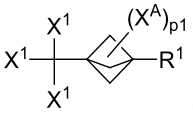
WHAT IS CLAIMED IS:

1. A compound selected from: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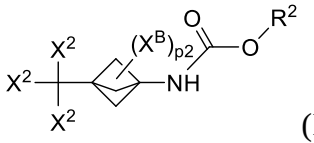
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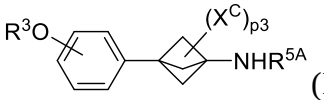
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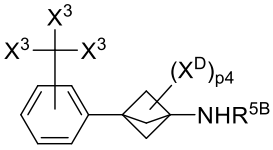
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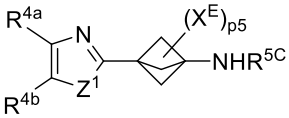
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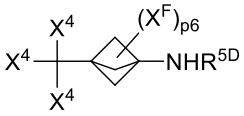
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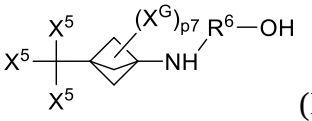
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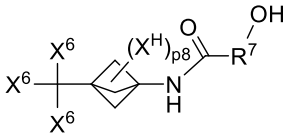
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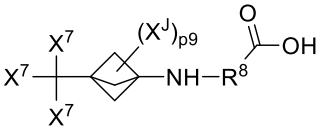
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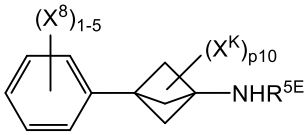
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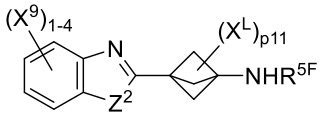
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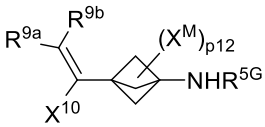
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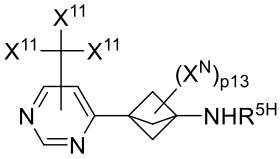
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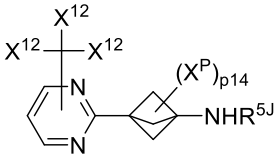
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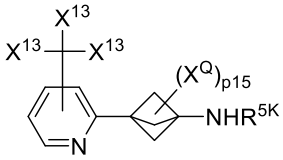
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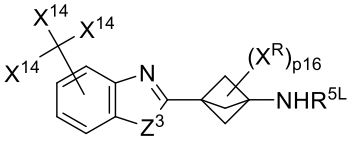
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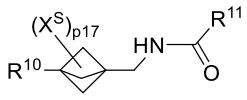
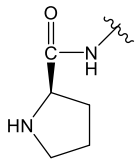

(It),

wherein:

each X^1, each X^2, each X^3, each X^4, each X^5, each X^6, each X^7, each X^{11}, each X^{12}, each X^{13} and each X^{14} are independently hydrogen, deuterium, an unsubstituted C_{1-4} alkyl, fluoro or chloro, provided that at least two of X^1 is fluoro or chloro, provided that at least two of X^2 is fluoro or chloro, provided that at least two of X^3 is fluoro or chloro, provided that at least two of X^4 is fluoro or chloro, provided that at least two of X^5 is fluoro or chloro, provided that at least two of X^6 is fluoro or chloro, provided that at least two of X^7 is fluoro or chloro, provided that at least two of X^{11} is fluoro or chloro, provided that at least two of X^{12} is fluoro or chloro, provided that at least two of X^{13} is fluoro or chloro, and provided that at

least two of X^{14} is fluoro or chloro;

each X^8, each X^9 and X^{10} are independently deuterium, fluoro or chloro;

R^1 is an unsubstituted H_2N-CH(R")-C(=O)-NH-, wherein R" is a side chain of an α-amino acid, or an unsubstituted 

;

R^2 and R^3 are independently an unsubstituted C_{1-4} alkyl;

R^{9a} and R^{9b} are independently an unsubstituted C_{1-4} alkyl;

R^{5A}, R^{5B}, R^{5D}, R^{5E}, R^{5F}, R^{5G}, R^{5H}, R^{5J}, R^{5K} and R^{5L} are independently hydrogen, deuterium or an unsubstituted C_{1-4} alkyl;

R^{5C} is selected from the group consisting of hydrogen, deuterium, an unsubstituted C_{1-4} alkyl and $C(=O)R^{12}$;

R^{4a} and R^{4b} are independently hydrogen, deuterium, an unsubstituted C_{1-4} alkyl, a hydroxy substituted C_{1-4} alkyl or $-C(X^{16})_3$, provided that at least one of R^{4a} and R^{4b} is $-C(X^{16})_3$;

R^6, R^7 and R^8 are independently $-(CH_2)_m-$;

R^{10} is selected from the group consisting of hydrogen, deuterium, halogen, hydroxy, an unsubstituted C_{1-8} alkyl, an unsubstituted C_{3-20} cycloalkyl and an unsubstituted C_{1-8} haloalkyl;

R^{11} is selected from the group consisting of hydrogen, deuterium, halogen, an unsubstituted C_{1-30} alkyl, an unsubstituted C_{2-30} alkenyl and an unsubstituted C_{1-8} haloalkyl;

R^{12} is selected from the group consisting of hydrogen, deuterium, an unsubstituted C_{1-30} alkyl and an unsubstituted C_{2-30} alkenyl;

each X^{16} is independently hydrogen, deuterium, an unsubstituted C_{1-4} alkyl, fluoro or chloro, provided that at least two of X^{16} is fluoro or chloro;

Z^1, Z^2 and Z^3 are independently nitrogen, oxygen or sulfur;

m is 1, 2, 3 or 4;

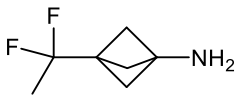
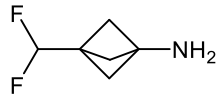
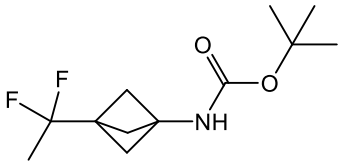
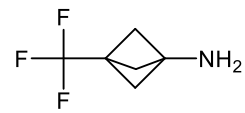
each X^A, each X^B, each X^C, each X^D, each X^E, each X^F, each X^G, each X^H, each X^J, each X^K, each X^L, each X^M, each X^N, each X^P, each X^Q, each X^R and each X^S are independently deuterium, chloro or fluoro;

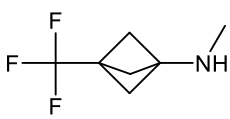
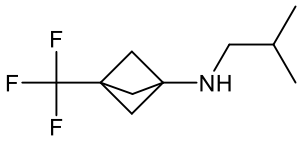
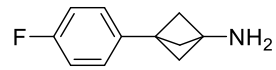
p1, p2, p3, p4, p5, p6, p7, p8, p9, p10, p11, p12, p13, p14, p15 and p16 are independently 0, 1, 2, 3, 4, 5 or 6;

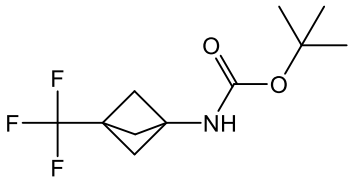
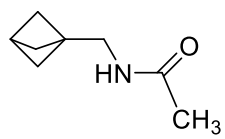
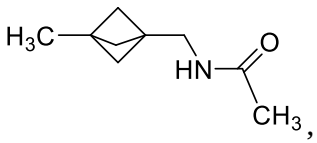
p17 is 1, 2, 3, 4, 5 or 6;

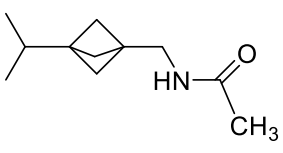
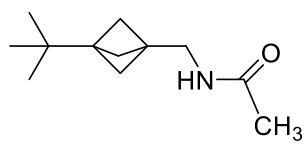
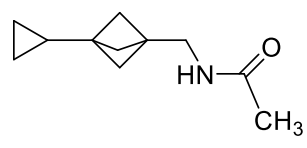
or a pharmaceutically acceptable salt thereof; and

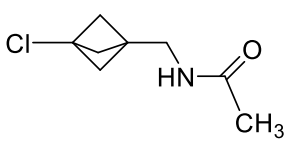
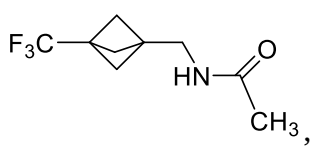
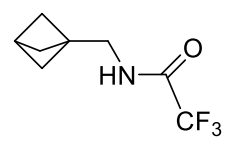
wherein the compound is not selected from the group consisting of:

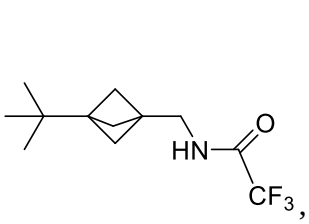
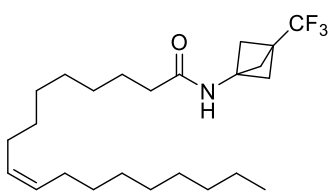
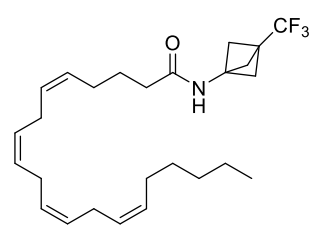
 ,  ,  ,  ,

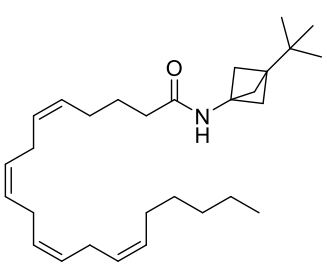
 ,  ,  ,

 ,  ,  ,

 ,  ,  ,

 ,  ,  ,

 ,  ,  ,



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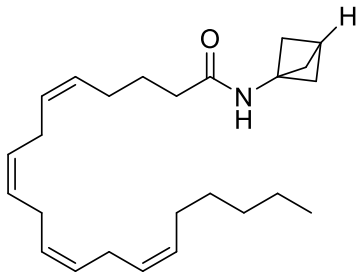


, or a pharmaceutically acceptable salt of any of the foregoing.

2. The compound of Claim 1, or a pharmaceutically acceptable salt thereof, of Formula (Ie).

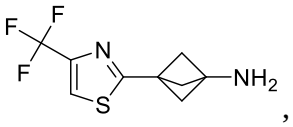
3. The compound of Claim 2, wherein Z^1 is sulfur.

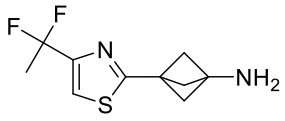
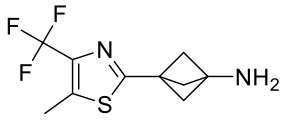
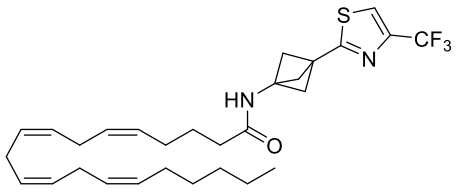
4. The compound of any one of Claims 1-3, or a pharmaceutically acceptable salt thereof, wherein R^{4a} is $-C(X^{16})_3$; and R^{4b} is hydrogen.

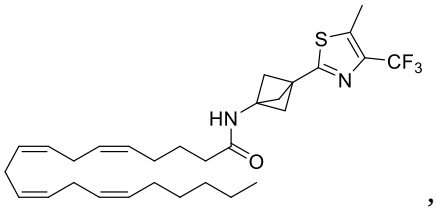
5. The compound of Claim 4, wherein each X^{16} is independently hydrogen, deuterium, fluoro or chloro, provided that at least two of X^{16} is fluoro or chloro.

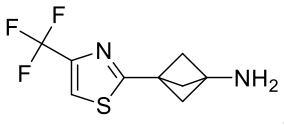
6. The compound of any one of Claims 1-5, wherein p5 is 0.

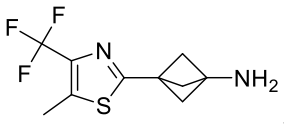
7. The compound of any one of Claims 1-6, wherein R^{5C} is hydrogen.

8. The compound of any one of Claims 1-7, or a pharmaceutically acceptable salt thereof, wherein the compound is selected from the group consisting of  ,

 ,  ,  and

 , or a pharmaceutically acceptable salt of any of the foregoing.

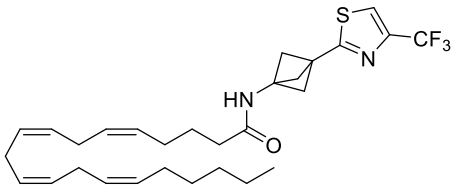
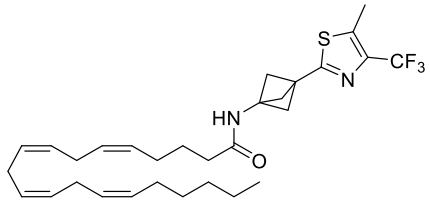
9. The compound of Claim 8, or a pharmaceutically acceptable salt thereof, wherein the compound is  , or a pharmaceutically acceptable salt thereof.

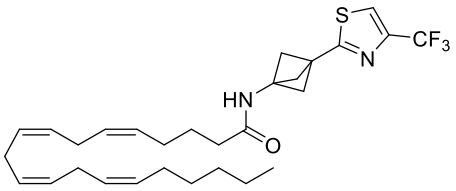
10. The compound of Claim 8, or a pharmaceutically acceptable salt thereof, wherein the compound is  , or a pharmaceutically acceptable salt thereof.

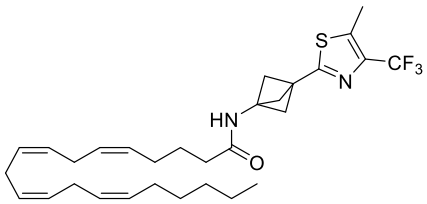
11. The compound of any one of Claims 1-6, or a pharmaceutically acceptable salt thereof, wherein R^{5C} is $C(=O)R^{12}$.

12. The compound of Claim 11, wherein R^{12} is an unsubstituted C_{1-30} alkyl or an unsubstituted C_{2-30} alkenyl.

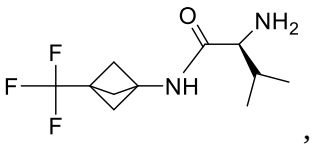
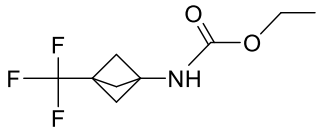
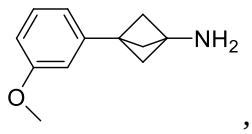
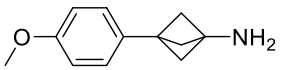
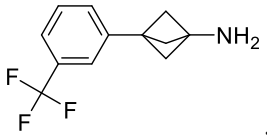
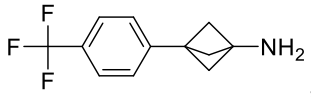
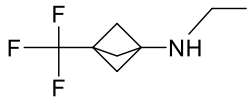
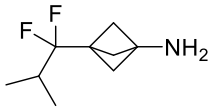
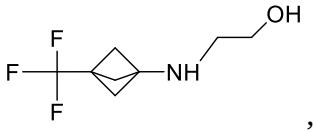
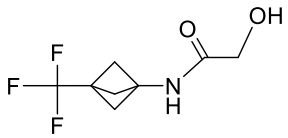
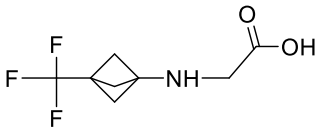
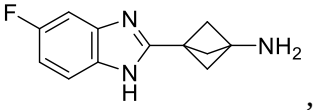
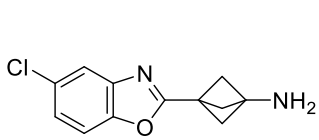
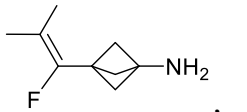
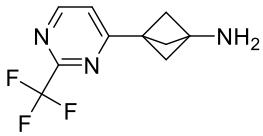
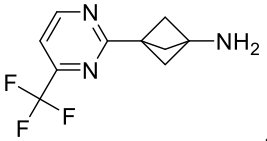
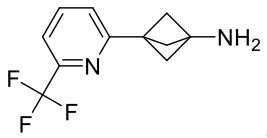
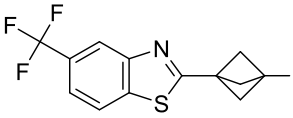
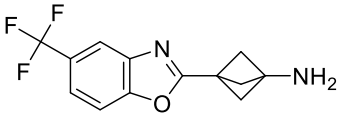
13. The compound of Claim 12, wherein R^{12} is an unsubstituted C_{2-30} alkenyl.

14. The compound of any one of Claims 11-13, or a pharmaceutically acceptable salt thereof, wherein the compound is selected from the group consisting of  and  , or a pharmaceutically acceptable salt of any of the foregoing.

15. The compound of Claim 14, or a pharmaceutically acceptable salt thereof, wherein the compound is  , or a pharmaceutically acceptable salt thereof.

16. The compound of Claim 14, or a pharmaceutically acceptable salt thereof, wherein the compound is  , or a pharmaceutically acceptable salt thereof.

17. The compound of Claim 1, or a pharmaceutically acceptable salt thereof, of Formula (Ia), Formula (Ib), Formula (Ic), Formula (Id), Formula (If), Formula (Ig), Formula (Ih), Formula (Ik), Formula (Im), Formula (In), Formula (Io), Formula (Ip), Formula (Iq), Formula (Ir), Formula (Is) or Formula (It) , or a pharmaceutically acceptable salt of any of the foregoing.

18. The compound of Claim 1, or a pharmaceutically acceptable salt thereof, wherein the compound is selected from the group consisting of  ,  ,  ,  ,  ,  ,  ,  ,  ,  ,  , ,  ,  ,  ,  ,  ,  ,  , and  , or a pharmaceutically acceptable salt of any of the foregoing.

19. A compound of any one of Claims 1-18, or a pharmaceutically acceptable salt of any of the foregoing, for use in reducing or at least partially preventing pain or fever.

20. Use of the compound of Claim 19, or a pharmaceutically acceptable salt of any of the foregoing, and an opioid analgesic and/or a non-steroidal anti-inflammatory drug (NSAID), in the manufacture of a medicament for reducing or at least partially preventing pain or fever.

21. The use of Claim 20, wherein the opioid analgesic is selected from the group consisting of morphine, codeine, hydrocodone, oxycodone, fentanyl, pethidine, methadone, pentazocine, sufentanil, levorphanol, dihydrocodeine, nalbuphine, butorphanol, tramadol, meptazinol, buprenorphine, dipipanone, alfentanil, remifentanil, oxymorphone, tapentadol, propoxyphene and hydromorphone; and wherein the NSAID is selected from the group consisting of celecoxib, ketorolac, ketoprofen, indomethacin, sulindac, etodolac, mefenamic acid, meclofenamic acid, meclofenamate sodium, flufenamic acid, tolmetin, diclofenac, diclofenac sodium, ibuprofen, naproxen, naproxen sodium, fenoprofen, flurbiprofen, oxaprozin, piroxicam, meloxicam, ampiroxicam, droxicam, lornoxicam, cinnoxicam, sudoxicam, and tenoxicam.

22. The use of Claim 20 or Claim 21, wherein the pain is selected from the group consisting of acute pain, post-operative pain, chronic pain, nociceptive pain, osteoarthritis, rheumatoid arthritis, neuropathic pain, migraine, visceral pain, mixed pain, lower back pain, cancer pain and fibromyalgia pain.

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