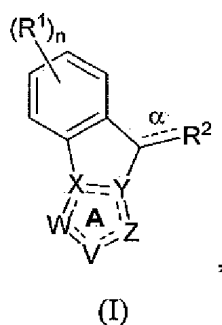


also be an important treatment strategy for patients with neurological or neuropsychiatric diseases or disorders such as depression. The compounds, compositions and methods herein help meet the current need for IDO1, IDO2 and TDO modulators.

[0014] In this invention, we describe novel structures related to imidazoisindoles, with different electronic and heme binding properties, that can specifically inhibit IDO1, IDO2 or TDO, or that can exert combined inhibition of tryptophan degradation mediated by any of these enzymes.

[0014a] In one aspect, there is provided a compound of the formula,

[Chemical formula 41]



its tautomers or a pharmaceutically acceptable salt thereof, wherein

n is 0, 1, 2, 3, or 4;

bond α is a single or double bond

ring A is an aromatic ring wherein

- i) V and X are N, W and Z are CH and Y is C; or
- ii) V, Y and Z are N, W is CH and X is C; or
- iii) V, W and Y are N, X is C and Z is CH; or
- iv) V and W are N or NH, X and Y are C and Z is CH;

each R^1 is independently halogen, cyano, nitro, C_{1-6} alkyl, C_{1-6} haloalkyl, -OR, -N(R)₂, -SR, -C(O)OR, -C(O)N(R)₂, -C(O)R, -S(O)R, -S(O)OR, -S(O)N(R)₂, -S(O)₂R, -S(O)₂OR, -S(O)₂N(R)₂, -OC(O)R, -OC(O)OR, -OC(O)N(R)₂, -N(R)C(O)R, -N(R)C(O)OR, or -N(R)C(O)N(R)₂;

R^2 is $-C_{1-4}$ alkyl- R^A or $-C_{2-4}$ alkenyl- R^3 when bond α is a single bond; and

R^2 is $=C(H)R^A$ when bond α is a double bond;

wherein

R^A is -CN, -C(O) R^3 , -C(O)OR³, -C(O)N(R^3)(R^C), -C(OR^B)(R^3)(R^C), -C(NHR^B)(R^3)(R^C), or -C(=N-OR^C) R^3 , wherein

- 5a -

R^B is hydrogen, C_{1-6} alkyl, C_{1-6} haloalkyl, $-C_{1-6}$ alkyl- R^{B1} , $-C(O)R^3$, $-S(O)_2R^3$, $-C(O)(CH_2)_{1-4}COOR$, $-C(O)CH(NH_2)(R^D)$, $-S(O)_2OR^3$, $-S(O)_2N(R^3)_2$, $-CH_2-OP(O)_2(OR)_2$, or $-P(O)(OR^3)_2$, wherein

R^{B1} is cyano, nitro, C_{1-6} alkyl, C_{1-6} haloalkyl, $-OR$, $-N(R)_2$, $-SR$, $-C(O)OR$, $-C(O)N(R)_2$, $-C(O)R$, $-S(O)R$, $-S(O)OR$, $-S(O)N(R)_2$, $-S(O)_2R$, $-S(O)_2OR$, $-S(O)_2N(R)_2$, $-OC(O)R$, $-OC(O)OR$, $-OC(O)N(R)_2$, $-N(R)C(O)R$, $-N(R)C(O)OR$, or $-N(R)C(O)N(R)_2$;

R^D is hydrogen, methyl, $-CH(CH_3)_2$, $-CH_2CH(CH_3)_2$, $-CH(CH_3)(CH_2CH_3)$, benzyl, 4-hydroxybenzyl, $-CH_2(3-indolyl)$, $-CH_2SH$, $-CH_2CH_2SCH_3$, $-CH_2OH$, $-CH(CH_3)OH$, $-(CH_2)_4-NH_2$, $-(CH_2)_3-N(H)C(=NH)NH_2$, $-CH_2(4-imidazolyl)$, $-CH_2COOH$, $-CH_2CH_2COOH$, $-CH_2CONH_2$, or $-CH_2CH_2CONH_2$;

each R^3 is independently hydrogen, C_{1-6} alkyl, C_{1-6} haloalkyl, aryl, heteroaryl, C_{3-8} cycloalkyl, C_{3-8} cycloalkenyl, 3-10 membered heterocyclyl, aryl C_{1-6} alkyl-, heteroaryl C_{1-6} alkyl-, C_{3-8} cycloalkyl C_{1-6} alkyl-, C_{3-8} cycloalkenyl C_{1-6} alkyl-, or (3-10 membered heterocyclyl) C_{1-6} alkyl-, wherein

the C_{1-6} alkyl, C_{3-8} cycloalkyl, C_{3-8} cycloalkenyl, 3-10 membered heterocyclyl, C_{3-8} cycloalkyl C_{1-6} alkyl-, C_{3-8} cycloalkenyl C_{1-6} alkyl-, and (3-10 membered heterocyclyl) C_{1-6} alkyl- are each optionally and independently substituted by one $=R^{32}$ group and each optionally substituted and independently by one, two, three, or four R^{31} groups; the aryl, heteroaryl, aryl C_{1-6} alkyl-, and heteroaryl C_{1-6} alkyl- groups, are each optionally substituted by one, two, three, or four R^{31} groups;

wherein

each R^{31} is independently halogen, cyano, nitro, C_{1-6} alkyl, $-C_{1-6}$ alkyl- R^{33} , C_{1-6} haloalkyl, $-OR$, $-N(R)_2$, $-SR$, $-C(O)OR$, $-C(O)N(R)_2$, $-C(O)R$, $-S(O)R$, $-S(O)OR$, $-S(O)N(R)_2$, $-S(O)_2R$,

- 5b -

-S(O)₂OR, -S(O)₂N(R)₂, -OC(O)R, -OC(O)OR, -OC(O)N(R)₂,
-N(R)C(O)R, -N(R)C(O)OR, or -N(R)C(O)N(R)₂, wherein
R³³ is cyano, -OR, -N(R)₂, -SR, -C(O)OR, -C(O)N(R)₂,
-C(O)R, -S(O)R, -S(O)OR, -S(O)N(R)₂, -S(O)₂R,
-S(O)₂OR, -S(O)₂N(R)₂, -OC(O)R, -OC(O)OR,
-OC(O)N(R)₂, -N(R)C(O)R, -N(R)C(O)OR, or
-N(R)C(O)N(R)₂;

R³² is =O, =S, =N(R), =N(OR), =C(R³⁴)₂, =(spiro-C₃₋₈cycloalkyl),
or =(spiro-(3-10 membered heterocyclyl)), wherein
each R³⁴ is independently hydrogen, halogen, C₁₋₆alkyl,
C₁₋₆haloalkyl, C₃₋₈cycloalkyl, or 3-10 membered
heterocyclyl;
or both R³⁴ taken together with the atom to which they are both
attached form a monocyclic C₃₋₈cycloalkyl or monocyclic
3-8 membered heterocyclyl;

R^C is hydrogen or C₁₋₆alkyl;

and

each R is independently hydrogen or R¹⁰, wherein

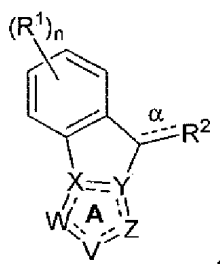
R¹⁰ is C₁₋₆alkyl, C₁₋₆haloalkyl, aryl, heteroaryl, C₃₋₈cycloalkyl, C₃₋₈cycloalkenyl, 3-
10 membered heterocyclyl, arylC₁₋₆alkyl, heteroarylC₁₋₆alkyl-, C₃₋₈ cycloalkylC₁₋₆
alkyl-, C₃₋₈cycloalkenylC₁₋₆alkyl-, or (3-10 membered heterocyclyl)C₁₋₆alkyl-,
each R¹⁰ optionally substituted by one, two, three, or four groups that are each
independently halogen, cyano, nitro, C₁₋₆alkyl, C₁₋₆haloalkyl, -OR¹¹, -N(R¹¹)₂,
-SR¹¹, -C(O)OR¹¹, -C(O)N(R¹¹)₂, -C(O)R¹¹, -S(O)R¹¹, -S(O)OR¹¹, -S(O)N(R¹¹)₂,
-S(O)₂R¹¹, -S(O)₂OR¹¹, -S(O)₂N(R¹¹)₂, -OC(O)R¹¹, -OC(O)OR¹¹, -OC(O)N(R¹¹)₂,
-N(R¹¹)C(O)R¹¹, -N(R¹¹)C(O)OR¹¹, or -N(R¹¹)C(O)N(R¹¹)₂, wherein each R¹¹ is
independently hydrogen or C₁₋₆alkyl.

[0014b] In one aspect, there is provided a pharmaceutical composition comprising a
compound as described herein, its tautomers or a pharmaceutically acceptable salt thereof, or a
compound as described herein, and a pharmaceutically acceptable diluent, excipient, or carrier.

[0014c] In one aspect, there is provided a pharmaceutical composition as described herein for
treating immunosuppression in a subject in need thereof.

[0014d] In one aspect, there is provided a composition comprising a compound as described herein, its tautomers or a pharmaceutically acceptable salt thereof, or a compound as described herein and a pharmaceutically acceptable diluent, excipient, or carrier for inhibiting IDO in a cell.

[0015] In one aspect, the invention comprises compounds according to the formula (I) and its tautomers,



wherein

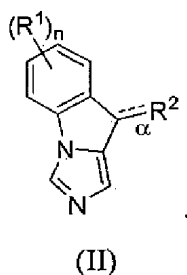
R^1 , R^2 , and n are each defined herein;

bond α is a simple or double bond

ring A is an aromatic ring wherein

- i) V and X are N, W and Z are CH and Y is C; or
- ii) V, Y and Z are N, W is CH and X is C; or
- iii) V, W and Y are N, X is C and Z is CH; or
- iv) V and W are N or NH, X and Y are C and Z is CH;

[0016] In a most preferred embodiment, the invention comprises compounds of Formula (I) where V and X are nitrogen (N), W and Z are CH and Y is carbon (C), to yield compounds of Formula (II),

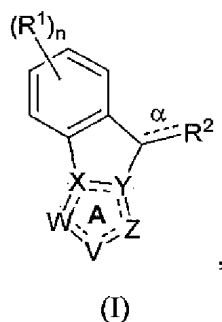


We Claim:

[Claim 1]

A compound of the formula,

[Chemical formula 41]



its tautomers or a pharmaceutically acceptable salt thereof, wherein

n is 0, 1, 2, 3, or 4;

bond α is a single or double bond

ring A is an aromatic ring wherein

- i) V and X are N, W and Z are CH and Y is C; or
- ii) V, Y and Z are N, W is CH and X is C; or
- iii) V, W and Y are N, X is C and Z is CH; or
- iv) V and W are N or NH, X and Y are C and Z is CH;

each R^1 is independently halogen, cyano, nitro, C_{1-6} alkyl, C_{1-6} haloalkyl, -OR, -N(R)₂, -SR, -C(O)OR, -C(O)N(R)₂, -C(O)R, -S(O)R, -S(O)OR, -S(O)N(R)₂, -S(O)₂R, -S(O)₂OR, -S(O)₂N(R)₂, -OC(O)R, -OC(O)OR, -OC(O)N(R)₂, -N(R)C(O)R, -N(R)C(O)OR, or -N(R)C(O)N(R)₂;

R^2 is $-C_{1-4}$ alkyl- R^A or $-C_{2-4}$ alkenyl- R^3 when bond α is a single bond; and

R^2 is $=C(H)R^A$ when bond α is a double bond;

wherein

R^A is -CN, -C(O)R³, -C(O)OR³, -C(O)N(R³)(R^C), -C(OR^B)(R³)(R^C), -C(NHR^B)(R³)(R^C), or -C(=N-OR^C)R³, wherein

R^B is hydrogen, C_{1-6} alkyl, C_{1-6} haloalkyl, $-C_{1-6}$ alkyl- R^{B1} , -C(O)R³, -S(O)₂R³, -C(O)(CH₂)₁₋₄COOR, -C(O)CH(NH₂)(R^D), -

S(O)₂OR³, -S(O)₂N(R³)₂, -CH₂-OP(O)₂(OR)₂, or -P(O)(OR³)₂, wherein

R^{B1} is cyano, nitro, C_{1-6} alkyl, C_{1-6} haloalkyl, -OR, -N(R)₂, -SR, -C(O)OR, -C(O)N(R)₂, -C(O)R, -S(O)R, -S(O)OR, -S(O)N(R)₂, -S(O)₂R, -S(O)₂OR, -S(O)₂N(R)₂, -OC(O)R,

-OC(O)OR, -OC(O)N(R)₂, -N(R)C(O)R, -N(R)C(O)OR, or
-N(R)C(O)N(R)₂;

R^D is hydrogen, methyl, -CH(CH₃)₂, -CH₂CH(CH₃)₂,
-CH(CH₃)(CH₂CH₃), benzyl, 4-hydroxybenzyl, -CH₂(3-
indolyl), -CH₂SH, -CH₂CH₂SCH₃, -CH₂OH, -CH(CH₃)OH,
-(CH₂)₄-NH₂, -(CH₂)₃-N(H)C(=NH)NH₂, -CH₂(4-
imidazolyl), -CH₂COOH, -CH₂CH₂COOH, -CH₂CONH₂,
or -CH₂CH₂CONH₂;

each R³ is independently hydrogen, C₁₋₆alkyl, C₁₋₆haloalkyl, aryl,
heteroaryl, C₃₋₈cycloalkyl, C₃₋₈cycloalkenyl, 3-10 membered
heterocyclyl, arylC₁₋₆alkyl-, heteroarylC₁₋₆alkyl-, C₃₋₈cycloalkylC₁₋₆
alkyl-, C₃₋₈cycloalkenylC₁₋₆alkyl-, or (3-10 membered
heterocyclyl)C₁₋₆alkyl-,

wherein

the C₁₋₆alkyl, C₃₋₈cycloalkyl, C₃₋₈cycloalkenyl, 3-10 membered
heterocyclyl, C₃₋₈cycloalkylC₁₋₆alkyl-,
C₃₋₈cycloalkenylC₁₋₆alkyl-, and (3-10 membered
heterocyclyl)C₁₋₆alkyl- are each optionally and
independently substituted by one =R³² group and each
optionally substituted and independently by one, two, three,
or four R³¹ groups;

the aryl, heteroaryl, arylC₁₋₆alkyl-, and heteroarylC₁₋₆alkyl-
groups, are each optionally substituted by one, two, three,
or four R³¹ groups;

wherein

each R³¹ is independently halogen, cyano, nitro, C₁₋₆alkyl,
-C₁₋₆alkyl-R³³, C₁₋₆haloalkyl, -OR, -N(R)₂, -SR, -C(O)OR,
-C(O)N(R)₂, -C(O)R, -S(O)R, -S(O)OR, -S(O)N(R)₂,
-S(O)₂R, -S(O)₂OR, -S(O)₂N(R)₂, -OC(O)R, -OC(O)OR,
-OC(O)N(R)₂, -N(R)C(O)R, -N(R)C(O)OR, or
-N(R)C(O)N(R)₂, wherein
R³³ is cyano, -OR, -N(R)₂, -SR, -C(O)OR, -C(O)N(R)₂,
-C(O)R, -S(O)R, -S(O)OR, -S(O)N(R)₂, -S(O)₂R,

-S(O)₂OR, -S(O)₂N(R)₂, -OC(O)R, -OC(O)OR,
-OC(O)N(R)₂, -N(R)C(O)R, -N(R)C(O)OR, or
-N(R)C(O)N(R)₂;

R³² is =O, =S, =N(R), =N(OR), =C(R³⁴)₂, =(spiro-C₃-
cycloalkyl), or =(spiro-(3-10 membered heterocyclyl)),
wherein

each R³⁴ is independently hydrogen, halogen, C₁₋₆alkyl,
C₁₋₆haloalkyl, C₃₋₈cycloalkyl, or 3-10 membered
heterocyclyl;

or both R³⁴ taken together with the atom to which they are
both attached form a monocyclic C₃₋₈cycloalkyl or
monocyclic 3-8 membered heterocyclyl;

R^C is hydrogen or C₁₋₆alkyl;

and

each R is independently hydrogen or R¹⁰, wherein

R¹⁰ is C₁₋₆alkyl, C₁₋₆haloalkyl, aryl, heteroaryl, C₃₋₈cycloalkyl, C₃₋₈cycloalkenyl,
3-10 membered heterocyclyl, arylC₁₋₆alkyl, heteroarylC₁₋₆alkyl-, C₃₋₈
cycloalkylC₁₋₆alkyl-, C₃₋₈cycloalkenylC₁₋₆alkyl-, or (3-10 membered
heterocyclyl)C₁₋₆alkyl-, each R¹⁰ optionally substituted by one, two, three, or
four groups that are each independently halogen, cyano, nitro, C₁₋₆alkyl,
C₁₋₆haloalkyl, -OR¹¹, -N(R¹¹)₂, -SR¹¹, -C(O)OR¹¹, -C(O)N(R¹¹)₂, -C(O)R¹¹,
-S(O)R¹¹, -S(O)OR¹¹, -S(O)N(R¹¹)₂, -S(O)₂R¹¹, -S(O)₂OR¹¹, -S(O)₂N(R¹¹)₂,
-OC(O)R¹¹, -OC(O)OR¹¹, -OC(O)N(R¹¹)₂, -N(R¹¹)C(O)R¹¹, -
N(R¹¹)C(O)OR¹¹, or -N(R¹¹)C(O)N(R¹¹)₂, wherein each R¹¹ is independently
hydrogen or C₁₋₆alkyl.

[Claim 2]

The compound of claim 1, its tautomers or a pharmaceutically acceptable salt thereof,
wherein bond α is a single bond.

[Claim 3]

The compound of claim 1, its tautomers or a pharmaceutically acceptable salt thereof,
wherein V and X are N, W and Z are CH and Y is C.

[Claim 4]

The compound of claim 1, its tautomers or a pharmaceutically acceptable salt thereof, wherein V, Y and Z are N, W is CH and X is C.

[Claim 5]

The compound of claim 1, its tautomers or a pharmaceutically acceptable salt thereof, wherein V, W and Y are N, X is C and Z is CH.

[Claim 6]

The compound of claim 1, its tautomers or a pharmaceutically acceptable salt thereof, wherein V and W are N or NH, X and Y are C and Z is CH.

[Claim 7]

The compound of claim 1, 2, 3, 4, 5 or 6, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^2 is $-C_{1-4}alkyl-R^A$.

[Claim 8]

The compound of claim 1, 2, 3, 4, 5 or 6, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^2 is $-CH_2-R^A$, $-CH_2CH_2-R^A$, $-C(H)(CH_3)CH_2-R^A$, or $-C(H)=C(H)R^3$.

[Claim 9]

The compound of claim 8, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^2 is $-CH_2-R^A$.

[Claim 10]

The compound of any one of claims 1 - 9, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^A is $-C(O)R^3$ or $-C(OR^B)(R^3)(R^C)$.

[Claim 11]

The compound of any one of claims 1 - 9, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^A is $-C(NHR^B)(R^3)(R^C)$, or $-C(=N-OR^C)R^3$.

[Claim 12]

The compound of any one of claims 1 - 9, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^A is $-C(NHR^B)(R^3)(R^C)$, wherein R^B is hydrogen, $C_{1-6}alkyl$, or $-C(O)C_{1-6}alkyl$.

[Claim 13]

The compound of any one of claims 1 - 9, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^A is $-C(OR^B)(R^3)(R^C)$.

[Claim 14]

The compound of any one of claims 1 - 9, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^A is $-CH(OH)(R^3)$.

[Claim 15]

The compound of any one of claims 1 - 14, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^3 is hydrogen.

[Claim 16]

The compound of any one of claims 1 - 14, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^3 is aryl, heteroaryl, C_{3-8} cycloalkyl, C_{3-8} cycloalkenyl, 3-10 membered heterocyclyl, or C_{3-8} cycloalkyl C_{1-6} alkyl-, wherein

the C_{3-8} cycloalkyl, C_{3-8} cycloalkenyl, 3-10 membered heterocyclyl, and C_{3-8} cycloalkyl C_{1-6} alkyl-, are each optionally and independently substituted by one $=R^{32}$ group and each optionally and independently substituted by one or two R^{31} groups; and
the aryl and heteroaryl groups, are each optionally substituted by one or two R^{31} groups.

[Claim 17]

The compound of any one of claims 1 - 14, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^3 is phenyl, a five or six membered heteroaryl, monocyclic C_{5-8} cycloalkyl, monocyclic C_{5-8} cycloalkenyl, a five or six membered monocyclic heterocyclyl, or (monocyclic C_{5-8} cycloalkyl) C_{1-6} alkyl-, wherein

the C_{5-8} cycloalkyl, C_{5-8} cycloalkenyl, 5 - 6 membered heterocyclyl, and C_{5-8} cycloalkyl C_{1-6} alkyl-, are each optionally and independently substituted by one $=R^{32}$ group and each optionally and independently substituted by one or two R^{31} groups; and
the phenyl and heteroaryl groups, are each optionally substituted by one or two R^{31} groups.

[Claim 18]

The compound of any one of claims 1 - 14, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^3 is phenyl or a five or six membered heteroaryl, each optionally substituted by one or two R^{31} groups.

[Claim 19]

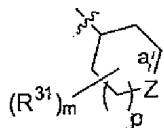
The compound of any one of claims 1 - 14, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^3 is monocyclic C_{5-8} cycloalkyl, monocyclic C_{5-8} cycloalkenyl, a five or six membered monocyclic heterocyclyl, or (monocyclic C_{5-8} cycloalkyl) C_{1-6} alkyl-,

cycloalkyl)C₁₋₆alkyl-, each optionally substituted by one =R³² group and one or two R³¹ groups.

[Claim 20]

The compound of any one of claims 1 - 14, its tautomers or a pharmaceutically acceptable salt thereof, wherein R³ is

[Chemical formula 42]



, wherein

bond a is a single bond or a double bond;

m is 0, 1, or 2;

p is 0 or 1; and

when bond "a" is a single bond, then Z is -C(R³⁶)₂-, -C(=R³²)-, -N(R³⁵)-, or -O-,

wherein

R³⁵ is hydrogen, C₁₋₆alkyl, -C(O)R, -S(O)₂R, -C(O)OR, -C(O)N(R)₂, -S(O)₂OR, or -S(O)₂N(R)₂; and

when bond a is a double bond, then Z is -C(R³⁶)= or -N=;

each R³⁶ is independently hydrogen or R³¹.

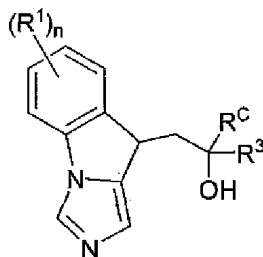
[Claim 21]

The compound of claim 20, its tautomers or a pharmaceutically acceptable salt thereof, wherein bond a is a single bond; and Z is -N(R³⁵)- or -O-.

[Claim 22]

The compound of claim 1 of the formula,

[Chemical formula 43]

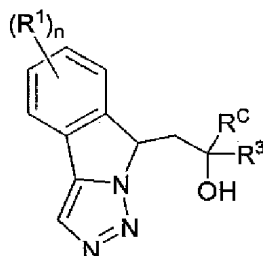


, its tautomers or a pharmaceutically acceptable salt thereof.

[Claim 23]

The compound of claim 1 of the formula,

[Chemical formula 44]

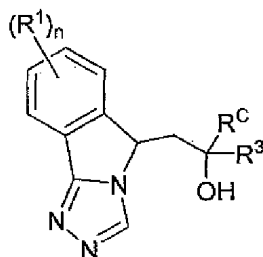


, its tautomers or a pharmaceutically acceptable salt thereof.

[Claim 24]

The compound of claim 1 of the formula,

[Chemical formula 45]

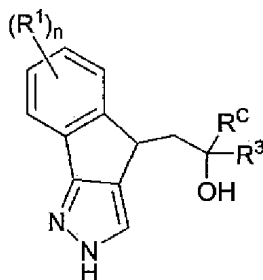


, its tautomers or a pharmaceutically acceptable salt thereof.

[Claim 25]

The compound of claim 1 of the formula,

[Chemical formula 46]



, its tautomers or a pharmaceutically acceptable salt thereof.

[Claim 26]

The compound of claim 22-25, its tautomers or a pharmaceutically acceptable salt thereof, wherein

R^c is hydrogen and

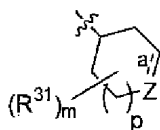
R^3 is aryl, heteroaryl, C_{3-8} cycloalkyl, C_{3-8} cycloalkenyl, or 3-10 membered heterocyclyl, wherein

the C_{3-8} cycloalkyl, C_{3-8} cycloalkenyl, and 3-10 membered heterocyclyl are each optionally substituted by one $=R^{32}$ group and one, two, three, or four R^{31} groups; and
the aryl and heteroaryl are each optionally substituted by one, two, three, or four R^{31} groups.

[Claim 27]

The compound of any one of claims 22 - 25, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^3 is

[Chemical formula 47]



, wherein

bond a is a single bond or a double bond;

m is 0, 1, or 2;

p is 0 or 1; and

when bond "a" is a single bond, then Z is $-C(R^{36})_2-$, $-C(=R^{32})-$, $-N(R^{35})-$, or $-O-$,
wherein

R^{35} is hydrogen, C_{1-6} alkyl, $-C(O)R$, $-S(O)_2R$, $-C(O)OR$, $-C(O)N(R)_2$, $-S(O)_2OR$,
or $-S(O)_2N(R)_2$; and

when bond a is a double bond, then Z is $-C(R^{36})=$ or $-N=$;

each R^{36} is independently hydrogen or R^{31} .

[Claim 28]

The compound according to claim 1, its tautomers or a pharmaceutically acceptable salt thereof, wherein

R^2 is $=C(H)R^A$ when bond a is a double bond;

R^A is $-C(O)R^3$ or $-C(OR^B)(R^3)(R^C)$;

R^B is hydrogen, C_{1-4} alkyl, or C_{1-4} haloalkyl; and

R^C is hydrogen or C_{1-4} alkyl.

[Claim 29]

The compound according to claim 1, its tautomers or a pharmaceutically acceptable salt thereof, wherein

α is a single bond and R^2 is $-C_{1-4}alkyl-R^A$ or $-C_{2-4}alkenyl-R^3$;

R^A is $-C(O)R^3$, $-C(OR^B)(R^3)(R^C)$, or $-C(NHR^B)(R^3)(R^C)$, or $-C(=N-OR^C)R^3$;

R^B is hydrogen, $C_{1-4}alkyl$, or $C_{1-4}haloalkyl$; and,

R^C is hydrogen or $C_{1-4}alkyl$.

[Claim 30]

The compound according to claim 22, 28, or 29, its tautomers or a pharmaceutically acceptable salt thereof, wherein

R^3 is aryl, heteroaryl, $C_{3-8}cycloalkyl$, $C_{3-8}cycloalkenyl$, or 3-10 membered heterocyclyl, wherein the $C_{3-8}cycloalkyl$, $C_{3-8}cycloalkenyl$, and 3-10 membered heterocyclyl are each optionally substituted by one $=R^{32}$ group and one, two, three, or four R^{31} groups; and

the aryl and heteroaryl are each optionally substituted by one, two, three, or four R^{31} groups

[Claim 31]

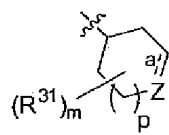
The compound according to claim 22, 28, or 29, its tautomers or a pharmaceutically acceptable salt thereof, wherein R^3 is phenyl, cyclopentyl, cyclohexyl, cyclohex-1-en-1-yl, cyclohex-3-en-1-yl, furan-2-yl, furan-3-yl, tetrahydropyran-3-yl, tetrahydropyran-4-yl, piperidin-3-yl, piperidin-4-yl, imidazol-2-yl, imidazol-4-yl, thiazol-2-yl, thiazol-4-yl, thiazol-5-yl, each optionally substituted by one or two R^{31} groups, and wherein the cyclopentyl, cyclohexyl, cyclohexenyl, and piperidinyl groups are each optionally substituted by one $=R^{32}$ group.

[Claim 32]

The compound according to claim 22, 28, or 29, its tautomers or a pharmaceutically acceptable salt thereof, wherein

R^3 is

[Chemical formula 10]



, wherein bond a is a single bond or a double bond; m is 0, 1, or 2; p is 0 or 1; and wherein

when bond a is a single bond, then Z is $-C(R^{36})_2-$, $-C(=R^{32})-$, $-N(R^{35})-$, or $-O-$, wherein each R^{36} is independently hydrogen or R^{31} ; and

R^{35} is hydrogen, C_{1-6} alkyl, $-C(O)R$, $-S(O)_2R$, $-C(O)OR$, $-C(O)N(R)_2$, $-S(O)_2OR$, or $-S(O)_2N(R)_2$;

and when bond a is a double bond, then Z is $-C(R^{36})=$ or $-N=$.

[Claim 33]

The compound according to claim 32, its tautomers or a pharmaceutically acceptable salt thereof, wherein bond a is a single bond; and Z is $-C(R^{36})_2$.

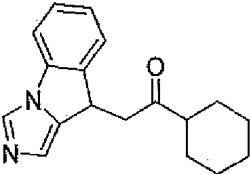
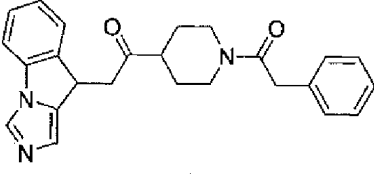
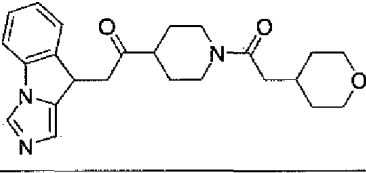
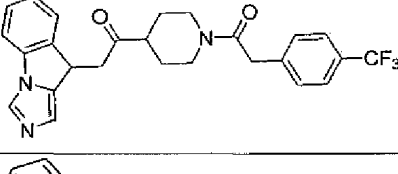
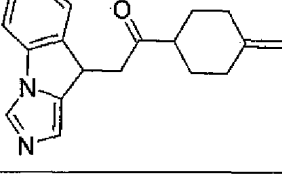
[Claim 34]

The compound according to any of claims 28 – 32, its tautomers or a pharmaceutically acceptable salt thereof, in which V and X are N, W and Z are CH and Y is C.

[Claim 35]

A compound that is

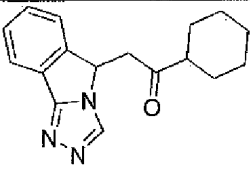
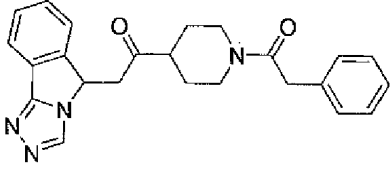
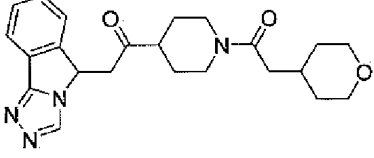
[Chemical formula 48]

27		1-cyclohexyl-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-one,
28		1-(4-(2-(9H-imidazo[1,5-a]indol-9-yl)acetyl)piperidin-1-yl)-2-phenylethan-1-one,
29		1-(4-(2-(9H-imidazo[1,5-a]indol-9-yl)acetyl)piperidin-1-yl)-2-(tetrahydro-2H-pyran-4-yl)ethan-1-one,
30		1-(4-(2-(9H-imidazo[1,5-a]indol-9-yl)acetyl)piperidin-1-yl)-2-(4-(trifluoromethyl)phenyl)ethan-1-one,
31		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(4-methylenecyclohexyl)ethan-1-one,

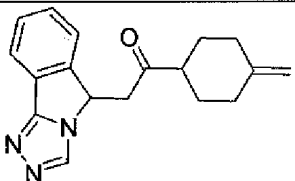
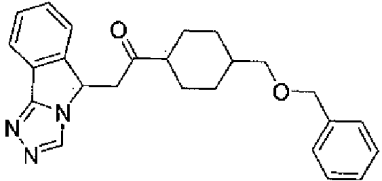
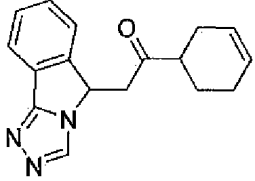
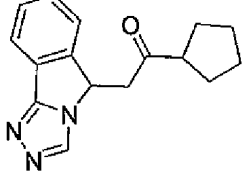
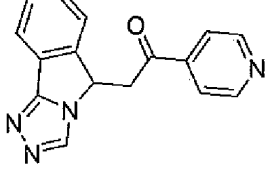
32		1-(4-((benzyloxy)methyl)cyclohexyl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-one,
33		1-(cyclohex-3-en-1-yl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-one,
34		1-cyclopentyl-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-one,

[Chemical formula 49]

35		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(pyridin-4-yl)ethan-1-one,
36		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(1,4-dioxaspiro[4.5]decan-8-yl)ethan-1-one,
37		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(thiazol-5-yl)ethan-1-one,
38		1-(2,4-dimethylfuran-3-yl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-one,
39		1-(3-chlorophenyl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-one,

43		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-cyclohexylethan-1-one,
44		1-(4-(2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)acetyl)piperidin-1-yl)-2-phenylethan-1-one,
45		1-(4-(2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)acetyl)piperidin-1-yl)-2-(tetrahydro-2H-pyran-4-yl)ethan-1-one,

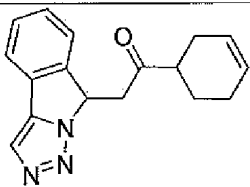
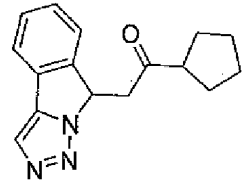
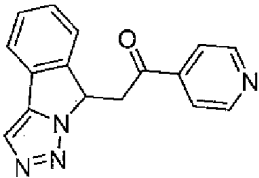
[Chemical formula 50]

46		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-(4-methylenecyclohexyl)ethan-1-one,
47		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-(4-((benzyloxy)methyl)cyclohexyl)ethan-1-one,
48		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-(cyclohex-3-en-1-yl)ethan-1-one,
49		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-cyclopentylethan-1-one,
50		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-(pyridin-4-yl)ethan-1-one,

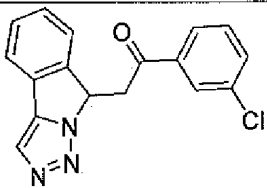
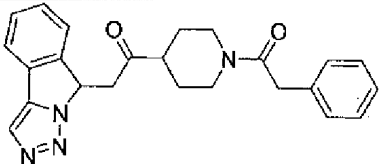
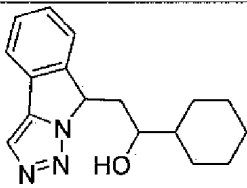
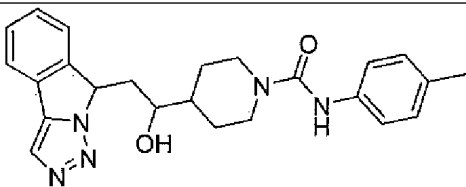
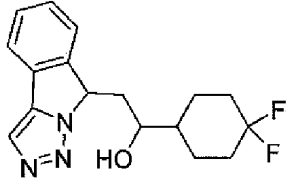
51		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-(3-chlorophenyl)ethan-1-one,
53		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-cyclohexylethan-1-one,
54		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(4,4-difluorocyclohexyl)ethan-1-one,

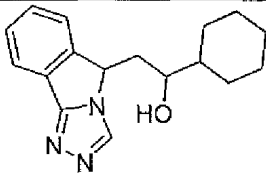
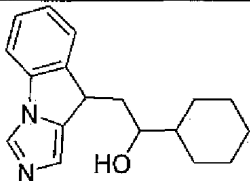
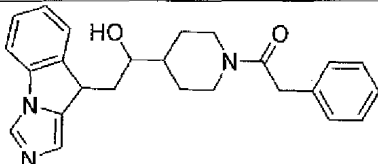
[Chemical formula 51]

55		1-(4-(2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)acetyl)piperidin-1-yl)-2-phenylethan-1-one,
56		1-(4-(2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)acetyl)piperidin-1-yl)-2-(tetrahydro-2H-pyran-4-yl)ethan-1-one,
57		1-(4-(2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)acetyl)piperidin-1-yl)-2-(4-(trifluoromethyl)phenyl)ethan-1-one,
58		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(4-methylenecyclohexyl)ethan-1-one,
59		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(4-((benzyloxy)methyl)cyclohexyl)ethan-1-one,

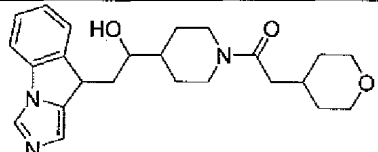
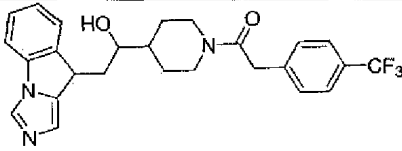
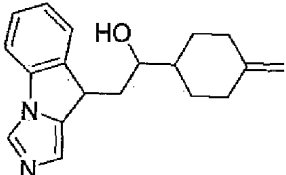
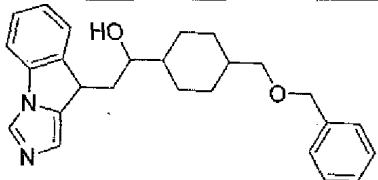
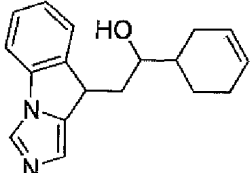
60		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(cyclohex-3-en-1-yl)ethan-1-one,
61		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-cyclopentylethan-1-one,
62		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(pyridin-4-yl)ethan-1-one,

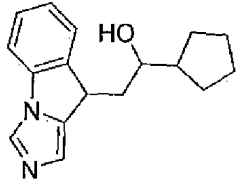
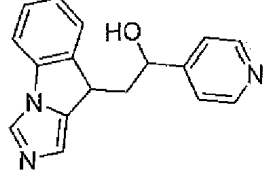
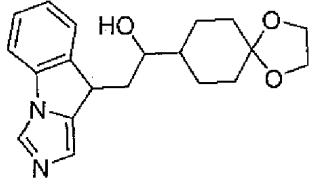
[Chemical formula 52]

63		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(3-chlorophenyl)ethan-1-one,
64		1-(4-(2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)acetyl)piperidin-1-yl)-2-phenylethan-1-one,
66		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-cyclohexylethan-1-ol,
67		4-(2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-hydroxyethyl)-N-(p-tolyl)piperidine-1-carboxamide,
68		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(4,4-difluorocyclohexyl)ethan-1-ol,

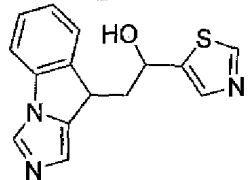
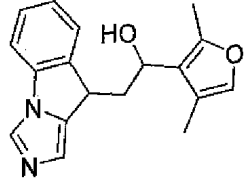
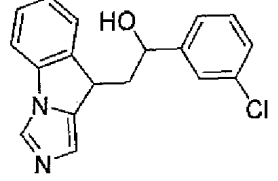
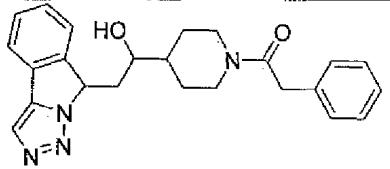
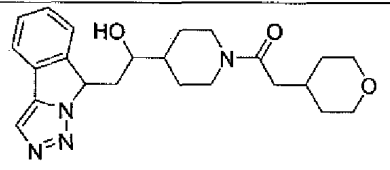
69		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-cyclohexylethan-1-ol,
70		1-cyclohexyl-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-ol,
71		1-(4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)piperidin-1-yl)-2-phenylethan-1-one,

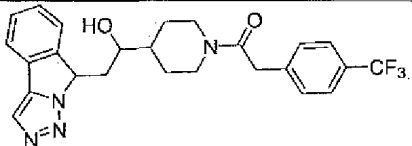
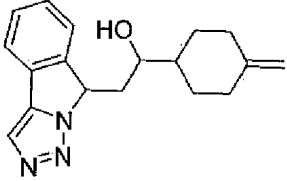
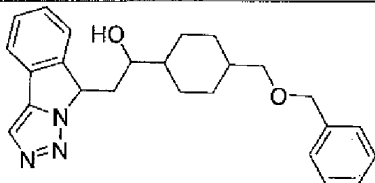
[Chemical formula 53]

72		1-(4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)piperidin-1-yl)-2-(tetrahydro-2H-pyran-4-yl)ethan-1-one,
73		1-(4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)piperidin-1-yl)-2-(4-(trifluoromethyl)phenyl)ethan-1-one,
74		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(4-methylenecyclohexyl)ethan-1-ol,
75		1-(4-((benzyloxy)methyl)cyclohexyl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-ol,
76		1-(cyclohex-3-en-1-yl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-ol,

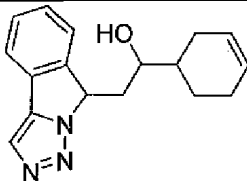
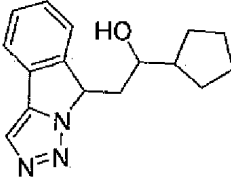
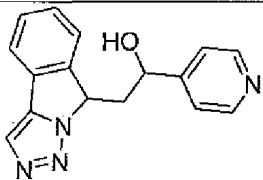
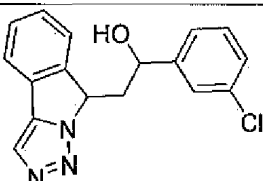
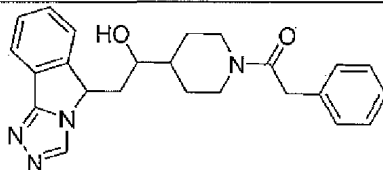
77		1-cyclopentyl-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-ol,
78		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(pyridin-4-yl)ethan-1-ol,
79		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(1,4-dioxaspiro[4.5]decan-8-yl)ethan-1-ol,

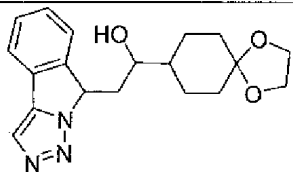
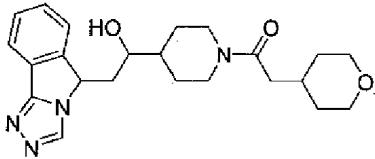
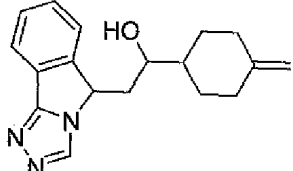
[Chemical formula 54]

80		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(thiazol-5-yl)ethan-1-ol,
81		1-(2,4-dimethylfuran-3-yl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-ol,
82		1-(3-chlorophenyl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-ol,
83		1-(4-(2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-hydroxyethyl)piperidin-1-yl)-2-phenylethan-1-one,
84		1-(4-(2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-hydroxyethyl)piperidin-1-yl)-2-(tetrahydro-2H-pyran-4-yl)ethan-1-one,

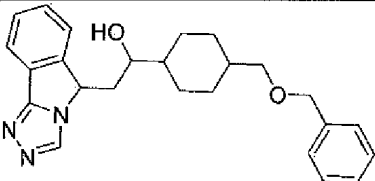
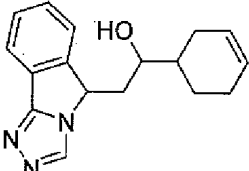
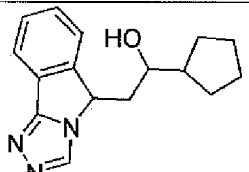
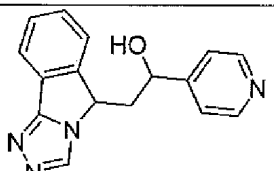
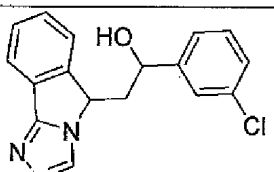
85		1-(4-(2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-hydroxyethyl)piperidin-1-yl)-2-(4-(trifluoromethyl)phenyl)ethan-1-one,
86		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(4-methylenecyclohexyl)ethan-1-ol,
87		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(4-((benzyloxy)methyl)cyclohexyl)ethan-1-ol,

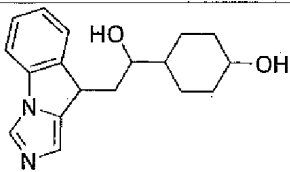
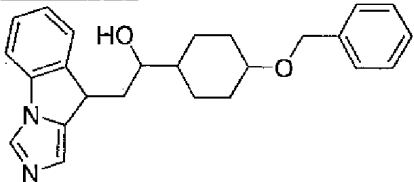
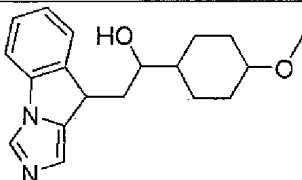
[Chemical formula 55]

88		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(cyclohex-3-en-1-yl)ethan-1-ol,
89		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-cyclopentylethan-1-ol,
90		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(pyridin-4-yl)ethan-1-ol,
91		2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-(3-chlorophenyl)ethan-1-ol,
92		1-(4-(2-(8H-[1,2,3]triazolo[5,1-a]isoindol-8-yl)-1-hydroxyethyl)piperidin-1-yl)-2-phenylethan-1-one,

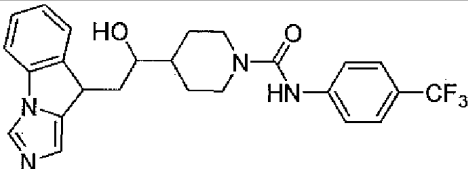
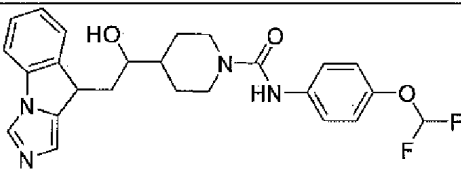
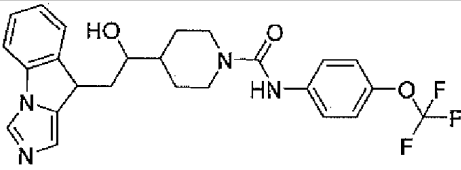
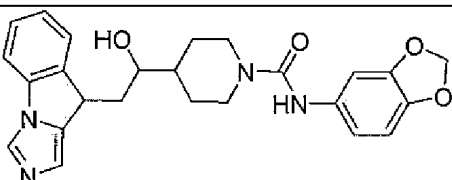
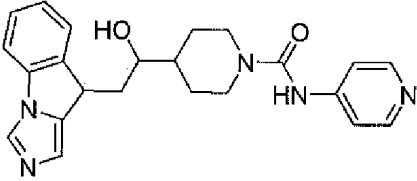
93		1-(4-(2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-hydroxyethyl)piperidin-1-yl)-2-phenylethan-1-one,
94		1-(4-(2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-hydroxyethyl)piperidin-1-yl)-2-(tetrahydro-2H-pyran-4-yl)ethan-1-one,
95		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-(4-methylenecyclohexyl)ethan-1-ol,

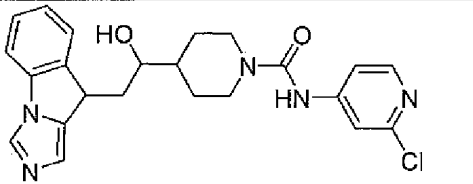
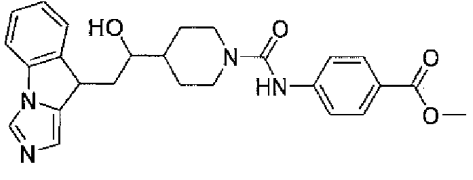
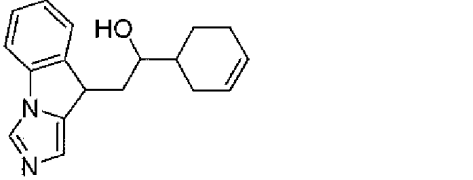
[Chemical formula 56]

96		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-(4-((benzyloxy)methyl)cyclohexyl)ethan-1-ol,
97		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-(cyclohex-3-en-1-yl)ethan-1-ol,
98		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-cyclopentylethan-1-ol,
99		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-(pyridin-4-yl)ethan-1-ol,
100		2-(5H-[1,2,4]triazolo[3,4-a]isoindol-5-yl)-1-(3-chlorophenyl)ethan-1-ol,

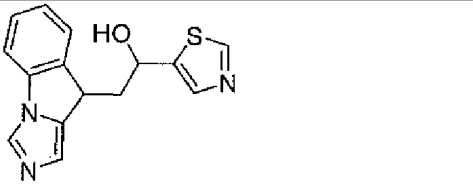
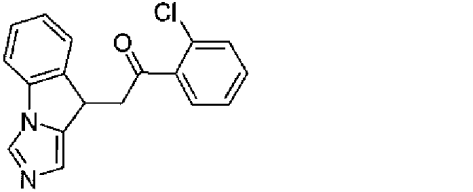
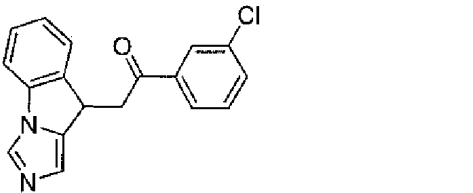
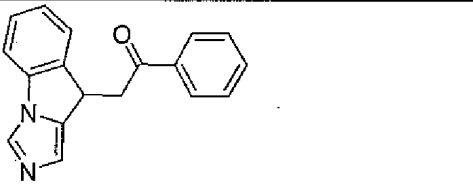
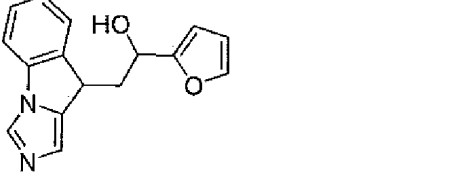
102		4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)cyclohexan-1-ol,
103		1-(4-(benzyloxy)cyclohexyl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-ol,
104		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(4-methoxycyclohexyl)ethan-1-ol,

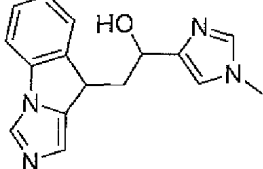
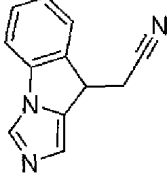
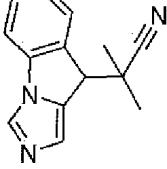
[Chemical formula 57]

105		4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)-N-(4-(trifluoromethyl)phenyl)piperidine-1-carboxamide,
106		N-(4-(difluoromethoxy)phenyl)-4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)piperidine-1-carboxamide,
107		4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)-N-(4-(trifluoromethoxy)phenyl)piperidine-1-carboxamide,
108		N-(benzo[d][1,3]dioxol-5-yl)-4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)piperidine-1-carboxamide,
109		4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)-N-(pyridin-4-yl)piperidine-1-carboxamide,

110		N-(2-chloropyridin-4-yl)-4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)piperidine-1-carboxamide,
111		methyl 4-(4-(1-hydroxy-2-(9H-imidazo[1,5-a]indol-9-yl)ethyl)piperidine-1-carboxamido)benzoate,
112		1-(cyclohex-3-en-1-yl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-ol,

[Chemical formula 58]

113		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(thiazol-5-yl)ethan-1-ol,
114		1-(2-chlorophenyl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-one,
115		1-(3-chlorophenyl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-one,
116		2-(9H-imidazo[1,5-a]indol-9-yl)-1-phenylethan-1-one,
117		1-(furan-2-yl)-2-(9H-imidazo[1,5-a]indol-9-yl)ethan-1-ol,

118		2-(9H-imidazo[1,5-a]indol-9-yl)-1-(1-methyl-1H-imidazol-4-yl)ethan-1-ol,
119		2-(9H-imidazo[1,5-a]indol-9-yl)acetonitrile, or
120		2-(9H-imidazo[1,5-a]indol-9-yl)-2-methylpropanenitrile.

[Claim 36]

A pharmaceutical composition comprising a compound according to any one of claims 1 - 34, its tautomers or a pharmaceutically acceptable salt thereof, or a compound of claim 35, and a pharmaceutically acceptable diluent, excipient, or carrier.

[Claim 37]

A pharmaceutical composition according to claim 36 for treating immunosuppression in a subject in need thereof.

[Claim 38]

The pharmaceutical composition of claim 37, wherein the immunosuppression is associated with an infectious disease, or cancer.

[Claim 39]

The pharmaceutical composition of claim 38, wherein the infectious disease is a viral infection selected from the group consisting of: influenza, hepatitis C virus (HCV), human papilloma virus (HPV), cytomegalovirus (CMV), Epstein-Barr virus (EBV), poliovirus, varicella zoster virus, coxsackie virus, human immunodeficiency virus (HIV).

[Claim 40]

A composition comprising a compound according to any one of claims 1-34, its tautomers or a pharmaceutically acceptable salt thereof, or a compound of claim 35 and a pharmaceutically acceptable diluent, excipient, or carrier for inhibiting IDO in a cell.