

CLAIMS

What is claimed is:

- 5 1. A topical composition, comprising:
 a combination of a therapeutically effective amount of superoxide
 dismutase (SOD) with a stabilizing carrier that is suitable for topical
 administration.
- 10 2. The topical composition of claim 1, wherein the stabilizing carrier comprises
 deoxygenated water.
3. The topical composition of claim 2, wherein the deoxygenated water has less than
 one or more of: about 10 ppm of oxygen, about 5 ppm of oxygen, about 3 ppm of
15 oxygen, about 1 ppm of oxygen, or combinations thereof.
4. The topical composition of claim 1, wherein the topical composition further
 comprises a support agent selected from the group consisting of: a stabilizing
 agent, a preservative, an emollient, an adjuvant, and combinations thereof.
- 20 5. The topical composition of claim 4, wherein the support agent is a stabilizing
 agent.
6. The topical composition of claim 5, wherein the stabilizing agent includes an
25 oxygen scavenger, a lipid, a synthetic polymer, a natural polymer, a
 polyelectrolyte, a protein, an amino acid, a co-polymer, an emulsion, a gel, an
 inert gas, or a combination thereof.
7. The topical composition of claim 5, wherein the stabilizing agent is an
30 encapsulating agent that forms a nanoemulsion, a liposome, a nanogel, or a
 combination thereof.

8. The topical composition of claim 7, wherein the encapsulating agent is selected from the group consisting essentially of: chitosan, cyclodextrin, dextran, starch, silicon, tragacanth, and combinations thereof.
- 5 9. The topical composition of claim 5, wherein the stabilizing agent forms a stabilizing complex with the SOD.
- 10 10. The topical composition of claim 5, wherein the stabilizing agent is a lipid selected from the group consisting essentially of: phospholipids, glycolipids, cholesterol, triglycerides, fatty acids, fatty acid glycerides, surfactants, or a combination thereof.
11. The topical composition of claim 10, wherein the lipid forms a liposome, an inverted micelle, or a combination thereof.
- 15 12. The topical composition of claim 1, wherein the topical composition further comprises an additional active agent.
- 20 13. The topical composition of claim 12, wherein the additional active agent is a member selected from the group consisting of: a catalase, an antioxidant, an anti-infective agent, an antibiotic, an anti-tumor agent, an anti-inflammatory agent, an anesthetic, an analgesic, an anti-rheumatic agent, a growth factor, a cytokine, an amino acid, a protein, a vaccine, a hormone, a vitamin, and combinations thereof.
- 25 14. The topical composition of claim 12, wherein the additional active agent is an anesthetic.
- 30 15. The topical composition of claim 14, wherein the anesthetic is a member selected from the group consisting of: articaine, bupivacaine, cinchocaine, etidocaine, levobupivacaine, lidocaine, mepivacaine, prilocaine, ropivacaine, trimecaine, and combinations thereof.
16. The topical composition of claim 14, wherein the anesthetic is lidocaine.

17. The topical composition of claim 12, wherein the additional active agent is in the same or a different composition as the SOD.
18. The topical composition of claim 12, wherein the additional active agent is present
5 at a concentration of from about 0.0001 wt% to about 10 wt%.
19. The topical composition of claim 1, wherein the topical composition has a pH of from about 6.8 to about 7.4.
- 10 20. The topical composition of claim 1, wherein the topical composition has a viscosity of from about 1 cP to about 1000 cP.
21. The topical composition of claim 1, wherein the topical composition is formulated as one of: a solution, a suspension, an emulsion, a gel, a hydrogel, a thermo-
15 responsive gel, a cream, an ointment, a paste, an adhesive, a liquid reservoir, a patch, or a combination thereof.
22. The topical composition of claim 1, wherein the SOD is present in the topical composition at a concentration of from about 0.0001 wt% to about 10 wt%.
- 20 23. The topical composition of claim 1, wherein when the SOD is present in the topical composition at a concentration of 0.02 wt%:
more than about 80% of the SOD remains in a reactive form after a period of about 2 weeks when stored at room temperature and 75% relative humidity; or
25 more than about 80% of the SOD remains in a reactive form after a period of about 2 weeks when stored at 40°C and 75% relative humidity; or
more than about 80% of the SOD remains in a reactive form after a period of about 4 weeks when stored at room temperature and 75% relative humidity; or
30 more than about 80% of the SOD remains in a reactive form after a period of about 4 weeks when stored at 40°C and 75% relative humidity.
24. The topical composition of claim 1, wherein the topical composition consists essentially of SOD and deoxygenated water.

25. The topical composition of claim 1, wherein the topical composition is substantially free of: ethanol, capsaicin, menthol, carbomer, and combinations thereof.
- 5 26. The topical composition of claim 1, wherein the SOD comprises human recombinant SOD (rhSOD), SOD-1, SOD-2, SOD-3, or a combination thereof.
27. A method of treating a condition in a subject that is responsive to treatment with superoxide dismutase (SOD), comprising:
- 10 administering a therapeutically effective amount of the topical composition of any of claims 1 to 26 to the subject.
28. The method of claim 27, wherein the condition is pain.
- 15 29. The method of claim 28, wherein the pain is one or more of chronic pain, back pain, leg pain, ankle pain, joint pain, arthritic pain, wound pain, or combinations thereof.
30. The method of claim 27, wherein the condition is edema.
- 20 31. The method of claim 27, wherein the condition is inflammation.
32. The method of claim 27, wherein the condition is erectile dysfunction.
- 25 33. The method of claim 27, wherein the condition is interstitial cystitis.
34. The method of claim 27, wherein the treatment provides a reduction in symptoms of at least 10% within a selected amount of time after administration.
- 30 35. The method of claim 27, wherein the topical composition comprises a dosage form having from about 25 ul to about 10 ml of SOD.
36. The method of claim 27, wherein a daily dose of the topical composition is from about 25 ul to about 10 ml of SOD.

37. The method of claim 27, further comprising administering the therapeutically effective amount of the topical composition to the subject 1 to 5 times per day.
- 5 38. The method of claim 27, further comprising administering the therapeutically effective amount of the topical composition to the subject according to a dosage regimen of at least once per day for a duration of from about a single day to about 3 months.
- 10 39. The method of claim 27, wherein symptoms of the condition are reduced by at least 10% since commencement of the treatment.
40. A method of stabilizing a superoxide dismutase (SOD) comprising:
combining an amount of SOD with deoxygenated water to form an SOD
15 solution; and
minimizing exposure of the SOD solution to reactive oxygen species (ROS).
41. The method of claim 40, wherein the deoxygenated water has less than one or
20 more of: about 10 ppm of oxygen, about 5 ppm of oxygen, about 3 ppm of oxygen, about 1 ppm of oxygen, or combinations thereof.
42. The method of claim 40, wherein the SOD is present in the SOD solution at a concentration of from about 0.0001 wt% to about 10 wt%.
- 25 43. The method of claim 40, wherein:
when the SOD is present in the SOD solution at a concentration of 0.02 wt%:
more than about 80% of the SOD remains in a reactive form after 2
30 weeks when stored at room temperature and 75% relative humidity; or
more than about 80% of the SOD remains in a reactive form after 4 weeks when stored at room temperature and 75% relative humidity; or
more than about 80% of the SOD remains in a reactive form after 2 weeks when stored at 40°C and 75% relative humidity; or

more than about 80% of the SOD remains in a reactive form after 4 weeks when stored at 40°C and 75% relative humidity.

44. The method of claim 40, wherein:

5 when the SOD is present in the SOD solution at a concentration of 0.02 wt%:

more than about 90% of the SOD remains in a reactive form after 2 weeks when stored at room temperature and 75% relative humidity; or

10 more than about 90% of the SOD remains in a reactive form after 4 weeks when stored at room temperature and 75% relative humidity; or

more than about 90% of the SOD remains in a reactive form after 2 weeks when stored at 40°C and 75% relative humidity; or

more than about 90% of the SOD remains in a reactive form after 4 weeks when stored at 40°C and 75% relative humidity.

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45. The method of claim 40, wherein more than about 80% of the SOD remains in a reactive form after 365 days when stored at room temperature and 75% relative humidity.

20 46. The method of claim 40, wherein the SOD solution further comprises lidocaine.

47. The method of claim 46, wherein:

when the SOD is present in the SOD solution at a concentration of 0.02 wt% and the lidocaine is present in the SOD solution at a concentration of 0.5 wt%:

25 more than about 80% of the SOD remains in a reactive form after 2 weeks when stored at room temperature and 75% relative humidity; or

more than about 80% of the SOD remains in a reactive form after 4 weeks when stored at room temperature and 75% relative humidity; or

30 more than about 80% of the SOD remains in a reactive form after 2 weeks when stored at 40°C and 75% relative humidity; or

more than about 80% of the SOD remains in a reactive form after 4 weeks when stored at 40°C and 75% relative humidity.

48. The method of claim 40, wherein:

when the SOD is present in the SOD solution at a concentration of 0.02 wt% and the lidocaine is present in the SOD solution at a concentration of 0.5 wt%:

- 5 more than about 90% of the SOD remains in a reactive form after 2 weeks when stored at room temperature and 75% relative humidity; or
 more than about 90% of the SOD remains in a reactive form after 4 weeks when stored at room temperature and 75% relative humidity; or
 more than about 90% of the SOD remains in a reactive form after 2 weeks when stored at 40°C and 75% relative humidity; or
10 more than about 90% of the SOD remains in a reactive form after 4 weeks when stored at 40°C and 75% relative humidity.
49. The method of claim 40, wherein the SOD solution is substantially free of ethanol, capsaicin, menthol, carbomer, and combinations thereof.
- 15 50. The method of claim 40, wherein the SOD is one or more of human recombinant SOD (rhSOD), SOD-1, SOD-2, SOD-3, or a combination thereof.
- 20 51. A topical composition for use in the treatment of a condition in a subject that is responsive to treatment with superoxide dismutase, comprising:
 administering a therapeutically effective amount of the topical composition of any of claims 1-26 to the subject.
- 25 52. The topical composition for use of claim 51, wherein the condition is pain.
53. The topical composition for use of claim 52, wherein the pain is one or more of chronic pain, back pain, leg pain, ankle pain, joint pain, arthritic pain, wound pain, or combinations thereof.
- 30 54. The topical composition for use of claim 51, wherein the condition is edema.
55. The topical composition for use of claim 51, wherein the condition is inflammation.

56. The topical composition for use of claim 51, wherein the condition is erectile dysfunction.
57. The topical composition for use of claim 51, wherein the condition is interstitial cystitis.
58. The use of a topical composition in the manufacture of a medicament for the treatment of a condition in a subject that is responsive to treatment with superoxide dismutase, comprising:
- 10 administering a therapeutically effective amount of the topical composition of any of claims 1-26 to the subject.
59. The use of claim 58, wherein the condition is pain.
60. The use of claim 59, wherein the pain is one or more of chronic pain, back pain, leg pain, ankle pain, joint pain, arthritic pain, wound pain, or combinations thereof
61. The use of claim 58, wherein the condition is edema.
62. The use of claim 58, wherein the condition is inflammation.
63. The use of claim 58, wherein the condition is erectile dysfunction.
64. The use of claim 58, wherein the condition is interstitial cystitis.