{1-6}alkyl$, $(haloC_{1-6}alkoxy)C_{1-6}alkyl$, $C_{1-6}alkoxy(halo)C_{1-6}alkyl$, $C_{3-7}cycloalkylC_{1-6}alkyl$, $C_{3-7}cycloalkylhaloC_{1-6}alkyl$, $C_{1-6}alkylcarbonyl$, $haloC_{1-6}alkylcarbonyl$, $C_{3-7}cycloalkylcarbonyl$, $haloC_{3-}$

$_7$cycloalkylcarbonyl, C_{1-6}alkoxycarbonyl, haloC_{1-6}alkoxycarbonyl, C_{3-7}cycloalkoxycarbonyl, haloC_{3-7}cycloalkoxycarbonyl, C_{1-6}alkylaminocarbonyl, $(C_{1-6}alkyl)_2$aminocarbonyl, C_{3-7}cycloalkylaminocarbonyl, $(C_{3-7}cycloalkyl)_2$aminocarbonyl, C_{1-6}alkylC_{3-7}cycloalkylaminocarbonyl, haloC_{1-6}alkylaminocarbonyl, halo$(C_{1-6}alkyl)_2$aminocarbonyl, haloC_{3-7}cycloalkylaminocarbonyl, halo$(C_{3-7}cycloalkyl)_2$aminocarbonyl, haloC_{1-6}alkylC_{3-7}cycloalkylaminocarbonyl, C_{1-6}alkylhaloC_{3-7}cycloalkylaminocarbonyl, $(C_{1-6}alkyl)_3$ammonio, C_{1-6}alkylsulfanyl, C_{1-6}alkylsulfinyl, C_{1-6}alkylsulfonyl, C_{3-7}cycloalkylsulfanyl, C_{3-7}cycloalkylsulfinyl, C_{3-7}cycloalkylsulfonyl, haloC_{1-6}alkylsulfanyl, haloC_{1-6}alkylsulfinyl, haloC_{1-6}alkylsulfonyl, haloC_{3-7}cycloalkylsulfanyl, haloC_{3-7}cycloalkylsulfinyl, haloC_{3-7}cycloalkylsulfonyl, haloC_{1-6}alkoxysulfonyl, C_{1-6}alkylaminosulfonyl, (haloC_{1-6}alkylamino)sulfonyl, $(C_{1-6}alkyl)_2$aminosulfonyl, $(haloC_{1-6}alkyl)_2$aminosulfonyl, $(C_{1-6}alkyl)_2$phosphoryl, $(C_{1-6}alkoxy)_2$phosphoryl, hydroxy(C_{1-6}alkoxy)phosphoryl, C_{1-6}alkoxyphosphoryl, hydroxy, hydroxyC_{1-6}alkyl, hydroxyhaloC_{1-6}alkyl, hydroxyC_{1-6}alkoxy, hydroxylhaloC_{1-6}alkoxy, hydroxyC_{3-7}cycloalkyl, hydroxyhaloC_{3-7}cycloalkyl and sulfonimidoyl optionally substituted by H, C_{1-6}alkyl, C_{1-6}alkoxy, aryl, heteroaryl and heterocyclyl;

M^2 is arylene, heterocyclylene or heteroarylene, said arylene, heterocyclylene or heteroarylene being optionally substituted by R^1;

M^3 is heterocyclylene which is unsubstituted or optionally substituted by R^3;

R^2 is deuterio, R, ROC_{1-6}alkyl, RSC_{1-6}alkyl, RSO_2C_{1-6}alkyl, RNHC_{1-6}alkyl or $(R)_2NC_{1-6}$alkyl;

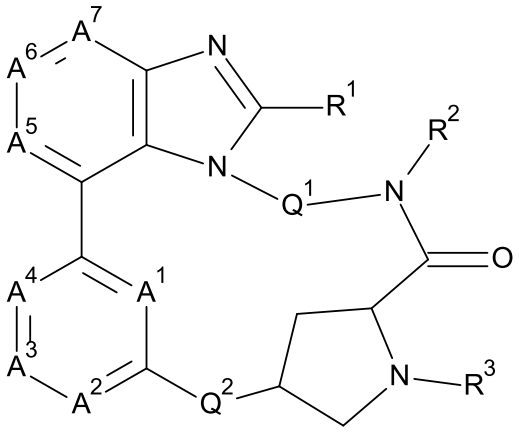
R^3 is heterocyclyl, heteroaryl, aryl, heterocyclylcarbonyl, heteroarylcarbonyl or arylcarbonyl, said heterocyclyl, heteroaryl, aryl, heterocyclylcarbonyl, heteroarylcarbonyl and arylcarbonyl being unsubstituted or optionally substituted by R^1;

Q^1 is -M-, -M-O-M-, -M-S-M-, -M-SO_2-M-, -M-SO(NH)-M-, -M-SO(NR)-M-, -M-SiR_2-M- or -M-N(R)-M-; wherein M is C_{1-6}alkylene unsubstituted or optionally substituted by the substituents independently selected from deuterio, halogen, cyano, carboxy, R, RO, RS, RNH, R_2N, RSO_2, ROC_{1-6}alkyl, RSC_{1-6}alkyl, RNHC_{1-6}alkyl, $(R)_2NC_{1-6}$alkyl and RSO_2C_{1-6}alkyl;

Q^2 is NH, O, S, SO, SO_2, P(O)R, PO(NR), Se, N(R), SiR_2, heterocyclyl, heteroaryl or aryl;

or a pharmaceutically acceptable salt thereof.

2. A compound of formula (I) according to claim 1,

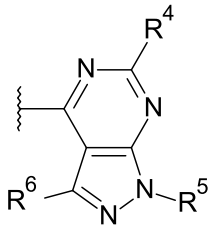
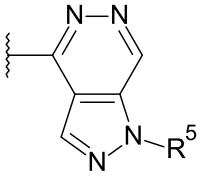
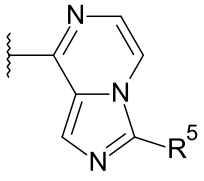
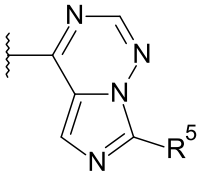
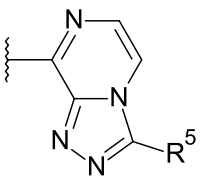
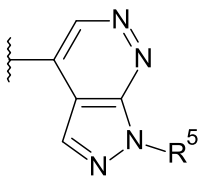
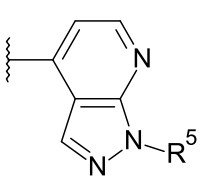
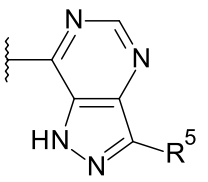


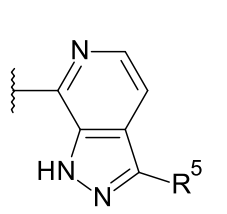
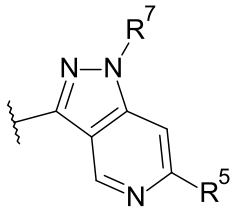
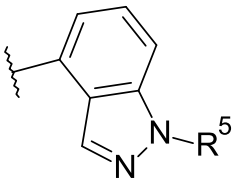
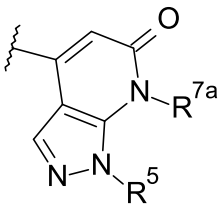
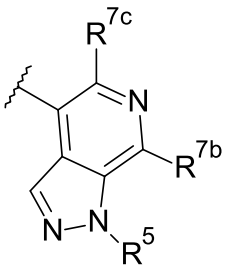
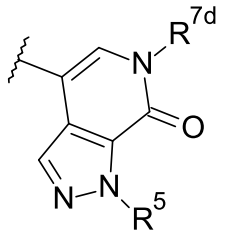
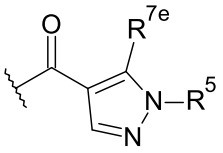
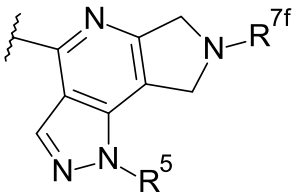
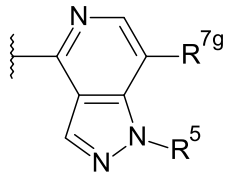
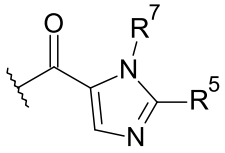
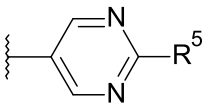
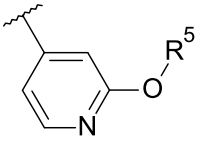
(I),

wherein

R^1 is deuterio, H, (2,3,4a,5,7,7a-hexahydro-[1,4]dioxino[2,3-c]pyrrolyl)C_{1-6}alkyl, (C_{1-6}alkylsulfonylpiperazinyl)C_{1-6}alkyl, amino, C_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkoxycarbonyl, C_{1-6}alkyl, C_{1-6}alkylaminocarbonyl, C_{1-6}alkylcarbonylamino, C_{3-7}cycloalkyl, haloC_{1-6}alkyl, hydroxyC_{1-6}alkyl, morpholinylC_{1-6}alkyl, hydroxy, pyrazolylC_{1-6}alkyl or triazolylC_{1-6}alkyl;

R^2 is H, (2-oxa-5-azabicyclo[2.2.1]heptanyl)C_{1-6}alkyl, (2-oxa-6-azaspiro[3.3]heptanyl)C_{1-6}alkyl, (2,3,4a,5,7,7a-hexahydro-[1,4]dioxino[2,3-c]pyrrolyl)C_{1-6}alkyl, ((C_{1-6}alkyl)$_3$ammonio)C_{1-6}alkyl, (C_{1-6}alkylsulfonylpiperazinyl)C_{1-6}alkyl, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkyl, C_{3-7}cycloalkyl, C_{3-7}cycloalkylC_{1-6}alkyl, haloC_{1-6}alkyl, hydroxyC_{1-6}alkyl, morpholinylC_{1-6}alkyl, phenylC_{1-6}alkyl, pyrimidinyloxyC_{1-6}alkyl, tetrazolylC_{1-6}alkyl or trideuterioC_{1-6}alkyl;

R^3 is , , , , , , , ,

, , , , , , , , , , , or ;

wherein

R^4 is H,

$(C_{1\text{-}6}alkyl)_2amino$,

$(C_{1\text{-}6}alkyl)_2aminocarbonyl$,

$(C_{1\text{-}6}alkyl)_2phosphoryl$,

1,1-dioxothianyl,

1,3,3a,4,6,6a-hexahydropyrrolo[3,4-c]pyrrolyl substituted by $C_{1\text{-}6}alkyl$,

1,6-diazaspiro[3.3]heptanyl substituted by halo$C_{1\text{-}6}alkyl$ or oxetanyl,

1,7-diazaspiro[3.5]nonanyl substituted by halo$C_{1\text{-}6}alkyl$ or oxetanyl,

2,2-dioxo-2λ⁶-thia-6-azaspiro[3.3]heptanyl,

2,3,4a,5,7,7a-hexahydropyrrolo[3,4-b][1,4]oxazinyl substituted by $C_{1\text{-}6}alkyl$,

2,5-diazabicyclo[2.2.1]heptanyl substituted by $C_{1\text{-}6}alkyl$, halo$C_{1\text{-}6}alkyl$ or oxetanyl,

2-azaspiro[3.3]heptanyl once or twice substituted by substituents independently selected from halo$C_{1\text{-}6}alkyl$ and hydroxy,

2-oxa-5-azabicyclo[2.2.1]heptanyl,

2-oxa-6-azaspiro[3.3]heptanyl,

3,4,6,7,8,8a-hexahydro-1H-pyrrolo[1,2-a]pyrazinyl,

3,4,6,7,9,9a-hexahydro-1H-pyrazino[2,1-c][1,4]oxazinyl,

3,6-diazabicyclo[3.1.1]heptanyl substituted by (halooxetanyl)C_{1-6}alkyl, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkyl, C_{3-7}cycloalkyl, C_{3-7}cycloalkylcarbonyl, cyanoC_{1-6}alkyl, haloC_{1-6}alkyl, haloC_{1-6}alkylcarbonyl, haloC_{3-7}cycloalkyl, hydroxyC_{1-6}alkyl, hydroxyC_{1-6}alkylcarbonyl, hydroxyC_{3-7}cycloalkylcarbonyl or oxetanyl,

3,6-diazabicyclo[3.2.0]heptanyl substituted by oxetanyl,

3,7-diazabicyclo[4.2.0]octanyl substituted by oxetanyl,

3,8-diazabicyclo[3.2.1]octanyl substituted by haloC_{1-6}alkyl,

3-oxa-6-azabicyclo[3.1.1]heptanyl,

3-oxa-7,9-diazabicyclo[3.3.1]nonanyl substituted by C_{1-6}alkyl,

3-oxa-8-azabicyclo[3.2.1]octanyl,

4,7-diazaspiro[2.5]octanyl unsubstituted or substituted by C_{1-6}alkyl,

4-oxo-6,7,9,9a-tetrahydro-1H-pyrazino[2,1-c][1,4]oxazinyl,

5-oxa-2,8-diazaspiro[3.5]nonanyl substituted by haloC_{1-6}alkyl or C_{1-6}alkoxyC_{1-6}alkyl,

6,6-dioxo-6λ^6-thia-1-azaspiro[3.3]heptanyl,

6-oxa-1-azaspiro[3.3]heptanyl,

6-oxa-3-azabicyclo[3.1.1]heptanyl,

8-oxa-2,5-diazaspiro[3.5]nonanyl substituted by haloC_{1-6}alkyl,

8-oxa-3-azabicyclo[3.2.1]octanyl,

amino,

aminocarbonyl,

azetidinyl unsubstituted or once or twice substituted by substituents independently selected from (C_{1-6}alkyl)$_2$amino, (C_{1-6}alkyl)$_2$aminoC_{1-6}alkyl, amino, C_{1-6}alkyl, C_{1-6}alkylcarbonyl, C_{1-6}alkylsulfonyl, C_{1-6}alkylsulfonylC_{1-6}alkyl, C_{3-7}cycloalkyl, cyano, cyanoC_{1-6}alkyl, haloC_{1-6}alkyl, hydroxyC_{1-6}alkyl, morpholinyl and morpholinylC_{1-6}alkyl,

azetidinyloxy substituted by haloC_{1-6}alkyl or C_{3-7}cycloalkyl,

C_{1-6}alkoxy unsubstituted or substituted by (haloC_{1-6}alkylcarbonyl)azetidinyl, C_{1-6}alkylazetidinyl, C_{1-6}alkylpiperazinyl, C_{1-6}alkylpyrrolidinyl, halo-1,2,3,5,6,7-hexahydropyrrolizinyl, haloC_{1-6}alkylazetidinyl, morpholinyl, oxetanyl or tetrahydrofuranyl,

C_{1-6}alkoxyC_{1-6}alkylamino,

C_{1-6}alkoxycarbonyl,

C_{1-6}alkyl unsubstituted or once or twice substituted by substituents independently selected from (C_{1-6}alkoxyC_{1-6}alkylcarbonyl)azetidinyl, (haloC_{1-6}alkylcarbonyl)pyrrolidinyl, 1,1-dioxothietanyl, 2-oxa-6-azaspiro[3.3]heptanyl, 3-oxa-6-azabicyclo[3.1.1]heptanyl, 4,7-diazaspiro[2.5]octanyl, C_{1-6}alkoxy, C_{1-6}alkylcarbonylazetidinyl, C_{1-6}alkylcarbonyloxy, C_{1-6}alkylcarbonylpyrrolidinyl, C_{1-6}alkylsulfonylazetidinyl, C_{1-6}alkylsulfonylpyrrolidinyl, C_{3-7}cycloalkylsulfonylazetidinyl, cyano, haloC_{1-6}alkylazetidinyl, hydroxy, hydroxyC_{1-6}alkyl, morpholinyl and oxetanyl,

C_{1-6}alkylamino,

C_{1-6}alkylsulfanyl unsubstituted or substituted by 1,1-dioxothietanyl,

C_{1-6}alkylsulfonyl,

C_{3-7}cycloalkyl unsubstituted or substituted by C_{1-6}alkoxy or hydroxy,

carboxy,

cyano,

haloC_{1-6}alkoxy,

haloC_{1-6}alkyl substituted by hydroxy,

hydroxy,

hydroxyC_{1-6}alkylcarbamoyl,

imidazolyl,

morpholinyl,

morpholinylcarbonyl,

oxetanyl,

oxomorpholinyl,

oxooxazolidinyl,

oxopiperazinyl substituted by C_{1-6}alkyl,

oxopyrrolidinyl unsubstituted or substituted by C_{1-6}alkyl or hydroxy,

piperazinyl once or twice substituted by substituents independently selected from C_{1-6}alkyl, carboxy, oxetanyl and phenylC_{1-6}alkyl,

piperidinyl substituted by C_{1-6}alkyl, 2H-tetrazolyl or oxazolyl,

pyrrolidinyl once or twice substituted by substituents independently selected from amino, C_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkylcarbonyl, C_{1-6}alkylsulfonyl, cyano, haloC_{1-6}alkylcarbonyl and halogen,

pyrrolidinylcarbonyl,

tetrahydrofuranyl, or

tetrahydropyranyl;

R^5 is 1,3-dihydroisobenzofuranyl,

1H-indazolyl substituted by halogen,

C_{2-6}alkynyl,

thienyl,

bicyclo[1.1.1]pentanyl unsubstituted or substituted by halogen or C_{1-6}alkyl,

phenyl substituted once, twice or three times by the substituents independently selected from (C_{1-6}alkoxyC_{1-6}alkyl)C_{3-7}cycloalkyloxy, (C_{1-6}alkyl)$_2$phosphoryl, (C_{1-6}alkylmorpholinyl)C_{1-6}alkoxy, (C_{1-6}alkylsulfonylazetidinyl)C_{1-6}alkoxy, 2-oxa-6-azaspiro[3.3]heptanyl, azetidinyl, C_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkoxyC_{2-6}alkynyl, C_{1-6}alkoxyC_{3-7}cycloalkyloxy, C_{1-6}alkoxypyrrolidinyl, C_{1-6}alkyl, C_{1-6}alkylamino, C_{1-6}alkylsulfanyl, C_{1-6}alkylsulfinyl, C_{1-6}alkylsulfonyl, C_{2-6}alkenyl, C_{3-7}cycloalkyl, C_{3-7}cycloalkyloxy, carboxy, cyano, cyanoC_{1-6}alkoxy, haloC_{1-6}alkoxy, haloC_{1-6}alkyl, halogen, hydroxy, morpholinyl, morpholinylC_{1-6}alkoxy, oxetanylC_{1-6}alkoxy, oxetanyloxy, oxopyrrolidinyl, phenyl, tetrahydrofuranyl, tetrahydrofuranyloxy and trideuterioC_{1-6}alkoxy,

pyridinyl unsubstituted or substituted once or twice by the substituents independently selected from halogen, hydroxy, C_{1-6}alkylsulfanyl and C_{1-6}alkoxy, or

pyrimidinyl unsubstituted or substituted by halogen;

R^6 is H or C_{1-6}alkyl;

R^7 is C_{1-6}alkyl;

R^{7a} is C_{1-6}alkyl, C_{3-7}cycloalkyl or C_{1-6}alkoxyC_{1-6}alkyl;

R^{7b} is H, C_{1-6}alkoxy or phenylC_{1-6}alkoxy;

R^{7c} is H or C_{1-6}alkoxy;

R^{7d} is C_{1-6}alkyl, C_{3-7}cycloalkylC_{1-6}alkyl or C_{1-6}alkoxyC_{1-6}alkyl;

R^{7e} is H, amino, phenylC_{1-6}alkylamino or morpholinylC_{1-6}alkylamino;

R^{7f} is C_{1-6}alkyl, phenylC_{1-6}alkyl or oxetanyl;

R^{7g} is H or C_{1-6}alkoxyC_{1-6}alkyl;

Q^1 is C_{1-6}alkylene unsubstituted or substituted once or twice by the substituents independently selected from C_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkoxy, C_{1-6}alkyl, C_{1-}

$_{6}$alkylcarbonyloxy, carboxy, hydroxy, hydroxyC_{1-6}alkoxy, hydroxyC_{1-6}alkyl, morpholinylC_{1-6}alkoxy and trideuterioC_{1-6}alkoxy, or

-M-O-M-, wherein M is C_{1-6}alkylene;

Q^2 is NH, O or S;

A^1 is CH or N;

A^2 is CH, CR^{7h} or N; wherein R^{7h} is C_{1-6}alkyl;

A^3 is CR^8 or N, wherein R^8 is H, C_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkyl, C_{3-7}cycloalkoxy, C_{3-7}cycloalkoxyamino, cyano, haloazetidinylC_{1-6}alkyl, haloC_{1-6}alkoxy, halogen, morpholinylC_{1-6}alkyl or oxetanyl;

A^4 is CR^9 or N, wherein R^9 is H; or R^8 and R^9 together with the atoms that they are attached to form a heterocyclyl;

A^5 is CR^{10} or N, wherein R^{10} is H or halogen;

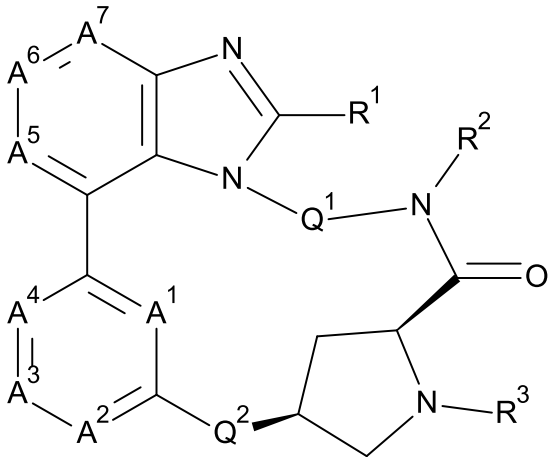
A^6 is CR^{11} or N, wherein R^{11} is H, deuterio, halogen, C_{1-6}alkyl or C_{1-6}alkoxy;

A^7 is CR^{12} or N, wherein R^{12} is H or halogen;

provided that no more than two of A^1, A^2, A^3 and A^4 are N simultaneously; and no more than one of A^5, A^6 and A^7 is N;

or a pharmaceutically acceptable salt thereof.

3. A compound of formula (Ib) according to claim 1,



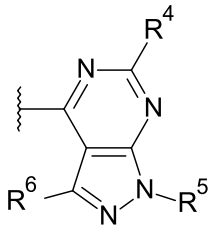
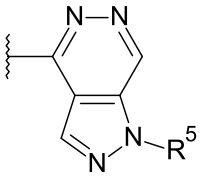
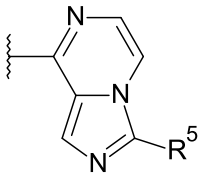
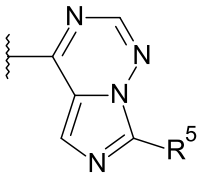
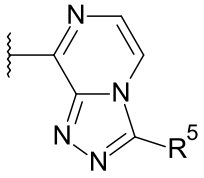
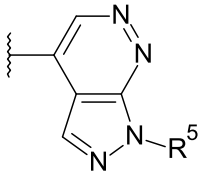
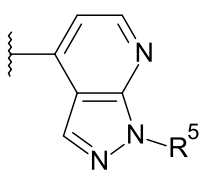
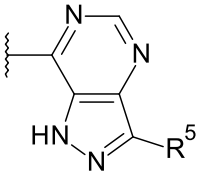
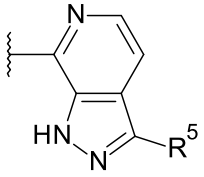
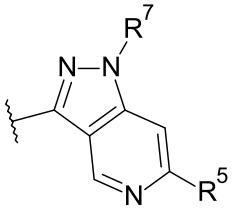
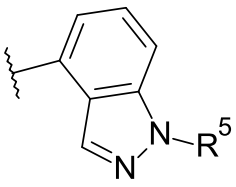
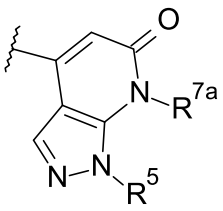
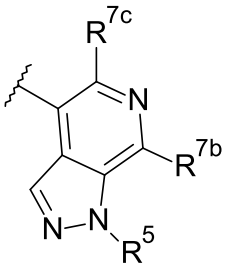
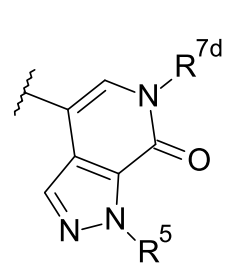
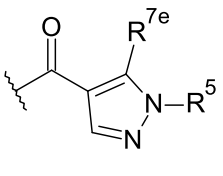
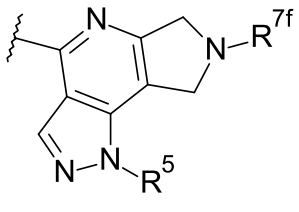
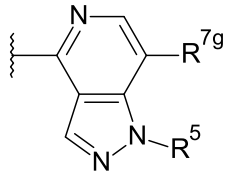
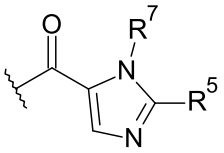
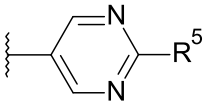
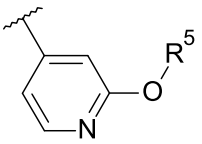
(Ib)

wherein

R^1 is deuterio, H, (2,3,4a,5,7,7a-hexahydro-[1,4]dioxino[2,3-c]pyrrolyl)C_{1-6}alkyl, (C_{1-6}alkylsulfonylpiperazinyl)C_{1-6}alkyl, amino, C_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkoxycarbonyl, C_{1-6}alkyl, C_{1-6}alkylaminocarbonyl, C_{1-6}alkylcarbonylamino, C_{3-}

$_7$cycloalkyl, haloC_{1-6}alkyl, hydroxyC_{1-6}alkyl, morpholinylC_{1-6}alkyl, hydroxy, pyrazolylC_{1-6}alkyl or triazolylC_{1-6}alkyl;

R^2 is H, (2-oxa-5-azabicyclo[2.2.1]heptanyl)C_{1-6}alkyl, (2-oxa-6-azaspiro[3.3]heptanyl)C_{1-6}alkyl, (2,3,4a,5,7,7a-hexahydro-[1,4]dioxino[2,3-c]pyrrolyl)C_{1-6}alkyl, ((C_{1-6}alkyl)$_3$ammonio)C_{1-6}alkyl, (C_{1-6}alkylsulfonylpiperazinyl)C_{1-6}alkyl, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkyl, C_{3-7}cycloalkyl, C_{3-7}cycloalkylC_{1-6}alkyl, haloC_{1-6}alkyl, hydroxyC_{1-6}alkyl, morpholinylC_{1-6}alkyl, phenylC_{1-6}alkyl, pyrimidinyloxyC_{1-6}alkyl, tetrazolylC_{1-6}alkyl or trideuterioC_{1-6}alkyl;

R^3 is , , , , , , , , , , , , , , , , , , , ;

wherein

R^4 is H,

$(C_{1-6}alkyl)_2amino$,
$(C_{1-6}alkyl)_2aminocarbonyl$,
$(C_{1-6}alkyl)_2phosphoryl$,
1,1-dioxothianyl,
1,3,3a,4,6,6a-hexahydropyrrolo[3,4-c]pyrrolyl substituted by $C_{1-6}alkyl$,
1,6-diazaspiro[3.3]heptanyl substituted by halo$C_{1-6}alkyl$ or oxetanyl,
1,7-diazaspiro[3.5]nonanyl substituted by halo$C_{1-6}alkyl$ or oxetanyl,
2,2-dioxo-2λ⁶-thia-6-azaspiro[3.3]heptanyl,
2,3,4a,5,7,7a-hexahydropyrrolo[3,4-b][1,4]oxazinyl substituted by $C_{1-6}alkyl$,
2,5-diazabicyclo[2.2.1]heptanyl substituted by $C_{1-6}alkyl$, halo$C_{1-6}alkyl$ or oxetanyl,
2-azaspiro[3.3]heptanyl once or twice substituted by substituents independently selected from halo$C_{1-6}alkyl$ and hydroxy,
2-oxa-5-azabicyclo[2.2.1]heptanyl,
2-oxa-6-azaspiro[3.3]heptanyl,
3,4,6,7,8,8a-hexahydro-1H-pyrrolo[1,2-a]pyrazinyl,
3,4,6,7,9,9a-hexahydro-1H-pyrazino[2,1-c][1,4]oxazinyl,
3,6-diazabicyclo[3.1.1]heptanyl substituted by (halooxetanyl)$C_{1-6}alkyl$, $C_{1-6}alkoxyC_{1-6}alkyl$, $C_{1-6}alkyl$, $C_{3-7}cycloalkyl$, $C_{3-7}cycloalkylcarbonyl$, cyano$C_{1-6}alkyl$, halo$C_{1-6}alkyl$, halo$C_{1-6}alkylcarbonyl$, halo$C_{3-7}cycloalkyl$, hydroxy$C_{1-6}alkyl$, hydroxy$C_{1-6}alkylcarbonyl$, hydroxy$C_{3-7}cycloalkylcarbonyl$ or oxetanyl,
3,6-diazabicyclo[3.2.0]heptanyl substituted by oxetanyl,
3,7-diazabicyclo[4.2.0]octanyl substituted by oxetanyl,
3,8-diazabicyclo[3.2.1]octanyl substituted by halo$C_{1-6}alkyl$,
3-oxa-6-azabicyclo[3.1.1]heptanyl,
3-oxa-7,9-diazabicyclo[3.3.1]nonanyl substituted by $C_{1-6}alkyl$,
3-oxa-8-azabicyclo[3.2.1]octanyl,
4,7-diazaspiro[2.5]octanyl unsubstituted or substituted by $C_{1-6}alkyl$,
4-oxo-6,7,9,9a-tetrahydro-1H-pyrazino[2,1-c][1,4]oxazinyl,
5-oxa-2,8-diazaspiro[3.5]nonanyl substituted by halo$C_{1-6}alkyl$ or $C_{1-6}alkoxyC_{1-6}alkyl$,
6,6-dioxo-6λ⁶-thia-1-azaspiro[3.3]heptanyl,
6-oxa-1-azaspiro[3.3]heptanyl,
6-oxa-3-azabicyclo[3.1.1]heptanyl,
8-oxa-2,5-diazaspiro[3.5]nonanyl substituted by halo$C_{1-6}alkyl$,

8-oxa-3-azabicyclo[3.2.1]octanyl,

amino,

aminocarbonyl,

azetidinyl unsubstituted or once or twice substituted by substituents independently selected from $(C_{1-6}alkyl)_2amino$, $(C_{1-6}alkyl)_2aminoC_{1-6}alkyl$, amino, $C_{1-6}alkyl$, $C_{1-6}alkylcarbonyl$, $C_{1-6}alkylsulfonyl$, $C_{1-6}alkylsulfonylC_{1-6}alkyl$, $C_{3-7}cycloalkyl$, cyano, $cyanoC_{1-6}alkyl$, $haloC_{1-6}alkyl$, $hydroxyC_{1-6}alkyl$, morpholinyl and $morpholinylC_{1-6}alkyl$,

azetidinyloxy substituted by $haloC_{1-6}alkyl$ or $C_{3-7}cycloalkyl$,

$C_{1-6}alkoxy$ unsubstituted or substituted by $(haloC_{1-6}alkylcarbonyl)azetidinyl$, $C_{1-6}alkylazetidinyl$, $C_{1-6}alkylpiperazinyl$, $C_{1-6}alkylpyrrolidinyl$, halo-1,2,3,5,6,7-hexahydropyrrolizinyl, $haloC_{1-6}alkylazetidinyl$, morpholinyl, oxetanyl or tetrahydrofuranyl,

$C_{1-6}alkoxyC_{1-6}alkylamino$,

$C_{1-6}alkoxycarbonyl$,

$C_{1-6}alkyl$ unsubstituted or once or twice substituted by substituents independently selected from $(C_{1-6}alkoxyC_{1-6}alkylcarbonyl)azetidinyl$, $(haloC_{1-6}alkylcarbonyl)pyrrolidinyl$, 1,1-dioxothietanyl, 2-oxa-6-azaspiro[3.3]heptanyl, 3-oxa-6-azabicyclo[3.1.1]heptanyl, 4,7-diazaspiro[2.5]octanyl, $C_{1-6}alkoxy$, $C_{1-6}alkylcarbonylazetidinyl$, $C_{1-6}alkylcarbonyloxy$, $C_{1-6}alkylcarbonylpyrrolidinyl$, $C_{1-6}alkylsulfonylazetidinyl$, $C_{1-6}alkylsulfonylpyrrolidinyl$, $C_{3-7}cycloalkylsulfonylazetidinyl$, cyano, $haloC_{1-6}alkylazetidinyl$, hydroxy, $hydroxyC_{1-6}alkyl$, morpholinyl and oxetanyl,

$C_{1-6}alkylamino$,

$C_{1-6}alkylsulfanyl$ unsubstituted or substituted by 1,1-dioxothietanyl,

$C_{1-6}alkylsulfonyl$,

$C_{3-7}cycloalkyl$ unsubstituted or substituted by $C_{1-6}alkoxy$ or hydroxy,

carboxy,

cyano,

$haloC_{1-6}alkoxy$,

$haloC_{1-6}alkyl$ substituted by hydroxy,

hydroxy,

$hydroxyC_{1-6}alkylcarbamoyl$,

imidazolyl,
morpholinyl,
morpholinylcarbonyl,
oxetanyl,
oxomorpholinyl,
oxooxazolidinyl,
oxopiperazinyl substituted by C_{1-6}alkyl,
oxopyrrolidinyl unsubstituted or substituted by C_{1-6}alkyl or hydroxy,
piperazinyl once or twice substituted by substituents independently selected from C_{1-6}alkyl, carboxy, oxetanyl and phenylC_{1-6}alkyl,
piperidinyl substituted by C_{1-6}alkyl, 2H-tetrazolyl or oxazolyl,
pyrrolidinyl once or twice substituted by substituents independently selected from amino, C_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkylcarbonyl, C_{1-6}alkylsulfonyl, cyano, haloC_{1-6}alkylcarbonyl and halogen,
pyrrolidinylcarbonyl,
tetrahydrofuranyl, or
tetrahydropyranyl;

R^5 is 1,3-dihydroisobenzofuranyl,
1H-indazolyl substituted by halogen,
C_{2-6}alkynyl,
thienyl,
bicyclo[1.1.1]pentanyl unsubstituted or substituted by halogen or C_{1-6}alkyl,
phenyl substituted once, twice or three times by the substituents independently selected from (C_{1-6}alkoxyC_{1-6}alkyl)C_{3-7}cycloalkyloxy, (C_{1-6}alkyl)$_2$phosphoryl, (C_{1-6}alkylmorpholinyl)C_{1-6}alkoxy, (C_{1-6}alkylsulfonylazetidinyl)C_{1-6}alkoxy, 2-oxa-6-azaspiro[3.3]heptanyl, azetidinyl, C_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkoxyC_{2-6}alkynyl, C_{1-6}alkoxyC_{3-7}cycloalkyloxy, C_{1-6}alkoxypyrrolidinyl, C_{1-6}alkyl, C_{1-6}alkylamino, C_{1-6}alkylsulfanyl, C_{1-6}alkylsulfinyl, C_{1-6}alkylsulfonyl, C_{2-6}alkenyl, C_{3-7}cycloalkyl, C_{3-7}cycloalkyloxy, carboxy, cyano, cyanoC_{1-6}alkoxy, haloC_{1-6}alkoxy, haloC_{1-6}alkyl, halogen, hydroxy, morpholinyl, morpholinylC_{1-6}alkoxy, oxetanylC_{1-6}alkoxy, oxetanyloxy, oxopyrrolidinyl, phenyl, tetrahydrofuranyl, tetrahydrofuranyloxy and trideuterioC_{1-6}alkoxy,

pyridinyl unsubstituted or substituted once or twice by the substituents independently selected from halogen, hydroxy, C_{1-6}alkylsulfanyl and C_{1-6}alkoxy, or

pyrimidinyl unsubstituted or substituted by halogen;

R^6 is H or C_{1-6}alkyl;

R^7 is C_{1-6}alkyl;

R^{7a} is C_{1-6}alkyl, C_{3-7}cycloalkyl or C_{1-6}alkoxyC_{1-6}alkyl;

R^{7b} is H, C_{1-6}alkoxy or phenylC_{1-6}alkoxy;

R^{7c} is H or C_{1-6}alkoxy;

R^{7d} is C_{1-6}alkyl, C_{3-7}cycloalkylC_{1-6}alkyl or C_{1-6}alkoxyC_{1-6}alkyl;

R^{7e} is H, amino, phenylC_{1-6}alkylamino or morpholinylC_{1-6}alkylamino;

R^{7f} is C_{1-6}alkyl, phenylC_{1-6}alkyl or oxetanyl;

R^{7g} is H or C_{1-6}alkoxyC_{1-6}alkyl;

Q^1 is C_{1-6}alkylene unsubstituted or substituted once or twice by the substituents independently selected from C_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkoxy, C_{1-6}alkyl, C_{1-6}alkylcarbonyloxy, carboxy, hydroxy, hydroxyC_{1-6}alkoxy, hydroxyC_{1-6}alkyl, morpholinylC_{1-6}alkoxy and trideuterioC_{1-6}alkoxy, or

-M-O-M-, wherein M is C_{1-6}alkylene;

Q^2 is NH, O or S;

A^1 is CH or N;

A^2 is CH, CR^{7h} or N; wherein R^{7h} is C_{1-6}alkyl;

A^3 is CR^8 or N, wherein R^8 is H, C_{1-6}alkoxy, C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkyl, C_{3-7}cycloalkoxy, C_{3-7}cycloalkoxyamino, cyano, haloazetidinylC_{1-6}alkyl, haloC_{1-6}alkoxy, halogen, morpholinylC_{1-6}alkyl or oxetanyl;

A^4 is CR^9 or N, wherein R^9 is H; or R^8 and R^9 together with the atoms that they are attached to form a heterocyclyl;

A^5 is CR^{10} or N, wherein R^{10} is H or halogen;

A^6 is CR^{11} or N, wherein R^{11} is H, deuterio, halogen, C_{1-6}alkyl or C_{1-6}alkoxy;

A^7 is CR^{12} or N, wherein R^{12} is H or halogen;

provided that no more than two of A^1, A^2, A^3 and A^4 are N simultaneously; and no more than one of A^5, A^6 and A^7 is N;

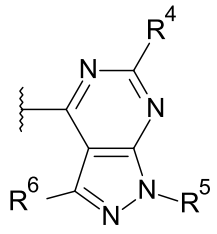
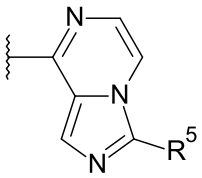
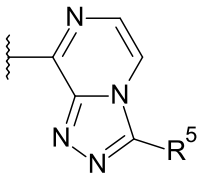
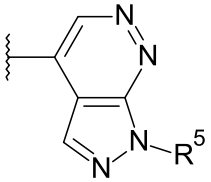
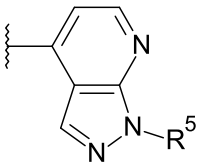
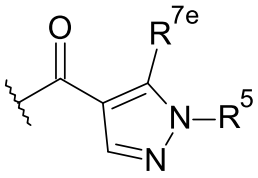
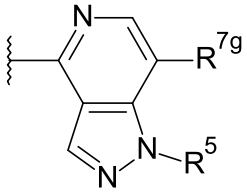
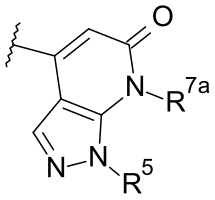
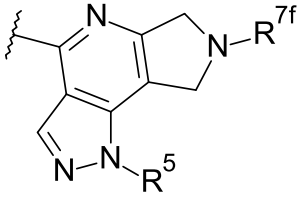
or a pharmaceutically acceptable salt thereof.

4. A compound according to claim 2 or 3, wherein R^1 is H or C_{1-6}alkyl.

5. A compound according to any one of claims 1-4, wherein R^1 is H or methyl.

6. A compound according to any one of claims 1-5, wherein R^2 is C_{1-6}alkyl or trideuterioC_{1-6}alkyl.

7. A compound according to any one of claims 1-6, wherein R^2 is methyl, ethyl or trideuteriomethyl.

8. A compound according to any one of claims 1-7, wherein R^3 is , , , , , , ,  or ;

wherein

R^4 is H,

2,5-diazabicyclo[2.2.1]heptanyl substituted by haloC_{1-6}alkyl or oxetanyl,

3,6-diazabicyclo[3.1.1]heptanyl substituted by C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkyl, C_{3-7}cycloalkylcarbonyl, haloC_{1-6}alkyl, haloC_{1-6}alkylcarbonyl, hydroxyC_{1-6}alkyl, hydroxyC_{1-6}alkylcarbonyl, hydroxyC_{3-7}cycloalkylcarbonyl or oxetanyl,

3-oxa-6-azabicyclo[3.1.1]heptanyl,

3-oxa-7,9-diazabicyclo[3.3.1]nonanyl substituted by C_{1-6}alkyl,

amino,

azetidinyl substituted by C_{1-6}alkylcarbonyl, C_{1-6}alkylsulfonyl or cyano,

C_{1-6}alkoxy,

C_{1-6}alkyl unsubstituted or substituted by C_{1-6}alkoxy, C_{1-6}alkylcarbonylazetidinyl, C_{1-6}alkylsulfonylazetidinyl, C_{3-7}cycloalkylsulfonylazetidinyl, haloC_{1-6}alkylazetidinyl, hydroxy, morpholinyl or oxetanyl,

oxetanyl,

oxooxazolidinyl, or

oxopyrrolidinyl unsubstituted or substituted by C_{1-6}alkyl or hydroxy;

R^5 is phenyl substituted once, twice or three times by the substituents independently selected from C_{1-6}alkoxy, C_{3-7}cycloalkyloxy, cyano, haloC_{1-6}alkoxy, halogen, hydroxy and trideuterioC_{1-6}alkoxy,

pyridinyl twice substituted by halogen, or

pyrimidinyl substituted by halogen;

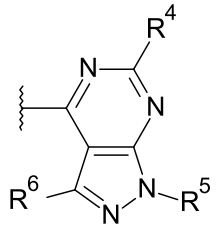
R^6 is H;

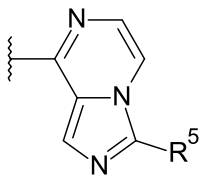
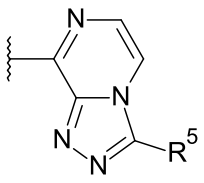
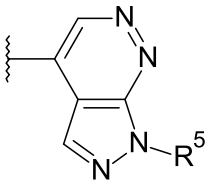
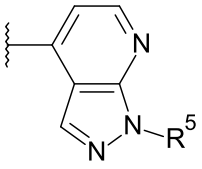
R^{7a} is C_{1-6}alkyl;

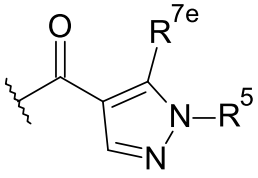
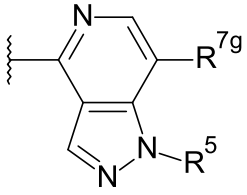
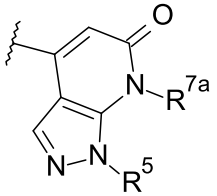
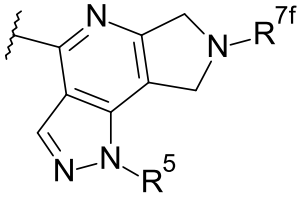
R^{7e} is amino;

R^{7f} is C_{1-6}alkyl;

R^{7g} is H or C_{1-6}alkoxyC_{1-6}alkyl.

9. A compound according to any one of claims 1-8, wherein R^3 is ,

, , , ,

, ,  or ;

wherein

R^4 is H,

R^4 is H, (1-acetylazetidin-3-yl)methyl, (1-cyclopropylsulfonylazetidin-3-yl)methyl, (1-ethylsulfonylazetidin-3-yl)methyl, (1-methylsulfonylazetidin-3-yl)methyl, [1-(2,2,2-trifluoroethyl)azetidin-3-yl]methyl, 1-acetylazetidin-3-yl, 1-methylsulfonylazetidin-3-yl, 2-oxooxazolidin-3-yl, 2-oxopyrrolidin-1-yl, 3-(1-hydroxycyclobutanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(1-hydroxycyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2,2-trifluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2-difluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-hydroxy-2-methyl-propanoyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-hydroxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(cyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-cyanoazetidin-1-yl, 3-hydroxy-2-oxo-pyrrolidin-1-yl, 3-methyl-2-oxo-pyrrolidin-1-yl, 3-methyl-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-oxa-6-azabicyclo[3.1.1]heptan-6-yl, 5-(2,2-difluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 5-(2-fluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 9-methyl-3-oxa-7,9-diazabicyclo[3.3.1]nonan-7-yl, amino, azetidin-1-yl, ethoxy, hydroxymethyl, methoxy, methoxymethyl, methyl, morpholinylmethyl, oxetan-2-ylmethyl or oxetan-3-yl;

R^5 is phenyl substituted once, twice or three times by the substituents independently selected from ethoxy, methoxy, cyclopropoxy, cyano, difluoromethoxy, fluoro, hydroxy and trideuteriomethoxy,

pyridinyl twice substituted by fluoro, or

pyrimidinyl substituted by fluoro;

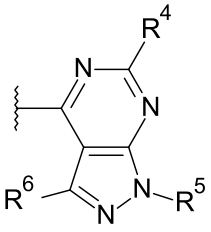
R^6 is H;

R^{7a} is methyl;

R^{7e} is amino;

R^{7f} is methyl;

R^{7g} is H or methoxymethyl.

10. A compound according to any one of claims 1-8, wherein R^3 is  ; wherein R^4 is H,

2,5-diazabicyclo[2.2.1]heptanyl substituted by haloC_{1-6}alkyl or oxetanyl,

3,6-diazabicyclo[3.1.1]heptanyl substituted by C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkyl, C_{3-7}cycloalkylcarbonyl, haloC_{1-6}alkyl, haloC_{1-6}alkylcarbonyl, hydroxyC_{1-6}alkyl, hydroxyC_{1-6}alkylcarbonyl, hydroxyC_{3-7}cycloalkylcarbonyl or oxetanyl,

3-oxa-6-azabicyclo[3.1.1]heptanyl,

3-oxa-7,9-diazabicyclo[3.3.1]nonanyl substituted by C_{1-6}alkyl,

amino,

azetidinyl substituted by C_{1-6}alkylcarbonyl, C_{1-6}alkylsulfonyl or cyano,

C_{1-6}alkoxy,

C_{1-6}alkyl unsubstituted or substituted by C_{1-6}alkoxy, C_{1-6}alkylcarbonylazetidinyl, C_{1-6}alkylsulfonylazetidinyl, C_{3-7}cycloalkylsulfonylazetidinyl, haloC_{1-6}alkylazetidinyl, hydroxy, morpholinyl or oxetanyl,

oxetanyl,

oxooxazolidinyl, or

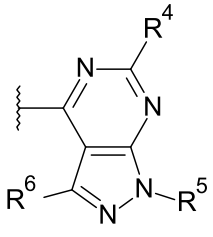
oxopyrrolidinyl unsubstituted or substituted by C_{1-6}alkyl or hydroxy;

R^5 is phenyl substituted once, twice or three times by the substituents independently selected from C_{1-6}alkoxy, C_{3-7}cycloalkyloxy, cyano, haloC_{1-6}alkoxy, halogen, hydroxy and trideuterioC_{1-6}alkoxy,

pyridinyl twice substituted by halogen, or

pyrimidinyl substituted by halogen;

R^6 is H.

11. A compound according to any one of claims 1-10, wherein R^3 is  ; wherein

R^4 is H, (1-acetylazetidin-3-yl)methyl, (1-cyclopropylsulfonylazetidin-3-yl)methyl, (1-ethylsulfonylazetidin-3-yl)methyl, (1-methylsulfonylazetidin-3-yl)methyl, [1-(2,2,2-trifluoroethyl)azetidin-3-yl]methyl, 1-acetylazetidin-3-yl, 1-methylsulfonylazetidin-3-yl, 2-oxooxazolidin-3-yl, 2-oxopyrrolidin-1-yl, 3-(1-hydroxycyclobutanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(1-hydroxycyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2,2-trifluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2-difluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-hydroxy-2-methyl-propanoyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-hydroxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(cyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-cyanoazetidin-1-yl, 3-hydroxy-2-oxo-pyrrolidin-1-yl, 3-methyl-2-oxo-pyrrolidin-1-yl, 3-methyl-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-oxa-6-azabicyclo[3.1.1]heptan-6-yl, 5-(2,2-difluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 5-(2-fluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 9-methyl-3-oxa-7,9-diazabicyclo[3.3.1]nonan-7-yl, amino, azetidin-1-yl, ethoxy, hydroxymethyl, methoxy, methoxymethyl, methyl, morpholinylmethyl, oxetan-2-ylmethyl or oxetan-3-yl;

R^5 is phenyl substituted once, twice or three times by the substituents independently selected from ethoxy, methoxy, cyclopropoxy, cyano, difluoromethoxy, fluoro, hydroxy and trideuteriomethoxy,

pyridinyl twice substituted by fluoro, or

pyrimidinyl substituted by fluoro;

R^6 is H.

12. A compound according to any one of claims 1-11, wherein Q^1 is C_{1-6}alkylene substituted once or twice by the substituents independently selected from C_{1-6}alkoxy, C_{1-6}alkylcarbonyloxy, hydroxy, C_{1-6}alkyl and trideuterioC_{1-6}alkoxy.

13. A compound according to any one of claims 1-12, wherein Q^1 is propylene substituted once or twice by the substituents independently selected from ethoxy, methoxy, hydroxy, methyl, acetoxy and trideuteriomethoxy.

14. A compound according to any one of claims 1-13, wherein Q^1 is C_{1-6}alkylene substituted by C_{1-6}alkoxy.

15. A compound according to any one of claim 1-14, wherein Q^1 is propylene substituted by methoxy or ethoxy.

16. A compound according to any one of claims 1-15, wherein Q^2 is NH or O.

17. A compound according to any one of claims 1-16, wherein A^1 is N.

18. A compound according to any one of claims 1-17, wherein A^3 is CR^8, wherein R^8 is H, C_{1-6}alkoxy, C_{1-6}alkyl, C_{3-7}cycloalkoxy, haloC_{1-6}alkoxy or halogen.

19. A compound according to any one of claims 18, wherein A^3 is CR^8, wherein R^8 is H, methoxy, methyl, cyclopropoxy, difluoromethoxy or fluoro.

20. A compound according to any one of claims 1-19, wherein A^3 is CR^8, wherein R^8 is H.

21. A compound according to any one of claims 1-20, wherein A^4 is CR^9, wherein R^9 is H.

22. A compound according to any one of claims 1-21, wherein A^5 is CR^{10} or N, wherein R^{10} is H or fluoro.

23. A compound according to any one of claims 1-22, wherein A^5 is CR^{10}, wherein R^{10} is H.

24. A compound according to any one of claims 1-22, wherein A^6 is CR^{11}, wherein R^{11} is H or halogen.

25. A compound according to any one of claims 1-24, wherein A^6 is CR^{11}, wherein R^{11} is H or fluoro.

26. A compound according to any one of claims 1-25, wherein A^6 is CR^{11}, wherein R^{11} is halogen.

27. A compound according to any one of claims 1-26, wherein A^6 is CR^{11}, wherein R^{11} is fluoro.

28. A compound according to any one of claims 1-27, wherein A^7 is CR^{12}, wherein R^{12} is H or halogen.

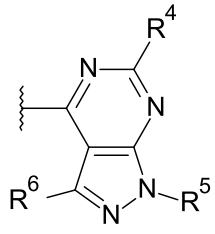
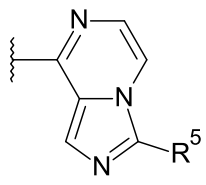
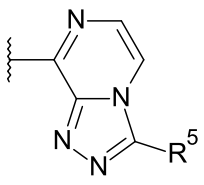
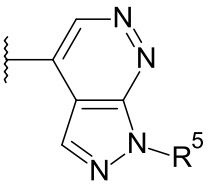
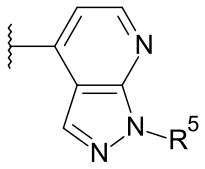
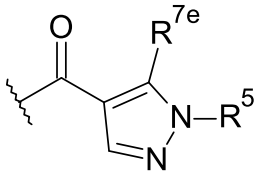
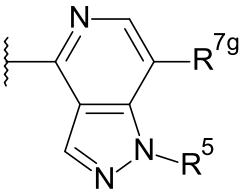
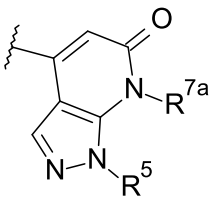
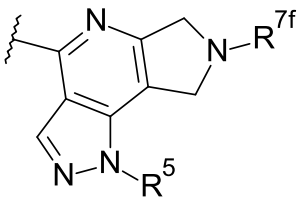
29. A compound according to any one of claims 1-28, wherein A^7 is CR^{12}, wherein R^{12} is H or fluoro.

30. A compound according to any one of claims 1-29, wherein A^7 is CR^{12}, wherein R^{12} is H.

31. A compound according to claim 2 or 3, wherein

R^1 is H or C_{1-6}alkyl;

R^2 is C_{1-6}alkyl or trideuterioC_{1-6}alkyl;

R^3 is , , , , , , , , or ; wherein

R^4 is H,

2,5-diazabicyclo[2.2.1]heptanyl substituted by haloC_{1-6}alkyl or oxetanyl,

3,6-diazabicyclo[3.1.1]heptanyl substituted by C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkyl, C_{3-7}cycloalkylcarbonyl, haloC_{1-6}alkyl, haloC_{1-6}alkylcarbonyl, hydroxyC_{1-6}alkyl, hydroxyC_{1-6}alkylcarbonyl, hydroxyC_{3-7}cycloalkylcarbonyl or oxetanyl,

3-oxa-6-azabicyclo[3.1.1]heptanyl,

3-oxa-7,9-diazabicyclo[3.3.1]nonanyl substituted by C_{1-6}alkyl,

amino,

azetidinyl substituted by C_{1-6}alkylcarbonyl, C_{1-6}alkylsulfonyl or cyano,

C_{1-6}alkoxy,

C_{1-6}alkyl unsubstituted or substituted by C_{1-6}alkoxy, C_{1-6}alkylcarbonylazetidinyl, C_{1-6}alkylsulfonylazetidinyl, C_{3-7}cycloalkylsulfonylazetidinyl, haloC_{1-6}alkylazetidinyl, hydroxy, morpholinyl or oxetanyl,

oxetanyl,

oxooxazolidinyl,

oxopyrrolidinyl unsubstituted or substituted by C_{1-6}alkyl or hydroxy;

R^5 is phenyl substituted once, twice or three times by the substituents independently selected from C_{1-6}alkoxy, C_{3-7}cycloalkyloxy, cyano, haloC_{1-6}alkoxy, halogen, hydroxy and trideuterioC_{1-6}alkoxy,

pyridinyl twice substituted by halogen, or

pyrimidinyl substituted by halogen;

R^6 is H;

R^{7a} is C_{1-6}alkyl;

R^{7e} is amino;

R^{7f} is C_{1-6}alkyl;

R^{7g} is H or C_{1-6}alkoxyC_{1-6}alkyl;

Q^1 is C_{1-6}alkylene substituted once or twice by the substituents independently selected from C_{1-6}alkoxy, C_{1-6}alkylcarbonyloxy, hydroxy, C_{1-6}alkyl and trideuterioC_{1-6}alkoxy;

Q^2 is NH or O;

A^1 is N;

A^2 is CH, CR^{7h} or N; wherein R^{7h} is C_{1-6}alkyl;

A^3 is CR^8, wherein R^8 is H, C_{1-6}alkoxy, C_{1-6}alkyl, C_{3-7}cycloalkoxy, haloC_{1-6}alkoxy or halogen;

A^4 is CR^9, wherein R^9 is H;

A^5 is CR^{10} or N, wherein R^{10} is H or halogen;

A^6 is CR^{11}, wherein R^{11} is H or halogen;

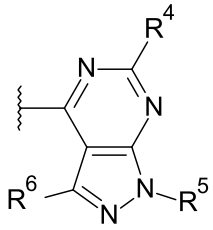
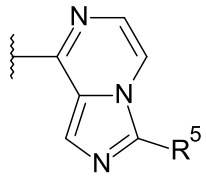
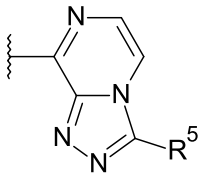
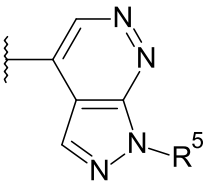
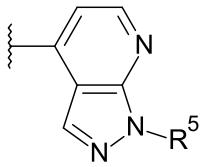
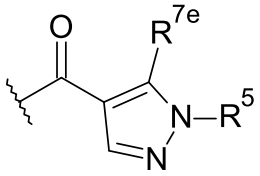
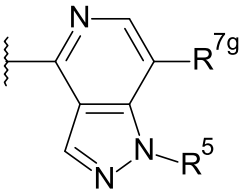
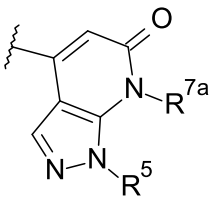
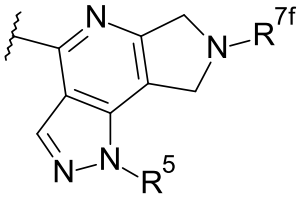
A^7 is CR^{12}, wherein R^{12} is H or halogen;

or a pharmaceutically acceptable salt thereof.

32. A compound according to claim 31, wherein

R^1 is H or methyl;

R^2 is methyl, ethyl or trideuteriomethyl;

R^3 is , , , , , , , , ; wherein

R^4 is H,

R^4 is H, (1-acetylazetidin-3-yl)methyl, (1-cyclopropylsulfonylazetidin-3-yl)methyl, (1-ethylsulfonylazetidin-3-yl)methyl, (1-methylsulfonylazetidin-3-yl)methyl, [1-(2,2,2-trifluoroethyl)azetidin-3-yl]methyl, 1-acetylazetidin-3-yl, 1-methylsulfonylazetidin-3-yl, 2-oxooxazolidin-3-yl, 2-oxopyrrolidin-1-yl, 3-(1-hydroxycyclobutanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(1-hydroxycyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2,2-trifluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2-difluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-hydroxy-2-methyl-propanoyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-hydroxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(cyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-cyanoazetidin-1-yl, 3-hydroxy-2-oxo-pyrrolidin-1-yl, 3-methyl-2-oxo-pyrrolidin-1-yl, 3-methyl-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-oxa-6-azabicyclo[3.1.1]heptan-6-yl, 5-(2,2-difluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 5-(2-fluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 5-(oxetan-3-yl)-2,5-

diazabicyclo[2.2.1]heptan-2-yl, 9-methyl-3-oxa-7,9-diazabicyclo[3.3.1]nonan-7-yl, amino, azetidin-1-yl, ethoxy, hydroxymethyl, methoxy, methoxymethyl, methyl, morpholinylmethyl, oxetan-2-ylmethyl or oxetan-3-yl;

R^5 is phenyl substituted once, twice or three times by the substituents independently selected from ethoxy, methoxy, cyclopropoxy, cyano, difluoromethoxy, fluoro, hydroxy and trideuteriomethoxy,

pyridinyl twice substituted by fluoro, or

pyrimidinyl substituted by fluoro;

R^6 is H;

R^{7a} is methyl;

R^{7e} is amino;

R^{7f} is methyl;

R^{7g} is H or methoxymethyl;

Q^1 is propylene substituted once or twice by the substituents independently selected from ethoxy, methoxy, hydroxy, methyl, acetoxy and trideuteriomethoxy;

Q^2 is NH or O;

A^1 is N;

A^2 is CH, CR^{7h} or N; wherein R^{7h} is methyl;

A^3 is CR^8, wherein R^8 is H, methoxy, methyl, cyclopropoxy, difluoromethoxy or fluoro;

A^4 is CR^9, wherein R^9 is H;

A^5 is CR^{10} or N, wherein R^{10} is H or fluoro;

A^6 is CR^{11}, wherein R^{11} is H or fluoro;

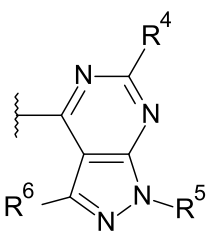
A^7 is CR^{12}, wherein R^{12} is H or fluoro;

or a pharmaceutically acceptable salt thereof.

33. A compound according to claim 31, wherein

R^1 is $C_{1\text{-}6}$alkyl;

R^2 is $C_{1\text{-}6}$alkyl;

R^3 is 

; wherein

R^4 is H,

2,5-diazabicyclo[2.2.1]heptanyl substituted by haloC_{1-6}alkyl or oxetanyl,

3,6-diazabicyclo[3.1.1]heptanyl substituted by C_{1-6}alkoxyC_{1-6}alkyl, C_{1-6}alkyl, C_{3-7}cycloalkylcarbonyl, haloC_{1-6}alkyl, haloC_{1-6}alkylcarbonyl, hydroxyC_{1-6}alkyl, hydroxyC_{1-6}alkylcarbonyl, hydroxyC_{3-7}cycloalkylcarbonyl or oxetanyl,

3-oxa-6-azabicyclo[3.1.1]heptanyl,

3-oxa-7,9-diazabicyclo[3.3.1]nonanyl substituted by C_{1-6}alkyl,

amino,

azetidinyl substituted by C_{1-6}alkylcarbonyl, C_{1-6}alkylsulfonyl or cyano,

C_{1-6}alkoxy,

C_{1-6}alkyl unsubstituted or substituted by C_{1-6}alkoxy, C_{1-6}alkylcarbonylazetidinyl, C_{1-6}alkylsulfonylazetidinyl, C_{3-7}cycloalkylsulfonylazetidinyl, haloC_{1-6}alkylazetidinyl, hydroxy, morpholinyl or oxetanyl,

oxetanyl,

oxooxazolidinyl,

oxopyrrolidinyl unsubstituted or substituted by C_{1-6}alkyl or hydroxy;

R^5 is phenyl substituted once, twice or three times by the substituents independently selected from C_{1-6}alkoxy, C_{3-7}cycloalkyloxy, cyano, haloC_{1-6}alkoxy, halogen, hydroxy and trideuterioC_{1-6}alkoxy,

pyridinyl twice substituted by halogen, or

pyrimidinyl substituted by halogen;

R^6 is H;

Q^1 is C_{1-6}alkylene substituted by C_{1-6}alkoxy;

Q^2 is NH or O;

A^1 is N;

A^2 is CH or N;

A^3 is CR^8, wherein R^8 is H;

A^4 is CR^9, wherein R^9 is H;

A^5 is CR^{10} or N, wherein R^{10} is H;

A^6 is CR^{11}, wherein R^{11} is halogen;

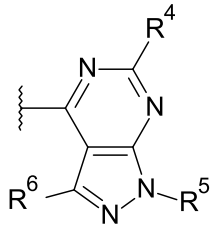
A^7 is CR^{12}, wherein R^{12} is H;

or a pharmaceutically acceptable salt thereof.

34. A compound according to claim 33, wherein

R^1 is methyl;

R^2 is methyl;

R^3 is  ; wherein

R^4 is H, (1-acetylazetidin-3-yl)methyl, (1-cyclopropylsulfonylazetidin-3-yl)methyl, (1-ethylsulfonylazetidin-3-yl)methyl, (1-methylsulfonylazetidin-3-yl)methyl, [1-(2,2,2-trifluoroethyl)azetidin-3-yl]methyl, 1-acetylazetidin-3-yl, 1-methylsulfonylazetidin-3-yl, 2-oxooxazolidin-3-yl, 2-oxopyrrolidin-1-yl, 3-(1-hydroxycyclobutanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(1-hydroxycyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2,2-trifluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2-difluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-hydroxy-2-methyl-propanoyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-hydroxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(cyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-cyanoazetidin-1-yl, 3-hydroxy-2-oxo-pyrrolidin-1-yl, 3-methyl-2-oxo-pyrrolidin-1-yl, 3-methyl-3,6-diazabicyclo[3.1.1]heptan-6-yl, 3-oxa-6-azabicyclo[3.1.1]heptan-6-yl, 5-(2,2-difluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 5-(2-fluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl, 9-methyl-3-oxa-7,9-diazabicyclo[3.3.1]nonan-7-yl, amino, azetidin-1-yl, ethoxy, hydroxymethyl, methoxy, methoxymethyl, methyl, morpholinylmethyl, oxetan-2-ylmethyl or oxetan-3-yl;

R^5 is phenyl substituted once, twice or three times by the substituents independently selected from ethoxy, methoxy, cyclopropoxy, cyano, difluoromethoxy, fluoro, hydroxy and trideuteriomethoxy,

pyridinyl twice substituted by fluoro, or

pyrimidinyl substituted by fluoro;

R^6 is H;

Q^1 is propylene substituted by methoxy or ethoxy;

Q^2 is NH or O;

A^1 is N;

A^2 is CH or N;

A^3 is CR^8, wherein R^8 is H;

A^4 is CR^9, wherein R^9 is H;

A^5 is CR^{10} or N, wherein R^{10} is H;

A^6 is CR^{11}, wherein R^{11} is fluoro;

A^7 is CR^{12}, wherein R^{12} is H;

or a pharmaceutically acceptable salt thereof.

35. A compound selected from:

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[2-(2,4-difluorophenyl)-3-methyl-imidazole-4-carbonyl]-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7-oxa-10,13,16,18-tetrazapentacyclo[14.6.1.1^{2,6}.1^{8,11}.0^{19,23}]pentacosa-1(22),2,4,6(25),17,19(23),20-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(2-hydroxyethyl)-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-13-benzyl-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(2-methoxyethyl)-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(3-morpholinopropyl)-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

3-[(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24), 21-heptaen-13-yl]propyl-trimethyl-ammonium;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyridazin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-13-methyl-10-[3-(3-pyridyl)imidazo[1,5-a]pyrazin-8-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[7-(2,4-difluorophenyl)imidazo[5,1-f][1,2,4]triazin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl] -15-(2-methoxyethoxy)-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl] -15-(2-methoxyethoxy)-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15- (2-methoxyethoxy)-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-4-(oxetan-3-yl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-22-fluoro-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(2-fluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(2-fluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-[4-fluoro-2-(trideuteriomethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluoro-6-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,6-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(1-bicyclo[1.1.1]pentanyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[3-(2,4-difluorophenyl)-[1,2,4]triazolo[4,3-a]pyrazin-8-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(3,4-difluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-15-methoxy-13,18-dimethyl-10-[1-(3-methyl-1-bicyclo[1.1.1]pentanyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(3-fluoro-1-bicyclo[1.1.1]pentanyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

5-fluoro-2-[4-[(8S,11S,15R)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-1-yl]benzonitrile;

[(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaene-4-carbonitrile;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaene-4-carbonitrile;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl) pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(cyclopropoxy) -4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18, 20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-4,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-4,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-4,10,13,17,19,26-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-4,10,13,17,19,26-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-4,5,10,13,17,19-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-4,5,10,13,17,19-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18, 20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-4,5,10,13,17,19-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-4,5,10,13,17,19-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-(hydroxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-(hydroxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-(4-fluoro-2-methoxy-phenyl)-4-[(8S,11S)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidine-6-carboxamide;

1-(4-fluoro-2-methoxy-phenyl)-N,N-dimethyl-4-[(8S,11S)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidine-6-carboxamide;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(methylamino)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-methoxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-hydroxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

8S,11S)-10-[1-(2,4-difluorophenyl)-6-ethoxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(2-methoxyethylamino)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(2-morpholinoethoxy)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-10,13,17,19,26-pentazapentacyclo-[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-10-[1-(4-fluoro-2-methylsulfanyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(difluoromethoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(difluoromethoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-[2-(difluoromethoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(difluoromethoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26 hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(6-fluoro-2-hydroxy-3-pyridyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,6-difluoro-3-pyridyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-13-methyl-10-[3-(2-thienyl)imidazo[1,5-a]pyrazin-8-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(5-fluoro-3-methoxy-2-pyridyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(3-fluoro-5-methoxy-2-pyridyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(3,5-difluoro-2-pyridyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[2-(2-chloro-4-fluoro-phenyl)pyrimidin-5-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(methylamino)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,22,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-13-(cyclopropylmethyl)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,21,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(2-morpholinoethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(2-morpholinoethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(2-methoxyethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[2-fluoro-4-(2-methoxyethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(oxetan-3-ylmethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[2-fluoro-4-(oxetan-3-ylmethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(oxetan-3-yloxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[2-fluoro-4-(oxetan-3-yloxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-[(4-methylmorpholin-2-yl)methoxy]phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-18-(2-methoxyethyl)-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-18-(2-morpholinoethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-18-(2-hydroxyethyl)-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-18-[2-[(4aR,7aS)-2,3,4a,5,7,7a-hexahydro-[1,4]dioxino[2,3-c]pyrrol-6-yl]ethyl]-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-18-[2-(4-methylsulfonylpiperazin-1-yl)ethyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-18-[2-(1,2,4-triazol-1-yl)ethyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-18-(2-pyrazol-1-ylethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-23-fluoro-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,19,21,28-hexazapentacyclo[17.6.1.1^{2,6}.1^{8,11}.0^{22,26}]octacosa-1(25),2(28),3,5,20,22(26),23-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-3-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[4-fluoro-2-(trifluoromethyl)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

5-fluoro-2-[4-[(8S,11S)-13-methyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-1-yl]benzonitrile;

5-fluoro-2-[4-[(8S,11S)-13-methyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-1-yl]benzoic acid;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-ethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-13-ethyl-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(2-methoxyethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-13-(2-morpholinoethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(3-hydroxypropyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-[3-(tetrazol-2-yl)propyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-[3-(tetrazol-1-yl)propyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-[3-(4-methylsulfonylpiperazin-1-yl)propyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-(2-hydroxyethoxy)-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-(2-hydroxyethoxy)-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-15-(2-morpholinoethoxy)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-15-(2-morpholinoethoxy)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-18,22-dideuterio-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-15-hydroxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-hydroxy-13,15,18-trimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-hydroxy-13,15,18-trimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-hydroxy-13,15,18-trimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-hydroxy-13,15,18-trimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

[(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

[(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-15-yl] acetate;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-dimethylphosphoryl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

1-(2,4-difluorophenyl)-4-[(8S,11S)-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidine-6-carboxylic acid;

1-(2,4-difluorophenyl)-4-[(8S,11S)-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidine-6-carbonitrile;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-c]pyridazin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-c]pyridazin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-b]pyridin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-b]pyridin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-b]pyridin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-b]pyridin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-23-chloro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(hydroxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-morpholino-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

Ethyl 1-(2,4-difluorophenyl)-4-[(8S,11S)-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidine-6-carboxylate;

(8S,11S)-10-[1-(4-fluoro-2-methylsulfanyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methylsulfinyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methylsulfonyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-dimethylphosphoryl-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-4-[(3,3-difluoroazetidin-1-yl)methyl]-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-4-(morpholinomethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-4,15-dimethoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-4,15-dimethoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-4,13-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-4,13-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[6-amino-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[6-amino-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-13-(cyclopropylmethyl)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-hydroxy-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-vinyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2-cyclopropyl-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2-bromo-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-18-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,22-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-4-(difluoromethoxy)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-4-(cyclopropoxy)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-13-cyclopropyl-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-(trideuteriomethoxy)-13-(trideuteriomethyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-[2-(difluoromethyl)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)-1H-pyrazolo[4,3-d]pyrimidin-7-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-18-ethyl-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-18-ethyl-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-(hydroxymethyl)-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[6-cyclobutyl-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4,5-difluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-[2-(difluoromethoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-[2-fluoro-4-(trifluoromethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-15-methoxy-10-[1-(2-methoxyphenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-chloro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(5-chloro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-4-(methoxymethyl)-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaene-4-carbonitrile;

(8S,11S)-10-[1-[4-fluoro-2-(trifluoromethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-18-(trifluoromethyl)-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-4,13-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13-ethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13-ethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)-6-methyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2-ethoxy-4-fluoro-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-4,13-dimethyl-3,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-3,5,7,10,13,17,19-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-thia-10,13,17,19,26-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(trideuteriomethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)triazolo[4,5-c]pyridin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(methoxymethyl)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluoro-6-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(5-fluoropyrimidin-2-yl)pyrazolo[4,3-c]pyridin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-3,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-3,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-3,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

8S,11S,15S)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-4,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-4,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-4,10,13,17,19-pentazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(1-bicyclo[1.1.1]pentanyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-10-[1-(5-fluoropyrimidin-2-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-fluoro-4-phenyl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(4,4,4-trifluorobutyl)-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-chloro-4-fluoro-phenyl)pyrazole-4-carbonyl]-22-fluoro-13-(3-pyrimidin-2-yloxypropyl)-4,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]-hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-22-chloro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-21-chloro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-21-fluoro-15-methoxy-13-methyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S,15R)-10-[1-(3,5-difluoro-2-pyridyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(3,5-difluoro-2-pyridyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-13-methyl-4,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-(3-methoxypropyl)-4,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-4,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13-methyl-4,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-16-oxa-7,10,13,19,21,28-hexazapentacyclo[17.6.1.1^{2,6}.1^{8,11}.0^{22,26}]octacosa-1(25),2(28),3,5,20,22(26),23-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluoro-6-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-22-fluoro-15-methoxy-13,18-dimethyl-10-(1-pyrimidin-5-ylpyrazolo[3,4-d]pyrimidin-4-yl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(11S,14S,18R)-13-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-25-fluoro-18-methoxy-16,21-dimethyl-5-oxa-10,13,16,20,22,29-hexazahexacyclo[18.6.1.1^{2,9}.1^{11,14}.0^{3,7}.0^{23,27}]nonacosa-1(26),2(29),3(7),8,21,23(27),24-heptaen-15-one;

(8S,11S,15R)-22-fluoro-15-methoxy-13,18-dimethyl-10-[1-methyl-6-(3-thienyl)pyrazolo[4,3-c]pyridin-3-yl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[6-(2,4-difluorophenyl)-1-methyl-pyrazolo[4,3-c]pyridin-3-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-21,23-difluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluoro-6-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[3-(2,4-difluorophenyl)imidazo[1,5-a]pyrazin-8-yl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-10-[2-(2,4-difluorophenoxy)-4-pyridyl]-15-ethoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S)-15-ethoxy-10-[1-[4-fluoro-2-(2-oxopyrrolidin-1-yl)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,15,18-trimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,15,18-trimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,15,18-trimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,15,18-trimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,21,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,21,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-21-fluoro-18-hydroxy-13-methyl-7-oxa-10,13,17,19-tetrazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaene-14-carboxylic acid;

(8S,11S,15R)-10-[6-ethoxy-1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-methoxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-ethoxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)-6-(morpholinomethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-18-isopropyl-13-methyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

methyl (8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-12-oxo-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaene-18-carboxylate;

(8S,11S)-18-(chloromethyl)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-18-hydroxy-13-methyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-18-amino-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-N,13-dimethyl-12-oxo-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaene-18-carboxamide;

(8S,11S,15R)-10-[1-[2-(cyclopropoxy)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

N-[(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-12-oxo-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-18-yl]acetamide;

(8S,11S,15R)-10-[1-(2,4-difluoro-6-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-18-cyclopropyl-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13-methyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S)-10-[1-(2-fluoro-4-tetrahydrofuran-3-yl-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22,23-difluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8S,11S)-10-[1-[2-fluoro-4-(2-oxopyrrolidin-1-yl)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2,4,6(26),18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-18-methyl-15-(trideuteriomethoxy)-13-(trideuteriomethyl)-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(oxetan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-tetrahydrofuran-3-yl-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-tetrahydropyran-4-yl-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(oxetan-3-ylmethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(oxetan-2-ylmethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1-methylsulfonylazetidin-3-yl)methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(1-acetylazetidin-3-yl)methyl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(oxetan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[2-hydroxy-1-(hydroxymethyl)ethyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[1-(hydroxymethyl)-2-methoxy-ethyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(1-methylsulfonylazetidin-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(1-acetylazetidin-3-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(1-methylsulfonylpyrrolidin-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[1-(2,2,2-trifluoroacetyl)pyrrolidin-3-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(1-acetylpyrrolidin-3-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1-methylsulfonylpyrrolidin-3-yl)methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[[1-(2,2,2-trifluoroacetyl)pyrrolidin-3-yl]methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(1-acetylpyrrolidin-3-yl)methyl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(1-methyl-4-piperidyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-18-methyl-13-(3-morpholinopropyl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-18-methyl-13-[3-(2-oxa-5-azabicyclo[2.2.1]heptan-5-yl)propyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-18-methyl-13-[3-(2-oxa-6-azaspiro[3.3]heptan-6-yl)propyl]-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-13-[3-[(4a*R*,7a*S*)-2,3,4a,5,7,7a-hexahydro-[1,4]dioxino[2,3-c]pyrrol-6-yl]propyl]-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-18-methyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1-ethylsulfonylazetidin-3-yl)methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(1-cyclopropylsulfonylazetidin-3-yl)methyl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[[1-(2,2,2-trifluoroethyl)azetidin-3-yl]methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-13,18-dimethyl-7,10,13,17,19,23,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)-6-(pyrrolidine-1-carbonyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-(2,4-difluorophenyl)-4-[(8S,11S)-22-fluoro-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]-N-(2-hydroxyethyl)pyrazolo[3,4-d]pyrimidine-6-carboxamide;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)-6-(morpholine-4-carbonyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-7-(methoxymethyl)pyrazolo[4,3-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-5,13,18-trimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(1,3-dihydroisobenzofuran-4-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-7-methyl-6-oxo-pyrazolo[3,4-b]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2,2,2-trifluoro-1-hydroxy-ethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[7-cyclopropyl-1-(2,4-difluorophenyl)-6-oxo-pyrazolo[3,4-b]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-7-(2-methoxyethyl)-6-oxo-pyrazolo[3,4-b]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-7-methoxy-pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26- hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-7-methoxy-pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(1-methoxycyclopropyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(1-hydroxycyclopropyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-[2-(azetidin-1-yl)-4-fluoro-phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-morpholino-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(2-fluoro-4-morpholino-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-[4-fluor-2-(2-oxa-6-azaspiro[3.3]heptan-6-yl)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluoro-6-methoxy-phenyl)-6-(methoxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-[4-fluoro-2-(3-methoxycyclobutoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-[4-fluoro-2-[1-(methoxymethyl)cyclopropoxy]phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(hydroxymethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-4,22-difluoro-10-[1-(4-fluoro-2-methoxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2,2-dioxo-2λ⁶-thia-6-azaspiro[3.3]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(hydroxymethyl)azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-oxa-6-azaspiro[3.3]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(1-hydroxy-1-methyl-ethyl)azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(3*R*,4*R*)-3-amino-4-fluoro-pyrrolidin-1-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-[(dimethylamino)methyl]azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-methylsulfonylazetidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(tetrahydrofuran-3-ylmethoxy)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(1-cyclopropylazetidin-3-yl)oxy-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[2-(4-methylpiperazin-1-yl)ethoxy]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[[1-(2,2-difluoroacetyl)azetidin-3-yl]methoxy]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(4-methylpiperazin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1-methylazetidin-3-yl)methoxy]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1-methylpyrrolidin-3-yl)methoxy]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[1-(2-fluoroethyl)azetidin-3-yl]oxy-pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(oxetan-3-ylmethoxy)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(3*S*,4*S*)-3-amino-4-methoxy-pyrrolidin-1-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(dimethylamino)azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(3-amino-3-methyl-azetidin-1-yl)-1-(2,4-difluorophenyl)pyrazolo [3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[1-(2,2-difluoroethyl)azetidin-3-yl]oxy-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-morpholinoazetidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[[1-(2,2-difluoroethyl)azetidin-3-yl]methoxy]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-

7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[1-(2,2-difluoroethyl)azetidin-3-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]ethyl acetate;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(5-fluoro-3-methylsulfanyl-2-pyridyl)-6-methylsulfanyl-pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*)-1-hydroxyethyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*S*)-1-hydroxyethyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-oxa-6-azaspiro[3.3]heptan-6-ylmethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(morpholinomethyl)azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

2-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]acetonitrile;

(8*S*,11*S*,15*R*)-10-[6-(4,7-diazaspiro[2.5]octan-7-ylmethyl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[4-(oxetan-3-yl)piperazin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(4,7-diazaspiro[2.5]octan-7-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[4-(2H-tetrazol-5-yl)-1-piperidyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(4-methyl-4,7-diazaspiro[2.5]octan-7-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(3,5-difluoro-2-pyridyl)-6-(3-morpholinoazetidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-methyl-3,6-diazabicyclo[3.1.1]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(3,4,6,7,8,8a-hexahydro-1H-pyrrolo[1,2-a]pyrazin-2-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[[(2*R*,8*S*)-2-fluoro-1,2,3,5,6,7-hexahydropyrrolizin-8-yl]methoxy]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(3a*R*,6a*S*)-2-methyl-1,3,3a,4,6,6a-hexahydropyrrolo[3,4-c]pyrrol-5-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-cyclopentyl-1-(3,5-difluoro-2-pyridyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-oxa-6-azabicyclo[3.1.1]heptan-6-ylmethyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(4a*R*,7a*R*)-4-methyl-2,3,4a,5,7,7a-hexahydropyrrolo[3,4-b][1,4]oxazin-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}] hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-benzyl-4-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]piperazine-2-carboxylic acid;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(9-methyl-3-oxa-7,9-diazabicyclo[3.3.1]nonan-7-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(4-oxazol-5-yl-1-piperidyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-methyl-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-methylsulfonyl-pyrazolo[3,4-*d*]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-10-(3-prop-1-ynylimidazo[1,5-*a*]pyrazin-8-yl)-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-methylsulfanyl-pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluoro-6-methoxy-phenyl)-6-[[1-(2-methoxyacetyl)azetidin-3-yl]methyl]pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(3-methoxy-3-methyl-but-1-ynyl)phenyl]pyrazolo[3,4-*d*]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(3-methoxypyrrolidin-1-yl)phenyl]pyrazolo[3,4-*d*]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluoro-6-tetrahydrofuran-3-yloxy-phenyl)pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

2-[2-[4-[(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-*d*]pyrimidin-1-yl]-3,5-difluoro-phenoxy]acetonitrile;

(8*S*,11*S*,15*S*)-10-[1-[2,4-difluoro-6-[(1-methylsulfonylazetidin-3-yl)methoxy]phenyl]pyrazolo[3,4-*d*]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-4-(cyclopropylamino)-10-[1-(2,4-difluorophenyl)pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(6-fluoro-1H-indazol-5-yl)pyrazolo[3,4-*d*]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)indazol-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(trideuteriomethoxy)phenyl]-6-(oxetan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-[4-fluoro-2-(trideuteriomethoxy)phenyl]-6-[3-oxa-6-azabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-methyl-7-oxo-pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-methoxyethyl)-7-oxo-pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[7-benzyloxy-1-(2,4-difluorophenyl)pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-10-[6-(cyclopropylmethyl)-1-(2,4-difluorophenyl)-7-oxo-pyrazolo[3,4-c]pyridin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[(1,1-dioxothietan-3-yl)methyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

2-[3-[1-(2,4-difluorophenyl)-4-[(8S,11S,15S)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]azetidin-1-yl]acetonitrile;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-(1,1-dioxothian-4-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(methylsulfonylmethyl)azetidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15S)-10-[6-(1-cyclopropylazetidin-3-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[(1,1-dioxothietan-3-yl)methylsulfanyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-[4-fluoro-2-(trideuteriomethoxy)phenyl]-6-(3-oxa-6-azabicyclo[3.1.1]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(azetidin-1-yl)-1-[4-fluoro-2-(trideuteriomethoxy)phenyl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-(azetidin-1-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-oxa-6-azabicyclo[3.1.1]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-oxa-6-azabicyclo[3.1.1]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(dimethylamino)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo [15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-oxa-5-azabicyclo[2.2.1]heptan-5-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(8-oxa-3-azabicyclo[3.2.1]octan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-oxa-8-azabicyclo[3.2.1]octan-8-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-imidazol-1-yl-pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2,2,2-trifluoroethoxy)pyrazolo[3,4-*d*]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-(oxetan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(3*S*)-1-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]pyrrolidine-3-carbonitrile;

8-[1-(2,4-difluorophenyl)-4-[(8S,11S,15R)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]-6,7,9,9a-tetrahydro-1*H*-pyrazino[2,1-c][1,4]oxazin-4-one;

(8*S*,11*S*,15*R*)-10-[6-(3,4,6,7,9,9a-hexahydro-1*H*-pyrazino[2,1-c][1,4]oxazin-8-yl)-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]azetidine-3-carbonitrile;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-oxa-6-azabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8S,11S,15R)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-oxa-6-azabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-methyl-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]azetidine-3-carbonitrile;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[8-(2-fluoroethyl)-5-oxa-2,8-diazaspiro[3.5]nonan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[8-(2-methoxyethyl)-5-oxa-2,8-diazaspiro[3.5]nonan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[5-(2-fluoroethyl)-8-oxa-2,5-diazaspiro[3.5]nonan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

4-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]morpholin-3-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-methyl-2-oxo-pyrrolidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,8-diazabicyclo[3.2.1]octan-8-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-

5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-5-(3-phenylpropylamino)pyrazole-4-carbonyl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(1*S*,5*S*)-3-cyclopropyl-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-oxopyrrolidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(6-oxa-3-azabicyclo[3.1.1]heptan-3-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[6-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-3-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-[(3-fluorooxetan-3-yl)methyl]-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-[(3-fluorooxetan-3-yl)methyl]-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[6-hydroxy-6-(trifluoromethyl)-2-azaspiro[3.3]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(6-oxa-1-azaspiro[3.3]heptan-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[6-(2-fluoroethyl)-1,6-diazaspiro[3.3]heptan-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-(3-oxa-6-azabicyclo[3.1.1]heptan-6-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(2-fluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(2-fluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(2-fluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2,2,2-trifluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-hydroxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

2-[3-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]azetidin-1-yl]acetonitrile;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-

dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

3-[6-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]-3,6-diazabicyclo[3.1.1]heptan-3-yl]propanenitrile;

(8*S*,11*S*,15*R*)-10-[6-[(1*R*,4*R*)-5-(2,2-difluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[(1*R*,4*R*)-5-(2,2-difluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(2,2,2-trifluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[6-[(1*R*,4*R*)-5-(2,2-difluoroethyl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-5-(2-morpholinoethylamino)pyrazole-4-carbonyl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[11-benzyl-3-(2,4-difluorophenyl)-3,4,8,11-tetrazatricyclo[7.3.0.0^{2,6}]dodeca-1(9),2(6),4,7-tetraen-7-yl]-22-fluoro-15-methoxy-13,18-

dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[3-(2,4-difluorophenyl)-11-methyl-3,4,8,11-tetrazatricyclo[7.3.0.0^{2,6}]dodeca-1(9),2(6),4,7-tetraen-7-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[5-amino-1-(2,4-difluorophenyl)pyrazole-4-carbonyl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(2*S*)-2-(methoxymethyl)pyrrolidin-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

2-[6-[1-(2,4-difluorophenyl)-4-[(8*S*,11S,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]-3,6-diazabicyclo[3.1.1]heptan-3-yl]acetonitrile;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-

7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

1-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]azetidine-3-carbonitrile;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-methyl-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[7-(2-fluoroethyl)-1,7-diazaspiro[3.5]nonan-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[7-(oxetan-3-yl)-1,7-diazaspiro[3.5]nonan-1-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[6-(oxetan-3-yl)-1,6-diazaspiro[3.3]heptan-1-

yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(6,6-dioxo-6λ^{6}-thia-1-azaspiro[3.3]heptan-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(oxetan-3-yl)-3,7-diazabicyclo[4.2.0]octan-7-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[3-(2,4-difluorophenyl)-11-(oxetan-3-yl)-3,4,8,11-tetrazatricyclo[7.3.0.0^{2,6}]dodeca-1(9),2(6),4,7-tetraen-7-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,5*S*)-3-(oxetan-3-yl)-3,6-diazabicyclo[3.2.0]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(2-methoxyethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[(1*R*,4*R*)-5-(oxetan-3-yl)-2,5-diazabicyclo[2.2.1]heptan-2-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-10-[1-(4-fluoro-2-hydroxy-phenyl)-6-[3-(oxetan-3-yl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(3-hydroxy-2-oxo-pyrrolidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(2-oxopyrrolidin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

3-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-15-ethoxy-22-fluoro-13,18-dimethyl-12-oxo-5,7,10,13,17,19,26-heptazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]oxazolidin-2-one;

3-[1-(2,4-difluorophenyl)-4-[(8*S*,11*S*,15*R*)-22-fluoro-15-methoxy-13,18-dimethyl-12-oxo-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-10-yl]pyrazolo[3,4-d]pyrimidin-6-yl]oxazolidin-2-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-(4-methyl-2-oxo-piperazin-1-yl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2,4,6(26),18,20,22-heptaen-12-one;

(8*S*,11*S*,15*S*)-22-fluoro-10-[6-[3-(2-fluoroethyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(4-fluoro-2-hydroxy-phenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[1-(2,4-difluorophenyl)-6-[3-(2,2,2-trifluoroacetyl)-3,6-

diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*S*)-10-[6-[3-(2,2-difluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(23),2(26),3,5,18,20(24),21-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2,2,2-trifluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[6-[3-(2,2-difluoroacetyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-15-ethoxy-22-fluoro-13,18-dimethyl-7-oxa-5,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(2-hydroxy-2-methyl-propanoyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(1-hydroxycyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(1-hydroxycyclobutanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

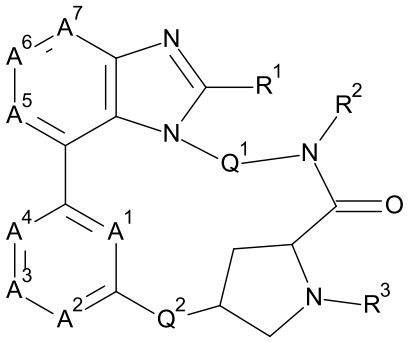
(8*S*,11*S*,15*R*)-10-[6-[3-(cyclopropanecarbonyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]-1-(2,4-difluorophenyl)pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one; and

(8*S*,11*S*,15*R*)-10-[1-(2,4-difluorophenyl)-6-[3-(3-fluorocyclobutyl)-3,6-diazabicyclo[3.1.1]heptan-6-yl]pyrazolo[3,4-d]pyrimidin-4-yl]-22-fluoro-15-methoxy-13,18-dimethyl-7,10,13,17,19,26-hexazapentacyclo[15.6.1.1^{2,6}.1^{8,11}.0^{20,24}]hexacosa-1(24),2(26),3,5,18,20,22-heptaen-12-one;

or a pharmaceutically acceptable salt thereof.

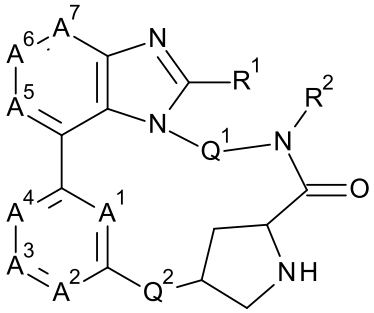
36. A process for the preparation of a compound according to any one of claims 1 to 35 comprising any of the following steps:

a) the formation of compound of formula (I),



(I),

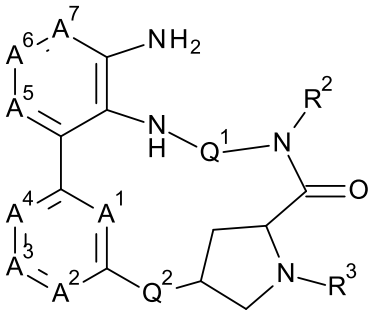
via nucleophilic substitution between compound of formula (VIII),

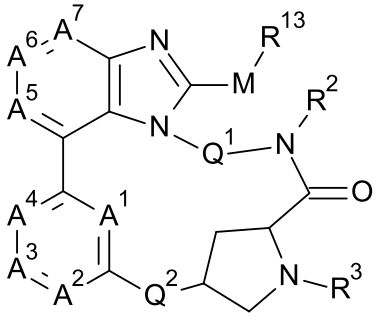
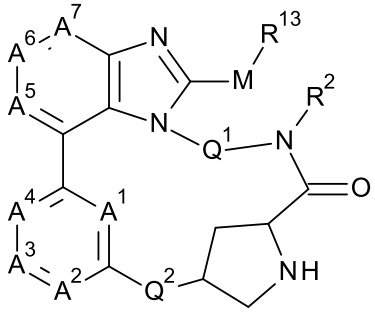


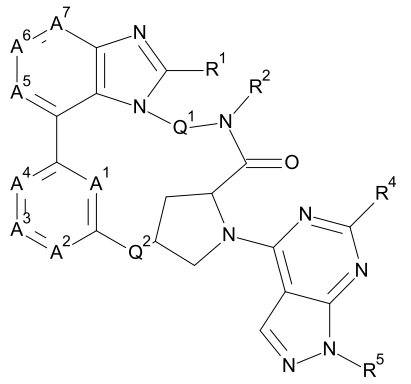
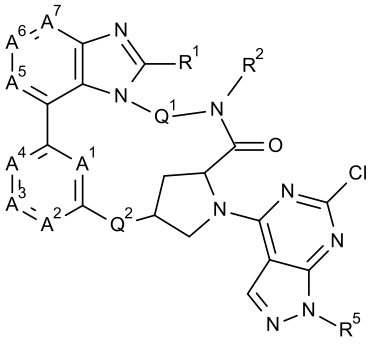
(VIII), and R^3X, in the presence of a base;

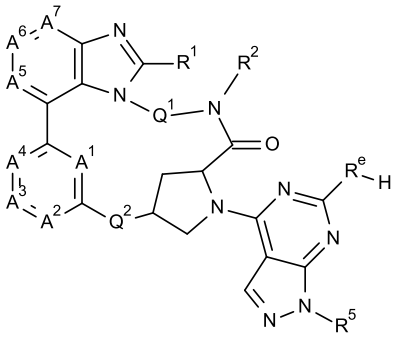
b) the formation of compound of formula (I) via Buchwald-Hartwig reaction between compound of formula (VIII) and R^3X, in the presence of a catalyst;

c) the formation of compound of formula (I) via condensation reaction between compound of formula (VIII) and R^3OH, in the presence of coupling reagent;

d) the formation of compound of formula (I) via cyclization reaction between compound of formula (XVIII) ,  (XVIII), and regents;

e) the formation of compound of formula (XXI),  (XXI), via nucleophilic substitution between compound of formula (XX),  (XX), and R^3X in the presence of a base;

f) the formation of compound of formula (XXII),  (XXII), via reaction between compound of formula (Ic),  (Ic), and alcohols or amines (R^4-X) in the presence of a base;

g) the amino group of compound of formula (XXIV),  (XXIV), reacts with halide via nucleophilic substitution or with ketone via reductive amination to provide final compound of formula (XXII);

wherein

in step a) and e) , the base is DIEA;

in step b), the catalyst is $Pd_2(dba)_3$;

in step c), the coupling reagent is HATU;

in step d), the regents is CDI, BrCN or $R^1(OCH_3)_3$;

in step e), the base is DIEA;

in step f), the base is DIPEA, K_2CO_3 or NaH;

X is halogen; R^{13} is R^a, R^b or R^c; M is $C_{1\text{-}6}$alkylene; R^e is a substituent bearing primary or secondary amino groups; R^1 to R^7, Q^1, Q^2, A^1 to A^7 are as defined as in any one of claims 1 to 34.

37. A compound or pharmaceutically acceptable salt according to any one of claims 1 to 35 for use as therapeutically active substance.

38. A pharmaceutical composition comprising a compound in accordance with any one of claims 1 to 35 and a therapeutically inert carrier.

39. The use of a compound according to any one of claims 1 to 35 in the manufacture of a medicament for the treatment or prophylaxis of autoimmune diseases, inflammatory diseases, neurological disorders diseases, metabolic diseases, cardiovascular diseases, ocular diseases, or selective types of cancers where overexpression or activation of STING is implicated.

40. The use of a compound according to any one of claims 1 to 35 in the manufacture of a medicament for the treatment to subjects suffered from an interferonopathy or auto-inflammatory diseases in which the STING activation are the root-cause of disease pathologies.

41. A compound according to any one of claims 1 to 35 for use in the inhibition of STING.

42. The use of a compound according to any one of claims 1 to 35 in the preparation of a medicament for the inhibition of STING.

43. A compound or pharmaceutically acceptable salt according to any one of claims 1 to 35, when manufactured according to a process of claim 36.

44. Use of a compound according to any one of claims 1 to 35 in the manufacture of a medicament for the treatment or prophylaxis of autoimmune diseases.