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(54) **INHIBITOR COMPOUNDS OF 11-BETA-HYDROXYSTEROID DEHYDROGENASE TYPE 1**

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(57) **ABSTRACT**

The compounds of formula (I) are derived from perhydroquinoline and perhydroisoquinoline and are useful as active pharmaceutical ingredients for the prophylaxis or treatment of diseases caused by 11-beta-hydroxysteroid dehydrogenase type 1 (11-beta-HSD1) enzyme-associated disorders, such as glaucoma, elevated ocular pressure, metabolic disorders, obesity, metabolic syndrome, dyslipidemia, hypertension, diabetes, atherosclerosis, Cushing's syndrome, psoriasis, rheumatoid arthritis, cognitive disorders, Alzheimer's disease or neurodegeneration.

INHIBITOR COMPOUNDS OF 11-BETA-HYDROXYSTEROID DEHYDROGENASE TYPE 1

[0001] This application is a continuation-in-part of U.S. Ser. No. 13/375,389, a §371 national stage of PCT International Application No. PCT/ES2010/000258, filed Jun. 4, 2010, claiming priority of Spanish Patent Application No. P200901402, filed Jun. 4, 2009, the content of each of which is hereby incorporated by reference into the present application.

[0002] The present invention relates to perhydroquinoline and perhydroisoquinoline derivatives and methods of treating certain diseases using such compounds.

STATE OF THE ART

[0003] Glucocorticoids (cortisol in humans, corticosterone in mice and rats) are an important adrenocorticosteroid group regulating many metabolic and homeostatic processes and form a key element of stress response. Glucocorticoids act through the intracellular glucocorticoid receptors and, in some tissues, through mineralocorticoid receptors, both being nuclear transcription factors. The action of glucocorticoids on the target tissues depends not only on circulating steroid concentrations and the cellular expression of the receptors, but also on the intracellular enzymes which critically determine up to what point the glucocorticoids will have active access to the receptors. The 11-beta-hydroxysteroid dehydrogenases (11-beta-HSD) catalyze the interconversion between the main active 11-hydroxy-glucocorticoid (cortisol in man) and its inactive 11-keto metabolites (cortisone in man).

[0004] The 11-beta-hydroxysteroid dehydrogenase type 1 (11-beta-HSD1) enzyme reconverts inactive glucocorticoids into active ones, thus playing an important role in modulating cellular agonist concentration and, therefore, in activating corticosteroid receptors in the target tissues. It has been described that the overexpression of 11-beta-HSD1 in mice adipocytes leads to visceral obesity and to the phenotype similar to that of the metabolic syndrome. Collectively, these data significantly confirm the important role of 11-beta-HSD1 in inducing obesity and the disequilibrium of glucose homeostasis and lipid parameters. Therefore, the selective inhibition of this enzyme could reduce the levels of blood glucose in type 2 diabetes patients, normalize the elevated lipid parameters and/or reduce the weight of obese subjects.

[0005] The first pharmacological indication that the inhibition of 11-beta-HSD1 in man could have beneficial effects has been achieved by using carbenoxolone, an anti-ulcer drug which inhibits both 11-beta-HSD1 and the similar 11-beta-HSD2 enzyme. Treatment with carbenoxolone increases the sensitivity to insulin, which indicates that the inhibition of 11-beta-HSD1 can reduce the levels of cortisol in the cells and therefore minimize some of its damaging effects.

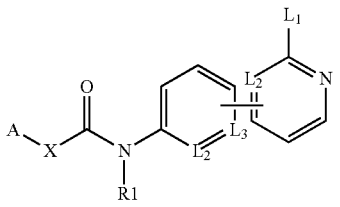
[0006] Studies conducted with the non-specific carbenoxolone inhibitor clearly show the importance of developing 11-beta-HSD1 specific inhibitors. Inhibition of the 11-beta-HSD2 enzyme is poorly tolerated and increases blood pressure. In contrast, inhibition of 11-beta-HSD1 would be well tolerated because it has been observed that the 11-beta-HSD1 knockout mice are healthy and resist hyperglycemia caused by obesity or stress (cf. Kotelevtsev et al., Proc. Natl. Acad. Sci. USA 1997, vol. 94, pp. 14924-14929). Other studies indicate that 11-beta-HSD1 inhibitors can also be beneficial for reducing high blood pressure (cf. Masuzaki et al., J. Clin. Invest. 2003, vol. 112, pp. 83-90), for reducing intraocular pressure (cf. Rauz et al., Q J Med 2003, vol. 96, pp. 481-490), for improving cognitive capacity (cf. Sandeep et al., Proc Natl Acad. Sci. USA 2004, vol. 101, pp. 6734-6739) or for improving deficiencies associated with Alzheimer's disease. Overall, inhibition of 11-beta-HSD1 can be a safe and effective strategy for treating symptoms of glaucoma, diabetes, obesity and other diseases.

[0007] Glucocorticoids increase the risk of glaucoma by increasing intraocular pressure when they are exogenously administered and in certain conditions of increased production such as Cushing's syndrome. The increase of intraocular pressure induced by corticosteroids is caused by an increased resistance to the aqueous efflux due to changes induced by glucocorticoids.

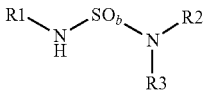
[0008] 11-beta-HSD1 is expressed in corneal epithelium basal cells and non-pigmented epithelial cells. The mRNA of the glucocorticoid receptor has been detected in the trabecular reticulum, whereas mRNA for the glucocorticoid receptor, the mineralocorticoid receptor and 11-beta-HSD1 was present in non-pigmented epithelial cells. The administration of carbenoxolone to patients resulted in a significant reduction in intraocular pressure (cf. Rauz et al., Invest. Ophtalmol. Vis. Sci. 2001, vol. 42, pp. 2037-2042), which suggests a role for HSD1 inhibitors in the treatment of glaucoma.

[0009] The expression of 11-beta-HSD isoenzymes in human and rodent eyes has been described (cf. Stokes et al., Invest Ophthalmol V is Sci. 2000, vol. 41, pp. 1629-1638), particularly 11-beta-HSD1 in ciliary epithelial cells, which suggests the possibility of a role in producing aqueous humor and in regulating intraocular pressure. In aqueous humor, cortisol concentrations are approximately 14 times greater than those of cortisone. This suggests to a large extent predominant 11-beta-reductase HSD1 activity. In a double blinded controlled study with glaucoma patients it was observed that treatment with carbenoxolone significantly reduces intraocular pressure therefore 11-beta-HSD1 inhibitors can represent a therapeutic strategy suitable for treating glaucoma.

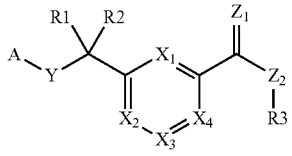
[0010] Document WO2007026920 describes N-arylamide compounds and related compounds as Rho kinase (ROCK) inhibitors, as well as pharmaceutical compositions and the use thereof in treating diseases related to ROCK.



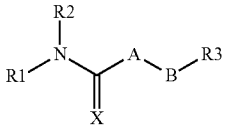
[0011] Document WO2006106423 describes N-pyridyl heterocyclylsulfonamide compounds and the use thereof as 11-beta-HSD1 modulators.



[0012] Document WO2006048330 describes N-benzylsulfonamide compounds and related derivatives as 11-beta-HSD1 inhibitors, pharmaceutical compositions and the use thereof in therapy.



[0013] Document WO2003045367 describes compounds derived from pyridylalkylurea, some of which are structurally similar to those of the present invention but with a different use.

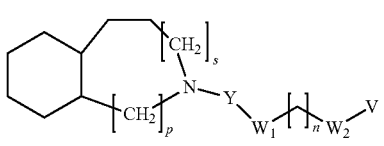


[0014] Nevertheless, providing new 11-beta-HSD1 inhibitor compounds is desirable.

DESCRIPTION OF THE INVENTION

[0015] The present invention provides new compounds derived from perhydroquinoline and perhydroisoquinoline of formula (I) which are effective as 11-beta-HSD1 inhibitors and have selectivity for 11-beta-HSD1 with respect to 11-beta-HSD2.

[0016] Thus, a first aspect of the invention refers to compounds of formula (I)



(I)

[0017] and pharmaceutically acceptable salts thereof, wherein:

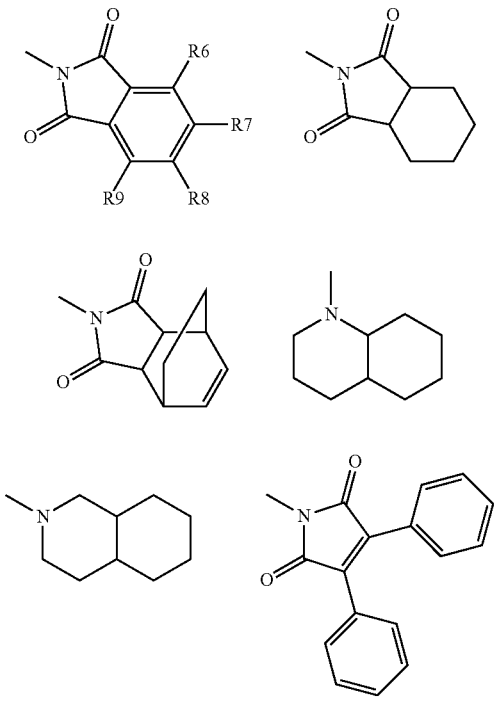
[0018] s and p are an integer selected in an opposite manner between 0 and 1, such that when s is 1, p is 0 (to form a perhydroquinoline) and when s is 0, p is 1 (to form a perhydroisoquinoline).

[0019] Y is a biradical selected from CO, CS and SO_2,

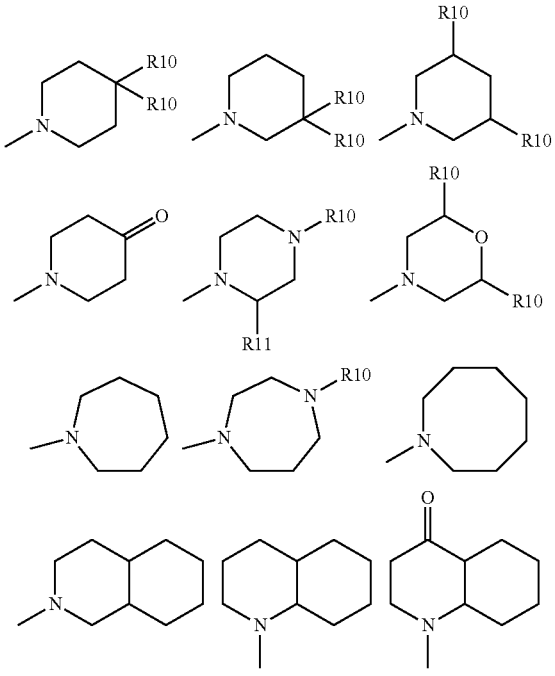
[0020] W1 and W2 can be independently a bond or a biradical selected from O, S and NR1, wherein R1 is optionally H, C_{1-4} alkyl or C_{3-10} cycloalkyl,

[0021] n is an integer selected from 0, 1, 2, 3 and 4,

[0022] V is a radical selected from —CO-T, —CS-T and —SO_2-T, or a radical selected from:



[0023] T is a group selected from NR2R3, R2, OR2 and SR2; or a group selected from



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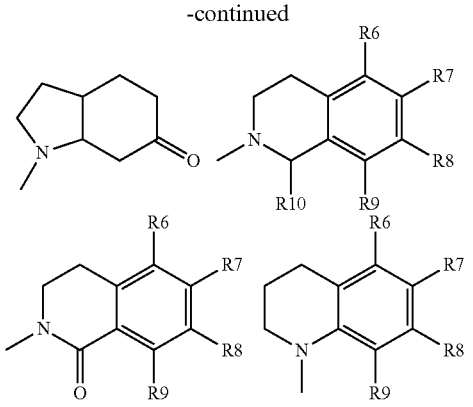


[0024] wherein R2 and R3 are independently selected from H, COR4, SO_2R4, C_{1-4} alkyl, aryl, benzyl, phenethyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{3-10} cycloalkyl or heterocycle

[0025] wherein when R2 or R3 is an alkyl or an alkenyl these can be optionally substituted with one or several substituents independently selected from F, OR4, NR4R5, COOR4, CONR4R5, C_{3-10} cycloalkyl, aryl and heterocycle;

[0026] wherein when R2 or R3 is an aryl, a benzyl, a phenethyl, a cycloalkyl or a heterocycle, these can be optionally substituted with one or several substituents independently selected from NH_2, F, Cl, CN, NO_2, COOH, R4, COOR4, OR4, OCF_3, SH, SR4, CONR4R5, SO_2NR4R5, COR4, NR1COR4, OCOR4, SOR4, SO_2R4 and heterocycle,

[0027] wherein when R2 or R3 is a cycloalkyl this can be optionally substituted with one or several benzene rings fused with the cycloalkyl, the benzene could be optionally substituted with one or several substituents independently selected from alkyl, alkoxide or halogen,

[0028] wherein R4 and R5 are independently selected from H, C_{1-4} alkyl, aryl, benzyl, phenethyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{3-10} cycloalkyl and heterocycle

[0029] wherein optionally R4 and R5 can be bound to one another forming a 3 to 8 membered cycle.

[0030] R6, R7, R8 and R9 are independently selected from H, OR4, F and Cl,

[0031] R10 is independently selected from H, OH, F, C_{1-4} alkyl, COOR11, COR11, phenyl, benzyl, benzhydryl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{3-10} cycloalkyl and heterocycle, and wherein the alkyl, phenyl, benzyl, benzhydryl, cycloalkyl or heterocycle can be optionally substituted with one or several substituents independently selected from NH_2, F, Cl, NO_2, COOH, COOR4, OR4, CF_3, SH, SR4, CONR4R5, SO_2NR4R5, COR4, NR1COR4, OCOR4, SOR4, SOR4 and C_{1-4} alkyl;

[0032] and R11 is selected from H, C_{1-4} alkyl, aryl and C_{3-10} cycloalkyl.

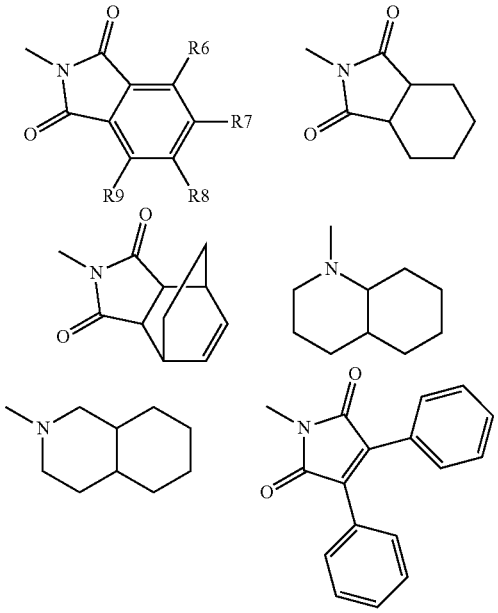
[0033] In a particular embodiment of the invention, s is 0 and p is 1. In another particular embodiment of the invention, s is 1 and p is 0.

[0034] In another particular embodiment of the invention, Y is selected from CO and SO_2.

[0035] In another particular embodiment of the invention, W1 and W2 are independently selected from a bond, S and NR1. In another particular embodiment of the invention, R1 is H.

[0036] In another particular embodiment of the invention, V is selected from —CO-T, —CS-T and —SO_2-T.

[0037] In another particular embodiment of the invention, V is selected from



[0038] In another particular embodiment of the invention, R2 and R3 are independently selected from H, COR4, SO_2R4, C_{1-4} alkyl, phenyl, naphthyl, benzyl, phenethyl, C_{2-4} alkenyl, C_{3-10} cycloalkyl, and heterocycle, particularly, 2-furanyl, 2-thiophenyl, 2-(1-methylindole), quinoline, isoquinoline, 2-benzofuranyl.

[0039] In another particular embodiment of the invention, when R2 or R3 are independently C_{1-4} alkyl or C_{2-4} alkenyl, R2 or R3 can be optionally substituted with one or several substituents independently selected from F, OR4, NR4R5, COOR4, CONR4R5, phenyl, C_{3-10} cycloalkyl, hexenyl, naphthyl and heterocycle, particularly pyridine, 3-(1-methylindole), 3-thiophenyl and 2-furanyl.

[0040] In another particular embodiment of the invention, when R2 or R3 are independently phenyl, benzyl, phenethyl or C_{3-10} cycloalkyl, R2 or R3 can be optionally substituted with one or several substituents independently selected from F, Cl and OR4.

[0041] In another particular embodiment of the invention, when R2 or R3 is a cycloalkyl this can be optionally substituted with one or several benzene rings fused with the cycloalkyl, the benzene could be optionally substituted with one or several substituents independently selected from alkyl, alkoxide or halogen.

[0042] In another particular embodiment of the invention, R4 and R5 are independently selected from C_{1-4} alkyl, benzyl, phenethyl and phenyl.

[0043] In another particular embodiment of the invention, R4 and R5 can be optionally bound to one another forming a 3 to 8 membered cycle.

[0044] In another particular embodiment of the invention, R6, R7, R8 and R9 are independently selected from H, OR4, F and Cl.

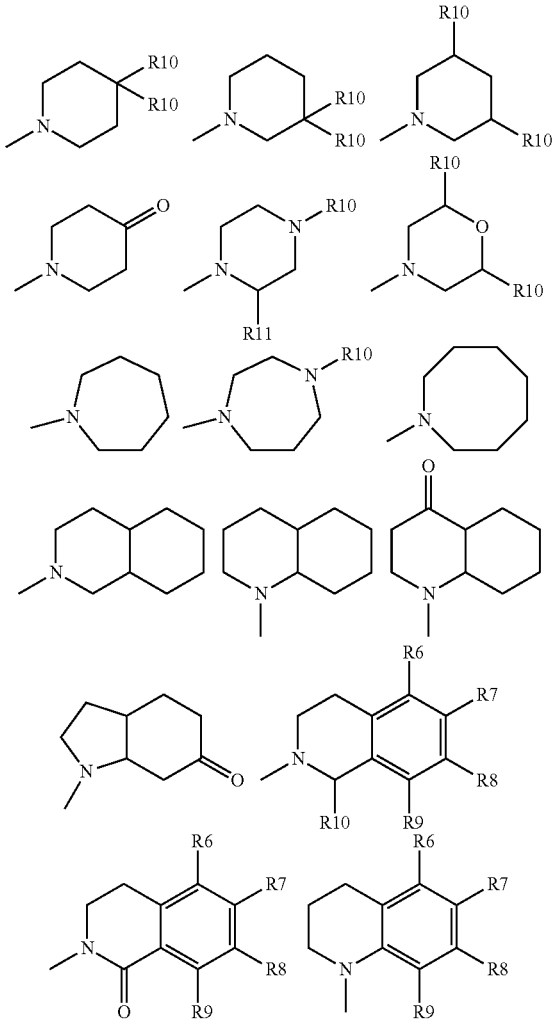
[0045] In another particular embodiment of the invention, R10 is selected from H, OH, F, C_{1-4} alkyl, COOR11, COR11, phenyl, benzyl and benzhydryl.

[0046] In another particular embodiment of the invention, R10 is selected from phenyl, benzyl and benzhydryl, all of them optionally substituted with one or several substituents, independently selected from F, OR4, CF_3, COR4 and C_{1-4} alkyl.

[0047] In another particular embodiment of the invention, R11 is selected from H and C_{3-10} cycloalkyl.

[0048] In another particular embodiment, T is selected from NR2R3, R2, OR2 and SR2.

[0049] In another particular embodiment, T is selected from



[0050] A second aspect of the present invention refers to a compound of formula (I) or a pharmaceutically acceptable salt thereof for use as a medicament, particularly for the prophylaxis or treatment of diseases caused by 11-beta-HSD1-associated disorders, particularly glaucoma, elevated ocular pressure, metabolic disorders, obesity, metabolic syndrome, dyslipidemia, hypertension, diabetes, particularly type II diabetes, atherosclerosis, Cushing's syndrome, psoriasis, rheumatoid arthritis, cognitive disorders, Alzheimer's disease or neurodegeneration, preferably for the prophylaxis or treatment of glaucoma or metabolic syndrome.

[0051] Another aspect of the present invention refers to the use of a compound of formula (I) or a pharmaceutically acceptable salt thereof in the manufacture of a medicaments intended for the prophylaxis or treatment of diseases caused by 11-beta-HSD1-associated disorders, particularly one of the disorders mentioned above.

[0052] Another aspect of the present invention refers to a method of prophylaxis or treatment of an individual who is suffering or is susceptible to suffering a disease caused by 11-beta-HSD1-associated disorders, particularly one of the disorders mentioned above, which comprises administrating to said individual a therapeutically effective amount of a compound of formula (I) or a pharmaceutically acceptable salt thereof together with one or more pharmaceutically acceptable excipients.

[0053] The compounds of formula (I) and the pharmaceutically acceptable salts thereof, particularly the compounds of formula (I) described as examples or as intermediates are preferred.

[0054] The compounds of the present invention can be used alone or in combination with one or more compounds which are useful for the prophylaxis or treatment of diseases such as glaucoma, elevated ocular pressure, metabolic disorders, such as obesity, metabolic syndrome, dyslipidemia, hypertension and/or diabetes, particularly type II diabetes, atherosclerosis, Cushing's syndrome, psoriasis, rheumatoid arthritis, cognitive disorders, Alzheimer's disease and/or neurodegeneration.

[0055] The term "C_{1-4} alkyl", alone or in combination, means a linear- or branched-chain alkyl group having 1 to 4 carbon atoms.

[0056] The terms "C_{2-4} alkenyl", and "C_{2-4} alkynyl", alone or in combination, mean a linear- or branched chain radical having 2 to 4 carbon atoms and having one or more unsaturated bonds.

[0057] The term "C_{3-10} cycloalkyl", alone or in combination, refers to a stable monocyclic, bicyclic or tricyclic radical of 3 to 10 members, which is saturated or partially saturated, and which only consists of carbon and hydrogen atoms. Examples of C_{3-10} cycloalkyl are the following: cyclopropyl, cyclopentyl, cyclohexyl, 1-cyclohexenyl, cycloheptyl, cyclooctyl, 1-tricyclo[3.3.1.$1^{3,7}$]decanyl, 2-tricyclo[3.3.1.$1^{3,7}$]decanyl and 2-bicyclo[2.2.1]heptanyl. Unless otherwise specifically established in the specification, the term "cycloalkyl" refers to that including cycloalkyl radicals which are optionally substituted with one or more substituents such as alkyl, halogen, hydroxyl, amino, cyano, nitro, alkoxyl, carboxyl, alkoxycarbonyl, phenyl, etc.

[0058] The term "aryl", alone or in combination refers to radicals of a single ring and multiple rings, including radicals of multiple rings containing separated and/or condensed aryl groups. The typical aryl groups contain 1 to 3 separated or condensed rings and from 6 to 18 carbon ring atoms, such as phenyl or naphthyl radicals, preferably a phenyl group optionally having one or several substituents, preferably from one to three, chosen independently from one another from halogen, trifluoromethyl, trifluoromethoxy, amino, alkyl, alkoxy, alkylcarbonyl, cyano, carbamoyl, alkoxycarbamoyl, methylenedioxy, carboxy, alkoxycarbonyl, aminocarbonyl, alkylaminocarbonyl, dialkylaminocarbonyl, hydroxy, nitro, alkyl-SO_2—, amino-SO_2—, cycloalkyl and the like. Phenyl or naphthyl is preferred, particularly phenyl optionally substituted from one to three times, preferably one or two times by substituents chosen independently from one another from

alkyl, halogen, alkoxy, trifluoromethoxy, nitro and trifluoromethyl. Phenyl is particularly preferred.

[0059] The terms "benzyl" and "phenethyl", can optionally have one or several substituents, chosen independently from one another from halogen, trifluoromethyl, trifluoromethoxy, amino, alkyl, alkoxy, alkylcarbonyl, cyano, carbamoyl, alkoxycarbamoyl, methylenedioxy, carboxy, alkoxycarbonyl, aminocarbonyl, alkylaminocarbonyl, dialkylaminocarbonyl, hydroxy, nitro, alkyl-SO_2—, amino-SO_2—, cycloalkyl and the like.

[0060] The term "heterocycle", alone or in combination, means a saturated, partially unsaturated or aromatic, 5 to 10 membered heterocycle, containing one or several heteroatoms chosen between nitrogen, oxygen and sulfur. For the purposes of this invention, the heterocycle can be a monocyclic, bicyclic or tricyclic ring system which can include condensed ring systems. The heterocycle can be substituted on one or several carbon atoms e.g. by halogen, alkyl, phenyl, alkoxy, oxo, etc. and/or on a secondary nitrogen atom (i.e., —NH—) by alkyl, cycloalkyl, aralkoxycarbonyl, alcanoyl, phenyl or phenylalkyl or on a tertiary nitrogen atom (i.e., =N—) by oxide, being especially preferred halogen, alkyl, cycloalkyl and alkoxy. Examples of heterocycle groups are pyrrolidinyl, piperidinyl, piperazinyl, azepine, morpholinyl, thiomorpholinyl, imidazolyl (e.g. imidazol-4-yl and 1-benzyloxycarbonylimidazol-4-yl), pyrazolyl, pyridyl, pyrazinyl, pyrimidinyl, hexahydropyrimidinyl, furyl, thienyl, thiazolyl, oxazolyl, indolyl (e.g. 2-indolyl), quinolyl (e.g. 2-quinolyl, 3-quinolyl and 1-oxide-2-quinolyl), isoquinolyl (e.g. 1-isoquinolyl and 3-isoquinolyl), tetrahydroquinolyl (e.g. 1,2,3,4-tetrahydro-2-quinolyl), 1,2,3,4-tetrahydrolsoquinolyl (e.g. 1,2,3,4-tetrahydro-1-oxoisoquinolyl)benzimidazoyl, benzothiazoyl and quinoxalinyl. Preferred examples are thiophenyl, quinolyl, piperidyl, morpholyl, thiomorpholyl, oxazolyl, pyridinyl, pyrimidinyl, pyrazolyl, imidazolyl and thiazolyl.

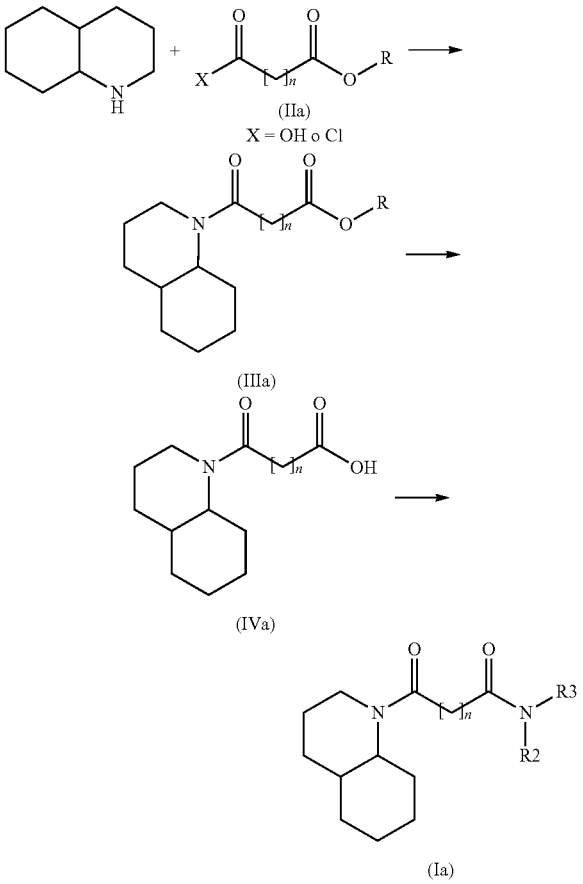
[0061] The term "pharmaceutically acceptable salts" means those salts which conserve the efficiency and the biological properties of the free bases or of the free acids and which are not disturbing in a biological sense or in any other sense.

[0062] According to the invention, the compounds of formula I and their pharmaceutically acceptable salts are useful for the prophylaxis or treatment of diseases caused by 11-beta-HSDl enzyme-associated disorders.

[0063] Unless defined otherwise, all the technical and scientific terms used herein have the same meaning as those commonly understood by a person skilled in the field of the invention. Methods and materials which are similar or equivalent to those described herein can be used in practicing the present invention. Throughout the description and claims the word "comprises" and its variants do not aim to exclude other technical features, additives, components, steps or stereoisomers of the compounds involved. For the persons skilled in the art, other objects, advantages and features of the invention will be inferred partially from the description and partially from practicing the invention.

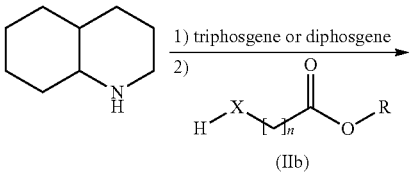
[0064] The compounds of formula (I) can be prepared following different methods known by any person skilled in the field of organic synthesis, particularly through the general processes shown in the following schemes. The starting materials for the preparative methods are commercially available or they can be prepared by means of methods of the literature. All of them started with perhydroquinoline but they are analogous for perhydroisoquinoline.

Scheme 1



[0065] According to this method an acid-ester (IIa) is treated with decahydroquinoline in the presence of a suitable coupling agent, such as for example the combination of 1-(3-dimethylaminopropyl)-3-ethylcarbodiimide EDC) and 1-hydroxybenzotriazole (HOBT), or by means of converting the acid to the corresponding acyl halide with a large variety of reagents such as thionyl chloride, sulfuryl chloride, oxalyl chloride, etc. In the presence of a tertiary base such as Et_3N (Elmore, Amino Acids Pep. Proteins 2001, vol. 32, pp. 107-162) for obtaining the amide-ester intermediate (IIIa). The diamide (Ia) is obtained by means of a prior saponification of the compound (IIIa) in aqueous medium with bases of the LiOH type, NaOH type, etc. and subsequent formation of the diamide with any of the methods previously described for the intermediate (IIIa).

Scheme 2



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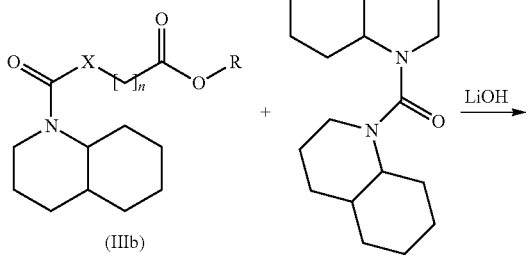

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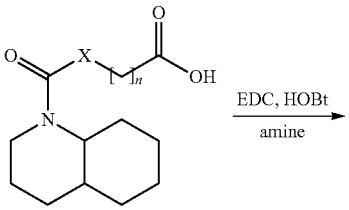
(Vb)



(IVb)

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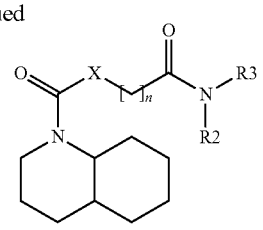
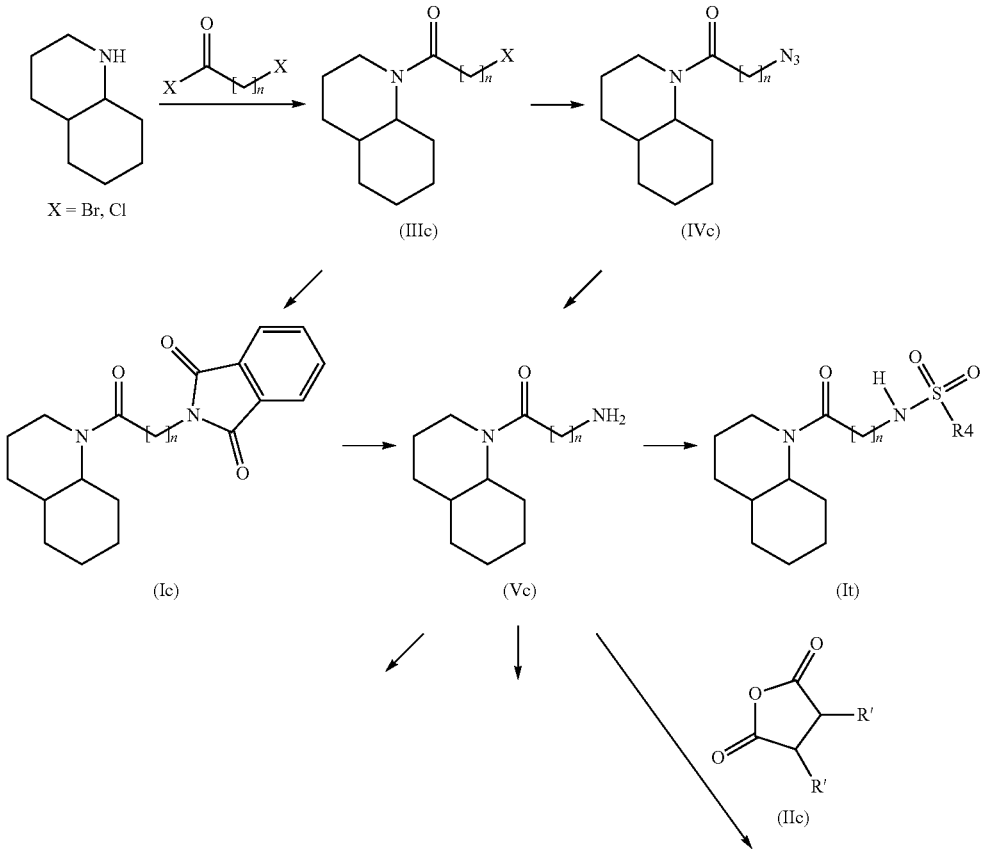


(Ib)

X: NR1, S or O

[0066] Scheme 2 shows the reaction of the decahydroquinoline with diphosgene or triphosgene and subsequent addition of amino-ester, hydroxy-ester or mercapto-ester type derivatives respectively providing the urea-ester, carbamate-ester or thiocarbamate-ester type intermediates. Subsequently a deprotection and coupling treatment similar to that described in scheme 1 for obtaining the amide derivatives is performed.

Scheme 3:



X = Br, Cl

(IIIc)

(IVc)

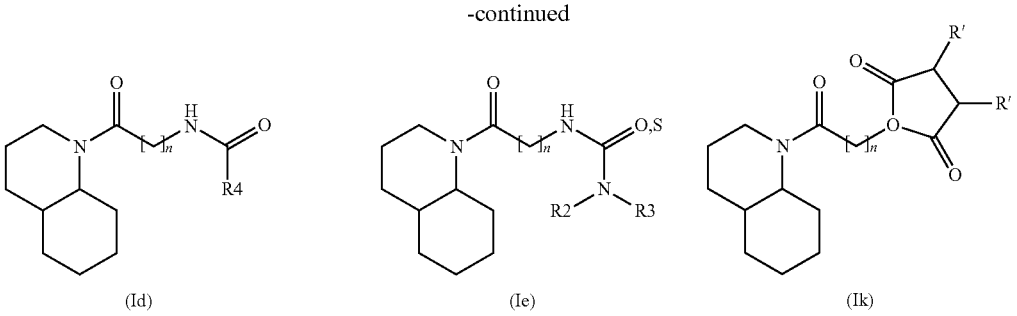
(Ic)

(Vc)

(It)

(IIc)

-continued

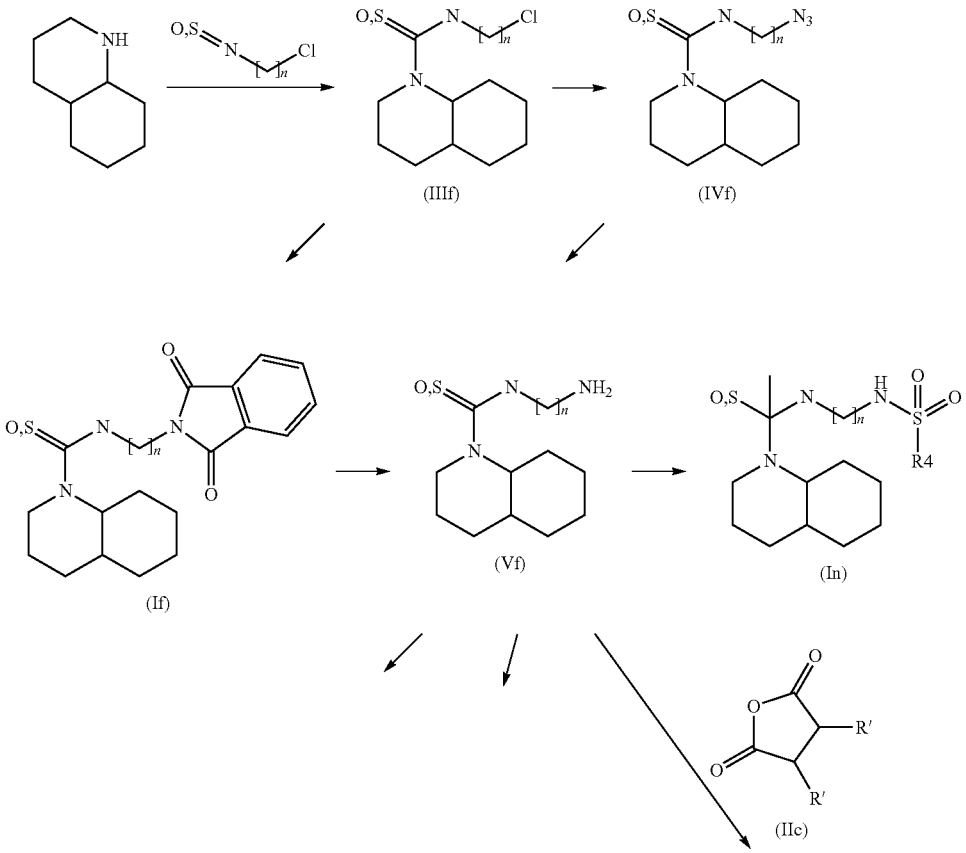


(Id) (Ie) (Ik)

[0067] Scheme 3 shows a method for preparing the compounds amido-succinimide (Ic and Ik), amido-amide (Id), amido-sulfonamide (It) and amido-urea or amido-thiourea (Ie) of the present invention. The intermediate amine (Vc) can be prepared by means of two alternative methods: on one hand Gabriel synthesis with potassium phthalimide from the intermediate (IIIc) providing the compound (Ic) and subsequent treatment of this compound with hydrazine under EtOH reflux yielding the mentioned amine (Vc), optionally the intermediate (IIIc) is reacted with sodium azide generating alkyl azide (IVc) which provides the amine (Vc) by reduction.

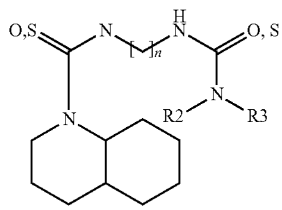
[0068] The intermediate amine (Vc) can be converted both into amide (Id) and into sulfamide (It) by any of the methods previously described or into urea or thiourea (Ie) by means of reaction with an isocyanate or thioisocyanate, respectively. The compound of formula (Ik) can easily be prepared by condensing an anhydride of a 1,2-dicarboxylic acid of general formula (IIc) and the intermediate amine (Vc) previously described.

Scheme 4

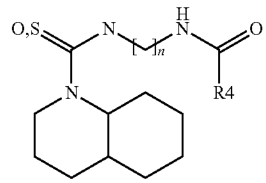


(IIIf) (IVf) (If) (Vf) (In) (IIc)

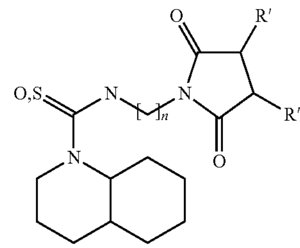
-continued



(Ig)



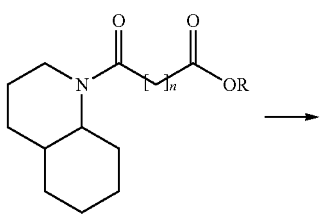
(Ih)



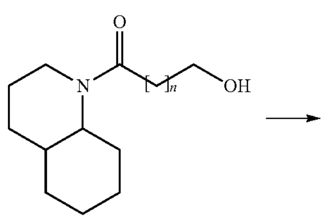
(Is)

[0069] Scheme 4 shows a method for preparing the compounds of the present invention applying a combination of methods previously described in the schemes above to provide ureas and thioureas from decahydroquinoline with different terminal functionalities already described previously in scheme 3.

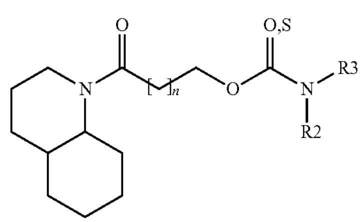
Scheme 5



(IIIa)



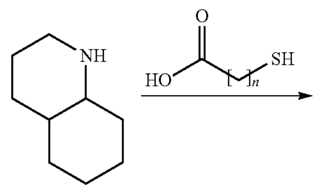
(IVj)

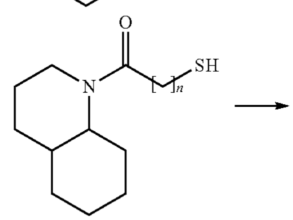


(Ij)

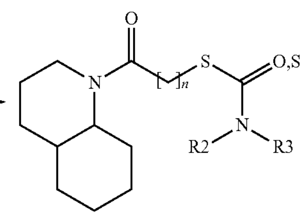
[0070] Scheme 5 shows a method for preparing the compounds of the present invention of amido-carbamate and amido-O-thiocarbamate type. The reduction of the intermediate ester (IIIa) to alcohol (IVj) by means of saponification of the ester, formation of a mixed anhydride and subsequent reduction thereof with sodium borohydride. The reaction of the alcohol (IVj) with isocyanate, thiocyanate, carbamoyl chloride or thiocarbamoyl chloride provides the carbamate or O-thiocarbamate (Ij).

Scheme 6





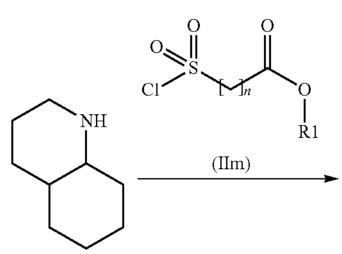
(IVp)

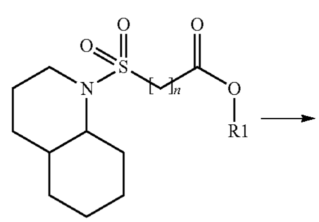


(Ip)

[0071] Scheme 6 shows a method for preparing the compounds of the present invention of amido-S-thiocarbamate and amido-dithiocarbamate type. After the formation of the intermediate amide (IVp) by methods previously described and its subsequent reaction with isocyanate, thiocyanate, carbamoyl chloride or thiocarbamoyl chloride provides S-thiocarbamate or dithiocarbamate (Ip).

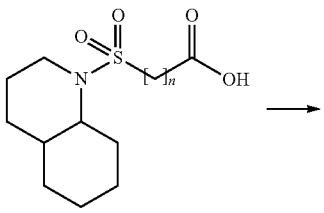
Scheme 7



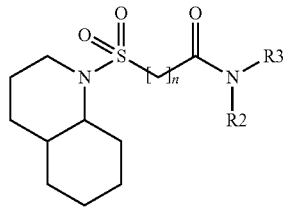


(IIIm)

-continued



(IVm)



(Im)

[0072] Scheme 7 shows a method for preparing the sulfonamide (Im) in which decahydroquinoline is reacted with the sulfonyl ester chloride (IIm). The deprotection and coupling treatment previously described in scheme 1 provides the sulfonamides (Im).

EXAMPLES

[0073] The following examples serve to better illustrate the invention but they must not be considered as limiting the same.

[0074] The nomenclature used in the present document is based on the Beilstein-Institut computer program known as AUTONOM (Automatic Nomenclature), which uses the systematic nomenclature of the IUPAC

ABBREVIATIONS

[0075] AcOEt ethyl acetate
[0076] Brine saturated NaCl solution
[0077] DCM dichloromethane
[0078] DMF dimethylformamide
[0079] DMSO dimethylsulfoxide
[0080] EDC 1-(3-dimethylaminopropyl)-3-ethylcarbodiimide
[0081] Et_3N triethylamine
[0082] EtOH ethanol
[0083] HOBT 1-hydroxybenzotriazole
[0084] HPLC High performance liquid chromatography
[0085] MeOH methanol
[0086] MS Mass spectrometry
[0087] m/z Mass/charge ratio
[0088] rt Room temperature
[0089] THF tetrahydrofuran
[0090] TLC Thin layer chromatography
[0091] tr Retention time
[0092] UV Ultraviolet

General Data:

[0093] The products were analyzed using Agilent HPLC-UV-MS equipment provided with a UV detector of variable wavelength and a mass spectrometer model 1100 VL. The wavelength used for detecting UV was 210 nm, whereas the MS detector has been operated in a positive electrospray ionization mode and a 100 to 700 m/z scan has been performed. Concerning chromatographic separation, the column used was a Kromasil 100 C18, 40×4.0 mm, 3.5 μm, and 2-5 μl have been injected. For the elution one of the two solvent gradients described below was followed:

[0094] Method A: 5-90% B, 0-8 min; 90% B, 8-11 min; 5% B, 9-11 min. The flow rate of the mobile phase is 0.7 ml/min.

[0095] Method B: 5-90% B, 0-4.5 min; 90% B, 4.5-6 min; 5% B, 6-7 min. The flow rate of the mobile phase is 1.4 ml/min.

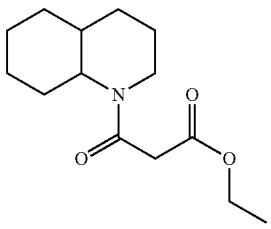
[0096] In both cases, the solvent A consists of 0.2% formic acid in water, whereas B is 0.2% formic acid in acetonitrile.

[0097] Alternatively, the analysis was conducted by means of Waters HPLC-UV-MS equipment provided with a detector having diodes in series and a mass spectrometer model EMD1000. The wavelength used for detecting UV was 210 nm, whereas the MS detector has been operated in a positive electrospray ionization mode and a 100 to 100 m/z scan has been performed. Concerning chromatographic separation, the column used was a Kromasil C18 2.1×50 mm, 3.5 μm and 2 μl have been injected. For the elution, the following gradient was followed: 5% B, 0.0-0.5 min; 5-100% B, 0.5-5.0 min; 100% B, 5.0-6.5 min; 100-5% B, 6.5-6.8 min; 5% B, 6.8-8.0 min.

[0098] The flow rate of the mobile phase is 0.5 ml/min.

Intermediate IIIa.1: ethyl 3-(octahydroquinolin-1-yl)-3-oxopropionate

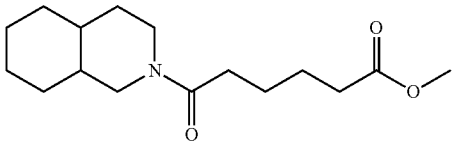
[0099]



[0100] 3.38 mL of Et_3N and 1.44 mL (11 mmol) of ethyl 3-chloro-3-oxopropionate are added to a solution of 1.5 mL (10 mmol) of decahydroquinoline in 100 mL of AcOEt. The resulting solution is kept under stirring at reflux for 10 h. Then water is added and the organic phase is separated, the aqueous phase is extracted once again with AcOEt. The organic phases are pooled and they are first washed with 5% solution of $NaHCO_3$ and subsequently with brine, they are dried over anhydrous Na_2SO_4, they are filtered and the solvent is evaporated under reduced pressure. 2.0 g of a yellow oil are obtained. It is identified as intermediate IIIa.1. Method A: tr: 6.76 min; m/z: 268.

Intermediate IIIa.2:
6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid methyl ester

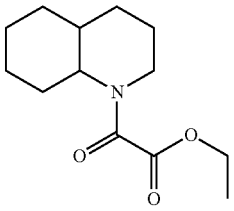
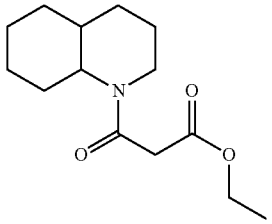
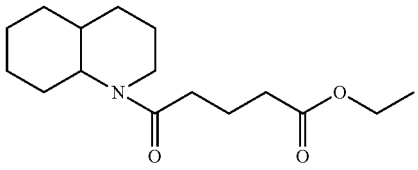
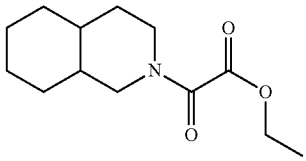
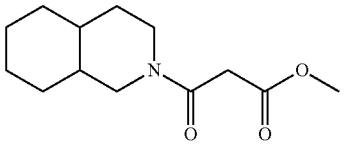
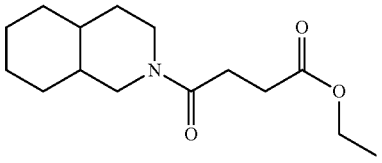
[0101]



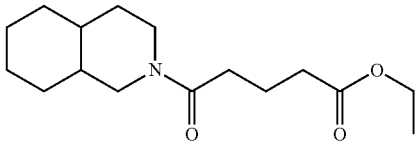
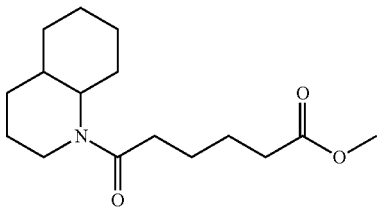
[0102] 80 mL of AcOEt and, then, 3 mL (22 mmol) of Et_3N are added to a mixture formed by 1.1 mL (8.32 mmol) of decahydroisoquinoline, 1.5 mL (10 mmol) of monomethyl adipate, 2 g (15 mmol) of HOBT and 2.9 g (15 mmol) of EDC. The solution formed is kept under stirring for 18 h. Then, it is treated with water and AcOEt, the organic phase is separated and the aqueous phase is extracted once more with AcOEt. The organic phases are pooled and they are washed successively with saturated $NaHCO_3$ solution, 1N HCl and brine. It is then dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. 1.5 g of an oil identified as intermediate IIIa.2 are obtained.

[0103] Method B: tr: 3.66 min/3.78 min; m/z: 282/282

[0104] The following intermediates were prepared in a manner similar to intermediates IIIa.1 or IIIa.2:

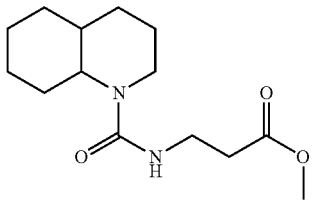
Int.	structure	name	method	rt (min)	m/z
IIIa.3		(Octahydroquinolin-1-yl)oxoacetic acid ethyl ester	A	6.83	240
IIIa.4		3-(octahydroquinolin-1-yl)-3-oxopropionic acid ethyl ester	A	6.37	254
IIIa.5		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid ethyl ester	A	6.96	282
IIIa.6		(Octahydroisoquinolin-2-yl)oxoacetic acid ethyl ester	A	6.83 7.05	240 240
IIIa.7		3-(octahydroisoquinolin-2-yl)-3-oxopropionic acid methyl ester	A	5.86 6.09	240 240
IIIa.8		4-(octahydroisoquinolin-2-yl)-4-oxobutyric acid ethyl ester	A	6.72 6.92	268 268

-continued

Int.	structure	name	method	rt (min)	m/z
IIIa.9		5-(octahydroisoquinolin-2-yl)-5-oxopentanoic acid ethyl ester	A	7.13 7.55	282 282
IIIa.10		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid methyl ester	B	3.67	282

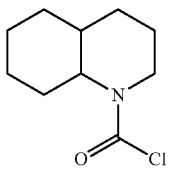
Intermediate IIIb.1: 3-[(octahydroquinoline-1-carbonyl)amino]propionic acid methyl ester

[0105]



Step 1:

[0106]



[0107] 2.5 g (18 mmol) of decahydroquinoline are dissolved in 150 mL of DCM and 2.7 g (9 mmol) of triphosgene are slowly added to the solution formed, preventing the temperature from exceeding 25° C. The resulting mixture is then refluxed for 18 h. It is then evaporated to dryness and the residue obtained is purified by means of silica gel filtration using AcOEt as eluent, obtaining 2 g of a yellowish oil identified as 1-chlorocarbonyloctahydroquinoline.

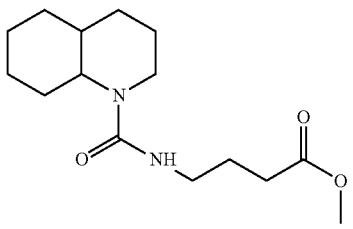
[0108] IR: 1729.

Step 2:

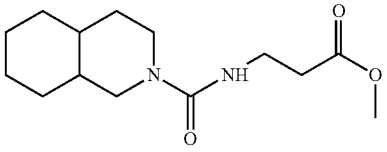
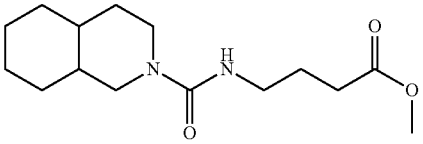
[0109] 0.5 g (3.5 mmol) of the 3-aminopropionic acid methyl ester hydrochloride are dissolved in 5 mL of anhydrous THF and 0.9 mL of Et_3N. Once dissolved, 650 mg of 1-chlorocarbonyloctahydroquinoline are slowly added and it is refluxed for 18 h. AcOEt is then added and the resulting solution is sequentially washed with water, 1N HCl and brine. The organic phase is dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure yielding 580 mg of intermediate IIIb.1.

[0110] Method B: tr: 3.17 min; m/z: 269.

[0111] The following intermediates were prepared in a manner similar to intermediate IIIb.1:

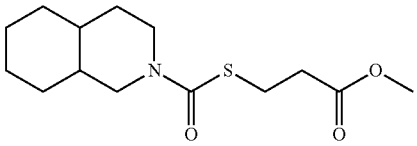
Int.	structure	name	method	rt (min)	m/z
IIIb.2		4-[(octahyrdoquinoline-1-carbonyl)amino]butyric acid methyl ester	B	3.29	283

-continued

Int.	structure	name	method	rt (min)	m/z
IIIb.3		3-[(octahydroisoquinoline-2-carbonyl)amino]propionic acid methyl ester	B	3.29	269
IIIb.4		4-[(octahydroisoquinoline-2-carbonyl)amino]butyric acid methyl ester	B	3.40	283

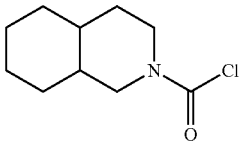
Intermediate IIIb.5: 3-(octahydroisoquinoline-2-carbonylsulfenyl)propionic acid methyl ester

[0112]



Step 1:

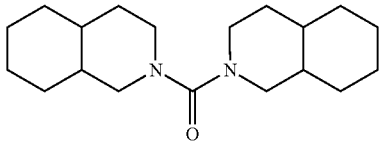
[0113]



[0114] 2-chlorocarbonyloctahydroisoquinoline: it was prepared in a manner similar to that described for step 1 of intermediate IIIb.1. IR: 1737.

Step 2:

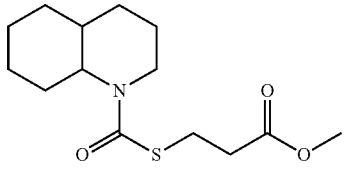
[0115] 1 g (5 mmol) of 2-chlorocarbonyloctahydroisoquinoline is added to a 0° C. solution of 0.55 mL (5 mmol) of 3-mercaptopropionic acid methyl ester in 10 mL of pyridine. Once added, it is left under stirring at rt for 18 h. AcOEt is then added to the reaction mixture and it is sequentially washed with a 5% solution of $NaHCO_3$, 1N HCl and brine. The organic phase is dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure being purified by means of silica gel column chromatography, using a (1:1) mixture of hexane:AcOEt as eluent, yielding 1.17 g of a residue identified as intermediate IIIb.5. and 210 mg of a solid identified as bis-(octahydroisoquinolin-2-yl)methanone (Vb.1)



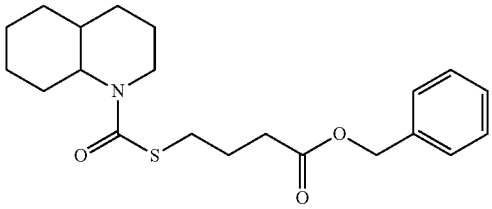
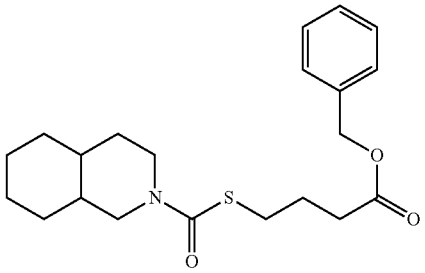
[0116] Method B: for IIIb.5 tr: 4.35 min; m/z: 286.

[0117] for Vb.1: tr: 5.34 min; m/z: 305.

[0118] The following intermediates were prepared in a manner similar to intermediate IIIb.5:

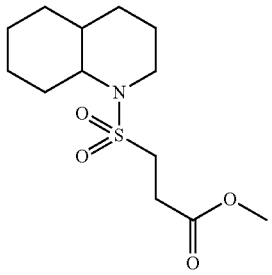
Int.	structure	name	method	rt (min)	m/z
IIIb.6		3-(octahydroquinoline-1-carbonylsulfenyl)propionic acid methyl ester	B	4.23	286

-continued

Int.	structure	name	method	rt (min)	m/z
IIIb.7		4-(octahydroquinoline-1-carbonylsulfenyl)butyric acid benzyl ester	B	5.04	376
IIIb.8		4-(octahydroisoquinoline-2-carbonylsulfenyl)butyric acid benzyl ester	B	5.14	377

Intermediate IIIm.1:
3-(octahydroquinoline-1-sulfonyl)propionic acid methyl ester

[0119]



[0120] Step 1

[0121] 4.48 mL (55.9 mmol) of sulfuryl chloride are added dropwise to a mixture formed by 2.47 mL (22.4 mmol) of methyl 3-mercaptopropionate and 5.64 g (55.9 mmol) of potassium nitrate cooled to 0° C. Once added it is left to reach rt keeping the stirring for 10 h. Saturated $NaHCO_3$ solution is added and the organic phase is separated, which is subsequently washed again with $NaHCO_3$ and then with brine. The organic phase is dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure yielding 2.6 g of an oil identified as the 3-chlorosulfonylpropionic acid methyl ester.

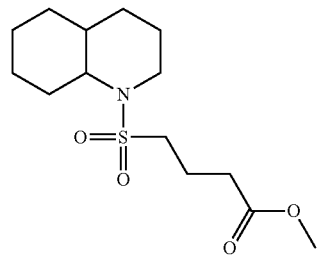
[0122] Method B: tr: 2.05 min; m/z: non ionizable.

[0123] Step 2

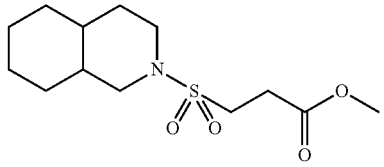
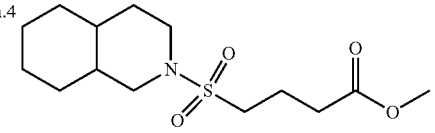
[0124] 2.1 mL of Et_3N (15.3 mmol) and subsequently, dropwise, 1.3 g (7 mmol) of the 3-chlorosulfonylpropionic acid methyl ester are added to a solution formed by 1.1 mL (7.6 mmol) of decahydroquinoline in 70 mL of AcOEt. The mixture is heated to 70° C. and that temperature is maintained for 12 h. It is then left to cool and water is added, the organic phase is separated and it is sequentially washed with 5% solution of sodium bicarbonate, 2N HCl and brine. The organic phase is dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure yielding 1.32 g of a reddish oil identified as intermediate IIIm.1.

[0125] Method B: tr: 3.36 min; m/z: 290

[0126] The following intermediates were prepared in a manner similar to intermediate IIIm.1:

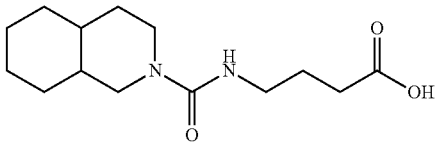
Int.	structure	name	method	rt (min)	m/z
IIIm.2		4-(octahydroquinoline-1-sulfonyl)butyric acid methyl ester	B	3.73	304

-continued

Int.	structure	name	method	rt (min)	m/z
IIIm.3		3-(octahydroisoquinoline-2-sulfonyl)propionic acid methyl ester	B	3.79	290
IIIm.4		4-(octahydroisoquinoline-2-sulfonyl)butyric acid methyl ester	B	3.83 3.91	304 304

Intermediate IVb.1:
4-[(octahydroisoquinoline-2-carbonyl)amino]butyric acid

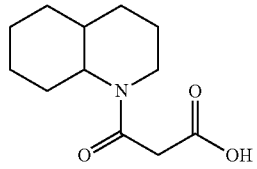
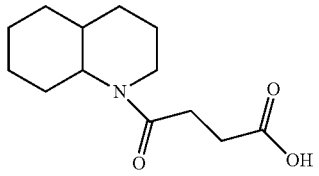
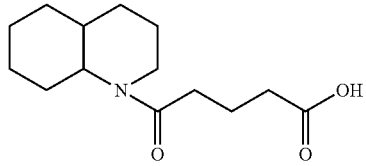
[0127]



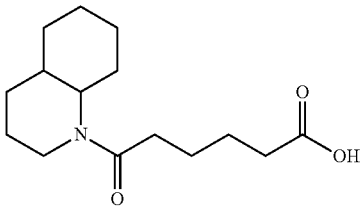
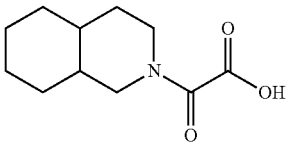
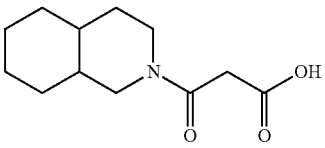
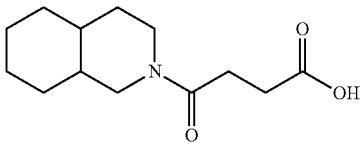
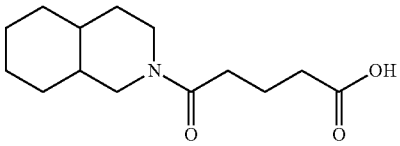
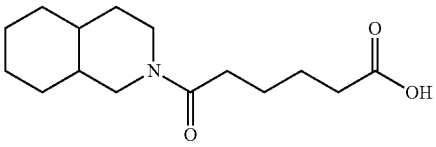
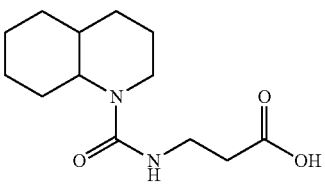
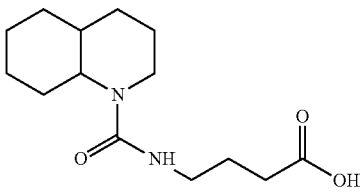
[0128] 1.2 g (4.21 mmol) of intermediate IIIb.4 are dissolved in a mixture formed by 37.5 mL of THF and 12.5 mL of MeOH, and 5.1 mL of a 1M LiOH solution in water are added to the solution formed. The resulting mixture is kept under stirring at rt for 18 h. Then it is diluted in AcOEt and washed with water, the aqueous phase is acidified with a 1N solution of HCl until pH=3 and it is extracted with AcOEt. Finally, the organic phases are pooled, they are dried over anhydrous Na_2SO_4, they are filtered and the solvent is evaporated under reduced pressure. 570 mg of a white solid are obtained.

[0129] Method B: tr: 2.85 min/2.96 min; m/z: 269/269

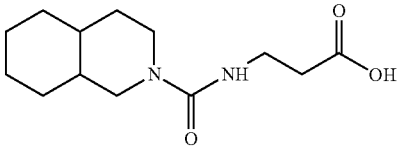
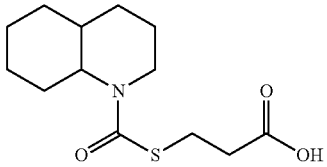
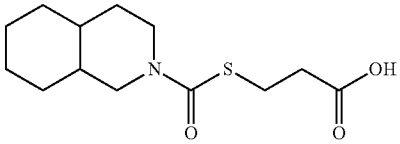
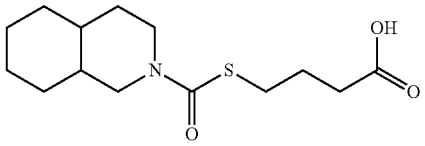
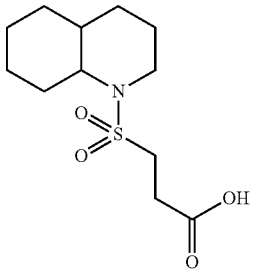
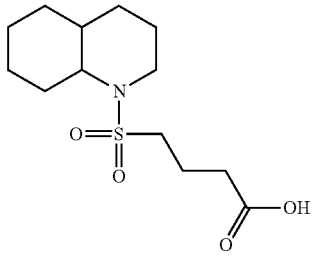
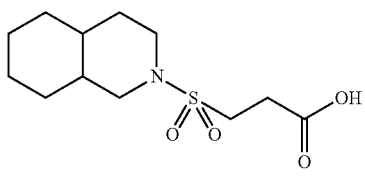
[0130] The following intermediates were prepared in a manner similar to intermediate IVb.1:

Int.	structure	name	method	rt (min)	m/z
IVa.1		3-(octahydroquinolin-1-yl)-3-oxopropionic acid	A	5.18	226
IVa.2		4-(octahydroquinolin-1-yl)-4-oxobutyric acid	A	5.32	240
IVa.3		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid	A	5.42	254

-continued

Int.	structure	name	method	rt (min)	m/z
IVa.4		6-(octahydroquinolin-2-yl)-6-oxobutyric acid	B	3.08	268
IVa.5		(octahydroisoquinolin-2-yl)oxoacetic acid	A	4.25 4.40	212 212
IVa.6		6-(octahydroquinolin-2-yl)-3-oxopropionic acid	A	5.19 5.40	226 226
IVa.7		4-(octahydroquinolin-2-yl)-4-oxobutyric acid	A	5.30 5.51	240 240
IVa.8		5-(octahydroquinolin-2-yl)-5-oxopentanoic acid	A	5.44 5.64	254 254
IVa.9		6-(octahydroquinolin-2-yl)-6-oxohexanoic acid	B	3.09 3.20	268 268
IVb.2		3-[(octahydroquinoline-1-carbonyl)amino]propionic acid	B	2.75	255
IVb.3		4-[(octahydroquinoline-1-carbonyl)amino]butyric acid	B	2.85	269

-continued

Int.	structure	name	method	rt (min)	m/z
IVb.4		3-[(octahydroquinoline-2-carbonyl)amino]propionic acid	B	2.87	255
IVb.5		3-(octahydroquinoline-1-carbonylsulfenyl)propionic acid	B	3.53	272
IVb.6		3-(octahydroquinoline-2-carbonylsulfenyl)propionic acid	B	3.66	272
IVb.7		4-(octahydroquinoline-2-carbonylsulfenyl)butyric acid	B	3.81	286
IVm.1		3-(octahydroquinoline-1-sulfonyl)propionic acid	B	3.10	276
IVm.2		4-(octahydroquinoline-1-sulfonyl)butyric acid	B	3.18	290
IVm.3		4-(octahydroquinoline-2-sulfonyl)propionic acid	B	3.18 3.27	276 276

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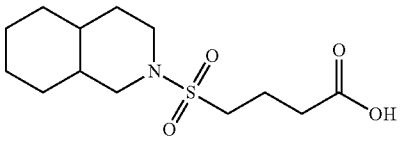
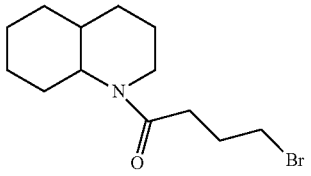
Int.	structure	name	method	rt (min)	m/z
IVm.4		4-(octahydroquinoline-2-sulfonyl)butyric acid	B	3.25 3.34	290 290

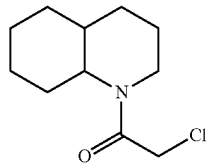
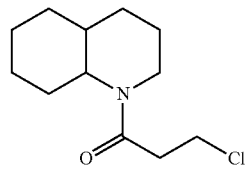
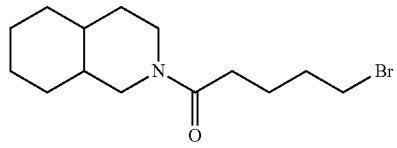
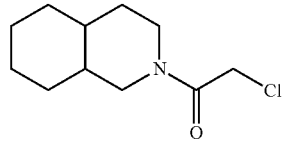
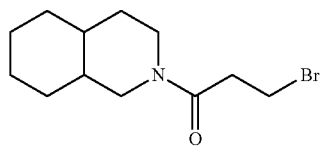
Intermediate IIIc.1:
4-bromo-1-(octahydroquinolin-1-yl)butan-1-one

[0131]

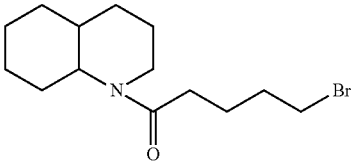


[0132] 1.37 mL (9.9 mmol) of Et_3N and 0.5 mL (4.5 mmol) of 5-bromovaleric acid chloride are added to a solution of 1 mL (4.0 mmol) of the decahydroquinoline in AcOEt. The resulting solution is kept under stirring for 18 h at rt. The solution is then washed with water, and the water re-extracted twice with AcOEt. The pooled organic phases are sequentially washed with 5% solution of $NaHCO_3$, 1N HCl and saturated ammonium chloride solution. Finally the organic phase is dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. 1.2 g of a yellow oil, which was used without subsequent purification, are obtained.

[0133] The following intermediates were prepared in a manner similar to intermediate IIIc.1:

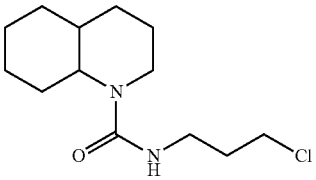
Int.	structure	name	method	rt (min)	m/z
IIIc.2		2-chloro-1-(octahydroquinolin-1-yl)ethanone	A	6.45	216, 218
IIIc.3		3-chloro-1-(octahydroquinolin-1-yl)propan-1-one	A	7.02	230, 232
IIIc.4		5-bromo-1-(octahydroquinolin-2-yl)pentan-1-one	A	7.60 7.78	(302, 304) (302, 304)
IIIc.5		2-chloro-1-(octahydroquinolin-2-yl)ethanone	A	6.44 6.68	(216, 218) (216, 218)
IIIc.6		3-bromo-1-(octahydroisoquinolin-2-yl)propan-1-one	A	7.00 7.21	(274, 276) (274, 276)

-continued

Int.	structure	name	method	rt (min)	m/z
IIIc.7		5-bromo-1-(octahydroisoquinolin-1-yl)pentan-1-one	A	—	—

Intermediate IIIf.1: Octahydrouinoline-1-carboxylic acid 3-chloropropylamide

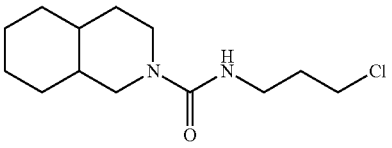
[0134]



[0135] 2.5 g (21 mmol) of 3-chloropropylisocyanate are dissolved in 150 mL of anhydrous THF and 6.4 mL (46 mmol) of Et_3N. Once dissolved 3.4 mL (23 mmol) of decahydroquinoline are slowly added and it is refluxed for 18 h. AcOEt is then added and the resulting solution is sequentially washed with water, 1N HCl and brine. The organic phase is dried over anhydrous Na_2SO_4 it is filtered and the solvent is evaporated under reduced pressure yielding 4.6 g of intermediate IIIf.1. Method B: tr: 3.58 min; m/z: 259.

Intermediate IIIf.2: Octahydroisoquinoline-2-carboxylic acid 3-(chloropropyl)amide

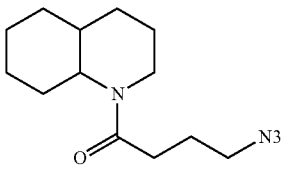
[0136]



[0137] It was prepared in a manner similar to that described for intermediate IIIf.1. Method B: tr: 3.57 min/3.68 min; m/z: 259, 259.

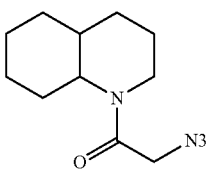
Intermediate IVc.1: 4-azido-1-(octahydroquinolin-1-yl)butan-1-one

[0138]

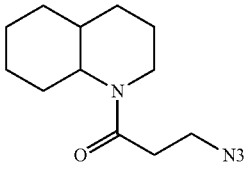
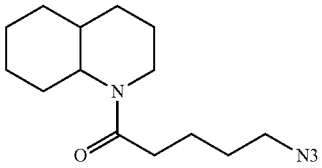
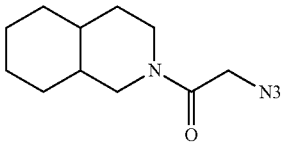
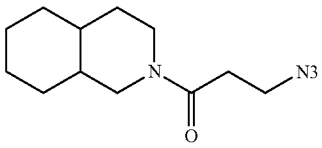
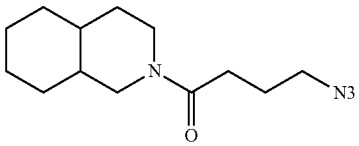
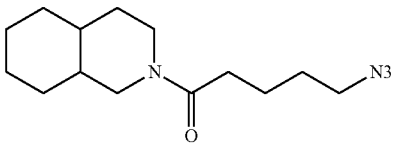
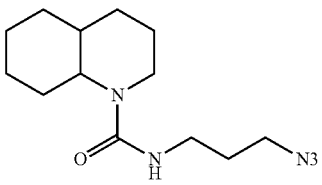
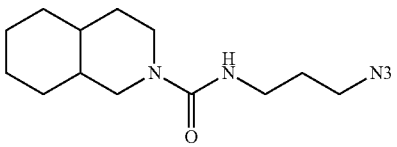


[0139] 710 mg (10.8 mmol) of sodium azide are added to a solution of 1.1 g (3.6 mmol) of intermediate IIIc.1 in 35 mL of anhydrous DMF. The resulting solution is kept under stirring for 18 h at a temperature of 90° C. The solution is then cooled and water is added, extracting three times with AcOEt. The pooled organic phases are washed with brine. Finally the organic phase is dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. 830 mg of a yellow oil, which was used without subsequent purification, are obtained. Method A: tr: 7.07 min; m/z: 251.

[0140] The following intermediates were prepared in a manner similar to intermediate IVc.1:

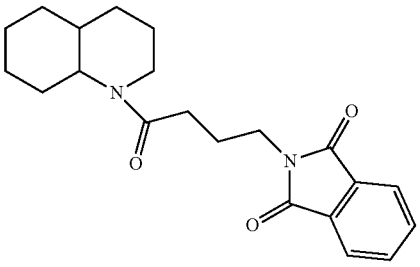
Int.	structure	name	method	rt (min)	m/z
IVc.2		2-azido-1-(octahydroquinolin-1-yl)ethanone	A	6.48	223

-continued

Int.	structure	name	method	rt (min)	m/z
IVc.3		3-azido-1-(octahydroquinolin-1-yl)propan-1-one	A	6.79	237
IVc.4		3-azido-1-(octahydroquinolin-1-yl)pentan-1-one	A	7.33	265
IVc.5		3-azido-1-(octahydroisoquinolin-2-yl)ethanone	A	6.46 6.70	223 223
IVc.6		3-azido-1-(octahydroisoquinolin-2-yl)propan-1-one	A	6.77 6.99	237 237
IVc.7		4-azido-1-(octahydroisoquinolin-2-yl)butan-1-one	A	7.07 7.27	251 251
IVc.8		5-azido-1-(octahydroisoquinolin-2-yl)pentan-1-one	A	7.33 7.51	265 265
IVf.1		Octahydroquinoline-1-carboxylic acid (3-azidopropyl)amide	B	3.56	266
IVf.2		Octahydroisoquinoline-2-carboxylic acid (3-azidopropyl)amide	B	3.55 3.66	266 266

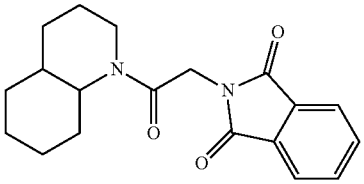
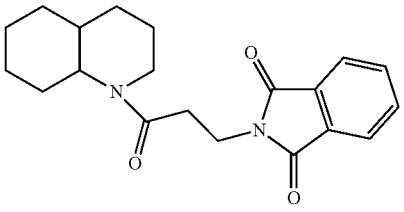
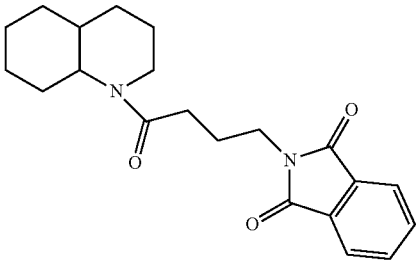
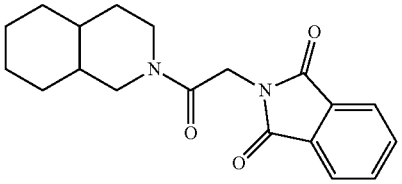
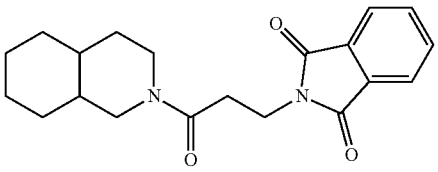
Intermediate Ic.1: 2-[4-(octahydroquinolin-1-yl)-4-oxobutyl]isoindole-1,3-dione

[0141]

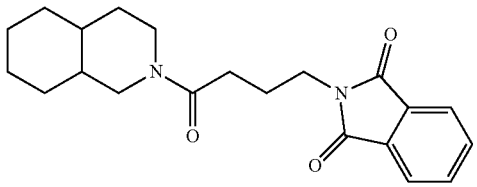
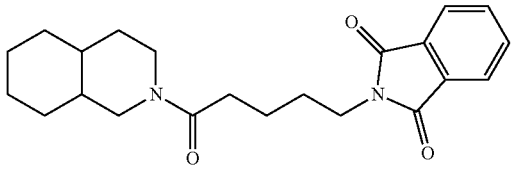


[0142] 200 mg (1.1 mmol) of potassium phthalimide are added to a solution of 307 mg (1.1 mmol) of intermediate IIIc.1 in 10 mL of anhydrous DMF. The resulting solution is kept under stirring for 18 h at a temperature of 90° C. The solution is then cooled and water is added, extracting it with AcOEt. The pooled organic phases are washed with brine. Finally the organic phase is dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. 360 mg of a paste, which was purified by means of silica gel column chromatography using a (1:1) mixture of hexane: AcOEt as eluent, yielding 180 mg of an oil identified as intermediate Ic.1, are obtained. Method A: tr: 7.17 min; m/z: 355.

[0143] The following intermediates were prepared in a manner similar to intermediate Ic.1:

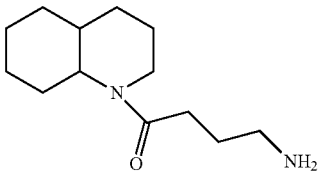
Ex.	structure	name	method	rt (min)	m/z
Ic.2		2-[2-(octahydroquinolin-1-yl)-2-oxoethyl]isoindole-1,3-dione	A	6.93	327
Ic.3		2-[3-(octahydroquinolin-1-yl)-3-oxopropyl]isoindole-1,3-dione	A	7.00	341
Ic.4		2-[4-(octahydroquinolin-1-yl)-4-oxobutyl]isoindole-1,3-dione	A	7.17	355
Ic.5		2-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]isoindole-1,3-dione	A	6.92 7.10	327 327
Ic.6		2-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]isoindole-1,3-dione	A	6.96 7.12	341 341

-continued

Ex.	structure	name	method	rt (min)	m/z
Ic.7		2-[4-(octahydroisoquinolin-2-yl)-4-oxobutyl]isoindole-1,3-dione	A	7.16 7.33	355 355
Ic.8		2-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]isoindole-1,3-dione	A	7.39 7.55	369 369

Intermediate Vc.1:
4-amino-1-(octahydroquinolin-1-yl)butan-1-one

[0144]

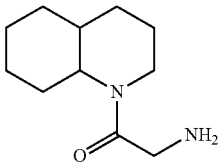
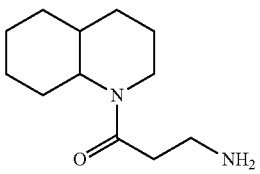


[0145] Option A: 0.1 mL of hydrazine (2.5 mmol) is added to a solution formed by 172 mg (0.48 mmol) of intermediate Ic.1 in 5 mL of EtOH and it is heated to reflux temperature for 2 h. The solution is then cooled and concentrated HCl is added until acidic pH is reached, stirring it for 2 h or more. The resulting suspension is filtered and the water is basified with 1N NaOH, extracting it with DCM. The pooled organic phases are washed with brine and are dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. The resulting residue is purified by means of silica gel column chromatography, using a (10:1) mixture of DCM:MeOH as eluent, yielding 90 mg of an oil identified as intermediate Vc.1.

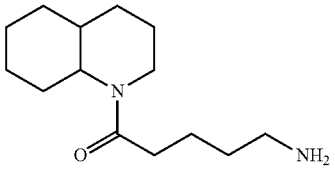
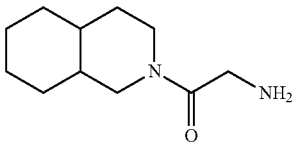
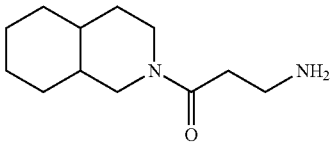
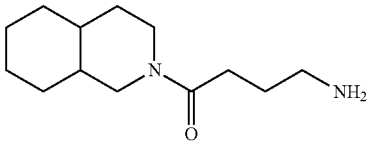
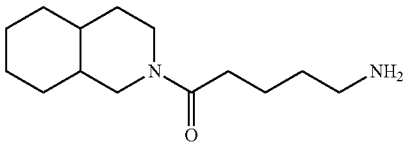
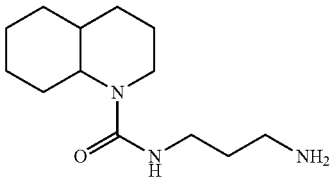
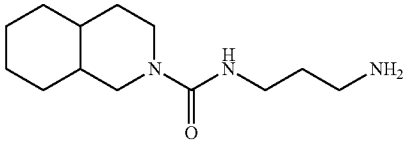
[0146] Option B: A suspension formed by 830 mg (3.32 mmol) of intermediate IVc.1 and 83 mg of 5% Pd/C in 35 mL of MeOH is stirred under hydrogen atmosphere until the disappearance of the starting product by monitoring by TLC. It is filtered through Celite and it is evaporated to dryness yielding 700 mg of an oil identified as intermediate Vc.1.

[0147] Method A: tr: 3.94 min; m/z: 225.

[0148] The following intermediates were prepared in a manner similar to intermediate Vc.1:

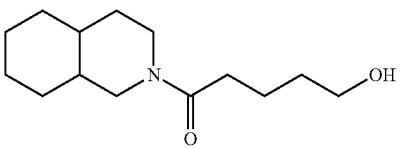
Ex.	structure	name	method	rt (min)	m/z
Vc.2		2-amino-1-(octahydroquinolin-1-yl)ethanone	A	3.41	197
Vc.3		3-amino-1-(octahydroquinolin-1-yl)propan-1-one	A	3.64	211

-continued

Ex.	structure	name	method	rt (min)	m/z
Vc.4		5-amino-1-(octahydroquinolin-1-yl)pentan-1-one	A	4.01	239
Vc.5		2-amino-1-(octahydroisoquinolin-2-yl)ethanone	A	3.69	197
Vc.6		3-amino-1-(octahydroisoquinolin-2-yl)propan-1-one	A	4.00	211
Vc.7		4-amino-1-(octahydroisoquinolin-2-yl)butan-1-one	A	4.03	225
Vc.8		5-amino-1-(octahydroisoquinolin-2-yl)pentan-1-one	A	4.27	239
Vc.9		Octahydroquinoline-1-carboxylic acid (3-aminopropyl)amide	B	2.11	240
Vc.10		Octahydroisoquinoline-2-carboxylic acid (3-aminopropyl)amide	B	2.22	240

Intermediate IVj.1: 5-hydroxyl-1-(octahydroisoquinolin-2-yl)pentan-1-one

[0149]



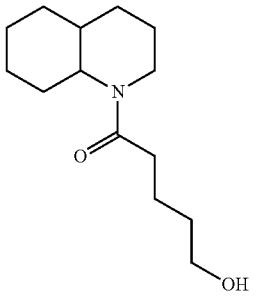
[0150] 0.63 mL (4.93 mmol) of isobutyl chloroformate are added dropwise to a solution of 1.25 g (4.93 mmol) of intermediate IVa.8 and 0.6 mL of N-methylmorpholine (5.4 mmol) in 50 mL of anhydrous THF and cooled to 0° C. The resulting solution is left to stir for 30 minutes at that temperature and 370 mg (9.86 mmol) of sodium borohydride are then added. Once added it is left to reach rt and it is stirred for 2 h. It is then partitioned between AcOEt and water. The aqueous phase is extracted twice with AcOEt, the organic phases are pooled, dried over anhydrous Na_2SO_4, filtered and evapo-

rated to dryness, 1 g of a yellow oil, which was used without subsequent purification, being obtained.

[0151] Method B: tr: 2.94 min/3.06 min; m/z: 240/240.

Intermediate LVj.2:
5-hydroxyl-1-(octahydroquinolin-1-yl)pentan-1-one

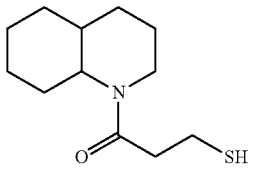
[0152]



[0153] It is prepared in a manner similar to that described for intermediate IVj.1. Method B: tr: 2.94 min; m/z: 240.

Intermediate IVp.1:
3-mercapto-1-(octahydroquinolin-1-yl)propan-1-one

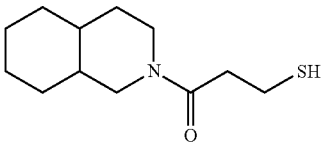
[0154]



[0155] 8.5 mL of Et_3N, 6.5 g of EDC (mmol) and 5.7 g of HOBT (mmol) are added to a solution formed by 3 g (28.2 mmol) of 3-mercaptopropionic acid in 45 mL of DMF and cooled to 0° C. and it is left for 15 minutes under stirring at that temperature. 4.2 g (28.3 mmol) of decahydroquinoline are then added and it is stirred at rt for 18 h. Then, it is treated with water and AcOEt, the organic phase is separated and the aqueous phase is extracted once more with AcOEt. The organic phases are pooled and successively washed with saturated $NaHCO_3$ solution, 1N HCl and brine. It is then dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. The resulting residue is purified by means of silica gel column chromatography, using a (50:1) mixture of DCM:MeOH as eluent, yielding 600 mg of the thiol intermediate IVp.1.

Intermediate IVp2: 3-mercapto-1-(octahydroisoquinolin-2-yl)propan-1-one

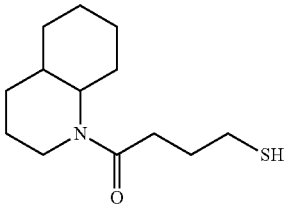
[0156]



[0157] It is prepared in a manner similar to that described for intermediate IVp.1, Method B: tr: 3.59 min/3.72 min; m/z: 228/228.

Intermediate IVp.3:
4-mercapto-1-(octahydroquinolin-1-yl)butan-1-one

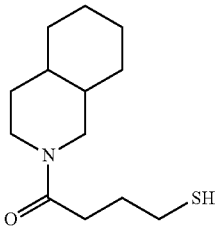
[0158]



[0159] 1.5 mL of decahydroquinoline (10 mmol) and 10 mg of camphorsulfonic acid are added to a solution formed by 0.9 mL (10 mmol) of γ-thiolactone in 80 mL of toluene. The resulting mixture is left to stir at 100° C. for 6 h. The solvent is then evaporated under reduced pressure and the resulting residue is purified by means of silica gel column chromatography, using a (50:1) mixture of DCM:MeOH as eluent, yielding 1.5 g of the thiol intermediate IVp.3. Method B: tr: 3.75 min; m/z: 242.

Intermediate IVp.4: 4-mercapto-1-(octahydroisoquinolin-2-yl)butan-1-one

[0160]

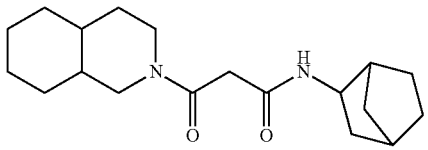


[0161] It is prepared in a manner similar to that described for intermediate IVp.3. Method B: tr: 3.75 min/3.87 min; m/z: 242/242

Example Ia.1

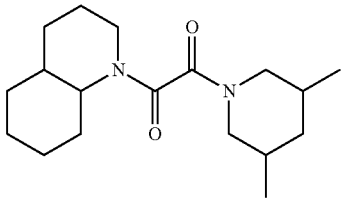
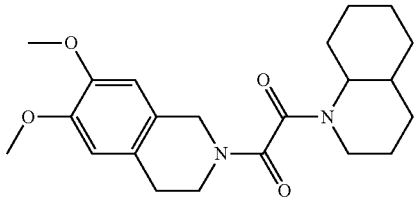
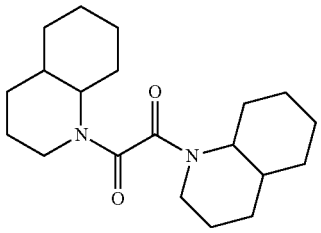
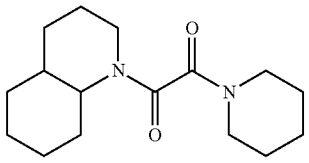
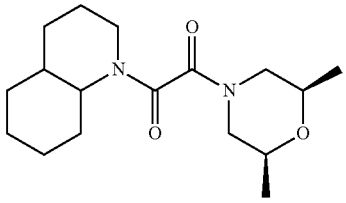
N-bicyclo[2.2.1]hept-2-yl-3-(octahydroisoquinolin-2-yl)-3-oxopropionamide

[0162]

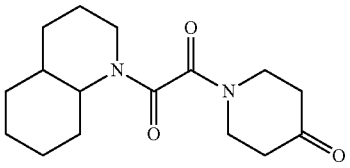
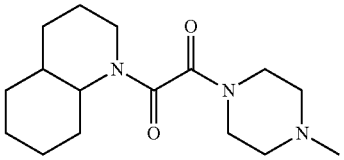
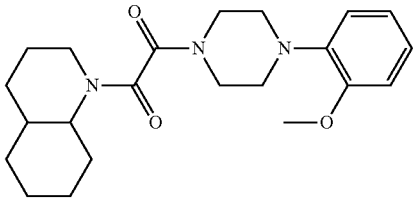
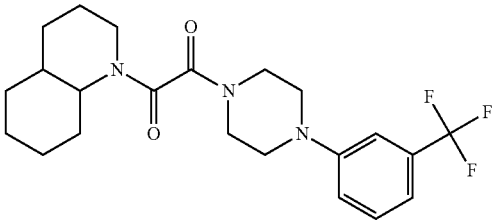
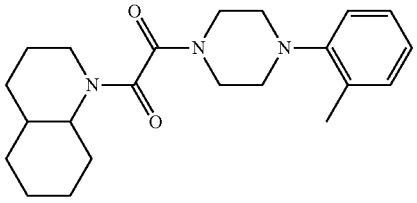
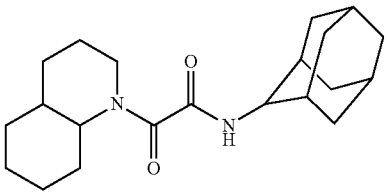
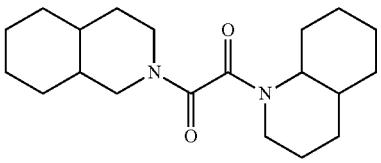


[0163] 43 μL (0.31 mmol) of Et_3N, 29 mg (0.21 mmol) of HOBT, 41 mg (0.21 mmol) of EDC and 18.6 μL (0.16 mmol) of 2-aminonorbornan are added to a solution of 32 mq (0.14 mmol) of the acid intermediate IVa.6 in 2 mL of AcOEt. The solution formed is kept under stirring for 18 h. It is then treated with water and more AcOEt is added, the organic phase is separated and the aqueous phase is extracted once more with more AcOEt. The organic phases are pooled and successively washed with saturated $NaHCO_3$ solution, 1N HCl and brine. It is then dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. 27 mg of the compound identified as example Ia.1 are obtained. Method A: tr: 6.76 min/6.84 min; m/z: 319/319.

[0164] The following examples were prepared in a manner similar to example Ia.1:

Ex.	structure	name	method	rt (min)	m/z
Ia.2		1-(3,5-dimethylpiperidin-1-yl)-2-(octahydroquinolin-1-yl)ethane-1,2-dione	A	7.22 7.35	307 307
Ia.3		1-(6,7-dimethoxy-3,4-dihydro-1-H-isoquinolin-2-yl)-2-(octahydroquinolin-1-yl)ethane-1,2-dione	A	6.54	387
Ia.4		1,2-bis-(octahydroquinolin-1-yl)ethane-1,2-dione	A	7.66 7.76	333 333
Ia.5		1-(octahydroquinolin-1-yl)-2-(piperidin-1-yl)ethane-1,2-dione	A	6.19	279
Ia.6		1-((2S,6R)-2,6-dimethylmorpholin-4-yl)-2-(octahydroquinolin-1-yl)ethane-1,2-dione	A	6.19	309

-continued

Ex.	structure	name	method	rt (min)	m/z
Ia.7		1-(octahydroquinolin-1-yl)-2-(4-oxopiperidin-1-yl)ethane-1,2-dione	A	5.10	293
Ia.8		1-(4-methylpiperazin-1-yl)-2-(octahydroquinolin-1-yl)ethane-1,2-dione	A	3.82	294
Ia.9		1-[4-(2-methoxyphenyl)piperazin-1-yl]-2-(octahydroquinolin-1-yl)ethane-1,2-dione	A	7.00	386
Ia.10		1-(octahydroquinolin-1-yl)-2-[4-(3-trifluoromethyl-phenyl)piperazin-1-yl]-ethane-1,2-dione	A	7.90	424
Ia.11		1-(octahydroquinolin-1-yl)-2-(4-o-tolylpiperazin-1-yl)ethane-1,2-dione	A	7.82	370
Ia.12		2-(octahydroquinolin-1-yl)-2-oxo-N-tricyclo[3.3.1.1^{3,7}]decan-2-ylacetamide	A	7.99 8.18	345 345
Ia.13		1-(octahydroisoquinolin-2-yl)-2-(octahydroquinolin-1-yl)ethane-1,2-dione	A	7.69 7.84	333 333

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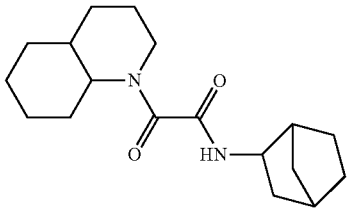
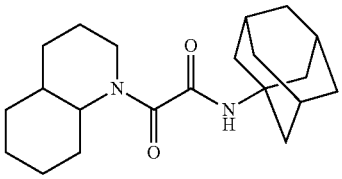
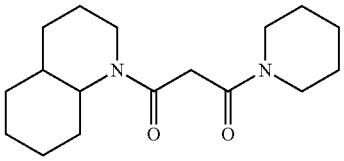
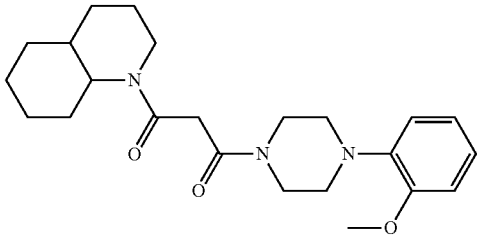
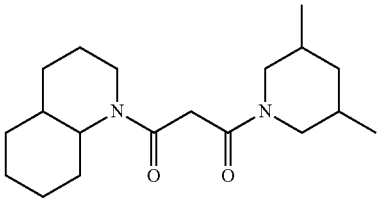
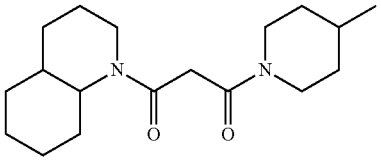
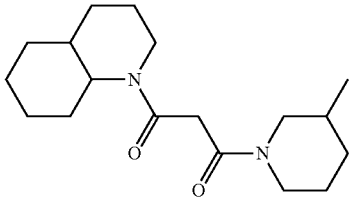
Ex.	structure	name	method	rt (min)	m/z
Ia.14		N-(bicyclo[2.2.1]hept-2-yl)-2-(octahydroquinolin-1-yl)-2-oxoacetamide	A	6.98 7.13	305 305
Ia.15		2-(octahydroquinolin-1-yl)-2-oxo-N-tricyclo[3.3.1.1^{3,7}]decan-1-ylacetamide	A	8.15 8.33	345 345
Ia.16		1-(octahydroquinolin-1-yl)-3-(piperidin-1-yl)propane-1,3-dione	A	6.09	293
Ia.17		1-[4-(2-methoxyphenyl)piperazin-1-yl]-3-(octahydroquinolin-1-yl)propane-1,3-dione	A	6.46	400
Ia.18		1-(3,5-dimethylpiperidin-1-yl)-3-(octahydroquinolin-1-yl)propane-1,3-dione	A	7.03 7.17	321 321
Ia.19		1-(4-methylpiperidin-1-yl)-3-(octahydroquinolin-1-yl)propane-1,3-dione	A	6.62	307
Ia.20		1-(3-methylpiperidin-1-yl)-3-(octahydroquinolin-1-yl)propane-1,3-dione	A	6.62	307

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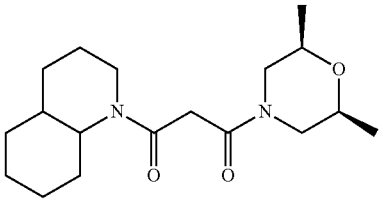
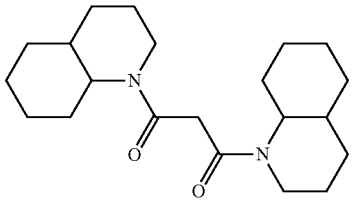
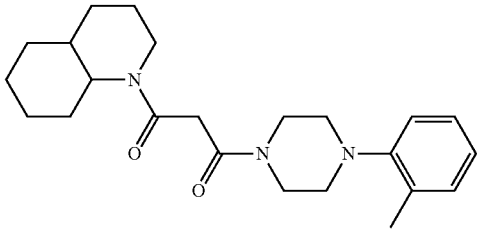
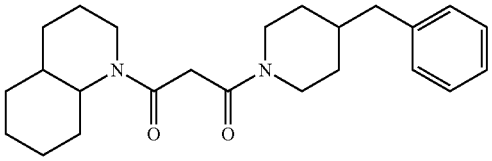
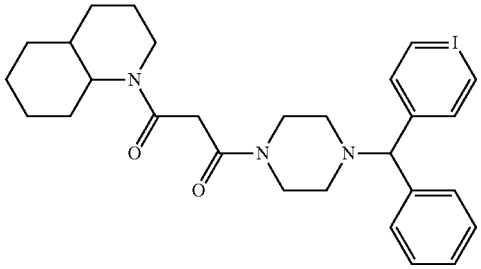
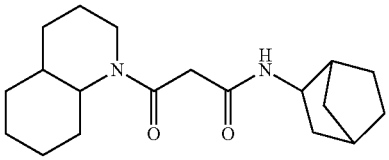
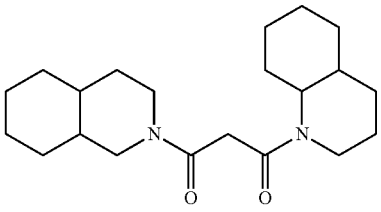
Ex.	structure	name	method	rt (min)	m/z
Ia.21		1-((2S,6R)-2,6-dimethylmorpholin-4-yl)-3-(octahydroquinolin-1-yl)propane-1,3-dione	A	5.90	323
Ia.22		1,3-bis-(octahydroquinolin-1-yl)propane-1,3-dione	A	7.53	347
Ia.23		1-(octahydroquinolin-1-yl)-3-(4-o-tolylpiperazin-1-yl)propane-1,3-dione	A	7.56	384
Ia.24		1-(4-benzylpiperidin-1-yl)-3-(octahydroquinolin-1-yl)propane-1,3-dione	A	6.31	460
Ia.25		1-(4-benzhydrylpiperazin-1-yl)-3-(octahydroquinolin-1-yl)propane-1,3-dione	A	7.72	383
Ia.26		N-bicyclo[2.2.1]hept-2-yl-3-(octahydroquinolin-1-yl)-3-oxopropionamide	A	6.76 6.84	319 319
Ia.27		1-(octahydroisoquinolin-2-yl)-3-(octahydroquinolin-1-yl)propane-1,3-dione	A	7.54 7.68	347 347

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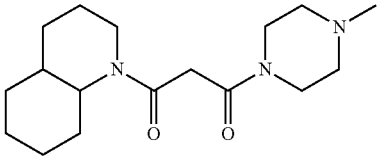
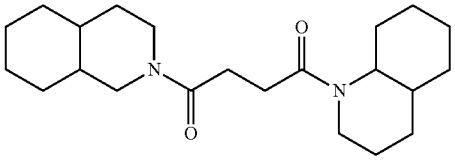
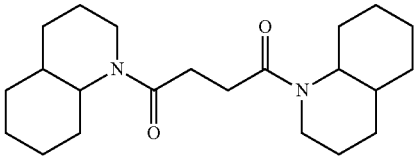
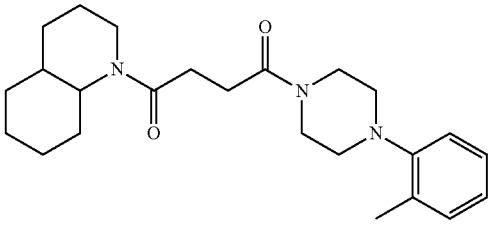
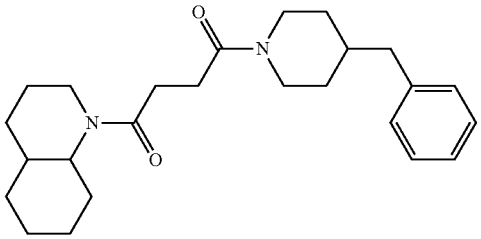
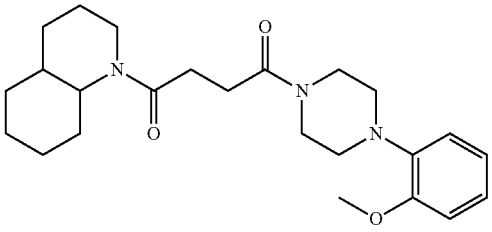
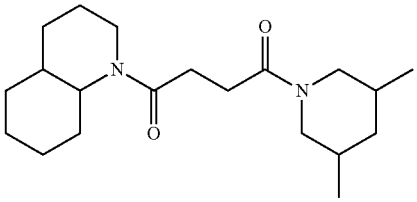
Ex.	structure	name	method	rt (min)	m/z
Ia.28		1-(4-methylpiperazin-1-yl)-3-(octahydroquinolin-1-yl)propane-1,3-dione	A	3.90	308
Ia.29		1-(octahydroisoquinolin-2-yl)-4-(octahydroquinolin-1-yl)butane-1,4-dione	A	7.75 7.90	361 361
Ia.30		1,4-bis-(octahydroquinolin-1-yl)butane-1,4-dione	A	7.84	361
Ia.31		1-(octahydroquinolin-1-yl)-4-(4-o-tolylpiperazin-1-yl)butane-1,4-dione	A	7.78	398
Ia.32		1-(4-benzylpiperidin-1-yl)-4-(octahydroquinolin-1-yl)butane-1,4-dione	A	7.99 8.11	397 397
Ia.33		1-[4-(2-methoxyphenyl)piperazin-1-yl]-4-(octahydroquinolin-1-yl)butane-1,4-dione	A	6.53	414
Ia.34		1-(3,5-dimethylpiperidin-1-yl)-4-(octahydroquinolin-1-yl)butane-1,4-dione	A	7.32 7.48	335 335

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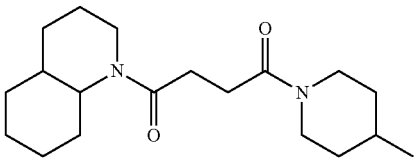
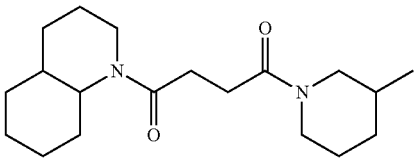
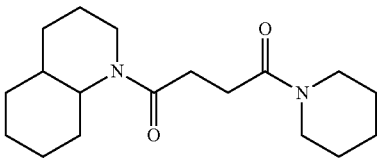
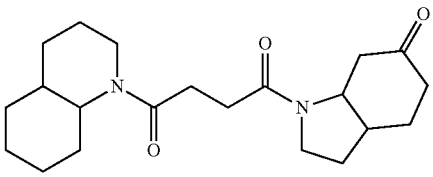
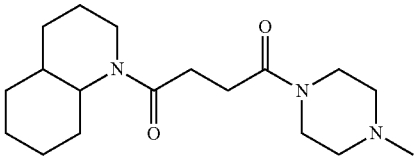
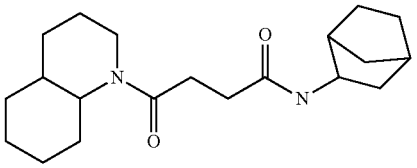
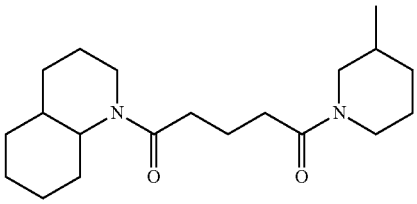
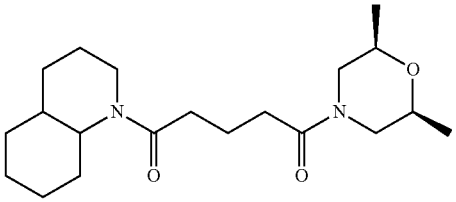
Ex.	structure	name	method	rt (min)	m/z
Ia.35		1-(4-methylpiperidin-1-yl)-4-(octahydroquinolin-1-yl)butane-1,4-dione	A	6.93	321
Ia.36		1-(3-methylpiperidin-1-yl)-4-(octahydroquinolin-1-yl)butane-1,4-dione	A	6.92	321
Ia.37		1-(octahydroquinolin-1-yl)-4-piperidin-1-ylbutane-1,4-dione	A	6.36	307
Ia.38		1-(octahydroquinolin-1-yl)-4-(6-oxooctahydroindol-1-yl)butane-1,4-dione	A	4.25	361
Ia.39		1-(4-methylpiperazin-1-yl)-4-(octahydroquinolin-1-yl)butane-1,4-dione	A	4.19	322
Ia.40		N-bicyclo[2.2.1]hept-2-yl-4-(octahydroquinolin-1-yl)-4-oxobutyramide	A	6.84	333
Ia.41		1-(3-methylpiperidin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	7.01	335
Ia.42		1-((2S,6R)-2,6-dimethylmorpholin-4-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	6.28	351

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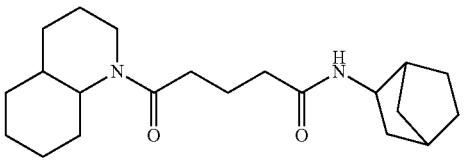
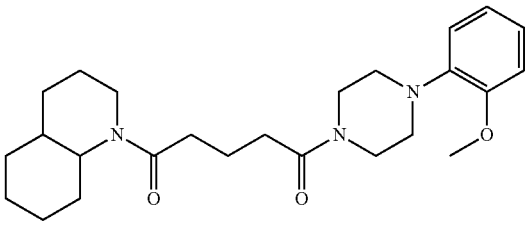
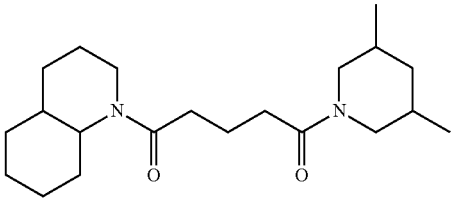
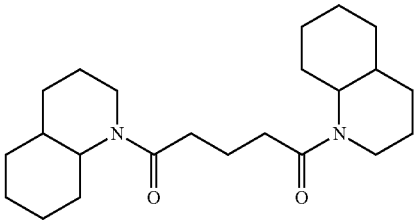
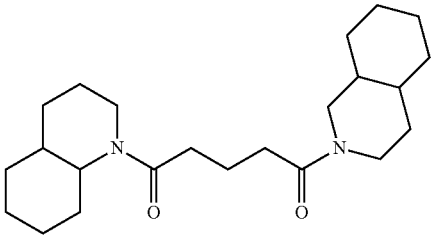
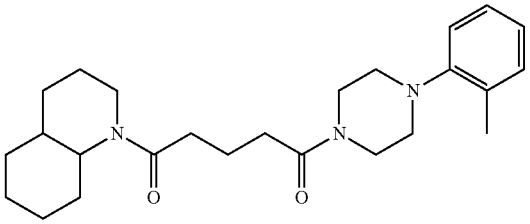
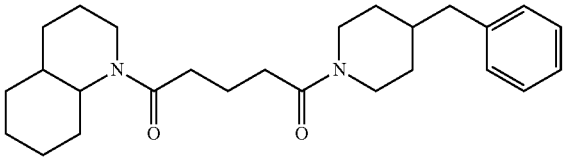
Ex.	structure	name	method	rt (min)	m/z
Ia.43		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid bicyclo[2.2.1]hept-2-ylamide	A	6.88	347
Ia.44		1-[4-(2-methoxyphenyl)piperazin-1-yl]-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	6.75	428
Ia.45		1-(3,5-dimethylpiperidin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	7.41 7.57	349 349
Ia.46		1,5-bis-(octahydroquinolin-1-yl)pentane-1,5-dione	A	7.92	375
Ia.47		1-(octahydroisoquinolin-2-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	7.91 8.08	375 375
Ia.48		1-(octahydroquinolin-1-yl)-5-(4-o-tolylpiperazin-1-yl)pentane-1,5-dione	A	7.89	412
Ia.49		1-(4-benzylpiperidin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	8.05	411

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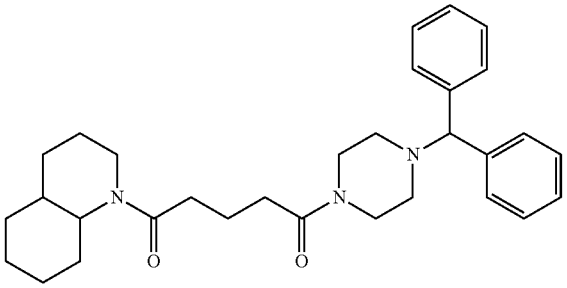
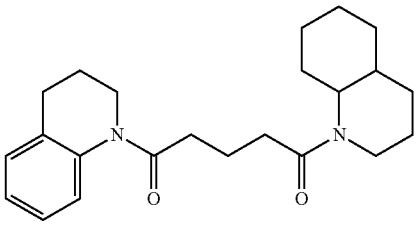
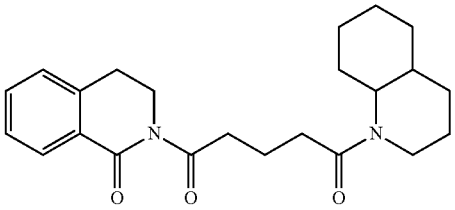
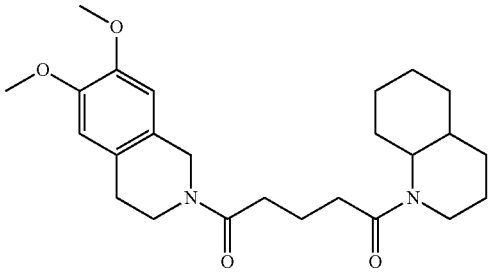
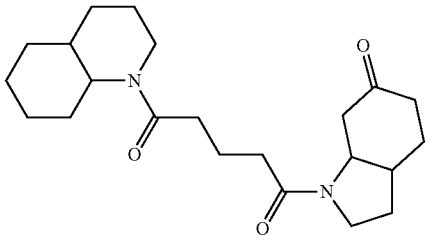
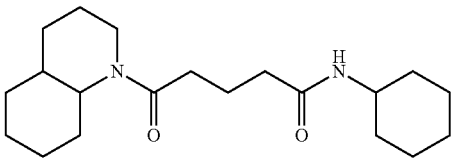
Ex.	structure	name	method	rt (min)	m/z
Ia.50		1-(4-benzhydrylpiperazin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	6.54	488
Ia.51		1-(3,4-dihydro-2-H-quinolin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	7.49	369
Ia.52		1-(octahydroquinolin-1-yl)-5-(1-oxo-3,4-dihydro-1-H-isoquinolin-2-yl)pentane-1,5-dione	A	7.75	383
Ia.53		1-(6,7-dimethoxy-3,4-dihydro-1-H-isoquinolin-2-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	6.58	429
Ia.54		1-(octahydroquinolin-1-yl)-5-(6-oxooctahydroindol-1-yl)pentane-1,5-dione	A	4.30	375
Ia.55		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid cyclohexylamide	B	3.68	335

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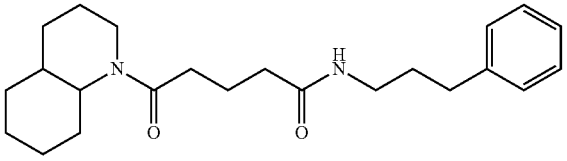
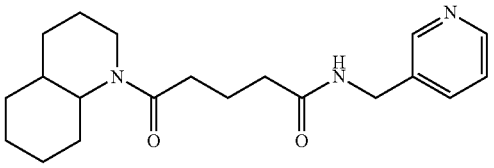
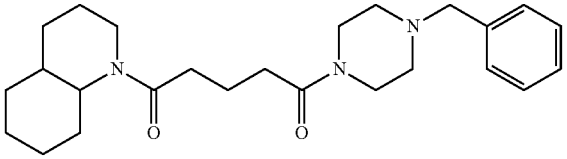
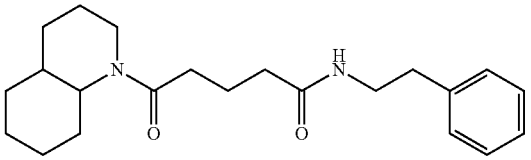
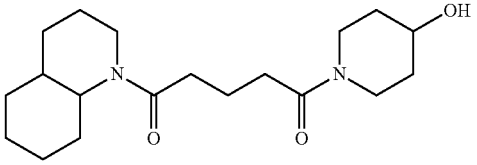
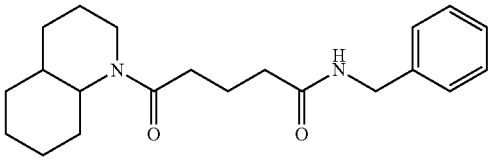
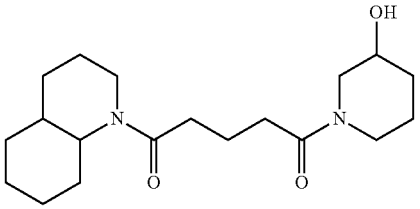
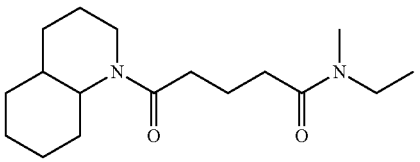
Ex.	structure	name	method	rt (min)	m/z
Ia.56		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid (3-phenylpropyl)amide	B	3.84	371
Ia.57		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid (pyridin-3-ylmethyl)amide	B	2.29	344
Ia.58		1-(4-benzylpiperazin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	B	2.63	412
Ia.59		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid phenethylamide	B	3.66	357
Ia.60		1-(4-hydroxypiperidin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	B	2.75	337
Ia.61		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid benzylamide	B	3.54	343
Ia.62		1-(3-hydroxypiperidin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	B	2.87	337
Ia.63		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid ethylmethylamide	B	3.24	295

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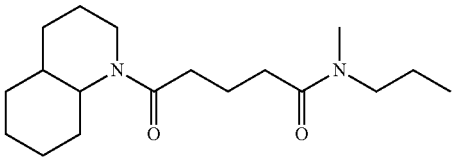
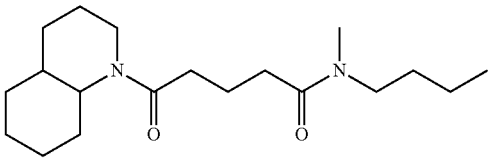
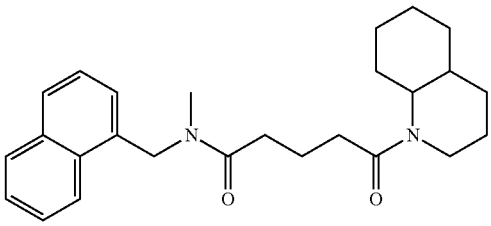
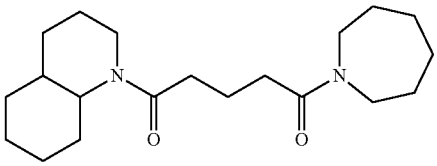
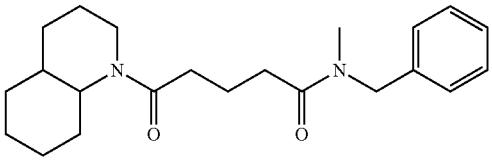
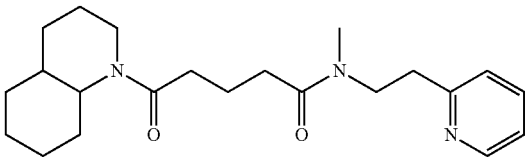
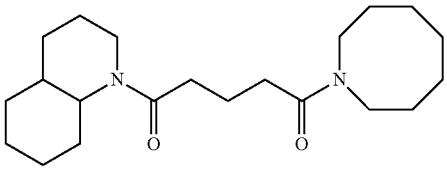
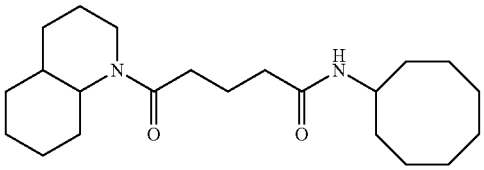
Ex.	structure	name	method	rt (min)	m/z
Ia.64		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid methylpropylamide	B	3.51	309
Ia.65		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid butylmethylamide	B	3.79	323
Ia.66		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid methylnaphthalen-1-ylmethylamide	B	4.24	407
Ia.67		5-(octahydroquinolin-1-yl)-1-perhydroazepin-1-ylpentane-1,5-dione	B	3.72	335
Ia.68		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid benzylmethylamide	B	3.84	357
Ia.69		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid methyl-(2-pyridin-2-ylethyl)amide	B	2.47	372
Ia.70		5-(octahydroquinolin-1-yl)-1-perhydroazocin-1-ylpentane-1,5-dione	B	3.84	349
Ia.71		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid cyclooctylamide	B	4.10	363

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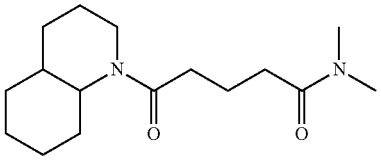
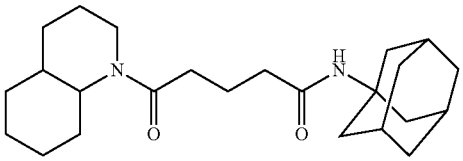
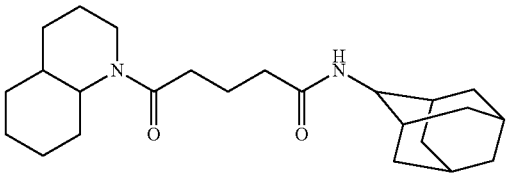
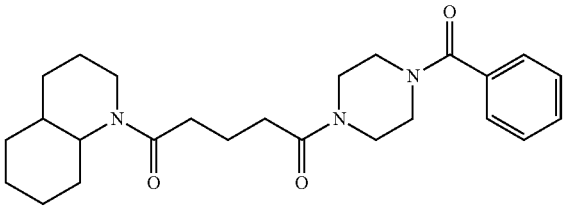
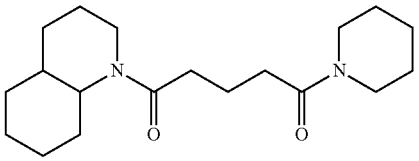
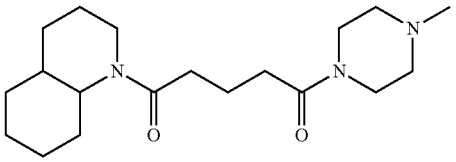
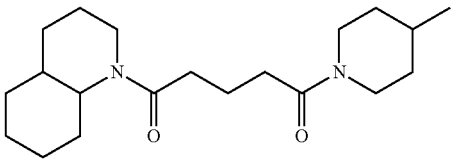
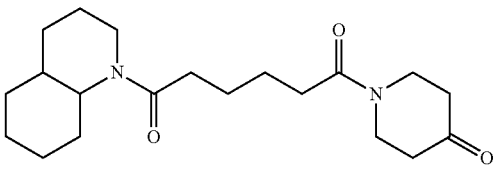
Ex.	structure	name	method	rt (min)	m/z
Ia.72		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid dimethylamide	B	3.01	381
Ia.73		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid tricyclo[3.3.1.1^{3,7}]decan-1-ylamide	B	4.35	387
Ia.74		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid tricyclo[3.3.1.1^{3,7}]decan-2-ylamide	B	4.28	387
Ia.75		1-(4-benzoylpiperazin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	B	3.34	426
Ia.76		1-(octahydroquinolin-1-yl)-5-piperidin-1-ylpentane-1,5-dione	A	6.52	321
Ia.77		1-(4-methylpiperazin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	4.20	336
Ia.78		1-(4-methylpiperidin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	A	7.02	335
Ia.79		1-(octahydroquinolin-1-yl)-6-(4-oxopiperidin-1-yl)hexane-1,6-dione	B	3.04	349

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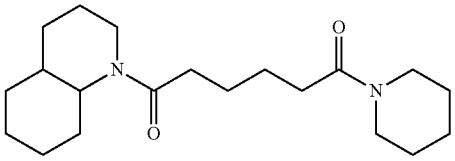
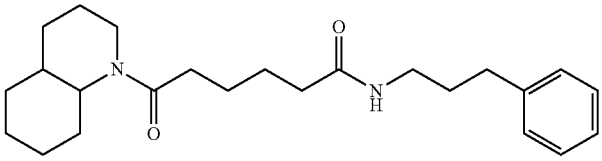
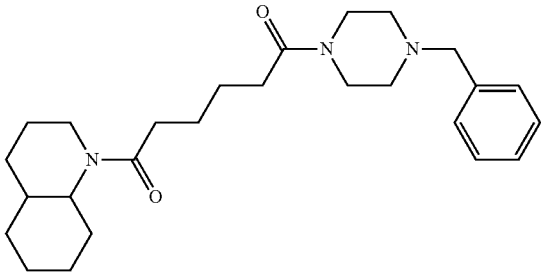
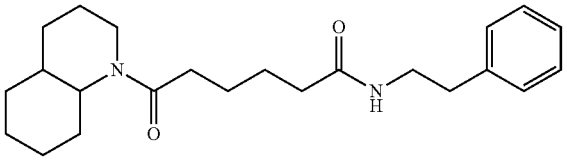
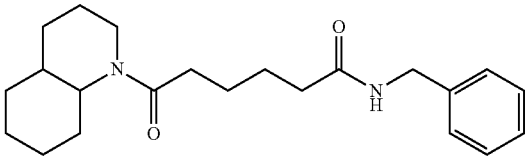
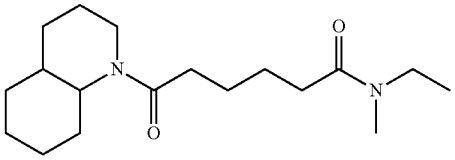
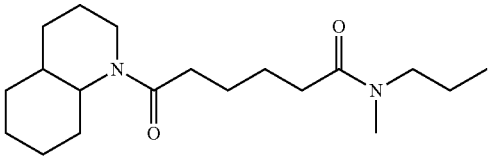
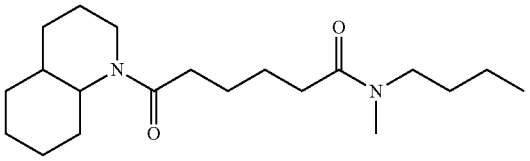
Ex.	structure	name	method	rt (min)	m/z
Ia.80		1-(octahydroquinolin-1-yl)-6-piperidin-1-ylhexane-1,6-dione	B	3.65	335
Ia.81		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid (3-phenylpropyl)amide	B	3.94	385
Ia.82		1-(4-benzylpiperazin-1-yl)-6-(octahydroquinolin-1-yl)hexane-1,6-dione	B	2.74	426
Ia.83		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid phenethylamide	B	3.76	371
Ia.84		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid benzylamide	B	3.64	357
Ia.85		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid ethylmethylamide	B	3.36	309
Ia.86		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid methylpropylamide	B	3.62	323
Ia.87		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid butylmethylamide	B	3.89	337

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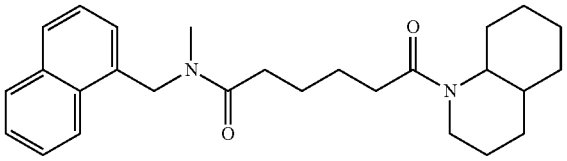
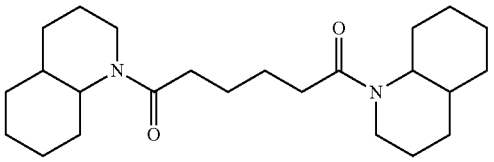
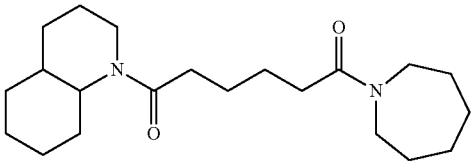
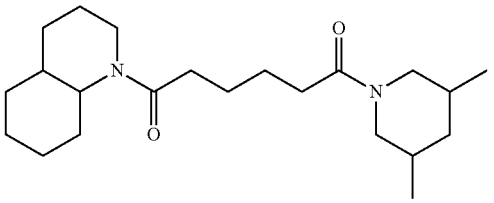
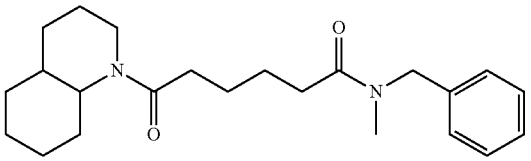
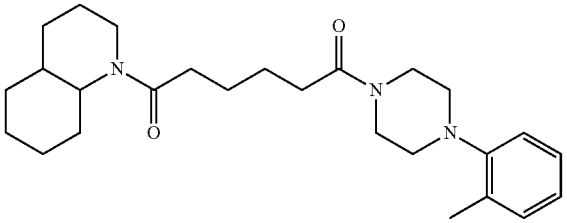
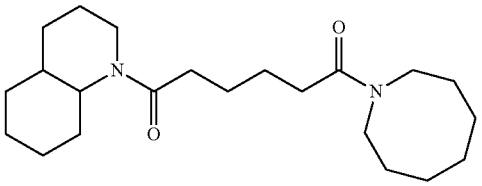
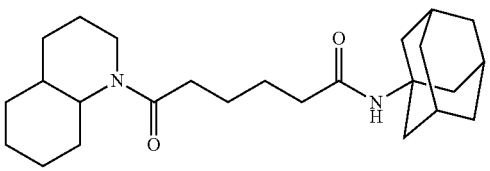
Ex.	structure	name	method	rt (min)	m/z
Ia.88		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid methylnaphthalen-1-ylmethylamide	B	4.24 4.31	421 421
Ia.89		1,6-bis-(octahydroquinolin-1-yl)hexane-1,6-dione	B	4.39	389
Ia.90		6-(octahydroquinolin-1-yl)-1-perhydroazepin-1-ylhexane-1,6-dione	B	3.83	349
Ia.91		1-(3,5-dimethylpiperidin-1-yl)-6-(octahydroquinolin-1-yl)hexane-1,6-dione	B	4.11 4.20	363 363
Ia.92		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid benzylmethylamide	B	3.93	371
Ia.93		1-(octahydroquinolin-1-yl)-6-(4-o-tolylpiperazin-1-yl)hexane-1,6-dione	B	4.37	426
Ia.94		6-(octahydroquinolin-1-yl)-1-perhydroazocin-1-ylhexane-1,6-dione	B	4.04	363
Ia.95		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid tricyclo[3.3.1.1^{3,7}]decan-1-ylamide	B	4.42	401

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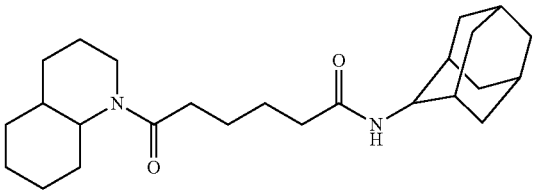
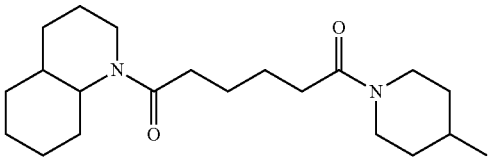
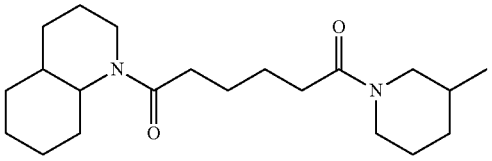
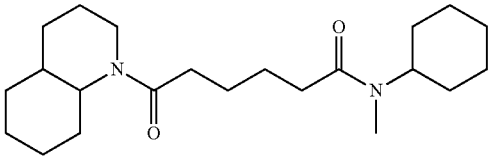
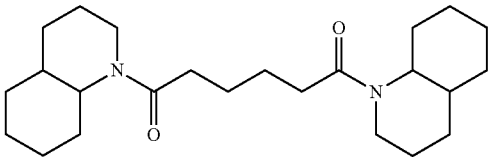
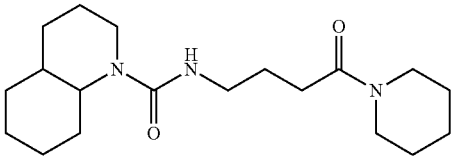
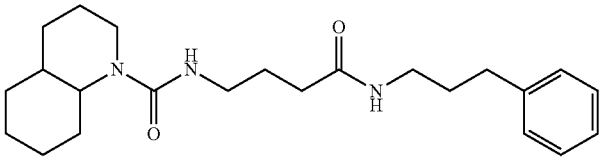
Ex.	structure	name	method	rt (min)	m/z
Ia.96		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid tricyclo[3.3.1.1^{3,7}]decan-2-ylamide	B	4.34	401
Ia.97		1-(4-methylpiperidin-1-yl)-6-(octahydroquinolin-1-yl)hexane-1,6-dione	B	3.92	349
Ia.98		1-(3-methylpiperidin-1-yl)-6-(octahydroquinolin-1-yl)hexane-1,6-dione	B	3.92	349
Ia.99		6-(octahydroquinolin-1-yl)-6-oxohexanoic acid cyclohexylmethylamide	B	4.13	363
Ia.100		1-(octahydroisoquinolin-2-yl)-6-(octahydroquinolin-1-yl)hexane-1,6-dione	B	4.38 4.47	389 389
Ib.1		Octahydroquinoline-1-carboxylic acid (4-oxo-4-piperidin-1-ylbutyl)amide	B	3.42	336
Ib.2		Octahydroquinoline-1-carboxylic acid [3-(3-phenylpropylcarbamoyl)propyl]amide	B	3.81	386

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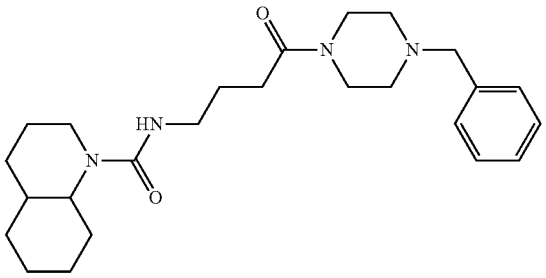
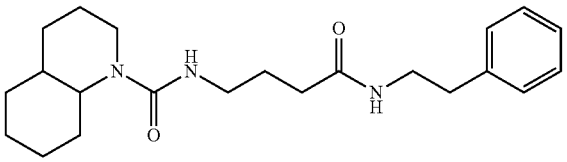
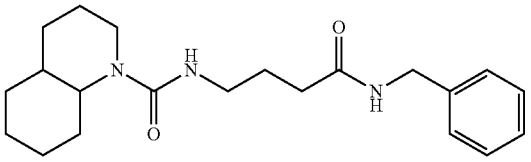
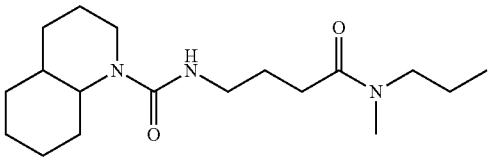
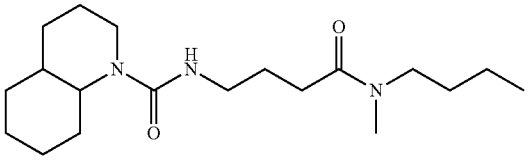
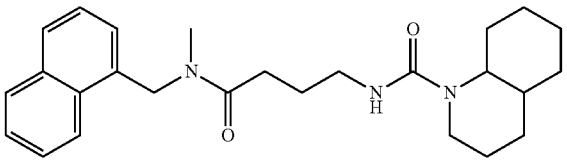
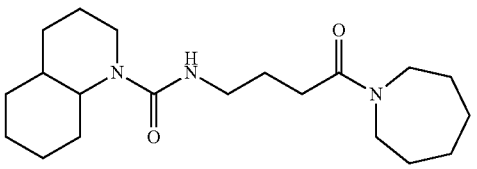
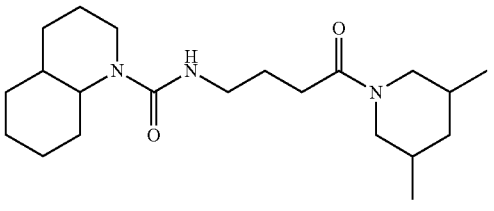
Ex.	structure	name	method	rt (min)	m/z
Ib.3		Octahydroquinoline-1-carboxylic acid [4-(4-benzylpiperazin-1-yl)-4-oxobutyl]amide	B	2.61	427
Ib.4		Octahydroquinoline-1-carboxylic acid (3-phenethylcarbamoyl propyl)amide	B	3.61	372
Ib.5		Octahydroquinoline-1-carboxylic acid (3-benzylcarbamoyl-propyl)amide	B	3.50	358
Ib.6		Octahydroquinoline-1-carboxylic acid [3-(methylpropyl-carbamoyl)propyl]amide	B	3.39	324
Ib.7		Octahydroquinoline-1-carboxylic acid [3-(butylmethylcar-bamoyl)propyl]amide	B	3.65	338
Ib.8		Octahydroquinoline-1-carboxylic acid [3-(methylnaphthalen-1-ylmethylcarbamoyl) propyl]amide	B	4.08	422
Ib.9		Octahydroquinoline-1-carboxylic acid (4-oxo-4-perhydroazepin-1-ylbutyl)amide	B	3.59	350
Ib.10		Octahydroquinoline-1-carboxylic acid [4-(3,5-dimethylpiperidin-1-yl)-4-oxobutyl]amide	B	3.87 3.96	364 364

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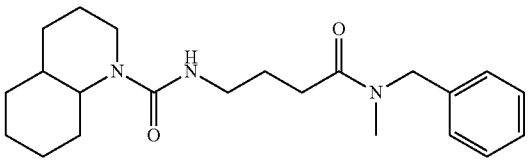
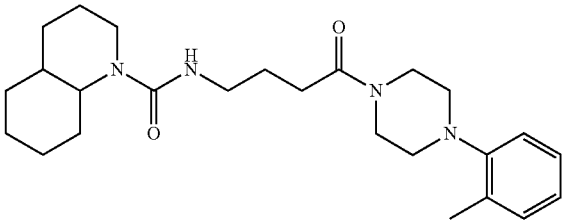
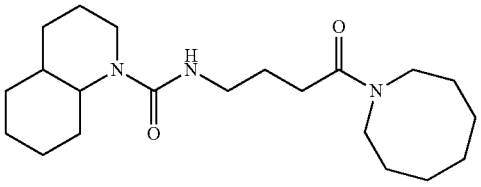
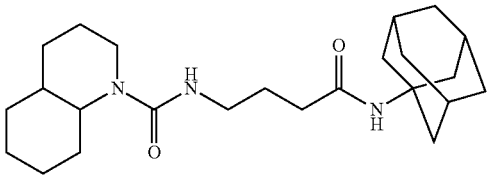
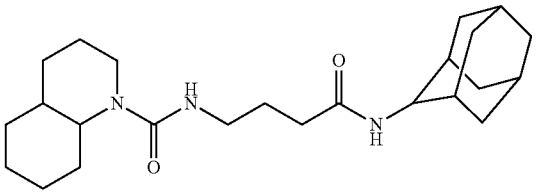
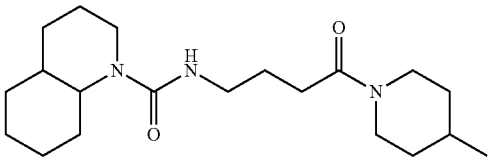
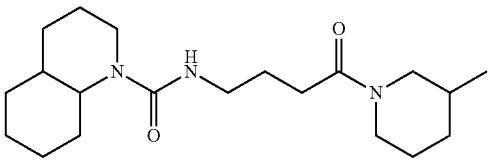
Ex.	structure	name	method	rt (min)	m/z
Ib.11		Octahydroquinoline-1-carboxylic acid [3-(benzylmethylcarbamoyl)propyl]amide	B	3.71	372
Ib.12		Octahydroquinoline-1-carboxylic acid [4-oxo-4-(4-o-tolylpiperazin-1-yl)butyl]amide	B	4.14	427
Ib.13		Octahydroquinoline-1-carboxylic acid (4-oxo-4-perhydroazocin-1-ylbutyl)amide	B	3.80	364
Ib.14		Octahydroquinoline-1-carboxylic acid [3-(tricyclo[3.3.1.1^{3,7}]decan-1-ylcarbamoyl)propyl]amide	B	4.31	402
Ib.15		Octahydroquinoline-1-carboxylic acid [3-(tricyclo[3.3.1.1^{3,7}]decan-2-ylcarbamoyl)propyl]amide	B	4.23	402
Ib.16		Octahydroquinoline-1-carboxylic acid [4-(4-methylpiperidin-1-yl)-4-oxobutyl]amide	B	3.69	350
Ib.17		Octahydroquinoline-1-carboxylic acid [4-(3-methylpiperidin-1-yl)-4-oxobutyl]amide	B	3.68	350

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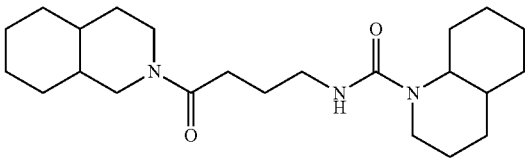
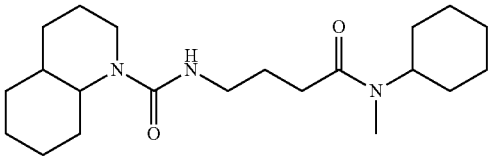
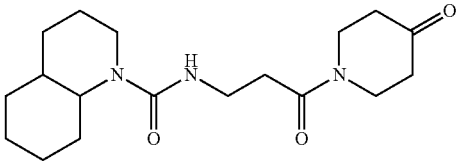
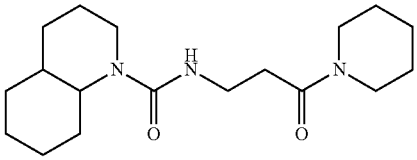
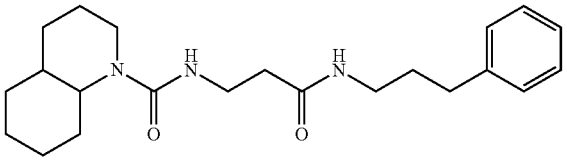
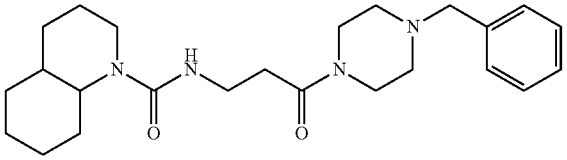
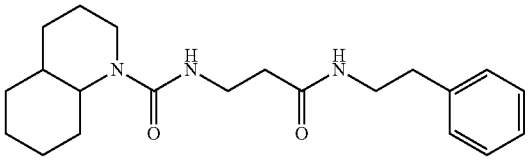
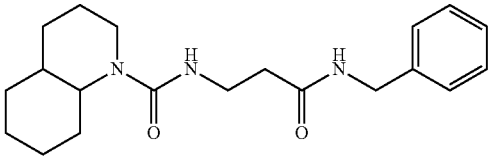
Ex.	structure	name	method	rt (min)	m/z
Ib.18		Octahydroquinoline-1-carboxylic acid [4-(octahydroisoquinolin-2-yl)-4-oxobutyl]amide	B	4.14 4.24	390 390
Ib.19		Octahydroquinoline-1-carboxylic acid [3-(cyclohexylmethylcarbamoyl)propyl]amide	B	3.89	364
Ib.20		Octahydroquinoline-1-carboxylic acid [3-oxo-3-(4-oxopiperidin-1-yl)propyl]amide	B	2.78	336
Ib.21		Octahydroquinoline-1-carboxylic acid (3-oxo-3-piperidin-1-ylpropyl)amide	B	3.35	322
Ib.22		Octahydroquinoline-1-carboxylic acid [2-(3-phenylpropylcarbamoyl)ethyl]amide	B	3.72	372
Ib.23		Octahydroquinoline-1-carboxylic acid [3-(4-benzylpiperazin-1-yl)-3-oxopropyl]amide	B	2.56	413
Ib.24		Octahydroquinoline-1-carboxylic acid (2-phenethylcarbamoylethyl)amide	B	3.56	358
Ib.25		Octahydroquinoline-1-carboxylic acid (2-benzylcarbamoylethyl)amide	B	3.42	344

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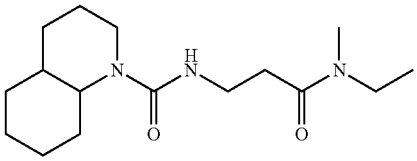
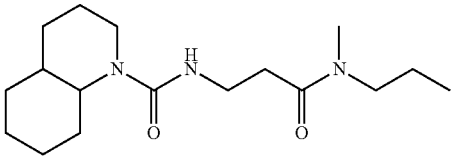
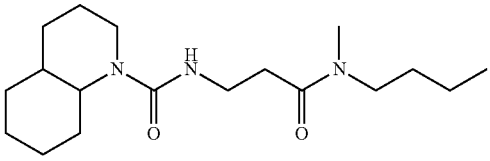
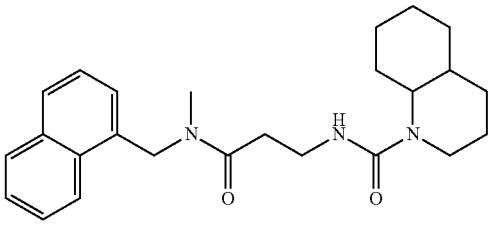
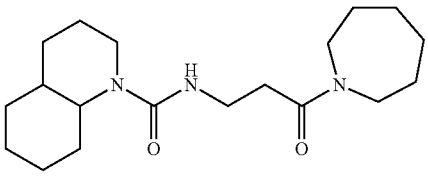
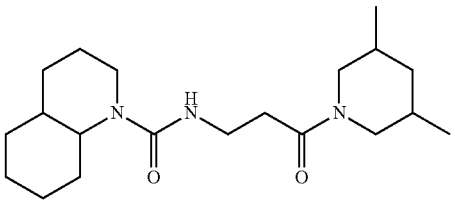
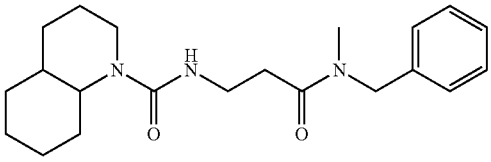
Ex.	structure	name	method	rt (min)	m/z
Ib.26		Octahydroquinoline-1-carboxylic acid [2-(ethylmethylcarbamoyl)ethyl]amide	B	3.07	296
Ib.27		Octahydroquinoline-1-carboxylic acid [2-(methylpropylcarbamoyl)ethyl]amide	B	3.33	310
Ib.28		Octahydroquinoline-1-carboxylic acid [2-(butylmethylcarbamoyl)ethyl]amide	B	3.60	324
Ib.29		Octahydroquinoline-1-carboxylic acid [2-(methylnaphthalen-1-ylmethylcarbamoyl)ethyl]amide	B	4.08	408
Ib.30		Octahydroquinoline-1-carboxylic acid (3-oxo-3-perhydroazepin-1-ylpropyl)amide	B	3.52	336
Ib.31		Octahydroquinoline-1-carboxylic acid [3-(3,5-dimethylpiperidin-1-yl)-3-oxopropyl]amide	B	3.82 3.90	350 350
Ib.32		Octahydroquinoline-1-carboxylic acid [2-(benzylmethylcarbamoyl)ethyl]amide	B	3.67	358

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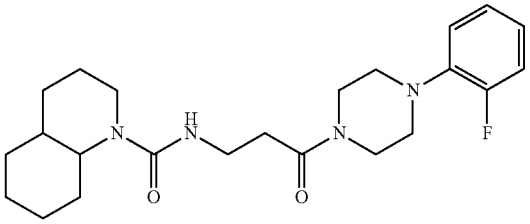
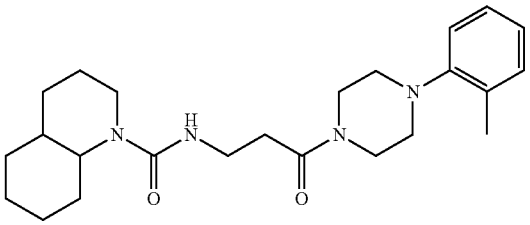
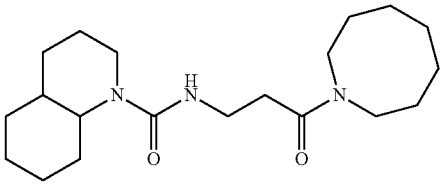
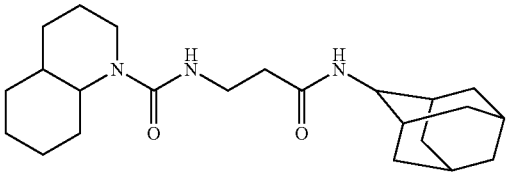
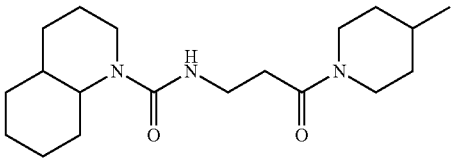
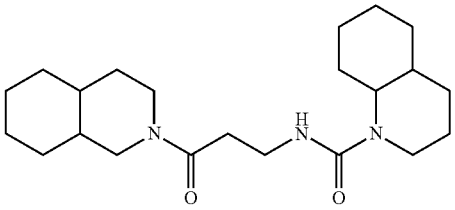
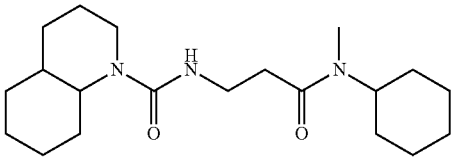
Ex.	structure	name	method	rt (min)	m/z
Ib.33		Octahydroquinoline-1-carboxylic acid {3-[4-(2-fluorophenyl)piperazin-1-yl]-3-oxopropyl}amide	B	3.81	417
Ib.34		Octahydroquinoline-1-carboxylic acid [3-oxo-3-(4-o-tolylpiperazin-1-yl)propyl]amide	B	4.09	413
Ib.35		Octahydroquinoline-1-carboxylic acid (3-oxo-3-perhydroazocin-1-ylpropyl)amide	B	3.74	350
Ib.36		Octahydroquinoline-1-carboxylic acid [2-(tricyclo[3.3.1.1^{3,7}]decan-2-ylcarbamoyl)ethyl] amide	B	4.11	388
Ib.37		Octahydroquinoline-1-carboxylic acid [3-(4-methylpiperidin-1-yl)-3-oxopropyl]amide	B	3.62	336
Ib.38		Octahydroquinoline-1-carboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	B	4.08 4.17	376 376
Ib.39		Octahydroquinoline-1-carboxylic acid[2-(cyclohexylmethyl-carbamoyl)ethyl]amide	B	3.83	350

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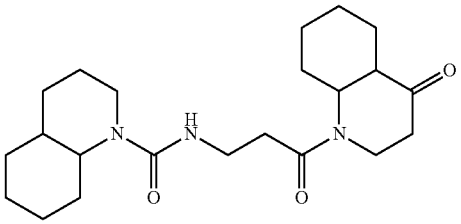
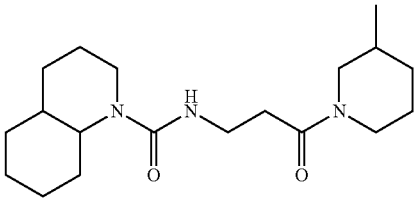
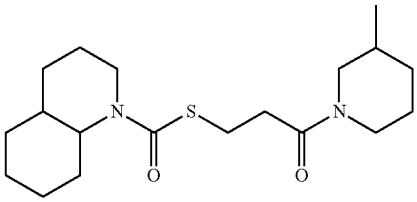
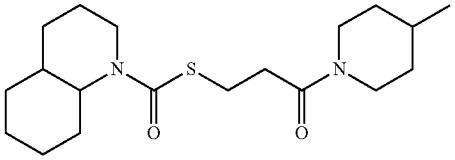
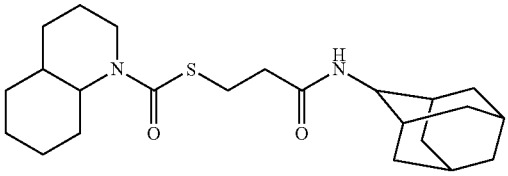
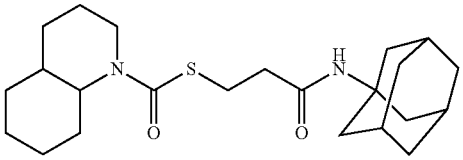
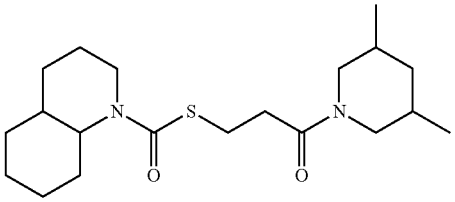
Ex.	structure	name	method	rt (min)	m/z
Ib.40		Octahydroquinoline-1-carboxylic acid [3-oxo-3-(4-oxooctahydroquinolin-1-yl)propyl]amide	B	3.44	390
Ib.41		Octahydroquinoline-1-carboxylic acid [3-(3-methylpiperidin-1-yl)-3-oxopropyl]amide	B	3.61	336
Ib.42		Octahydroquinoline-1-carbothioic acid S-[3-(3-methylpiperidin-1-yl)-3-oxopropyl] ester	B	4.55	353
Ib.43		Octahydroquinoline-1-carbothioic acid S-[3-(4-methylpiperidin-1-yl)-3-oxopropyl] ester	B	3.57	353
Ib.44		Octahydroquinoline-1-carbothioic acid S-[2-(tricyclo[3.3.1.1^{3,7}]decan-2-ylcarbamoyl)ethyl) ester	B	4.79	405
Ib.45		Octahydroquinoline-1-carbothioic acid S-[2-(tricyclo[3.3.1.1^{3,7}]decan-ylcarbamoyl)ethyl] ester	B	4.88	405
Ib.46		Octahydroquinoline-1-carbothioic acid S-[3-(3,5-dimethylpiperidin-1-yl)-3-oxopropyl] ester	B	4.75	367

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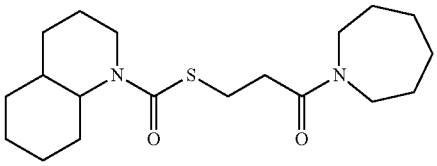
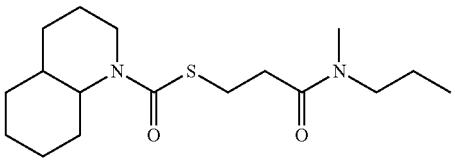
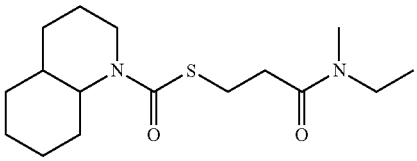
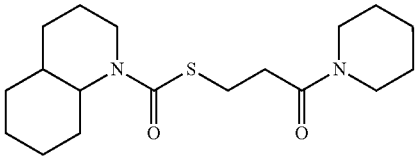
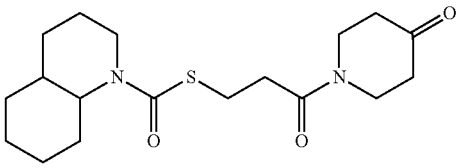
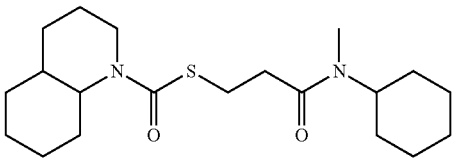
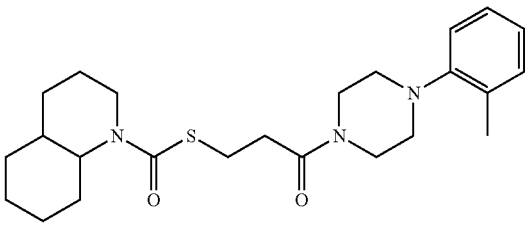
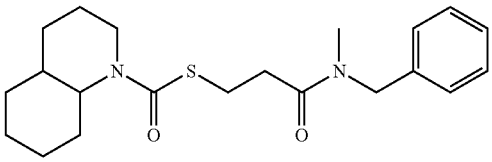
Ex.	structure	name	method	rt (min)	m/z
Ib.47		Octahydroquinoline-1-carbothioic acid S-(3-oxo-3-perhydroazepin-1-ylpropyl) ester	B	4.56	353
Ib.48		Octahydroquinoline-1-carbothioic acid S-[2-(methylpropylcarbamoyl)ethyl] ester	B	4.14	327
Ib.49		Octahydroquinoline-1-carbothioic acid S-[2-(ethylmethylcarbamoyl)ethyl] ester	B	3.88	313
Ib.50		Octahydroquinoline-1-carbothioic acid S-(3-oxo-3-piperidin-1-ylpropyl) ester	B	4.20	339
Ib.51		Octahydroquinoline-1-carbothioic acid S-[3-oxo-3-(4-oxopiperidin-1-yl)propyl] ester	B	3.49	353
Ib.52		Octahydroquinoline-1-carbothioic acid S-[2-(cyclohexylmethylcarbamoyl)ethyl] ester	B	4.67	364
Ib.53		Octahydroquinoline-1-carbothioic acid S-[3-oxo-3-(4-o-tolylpiperazin-1-yl]propyl] ester	B	4.86	430
Ib.54		Octahydroquinoline-1-carbothioic acid S-[2-(benzylmethylcarbamoyl)ethyl] ester	B	4.42	375

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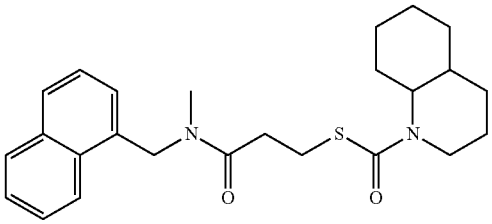
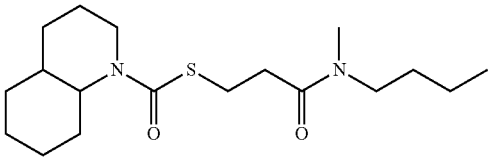
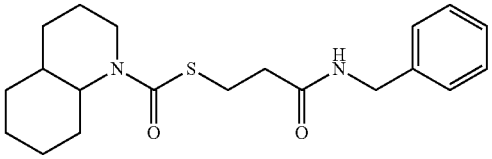
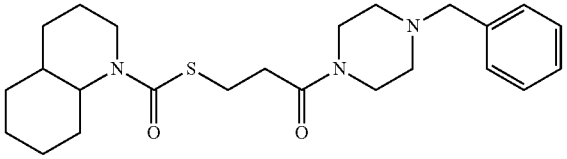
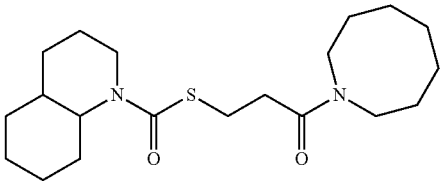
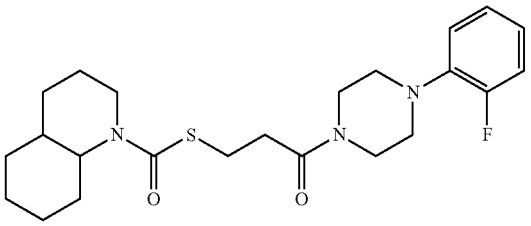
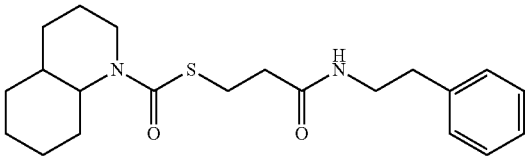
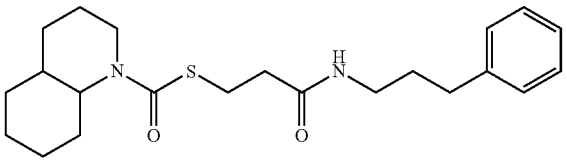
Ex.	structure	name	method	rt (min)	m/z
Ib.55		Octahydroquinoline-1-carbothioic acid S-[2-(methylnaphthalen-1-ylmethylcarbamoyl) ethyl] ester	B	4.78	425
Ib.56		Octahydroquinoline-1-carbothioic acid S-[2-(butylmethylcar-bamoyl)ethyl] ester	B	4.41	341
Ib.57		Octahydroquinoline-1-carbothioic acid S-(2-benzylcarbamoyl-ethyl) ester	B	4.06	361
Ib.58		Octahydroquinoline-1-carbothioic acid S-[3-(4-benzylpiperazin-1-yl)-3-oxopropyl] ester	B	2.98	430
Ib.59		Octahydroquinoline-1-carbothioic acid S-(3-oxo-3-perhydroazocin-1-ylpropyl) ester	B	4.58	367
Ib.60		Octahydroquinoline-1-carbothioic acid S-{3-[4-(2-fluorophenyl)piperazin-1-yl]-3-oxopropyl} ester	B	4.59	434
Ib.61		Octahydroquinoline-1-carbothioic acid S-(2-phenethylcarbamoyl ethyl) ester	B	4.20	375
Ib.62		Octahydroquinoline-1-carbothioic acid S-[2-(3-phenylpropylcar-bamoyl)ethyl] ester	B	4.34	389

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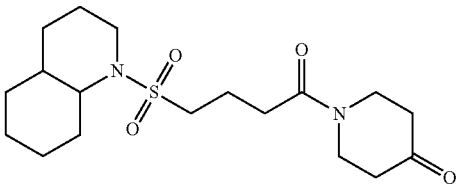
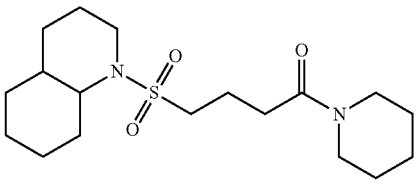
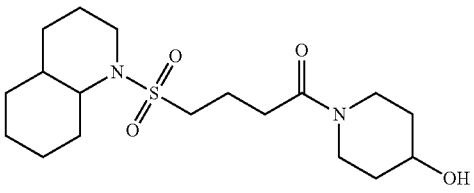
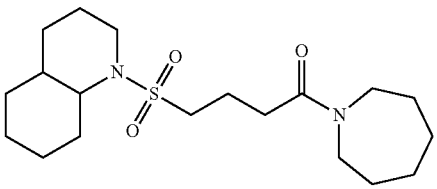
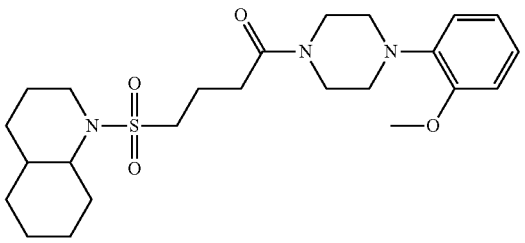
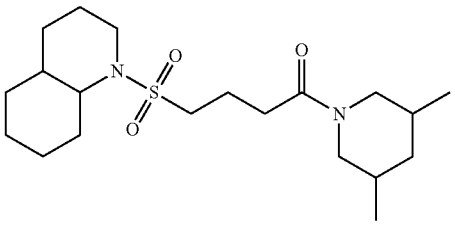
Ex.	structure	name	method	rt (min)	m/z
Im.1		1-[4-(octahydroquinoline-1-sulfonyl)butyryl]piperidin-4-one	B	3.16	371
Im.2		4-(octahydroquinoline-1-sulfonyl)-1-piperidin-1-ylbutan-1-one	B	3.79	357
Im.3		1-(4-hydroxypiperidin-1-yl)-4-(octahydroquinoline-1-sulfonyl)butan-1-one	B	2.96	373
Im.4		4-(octahydroquinoline-1-sulfonyl)-1-perhydroazepin-1-ylbutan-1-one	B	3.96	371
Im.5		1-[4-(2-methoxyphenyl)piperazin-1-yl]-4-(octahydroquinoline-1-sulfonyl)butan-1-one	B	3.92	464
Im.6		1-(3,5-dimethylpiperidin-1-yl)-4-(octahydroquinoline-1-sulfonyl)butan-1-one	B	4.34	385

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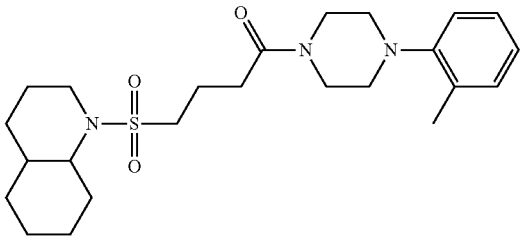
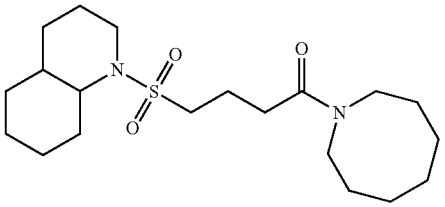
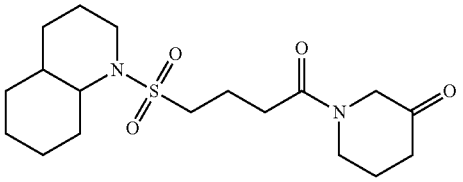
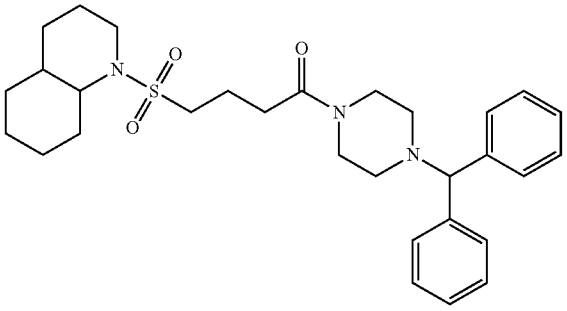
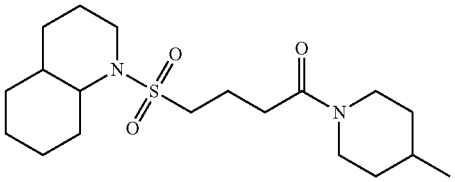
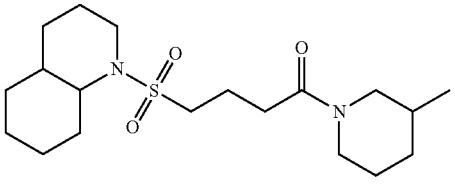
Ex.	structure	name	method	rt (min)	m/z
Im.7		4-(octahydroquinoline-1-sulfonyl)-1-(4-o-tolylpiperazin-1-yl)butan-1-one	B	4.49	448
Im.8		4-(octahydroquinoline-1-sulfonyl)-1-perhydroazocin-1-ylbutan-1-one	B	4.17	385
Im.9		1-((R)-3-hydroxypiperidin-1-yl)-4-(octahydroquinoline-1-sulfonyl)butan-1-one	B	3.07	373
Im.10		1-(4-benzhydrylpiperazin-1-yl)-4-(octahydroquinoline-1-sulfonyl)butan-1-one	B	3.82	524
Im.11		1-(4-methylpiperidin-1-yl)-4-(octahydroquinoline-1-sulfonyl)butan-1-one	B	4.06	371
Im.12		1-(3-methylpiperidin-1-yl)-4-(octahydroquinoline-1-sulfonyl)butan-1-one	B	4.05	371

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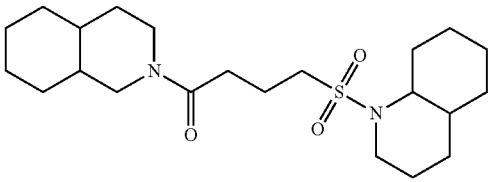
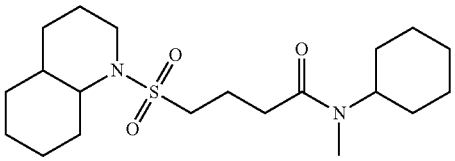
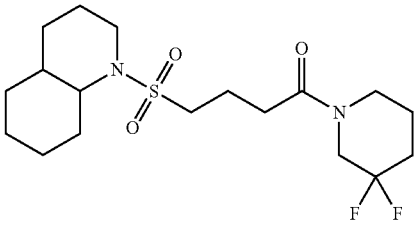
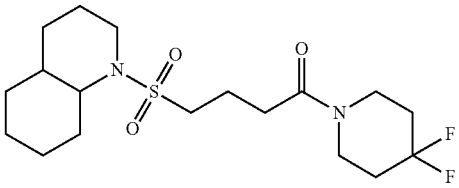
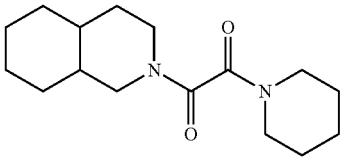
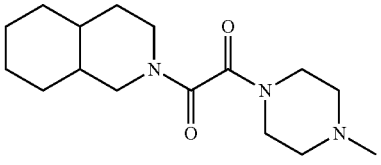
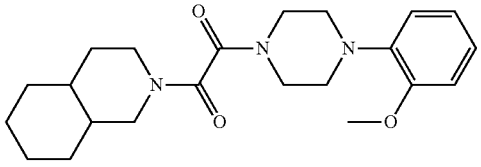
Ex.	structure	name	method	rt (min)	m/z
Im.13		1-(octahydroisoquinolin-2-yl)-4-(octahydroquinoline-1-sulfonyl)butan-1-one	B	4.50 4.59	411 411
Im.14		N-cyclohexyl-N-methyl-4-(octahydroquinoline-1-sulfonyl)butyramide	B	4.26	385
Im.15		1-(3,3-difluoropiperidin-1-yl)-4-(octahydroquinoline-1-sulfonyl)butan-1-one	B	3.80	393
Im.16		1-(4,4-difluoropiperidin-1-yl)-4-(octahydroquinoline-1-sulfonyl)butan-1-one	B	3.81	393
Ia.101		1-(octahydroisoquinolin-2-yl)-2-piperidin-1-ylethane-1,2-dione	A	6.28 6.45	279 279
Ia.102		1-(4-methylpiperazin-1-yl)-2-(octahydroisoquinolin-2-yl)ethane-1,2-dione	A	4.04	294
Ia.103		1-[4-(2-methoxyphenyl)piperazin-1-yl]-2-(octahydroisoquinolin-2-yl)ethane-1,2-dione	A	7.03 7.17	386 386

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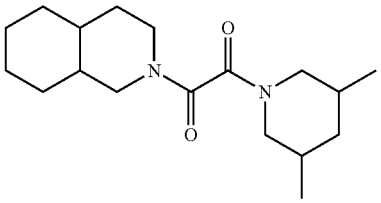
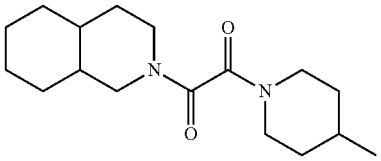
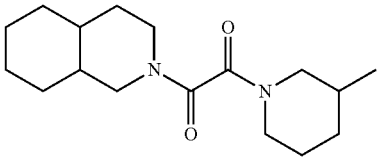
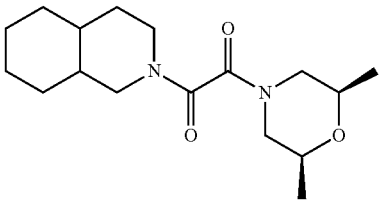
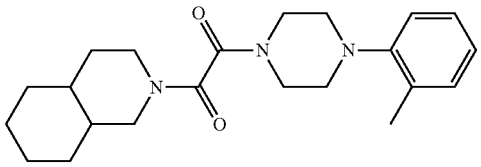
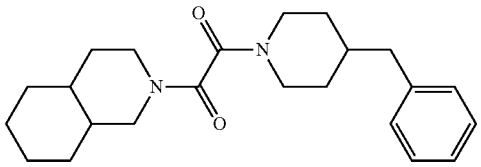
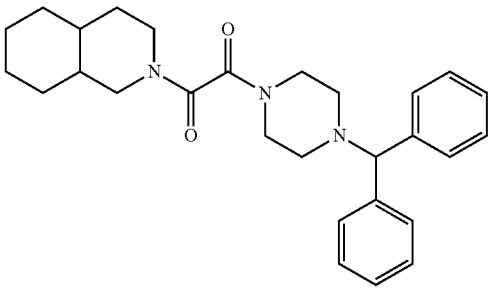
Ex.	structure	name	method	rt (min)	m/z
Ia.104		1-(3,5-dimethylpiperidin-1-yl)-2-(octahydroisoquinolin-2-yl)ethane-1,2-dione	A	7.41 7.53	307 307
Ia.105		1-(4-methylpiperidin-1-yl)-2-(octahydroisoquinolin-2-yl)ethane-1,2-dione	A	6.86 7.01	293 293
Ia.106		1-(3-methylpiperidin-1-yl)-2-(octahydroisoquinolin-2-yl)ethane-1,2-dione	A	6.83 6.99	293 293
Ia.107		1-((2S,6R)-2,6-dimethylmorpholin-4-yl)-2-(octahydroisoquinolin-2-yl)ethane-1,2-dione	A	6.20 6.38	309 309
Ia.108		1-(octahydroisoquinolin-2-yl)-2-(4-o-tolylpiperazin-1-yl)ethane-1,2-dione	A	7.98	370
Ia.109		1-(4-benzylpiperidin-1-yl)-2-(octahydroisoquinolin-2-yl)ethane-1,2-dione	A	8.04	369
Ia.110		1-(4-benzhydrylpiperazin-1-yl)-2-(octahydroisoquinolin-2-yl)ethane-1,2-dione	A	7.63 7.80	446 446

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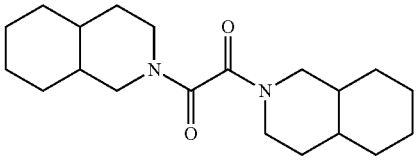
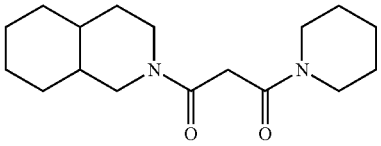
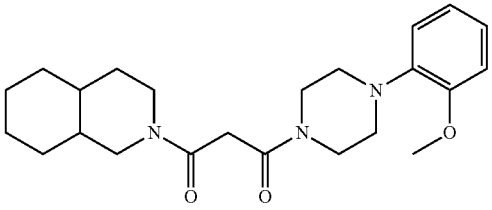
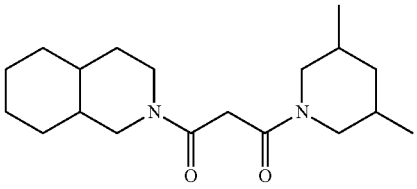
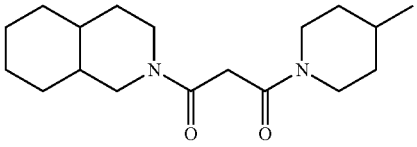
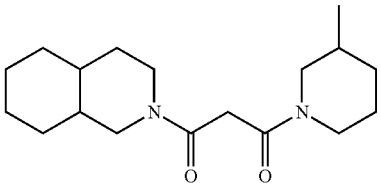
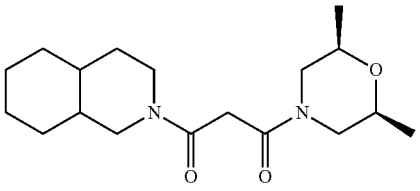
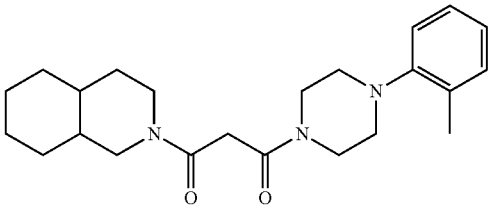
Ex.	structure	name	method	rt (min)	m/z
Ia.111		1,2-bis-(octahydroisoquinolin-2-yl)ethane-1,2-dione	A	8.06	333
Ia.112		1-(octahydroisoquinolin-2-yl)-3-piperidin-1-ylpropane-1,3-dione	A	6.11 6.28	293 293
Ia.113		1-[4-(2-methoxyphenyl)piperazin-1-yl]-3-(octahydroisoquinolin-2-yl)propane-1,3-dione	A	6.47 6.64	400 400
Ia.114		1-(3,5-dimethylpiperidin-1-yl)-3-(octahydroisoquinolin-2-yl)propane-1,3-dione	A	7.20 7.33	321 321
Ia.115		1-(4-methylpiperidin-1-yl)-3-(octahydroisoquinolin-2-yl)propane-1,3-dione	A	6.66 6.81	307 307
Ia.116		1-(3-methylpiperidin-1-yl)-3-(octahydroisoquinolin-2-yl)propane-1,3-dione	A	6.63 6.79	307 307
Ia.117		1-((2S,6R)-2,6-dimethylmorpholin-4-yl)-3-(octahydroisoquinolin-2-yl)propane-1,3-dione	A	5.91 6.09	323 323
Ia.118		1-(octahydroisoquinolin-2-yl)-3-(4-o-tolylpiperazin-1-yl)propane-1,3-dione	A	7.57 7.70	384 384

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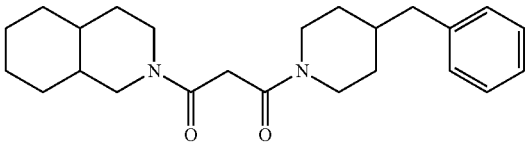
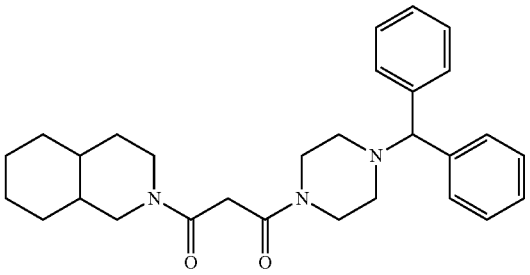
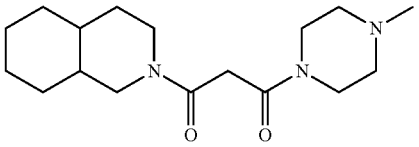
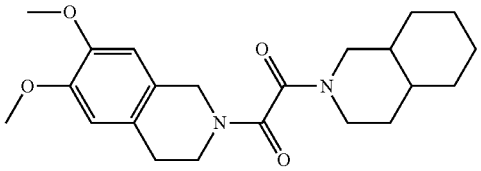
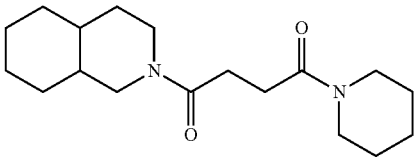
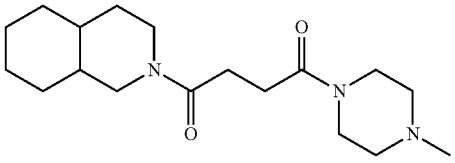
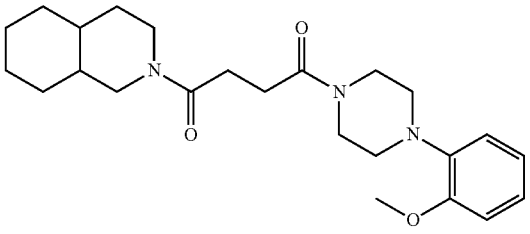
Ex.	structure	name	method	rt (min)	m/z
Ia.119		1-(4-benzylpiperidin-1-yl)-3-(octahydroisoquinolin-2-yl)propane-1,3-dione	A	7.73 7.85	383 383
Ia.120		1-(4-benzhydrylpiperazin-1-yl)-3-(octahydroisoquinolin-2-yl)propane-1,3-dione	A	6.34 6.48	460 460
Ia.121		1-(4-methylpiperazin-1-yl)-3-(octahydroisoquinolin-2-yl)propane-1,3-dione	A	4.06	308
Ia.122		1-(6,7-dimethoxy-3,4-dihydro-1-H-isoquinolin-2-yl)-2-(octahydroisoquinolin-2-yl)ethane-1,2-dione	A	6.65	387
Ia.123		1-(octahydroisoquinolin-2-yl)-4-piperidin-1-ylbutane-1,4-dione	A	6.37 6.56	307 307
Ia.124		1-(4-methylpiperazin-1-yl)-4-(octahydroisoquinolin-2-yl)butane-1,4-dione	A	4.32	322
Ia.125		1-[4-(2-methoxyphenyl)piperazin-1-yl]-4-(octahydroisoquinolin-2-yl)butane-1,4-dione	A	6.52 6.71	414 414

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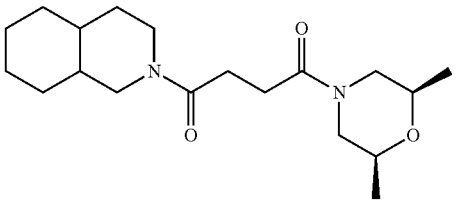
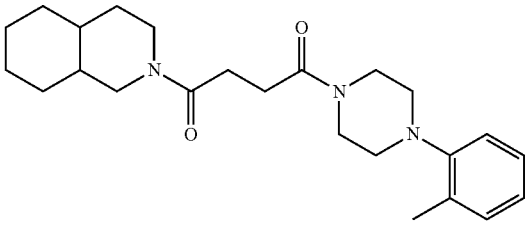
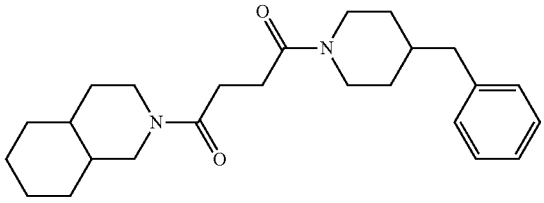
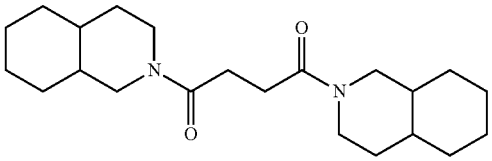
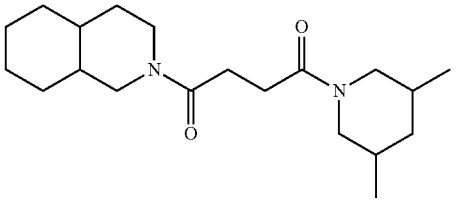
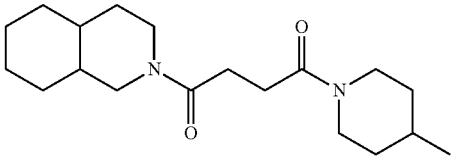
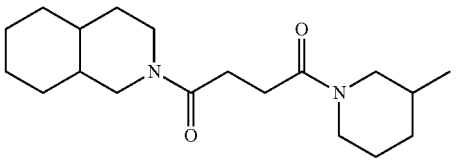
Ex.	structure	name	method	rt (min)	m/z
Ia.126		1-((2S,6R)-2,6-dimethylmorpholin-4-yl)-4-(octahydroisoquinolin-2-yl)butane-1,4-dione	A	6.15 6.34	337 337
Ia.127		1-(octahydroisoquinolin-2-yl)-4-(4-o-tolylpiperazin-1-yl)butane-1,4-dione	A	7.66 7.81	398 398
Ia.128		1-(4-benzylpiperidin-1-yl)-4-(octahydroisoquinolin-2-yl)butane-1,4-dione	A	7.90 8.03	397 397
Ia.129		1,4-bis-(octahydroisoquinolin-2-yl)butane-1,4-dione	A	7.94 8.09	361 361
Ia.130		1-(3,5-dimethylpiperidin-1-yl)-4-(octahydroisoquinolin-2-yl)butane-1,4-dione	A	7.44 7.59	335 335
Ia.131		1-(4-methylpiperidin-1-yl)-4-(octahydroisoquinolin-2-yl)butane-1,4-dione	A	6.91 7.08	321 321
Ia.132		1-(3-methylpiperidin-1-yl)-4-(octahydroisoquinolin-2-yl)butane-1,4-dione	A	6.89 7.09	321 321

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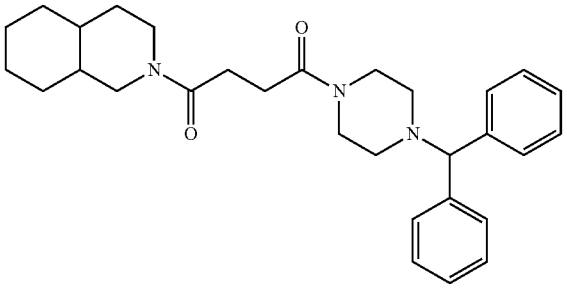
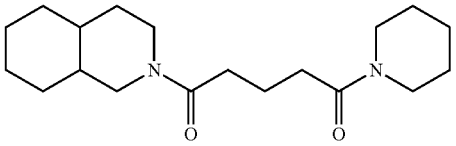
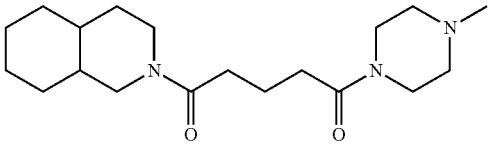
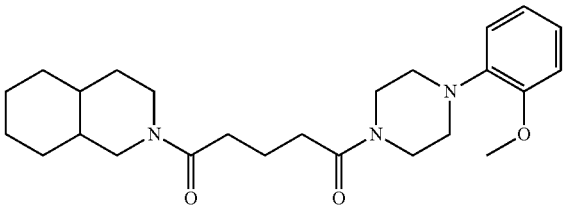
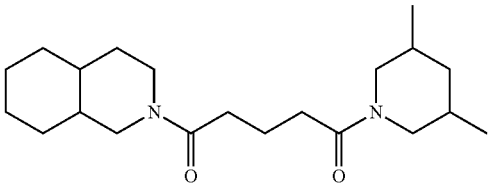
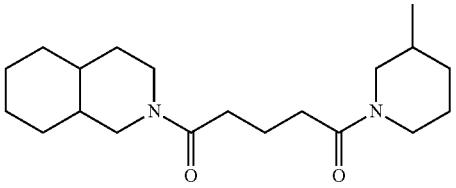
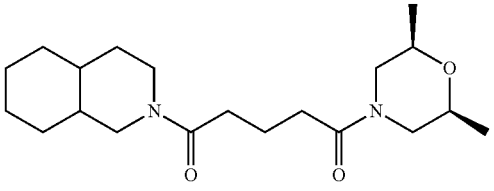
Ex.	structure	name	method	rt (min)	m/z
Ia.133		1-(4-benzhydrylpiperazin-1-yl)-4-(octahydroisoquinolin-2-yl)butane-1,4-dione	A	6.24 6.34	474 474
Ia.134		1-(octahydroisoquinolin-2-yl)-5-piperidin-1-ylpentane-1,5-dione	A	6.51 6.70	321 321
Ia.135		1-(4-methylpiperazin-1-yl)-5-(octahydroisoquinolin-2-yl)pentane-1,5-dione	A	4.27	336
Ia.136		1-[4-(2-methoxyphenyl)piperazin-1-yl]-5-(octahydroisoquinolin-2-yl)pentane-1,5-dione	A	6.74 6.93	428 428
Ia.137		1-(3,5 dimethylpiperidin-1-yl)-5-(octahydroisoquinolin-2-yl)pentane-1,5-dione	A	7.52 7.68	349 349
Ia.138		1-(3-methylpiperidin-1-yl)-5-(octahydroisoquinolin-2-yl)pentane-1,5-dione	A	7.01 7.19	335 335
Ia.139		1-((2R, 6R)-2,6-dimethylmorpholin-4-yl)-5-(octahydroisoquinolin-2-yl)pentane-1,5-dione	A	6.26 6.45	351 351

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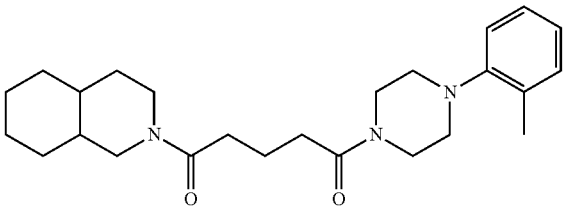
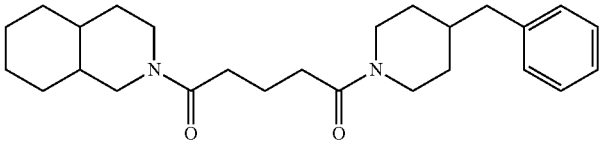
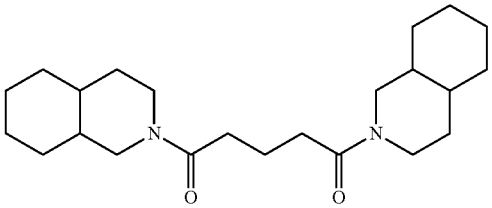
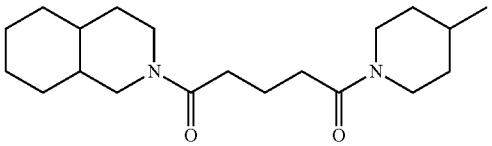
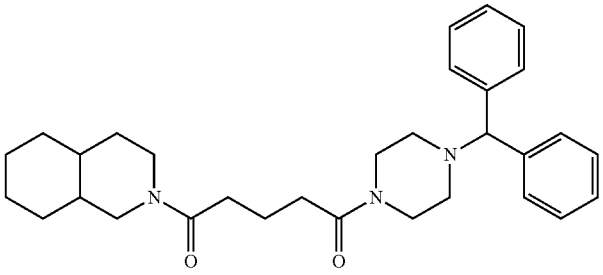
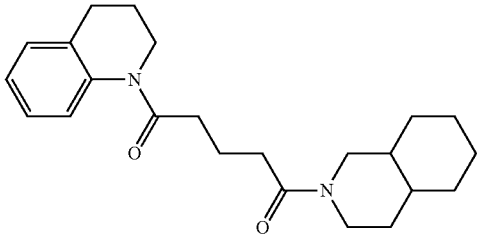
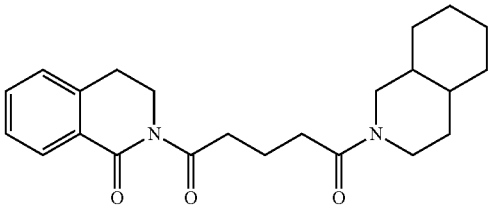
Ex.	structure	name	method	rt (min)	m/z
Ia.140		1-(octahydroisoquinolin-2-yl)-5-(4-o-tolylpiperazin-1-yl)pentane-1,5-dione	A	7.85 8.01	412 412
Ia.141		1-(4-benzylpiperidin-1-yl)-5-(octahydroisoquinolin-2-yl)pentane-1,5-dione	A	8.01 8.16	411 411
Ia.142		1,5-bis-(octahydroisoquinolin-2-yl)pentane-1,5-dione	A	8.04 8.21	375 375
Ia.143		1-(4-methylpiperidin-1-yl)-5-(octahydroisoquinolin-2-yl)pentane-1,5-dione	A	7.03 7.22	335 335
Ia.144		1-(4-benzhydrylpiperazin-1-yl)-5-(octahydroisoquinolin-2-yl)pentane-1,5-dione	A	6.53 6.67	488 448
Ia.145		1-(3,4-dihydro-2-H-quinolin-1-yl)-5-(octahydroisoquinolin-2-yl)pentane-1,5-dione	A	7.49 7.66	369 370
Ia.146		1-(octahydroisoquinolin-2-yl)-5-(1-oxo-3,4-dihydro-1-H-isoquinolin-2-yl)pentane-1,5-dione	A	7.73 7.90	383 383

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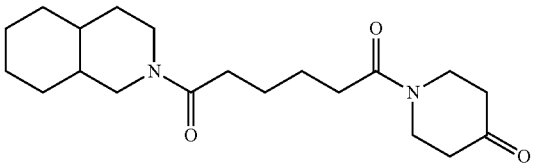
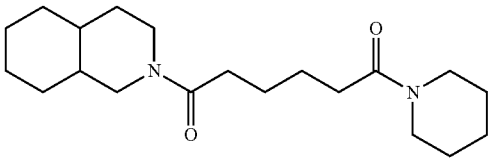
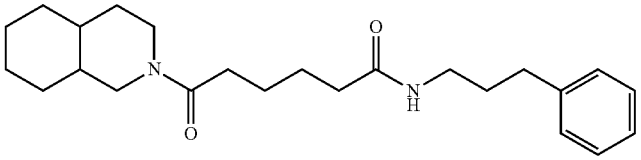
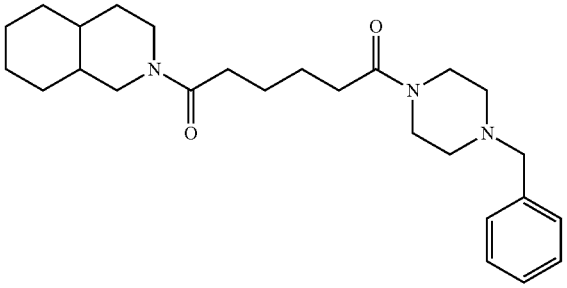
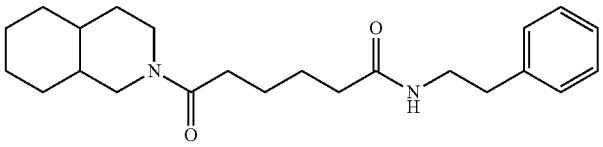
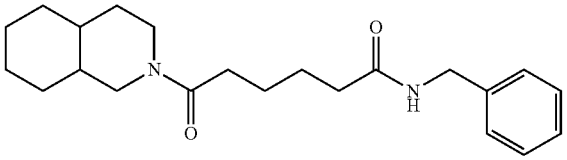
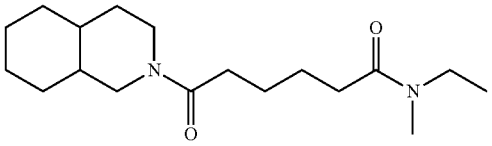
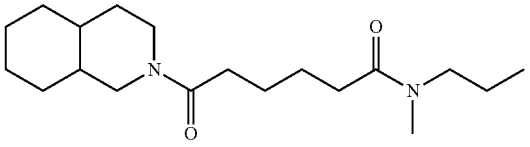
Ex.	structure	name	method	rt (min)	m/z
Ia.147		1-(octahydroisoquinolin-2-yl)-6-(4-oxopiperidin-1-yl)hexane-1,6-dione	B	3.04 3.14	349 349
Ia.148		1-(octahydroisoquinolin-2-yl)-6-piperidin-1-ylhexane-1,6-dione	B	3.64 3.75	335 335
Ia.149		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid (3-phenylpropyl)amide	B	3.93 4.02	385 385
Ia.150		1-(4-benzylpiperazin-1-yl)-6-(octahydroisoquinolin-2-yl)hexane-1,6-dione	B	2.77 2.83	426 426
Ia.151		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid phenethylamide	B	3.75 3.84	371 371
Ia.152		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid benzylamide	B	3.63 3.73	357 357
Ia.153		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid ethylmethylamide	B	3.36 3.46	309 309
Ia.154		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid methylpropylamide	B	3.61 3.72	323 323

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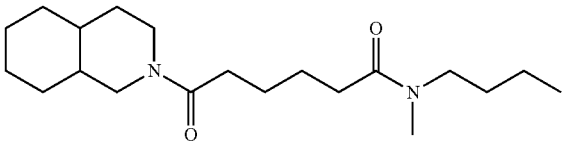
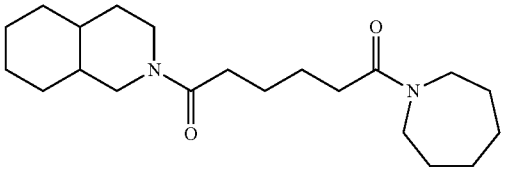
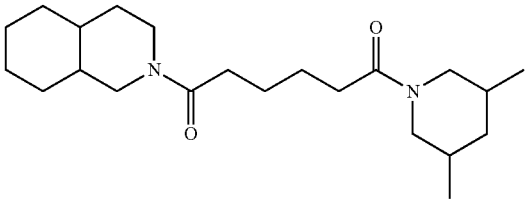
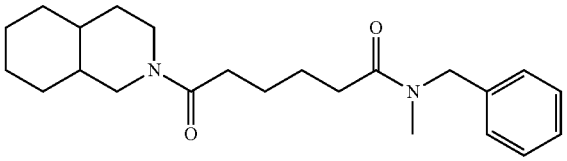
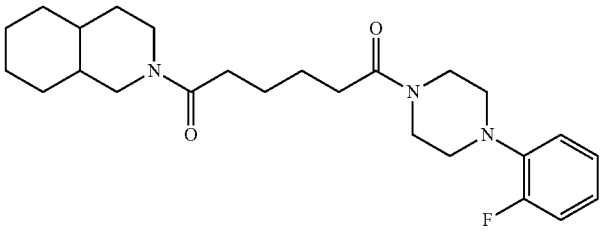
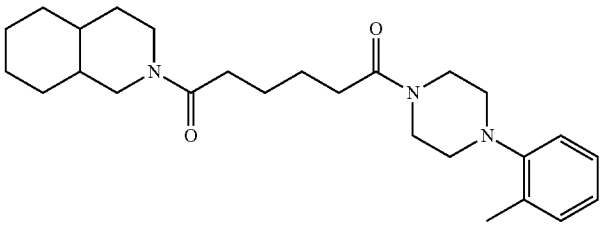
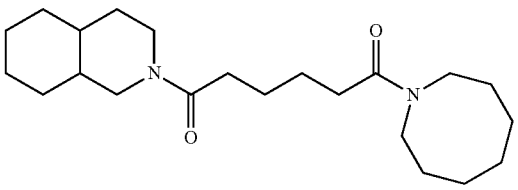
Ex.	structure	name	method	rt (min)	m/z
Ia.155		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid butylmethylamide	B	3.88 3.98	337 337
Ia.156		6-(octahydroisoquinolin-2-yl)-1-perhydroazepin-1-ylhexane-1,6-dione	B	3.82 3.92	349 349
Ia.157		1-(3,5-dimethylpiperidin-1-yl)-6-(octahydroisoquinolin-2-yl)hexane-1,6-dione	B	4.20 4.29	363 363
Ia.158		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid benzylmethylamide	B	3.92 4.02	371 371
Ia.159		1-[4-(2-fluorophenyl)piperazin-1-yl]-6-(octahydroisoquinolin-2-yl)hexane-1,6-dione	B	4.08 4.17	430 430
Ia.160		1-(octahydroisoquinolin-2-yl)-6-(4-o-tolylpiperazin-1-yl)hexane-1,6-dione	B	4.35 4.44	426 426
Ia.161		6-(octahydroisoquinolin-2-yl)-1-perhydroazocin-1-ylhexane-1,6-dione	B	4.03 4.13	363 363

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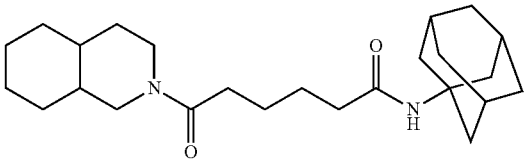
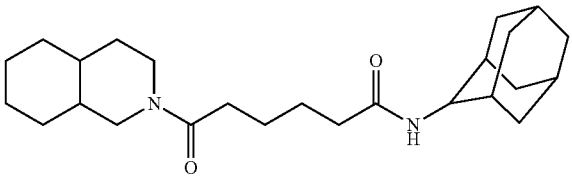
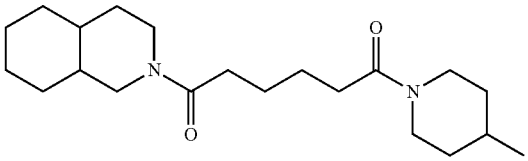
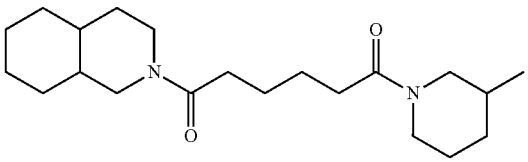
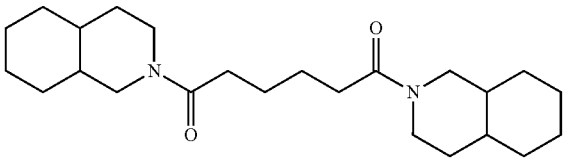
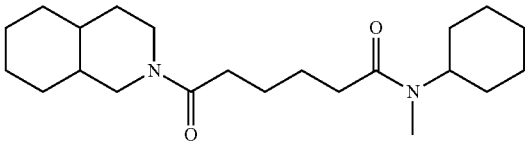
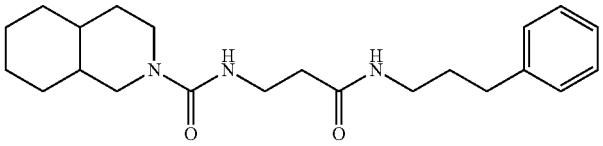
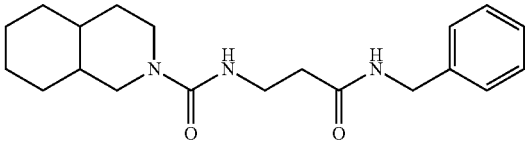
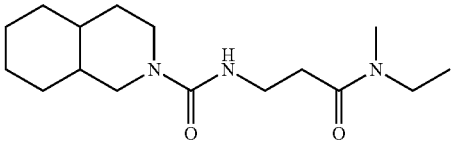
Ex.	structure	name	method	rt (min)	m/z
Ia.162		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid tricyclo [3.3.1.1^{3,7}]decan-1-ylamide	B	4.41 4.50	401 401
Ia.163		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid tricyclo[3.3.1.1^{3,7}]decan-2-ylamide	B	4.32 4.41	401 401
Ia.164		1-(4-methylpiperidin-1-yl)-6-(octahydroisoquinolin-2-yl)hexane-1,6-dione	B	3.92 4.02	349 349
Ia.165		1-(3-methylpiperidin-1-yl)-6-(octahydroisoquinolin-2-yl)hexane-1,6-dione	B	3.91 4.01	349 349
Ia.166		1,6-bis-(octahydroisoquinolin-2-yl)hexane-1,6-dione	B	4.47 4.56	389 389
Ia.167		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid cyclohexylmethylamide	B	4.12 4.21	363 363
Ib.63		Octahydroisoquinoline-2-carboxylic acid [2-(3-phenylpropylcarbamoyl)ethyl]amide	B	3.72 3.80	372 372
Ib.64		Octahydroisoquinoline-2-carboxylic acid (2-benzylcarbamoyl-ethyl)amide	B	3.42 3.50	344 344
Ib.65		Octahydroisoquinoline-2-carboxylic-acid [2-(ethylmethylcarbamoyl)ethyl]amide	B	3.06 3.17	296 296

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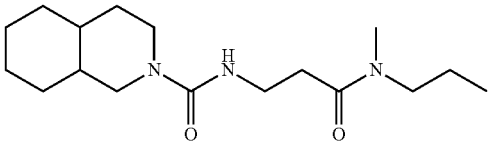
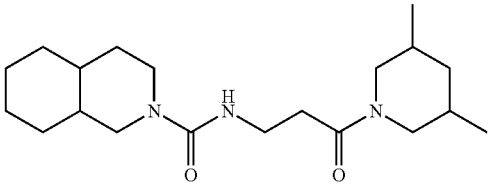
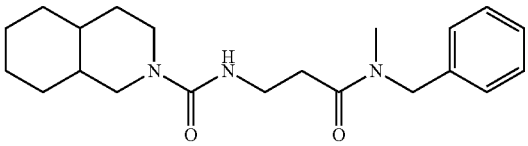
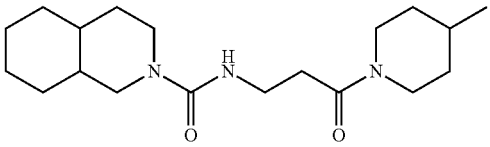
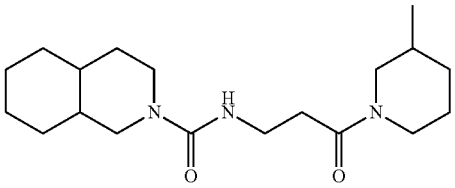
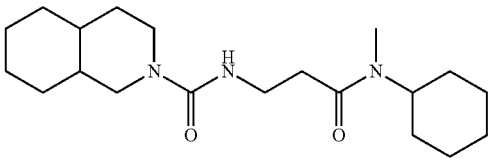
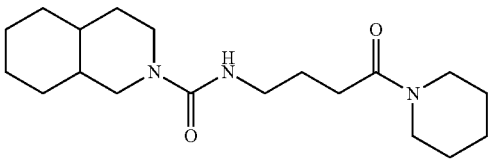
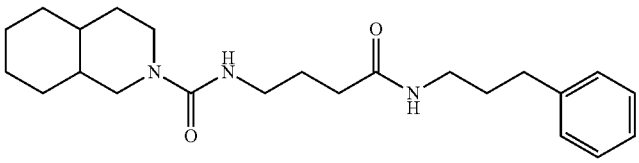
Ex.	structure	name	method	rt (min)	m/z
Ib.66		Octahydroisoquinoline-2-carboxylic acid [2-(methylpropylcarbamoyl)ethyl]amide	B	3.32 3.42	310 310
Ib.67		Octahydroisoquinoline-2-carboxylic acid [3-(3,5-dimethylpiperidin-1-yl)-3-oxopropyl]amide	B	3.91 3.98	350 350
Ib.68		Octahydroisoquinoline-2-carboxylic acid [2-(benzylmethylcarbamoyl)ethyl]amide	B	3.66 3.75	358 358
Ib.69		Octahydroisoquinoline-2-carboxylic acid [3-(4-methylpiperidin-1-yl)-3-oxopropyl]amide	B	3.62 3.71	336 336
Ib.70		Octahydroisoquinoline-2-carboxylic acid [3-(3-methylpiperidin-1-yl)-3-oxopropyl]amide	B	3.61 3.70	336 336
Ib.71		Octahydroisoquinoline-2-carboxylic acid [2-(cyclohexylmethylcarbamoyl)ethyl]amide	B	3.83 3.91	350 350
Ib.72		Octahydroisoquinoline-2-carboxylic acid (4-oxo-4-piperidin-1-ylbutyl)amide	B	3.41 3.51	336 336
Ib.73		Octahydroisoquinoline-2-carboxylic acid [3-(3-phenylpropylcarbamoyl)propyl]amide	B	3.88	386

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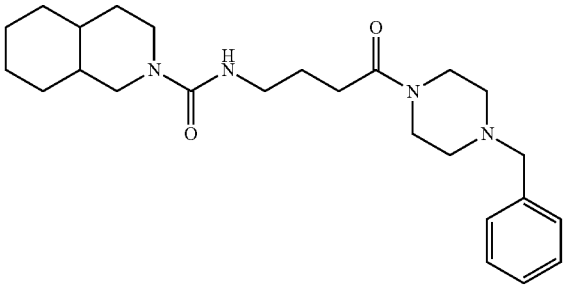
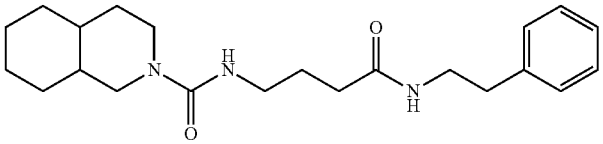
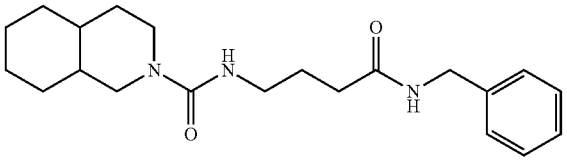
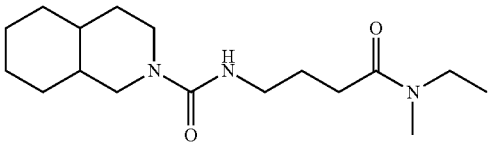
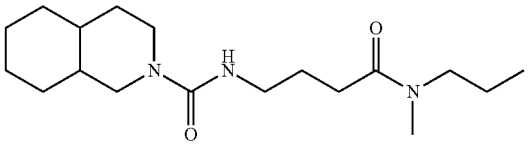
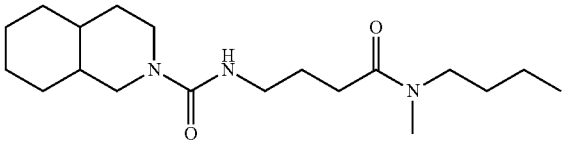
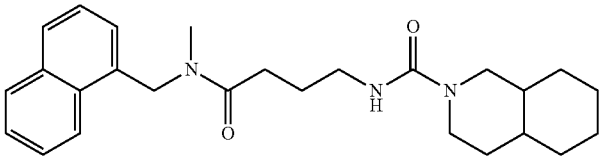
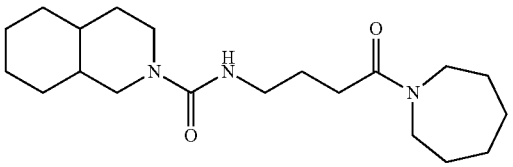
Ex.	structure	name	method	rt (min)	m/z
Ib.74		Octahydroisoquinoline-2-carboxylic acid [4-(4-benzylpiperazin-1-yl)-4-oxobutyl]amide	B	2.67	427
Ib.75		Octahydroisoquinoline-2-carboxylic acid (3-phenethylcarbamoylpropyl)amide	B	3.70	372
Ib.76		Octahydroisoquinoline-2-carboxylic acid (3-benzylcarbamoylpropyl)amide	B	3.58	358
Ib.77		Octahydroisoquinoline-2-carboxylic acid [3-(ethylmethylcarbamoyl)propyl]amide	B	3.24	310
Ib.78		Octahydroisoquinoline-2-carboxylic acid [3-(methylpropylcarbamoyl)propyl]amide	B	3.37 3.47	324 324
Ib.79		Octahydroisoquinoline-2-carboxylic acid [3-(butylmethylcarbamoyl)propyl]amide	B	3.74	338
Ib.80		Octahydroisoquinoline-2-carboxylic acid [3-(methylnaphthalen-1-ylmethylcarbamoyl)propyl]amide	B	4.15	422
Ib.81		Octahydroisoquinoline-2-carboxylic acid (4-oxo-4-perhydroazepin-1-ylbutyl)amide	B	3.57 3.67	350 350

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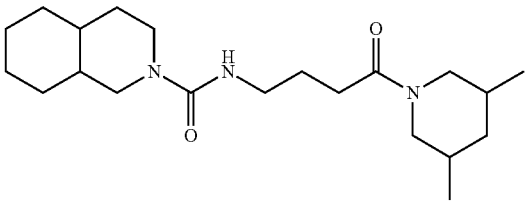
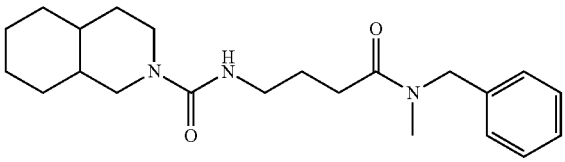
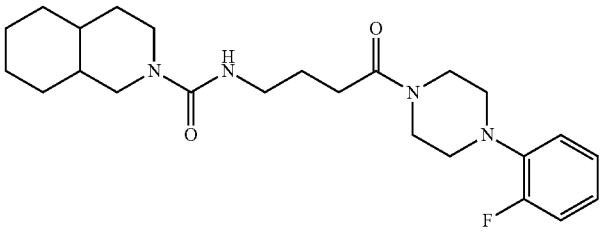
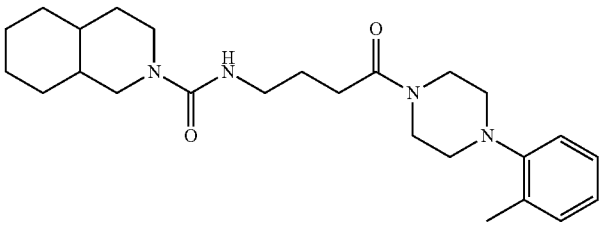
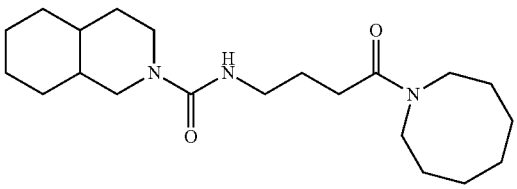
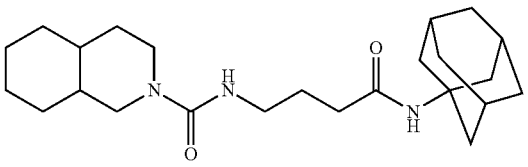
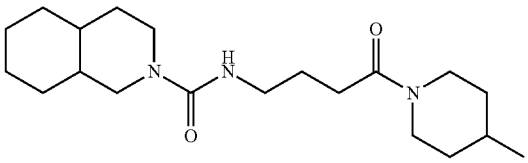
Ex.	structure	name	method	rt (min)	m/z
Ib.82		Octahydroisoquinoline-2-carboxylic acid [4-(3,5-dimethylpiperidin-1-yl)-4-oxobutyl]amide	B	3.95 4.04	364 364
Ib.83		Octahydroisoquinoline-2-carboxylic acid [3-(benzylmethylcarbamoyl)propyl]amide	B	3.69 3.78	372 372
Ib.84		Octahydroisoquinoline-2-carboxylic acid {4-[4-(2-fluorophenyl)piperazin-1-yl]-4-oxobutyl}amide	B	3.85 3.94	431 431
Ib.85		Octahydroisoquinoline-2-carboxylic acid [4-oxo-4-(4-o-tolylpiperazin-1-yl)butyl]amide	B	4.12 4.20	427 427
Ib.86		Octahydroisoquinoline-2-carboxylic acid (4-oxo-4-perhydroazocin-1-ylbutyl)amide	B	3.78 3.87	364 364
Ib.87		Octahydroisoquinoline-2-carboxylic acid [3-(tricyclo(3.3.1.1^{3,7}]decan-1-ylcarbamoyl)propyl]amide	B	4.38	402
Ib.88		Octahydroisoquinoline-2-carboxylic acid [4-(4-methylpiperidin-1-yl)-4-oxobutyl]amide	B	3.67 3.77	350 350

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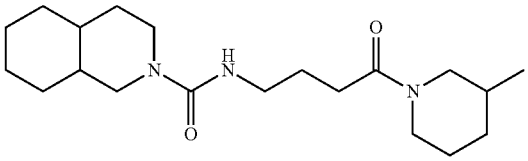
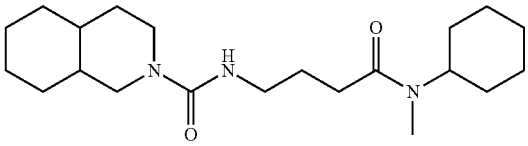
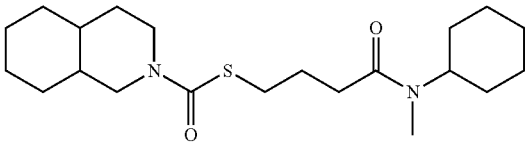
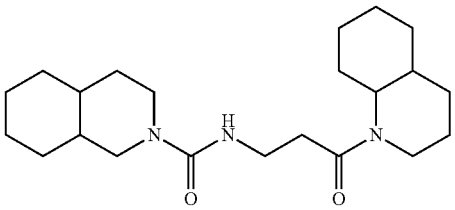
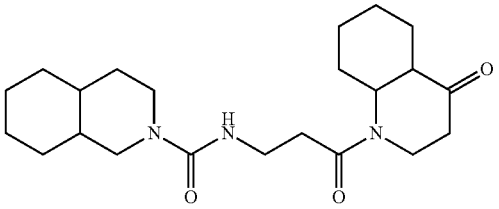
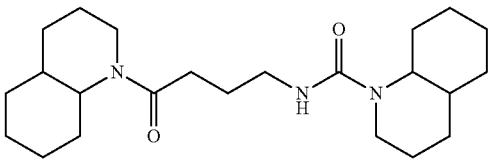
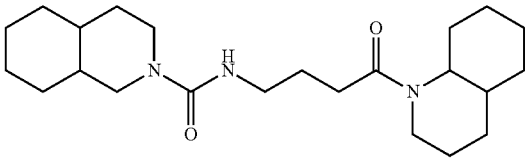
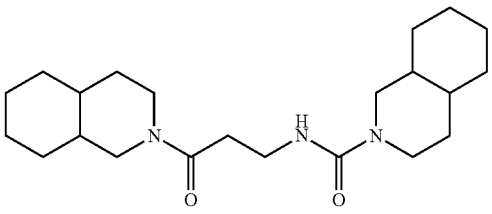
Ex.	structure	name	method	rt (min)	m/z
Ib.89		Octahydroisoquinoline-2-carboxylic acid [4-(3-methylpiperidin-1-yl)-4-oxobutyl]amide	B	3.66 3.76	350 350
Ib.90		Octahydroisoquinoline-2-carboxylic acid [3-(cyc1ohexylmethylcarbamoyl)propyl]amide	B	3.87 3.96	364 364
Ib.91		Octahydroisoquinoline-2-carbothioic acid S-[3-(cyclohexylmethylcarbamoyl)propyl] ester	B	4.49	381
Ib.92		Octahydroisoquinoline-2-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	B	4.10 4.17	376 376
Ib.93		Octahydroisoquinoline-2-carboxylic acid [3-oxo-3-(4-oxooctahydroquinolin-1-yl)propyl]amide	B	3.46 3.53	390 390
Ib.94		Octahydroquinoline-1-carboxylic acid (octahydroquinolin-1-yl)-4-oxobutyl]amide	B	4.15	390
Ib.95		Octahydroisoquinoline-2-carboxylic acid [4-(octahydroquinolin-1-yl)-4-oxobutyl]amide	B	4.22	390
Ib.96		Octahydroisoquinoline-2-carboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	B	4.17 4.25	376 376

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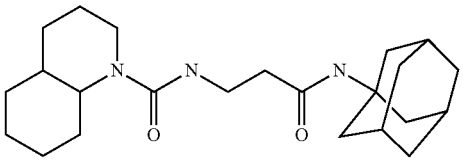
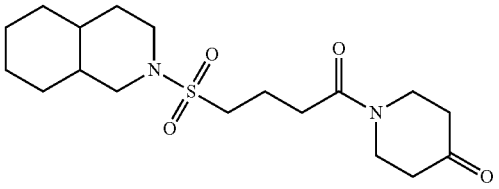
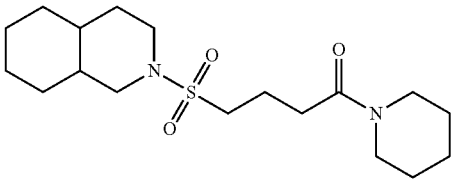
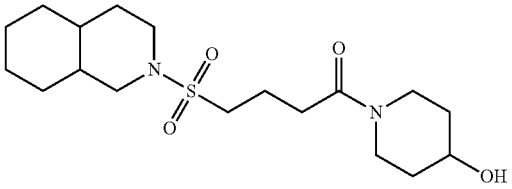
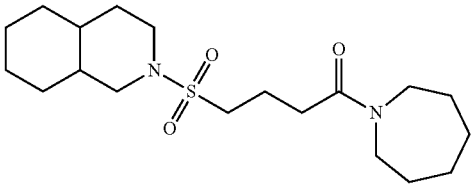
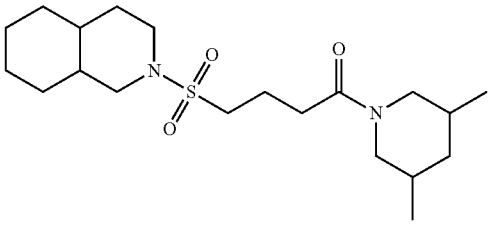
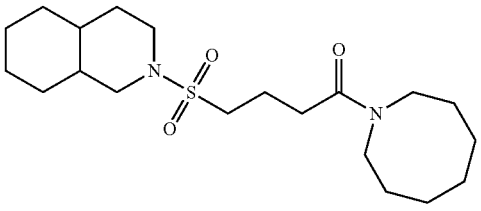
Ex.	structure	name	method	rt (min)	m/z
Ib.97		Octahydroquinoline-1-carboxylic acid N-[3-[(3s,5s,7s)-adamantan-1-ylamino-3-oxopropyl] amide	B	4.18	388
Im.17		1-[4-(octahydroisoquinoline-2-sulfonyl)butyryl]piperidin-4-one	B	3.25 3.31	371 371
Im.18		4-(octahydroisoquinoline-2-sulfonyl)-1-piperidin-1-ylbutan-1-one	B	3.93	357
Im.19		1-(4-hydroxypiperidin-1-yl)-4-(octahydroisoquinoline-2-sulfonyl)butan-1-one	B	3.04 3.11	373 373
Im.20		4-(octahydroisoquinoline-2-sulfonyl)-1-perhydroazepin-1-ylbutan-1-one	B	4.02 4.09	371 371
Im.21		1-(3,5-dimethylpiperidin-1-yl)-4-(octahydroisoquinoline-2-sulfonyl)butan-1-one	B	4.37 4.45	385 385
Im.22		4-(octahydroisoquinoline-2-sulfonyl)-1-perhydroazocin-1-ylbutan-1-one	B	4.23 4.30	385 385

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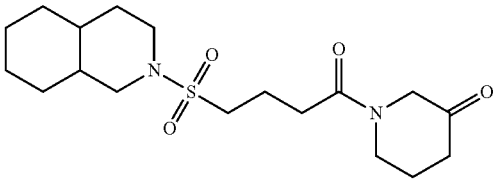
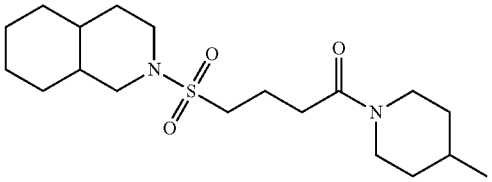
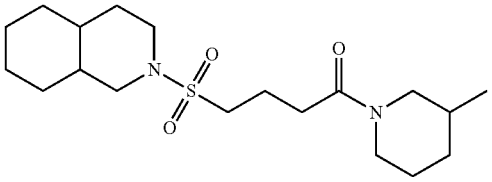
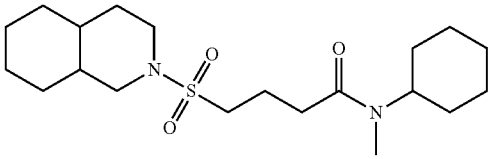
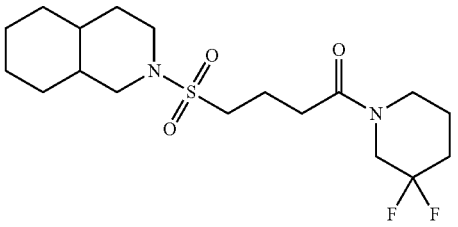
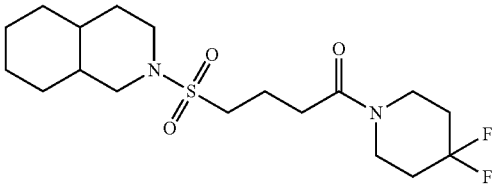
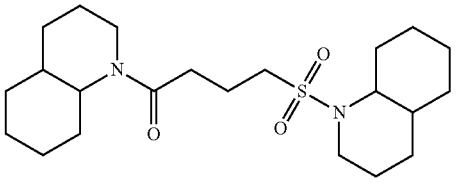
Ex.	structure	name	method	rt (min)	m/z
Im.23		1-((R-3-hydroxypiperidin-1-yl)-4-(octahydroisoquinoline-2-sulfonyl)butan-1-one	B	3.15 3.22	373 373
Im.24		1-(4-methylpiperidin-1-yl)-4-(octahydroisoquinoline-2-sulfonyl)butan-1-one	B	4.12 4.19	371 371
Im.25		1-(3-methylpiperidin-1-yl)-4-(octahydroisoquinoline-2-sulfonyl)butan-1-one	B	4.11 4.18	371 371
Im.26		N-cyclohexyl-N-methyl-4-(octahydroisoquinoline-2-sulfonyl)butyramide	B	4.31 4.37	385 385
Im.27		1-(3,3-difluoropiperidin-1-yl)-4-(octahydroisoquinoline-2-sulfonyl)butan-1-one	B	3.87 3.93	393 393
Im.28		1-(4,4-difluoropiperidin-1-yl)-4-(octahydroisoquinoline-2-sulfonyl)butan-1-one	B	3.87 3.94	393 393
Im.29		4-(octahydroquinoline-1-sulfonyl)-1-(octahydroquinolin-1-yl)butan-1-one	B	4.51	411

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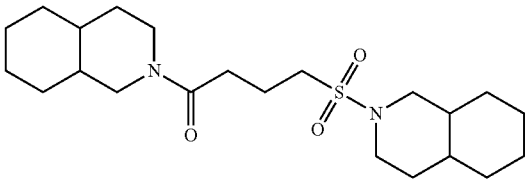
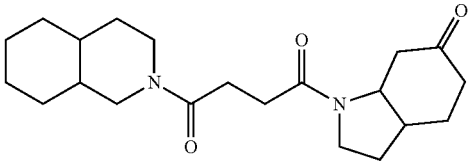
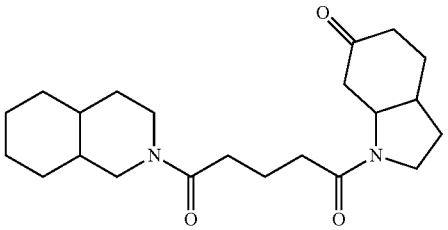
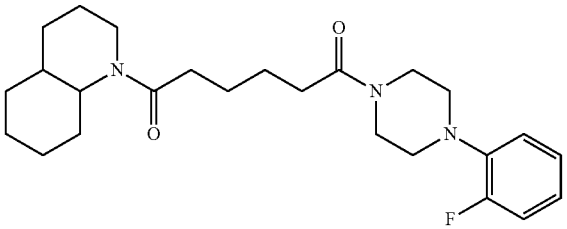
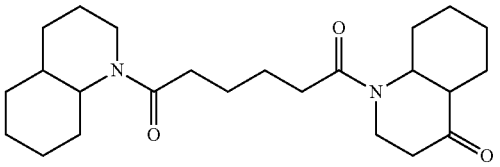
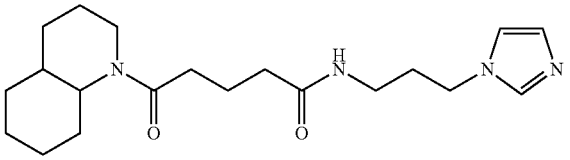
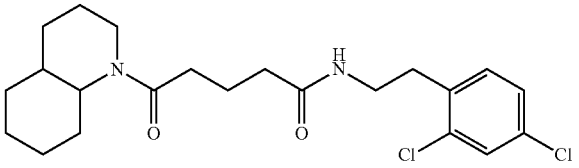
Ex.	structure	name	method	rt (min)	m/z
Im.30		4-(octahydroisoquinoline-2-sulfonyl)-1-(octahydroisoquinolin-2-yl)butan-1-one	B	4.63 4.70	411 411
Ia.168		1-(octahydroisoquinolin-2-yl)-4-(6-oxooctahydroindol-1-yl)butane-1,4-dione	B	4.33 4.41	361 361
Ia.169		1-(octahydroisoquinolin-2-yl)-5-(6-oxooctahydroindol-1-yl)pentane-1,5-dione	B	4.38 4.48	375 375
Ia.170		1-[4-(2-fluorophenyl)piperazin-1-yl]-6-(octahydroquinolin-1-yl)hexane-1,6-dione	B	4.09	430
Ia.171		1-(octahydroquinolin-1-yl)-6-(4-oxooctahydroquinolin-1-yl)hexane-1,6-dione	B	3.72	403
Ia.172		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[3-(1H-imidazol-1-yl)propyl]amide	B	2.29	361
Ia.173		(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2,4-dichlorophenethyl) amide	B	4.20	425, 427

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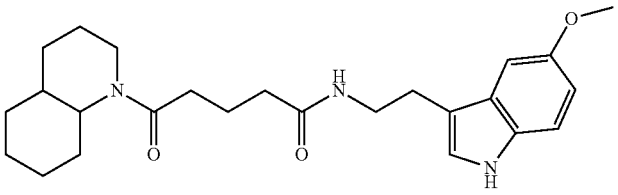
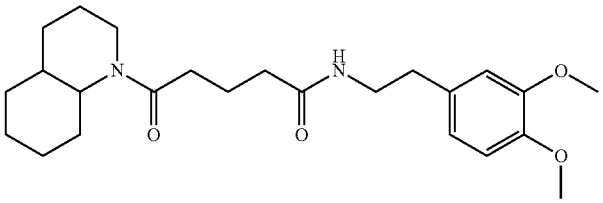
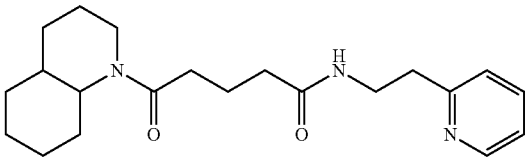
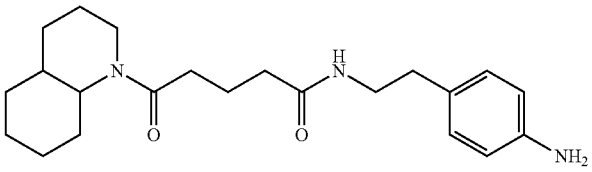
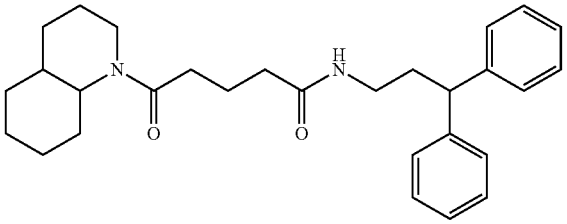
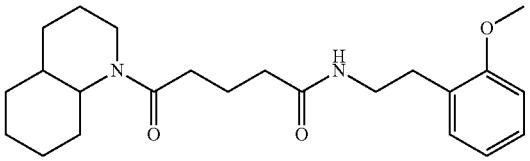
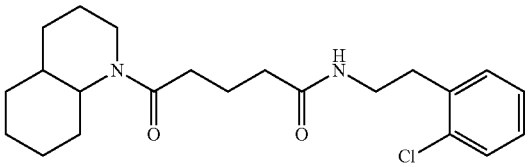
Ex.	structure	name	method	rt (min)	m/z
Ia.174		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[2-(5-methoxy-1H-indol-3-yl)ethyl]amide	B	3.47	426
Ia.175		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(3,4-dimethoxyphen-ethyl)amide	B	3.41	417
Ia.176		(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[2-(pyridin-2-yl)ethyl]amide	B	2.34	358
Ia.177		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(4-aminophenethyl)amide	B	2.44	372
Ia.178		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(3,3-diphenylpropyl)amide	B	4.28	447
Ia.179		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2-methoxyphenethyl)amide	B	3.74	387
Ia.180		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2-chlorophenethyl)amide	B	3.88	391

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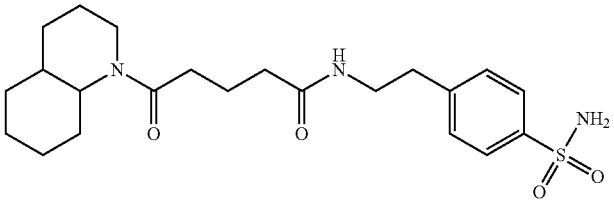
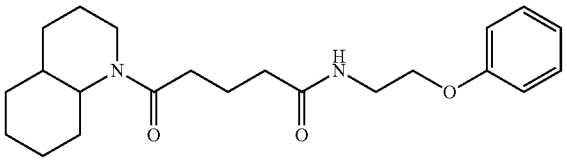
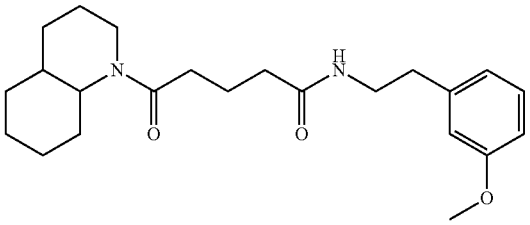
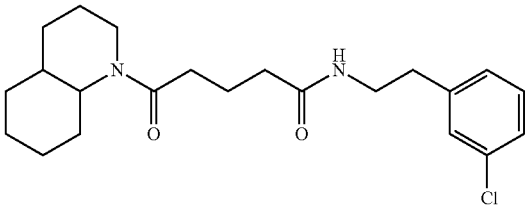
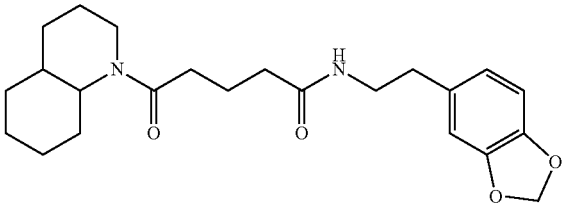
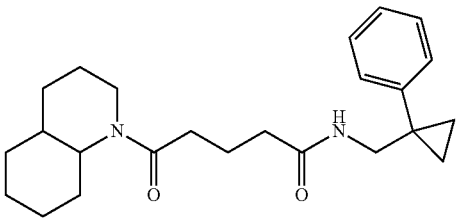
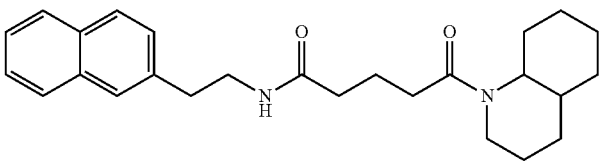
Ex.	structure	name	method	rt (min)	m/z
Ia.181		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(4-sulfamoylphen-ethyl)amide	B	2.97	436
Ia.182		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2-phenoxyethyl)amide	B	3.67	373
Ia.183		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(3-methoxyphenethyl)amide	B	3.65	387
Ia.184		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(3-chlorophenethyl)amide	B	3.92	391
Ia.185		5-(octahydroquinolin-1)-yl)-5-oxopentanoic acid N-[2-(benzo[d][1,3]dioxol-5-yl)ethyl]amide	B	3.56	401
Ia.186		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(1-phenylcyclopro-pyl)methyl]amide	B	3.92	383
Ia.187		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[2-(naphthalen-2-yl)ethyl]amide	B	4.05	407

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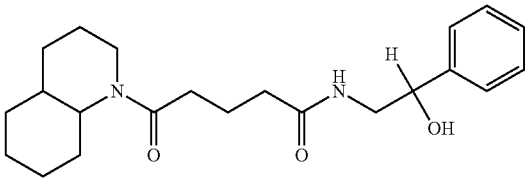
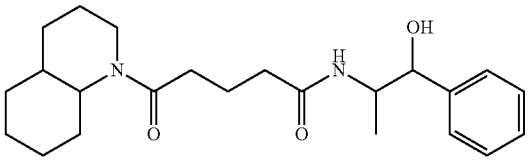
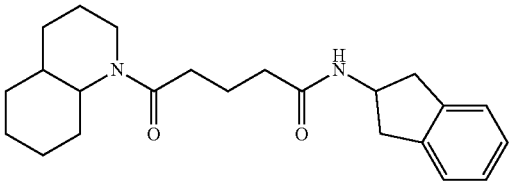
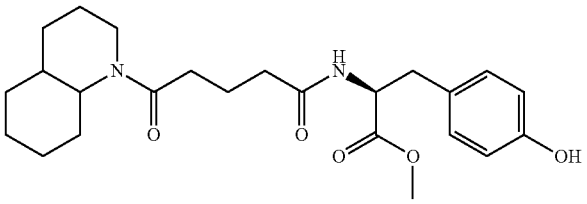
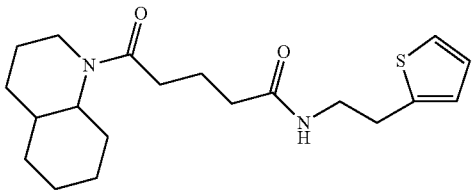
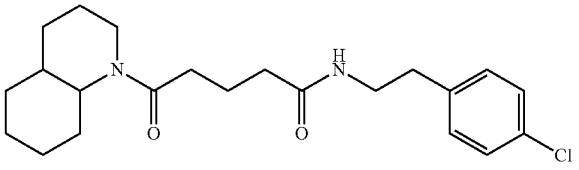
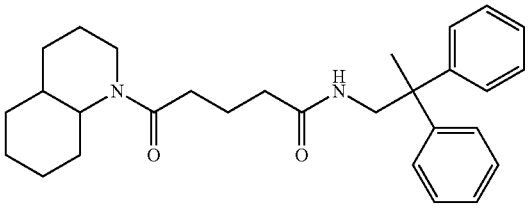
Ex.	structure	name	method	rt (min)	m/z
Ia.188		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(R)-2-phenyl-2-hydroxyethyl]amide	B	3.19	373
Ia.189		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(1R,2S)-1-phenyl-1-hydroxypropan-2-yl]amide	B	3.37	387
Ia.190		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2,3-dihydro-1H-inden-2-yl)amide	B	3.77	369
Ia.191		(2S)-3-(4-hydroxyphenyl)-2-[5-(octahydroquinolin-1-yl)-5-oxopentanamide]propanoic acid methyl ester	B	3.21	431
Ia.192		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[2-(thiophen-2-yl)ethyl]amide	B	3.58	363
Ia.193		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(4-chlorophenethyl)amide	B	3.93	391
Ia.194		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2,2-diphenylpropyl)amide	B	4.39	447

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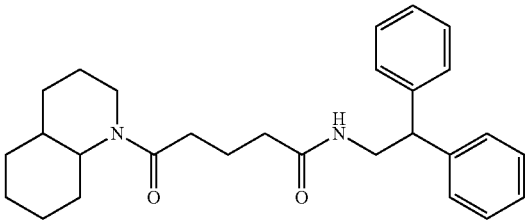
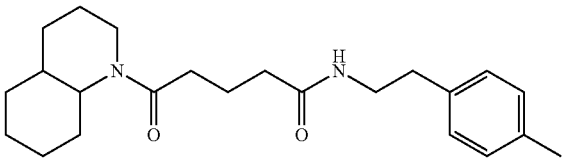
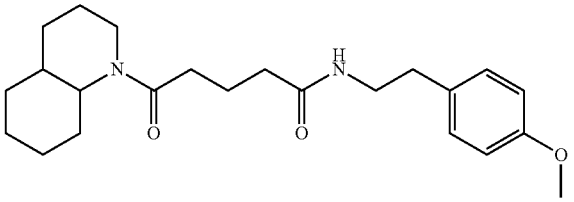
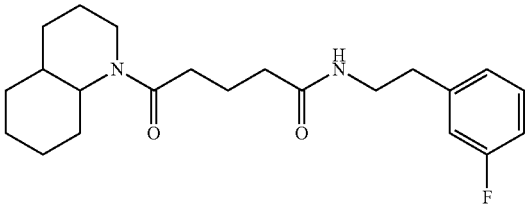
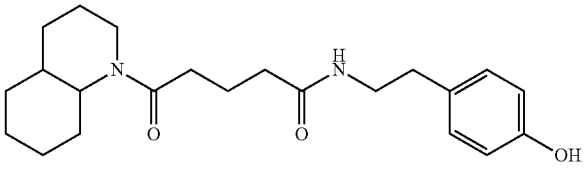
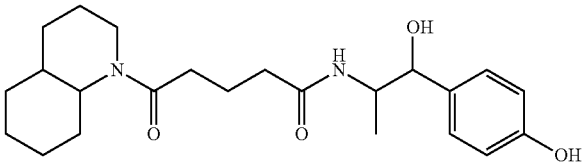
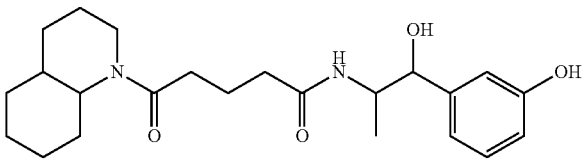
Ex.	structure	name	method	rt (min)	m/z
Ia.195		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2,2-diphenylethyl)amide	B	4.16	433
Ia.196		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(4-methylphenethyl)amide	B	3.88	371
Ia.197		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(4-methoxyphenethyl)amide	B	3.61	387
Ia.198		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(3-fluorophenethyl)amide	B	3.72	375
Ia.199		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(4-hydroxyphenethyl)amide	B	3.10	373
Ia.200		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(1-hydroxyl-1-(4-hydroxyphenyl)propan-2-yl)amide	B	2.88	403
Ia.201		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(1-hydroxyl-1-(3-hydroxyphenyl)propan-2-yl)amide	B	3.00	403

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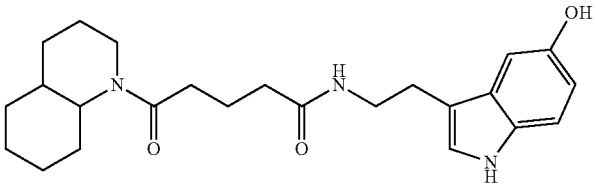
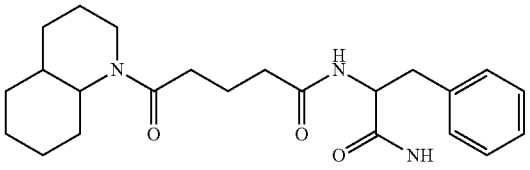
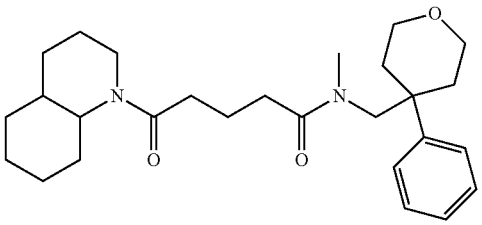
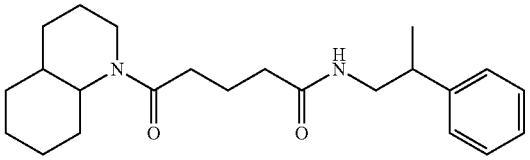
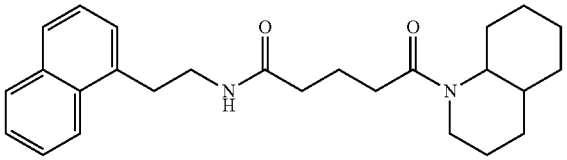
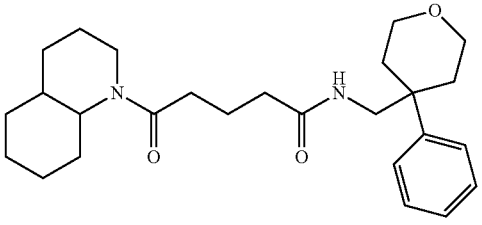
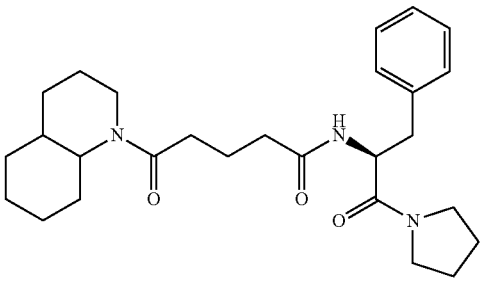
Ex.	structure	name	method	rt (min)	m/z
Ia.202		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[2-(5-hydroxyl-1H-indol-3-yl)ethyl]amide	B	3.04	412
Ia.203		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(R)-1-amino-3-phenyl-1-oxopropan-2-yl]amide	B	3.19	400
Ia.204		N-methyl-5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-((4-phenyltetrahydro-2H-pyran-4-yl)methyl)amide	B	3.89	441
Ia.205		5-octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2-phenylpropyl)amide	B	4.02	385
Ia.206		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[2-(naphthalen-1-yl)ethyl]amide	B	4.06	407
Ia.207		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(4-phenyltetrahydro-2H-pyran-4-yl)methyl]amide	B	3.06	427
Ia.208		5-(octahydroquinolin-1-yl)-5-oxopropanoic acid N-([S)-3-phenyl-1-oxo-1-(pyrrolidin-1-yl)propan-2-yl]amide	B	3.65	454

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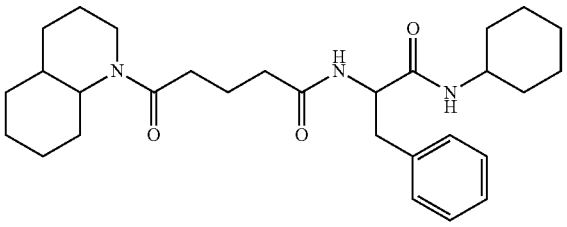
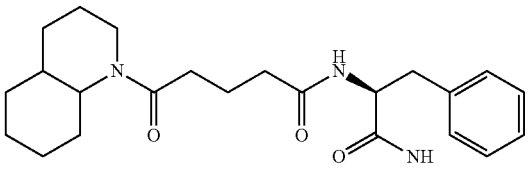
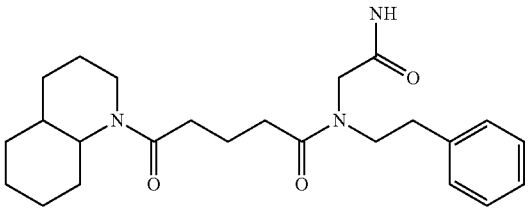
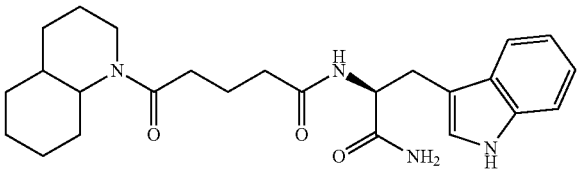
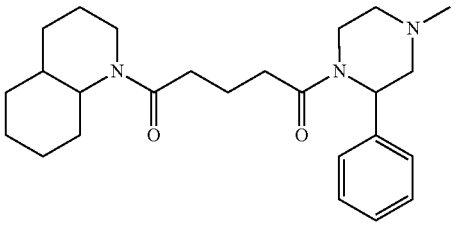
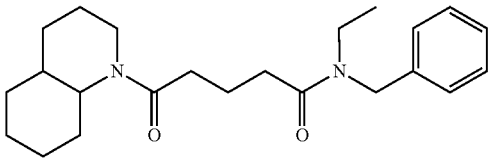
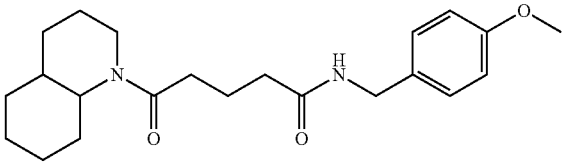
Ex.	structure	name	method	rt (min)	m/z
Ia.209		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(S)-1-(cyclohexylamino)-3-phenylpropane-1-oxo-2-yl]amide	B	4.06	482
Ia.210		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(S)-1-amino-3-phenyl-1-oxopropan-2-yl]amide	B	3.19	400
Ia.211		N-(2-amino-2-oxoethyl)-5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-phenethylamide	B	3.41	414
Ia.212		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(S)-1-amino-3-(1H-indol-3-yl)-1-oxopropan-2-yl]amide	B	3.15	439
Ia.213		1-(2-phenyl-4-methylpiperazin-1-yl)-5-(octahydroquinolin-1-yl)pentane-1,5-dione	B	2.58	412
Ia.214		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-benzyl-N-ethylamide	B	4.06	371
Ia.215		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(4-methoxybenzyl)amide	B	3.50	373

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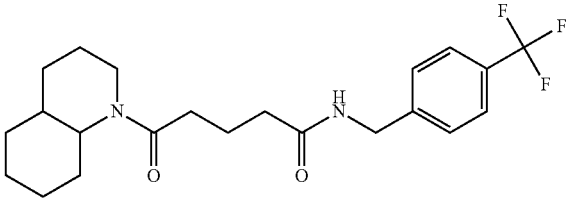
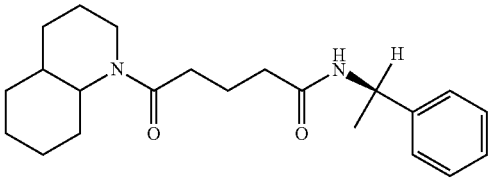
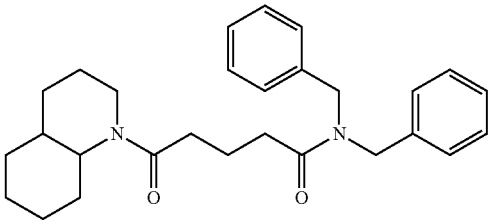
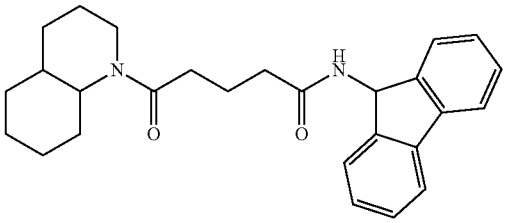
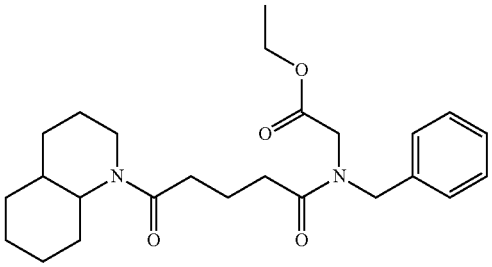
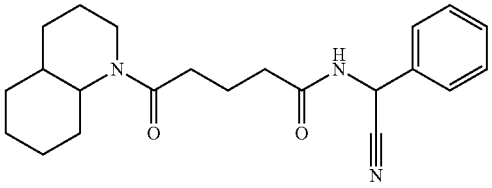
Ex.	structure	name	method	rt (min)	m/z
Ia.216		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(4-(trifluoromethyl)benzyl)amide	B	3.97	411
Ia.217		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(R)-1-phenylethyl]amide	B	3.70	357
Ia.218		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N,N-dibenzylamide	B	4.53	433
Ia.219		5-octahydroquinolin-1-yl)-5-oxopentanoic acid N-(9H-fluoren-9-yl)amide	B	4.18	417
Ia.220		2-(N-benzyl-5-(octahydroquinolin-1-yl)-5-oxopentanamide)acetic acid ethyl ester	B	4.12	429
Ia.221		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[cyano(phenyl)methyl)amide	B	3.68	368

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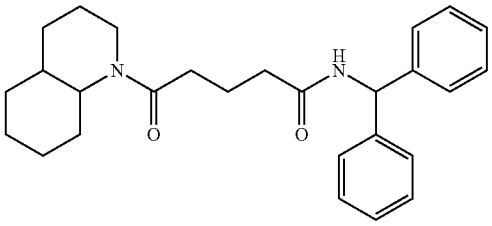
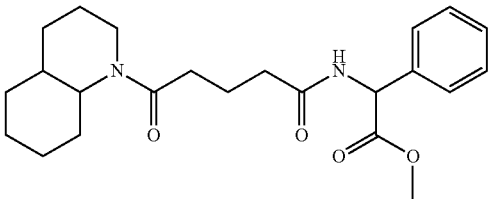
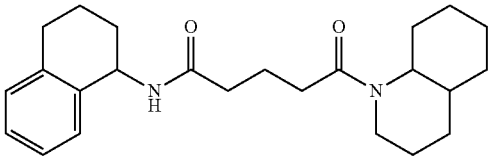
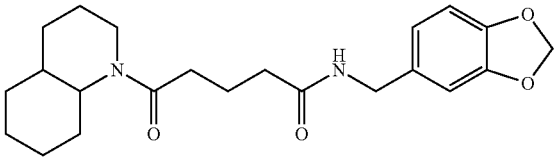
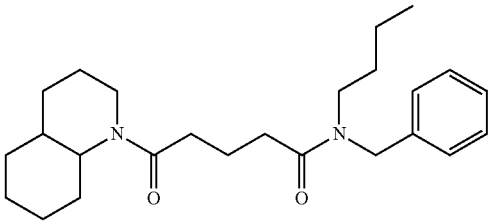
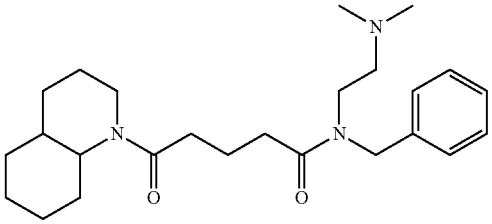
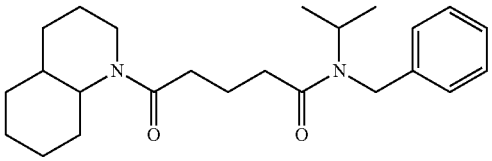
Ex.	structure	name	method	rt (min)	m/z
Ia.222		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-benzhydrylamide	B	4.15	419
Ia.223		(2R)-2-(5-(octahydroquinolin-1-yl)-5-oxopentanamido)-2-phenylacetic acid methyl ester	B	3.67	401
Ia.224		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(1,2,3,4-tetrahydronaphthalen-1-yl)amide	B	3.96	383
Ia.225		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(benzo[d][1,3]dioxol-5-yl methyl)amide	B	3.46	387
Ia.226		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-benzyl-N-butylamide	B	4.51	399
Ia.227		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-benzyl-N-[2-(dimethylamino)ethyl]amide	B	2.83	414
Ia.228		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-benzyl-N-isopropylamide	B	4.22	385

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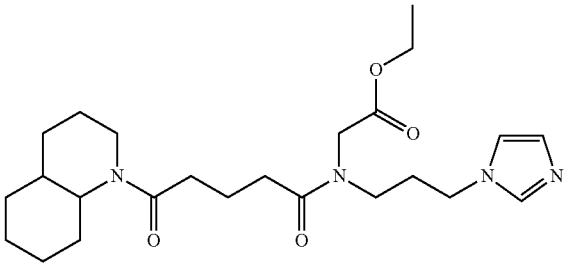
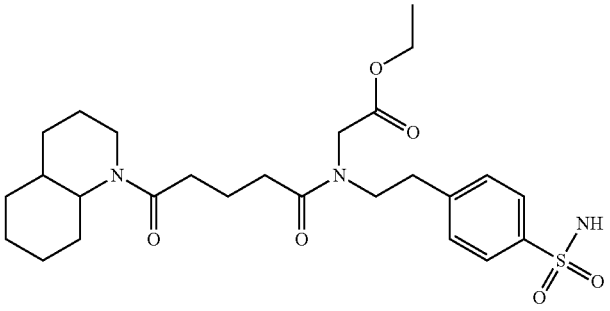
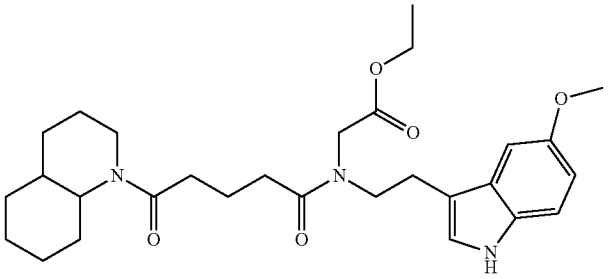
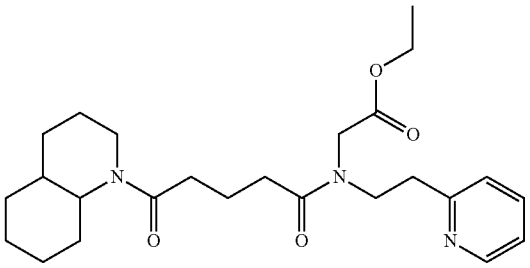
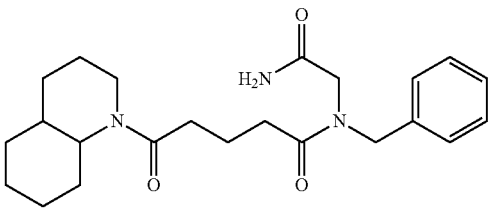
Ex.	structure	name	method	rt (min)	m/z
Ia.229		2-[N-[3-(1H-imidazol-1-yl)propyl)-5-(octahydroquinolin-1-yl)-5-oxopentanamido]acetic acid ethyl ester	B	2.67	447
Ia.230		2-[5-(octahydroquinolin-1-yl)-5-oxo-N-(4-sulfamoylphen-ethyl)pentanamido]acetic acid ethyl ester	B	3.43	522
Ia.231		2-[N-[2-(5-methoxy-1H-indol-3-yl)ethyl]-5-(octahydroquinolin-1-yl)-5-oxopentanamido]acetic acid ethyl ester	B	3.92	512
Ia.232		2-[5-(octahydroquinolin-1-yl)-5-oxo-N-[2-(pyridin-2-yl)ethyl]pentan-amido]acetic acid ethyl ester	B	2.79	444
Ia.233		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2-amino-2-oxoethyl)-N-benzylamide	B	3.28	400

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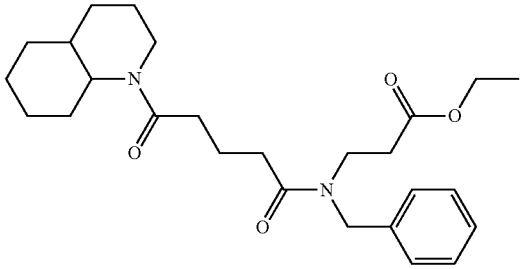
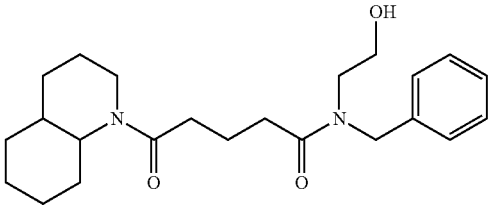
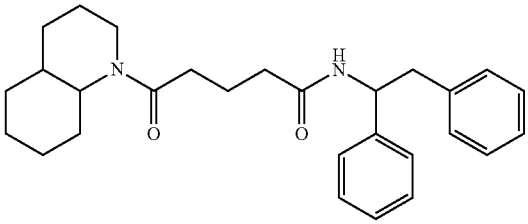
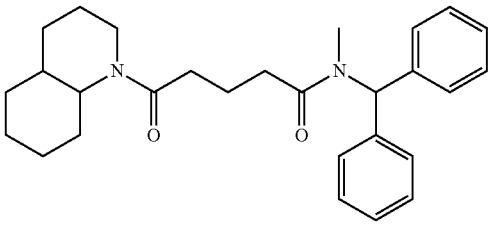
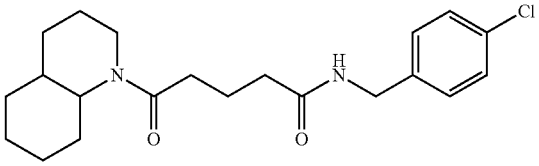
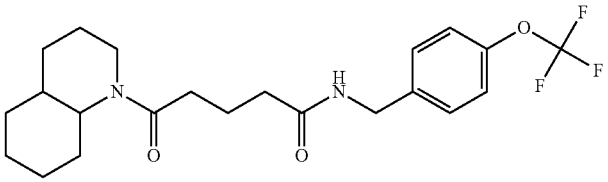
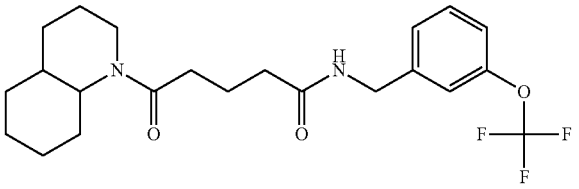
Ex.	structure	name	method	rt (min)	m/z
Ia.234		3-[N-benzyl-5-(octahydroquinolin-1-yl)-5-oxopentanamido]propanoic acid ethyl ester	B	4.15	443
Ia.235		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-benzyl-N-(2-hydroxyethyl)amide	B	3.47	387
Ia.236		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(1,2-diphenylethyl)amide	B	4.21	433
Ia.237		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-benzhydryl-N-methylamide	B	4.53	433
Ia.238		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(4-chlorobenzyl)amide	B	3.82	377
Ia.239		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[4-(trifluoromethoxy)benzyl]amide	B	4.06	427
Ia.240		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[3-(trifluoromethoxy)benzyl]amide	B	4.06	427

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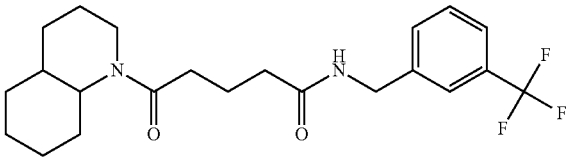
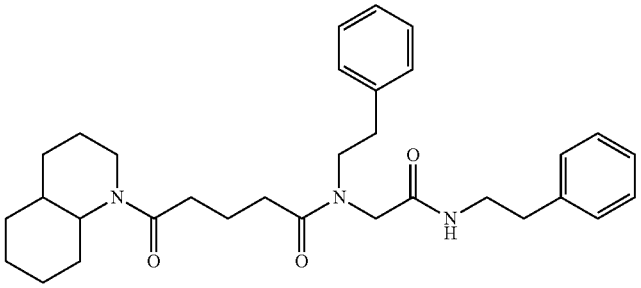
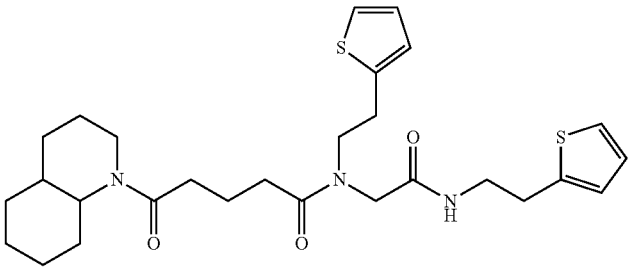
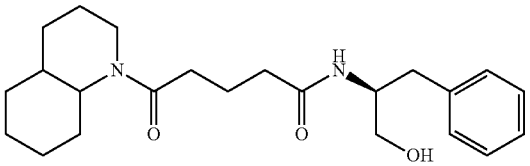
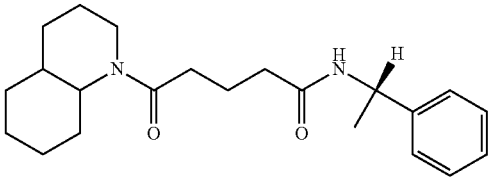
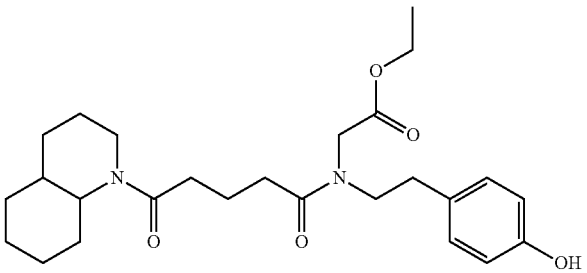
Ex.	structure	name	method	rt (min)	m/z
Ia.241		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[3-(trifluoromethyl)benzyl]amide	B	3.97	411
Ia.242		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-phenethyl-N-[2-(phenethylamino)-2-oxoethyl]amide	B	4.25	518
Ia.243		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[2-oxo-2-{[2-(thiophen-2-yl)ethyl]amino}ethyl]-N-[2-(thiophen-2-yl)ethyl]amide	B	4.11	530
Ia.244		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(S)-3-phenyl-1-hydroxypropan-2-yl]amide	B	3.31	387
Ia.245		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-[(S)-1-phenylethyl]amide	B	3.70	357
Ia.246		2-[N-(4-hydroxyphenethyl)-5-(octahydroquinolin-1-yl)-5-oxopentanamido]acetic acid ethyl ester	B	3.59	459

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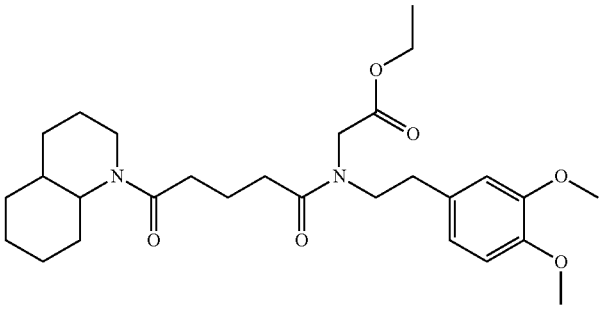
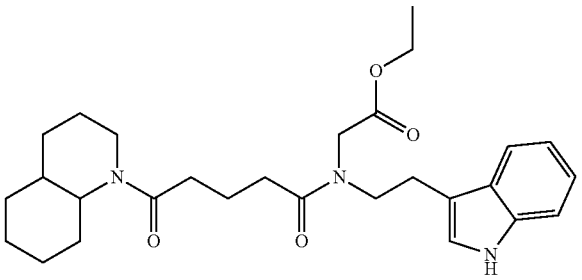
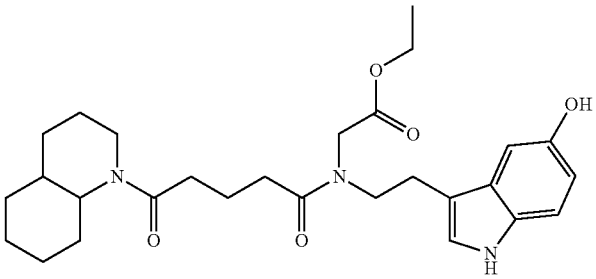
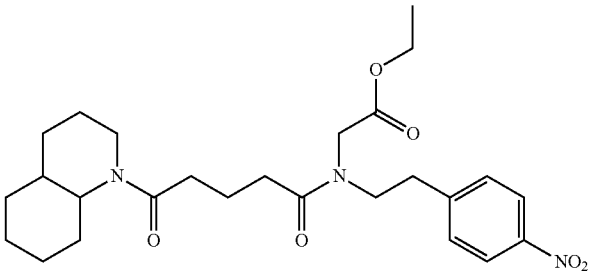
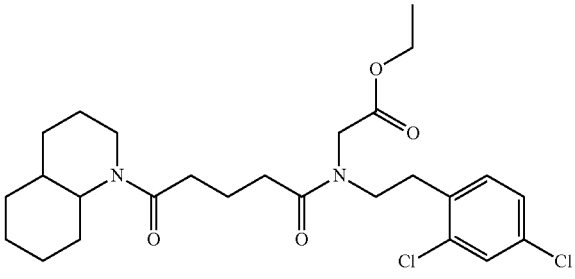
Ex.	structure	name	method	rt (min)	m/z
Ia.247		2-(N-(3,4-dimethoxyphenethyl)-5-(octahydroquinolin-1-yl)-5-oxopentanamido]acetic acid ethyl ester	B	3.91	503
Ia.248		2-[N-[2-(1H-indol-3-yl)ethyl]-5-(octahydroquinolin-1-yl)-5-oxopentanamido)acetic acid ethyl ester	B	4.04	482
Ia.249		2-[N-[2-(5-hydroxy-1H-indol-3-yl)ethyl]-5-(octahydroquinolin-1-yl)-5-oxopentanamido]acetic acid ethyl ester	B	3.47	498
Ia.250		2-[N-(4-nitrophenethyl)-5-(octahydroquinolin-1-yl)-5-oxopentanamido]acetic acid ethyl ester	B	4.10	488
Ia.251		2-(N-(2,4-dichlorophenethyl)-5-(octahydroquinolin-1-yl)-5-oxopentanamido]acetic acid ethyl ester	B	4.73	511

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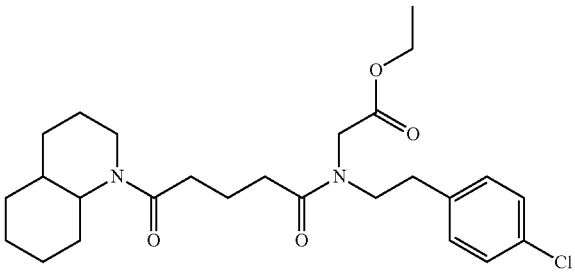
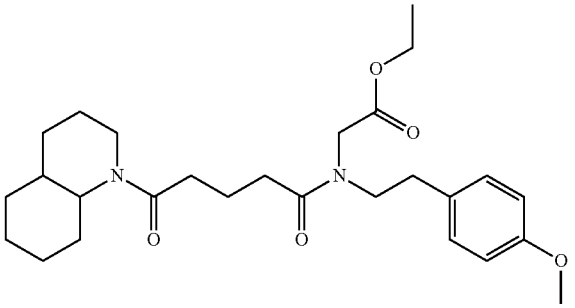
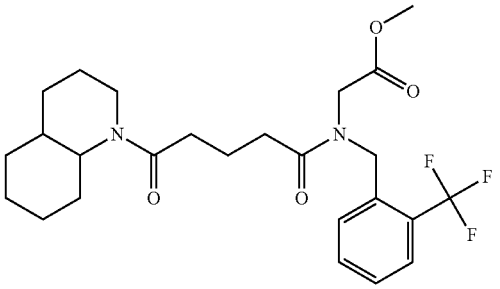
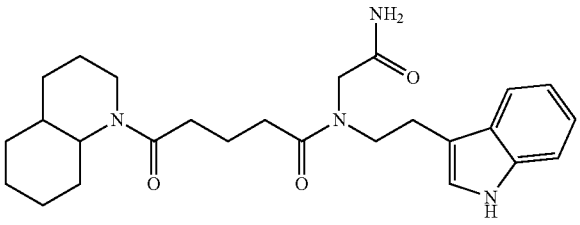
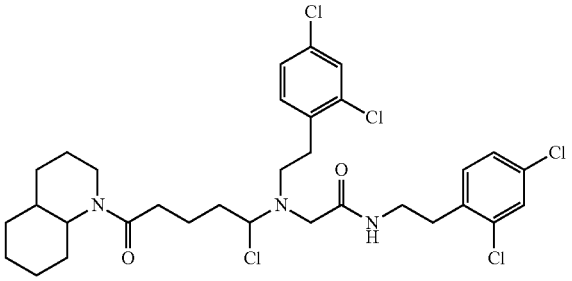
Ex.	structure	name	method	rt (min)	m/z
Ia.252		2-[N-(4-chlorophenethyl)-5-(octahydroquinolin-1-yl)-5-oxopentanamido]acetic acid ethyl ester	B	4.45	477
Ia.253		2-[N-(4-methoxyphenethyl)-5-(octahydroquinolin-1-yl)-5-oxopentanamido]acetic acid ethyl ester	B	4.14	473
Ia.254		2-[5-(octahydroquinolin-1-yl)-5-oxo-N-[2-(trifluoromethyl)benzyl]pentanamido]acetic acid methyl ester	B	4.27	483
Ia.255		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2-amino-2-oxoethyl)-N-[2-(1H-indol-3-yl)ethyl]amide	B	3.34	453
Ia.256		5-(octahydroquinolin-1-yl)-5-oxopentanoic acid N-(2,4-dichlorophenethyl)-N-[2-[(2,4-dichlorophenethyl)amino]-2-oxoethyl]amide	B	5.13	654, 656

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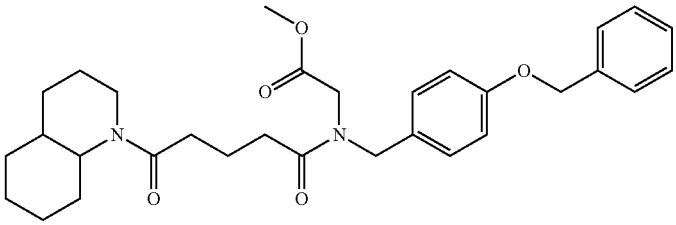
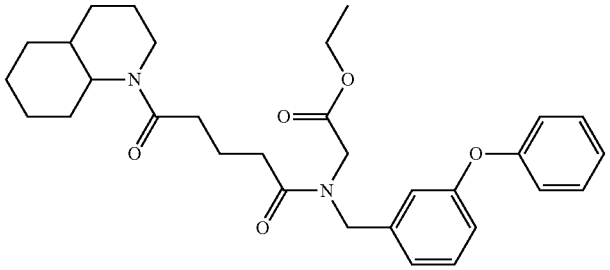
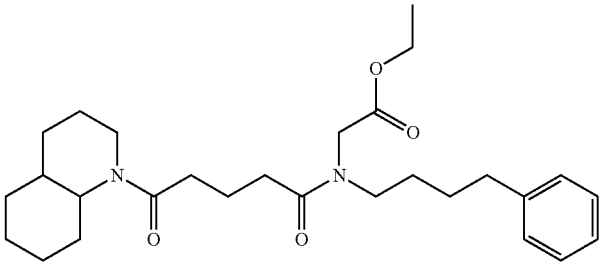
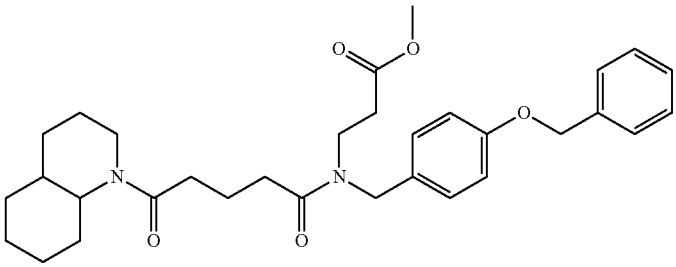
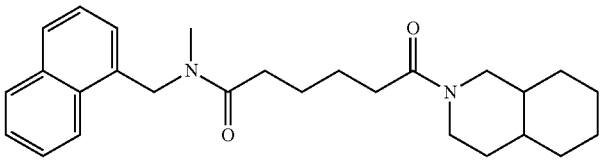
Ex.	structure	name	method	rt (min)	m/z
Ia.257		2-[N-[4-(benzyloxy)benzyl]-5-(octahydroquinolin-1-yl)-5-oxopentanamido)acetic acid methyl ester	B	4.46	521
Ia.258		2-[5-(octahydroquinolin-1-yl)-5-oxo-N-(3-phenoxybenzyl)pentan-amido]acetic acid ethyl ester	B	4.65	521
Ia.259		2-[5-(octahydroquinolin-1-yl)-5-oxo-N-(4-phenylbutyl)pentan-amido]acetic acid ethyl ester	B	4.54	471
Ia.260		3-(N-[4-(benzyloxy)benzyl]-5-(octahydroquinolin-1-yl)-5-oxopentanamido]pro-panoic acid methyl ester	B	4.48	535
Ia.261		6-(octahydroisoquinolin-2-yl)-6-oxohexanoic acid N-methyl-N-(naphthalen-1-ylmethyl)amide	B	4.31 4.40	421 421

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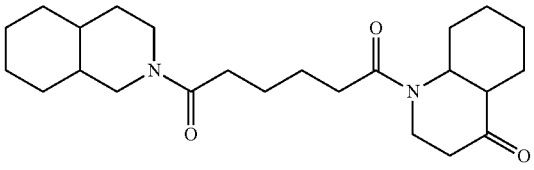
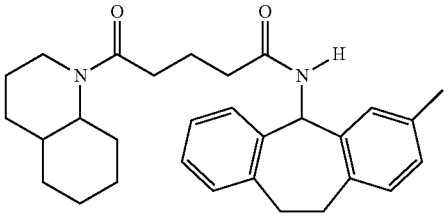
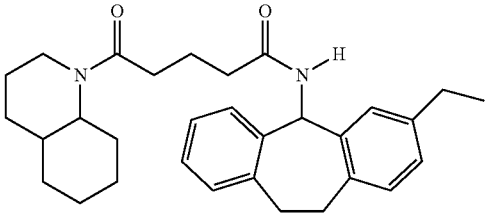
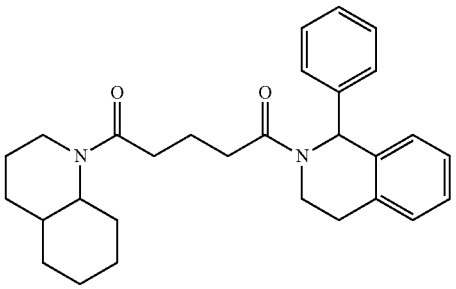
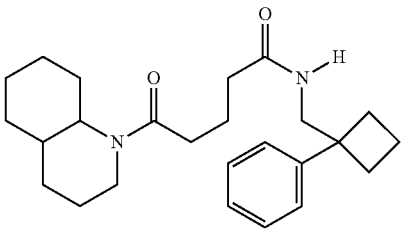
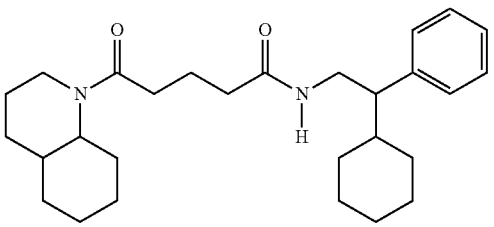
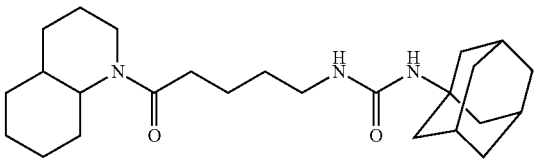
Ex.	structure	name	method	rt (min)	m/z
Ia.262		1-(octahydroisoquinolin-2-yl)-6-(4-oxooctahydroquinolin-1-yl)hexane-1,6-dione	B	3.72 3.81	403 403
Ia.263		5-(octahydro-quinolin-1-yl)-5-oxo-pentanoic acid (3-methyl-10,11-dihydro-5H-dibenzo[a,d]cyclohepten-5-yl)amide	C	5.01	459
Ia.264		5-(octahydro-quinolin-1-yl)-5-oxo-pentanoic acid (3-ethyl-10,11-dihydro-5H-dibenzo[a,d]cyclohepten-5-yl)amide	C	5.20	473
Ia.265		5-[(R)-1-phenyl-3,4-dihydro-1H-isoquinolin-2-yl]-1-(octahydro-quinolin-1-yl)-pentane-1,5-dione	C	5.02	445
Ia.266		5-(octahydro-quinolin-1-yl)-5-oxo-pentanoic acid (1-phenyl-cyclobutylmethyl)amide	C	4.64	397
Ia.267		5-(octahydro-quinolin-1-yl)-5-oxo-pentanoic acid (2-cyclohexyl-2-phenyl-ethyl)amide	C	5.19	439

Ex. Ie.1

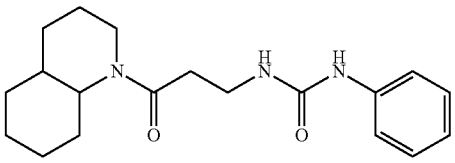
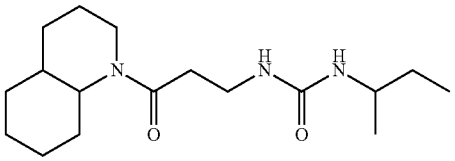
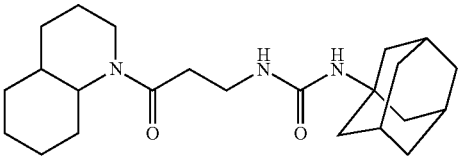
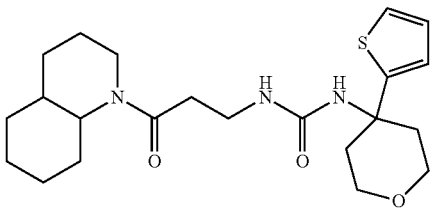
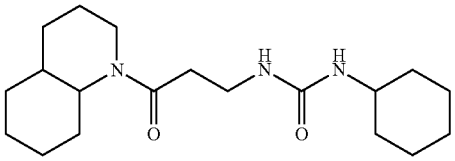
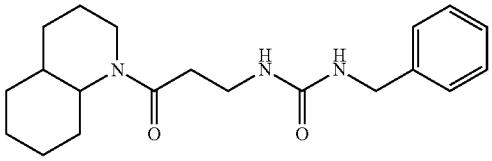
1-[5-(octahydroquinolin-1-yl)-5-oxopentyl]-3-tricyclo[3.3.1.1^{3,7}]decan-1-ylurea

[0165]



[0166] 35 mg (0.15 mmol) of intermediate Vc.4 are dissolved in 2 mL of anhydrous THF and 23 μL (0.25 mmol) of Et_3N. Once dissolved 17.3 mg (0.1 mmol) of 1-adamantyl isocyanate are slowly added and it is refluxed for 2 days. AcOEt is then added and the resulting solution is sequentially washed with water, 1N HCl and brine. The organic phase is dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure yielding 47.5 mg of a yellowish oil identified as example Ie.1. Method B: tr: 4.27 min; m/z: 416.

[0167] The following examples were prepared in a manner similar to example Ie.1.

Ex.	structure	name	method	rt (min)	m/z
Ie. 2		1-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-3-phenylurea	B	3.49	330
Ie. 3		1-s-butyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]urea	B	3.28 3.56	310 310
Ie. 4		1-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-3-tricyclo[3.3.1.1^{3,7}]decan-1-ylurea	B	4.18	388
Ie. 5		1-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-3-(4-thiophen-2-yltetrahydropyran-4-yl)urea	B	3.40	420
Ie. 6		1-cyclohexyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]urea	A	6.55	336
Ie. 7		1-benzyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]urea	A	6.28	344

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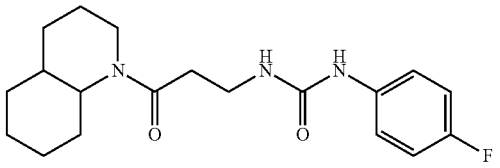
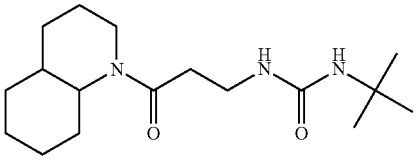
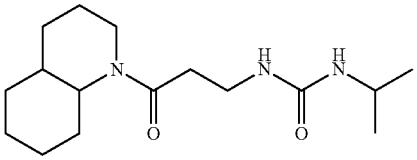
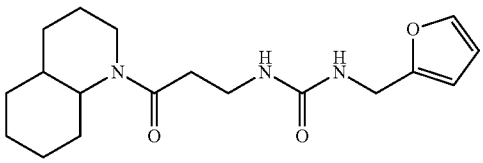
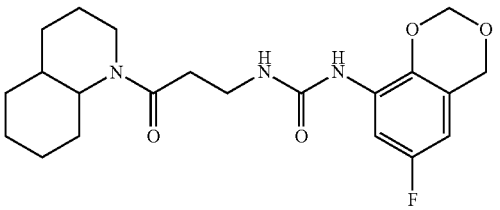
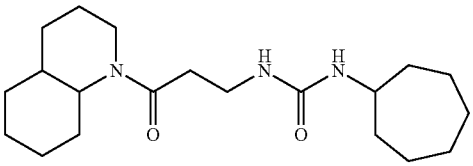
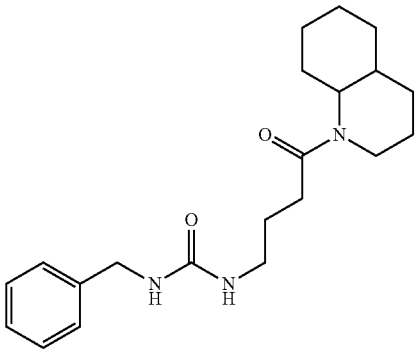
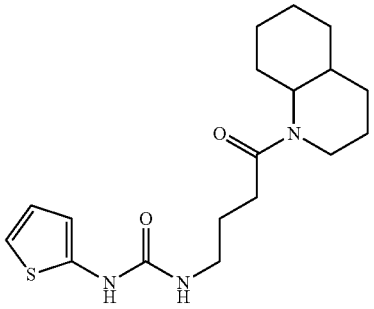
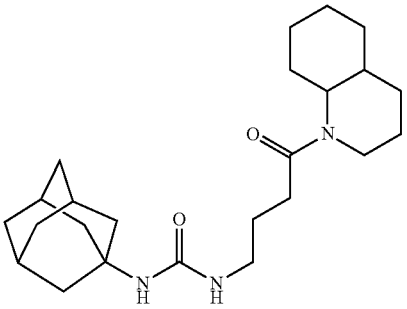
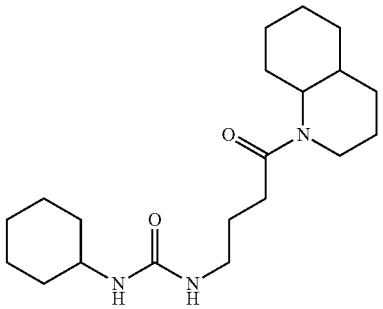
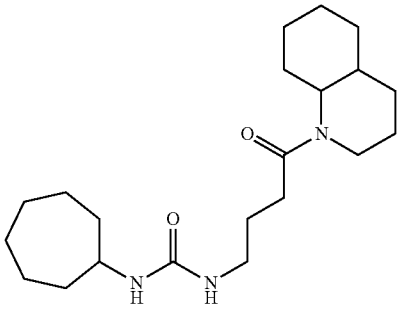
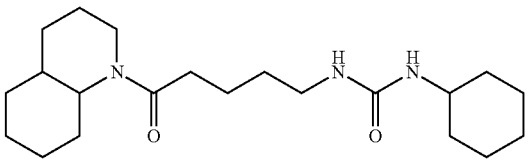
Ex.	structure	name	method	rt (min)	m/z
Ie. 8		1-(4-fluorophenyl)-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]urea	A	6.56	348
Ie. 9		1-t-butyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]urea	A	6.25	310
Ie. 10		1-isopropyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]urea	A	5.65	296
Ie. 11		1-furan-2-ylmethyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]urea	A	5.85	334
Ie. 12		1-(6-fluoro-4-H-1,3-benzodioxin-8-yl)-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]urea	A	6.75	406
Ie. 13		1-cycloheptyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]urea	A	6.93	350
Ie. 18		1-benzyl-3-[4-(octahydroquinolin-1-yl)-4-oxobutyl]urea	B	3.44	358

-continued

Ex.	structure	name	method	rt (min)	m/z
Ie. 19		1-[4-(octahydroquinolin-1-yl)-4-oxobutyl]-3-thiophen-2-ylurea	B	3.41	350
Ie. 20		1-[4-(octahydroquinolin-1-yl)-4-oxobutyl]-3-tricyclo[3.3.1.1^{3,7}]decan-1-ylurea	B	4.18	402
Ie. 21		1-cyclohexyl-3-[4-(octahydroquinolin-1-yl)-4-oxobutyl]urea	B	3.59	350
Ie. 22		1-cycloheptyl-3-[4-(octahydroquinolin-1-yl)-4-oxobutyl]urea	B	3.80	364
Ie. 23		1-cyclohexyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]urea	B	3.69	364

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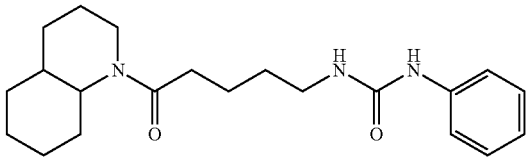
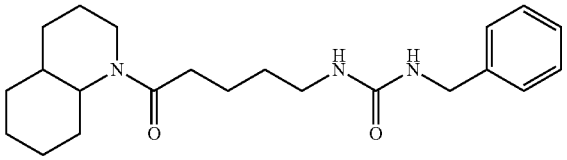
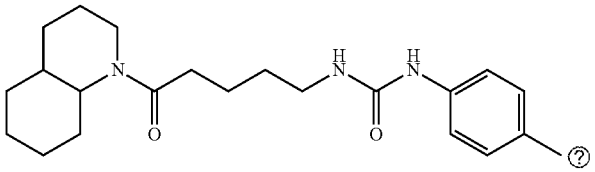
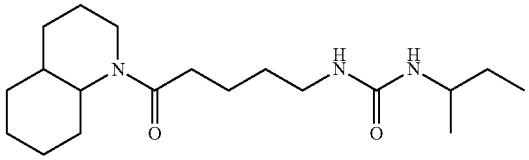
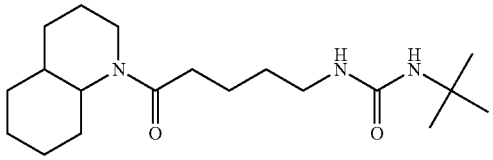
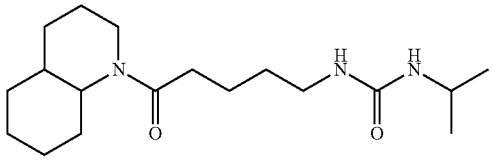
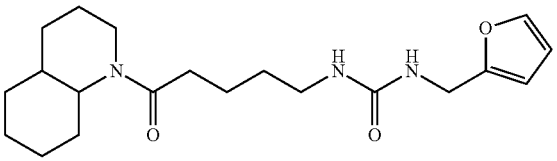
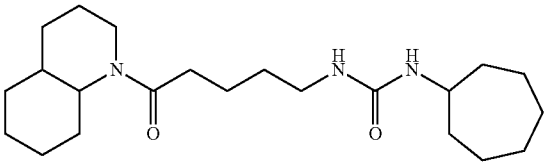
Ex.	structure	name	method	rt (min)	m/z
Ie. 24		1-[5-(octahydroquinoln-1-yl)-5-oxopentyl]-3-phenylurea	B	3.61	358
Ie. 25		1-benzyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]urea	B	3.55	372
Ie. 26		1-(4-fluorophenyl)-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]urea	B	3.67	376
Ie. 27		1-s-butyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]urea	B	3.44	338
Ie. 28		1-t-butyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]urea	B	3.55	338
Ie. 29		1-isopropyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]urea	B	3.24	324
Ie. 30		1-furan-2-ylmethyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]urea	B	3.33	362
Ie. 31		1-cycloheptyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]urea	B	3.88	378

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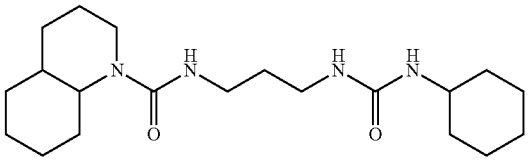
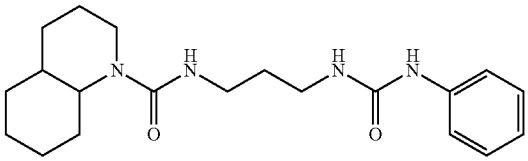
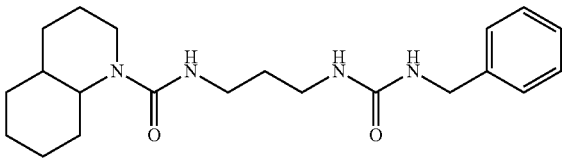
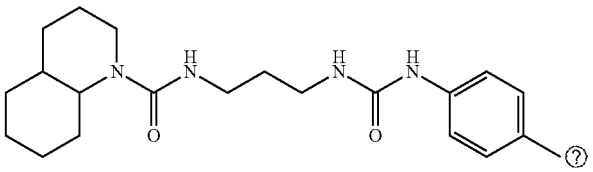
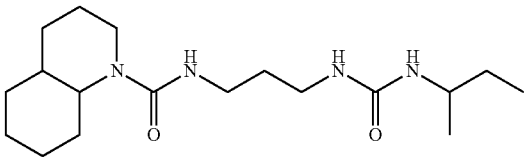
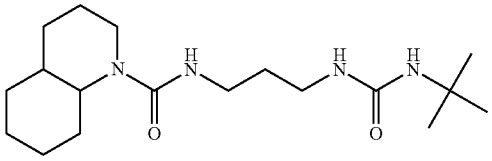
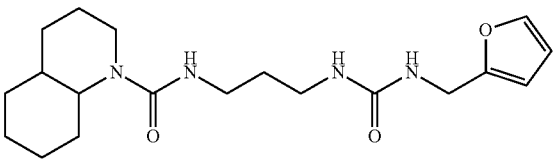
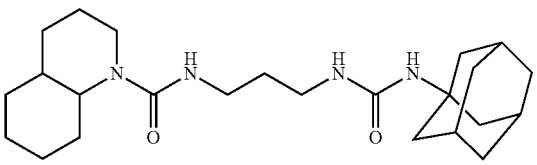
Ex.	structure	name	method	rt (min)	m/z
Ig. 1		Octahydroquinoline-1-carboxylic acid [3-(3-cyclohexylureido) propyl]amide	B	3.60	365
Ig. 2		Octahydroquinoline-1-carboxylic acid [3-(3-phenylureido)propyl]amide	B	3.52	359
Ig. 3		Octahydroquinoline-1-carboxylic acid [3-(3-benzylureido)propyl]amide	B	3.45	373
Ig. 4		Octahydroquinoline-1-carboxylic acid {3-[3-(4-fluorophenyl)ureido]propyl}amide	B	3.58	377
Ig. 5		Octahydroquinoline-1-carboxylic acid [3-(3-s-butylureido)propyl]amide	B	3.33	339
Ig. 6		Octahydroquinoline-1-carboxylic acid [3-(3-t-butylureido)propyl]amide	B	3.48	339
Ig. 7		Octahydroquinoline-1-carboxylic acid [3-(3-furan-2-ylmethylureido)propyl]amide	B	3.22	363
Ig. 8		Octahydroquinoline-1-carboxylic acid [3-(3-tricyclo[3.3.1.1^{3,7}]decan-1-ylureido)propyl]amide	B	4.26	417

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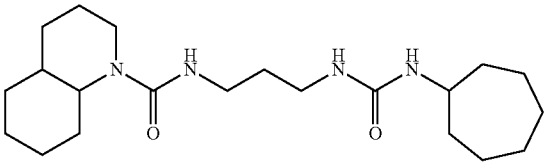
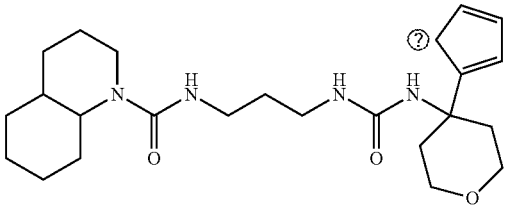
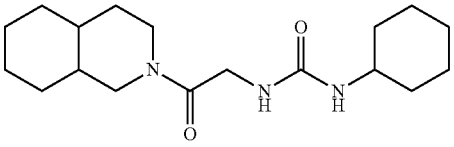
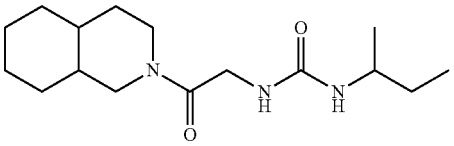
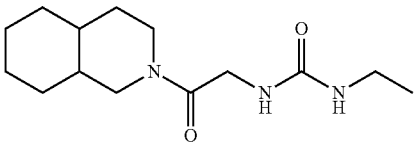
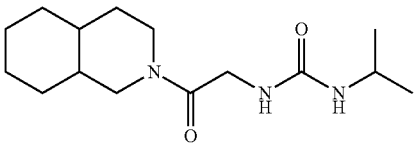
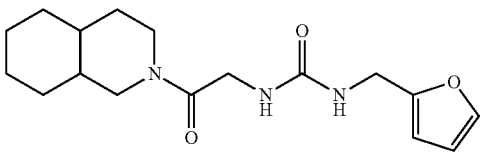
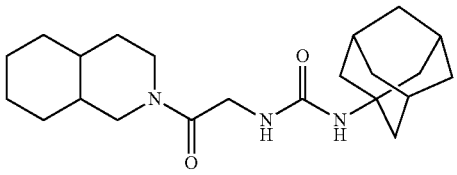
Ex.	structure	name	method	rt (min)	m/z
Ig. 9		Octahydroquinoline-1-carboxylic acid [3-(3-cycloheptylureido)propyl]amide	B	3.80	379
Ig. 10		Octahydroquinoline-1-carboxylic acid {3-[3-(4-thiophen-2-yltetrahydropyran-4-yl)ureido]propyl}amide	B	3.47	449
Ie. 32		1-cyclohexyl-3-[2-(octahydroisoquiolin-2-yl)-2-oxoethyl]urea	B	3.52 3.62	322 322
Ie. 33		1-s-butyl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]urea	B	3.35	296
Ie. 34		1-ethyl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]urea	B	2.78 2.90	268 268
Ie. 35		1-isopropyl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]urea	B	3.01 3.12	282 282
Ie. 36		1-furan-2-ylmethyl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]urea	B	3.15 3.26	320 320
Ie. 37		1-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]-3-tricyclo[3.3.1.1^{3,7}]decan-1-ylurea	B	4.25	374

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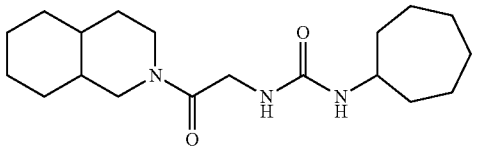
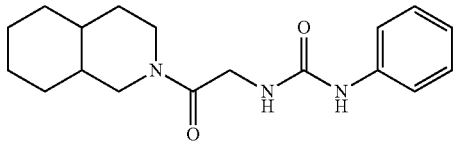
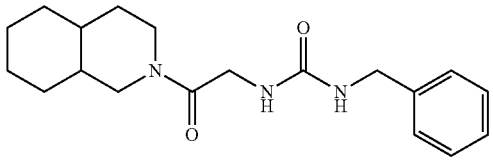
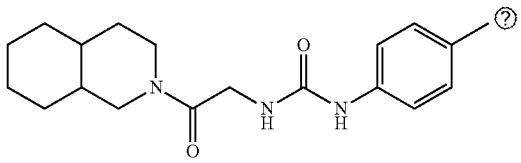
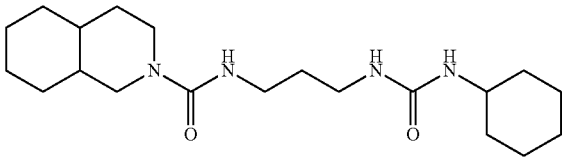
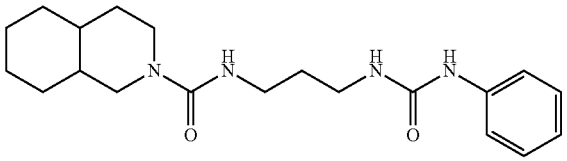
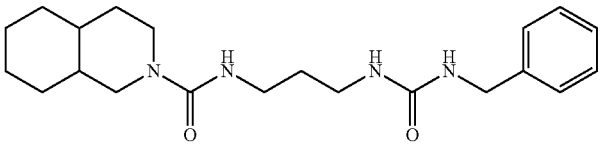
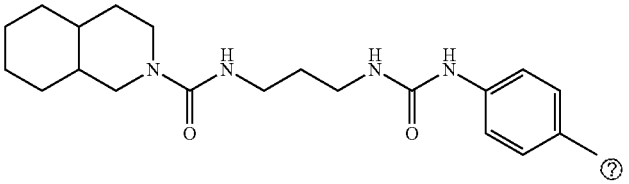
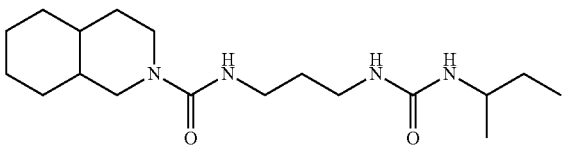
Ex.	structure	name	method	rt (min)	m/z
Ie. 38		1-cycloheptyl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]urea	B	3.83	336
Ie. 39		1-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]-3-phenylurea	B	3.60	316
Ie. 40		1-benzyl-3-[2-(octahydroisoquiolin-2-yl)-2-oxoethyl]urea	B	3.41 3.50	330 330
Ie. 41		1-(4-fluorophenyl)-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]urea	B	3.57 3.67	334 334
Ig. 11		octahydroisoquinoline-2-carboxylic acid [3-(3-cyclohexylureido)propyl]amide	B	3.59 3.68	365 365
Ig. 12		Octahydroisoquinoline-2-carboxylic acid [3-(3-phenylureido)propyl]amide	B	3.51 3.60	359 359
Ig. 13		Octahydroisoquinoline-2-carboxylic acid [3-(3-benzylureido)propyl)amide	B	3.45 3.54	373 373
Ig. 14		Octahydroisoquinoline-2-carboxylic acid {3-[3-(4-fluorophenyl)ureido]propyl}amide	B	3.57 3.66	377 377
Ig. 15		Octahydroisoquinoline-2-carboxylic acid [3-(3-s-butylureido)propyl]amide	B	3.33 3.42	339 339

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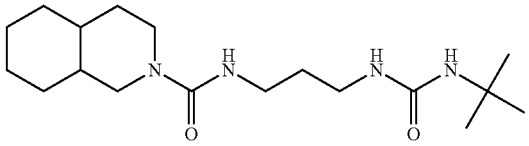
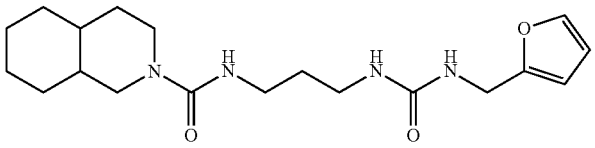
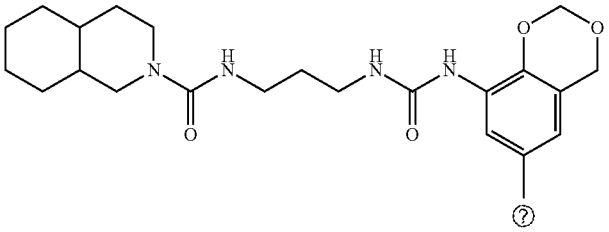
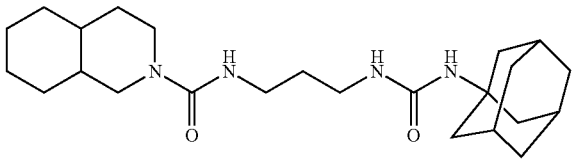
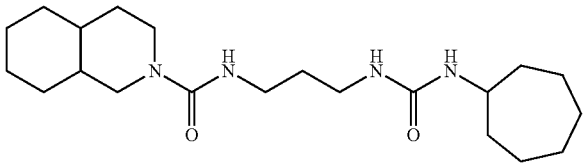
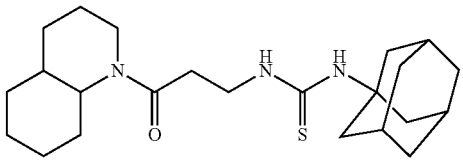
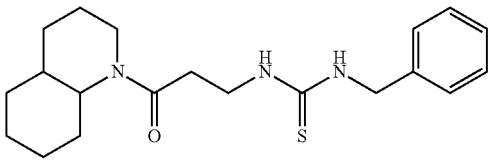
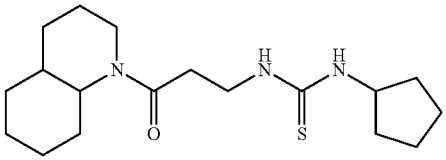
Ex.	structure	name	method	rt (min)	m/z
Ig. 16		Octahydroisoquinoline-2-carboxylic acid [3-(3-t-butylureido)propyl]amide	B	3.46 3.56	339 339
Ig. 17		Octahydroisoquinoline-2-carboxylic acid [3-(3-furan-2-ylmethylureido)propyl]amide	B	3.22 3.32	363 363
Ig. 18		Octahydroisoquinoline-2-carboxylic acid {3-[3-(6-fluoro-4H-1,3-benzodioxin-8-yl)ureido]propyl}amide	B	3.67 3.75	435 435
Ig. 19		Octahydroisoquinoline-2-carboxylic acid [3-(3-tricyclo[3.3.1.1^{3,7}] decan-1-ylureido)propyl]amide	B	4.23 4.31	417 417
Ig. 20		Octahydroisoquinoline-2-carboxylic acid [3-(3-cycloheptylureido)propyl]amide	B	3.79 3.88	379 379
Ie. 43		1-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-3-tricyclo[3.3.1.1^{3,7}]decan-1-ylthiourea	B	4.68	404
Ie. 44		1-benzyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]thiourea	B	3.88	360
Ie. 45		1-cyclopentyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]thiourea	B	3.84	338

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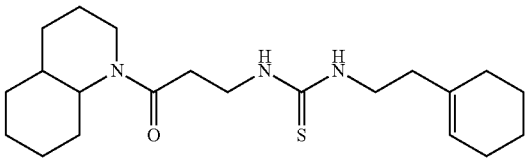
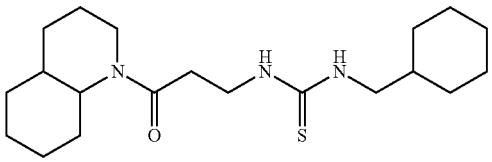
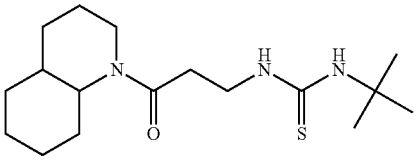
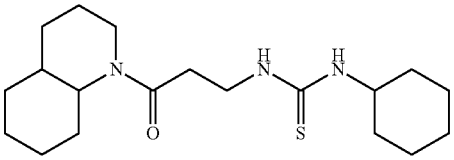
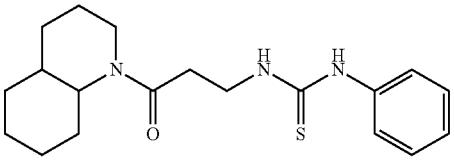
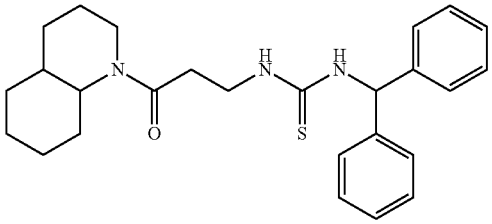
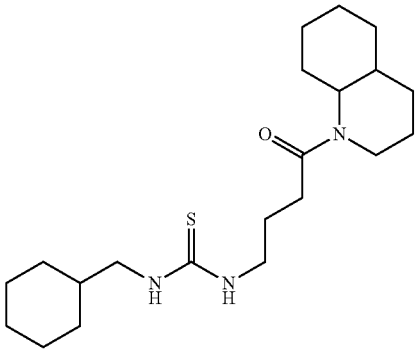
Ex.	structure	name	method	rt (min)	m/z
Ie. 46		1-(2-cyclohex-1-enylethyl)-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]thiourea	B	4.40	378
Ie. 47		1-cyclohexylmethyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]thiourea	A	7.92	366
Ie. 48		1-t-butyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]thiourea	A	7.11	326
Ie. 49		1-cyclohexyl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]thiourea	A	7.47	352
Ie. 50		1-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-3-phenylthiourea	A	6.94	346
Ie. 51		1-benzhydryl-3-[3-(octahydroquinolin-1-yl)-3-oxopropyl]thiourea	A	8.22	436
Ie.52		1-cyclohexylmethyl-3-[4-(octahydroquinolin-1-yl)-4-oxobutyl]thiourea	B	4.33	380

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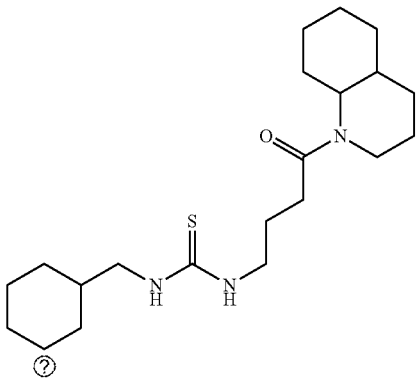
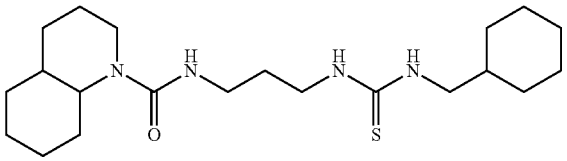
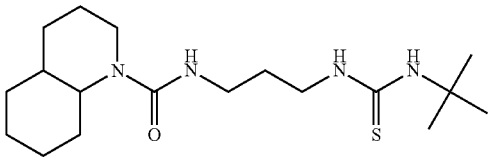
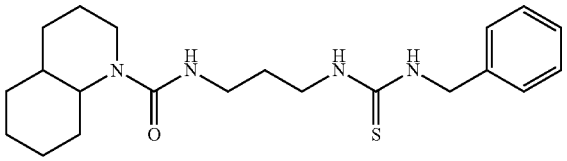
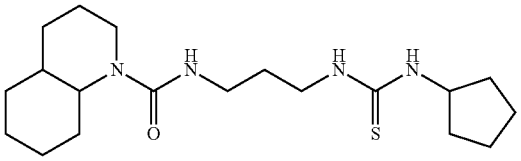
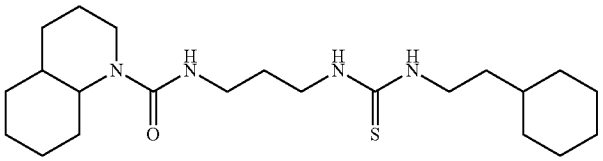
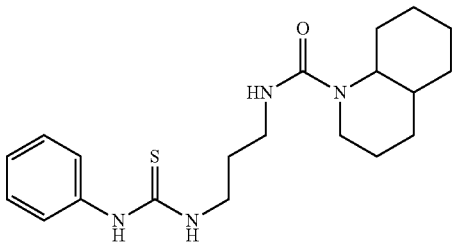
Ex.	structure	name	method	rt (min)	m/z
Ie. 53		1-cyclohexyl-3-[4-(octahydroquinolin-1-yl)-4-oxobutyl]thiourea	B	4.10	366
Ig. 21		Octahydroquinoline-1-carboxylic acid [3-(3-cyclohexylmethyl-thioureido)pro-pyl]amide	B	4.23	395
Ig. 22		Octahydroquinoline-1-carboxylic acid [3-(3-t-butylthioureido)pro-pyl]amide	B	3.79	355
Ig. 23		Octahydroquinoline-1-carboxylic acid [3-(3-benzylthioureido)pro-pyl]amide	B	3.82	389
Ig. 24		Octahydroquinoline-1-carboxylic acid [3-(3-cyclopentyl-thioureido)pro-pyl]amide	B	3.79	367
Ig. 25		Octahydroquinoline-1-carboxylic acid {3-[3-(2-cyclohex-1-enyleth-yl)thioureido]pro-pyl}amide	B	4.32	407
Ig. 26		Octahydroquinoline-1-carboxylic acid [3-(3-phenylthioureido)pro-pyl]amide	B	3.66	375

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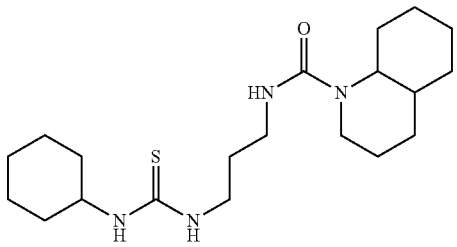
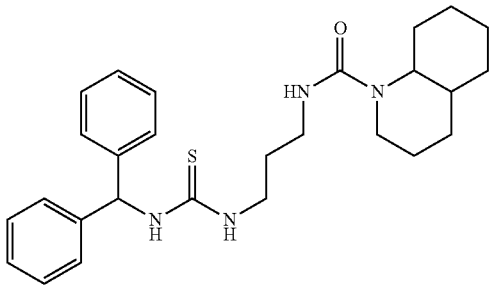
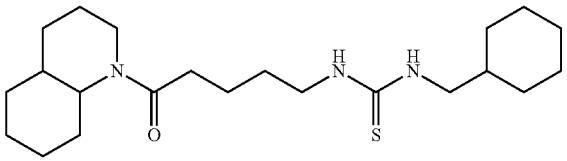
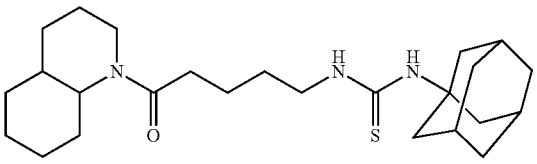
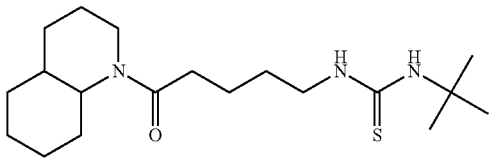
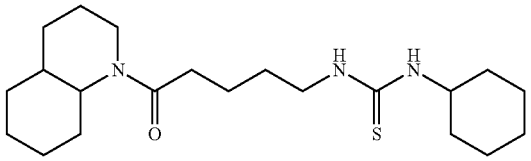
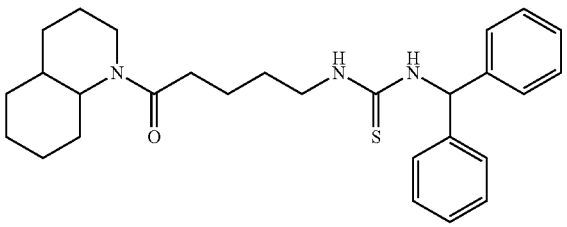
Ex.	structure	name	method	rt (min)	m/z
Ig. 27		Octahydroquinoline-1-carboxylic acid [3-(3-cyclohexylthioureido)propyl]amide	B	4.00	381
Ig. 28		Octahydroquinoline-1-carboxylic acid [3-(3-benzhydrylthioureido)propyl]amide	B	4.41	465
Ie. 54		1-cyclohexylmethyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]thiourea	B	4.32	394
Ie. 55		1-[5-(octahydroquinolin-1-yl)-5-oxopentyl]-3-tricyclo[3.3.1.1^{3,7}]decan-1-ylthiourea	B	4.67	432
Ie. 56		1-t-butyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]thiourea	B	3.90	354
Ie. 57		1-cyclohexyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]thiourea	B	4.10	380
Ie. 58		1-benzhydryl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]thiourea	B	4.49	464

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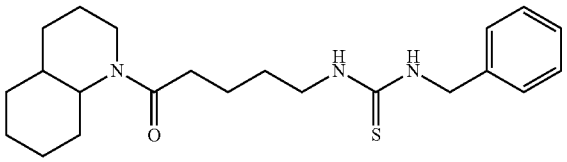
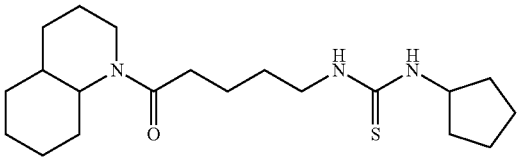
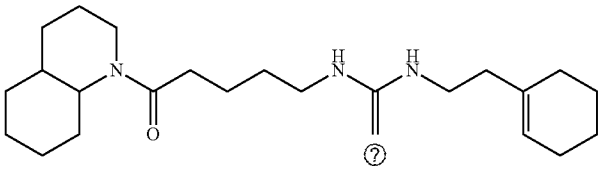
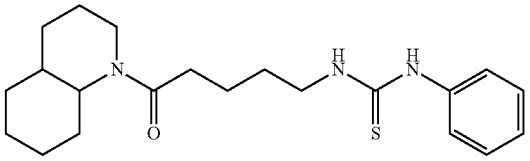
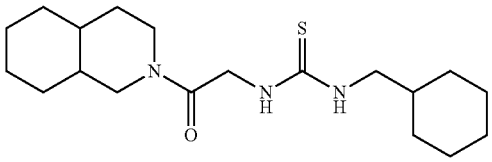
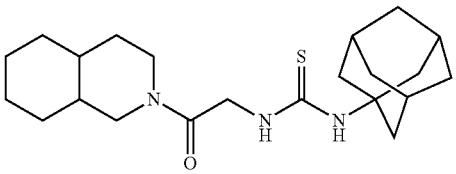
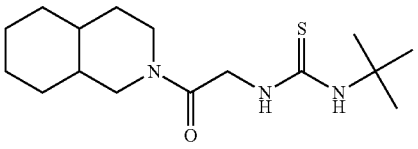
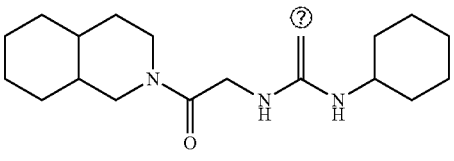
Ex.	structure	name	method	rt (min)	m/z
Ie. 59		1-benzyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]thiourea	B	3.93	388
Ie. 60		1-cyclopentyl-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]thiourea	B	3.89	366
Ie. 61		1-(2-cyclohex-1-enylethyl)-3-[5-(octahydroquinolin-1-yl)-5-oxopentyl]thiourea	B	4.42	406
Ie. 62		1-[5-(octahydroquinolin-1-yl)-5-oxopentyl]-3-phenylthiourea	B	3.81	374
Ie. 63		1-cyclohexylmethyl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]thiourea	B	4.35 4.44	352 352
Ie. 64		1-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]-3-tricyclo[3.3.1.1^{3,7}]decan-1-ylthiourea	B	4.77 4.81	390 390
Ie. 65		1-t-butyl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]thiourea	B	3.90 4.01	312 312
Ie. 66		1-cyclohexyl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]thiourea	B	4.10 4.19	338 338

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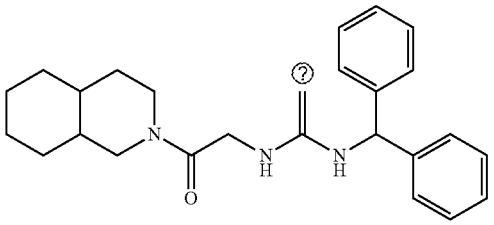
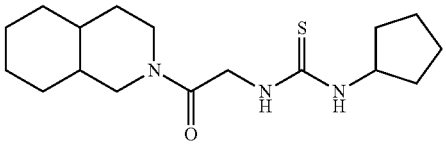
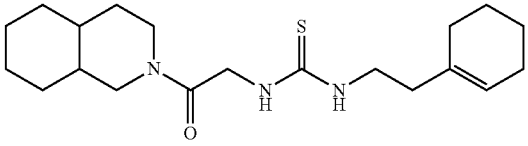
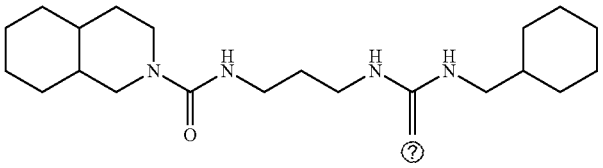
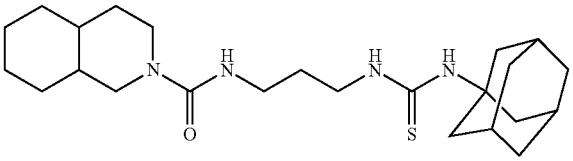
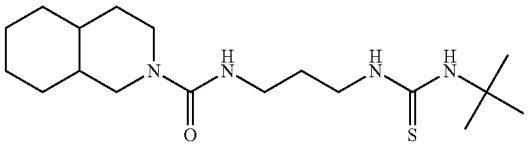
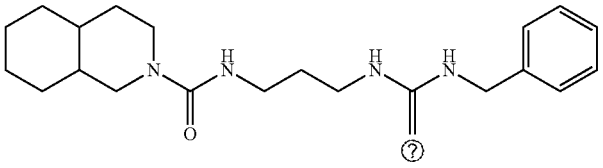
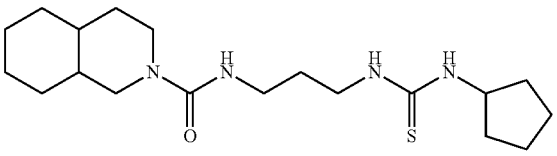
Ex.	structure	name	method	rt (min)	m/z
Ie. 67		1-benzhydryl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]thiourea	B	4.52 4.59	nd nd
Ie. 68		1-cyclopentyl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]thiourea	B	3.89 3.99	324 324
Ie. 69		1-(2-cyclohex-1-enylethyl)-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]thiourea	B	4.45 4.53	364 364
Ig. 29		Octahydroisoquinoline-2-carboxylic acid [3-(3-cyclohexylmethyl-thioureido)pro-pyl]amide	B	4.22 4.30	395 395
Ig. 30		Octahydroisoquinoline-2-carboxylic acid [3-(3-tricyclo[3.3.1.1^{3,7}]decan-1-ylthioureido)pro-pyl]amide	B	4.65 4.71	433 433
Ig. 31		Octahydroisoquinoline-2-carboxylic acid [3-(3-t-butylthioureido)pro-pyl]amide	B	3.78 3.87	355 355
Ig. 32		Octahydroisoquinoline-2-carboxylic acid [3-(3-benzylthioureido)pro-pyl]amide	B	3.81 3.89	389 389
Ig. 33		Octahydroisoquinoline-2-carboxylic acid [3-(3-cyclopentyl-thioureido)pro-pyl]amide	B	3.78 3.87	367 367

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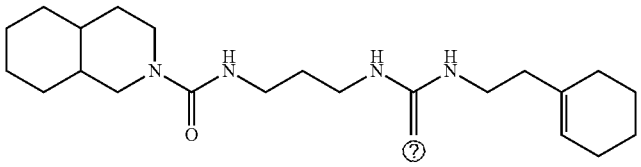
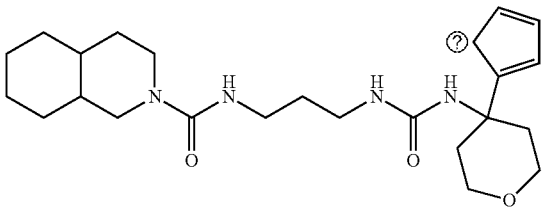
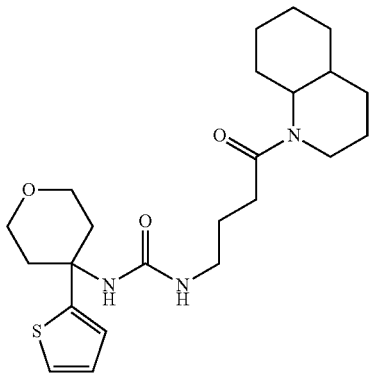
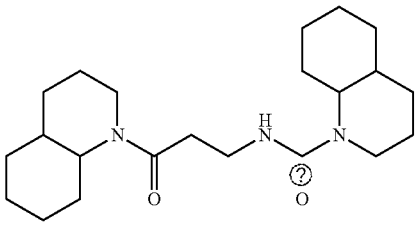
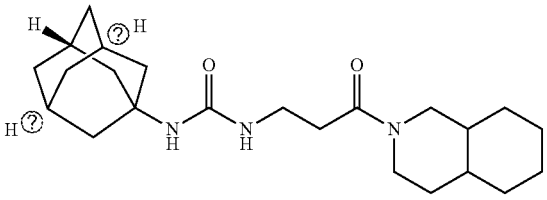
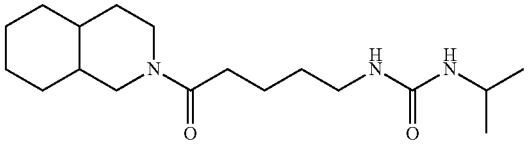
Ex.	structure	name	method	rt (min)	m/z
Ig. 34		Octahydroisoquinoline-2-carboxylic acid {3-[3-(2-cyclohex-1-enyleth-yl)thioureido]pro-pyl}amide	B	4.31 4.39	407 407
Ig. 35		Octahydroisoquinoline-2-carboxylic acid{3-[3-(4-thiophen-2-yltetrahydropyran-4-yl)ureido]pro-pyl}amide	B	3.45 3.54	449 449
Ie. 70		1-[4-(octahydroquinolin-1-yl)-4-oxobutyl]-3-[4-(thiophen-2-yl)tetrahydropyran-4-yl]urea	B	3.43	434
Ie. 71		Octahydroquinoline-1-carboxylic acid N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	B	4.09	376
Ie. 72		1-[(3s,5s,7s)-adamantan-1-yl]-3-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]urea	B	4.17 4.26	388 388
Ie. 73		1-isopropyl-3-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]urea	B	4.60 4.68	324 324

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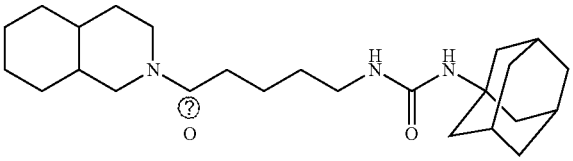
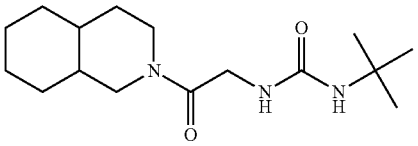
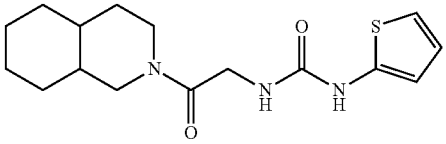
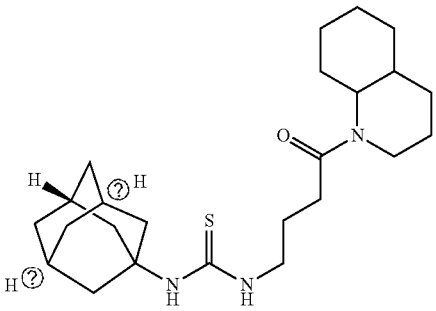
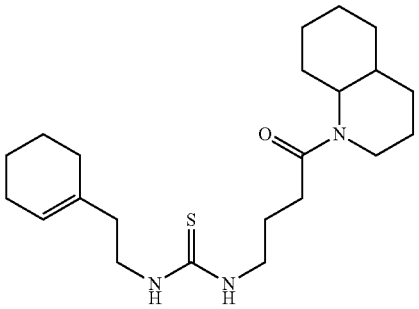
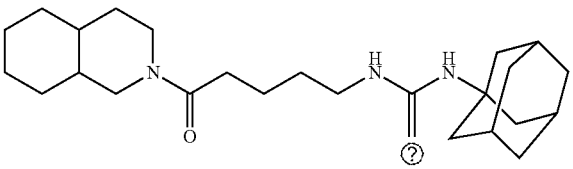
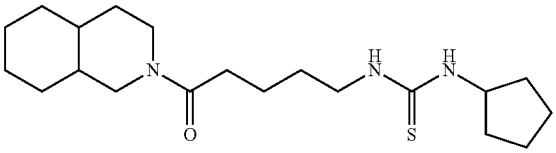
Ex.	structure	name	method	rt (min)	m/z
Ie. 74		1-[(3s,5s,7s)-adamantan-1-yl]-3-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]urea	B	4.27 4.35	416 416
Ie. 75		1-t-butyl-3-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]urea	B	3.47	296
Ie. 76		1-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]-3-(thiophen-2-yl)urea	B	3.50	322
Ie. 77		1-[(3s,5s,7s)-adamantan-1-yl]-3-[4-(octahydroquinolin-1-yl)-4-oxobutyl]thiourea	B	4.68	418
Ie. 78		1-[2-(cyclohex-1-en-1-yl)ethyl]-3-[4-(octahydroquinolin-1-yl)-4-oxobutyl]thiourea	B	4.44	392
Ie. 79		1-[(3s,5s,7s)-adamantan-1-yl]-3-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]thiourea	B	4.67 4.74	432 432
Ie. 80		1-cyclopentyl-3-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]thiourea	B	3.88 3.98	366 366

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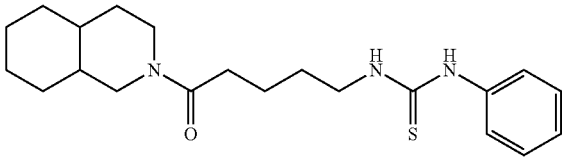
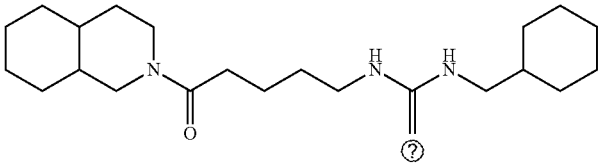
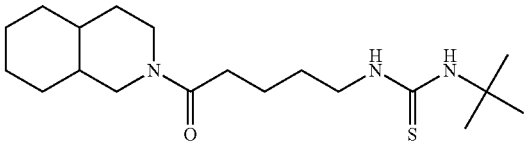
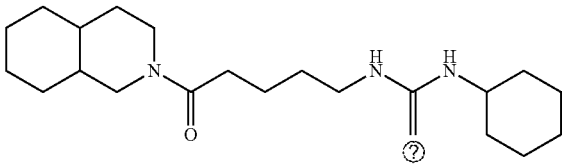
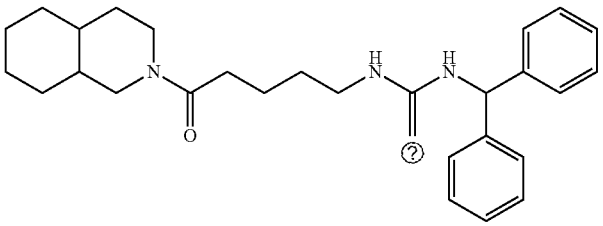
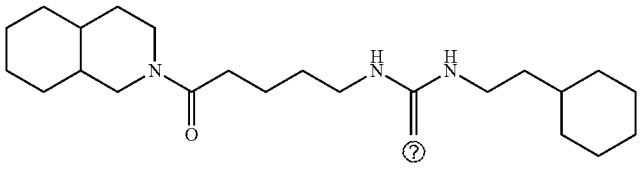
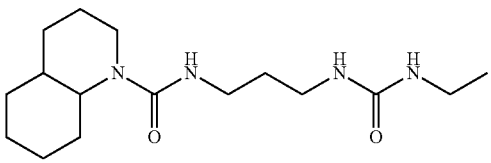
Ex.	structure	name	method	rt (min)	m/z
Ie. 81		3-phenyl-1-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]-thiourea	B	3.81 3.87	374 374
Ie. 82		1-(cyclohexylmethyl)-3-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]thiourea	B	4.32 4.40	394 394
Ie. 83		1-t-butyl-3-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]thiourea	B	3.89 3.99	354 354
Ie. 84		1-cyclohexyl-3-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]thiourea	B	4.09 4.18	380 380
Ie. 85		1-benzhydryl-3-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]thiourea	B	4.49 4.56	464 464
Ie. 86		1-[2-(cyclohex-1-en-1-yl)ethyl]-3-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]thiourea	B	4.41 4.49	406 406
Ig. 36		Octahydroquinoline-1-carboxylic acid N-[3-(3-ethylureido)propyl]amide	B	2.91	311

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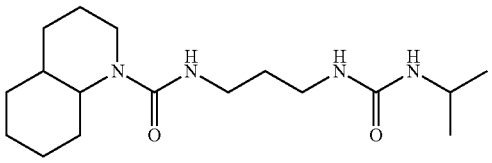
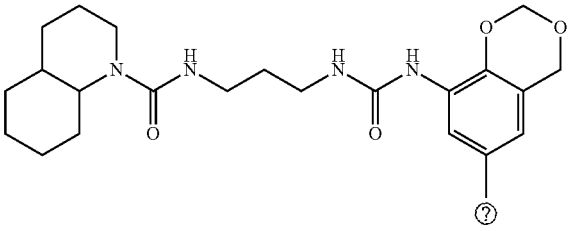
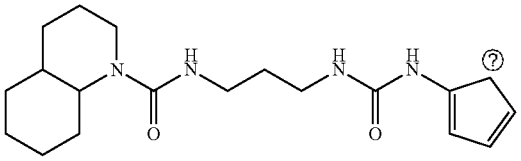
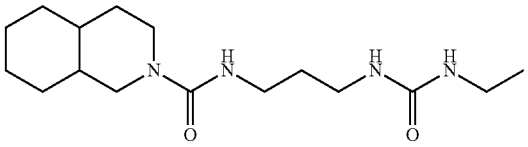
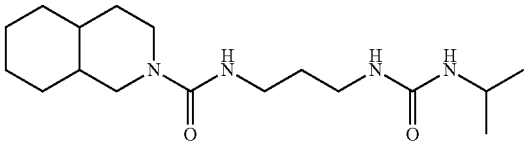
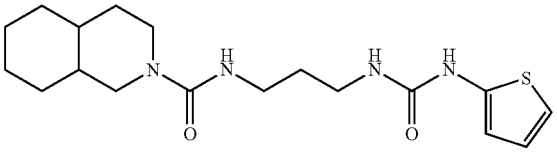
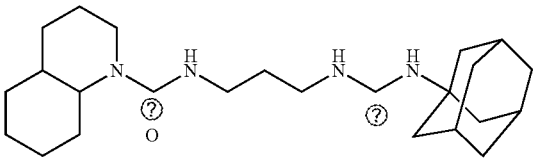
Ex.	structure	name	method	rt (min)	m/z
Ig. 37		Octahydroquinoline-1-carboxylic acid N-[3-(3-isopropylureido)propyl]amide	B	3.13	325
Ig. 38		Octahydroquinoline-1-carboxylic acid N-[3-[3-(6-fluoro-4H-benzo[d][1,3]dioxin-8-yl)ureido]propyl]amide	B	3.69	435
Ig. 39		Octahydroquinoline-1-carboxylic acid N-[3-[3-thiophen-2-yl)ureido]propyl]amide	B	3.34	365
Ig. 40		Octahydroisoquinoline-2-carboxylic acid N-[3-(3-ethylureido)propy 1]amide	B	3.02	311
Ig. 41		Octahydroisoquinoline-2-carboxylic acid N-[3-(3-isopropylureido)propyl]amide	B	3.23	325
Ig. 42		Octahydroisoquinoline-2-carboxylic acid N-[3-[3-(thiophen-2-yl)ureido]propyl]amide	B	3.51	365
Ig. 43		Octahydroquinoline-1-carboxylic acid N-[3-[3-[(3s,5s,7s)-adamantan-1-yl)thioureido]propyl]amide	B	4.67	433

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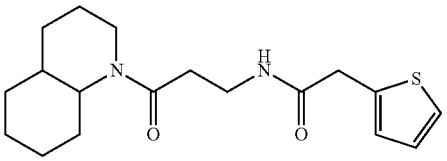
Example Id.1

N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-2-thiophen-2-ylacetamide

[0168]

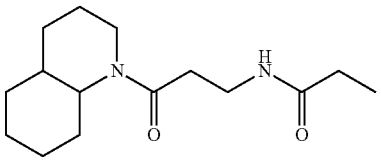


[0169] 46 μL of Et_3N, 30.4 mg (0.22 mmol) of HOBT and 43.1 mg (0.22 mmol) of EDC are added to a solution of 21.3 mg (0.15 mmol) of 2-thienylacetic acid in 2 mL of AcOEt; 35 mg (0.17 mmol) of the intermediate amine Vc.3 are added to it. The suspension formed is kept under stirring for 18 h. It is then treated with water and more AcOEt is added, the organic phase is separated and the aqueous phase is extracted once more with more AcOEt. The organic phases are pooled and successively washed with saturated $NaHCO_3$ solution, 1N HCl and brine. It is then dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. The residue is purified by means of silica gel column chromatography, using a (30:1) mixture of DCM-MeOH as eluent, yielding 5 mg of an oil identified as example Id.1. Method A: tr: 6.30 min; m/z: 335.

Example Id.2

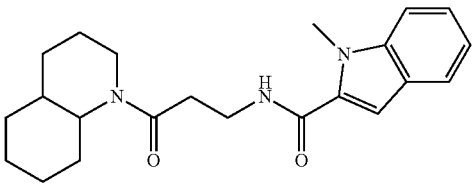
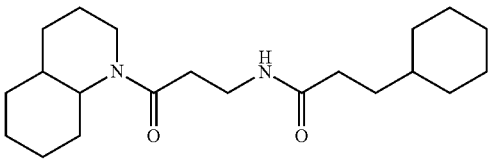
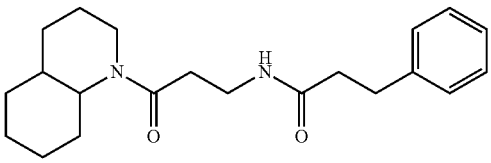
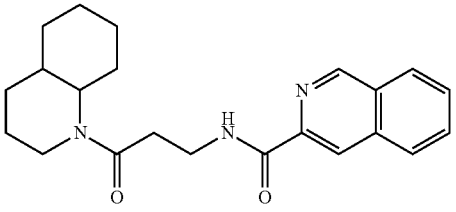
N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]propionamide

[0170]



[0171] 35 mg (0.17 mmol) of the intermediate amine Vc.3 and 46 μL of Et_3N are added to a solution of 13 μL (0.15 mmol) of propionyl chloride in 2 mL of AcOEt. The solution formed is kept under stirring for 18 h. It is then treated with water and more AcOEt is added, the organic phase is separated and the aqueous phase is extracted once more with more AcOEt. The organic phases are pooled and successively washed with saturated $NaHCO_3$ solution, 1N HCl and brine. It is then dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure, 11 mg of example Id.2 being obtained. Method A: tr: 5.35 min; m/z: 267.

[0172] The following examples were prepared in a manner similar to examples Id.1 or Id.2:

Ex.	structure	name	method	rt (min)	m/z
Id.3		1-methyl-1H-indole-2-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	7.41	368
Id.4		3-cyclohexyl-N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]propionamide	A	7.70	349
Id.5		N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-3-phenylpropionamide	A	6.71	343
Id.6		Isoquinoline-3-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	7.05	366

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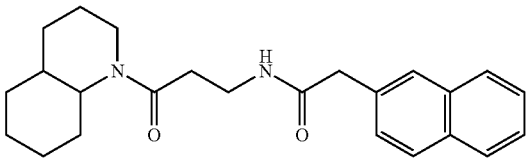
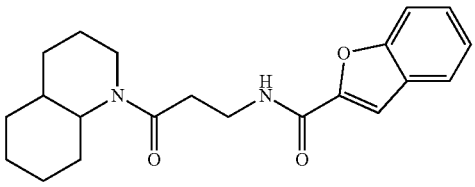
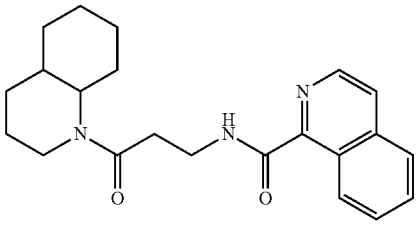
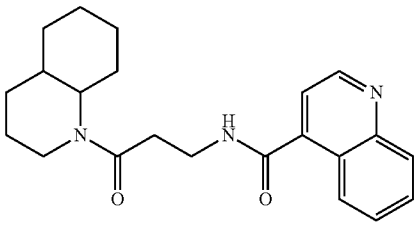
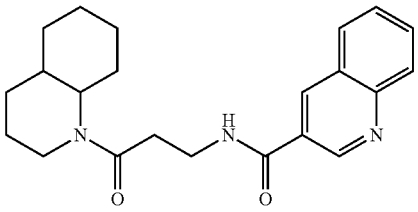
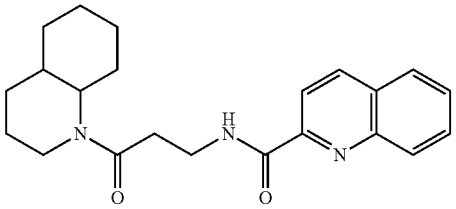
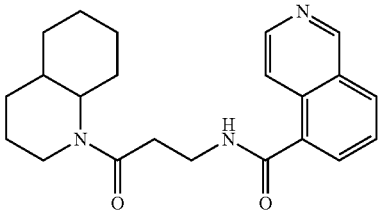
Ex.	structure	name	method	rt (min)	m/z
Id.7		2-naphthalen-2-yl-N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]acetamide	A	7.16	379
Id.8		Benzofuran-2-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	7.03	355
Id.9		Isoquinoline-1-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	7.11	366
Id.10		Quinoline-4-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	5.65	366
Id.11		Quinoline-3-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	6.00	366
Id.12		Quinoline-2-carboxylic acid[3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	7.29	366
Id.13		Isoquinoline-5-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	4.68	366

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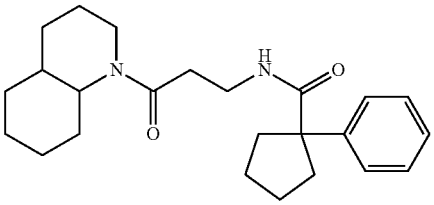
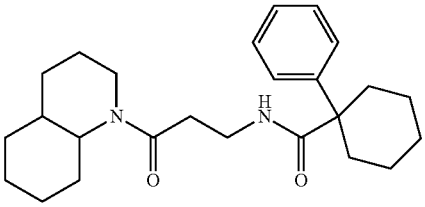
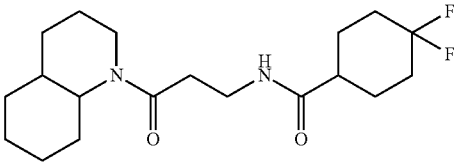
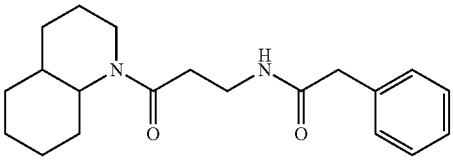
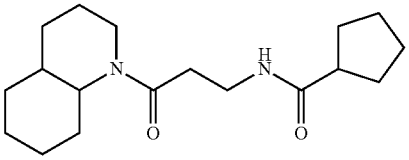
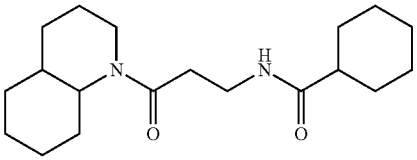
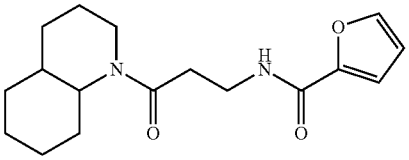
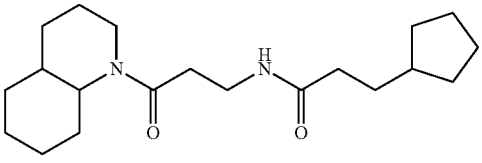
Ex.	structure	name	method	rt (min)	m/z
Id.14		1-phenylcyclopentane-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	7.87	383
Id.15		1-phenylcyclohexane-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	8.21	397
Id.16		4,4-difluorocyclohexane-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	6.49	357
Id.17		N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-2-phenylacetamide	A	6.42	329
Id.18		Cyclopentanecarboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	6.39	307
Id.19		Cyclohexanecarboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	6.76	321
Id.20		Furan-2-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	5.89	305
Id.21		3-cyclopentyl-N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]propionamide	A	7.28	335

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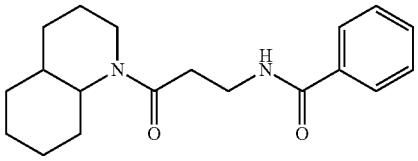
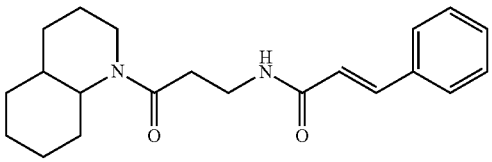
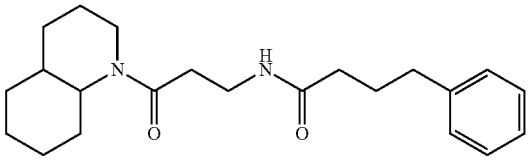
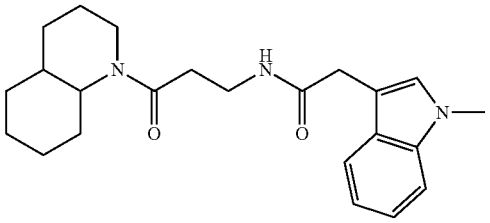
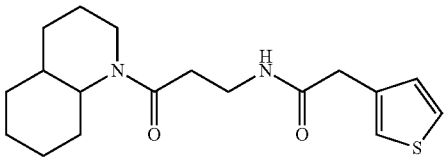
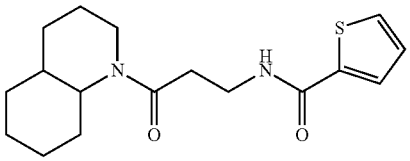
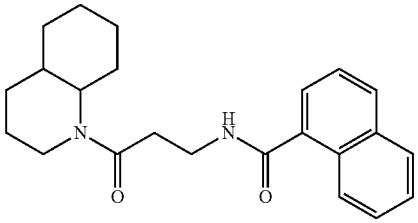
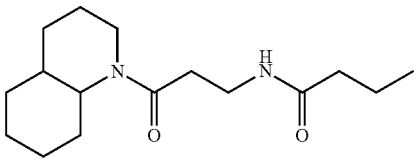
Ex.	structure	name	method	rt (min)	m/z
Id.22		N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]benzamide	A	6.39	315
Id.23		(E)-N-[3 (octahydroquinolin-1-yl)-3-oxopropyl]-3-phenylacrylamide	A	6.77	341
Id.24		N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-4-phenylbutyramide	A	7.05	357
Id.25		2-(1-methyl-1H-indol-3-yl)-N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]acetamide	A	6.83	382
Id.26		N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-2-thiophen-3-ylacetamide	A	6.27	335
Id.27		Thiophene-2-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	6.26	321
Id.28		Naphthalene-1-carboxylic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	7.09	365
Id.29		N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]butyramide	A	5.79	281

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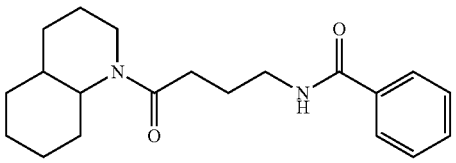
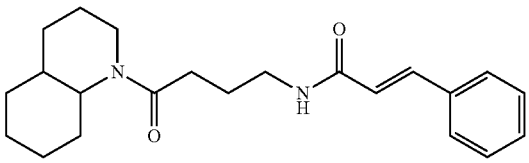
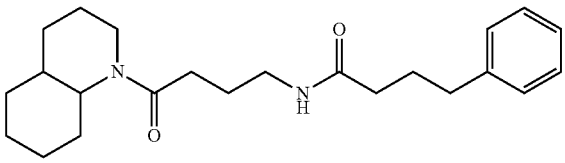
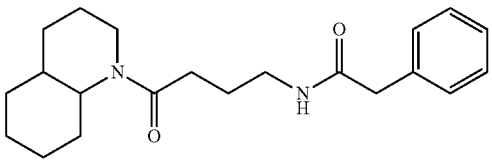
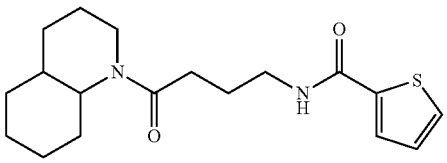
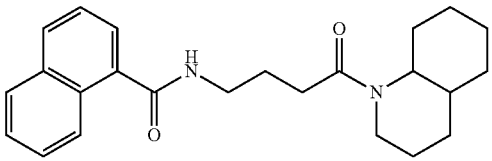
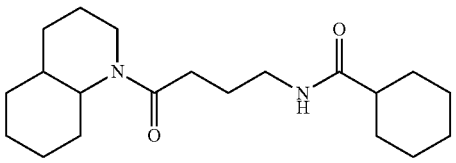
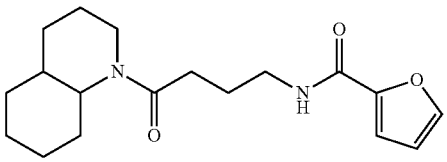
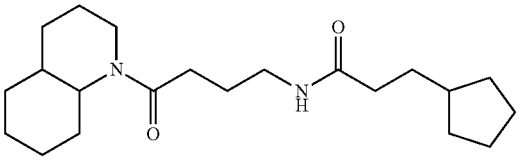
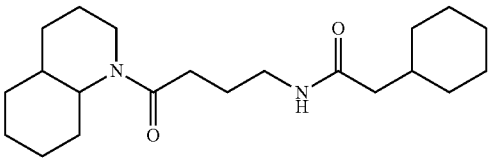
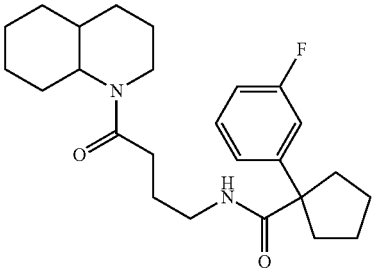
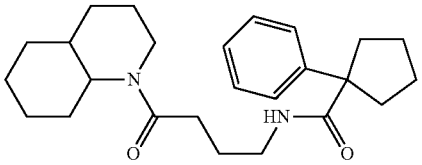
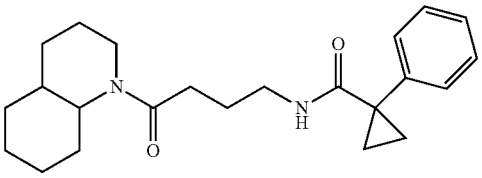
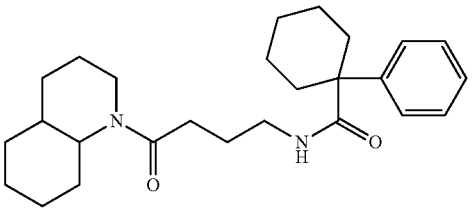
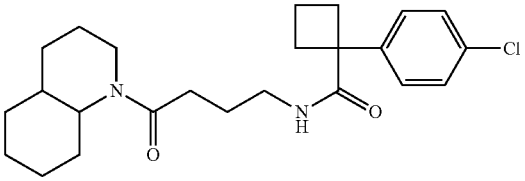
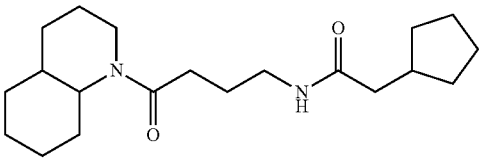
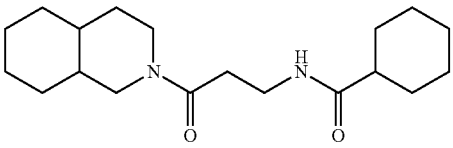
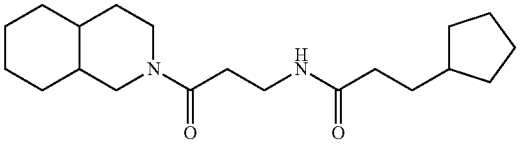
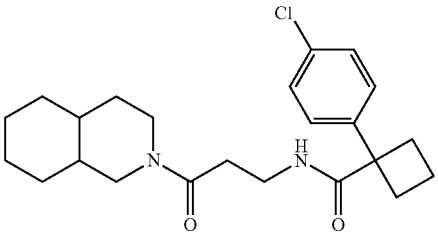
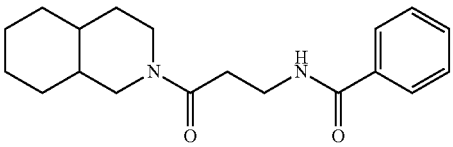
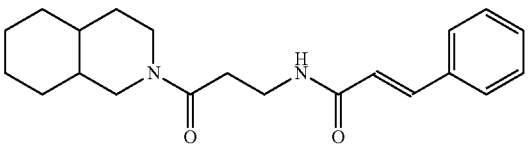
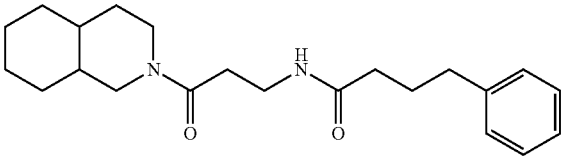
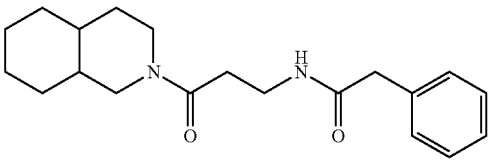
Ex.	structure	name	method	rt (min)	m/z
Id.30		N-[4-(octahydroquinolin-1-yl)-4-oxobutyl]benzamide	A	6.43	329
Id.31		(E)-N-[4-(octahydroquinolin-1-yl)-4-oxobutyl]-3-phenylacrylamide	A	6.75	355
Id.32		N-[4-(octahydroquinolin-1-yl)-4-oxobutyl]-4-phenylbutyramide	A	7.03	371
Id.33		N-[4-(octahydroquinolin-1-yl)-4-oxobutyl]-2-phenylacetamide	A	6.44	343
Id.34		Thiophene-2-carboxylic acid [4-(octahydroquinolin-1-yl)-4-oxobutyl]amide	A	6.32	335
Id.35		Naphthalene-1-carboxylic acid [4-(octahydroquinolin-1-yl)-4-oxobutyl]amide	A	7.04	379
Id.36		Cyclohexanecarboxylic acid [4-(octahydroquinolin-1-yl)-4-oxobutyl]amide	A	6.77	335
Id.37		Furan-2-carboxylic acid [4-octahydroquinolin-1-yl)-4-oxobutyl]amide	A	5.87	319

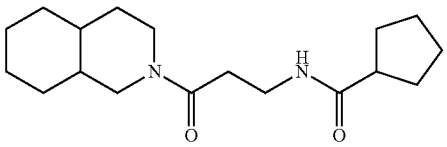
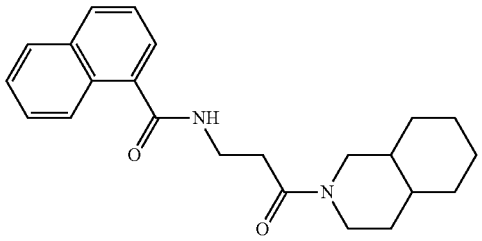
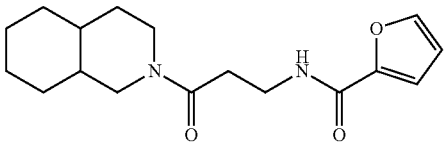
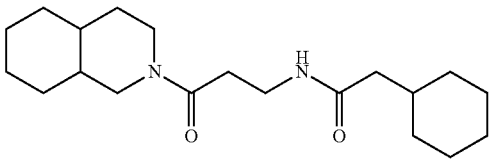
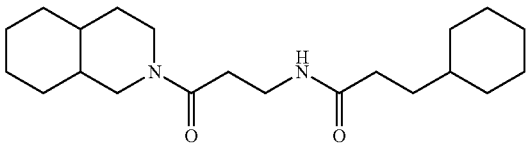
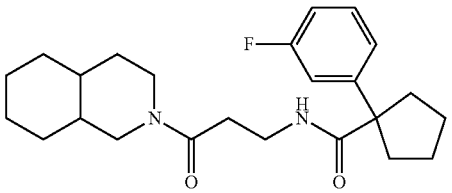
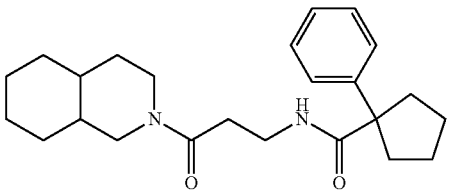
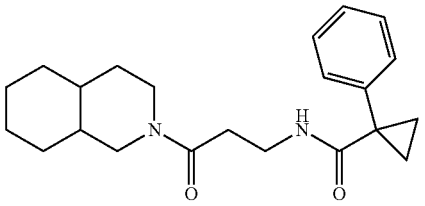
-continued

Ex.	structure	name	method	rt (min)	m/z
Id.38		3-cyclopentyl-N-[4-(octahydroquinolin-1 yl)-4-oxobutyl]propionamide	A	7.27	349
Id.39		2-cyclohexyl-N-[4-(octahydroquinolin-1-yl)-4-oxobutyl]acetamide	A	7.14	349
Id.40		1-(3-fluorophenyl)cyclo-pentanecarboxylic acid [4-(octahydroquinolin-1-yl)-4-oxobutyl]amide	A	7.95	415
Id.41		1-phenylcyclopentane-carboxylic acid [4-(octahydroquinolin-1-yl)-4-oxobutyl]amide	A	7.83	397
Id.42		1-phenylcylopropane-carboxylic acid [4-(octahydroquinolin-1 yl)-4-oxobutyl]amide	A	7.23	369
Id.43		1-phenylcyclohexane-carboxylic acid [4-(octahydroquinolin-1-yl)-4-oxobutyl]amide	A	8.16	411
Id.44		1-(4-chlorophenyl)cyclo-butanecarboxylic acid [4-(octahydroquinolin-1-yl)-4-oxobutyl]amide	A	7.94	417

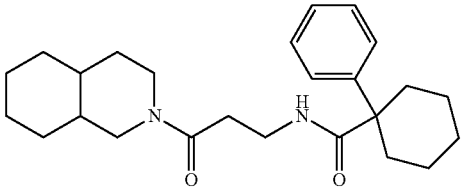
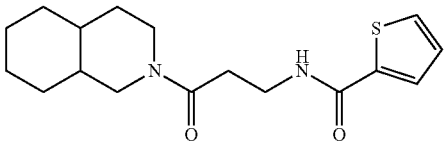
-continued

Ex.	structure	name	method	rt (min)	m/z
Id.45		2-cyclopentyl-N-[4-(octahydroquinolin-1-yl)-4-oxobutyl]acetamide	A	6.77	335
Id.46		Cyclohexanecarboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	A	6.74 6.93	321 321
Id.47		3-cyclopentyl-N-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]propionamide	A	7.27 7.44	335 335
Id.48		1-(4-chlorophenyl)cyclobutanecarboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	A	7.95 8.09	403, 405 403, 405
Id.49		N-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]benzamide	A	6.34 6.53	315 315
Id.50		(E)-N-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]-3-phenylacrylamide	A	6.71 6.88	341 341
Id 51		N-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]-4-phenylbutyramide	A	7.02 7.19	357 357
Id.52		N-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]-2-phenylacetamide	A	6.41 6.59	329 329

-continued

Ex.	structure	name	method	rt (min)	m/z
Id.53		Cyclopentanecarboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	A	6.38 6.57	307 307
Id.54		Naphthalene-1-carboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	A	7.04 7.21	365 365
Id.55		Furan-2-carboxylic acid [3-octahydroisoquinolin-2-yl)-3-oxopropyl]amide	A	5.84 6.05	305 305
Id.56		2-cyclohexyl-N-[3-octahydroisoquinolin-2-yl)-3-oxopropyl]acetamide	A	7.14 7.30	335 335
Id.57		3-cyclohexyl-N-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]propionamide	A	7.67 7.83	349 349
Id.58		1-(3-fluorophenyl)cyclo-pentanecarboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	A	7.95 8.09	401 401
Id.59		1-phenylcyclopentane-carboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	A	7.84 7.99	383 383
Id.60		1-phenylcyclopropane-carboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	A	7.25 7.41	355 355

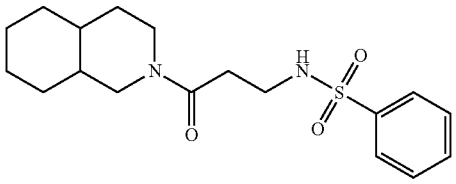
-continued

Ex.	structure	name	method	rt (min)	m/z
Id.61		1-phenylcyclohexane-carboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	A	8.17 8.31	397 397
Id.62		Thiophene-2-carboxylic acid [3-(octahydroisoquinolin-2-yl)-3-oxopropyl]amide	A	6.21 6.39	321 321

Example It.1

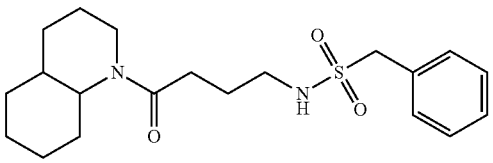
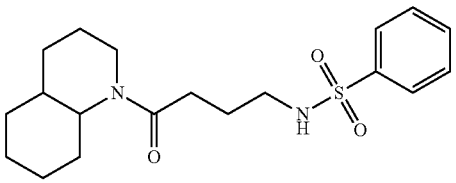
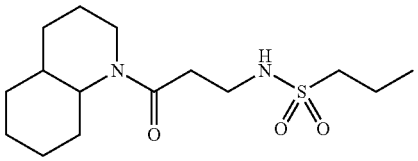
N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]propionamide

[0173]

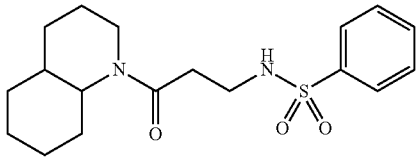
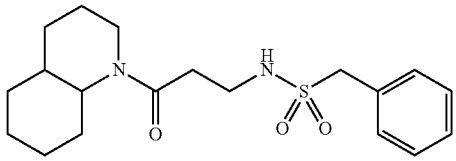
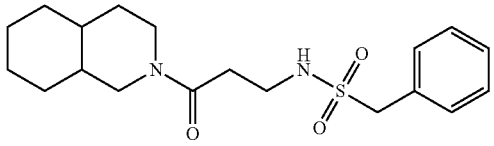


[0174] 23.4 mg (0.11 mmol) of the intermediate amine Vc.6 and 29 μL of Et_3N are added to a solution of 18.1 mg (0.10 mmol) of benzenesulfonyl chloride in 2 mL of AcOEt. The solution formed is kept under stirring at rt for 18 h. It is then treated with water and more AcOEt is added, the organic phase is separated and the aqueous phase is extracted once more with more AcOEt. The organic phases are pooled and successively washed with saturated $NaHCO_3$ solution, 1N HCl and brine. It is then dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. The residue is purified by means of silica gel column chromatography, using a (1:1) mixture of hexane-AcOEt as eluent, yielding 23 mg of a yellowish paste identified as example It.1. Method A: tr: 6.76 min/6.97 min; m/z: 351/351.

[0175] The following examples were prepared in a manner similar to example It.1:

Ex.	structure	name	method	rt (min)	m/z
It.2		N-[4-(octahydroquinolin-1-yl)-4-oxobutyl]-C-phenylmethanesulfonamide	A	6.84	379
It.3		N-[4-(octahydroquinolin-1-yl)-4-oxobutyl]bonzosulfonamide	A	6.79	365
It.4		Propane-1-sulfonic acid [3-(octahydroquinolin-1-yl)-3-oxopropyl]amide	A	6.26	317

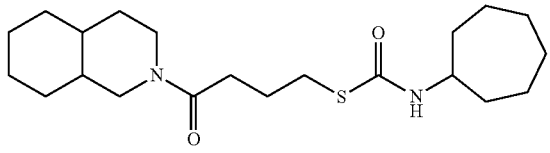
-continued

Ex.	structure	name	method	rt (min)	m/z
It.5		N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]benzesulfonamide	A	6.82	351
It.6		N-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-C-phenylmethanesulfonamide	A	6.85	365
It.7		N-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]-C-phenylmethanesulfonamide	A	6.84 7.01	365 365

Example Ip.1

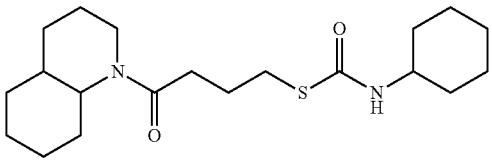
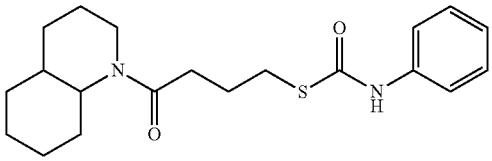
Cycloheptylthiocarbamic acid S-[4-(octahydroisoquinolin-2-yl)-4-oxobutyl]ester

[0176]

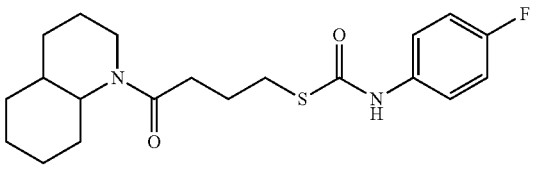
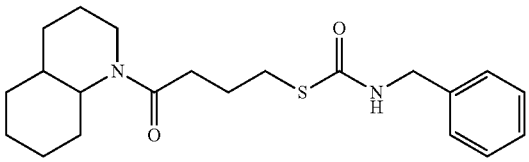
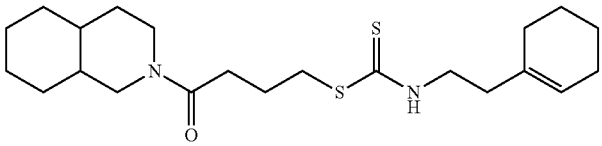
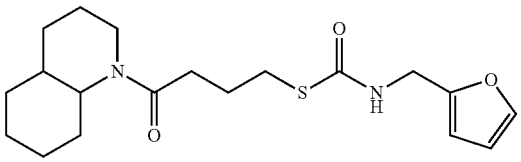
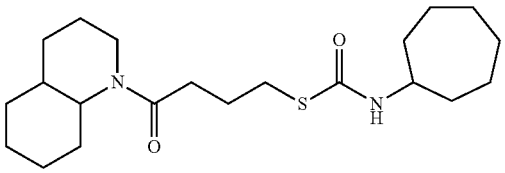
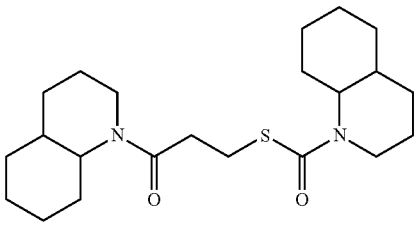
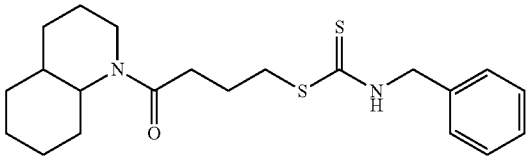


[0177] 12.8 μL of cycloheptylisocyanate (0.1 mmol) are added to a solution of 35 mg (0.15 mmol) of intermediate IVp.4 and 23 μL of Et_3N in 2 mL of THF at 0° C. Once added it is left to stir at reflux for 24 h. AcOEt is then added to the reaction mixture and it is sequentially washed with water and brine. The organic phase is dried over anhydrous Na_2S_4, it is filtered and the solvent is evaporated under reduced pressure. The residue is purified by means of silica gel column chromatography, using a (200:3) mixture of DCM-MeOH as eluent, yielding 23 mg of a colorless paste identified as example Ip.1. Method B: tr: 4.63 min; m/z: 381.

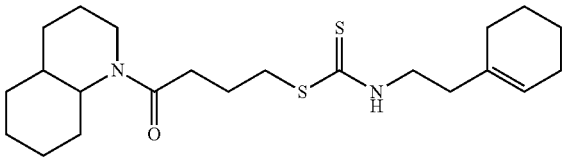
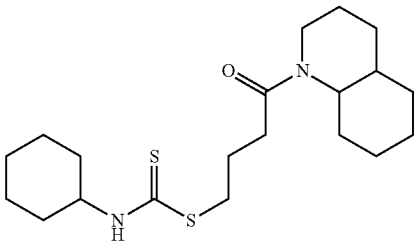
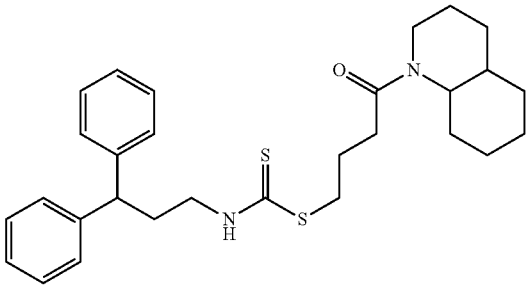
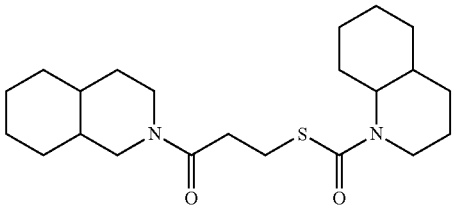
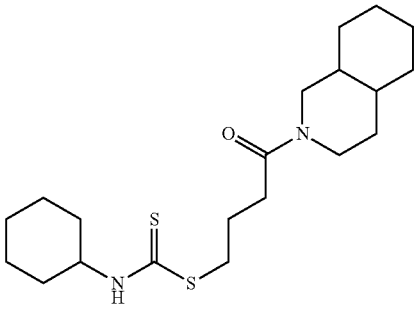
[0178] The following compounds were prepared in a manner similar to example Ip.1:

Ex.	structure	name	method	rt (min)	m/z
Ip.2		Cyclohexylthiocarbamic acid S-[4-(octahydroquinolin-1-yl)-4-oxobutyl]ester	B	4.34	367
Ip.3		Phenylthiocarbamic acid S-[4-(octahydroquinolin-1-yl)-4-oxobutyl]ester	B	4.14	361

-continued

Ex.	structure	name	method	rt (min)	m/z
Ip.4		(4-fluorophenyl)thiocarbamic acid S-[4-(octahydroquinolin-1-yl)-4-oxobutyl]ester	B	4.19	379
Ip.5		Benzylthiocarbamic acid S-[4-(octahydroquinolin-1-yl)-4-oxobutyl]ester	B	4.08	375
Ip.6		(2-cyclohex-1-enylethyl)dithiocarbamic acid 4-(octahydroisoquinolin-2-yl)-4-oxobutyl ester	B	5.19	409
Ip.7		(Furan-2-yl-methyl)carbamic acid S-[4-(octahydroquinolin-1-yl)-4-oxobutyl]ester	B	3.82	365
Ip.8		Cycloheptylthiocarbamic acid S-[4-(octahydroquinolin-1-yl)-4-oxobutyl]ester	B	4.54	381
Ip.9		Octahydroquinoline-1-thiocarbamic acid S-[3-(octahydroquinolin-1-yl)-3-oxopropyl]ester	B	4.94	393
Ip.10		Benzyldithiocarbamic acid 4-(octahydroquinolin-1-yl)-4-oxobutyl ester	B	4.54	391

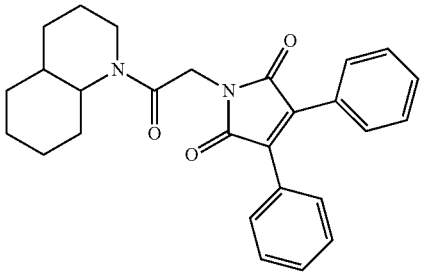
-continued

Ex.	structure	name	method	rt (min)	m/z
Ip.11		[2-(cyclohex-1-en-1-yl)ethyl]dithiocarbamic acid 4-(octahydroquinolin-1-yl)-4-oxobutyl ester	B	5.13	409
Ip.12		Cyclohexyldithiocarbamic acid 4-(octahydroquinolin-1-yl)-4-oxobutyl ester	B	4.82	383
Ip.13		Benzhydryldithiocarbamic acid 4-(octahydroquinolin-1 yl)-4-oxobutyl ester	B	4.98	467
Ip.14		Octahydroquinoline-1-thiocarbamic acid S-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl] ester	B	4.93 5.02	393 393
Ip.15		Cyclohexyldithiocarbamic acid 4-(octahydroisoquinolin-2-yl)-4-oxobutyl ester	B	4.83 4.89	383 383

Example Ik.1

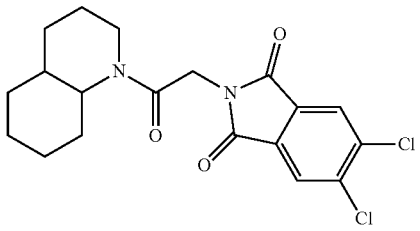
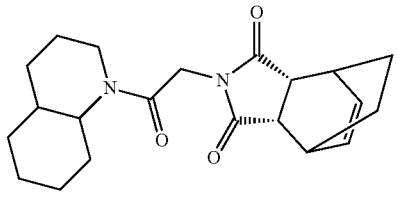
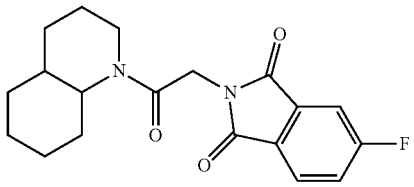
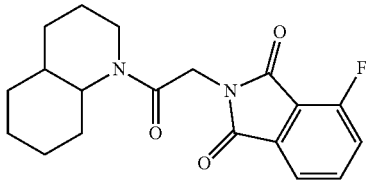
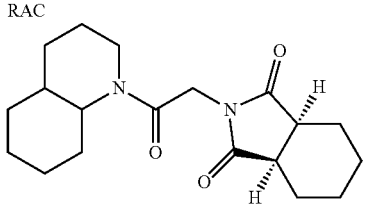
1-[2-(octahydroquinolin-1-yl)-2-oxoethyl]-3,4-diphenylpyrrole-2,5-dione

[0179]



[0180] 22 mg of the intermediate amine Vc.2 (0.11 mmol) are added to a solution of 34 mg (0.13 mmol) of 3,4-diphenylfuran-2,5-dione in 2 mL of DMF. Once added it is left to stir at 100° C. for 48 h. AcOEt is then added to the reaction mixture and it is sequentially washed with water, 5% $NaHCO_3$ solution and brine. The organic phase is dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. The residue is purified by means of silica gel column chromatography, using a (3:2) mixture of hexane-AcOEt as eluent, yielding 7.5 mg of a paste identified as example Ik.1. Method A: tr: 8.55 min; m/z: 429.

[0181] The following compounds were prepared in a manner similar to example Ik.1:

Ex.	structure	name	method	rt (min)	m/z
Ik.2		5,6-dichloro-2-[2-(octahydroquinolin-1-yl)-2-oxoethyl]isoindole-1,3-dione	A	8.12	395, 397, 399
Ik.3	RAC 	(2R,6S)-4-[2-(octahydroquinolin-1-yl)-2-oxoethyl]-4-azatricyclo[5.2.2.0$^{(2,6)}$]undec-8-ene-3,5-dione	A	6.85	357
Ik.4		5-fluoro-2-[2-(octahydroquinin-1-yl)-2-oxoethyl]isoindole-1,3-dione	A	7.16	345
Ik.5		4-fluoro-2-[2-(octahydroquinolin-1-yl)-2-oxoethyl]isoindole-1,3-dione	A	7.03	345
Ik.6	RAC 	(3aR,7aS)-2-[2-(octahydroquinolin-1-yl)-2-oxoethyl]hexahydroisoindole-1,3-dione	A	6.76	333

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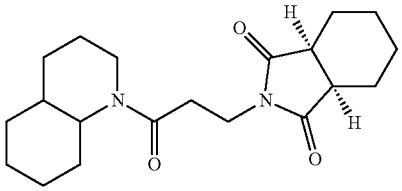
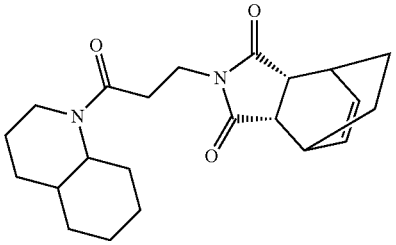
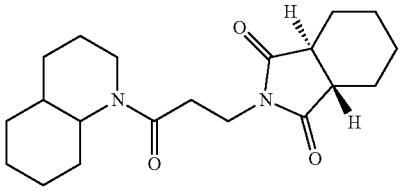
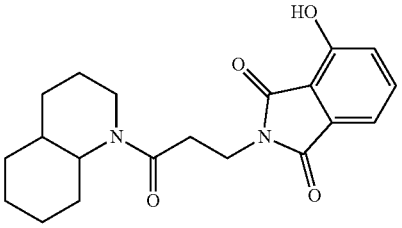
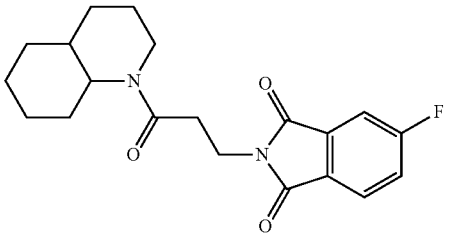
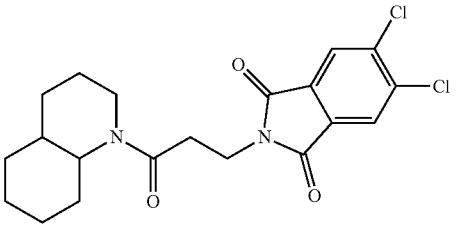
Ex.	structure	name	method	rt (min)	m/z
Ik.7	RAC 	(3aS,7aR)-2-[3-(octahydroquinolin-1-yl)-3-oxopropyl]hexahydro-isoindole-1,3-dione	A	6.89	347
Ik.8	RAC 	(2R,6S)-4-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-4-azatricyclo[5.2.2.0$^{(2,6)}$]undec-8-ene-3,5-dione	A	7.07	371
Ik.9	RAC 	(3aS,7aS)-2-[3-(octahydroquinolin-1-yl)-3-oxopropyl]hexahydro-isoindole-1,3-dione	A	6.89	347
Ik.10		4-hydroxy-2-[3-(octahydroquinolin-1-yl)-3-oxopropyl]isoindole-1,3-dione	A	6.28	357
Ik.11		5-fluoro-2-[3-(octahydroquinolin-1-yl)-3-oxopropyl]isoindole-1,3-dione	A	7.25	359
Ik.12		5,6-dichloro-2-[3-(octahydroquinolin-1-yl)-3-oxopropyl]isoindole-1,3-dione	A	8.27	409, 411, 413

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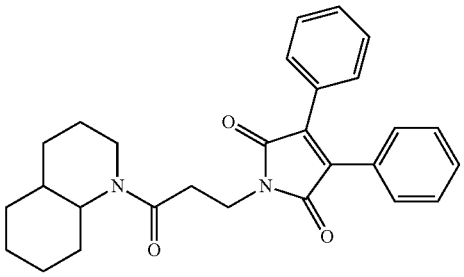
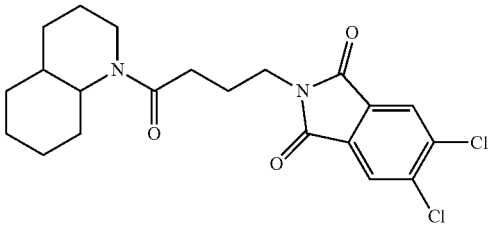
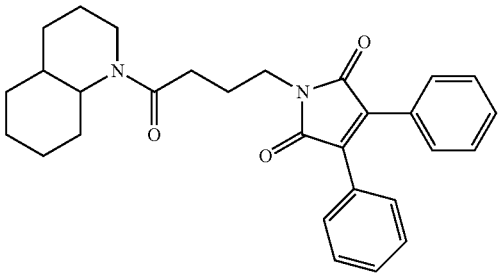
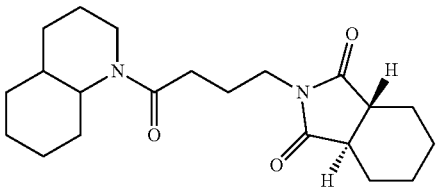
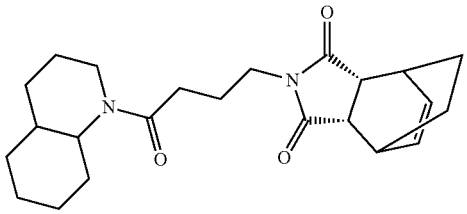
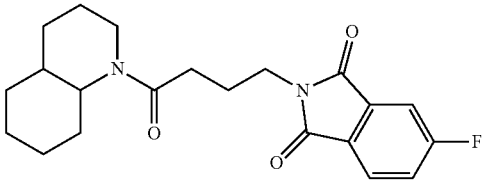
Ex.	structure	name	method	rt (min)	m/z
Ik.13		1-[3-(octahydroquinolin-1-yl)-3-oxopropyl]-3,4-diphenylpyrrole-2,5-dione	A	8.62	443
Ik.14		5,6-dichloro-2-[4-octahydroquinolin-1-yl)-4-oxobutyl]isoindole-1,3-dione	A	8.43	423, 425, 427
Ik.15		1-[4-(octahydroquinolin-1-yl)-4-oxobutyl]-3,4-diphenylpyrrole-2,5-dione	A	8.75	457
Ik.16	RAC 	(3aR,7aR)-2-[4-(octahydroquinolin-1-yl)-4-oxobutyl]hexahydroisoindole-1,3-dione	A	7.04	361
Ik.17	RAC 	(2R,6S)-4-[4-(octahydroquinolin-1-yl)-4-oxobutyl]-4-azatricyclo[5.2.2.0$^{(2,6)}$]undec-8-ene-3,5-dione	A	7.14	385
Ik.18		5-fluoro-2-[4-(octahydroquinolin-1-yl)-4-oxobutyl]isoindole-1,3-dione	A	7.38	373

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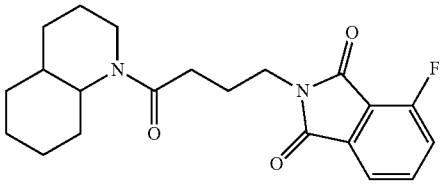
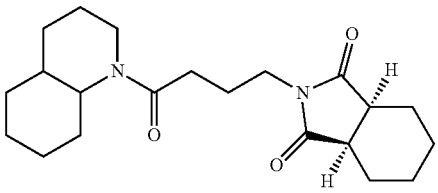
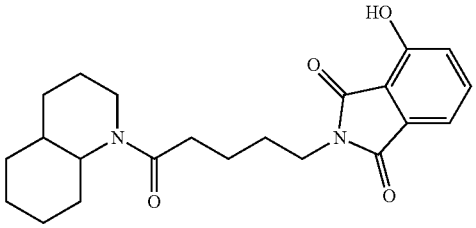
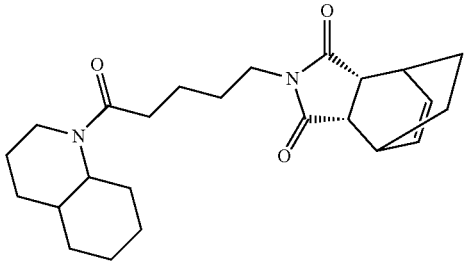
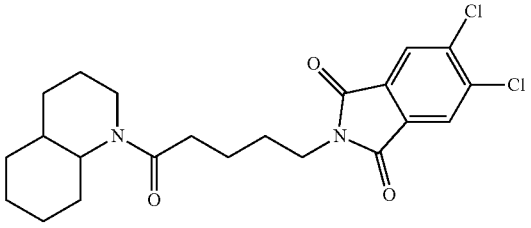
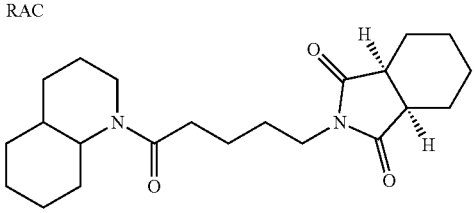
Ex.	structure	name	method	rt (min)	m/z
Ik.19		4-fluoro-2-[4-(octahydroquinolin-1-yl)-4-oxobutyl]isoindole-1,3-dione	A	7.19	373
Ik.20	RAC 	(3aR,7aS)-2-[4-(octahydroquinolin-1-yl)-4-oxobutyl]hexahydroisoindole-1,3-dione	A	7.13	361
Ik.21		4-hydroxy-2-[5-(octahydroquinolin-1-yl)-5-oxopentyl]isoindole-1,3-dione	A	6.64 6.69	385 385
Ik.22	RAC 	(2R,6S)-4-[5-(octahydroquinolin-1-yl)-5-oxopentyl]-4-azatricyclo[5.2.2.0$^{(2,6)}$]undec-8-ene-3,5-dione	A	7.32	399
Ik.23		5,6-dichloro-2-[5-(octahydroquinolin-1-yl)-5-oxopentyl]isoindole-1,3-dione	A	8.62	437, 439, 441
Ik.24	RAC 	(3aS,7aR)-2-[5-(octahydroquinolin-1-yl)-5-oxopentyl]hexahydroisoindole-1,3-dione	A	7.22	375

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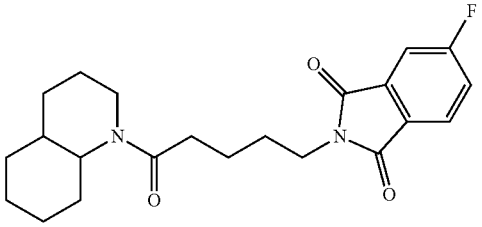
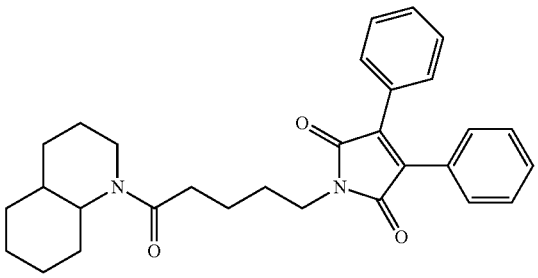
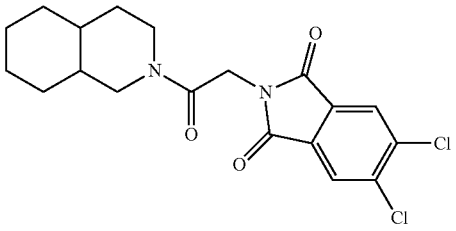
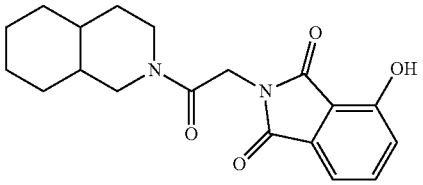
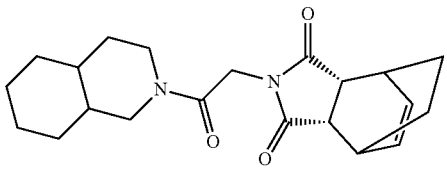
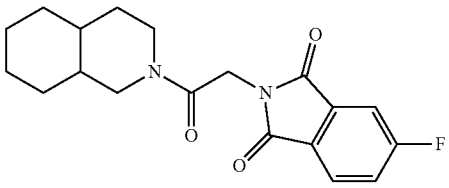
Ex.	structure	name	method	rt (min)	m/z
Ik.25		5-fluoro-2-[5-(octahydroquinolin-1-yl)-5-oxopentyl]isoindole-1,3-dione	A	7.61	387
Ik.26		1-[5-(octahydroquinolin-1-yl)-5-oxopentyl]-3,4-diphenylpyrrole-2,5-dione	A	8.90	471
Ik.27		5,6-dichloro-2-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]isoindole-1,3-dione	A	8.24	395, 397, 399
Ik.28		4-hydroxy-2-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]isoindole-1,3-dione	A	6.45	343
Ik.29	RAC 	(2R,6S)-4-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]-4-azatricyclo[5.2.2.0$^{(2,6)}$]undec-8-ene-3,5-dione	A	7.02	357
Ik.30		5-fluoro-2-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]isoindole-1,3-dione	A	7.14 7.31	345 345

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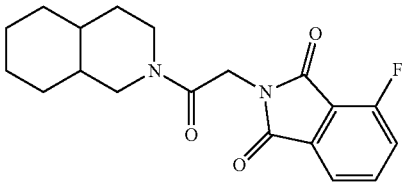
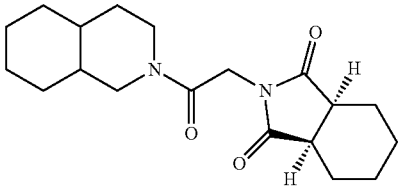
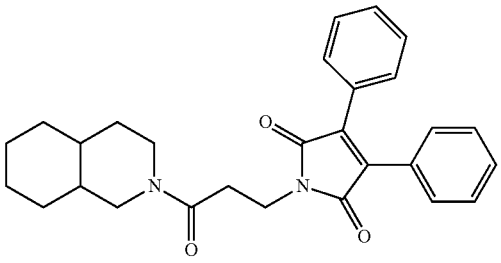
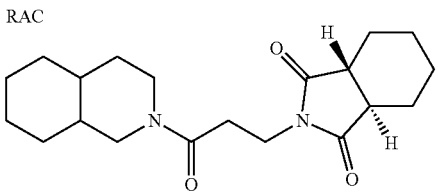
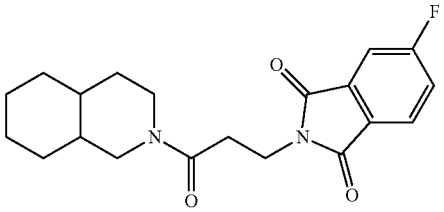
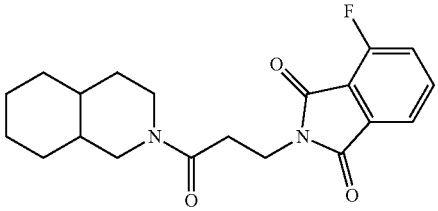
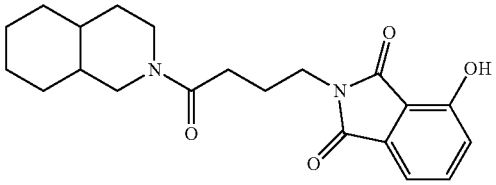
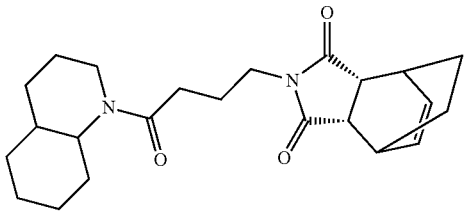
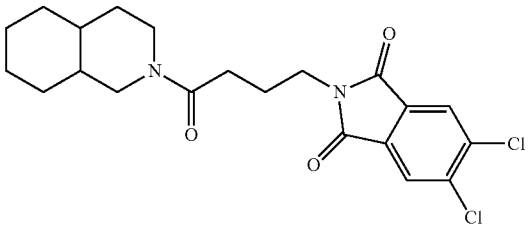
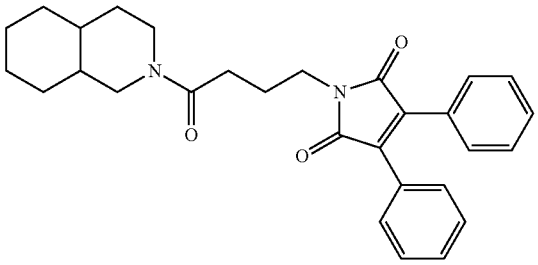
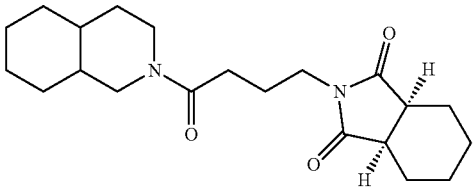
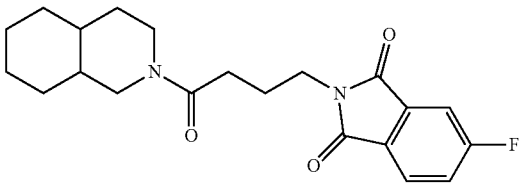
Ex.	structure	name	method	rt (min)	m/z
Ik.31		4-fluoro-2-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]isoindole-1,3-dione	A	7.19	345
Ik.32	RAC 	(3aR,7aS)-2-[2-(octahydroisoquinolin-2-yl)-2-oxoethyl]hexahydroisoindole-1,3-dione	A	6.76 6.94	333 333
Ik.33		1-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]-3,4-diphenylpyrrole-2,5-dione	A	8.61 8.73	443 443
Ik.34	RAC 	(3aR,7aR)-2-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]hexahydroisoindole-1,3-dione	A	6.89 7.07	347 347
Ik.35		5-fluoro-2-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]isoindole-1,3-dione	A	7.36	359
Ik.36		4-fluoro-2-[3-(octahydroisoquinolin-2-yl)-3-oxopropyl]isoindole-1,3-dione	A	7.01 7.18	359 359

-continued

Ex.	structure	name	method	rt (min)	m/z
Ik.37		4-hydroxy-2-[4-(octahydroisoquinolin-2-yl)-4-oxobutyl]isoindole-1,3-dione	A	6.45 6.61	371 371
Ik.38	RAC 	(2R,6S)-4-[4-(octahydroisoquinolin-2-yl)-4-oxobutyl]-4-azatricyclo[5.2.2.0$^{(2,6)}$]undec-8-ene-3,5-dione	A	7.14 7.32	385 385
Ik.39		5,6-dichloro-2-[4-(octahydroisoquinolin-2-yl)-4-oxohutyl]isoindole-1,3-dione	A	8.43 8.56	423, 425, 427 423, 425, 427
Ik.40		1-[4-(octahydroisoquinolin-2-yl)-4-oxobutyl]-3,4-diphenylpyrrole-2,5-dione	A	8.77 8.89	457 457
Ik.41	RAC 	(3aS,7aR)-2-[4-(octahydroisoquinolin-2-yl)-4-oxotbutyl]hexahydroiso-indole-1,3-dione	A	7.04 7.22	361 361
Ik.42		5-fluoro-2-[4-(octahydroisoquinolin-2-yl)-4-oxobutyl]isoindole-1,3-dione	A	7.39 7.55	373 373

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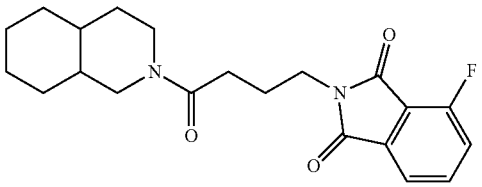
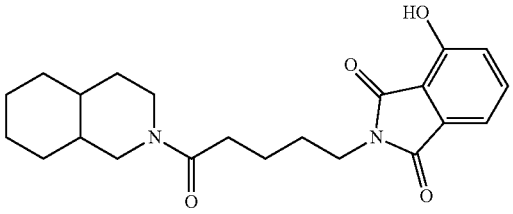
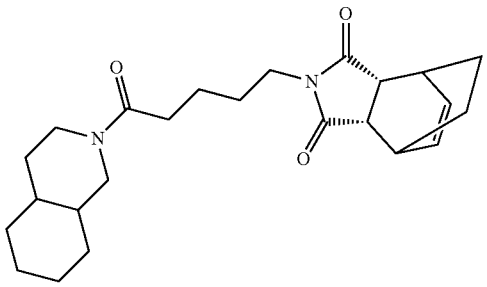
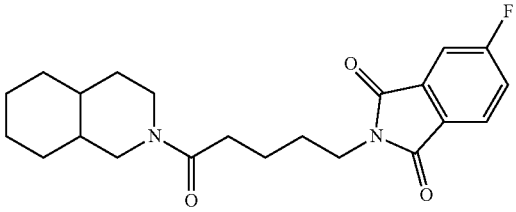
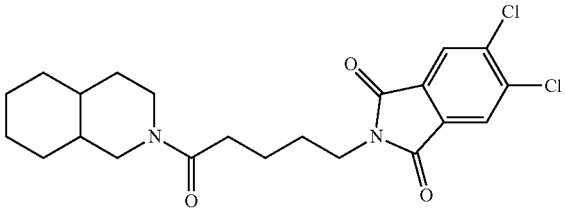
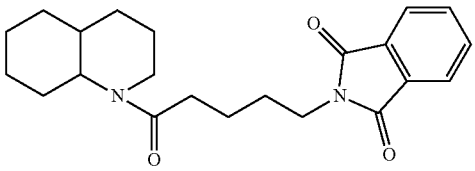
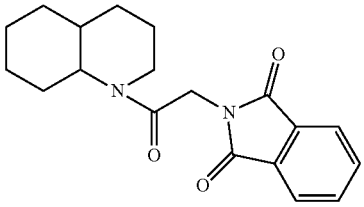
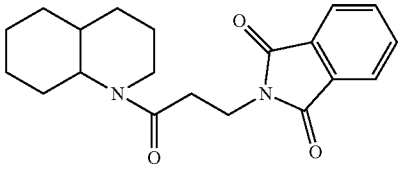
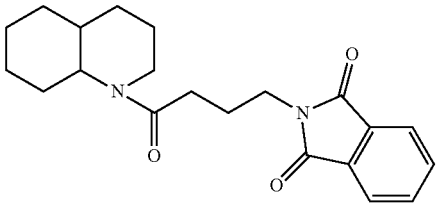
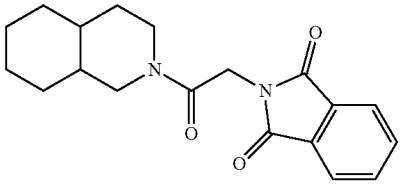
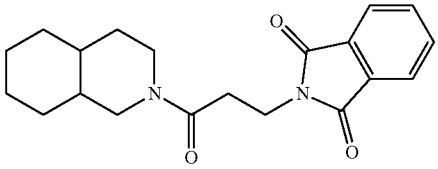
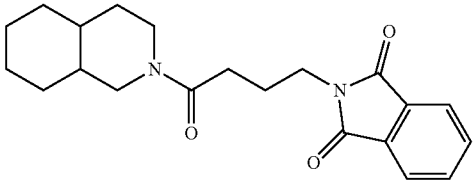
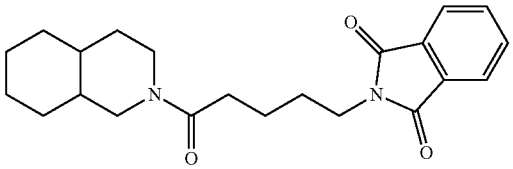
Ex.	structure	name	method	rt (min)	m/z
Ik.43		4-fluoro-2-[4-(octahydroisoquinolin-2-yl)-4-oxobutyl]isoindole-1,3-dione	A	7.20 7.36	373 373
Ik.44		4-hydroxy-2-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]isoindole-1,3-dione	A	6.64 6.80	385 385
Ik.45	RAC 	(2R,6S)-4-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]-4-azatricyclo[5.2.2.0$^{(2,6)}$]undec-8-ene-3,5-dione	A	7.50	399
Ik.46		5-fluoro-2-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]isoindole-1,3-dione	A	7.59 7.75	387 387
Ik.47		5,6-dichloro-2-[5-(octahydroisoquinolin-2-yl)-5-oxopentyl]isoindole-1,3-dione	A	8.60 8.72	437, 439, 441 437, 439, 441
If.1		2-(5-(octahydroquinolin-1-yl)-5-oxopentyl)isoindoline-1,3-dione	A	7.39	369

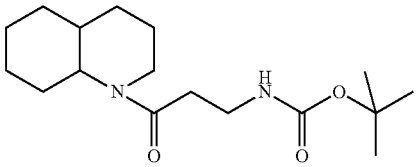
-continued

Ex.	structure	name	method	rt (min)	m/z
If.2		2-(2-(octahydroquinolin-1-yl)-2-oxoethyl)isoindoline-1,3-dione	A	6.94	327
If.3		2-(3-(octahydroquinolin-1-yl)-3-oxopropyl)isoindoline-1,3-dione	A	7.01	341
If.4		2-(4-(octahydroquinolin-1-yl)-4-oxobutyl)isoindoline-1,3-dione	A	7.18	355
If.5		2-(2-(octahydroisoquinolin-2-yl)-2-oxoethyl) isoindoline-1,3-dione	A	6.92 7.11	327 327
If.6		2-(3-(octahydroisoquinolin-2-yl)-3-oxopropyl) isoindoline-1,3-dione	A	6.96 7.13	341 341
If.7		2-(4-(octahydroisoquinolin-2-yl)-4-oxobutyl) isoindoline-1,3-dione	A	7.17 7.33	355 355
If.8		2-(5-(octahydroisoquinolin-2-yl)-5-oxopentyl) isoindoline-1,3-dione	A	7.39 7.55	369 369

Example Ij.1

[3-(octahydroquinolin-1-yl)-3-oxopropyl]carbamic acid t-butylester

[0182]



[0183] 0.12 mg (1 μmol) of 4-dimethylaminopyridine and 43 mg (0.2 mmol) of di-tertbutyl dicarbonate are added to a solution of 21 mq (0.1 mmol) of intermediate Vc.3 in 2 mL of DCM. The mixture is stirred for 18 h at rt. The solvent is evaporated and water and AcOEt are then added, the aqueous phase is separated and extracted again with AcOEt, the pooled organic phases are sequentially washed with 1N HCl and brine. The organic phase is dried over anhydrous Na_2SO_4, it is filtered and the solvent is evaporated under reduced pressure. The residue is purified by means of silica gel column chromatography, using a (20:1) mixture of DCM-MeOH as eluent, yielding 6.5 mg of a colorless paste identified as example Ij.1. Method A: tr: 7.15 min; m/z: 311.

Pharmacological Examples

Determination of the Inhibitory Activity of 11-Beta-HSD1 in The Microsomal Fraction of Transfected HEK293 Cells

[0184] 99 μL of a reaction mixture which contained 80 nM [^{3}H]-cortisone, 1 mM NADPH and 40 μg/mL of the microsome preparations of HEK-293 cells stably transfected with the 11-beta-HSD1 clone, dissolved in assay buffer (50 mM HEPES, 100 mM KCl, 5 mM NaCl, 2 mM $MgCl_2$/100 mL H_2O) were added to a 96-well plate. The inhibitors to be analyzed were dissolved in 100% DMSO, and the final concentration in the reaction was 1%.

[0185] The reaction mixture was incubated for 2 hours at 37° C. under stirring. The total reaction volume per well was 100 μL. The stop solution (5 mg/mL ProteinA Spa bead, Superblock Blocking Buffer, 30 μM glycyrrhetinic acid, 1 μg/mL anticortisol monoclonal antibody) was simultaneously prepared, being incubated under stirring for two hours at rt and protected from light. Once the two hours lapsed, 50 μL of the stop solution were dispensed to each well in the reaction plate and was left to incubate for two hours at rt under stirring and protected from light. Once the second incubation ended, the plate was read in a 1450 Microbeta Tritalux (Wallac®) scintillation counter for 30 seconds per well.

[0186] Determination of the Inhibitory Activity of 11-Beta-HSD2 In the Microsomal Fraction of Transfected HEK293 Cells

[0187] 59 μL of the reaction mixture were added to a 96-well plate. This contained 1 mM NAD^+ and 80 μg/mL of the microsome preparations of HEK-293 cells stably transfected with the 11-beta-HSD2 clone, dissolved in assay buffer (50 mM HEPES, 100 mM KCl, 5 mM NaCl, 2 mM $MgCl_2$/100 mL H_2O). The inhibitors to be analyzed were dissolved in 100% DMSO, and the final concentration in the reaction was 1%. The reaction mixture was pre-incubated for 30 minutes at 37° C. under stirring. 40 μL of the reaction substrate, 3.2 nM [^{3}H]-cortisol dissolved in assay buffer were added. It was incubated at 37° C. for two hours under stirring. The stop solution was simultaneously prepared, being incubated for two hours at rt and protected from light. After this time 50 μL of the stop solution were added to end the reaction, and it was incubated for two hours at rt under stirring and protected from light. The signal emitted by SPA beads-cortisol complex was measured in the 1450 Microbeta Tritalux (Wallac®) scintillation counter for 30 seconds per well. This case measures the binding to the substrate, and not to the product as in the assay for the type 1 isoenzyme.

[0188] The following table indicates the activity values of some compounds described in the examples expressed as percentage of inhibition at a 10 μM concentration.

	% Inhibition (10 μM)	
Example	11-beta-HSD1	11-beta-HSD2
Ia.59	100	0
Ia.61	93	0
Ia.66	100	6
Ib.8	100	5
Ib.18	100	10
Ib.80	100	13
Ie.16	100	5
Ib.8	100	0
Id.6	100	0
Id.12	100	0
If.1	100	0
Ik.25	100	6
If.8	100	4

In Vivo Efficacy of 11beta-HSD1 Inhibitors

[0189] In vivo efficacy of 11beta-HSD1 inhibitors was evaluated in an acute glaucoma model based on a glucose-induced ocular hypertension model in rabbit.

[0190] Male Chinchilla Bastard rabbits (1.5-2.5 kg) were obtained from Charles River (Germany) and were kept under controlled environmental conditions (12-hour light/12-hour dark cycle) with food and drinking water ad libitum. Animals were treated in accordance with the ARVO Statement for the Use of Animals in Ophthalmic and Vision Research.

[0191] In order to measure intraocular pressure (IOP) according to the glucose-induced ocular hypertension model, rabbits were restrained and basal IOP was determined using a Tonolab tonometer (Tiolat Oy) at least 1 hour after initial immobilization. Acute ocular hypertension was induced by a bolus of 5% glucose solution at 20 ml/kg into the marginal ear vein and IOP was determined 5, 10, 20, 30 and 40 minutes later. Eye treatment instillation was performed 15 minutes before the hypertension induction and both eyes of each animal were instilled with the same solution, either the vehicle, the 11beta-HSD1 inhibitor suspension at 0.5% or Timolol maleate solution at 0.5% as a positive control.

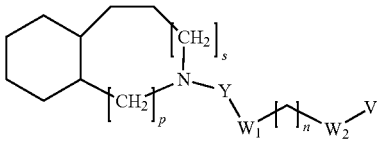
[0192] The results obtained showed that the inhibitors markedly reduced ocular hypertension (% inhibition). The intraocular pressure (IOP) increment time-course was statistically lower after 11beta-HSD1 inhibitors treatment compared to vehicle. On the other hand, it was noteworthy that 11beta-HSD1 inhibitors efficacy regulating IOP was compa-

rable to Timolol (expressed as Ratio inhibitor/Timolol), suggesting that 11beta-HSD1 inhibitors can be as effective as other glaucoma treatments.

[0193] The following table indicates the inhibition percentage of several compounds described in the examples as well as their efficacy in comparison with the standard.

	Glucose-induced Hypertension	
Example	% Inhibition	Ratio Inhibitor/Timolol
If.1	51	0.90
Id.12	50	0.87
Ia.96	46	0.84
Ia.198	41	0.65
Ia.224	16	0.25

1. A method of treatment of an individual who is suffering or is susceptible to suffering a disease caused by 11-beta-HSD1-associated disorders which comprises administering to said individual a pharmaceutical composition comprising one or more pharmaceutically acceptable excipient and a therapeutically effective amount of a compound of formula (I)



(I)

or pharmaceutically acceptable salts thereof, wherein:

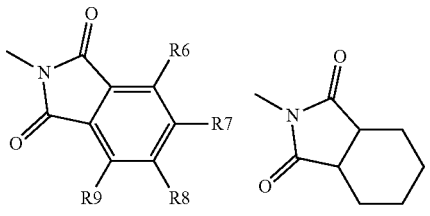
s and p are an integer selected in an opposite manner between 0 and 1, such that when s is 1, p is 0 (to form a perhydroquinoline) and when s is 0, p is 1 (to form a perhydroisoquinoline),

Y is a biradical selected from the group consisting of CO, CS and SO_2,

W1 and W2 is each independently a bond or a biradical selected from O, S and NR1, wherein R1 is selected from H, C_{1-4} alkyl and C_{3-10} cycloalkyl,

n is an integer selected from 0, 1, 2, 3 and 4,

V is a radical selected from the groups consisting of —CO-T, —CS-T and —SO_2-T, or a radical selected from the group consisting of:



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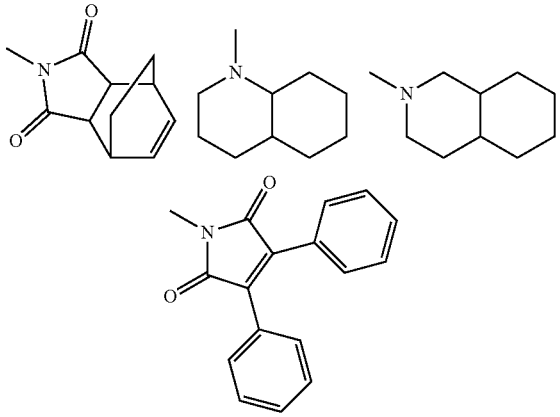
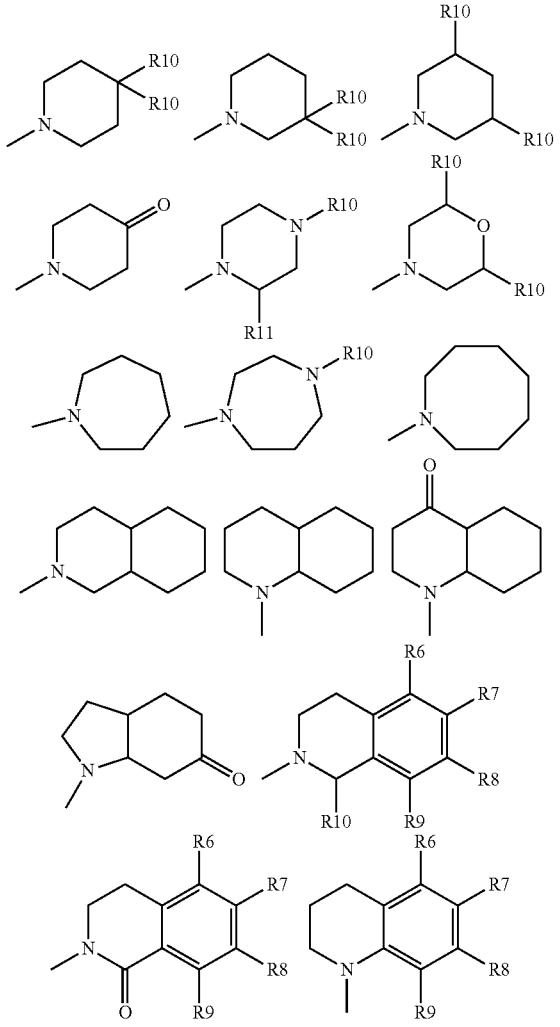


where T is a group selected from the groups consisting of NR2R3, R2, OR2 and SR2; or a group selected from the group consisting of



wherein R2 and R3 is each independently H, COR4, SO_2R4, C_{1-4} alkyl, aryl, benzyl, phenethyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{3-10} cycloalkyl or heterocycle

wherein when R2 or R3 is an alkyl, or an alkenyl these can be optionally substituted with one or several substituents independently selected from the groups consisting of F, OR4, NR4R5, COOR4, CONR4R5, C_{3-10} cycloalkyl, aryl and heterocycle;

wherein when R2 or R3 is an aryl, a benzyl, a phenethyl, a cycloalkyl or a heterocycle, these can be optionally substituted with one or several substituents independently selected from the group consisting of NH_2, F, Cl, CN, NO_2, COOH, R4, COOR4, OR4, OCF_3, SH, SR4, CONR4R5, SO_2NR4R5, COR4, NR1COR4, OCOR4, SOR4, SO_2R4 and heterocycle; and

wherein when R2 or R3 is a cycloalkyl, this can be optionally substituted with one or several fused benzene rings, the benzene could be optionally substituted with one or several substituents independently selected from the group consisting of alkyl, alkoxide and halogen,

wherein R4 and R5 is each independently selected from the group consisting of H, C_{1-4} alkyl, aryl, benzyl, phenethyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{3-10} cycloalkyl and heterocycle, or optionally R4 and R5 can be bound to one another forming a 3 to 8 membered cycle,

wherein R6, R7, R8 and R9 is each independently selected from the groups consisting of H, OR4, F and Cl, and

wherein R10 is independently selected from the group consisting of H, OH, F, C_{1-4} alkyl, COOR11, COR11, phenyl, benzyl, benzhydryl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{3-10} cycloalkyl and heterocycle, and wherein the alkyl, phenyl, benzyl, benzhydryl, cycloalkyl or heterocycle can be optionally substituted with one or several substituents independently selected from the group consisting of NH_2, F, Cl, NO_2, COOH, COOR4, OR4, CF, SH, SR4, CONR4R5, SO_2NR4R5, COR4, NR1COR4, OCOR4, SOR4, SO_2R4 and C_{1-4} alkyl where R11 is selected from the group consisting of H, C_{1-4} alkyl and C_{3-10} cycloalkyl,

so as to thereby treat the individual.

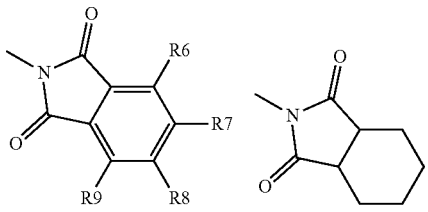
2. The method according to claim **1**, wherein in the compound Y is CO or SO_2.

3. The method according to claim **1**, wherein in the compound W1 and W2 is each independently selected from the group consisting of a bond, S and NR1.

4. The method according to claim **3**, wherein in the compound R1 is H

5. The method according to claim **1**, wherein in the compound V is —CO-T, —CS-T or —SO_2-T.

6. The method according to claim **1**, wherein in the compound V is selected from the group consisting of:



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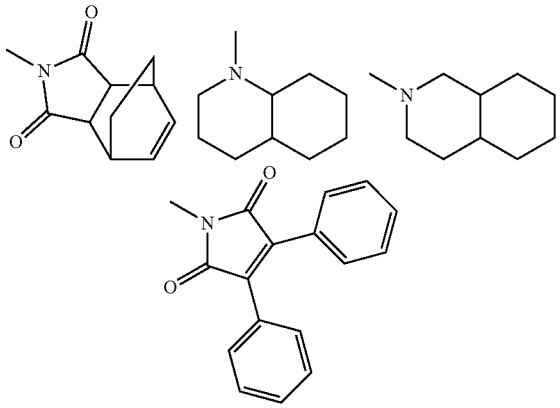


7. The method according to claim **1**, wherein in the compound T is NR2R3, R2, OR2 or SR2.

8. The method according to claim **1**, wherein in the compound R2 and R3 is each independently selected from the group consisting of H, COR4, SO_2R4, C_{1-4} alkyl, phenyl, naphthyl, benzyl, phenethyl, C_{2-4} alkenyl, C_{3-10} cycloalkyl, and heterocycle, particularly, 2-furanyl, 2-thiophenyl, 2-(1-methylindole), quinoline, isoquinoline and 2-benzofuranyl.

9. The method according to claim **8**, wherein in the compound R2 and R3 is each independently selected from the groups consisting of C_{1-4} alkyl and C_{2-4} alkenyl.

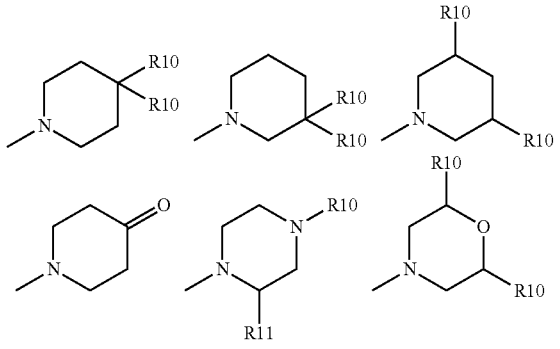
10. The method according to claim **9**, wherein in the compound R2 or R3 are optionally substituted with one or several substituents independently selected from the groups consisting of F, OR4, NR4R5, COOR4, CONR4R5, phenyl, C_{3-10} cycloalkyl, hexenyl, naphthyl and heterocycle, particularly pyridine, 3-(1-methylindole), 3-thiophenyl and 2-furanyl.

11. The method according to claim **8**, wherein in the compound R2 and R3 is each independently selected from the group consisting of phenyl, benzyl, phenethyl and C_{3-10} cycloalkyl.

12. The method according to claim **11**, wherein in the compound R2 or R3 are optionally substituted with one or several substituents independently selected from the group consisting of F, Cl and OR4.

13. The method according to claim **1**, wherein in the compound R4 and R5 is each independently selected from the groups consisting of C_{1-4} alkyl, benzyl, phenethyl and phenyl.

14. The method according to claim **1**, wherein in the compound T is selected from the group consisting of:



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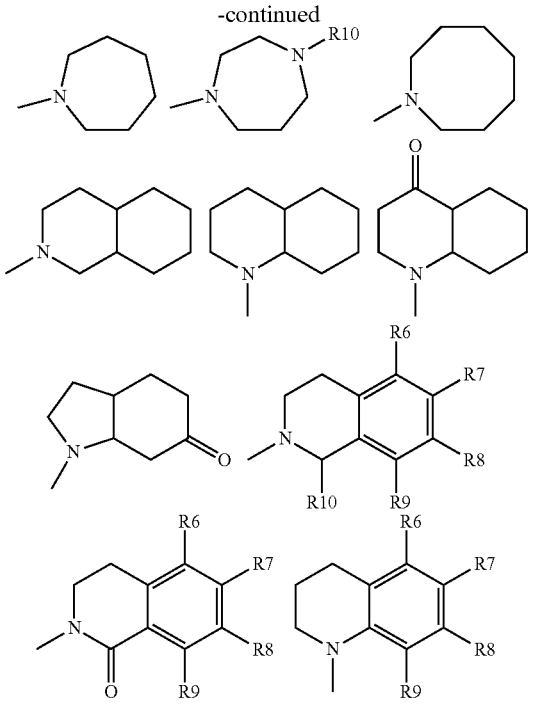

15. The method according to claim **1**, wherein in the compound R6, R7, R8 and R9 is each independently selected from the group consisting of H, OR4, F and Cl, and wherein R10 is independently selected from H, OH, F, C_{1-4} alkyl, COOR11, COR11, phenyl, benzyl and benzhydryl.

16. The method according to claim **15**, wherein in the compound R10 is phenyl, benzyl or benzhydryl.

17. The method according to claim **16**, wherein in the compound R10 is optionally substituted with one or several substituents, independently selected from the group consisting of F, OR4, CF3, COR4 and C_{1-4} alkyl.

18. The method according to claim **1**, wherein in the compound R11 is H or C_{3-10} cycloalkyl.

19. The method according to claim **1**, wherein in the compound s is 0 and p is 1.

20. The method according to claim **1**, wherein s is 1 and p is 0.

21. The method according to claim **1**, wherein the disease caused by 11-beta-HSD1-associated disorders is glaucoma, elevated ocular pressure, metabolic disorders, obesity, metabolic syndrome, dyslipidemia, hypertension, diabetes, atherosclerosis, Cushing's syndrome, psoriasis, rheumatoid arthritis, cognitive disorders, Alzheimer's disease or neurodegeneration.

22. The method according to claim **1**, wherein the disease caused by 11-beta-HSD1-associated disorders is glaucoma or metabolic syndrome.

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