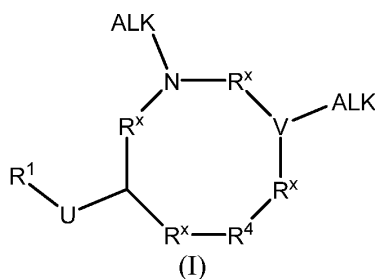


WHAT IS CLAIMED IS:

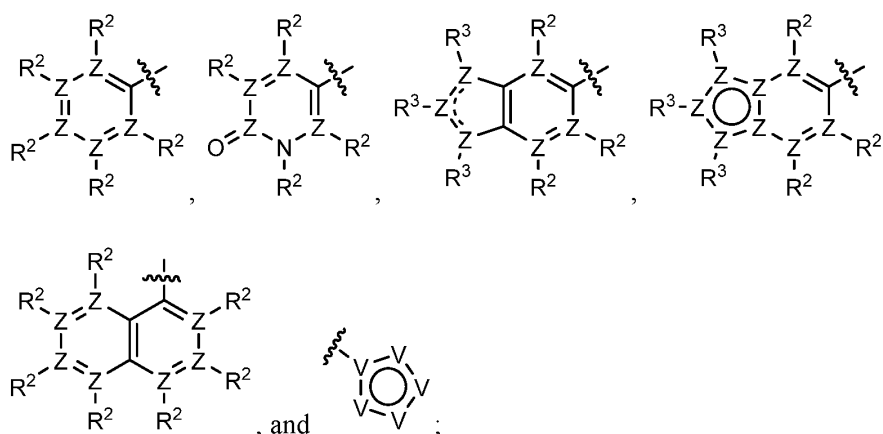
1. A compound of formula I:



or a pharmaceutically acceptable salt or solvate thereof,

wherein

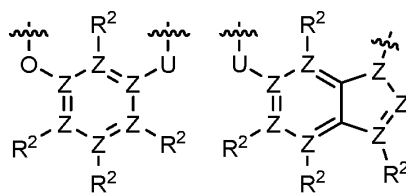
R¹ is selected from the group consisting of



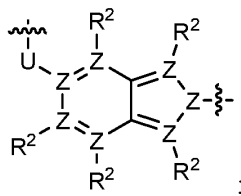
each R² is independently selected from the group consisting of absent, halogen, -NHR³, -OR³, -R³, -C(O)R³, -CO₂R³, C(O)N(R³)₂, -CH₂C(O)N(R³)₂, -S(O)₂R³, and -CN;

or two instances of R² are taken together with the atoms on which they are attached to form a heterocyclyl or heteroaryl ring;

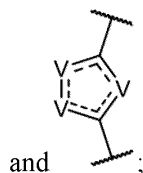
each R³ is independently selected from absent, -H, oxo, ALK, phenyl, heterocyclyl, and heteroaryl;



R^4 is selected from the group consisting of



U is selected from the group consisting of -C(O)-, alkylene, -O-, -N(R³)-, -C(O)O-, -C(O)N(R³)-,



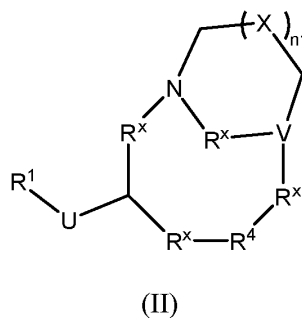
each R^x is independently selected from alkylene,

each V is independently selected from -C(R³)-, -N(R³)-, -N-, and -O-; and

Z is C or N ,

wherein ALK is unsubstituted alkyl or substituted alkyl, or two instances of ALK may be joined together with their intervening atoms to form a cycloalkyl or heterocyclyl ring.

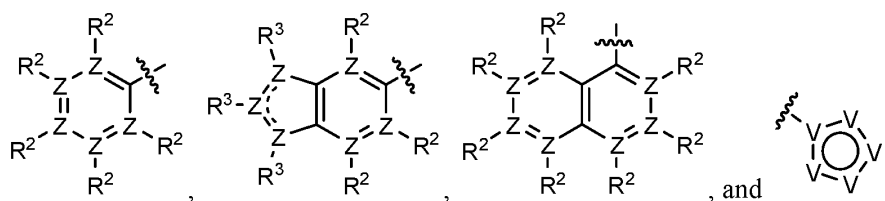
2. A compound of formula II:



or a pharmaceutically acceptable salt or solvate thereof,

wherein

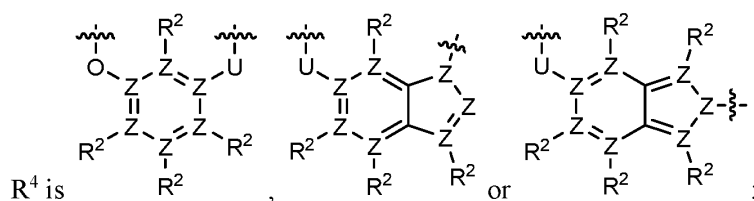
R¹ is selected from the group consisting of ALK, cycloalkyl, heterocyclyl,



each R^2 is independently selected from the group consisting of absent, halogen, $-OR^3$, $-R^3$, $-CO_2R^3$, $C(O)N(R^3)_2$, $-CH_2C(O)N(R^3)_2$, $-S(O)_2R^3$, and $-CN$;

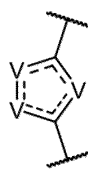
or two instances of R^2 are taken together with the atoms on which they are attached to form a heterocyclyl or heteroaryl ring;

each R^3 is independently selected from absent, $-H$, ALK , phenyl, and heteroaryl;



X is selected from the group consisting of $-CH_2-$, $-CHF-$, $-CF_2-$;

each U is indendently selected from the group consisting of $-O-$, $-N(R^3)-$, $-C(O)O-$, $-C(O)N(R^3)-$,



, $-C(O)-$, $-O-N=C(H)-$ and alkylene;

each R^x is independently selected from alkylene;

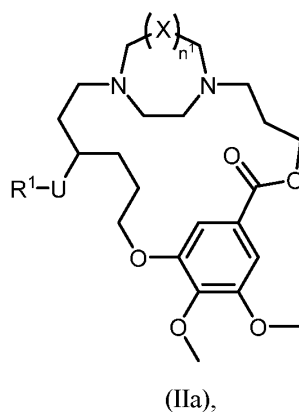
each V is independently selected from $-C(R^3)-$, $-N(R^3)-$, $-N=$, and $-O-$;

each Z is independently C or N ; and

n^1 is a number of 0 or 1,

wherein ALK is unsubstitued alkyl or substituted alkyl, or two instances of ALK may be joined together with their intervening atoms to form a cycloalkyl or heterocyclyl ring.

3. The compound according to claim 2, of formula IIa:

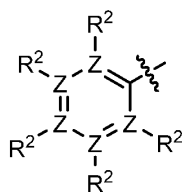


or a pharmaceutically acceptable salt or solvate thereof

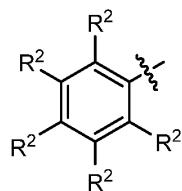
wherein

$X = CH_2, CHF, CF_2$.

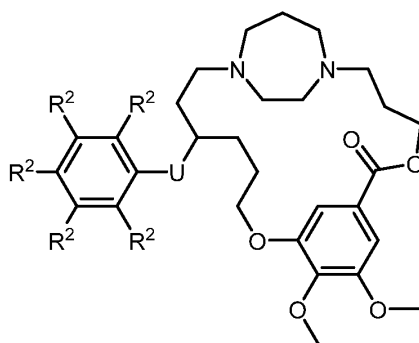
4. The compound of any of claims 1-3, wherein R^1 is



5. The compound of claims 4, wherein R^1 is



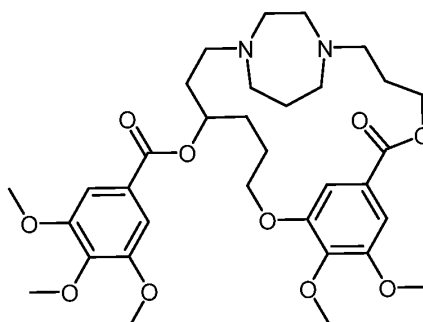
6. The compound of any of claims 1-5, wherein the compound is a compound of formula (IIb):



(IIb),

or a pharmaceutically acceptable salt or solvate thereof.

7. The compound of any of claims 1-6, wherein U is -C(O)O-.
8. The compound according to claim 2, of formula IIa1:



(IIa1)

or a pharmaceutically acceptable salt or solvate thereof.

9. The compound according to any one of the claims 2-8, selected from the group consisting of:
 - (12S)-74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate
 - (12R)-74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate
 - 16,16-difluoro-74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate
 - (12S)-16,16-difluoro-74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate
 - (12R)-16,16-difluoro-74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate
 - N-(74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl)-3,4,5-trimethoxybenzamide

74,75-dimethoxy-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

(12S)-74,75-dimethoxy-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

(12R)-74,75-dimethoxy-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

74,75-dimethoxy-5-methyl-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

(12S)-74,75-dimethoxy-5-methyl-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

(12R)-74,75-dimethoxy-5-methyl-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

(11R)-74,75-dimethoxy-6-oxo-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotridecaphane-11-yl 3,4,5-trimethoxybenzoate

(10S)-14-chloro-2-oxo-11H-3-aza-1(6,1)-indazola-7(1,4)-diazepanacyclotridecaphane-10-yl 3,4,5-trimethoxybenzoate

(10R)-14-chloro-2-oxo-11H-3-aza-1(6,1)-indazola-7(1,4)-diazepanacyclotridecaphane-10-yl 3,4,5-trimethoxybenzoate

(12S)-6-oxo-5,8-dioxa-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

(12R)-6-oxo-5,8-dioxa-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

(12S)-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl benzoate

(12R)-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl benzoate

74,75-dichloro-6-oxo-5,8-dioxa-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl
3,4,5-trimethoxybenzoate

(12S)-74,75-dichloro-6-oxo-5,8-dioxa-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-
12-yl 3,4,5-trimethoxybenzoate

(12R)-74,75-dichloro-6-oxo-5,8-dioxa-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-
12-yl 3,4,5-trimethoxybenzoate

75-carbamoyl-74-chloro-6-oxo-5,8-dioxa-1(1,4)-diazepana-7(1,3)-
benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

(11Z,16E,10S)-14-chloro-2-oxo-12H-3-aza-1(6,2)-indazola-7(1,4)-
diazepanacyclotridecaphane-10-yl 3,4,5-trimethoxybenzoate

(11Z,16E,10R)-14-chloro-2-oxo-12H-3-aza-1(6,2)-indazola-7(1,4)-
diazepanacyclotridecaphane-10-yl 3,4,5-trimethoxybenzoate

(12S)-74-carbamoyl-75-chloro-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-
benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

(12R)-74-carbamoyl-75-chloro-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-
benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

74-bromo-75-chloro-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-
benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

75-chloro-74-cyano-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-
benzenacyclotetradecaphane-12-yl 3,4,5-trimethoxybenzoate

(12R)-74,75-dimethoxy-6-oxo-5,8-dioxa-1(1,4)-diazepana-7(1,3)-
benzenacyclotetradecaphane-12-yl benzoate

(12R)-74,75-dimethoxy-6-oxo-5,8-dioxa-1(1,4)-diazepana-7(1,3)-
benzenacyclotetradecaphane-12-yl benzoate

(12S)-74,75-dimethoxy-6-oxo-5,8-dioxa-1(1,4)-diazepana-7(1,3)-
benzenacyclotetradecaphane-12-yl benzoate

(Z)-benzaldehyde O-(74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl) oxime

12-hydroxy-74,75-dimethoxy-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphan-6-one

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-hydroxybenzoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-fluorobenzoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-isopropoxybenzoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-(trifluoromethyl)benzoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-(methylsulfonyl)benzoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-phenoxybenzoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 2-fluorobenzoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-bromo-3-cyanobenzoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-methyl-5-(trifluoromethyl)benzoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 2-fluoro-4-methoxybenzoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-methoxy-2-(trifluoromethoxy)benzoate

- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl picolinate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl nicotinate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl pyrazine-2-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 6-hydroxynicotinate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl quinoline-5-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl oxazole-4-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 1H-1,2,3-triazole-4-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl acetate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl cyclopropanecarboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-methylbutanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4,4,4-trifluorobutanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl cyclohexanecarboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 1-methylpiperidine-4-carboxylate

- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,3-dimethylcyclobutane-1-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 2-(oxetan-3-yl)acetate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl (1R,5S,6r)-3-oxabicyclo[3.1.0]hexane-6-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 5-oxopyrrolidine-3-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 1-benzyl-5-oxopyrrolidine-3-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-methoxycyclohexane-1-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 2,6-difluorobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-(trifluoromethoxy)benzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-cyanobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 2-oxo-1,2,3,4-tetrahydroquinoline-6-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-(difluoromethoxy)benzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,5-dichlorobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4-dichlorobenzoate

- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 2,3-dichlorobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 2-chloro-6-fluoro-3-methylbenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-fluoro-5-(trifluoromethyl)benzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-fluoro-3-(trifluoromethyl)benzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-cyano-3-fluorobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-(trifluoromethyl)benzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,5-difluorobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4-difluorobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-cyano-4-fluorobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-cyanobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-chloro-4-fluorobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 1-methyl-1H-benzo[d]imidazole-5-carboxylate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-(oxazol-5-yl)benzoate

- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4,5-dichloro-2-fluorobenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3,4,5-triethoxybenzoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-methoxypropanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-(1H-pyrazol-1-yl)propanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-cyanopropanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-cyanobutanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-acetamidobutanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-(1H-tetrazol-1-yl)propanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-(dimethylamino)-4-oxobutanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-acetamidopropanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-(methylamino)-4-oxobutanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-(1H-1,2,4-triazol-1-yl)propanoate
- 74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-morpholino-4-oxobutanoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-(4-fluorophenoxy)propanoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4,4-difluorocyclohexane-1-carboxylate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 4-(trifluoromethyl)cyclohexane-1-carboxylate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-(2,5-dioxopyrrolidin-1-yl)propanoate

74,75-dimethoxy-6-oxo-5,8-dioxo-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl 3-methoxycyclohexane-1-carboxylate

74,75-dimethoxy-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl benzoate

(E)-benzaldehyde O-(74,75-dimethoxy-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl) oxime

(E)-benzaldehyde O-((12R)-74,75-dimethoxy-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl) oxime

(E)-benzaldehyde O-((12S)-74,75-dimethoxy-6-oxo-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphane-12-yl) oxime

12-hydroxy-74,75-dimethoxy-8-oxa-5-aza-1(1,4)-diazepana-7(1,3)-benzenacyclotetradecaphan-6-one,

and pharmaceutically acceptable salts or solvates thereof.

10. The compound according to any one of the claims preceding, wherein the compound comprises only one chiral center.
11. The compound according to claim 10, wherein the compound is racemic mixture containing 'R' isomer and 'S' isomer.

12. The compound according to claim 10, wherein the compound is 'R' isomer.
13. The compound according to claim 10, wherein the compound is 'S' isomer.
14. The compound according to any one of the claims 1-9, wherein the compound comprises more than one chiral center.
15. The compound according to claim 14, wherein the chiral centers comprise 'R' or 'S' configurations independently.
16. The compound according to claim 14, wherein the chiral centers comprise the same configuration.
17. A pharmaceutical composition comprising a compound according to any one of the claims preceding and at least one pharmaceutically acceptable excipient.
18. The pharmaceutical composition according to claim 17, further comprising an adenosine receptor antagonist.
19. The pharmaceutical composition according to claim 18, wherein the adenosine receptor antagonist is an A2A or A2B receptor antagonist.
20. The pharmaceutical composition according to claim 18, wherein the adenosine receptor antagonist is selected from:

5-bromo-2,6-di-(1H-pyrazol-1-yl)pyrimidin-4-amine;

(S)-7-(5-methylfuran-2-yl)-3-((6-(((tetrahydrofuran-3-yl)oxy)methyl)pyridin-2-yl)methyl)-3H-[1,2,3]triazolo[4,5-d]pyrimidin-5-amine;

6-(2-chloro-6-methylpyridin-4-yl)-5-(4-fluorophenyl)-1,2,4-triazin-3-amine;

3-(2-amino-6-(1-((6-(2-hydroxypropan-2-yl)pyridin-2-yl)methyl)-1H-1,2,3-triazol-4-yl)pyrimidin-4-yl)-2-methylbenzonitrile;

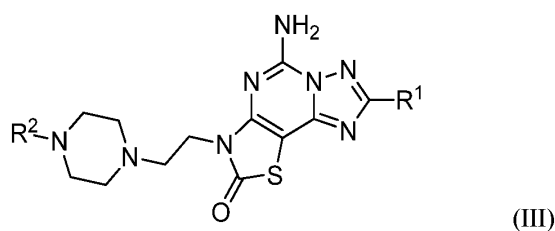
2-(2-furanyl)-7-(2-(4-(4-(2-methoxyethoxy)phenyl)-1-piperazinyl)ethyl)-7H-pyrazolo(4,3-e)(1,2,4)triazolo(1,5-c)pyrimidine-5-amine;

3-(4-amino-3-methylbenzyl)-7-(2-furyl)-3H-(1,2,3)triazolo(4,5-d)pyrimidine-5-amine;

and

4-hydroxy-N-(4-methoxy-7-morpholinobenzo[d]thiazol-2-yl)-4-methylpiperidine-1-carboxamide.

21. The pharmaceutical composition according to claim 18, wherein the adenosine receptor antagonist is a compound of Formula (III):



or a pharmaceutically acceptable salt or solvate thereof, wherein:

R^1 = 5- or 6-membered heteroaryl or 5- or 6-membered aryl, wherein heteroaryl or aryl groups are optionally substituted by one or more substituent selected from C1-C6 alkyl and halo;

R^2 = 6-membered aryl or 6-membered heteroaryl,

wherein heteroaryl or aryl groups are optionally substituted by one or more substituent selected from halo, alkyl, heterocyclyl, alkoxy, cycloalkyloxy, heterocyclyloxy, carbonyl, alkylcarbonyl, aminocarbonyl, hydroxycarbonyl, heterocyclylcarbonyl, alkylsulfoxide, alkylsulfonyl, aminosulfonyl, heterocyclylsulfonyl, alkylsulfonimidoyl, carbonylamino, sulfonylamino and alkylsulfonealkyl;

said substituents being optionally substituted by one or more substituent selected from oxo, halo, hydroxy, cyano, alkyl, alkenyl, aldehyde, heterocyclylalkyl, hydroxyalkyl, dihydroxyalkyl, hydroxyalkylaminoalkyl, aminoalkyl, alkylaminoalkyl, dialkylaminoalkyl, (heterocyclyl)(alkyl)aminoalkyl, heterocyclyl, heteroaryl, alkylheteroaryl, alkyne, alkoxy, amino, dialkylamino, aminoalkylcarbonylamino, aminocarbonylalkylamino, (aminocarbonylalkyl)(alkyl)amino, alkenylcarbonylamino, hydroxycarbonyl, alkyloxycarbonyl, aminocarbonyl, aminoalkylaminocarbonyl, alkylaminoalkylaminocarbonyl, dialkylaminoalkylaminocarbonyl, heterocyclylalkylaminocarbonyl,

(alkylaminoalkyl)(alkyl)aminocarbonyl, alkylaminoalkylcarbonyl, dialkylaminoalkylcarbonyl, heterocyclylcarbonyl, alkenylcarbonyl, alkynylcarbonyl, alkylsulfoxide, alkylsulfoxidealkyl, alkylsulfonyl and alkylsulfonealkyl;

or the heteroaryl or aryl groups are optionally substituted with two substituents that form together with the atoms to which they are attached a 5- or 6-membered aryl ring, a 5- or 6-membered heteroaryl ring, a 5- or 6-membered cycloalkyl ring or a 5- or 6-membered heterocyclyl ring; optionally substituted by one or more substituent selected from oxo, halo, hydroxy, cyano, alkyl, alkenyl, aldehyde, heterocyclylalkyl, hydroxyalkyl, dihydroxyalkyl, hydroxyalkylaminoalkyl, aminoalkyl, alkylaminoalkyl, dialkylaminoalkyl, (heterocyclyl)(alkyl)aminoalkyl, heterocyclyl, heteroaryl, alkylheteroaryl, alkyne, alkoxy, amino, dialkylamino, aminoalkylcarbonylamino, aminocarbonylalkylamino, (aminocarbonylalkyl)(alkyl)amino, alkenylcarbonylamino, hydroxycarbonyl, alkyloxycarbonyl, aminocarbonyl, aminoalkylaminocarbonyl, alkylaminoalkylaminocarbonyl, dialkylaminoalkylaminocarbonyl, heterocyclylalkylaminocarbonyl, (alkylaminoalkyl)(alkyl)aminocarbonyl, alkylaminoalkylcarbonyl, dialkylaminoalkylcarbonyl, heterocyclylcarbonyl, alkenylcarbonyl, alkynylcarbonyl, alkylsulfoxide, alkylsulfoxidealkyl, alkylsulfonyl and alkylsulfonealkyl.

22. A method of inhibiting ENT1 in a patient need thereof, comprising: administering to said patient an effective amount of a compound according to any one of the claims 1 to 16.
23. A method of treating cancer in a patient need thereof, comprising: administering to said patient an effective amount of a compound according to any one of the claims 1 to 16.
24. A method of treating cancer in a patient need thereof, comprising: administering to said patient a combination of a compound according to any one of the claims 1 to 16 and an adenosine receptor antagonist.

25. The method according to claim 24, wherein the compound according to any one of the claims 1 to 16 is administered prior to, concomitant with, or subsequent to administration of the adenosine receptor antagonist.
26. The method according to any one of the claims 24-25, wherein the adenosine receptor antagonist is an A2A or A2B receptor antagonist.
27. The method according to claim 24, wherein the adenosine receptor antagonist is selected from:

5-bromo-2,6-di-(1H-pyrazol-1-yl)pyrimidin-4-amine;

(S)-7-(5-methylfuran-2-yl)-3-((6-((tetrahydrofuran-3-yl)oxy)methyl)pyridin-2-yl)methyl)-3H-[1,2,3]triazolo[4,5-d]pyrimidin-5-amine;

6-(2-chloro-6-methylpyridin-4-yl)-5-(4-fluorophenyl)-1,2,4-triazin-3-amine;

3-(2-amino-6-(1-((6-(2-hydroxypropan-2-yl)pyridin-2-yl)methyl)-1H-1,2,3-triazol-4-yl)pyrimidin-4-yl)-2-methylbenzonitrile;

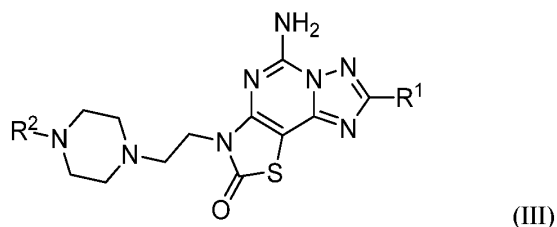
2-(2-furanyl)-7-(2-(4-(4-(2-methoxyethoxy)phenyl)-1-piperazinyl)ethyl)-7H-pyrazolo(4,3-e)(1,2,4)triazolo(1,5-c)pyrimidine-5-amine;

3-(4-amino-3-methylbenzyl)-7-(2-furyl)-3H-(1,2,3)triazolo(4,5-d)pyrimidine-5-amine;

and

4-hydroxy-N-(4-methoxy-7-morpholinobenzo[d]thiazol-2-yl)-4-methylpiperidine-1-carboxamide.

28. The method according to claim 24, wherein the adenosine receptor antagonist is a compound of Formula (III):



or a pharmaceutically acceptable salt or solvate thereof, wherein:

R^1 = 5- or 6-membered heteroaryl or 5- or 6-membered aryl, wherein heteroaryl or aryl groups are optionally substituted by one or more substituent selected from C1-C6 alkyl and halo;

R^2 = 6-membered aryl or 6-membered heteroaryl,

wherein heteroaryl or aryl groups are optionally substituted by one or more substituent selected from halo, alkyl, heterocyclyl, alkoxy, cycloalkyloxy, heterocyclyloxy, carbonyl, alkylcarbonyl, aminocarbonyl, hydroxycarbonyl, heterocyclylcarbonyl, alkylsulfoxide, alkylsulfonyl, aminosulfonyl, heterocyclylsulfonyl, alkylsulfonimidoyl, carbonylamino, sulfonylamino and alkylsulfonealkyl;

said substituents being optionally substituted by one or more substituent selected from oxo, halo, hydroxy, cyano, alkyl, alkenyl, aldehyde, heterocyclylalkyl, hydroxyalkyl, dihydroxyalkyl, hydroxyalkylaminoalkyl, aminoalkyl, alkylaminoalkyl, dialkylaminoalkyl, (heterocyclyl)(alkyl)aminoalkyl, heterocyclyl, heteroaryl, alkylheteroaryl, alkyne, alkoxy, amino, dialkylamino, aminoalkylcarbonylamino, aminocarbonylalkylamino, (aminocarbonylalkyl)(alkyl)amino, alkenylcarbonylamino, hydroxycarbonyl, alkyloxycarbonyl, aminocarbonyl, aminoalkylaminocarbonyl, alkylaminoalkylaminocarbonyl, dialkylaminoalkylaminocarbonyl, heterocyclylalkylaminocarbonyl, (alkylaminoalkyl)(alkyl)aminocarbonyl, alkylaminoalkylcarbonyl, dialkylaminoalkylcarbonyl, heterocyclylcarbonyl, alkenylcarbonyl, alkynylcarbonyl, alkylsulfoxide, alkylsulfoxidealkyl, alkylsulfonyl and alkylsulfonealkyl;

or the heteroaryl or aryl groups are optionally substituted with two substituents that form together with the atoms to which they are attached a 5- or 6-membered aryl ring, a 5- or 6-membered heteroaryl ring, a 5- or 6-membered cycloalkyl ring or a 5- or 6-membered heterocyclyl ring; optionally substituted by one or more substituent selected from oxo, halo, hydroxy, cyano, alkyl, alkenyl, aldehyde, heterocyclylalkyl, hydroxyalkyl, dihydroxyalkyl, hydroxyalkylaminoalkyl, aminoalkyl, alkylaminoalkyl, dialkylaminoalkyl, (heterocyclyl)(alkyl)aminoalkyl, heterocyclyl, heteroaryl, alkylheteroaryl, alkyne, alkoxy, amino,

dialkylamino, aminoalkylcarbonylamino, aminocarbonylalkylamino, (aminocarbonylalkyl)(alkyl)amino, alkenylcarbonylamino, hydroxycarbonyl, alkyloxycarbonyl, aminocarbonyl, aminoalkylaminocarbonyl, alkylaminoalkylaminocarbonyl, dialkylaminoalkylaminocarbonyl, heterocyclalkylaminocarbonyl, (alkylaminoalkyl)(alkyl)aminocarbonyl, alkylaminoalkylcarbonyl, dialkylaminoalkylcarbonyl, heterocyclcarbonyl, alkenylcarbonyl, alkynylcarbonyl, alkylsulfoxide, alkylsulfoxidealkyl, alkylsulfonyl and alkylsulfonealkyl.

29. A kit of parts comprising:

(a) a first part comprising an effective amount of a compound according to any one of the claims 1 to 16; and

(b) a second part comprising an effective amount of an adenosine receptor antagonist.

30. The kit of parts according to claim 29, wherein the adenosine receptor antagonist is an A2A or A2B receptor antagonist.

31. The kit of parts according to claim 29, wherein the adenosine receptor antagonist is selected from:

5-bromo-2,6-di-(1H-pyrazol-1-yl)pyrimidin-4-amine;

(S)-7-(5-methylfuran-2-yl)-3-((6-([tetrahydrofuran-3-yl]oxy)methyl)pyridin-2-yl)methyl)-3H-[1,2,3]triazolo[4,5-d]pyrimidin-5-amine;

6-(2-chloro-6-methylpyridin-4-yl)-5-(4-fluorophenyl)-1,2,4-triazin-3-amine;

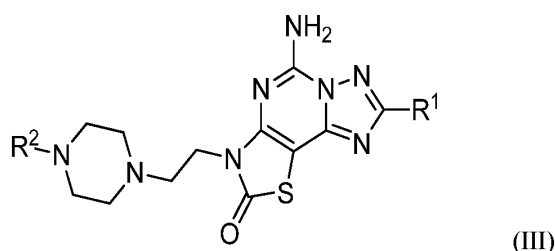
3-(2-amino-6-(1-((6-(2-hydroxypropan-2-yl)pyridin-2-yl)methyl)-1H-1,2,3-triazol-4-yl)pyrimidin-4-yl)-2-methylbenzonitrile;

2-(2-furanyl)-7-(2-(4-(4-(2-methoxyethoxy)phenyl)-1-piperazinyl)ethyl)-7H-pyrazolo(4,3-e)(1,2,4)triazolo(1,5-c)pyrimidine-5-amine;

3-(4-amino-3-methylbenzyl)-7-(2-furyl)-3H-(1,2,3)triazolo(4,5-d)pyrimidine-5-amine;
and

4-hydroxy-N-(4-methoxy-7-morpholinobenzo[d]thiazol-2-yl)-4-methylpiperidine-1-carboxamide.

32. The kit of parts according to claim 29, wherein the adenosine receptor antagonist is a compound of Formula (III):



or a pharmaceutically acceptable salt or solvate thereof, wherein:

R^1 = 5- or 6-membered heteroaryl or 5- or 6-membered aryl, wherein heteroaryl or aryl groups are optionally substituted by one or more substituent selected from C1-C6 alkyl and halo;

R^2 = 6-membered aryl or 6-membered heteroaryl,

wherein heteroaryl or aryl groups are optionally substituted by one or more substituent selected from halo, alkyl, heterocyclyl, alkoxy, cycloalkyloxy, heterocyclyloxy, carbonyl, alkylcarbonyl, aminocarbonyl, hydroxycarbonyl, heterocyclylcarbonyl, alkylsulfoxide, alkylsulfonyl, aminosulfonyl, heterocyclylsulfonyl, alkylsulfonimidoyl, carbonylamino, sulfonylamino and alkylsulfonealkyl;

said substituents being optionally substituted by one or more substituent selected from oxo, halo, hydroxy, cyano, alkyl, alkenyl, aldehyde, heterocyclylalkyl, hydroxyalkyl, dihydroxyalkyl, hydroxyalkylaminoalkyl, aminoalkyl, alkylaminoalkyl, dialkylaminoalkyl, (heterocyclyl)(alkyl)aminoalkyl, heterocyclyl, heteroaryl, alkylheteroaryl, alkyne, alkoxy, amino, dialkylamino, aminoalkylcarbonylamino, aminocarbonylalkylamino, (aminocarbonylalkyl)(alkyl)amino, alkenylcarbonylamino, hydroxycarbonyl, alkylloxycarbonyl,

aminocarbonyl, aminoalkylaminocarbonyl, alkylaminoalkylaminocarbonyl, dialkylaminoalkylaminocarbonyl, heterocyclalkylaminocarbonyl, (alkylaminoalkyl)(alkyl)aminocarbonyl, alkylaminoalkylcarbonyl, dialkylaminoalkylcarbonyl, heterocyclcarbonyl, alkenylcarbonyl, alkynylcarbonyl, alkylsulfoxide, alkylsulfoxidealkyl, alkylsulfonyl and alkylsulfonealkyl;

or the heteroaryl or aryl groups are optionally substituted with two substituents that form together with the atoms to which they are attached a 5- or 6-membered aryl ring, a 5- or 6-membered heteroaryl ring, a 5- or 6-membered cycloalkyl ring or a 5- or 6-membered heterocycl ring; optionally substituted by one or more substituent selected from oxo, halo, hydroxy, cyano, alkyl, alkenyl, aldehyde, heterocyclalkyl, hydroxyalkyl, dihydroxyalkyl, hydroxyalkylaminoalkyl, aminoalkyl, alkylaminoalkyl, dialkylaminoalkyl, (heterocycl)(alkyl)aminoalkyl, heterocycl, heteroaryl, alkylheteroaryl, alkyne, alkoxy, amino, dialkylamino, aminoalkylcarbonylamino, aminocarbonylalkylamino, (aminocarbonylalkyl)(alkyl)amino, alkenylcarbonylamino, hydroxycarbonyl, alkyloxycarbonyl, aminocarbonyl, aminoalkylaminocarbonyl, alkylaminoalkylaminocarbonyl, dialkylaminoalkylaminocarbonyl, heterocyclalkylaminocarbonyl, (alkylaminoalkyl)(alkyl)aminocarbonyl, alkylaminoalkylcarbonyl, dialkylaminoalkylcarbonyl, heterocyclcarbonyl, alkenylcarbonyl, alkynylcarbonyl, alkylsulfoxide, alkylsulfoxidealkyl, alkylsulfonyl and alkylsulfonealkyl.