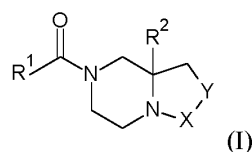


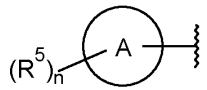
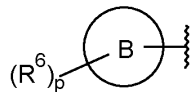
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CLAIMS

1 A compound of formula (I)

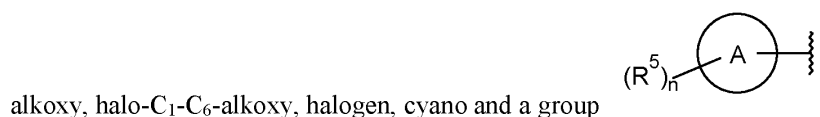


or a pharmaceutically acceptable salt thereof, wherein:

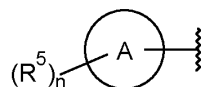
- 5 X is $-\text{CH}_2\text{CR}^3\text{R}^4-$ or $-\text{CH}=\text{CR}^3-$;
 Y is $-\text{O}-$, $-\text{S}-$, $-\text{SO}-$, or $-\text{SO}_2-$;
 R^1 is C_6 - C_{14} -aryl or 5- to 14-membered heteroaryl, wherein said C_6 - C_{14} -aryl or 5- to 14-membered heteroaryl are optionally substituted with 1-5 substituents independently selected from C_1 - C_6 -alkyl, C_1 - C_6 -alkoxy, halo- C_1 - C_6 -alkyl,
- 10 halo- C_1 - C_6 -alkoxy, halogen, cyano, hydroxy, and a group  ;
 R^2 is hydrogen or C_1 - C_6 -alkyl;
- R^3 is a group  ;
 R^4 is hydrogen, halogen or hydroxy;
 R^5 is hydrogen, halogen, hydroxy, amino, halo- C_1 - C_6 -alkyl, C_1 - C_6 -alkoxy, C_1 - C_6 -alkyl, halo- C_1 - C_6 -alkoxy, cyano or oxo;
- 15 R^6 is hydrogen, halogen, C_6 - C_{14} -aryl, 5- to 14-membered heteroaryl, halo- C_1 - C_6 -alkoxy, halo- C_1 - C_6 -alkyl, oxo, C_1 - C_6 -alkyl, C_1 - C_6 -alkoxy, C_2 - C_6 -alkenyl, C_2 - C_6 -alkynyl, C_3 - C_{14} -cycloalkyl or cyano; wherein said 5- to 14-membered heteroaryl or C_6 - C_{14} -aryl is optionally substituted with 1-2 substituents
- 20 selected from halogen, cyano, amino, hydroxy, C_1 - C_6 -alkyl and C_1 - C_6 -alkoxy;
 A and B are each independently 5- to 14-membered heteroaryl, C_6 - C_{14} -aryl, 3- to 14-membered heterocyclyl or C_3 - C_{14} -cycloalkyl; and
 n and p are each independently 1, 2, 3, 4 or 5.
- 2 The compound of formula (I) according to claim 1, or a pharmaceutically acceptable
- 25 salt thereof, wherein X is $-\text{CH}_2\text{CR}^3\text{R}^4-$.

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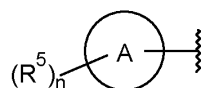
- 3 The compound of formula (I) according to claim 1 or 2, or a pharmaceutically acceptable salt thereof, wherein Y is $-\text{O}-$ or $-\text{S}-$.
- 4 The compound of formula (I) according to claim 1 or 2, or a pharmaceutically acceptable salt thereof, wherein Y is $-\text{O}-$.
- 5 5 The compound of formula (I) according to any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein R^1 is $\text{C}_6\text{-C}_{14}$ -aryl or 5- to 14-membered heteroaryl, wherein said $\text{C}_6\text{-C}_{14}$ -aryl or 5- to 14-membered heteroaryl are substituted with 1-3 substituents independently selected from $\text{C}_1\text{-C}_6$ -



- 10 6 The compound of formula (I) according to any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein R^1 is $\text{C}_6\text{-C}_{14}$ -aryl substituted with 2-3 substituents independently selected from $\text{C}_1\text{-C}_6$ -alkoxy, halogen, and a group



- 7 The compound of formula (I) according to any one of claims 1 to 4, or a pharmaceutically acceptable salt thereof, wherein R^1 is phenyl substituted with 2-3 substituents independently selected from chloro, fluoro, methoxy and a group
- 15



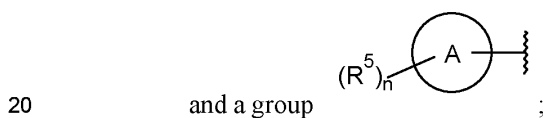
- 8 The compound of formula (I) according to any one of claims 1 to 7, or a pharmaceutically acceptable salt thereof, wherein R^2 is hydrogen.
- 20 9 The compound of formula (I) according to any one of claims 1 to 8, or a pharmaceutically acceptable salt thereof, wherein R^4 is hydrogen or hydroxy.
- 10 The compound of formula (I) according to any one of claims 1 to 9, or a pharmaceutically acceptable salt thereof, wherein R^5 is hydrogen, halogen, hydroxy, halo- $\text{C}_1\text{-C}_6$ -alkyl, $\text{C}_1\text{-C}_6$ -alkoxy, $\text{C}_1\text{-C}_6$ -alkyl, cyano or oxo.

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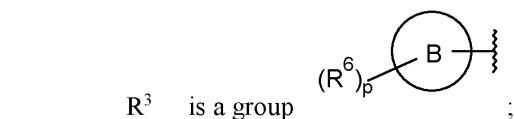
- 11 The compound of formula (I) according to any one of claims 1 to 9, or a
pharmaceutically acceptable salt thereof, wherein R⁵ is hydrogen, halogen, cyano,
hydroxy or oxo.
- 12 The compound of formula (I) according to any one of claims 1 to 9, or a
5 pharmaceutically acceptable salt thereof, wherein R⁵ is hydrogen, fluoro, cyano,
hydroxy or oxo.
- 13 The compound of formula (I) according to any one of claims 1 to 12, or a
pharmaceutically acceptable salt thereof, wherein R⁶ is hydrogen, halogen, C₆-C₁₄-
aryl, 5- to 14-membered heteroaryl, halo-C₁-C₆-alkoxy, halo-C₁-C₆-alkyl, oxo, C₁-
10 C₆-alkyl, C₁-C₆-alkoxy, C₂-C₆-alkenyl, C₂-C₆-alkynyl, C₃-C₁₄-cycloalkyl or cyano,
wherein said 5- to 14-membered heteroaryl or C₆-C₁₄-aryl is optionally substituted
with a substituent selected from halogen, C₁-C₆-alkyl and C₁-C₆-alkoxy.
- 14 The compound of formula (I) according to any one of claims 1 to 12, or a
pharmaceutically acceptable salt thereof, wherein R⁶ is halogen, oxo, halo-C₁-C₆-
15 alkyl or halo-C₁-C₆-alkoxy.
- 15 The compound of formula (I) according to any one of claims 1 to 12, or a
pharmaceutically acceptable salt thereof, wherein R⁶ is chloro, fluoro, oxo,
trifluoromethyl or difluoromethoxy.
- 16 The compound of formula (I) according to any one of claims 1 to 15, or a
20 pharmaceutically acceptable salt thereof, wherein n is 1 or 2.
- 17 The compound of formula (I) according to any one of claims 1 to 16, or a
pharmaceutically acceptable salt thereof, wherein p is 1, 2 or 3.
- 18 The compound of formula (I) according to any one of claims 1 to 16, or a
pharmaceutically acceptable salt thereof, wherein p is 1 or 2.
- 25 19 The compound of formula (I) according to any one of claims 1 to 18, or a
pharmaceutically acceptable salt thereof, wherein A is 5- to 14-membered
heteroaryl, C₆-C₁₄-aryl or 3- to 14-membered heterocyclyl.

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- 20 The compound of formula (I) according to any one of claims 1 to 18, or a pharmaceutically acceptable salt thereof, wherein A is 5- to 14-membered heteroaryl or 3- to 14-membered heterocyclyl.
- 21 The compound of formula (I) according to any one of claims 1 to 18, or a pharmaceutically acceptable salt thereof, wherein A is imidazolyl, pyrazolyl, pyrrolyl, or 1,2-dihydropyridinyl.
- 22 The compound of formula (I) according to any one of claims 1 to 21, or a pharmaceutically acceptable salt thereof, wherein B is C₆-C₁₄-aryl, 5- to 14-membered heteroaryl or 3- to 14-membered heterocyclyl.
- 23 The compound of formula (I) according to any one of claims 1 to 21, or a pharmaceutically acceptable salt thereof, wherein B is phenyl, 1,2-dihydropyridinyl or pyridyl.
- 24 The compound of formula (I) according to claim 1, or a pharmaceutically acceptable salt thereof, wherein:
- 15 X is $-\text{CH}_2\text{CR}^3\text{R}^4-$ or $-\text{CH}=\text{CR}^3-$;
- Y is $-\text{O}-$ or $-\text{S}-$;
- R¹ is C₆-C₁₄-aryl or 5- to 14-membered heteroaryl, wherein said C₆-C₁₄-aryl or 5- to 14-membered heteroaryl are substituted with 1-3 substituents independently selected from C₁-C₆-alkyl, C₁-C₆-alkoxy, halo-C₁-C₆-alkoxy, halogen, cyano



R² is hydrogen or C₁-C₆-alkyl;



R³ is a group

R⁴ is hydrogen, halogen or hydroxy;

25 R⁵ is hydrogen, halogen, hydroxy, halo-C₁-C₆-alkyl, C₁-C₆-alkoxy, C₁-C₆-alkyl, cyano or oxo;

R⁶ is hydrogen, halogen, C₆-C₁₄-aryl, 5- to 14-membered heteroaryl, halo-C₁-C₆-alkoxy, halo-C₁-C₆-alkyl, oxo, C₁-C₆-alkyl, C₁-C₆-alkoxy, C₂-C₆-alkenyl, C₂-

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C₆-alkynyl, C₃-C₁₄-cycloalkyl or cyano, wherein said 5- to 14-membered heteroaryl or C₆-C₁₄-aryl is optionally substituted with a substituent selected from halogen, C₁-C₆-alkyl and C₁-C₆-alkoxy;

A and B are each independently 5- to 14-membered heteroaryl, C₆-C₁₄-aryl or 3- to 14-membered heterocyclyl;

n is 1 or 2; and

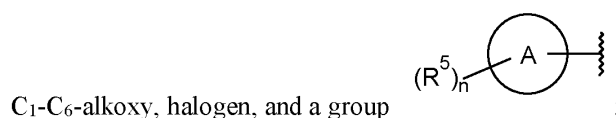
p is 1, 2 or 3.

25 The compound of formula (I) according to claim 1, or a pharmaceutically acceptable salt thereof, wherein:

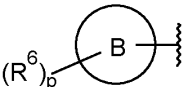
10 X is $-\text{CH}_2\text{CR}^3\text{R}^4-$;

Y is $-\text{O}-$;

R¹ is C₆-C₁₄-aryl substituted with 1-2 substituents independently selected from



R² is hydrogen;

15 R³ is a group  ;

R⁴ is hydrogen or hydroxy;

R⁵ is hydrogen, halogen, cyano, hydroxy or oxo;

R⁶ is halogen, oxo, halo-C₁-C₆-alkyl or halo-C₁-C₆-alkoxy;

A is 5- to 14-membered heteroaryl or 3- to 14-membered heterocyclyl;

20 B is C₆-C₁₄-aryl or 5- to 14-membered heteroaryl; and

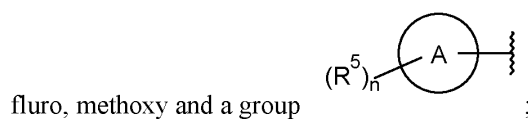
n and p are each independently 1 or 2.

26 The compound of formula (I) according to claim 1, or a pharmaceutically acceptable salt thereof, wherein:

X is $-\text{CH}_2\text{CR}^3\text{R}^4-$;

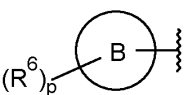
25 Y is $-\text{O}-$;

R¹ is phenyl substituted with 2-3 substituents independently selected from chloro,



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R^2 is hydrogen;

R^3 is a group ;

R^4 is hydrogen or hydroxy;

R^5 is hydrogen, fluoro, cyano, hydroxy or oxo;

5 R^6 is chloro, fluoro, oxo, trifluoromethyl or difluoromethoxy;

A is imidazolyl, pyrazolyl, pyrrolyl, or 1,2-dihydropyridinyl;

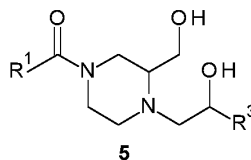
B is phenyl, 1,2-dihydropyridinyl or pyridyl; and

n and p are each independently 1 or 2.

27 The compound of formula (I) according to claim 1, or a pharmaceutically acceptable
10 salt thereof, wherein the compound of formula (I) is selected from the compounds of
Table 1.

28 A process of manufacturing the compounds of formula (I) according to any one of
claims 1 to 27, comprising:

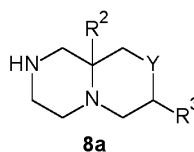
(a) reacting a diol **5**, wherein R^1 and R^3 are as defined herein,



15

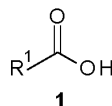
with an etherification reagent, such as (cyanomethylene)tributylphosphorane
(CMBP, Tsunoda reagent) to yield said compound of formula (I), wherein X is
-CH₂CHR³-, Y is -O-, R^1 and R^3 are as defined herein, and R^2 is hydrogen; or

(b) reacting an amine **8a**, wherein Y, R^2 and R^3 are as defined herein,



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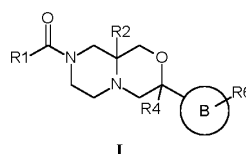
with a carboxylic acid **1**, wherein R^1 is as defined herein,



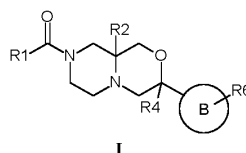
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in the presence of a base, such as trimethylamine or diisopropylamine, and a coupling reagent, such as HATU, T3P or EDC*HCl, to yield said compound of formula (I), wherein X is $-\text{CH}_2\text{CHR}^3-$, Y, R^1 and R^3 are as defined herein, and R^2 is hydrogen; or

- 5 (c) submitting a compound of formula (I), wherein Y is as defined herein, X is $-\text{CH}_2\text{CR}^3\text{R}^4-$, R^1 to R^3 are as defined herein and R^4 is hydroxy to dehydrating conditions, for example using TES and TFA at elevated temperatures, to yield a compound of formula (I) wherein Y is as defined herein, X is $-\text{CH}_2\text{CHR}^3-$ and R^1 to R^3 are as defined herein and/or a compound of formula (I) wherein Y is as defined herein, X is $-\text{CH}=\text{CR}^3-$ and R^1 to R^3 are as defined herein; or
- 10 (d) reacting a compound of formula (I), wherein Y is as defined herein, X is $-\text{CH}_2\text{CR}^3\text{R}^4-$, R^1 to R^3 are as defined herein and R^4 is hydroxy, with a nucleophilic fluorinating agent, such as DAST, to yield a compound of formula (I) wherein X is $-\text{CH}_2\text{CFR}^3-$ and R^1 to R^3 are as defined herein; or
- 15 (e) submitting a compound of formula (I),



wherein R^1 , R^2 , R^4 , and B are as defined herein, and R^6 is bromo, to a transition metal catalyzed cross-coupling reaction, such as a Suzuki–Miyaura reaction or a Sonogashira reaction, to afford compound of formula (I)

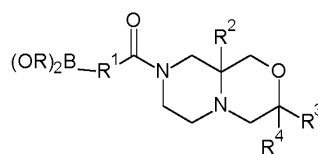


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wherein R^1 , R^2 , R^4 , and B are as defined herein, and R^6 is selected from aryl, alkenyl, alkynyl, alkyl, cyano, cycloalkyl, and heteroaryl; or

- (f) submitting a compound **12**, wherein R^1 is aryl or heteroaryl optionally substituted with 1-4 substituents independently selected from C_1 - C_6 -alkyl, C_1 - C_6 -alkoxy, halo- C_1 - C_6 -alkyl, halo- C_1 - C_6 -alkoxy, halogen, and cyano; and R^2 to R^4 are as defined herein,
- 25

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$B(OR)_2 = B(OH)_2$ or
boronate pinacol ester

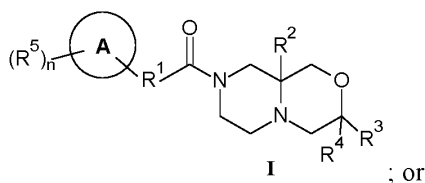
to a Suzuki-Miyaura cross coupling reaction with a compound **13**, wherein A, R⁵, and n are as defined herein



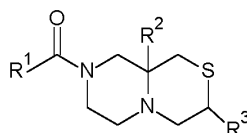
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X = Cl, Br, I, OTf

in the presence of a transition metal catalyst, e.g. $\text{PdCl}_2(\text{dppf})\text{-CH}_2\text{Cl}_2$, $\text{Pd}_2(\text{dba})_3 + \text{Xantphos}$, $(\text{A-}^t\text{Phos})_2\text{PdCl}_2$, or *cataCXium A Pd G2*, and an organic or inorganic base such as sodium carbonate, triethylamine, TMEDA or cesium carbonate; to afford said compound of formula (I), wherein R^1 is aryl or heteroaryl optionally substituted with 1-4 substituents independently selected from $\text{C}_1\text{-C}_6\text{-alkyl}$, $\text{C}_1\text{-C}_6\text{-alkoxy}$, halo- $\text{C}_1\text{-C}_6\text{-alkyl}$, halo- $\text{C}_1\text{-C}_6\text{-alkoxy}$, halogen, and cyano; and R^2 to R^5 and n are as defined herein



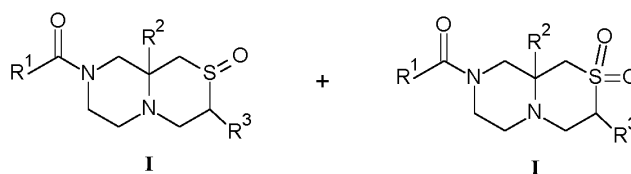
(g) reacting a compound of formula (I), wherein Y is sulfur and R¹, R², and R³ are as defined herein,



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with an oxidizing agent, such as mCPBA, hydrogen peroxide or sodium periodate, to afford a compound of formula (I), wherein Y is a sulfoxide or a sulfone and R¹, R², and R³ are as defined herein.

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- 29 A compound of formula (I) according to any one of claims 1 to 27, when manufactured according to the process of claim 28.
- 30 The compound of formula (I) according to any one of claims 1 to 27 and 29,
5 wherein said compound of formula (I) has an IC₅₀ for monoacylglycerol lipase below 10 μM.
- 31 A compound of formula (I) according to any one of claims 1 to 27, 29 and 30 for use as therapeutically active substance.
- 32 A pharmaceutical composition comprising a compound of formula (I) according
10 to any one of claims 1 to 27, 29 and 30 and a therapeutically inert carrier.
- 33 The use of a compound of formula (I) according to any one of claims 1 to 27, 29 and 30 for the treatment or prophylaxis of neuroinflammation, neurodegenerative diseases, pain, cancer and/or mental disorders in a mammal.
- 34 The use of a compound of formula (I) according to any one of claims 1 to 27, 29
15 and 30 for the treatment or prophylaxis of multiple sclerosis, Alzheimer's disease, Parkinson's disease, amyotrophic lateral sclerosis, traumatic brain injury, neurotoxicity, stroke, epilepsy, anxiety, migraine, depression, hepatocellular carcinoma, colon carcinogenesis, ovarian cancer, neuropathic pain, chemotherapy induced neuropathy, acute pain, chronic pain and/or
20 spasticity associated with pain in a mammal.
- 35 A compound of formula (I) according to any one of claims 1 to 27, 29 and 30 for use in the treatment or prophylaxis of neuroinflammation, neurodegenerative diseases, pain, cancer and/or mental disorders in a mammal.
- 36 A compound of formula (I) according to any one of claims 1 to 27, 29 and 30 for
25 use in the treatment or prophylaxis of multiple sclerosis, Alzheimer's disease, Parkinson's disease, amyotrophic lateral sclerosis, traumatic brain injury, neurotoxicity, stroke, epilepsy, anxiety, migraine, depression, hepatocellular

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carcinoma, colon carcinogenesis, ovarian cancer, neuropathic pain, chemotherapy induced neuropathy, acute pain, chronic pain and/or spasticity associated with pain in a mammal.

- 37 The use of a compound of formula (I) according to any one of claims 1 to 27, 29
5 and 30 for the preparation of a medicament for the treatment or prophylaxis of neuroinflammation, neurodegenerative diseases, pain, cancer and/or mental disorders in a mammal.
- 38 The use of a compound of formula (I) according to any one of claims 1 to 27, 29
10 and 30 for the preparation of a medicament for the treatment or prophylaxis of multiple sclerosis, Alzheimer's disease, Parkinson's disease, amyotrophic lateral sclerosis, traumatic brain injury, neurotoxicity, stroke, epilepsy, anxiety, migraine, depression, hepatocellular carcinoma, colon carcinogenesis, ovarian cancer, neuropathic pain, chemotherapy induced neuropathy, acute pain, chronic pain and/or spasticity associated with pain in a mammal.
- 15 39 A method for the treatment or prophylaxis of neuroinflammation, neurodegenerative diseases, pain, cancer and/or mental disorders in a mammal, which method comprises administering an effective amount of a compound of formula (I) according to any one of claims 1 to 27, 29 and 30 to the mammal.
- 20 40 A method for the treatment or prophylaxis of multiple sclerosis, Alzheimer's disease, Parkinson's disease, amyotrophic lateral sclerosis, traumatic brain injury, neurotoxicity, stroke, epilepsy, anxiety, migraine, depression, hepatocellular carcinoma, colon carcinogenesis, ovarian cancer, neuropathic pain, chemotherapy induced neuropathy, acute pain, chronic pain and/or spasticity associated with pain in a mammal, which method comprises
25 administering an effective amount of a compound of formula (I) according to any one of claims 1 to 27, 29 and 30 to the mammal.
- 41 The invention as described hereinbefore.