

CLAIMS

What is claimed is:

- 5 1. A method of treating alopecia areata (AA) in an individual in need thereof comprising:
administering to the individual in need thereof a pharmaceutical dosage form comprising a
therapeutically effective amount of (*R*)-2-(7-(4-cyclopentyl-3-(trifluoromethyl)benzyloxy)-
1,2,3,4-tetrahydrocyclopenta[*b*]indol-3-yl)acetic acid (Compound 1), or a pharmaceutically
acceptable salt thereof.
- 10 2. The method of claim 1, further comprising detecting in the individual an AA-associated
biomarker.
- 15 3. The method of claim 2, wherein the AA-associated biomarker is indicative of severity of
alopecia areata.
4. The method of claim 2, wherein the AA-associated biomarker is indicative of the
propensity of the individual to respond to treatment with Compound 1, or a
pharmaceutically acceptable salt thereof.
- 20 5. The method of any one of the above claims, wherein prior to administering Compound 1,
or a pharmaceutically acceptable salt thereof, to the individual, the method further
comprises selecting the individual based on a level of an AA-associated biomarker.
- 25 6. The method of any one of the preceding claims, wherein the administration of Compound
1, or a pharmaceutically acceptable salt thereof, results in a change in a level of an AA-
associated biomarker in the individual.
- 30 7. The method of any one of the preceding claims, wherein the individual has severe alopecia
areata.
8. The method of any one of the preceding claims, wherein the individual has moderate
alopecia areata.

9. The method of any one of claims 1 to 8, wherein the individual has diffuse alopecia areata.
10. The method of any one of claims 1 to 8, wherein the individual has alopecia areata
5 monolocularis.
11. The method of any one of claims 1 to 8, wherein the individual has alopecia areata
multilocularis.
- 10 12. The method of any one of claims 1 to 8, wherein the individual has ophiasis.
13. The method of any one of claims 1 to 8, wherein the individual has alopecia areata barbae.
14. The method of any one of claims 1 to 7, wherein the individual has alopecia areata totalis.
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15. The method of any one of claims 1 to 7, wherein the individual has alopecia areata
universalis.
16. The method of any one of the preceding claims, wherein the Compound 1, or a
20 pharmaceutically acceptable salt thereof, is administered in combination with a second
therapeutic agent or therapy.
17. The method of any one of the preceding claims, wherein the individual has demonstrated
an inadequate response to, loss of response to, or intolerance of to at least one therapeutic
25 agent or therapy.
18. The method of any one of the preceding claims, wherein the dosage form is administered
under fasted conditions.
- 30 19. The method of any one of claims 1 to 17, wherein the dosage form is administered
under fed conditions.

20. The method of any one of the preceding claims, wherein the therapeutically effective amount is equivalent to about 0.5 to about 5.0 mg of Compound 1.
21. The method of claim 20, wherein the therapeutically effective amount is in an amount
5 equivalent to 2 mg of Compound 1.
22. The method of claim 20, wherein the individual is administered an amount equivalent to 2 mg of Compound 1 for a first time period and subsequently an amount equivalent to 3 mg of Compound 1 for a second time period.
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23. The method of claim 20, wherein the therapeutically effective amount is in an amount equivalent to 3 mg of Compound 1.
24. The method of any one of the preceding claims, wherein the dosage form is
15 administered without titration.
25. The method of any one of the preceding claims, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is administered orally.
- 20 26. The method of any one of the preceding claims, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is formulated as a capsule or tablet suitable for oral administration.
27. The method of any one of the preceding claims, wherein the Compound 1, or a
25 pharmaceutically acceptable salt thereof, is selected from:
Compound 1;
a calcium salt of Compound 1; and
an L-arginine salt of Compound 1.
- 30 28. The method of claim 27, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is an L-arginine salt of Compound 1.

29. The method of claim 28, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is an anhydrous, non-solvated crystalline form of an L-arginine salt of Compound 1.
- 5 30. The method of claim 27, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is an anhydrous, non-solvated crystalline form of Compound 1.
31. The method of any one of the preceding claims, wherein the therapeutically effective amount of Compound 1, or a pharmaceutically acceptable salt thereof is administered once
10 daily to the individual.
32. The method of any one of the preceding claims, wherein the method is non-gender specific.
- 15 33. The method of any one of the preceding claims, wherein the individual was previously administered at least one therapeutic agent or therapy.
34. The method of claim 33, wherein the individual had an inadequate response with, lost response to, or was intolerant to the at least one therapeutic agent or therapy.
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35. The method of any one of the preceding claims, wherein treating comprises inducing and/or maintaining clinical response and/or inducing and/or maintaining clinical remission.
36. The method of any one of the preceding claims, wherein said administering results in no
25 serious adverse events.
37. The method of any one of the preceding claims, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is administered without substantially inducing an acute heart rate reduction or heart block in the individual.
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38. The method of any one of the preceding claims, further comprising monitoring for adverse events during the administration of the Compound 1, or a pharmaceutically acceptable salt

thereof, and optionally, interrupting or terminating the administration of the Compound 1, or a pharmaceutically acceptable salt thereof.

39. The method of any one of Claims 2-38, wherein the AA-associated biomarker is selected from at least one of IL-2, IL-10, IL-12, IL-13, IL-17, IL-17A, IL-22, and IFN- γ .
40. The method of any one of the preceding claims, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is the sole active ingredient in the pharmaceutical dosage form.
41. The method of any one of the preceding claims, wherein the individual has greater than or equal to 50% scalp hair loss.
42. The method of any one of the preceding claims, wherein the method is therapeutically effective to achieve at least a 50% improvement from the individual's baseline Severity of Alopecia Tool (SALT) score.
43. The method of claim 42, wherein the method is therapeutically effective to achieve at least a 50% improvement from the individual's baseline SALT score in a period of time of at least about 24 weeks.
44. The method of claim 43, wherein the method is therapeutically effective to achieve at least a 50% improvement from the individual's baseline SALT score in a period of time of about 24 weeks.
45. A compound that is (R)-2-(7-(4-cyclopentyl-3-(trifluoromethyl)benzyloxy)-1,2,3,4-tetrahydrocyclopenta[b]indol-3-yl)acetic acid (Compound 1), or a pharmaceutically acceptable salt thereof, for use in a method of treatment of alopecia areata in an individual.
46. The compound for use according to claim 45, wherein the alopecia areata is moderate to severe alopecia areata.
47. The compound for use according to claim 45 or claim 46, wherein the alopecia areata is moderate alopecia areata.

48. The compound for use according to any one of claims 45-47, wherein the use further comprises administration of a therapeutically effective amount of Compound 1 in an amount equivalent to about 0.5 to about 5.0 mg of Compound 1.
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49. The compound for use according to claim 48, wherein the therapeutically effective amount is in an amount equivalent to 2 mg of Compound 1.
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50. The compound for use according to claim 48, wherein the therapeutically effective amount is in an amount equivalent to 3 mg of Compound 1.
51. The compound for use according to any one of claims 45-50, wherein the Compound 1 is administered to the individual in need thereof at a frequency of once daily.
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52. The compound for use according to any one of claims 45-51, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is selected from:
- Compound 1;
a calcium salt of Compound 1; and
20 an L-arginine salt of Compound 1.
53. The compound for use according to claim 52, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is an L-arginine salt of Compound 1.
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54. The compound for use according to claim 52, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is an anhydrous, non-solvated crystalline form of an L-arginine salt of Compound 1.
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55. The compound for use according to claim 52, wherein the Compound 1, or a pharmaceutically acceptable salt thereof, is an anhydrous, non-solvated crystalline form of Compound 1.