

A Systematic Review of the Clinical Use of Withania Somnifera (Ashwagandha) to Ameliorate Cognitive Dysfunction

[Qin Xiang Ng ^{1,2}](#), [Wayren Loke ¹](#), [Nadine Xinhui Foo ¹](#), [Weng Jun Tan ^{1,2}](#), [Hwei Wuen Chan ^{3,4}](#),
[Donovan Yutong Lim ^{2,4}](#), [Wee Song Yeo ^{3,4}](#)

Affiliations

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Abstract

Many developed countries are experiencing a rapidly "greying" population, and cognitive decline is common in the elderly. There is no cure for dementia, and pharmacotherapy options to treat cognitive dysfunction provide limited symptomatic improvements. *Withania somnifera* (Ashwagandha), a popular herb highly valued in Ayurvedic medicine, has often been used to aid memory and cognition. This systematic review thus aimed to evaluate the clinical evidence base and investigate the potential role of *W. somnifera* in managing cognitive dysfunction. Using the following keywords [withania somnifera OR indian ginseng OR Ashwagandha OR winter cherry] AND [brain OR cognit* OR mental OR dementia OR memory], a comprehensive search of PubMed, EMBASE, Medline, PsycINFO and Clinicaltrials.gov databases found five clinical studies that met the study's eligibility criteria. Overall, there is some early clinical evidence, in the form of randomized, placebo-controlled, double-blind trials, to support the cognitive benefits of *W. somnifera* supplementation. However, a rather heterogeneous study population was sampled, including older adults with mild cognitive impairment and adults with schizophrenia, schizoaffective disorder, or bipolar disorder. In most instances, *W. somnifera* extract improved performance on cognitive tasks, executive function, attention, and reaction time. It also appears to be well tolerated, with good adherence and minimal side effects.

Keywords: Aryurveda; Ashwagandha; Indian ginseng; *Withania somnifera*; ageing; cognitive decline; dementia.

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- [Antihyperalgesic effects of ashwagandha \(Withania somnifera root extract\) in rat models of postoperative and neuropathic pain.](#) Lim DW, Kim JG, Lim EY, Kim YT. *Inflammopharmacology*. 2018 Feb;26(1):207-215. doi: 10.1007/s10787-017-0389-1. Epub 2017 Aug 28. PMID: 2849547
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Young V, Morley CP. J Altern Complement Med. 2014 Dec;20(12):901-8. doi: 10.1089/acm.2014.0177. PMID: 25405876 Free PMC article. Review.

- [Adjunctive Use of a Standardized Extract of Withania somnifera \(Ashwagandha\) to Treat Symptom Exacerbation in Schizophrenia: A Randomized, Double-Blind, Placebo-Controlled Study.](#) Chengappa KNR, Brar JS, Gannon JM, Schlicht PJ. J Clin Psychiatry. 2018 Jul 10;79(5):17m11826. doi: 10.4088/JCP.17m11826. PMID: 29995356 Clinical Trial.
- [Examining the effect of Withania somnifera supplementation on muscle strength and recovery: a randomized controlled trial.](#) Wankhede S, Langade D, Joshi K, Sinha SR, Bhattacharyya S. J Int Soc Sports Nutr. 2015 Nov 25;12:43. doi: 10.1186/s12970-015-0104-9. eCollection 2015. PMID: 26609282 Free PMC article. Clinical Trial.
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- [Scientific basis for the therapeutic use of Withania somnifera \(ashwagandha\): a review.](#) Mishra LC, Singh BB, Dagenais S. Altern Med Rev. 2000 Aug;5(4):334-46. PMID: 10956379 Review.
- [Effect of standardized aqueous extract of Withania somnifera on tests of cognitive and psychomotor performance in healthy human participants.](#) Pingali U, Pilli R, Fatima N. Pharmacognosy Res. 2014 Jan;6(1):12-8. doi: 10.4103/0974-8490.122912. PMID: 24497737 Free PMC article.
- [Safety and clinical effectiveness of Withania Somnifera \(Linn.\) Dunal root in human ailments.](#) Tandon N, Yadav SS. J Ethnopharmacol. 2020 Mar 19;112768. doi: 10.1016/j.jep.2020.112768. Online ahead of print. PMID: 32201301 Review.
- [Neuropharmacological Properties of Withania somnifera - Indian Ginseng: An Overview on Experimental Evidence with Emphasis on Clinical Trials and Patents.](#) Yenisetti SC, Manjunath MJ, Muralidhara C. Recent Pat CNS Drug Discov. 2016;10(2):204-215. doi: 10.2174/1574889810666160615014106. PMID: 27316579 Review.
- [Systemic administration of defined extracts from Withania somnifera \(Indian Ginseng\) and Shilajit differentially affects cholinergic but not glutamatergic and GABAergic markers in rat brain.](#) Schliebs R, Liebmann A, Bhattacharya SK, Kumar A, Ghosal S, Bigl V. Neurochem Int. 1997 Feb;30(2):181-90. doi: 10.1016/s0197-0186(96)00025-3. PMID: 9017665
- [Withania somnifera \(Indian ginseng\) in diabetes mellitus: A systematic review and meta-analysis of scientific evidence from experimental research to clinical application.](#) Durg S,

Bavage S, Shivaram SB. Phytother Res. 2020 Jan 23. doi: 10.1002/ptr.6589. Online ahead of print. PMID: 31975514 Review.

- [In vitro screening of neuroprotective activity of Indian medicinal plant *Withania somnifera*.](#) Singh M, Ramassamy C. J Nutr Sci. 2017 Oct 18;6:e54. doi: 10.1017/jns.2017.48. eCollection 2017. PMID: 29152258 Free PMC article.

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References

REFERENCES

1. Anbalagan, K. (1982). Influence of an Indian medicine (Ashwagandha) on acute-phase reactants in inflammation. Indian Journal of Experimental Biology, 19, 245-249.
2. Anbalagan, K., & Sadique, J. (1984 Apr 1). Role of prostaglandins in acute phase proteins in inflammation. Biochemical Medicine, 31(2), 236-245. [https://doi.org/10.1016/0006-2944\(84\)90028-0](https://doi.org/10.1016/0006-2944(84)90028-0)
3. Audyy, B., Hazra, J., Mitra, A., Abedon, B., & Ghosal, S. (2008). A standardized Withania somnifera extract significantly reduces stress related parameters in chronically stressed humans: A double blind, randomized, placebo controlled study. [Journal of American Nutraceutical Association, 11, 50-56. Available from: http://www.lifeforce.net/pdfs/withania_review.pdf
4. Chengappa, K. R., Bowie, C. R., Schlicht, P. J., Fleet, D., Brar, J. S., & Jindal, R. (2013). Randomized placebo-controlled adjunctive study of an extract of Withania somnifera for cognitive dysfunction in bipolar disorder. The Journal of Clinical Psychiatry., 74(11), 1076-1083. <https://doi.org/10.4088/JCP.13m08413>
5. Chengappa, K. R., Brar, J. S., Gannon, J. M., & Schlicht, P. J. (2018). Adjunctive use of a standardized extract of Withania somnifera (Ashwagandha) to treat symptom exacerbation in schizophrenia: A randomized, double-blind, placebo-controlled study. The Journal of Clinical Psychiatry, 79(5), 17m11826.
6. Choudhary, D., Bhattacharyya, S., & Bose, S. (2017). Efficacy and safety of Ashwagandha (Withania somnifera (L.) Dunal) root extract in improving memory and cognitive functions. Journal of Dietary Supplements., 14(6), 599-612. <https://doi.org/10.1080/19390211.2017.1284970>
7. Corson, T. W., & Crews, C. M. (2007 Sep 7). Molecular understanding and modern application of traditional medicines: Triumphs and trials. Cell, 130(5), 769-774. <https://doi.org/10.1016/j.cell.2007.08.021>
8. Dave, U. P., Chauhan, V., & Dalvi, J. (1993 May 1). Evaluation of BR-16 a (Mentat) in cognitive and behavioural dysfunction of mentally retarded children- A placebo-controlled study. The Indian Journal of Pediatrics., 60(3), 423-428. <https://doi.org/10.1007/bf02751207>

9. Erhart, S. M., Marder, S. R., & Carpenter, W. T. (2006 Feb 21). Treatment of schizophrenia negative symptoms: Future prospects. *Schizophrenia Bulletin*, 32(2), 234-237. <https://doi.org/10.1093/schbul/sbj055>
10. Higgins, J. P., Altman, D. G., Gøtzsche, P. C., Jüni, P., Moher, D., Oxman, A. D., ... Sterne, J. A. (2011 Oct 18). The Cochrane Collaboration's tool for assessing risk of bias in randomised trials. *BMJ*, 343, d5928.
11. Hill, S. K., Reilly, J. L., Harris, M. S., Rosen, C., Marvin, R. W., DeLeon, O., & Sweeney, J. A. (2009 Sep 1). A comparison of neuropsychological dysfunction in first-episode psychosis patients with unipolar depression, bipolar disorder, and schizophrenia. *Schizophrenia Research*, 113(2-3), 167-175. <https://doi.org/10.1016/j.schres.2009.04.020>
12. Hurd, M. D., Martorell, P., Delavande, A., Mullen, K. J., & Langa, K. M. (2013 Apr 4). Monetary costs of dementia in the United States. *The New England Journal of Medicine*, 368(14), 1326-1334. <https://doi.org/10.1056/NEJMsa1204629>
13. Jayaprakasam, B., & Nair, M. G. (2003 Feb 3). Cyclooxygenase-2 enzyme inhibitory withanolides from *Withania somnifera* leaves. *Tetrahedron*, 59(6), 841-849.
14. Joyashiki, E., Matsuya, Y., & Tohda, C. (2011 Mar 2). Sominone improves memory impairments and increases axonal density in Alzheimer's disease model mice, 5XFAD. *The International Journal of Neuroscience*, 121(4), 181-190. <https://doi.org/10.3109/00207454.2010.541571>
15. Knopman, D. S., & Petersen, R. C. (2014). Mild cognitive impairment and mild dementia: A clinical perspective. *Mayo Clinic Proceedings*, 89, 1452-1459. <https://doi.org/10.1016/j.mayocp.2014.06.019>
16. Konar, A., Shah, N., Singh, R., Saxena, N., Kaul, S. C., Wadhwa, R., & Thakur, M. K. (2011 Nov 11). Protective role of Ashwagandha leaf extract and its component withanone on scopolamine-induced changes in the brain and brain-derived cells. *PLoS ONE*, 6(11), e27265. <https://doi.org/10.1371/journal.pone.0027265>
17. Kuboyama, T., Tohda, C., & Komatsu, K. (2006 Mar). Withanoside IV and its active metabolite, sominone, attenuate A β (25-35)-induced neurodegeneration. *The European Journal of Neuroscience*, 23(6), 1417-1426. <https://doi.org/10.1111/j.1460-9568.2006.04664.x>
18. Kuboyama, T., Tohda, C., Zhao, J., Nakamura, N., Hattori, M., & Komatsu, K. (2002 Oct 7). Axon-or dendrite-predominant outgrowth induced by constituents from Ashwagandha. *Neuroreport*, 13(14), 1715-1720. <https://doi.org/10.1097/00001756-200210070-00005>
19. Manchanda, S., & Kaur, G. (2017 Dec). *Withania somnifera* leaf alleviates cognitive dysfunction by enhancing hippocampal plasticity in high fat diet induced obesity model. *BMC Complementary and Alternative Medicine*, 17(1), 136. <https://doi.org/10.1186/s12906-017-1652-0>
20. Marell, P., Murphy, T., Brar, J., Schlicht, P., Haigh, S., Coffman, B., ... Salisbury, D. (2017). Pilot study on the effects of *Withania somnifera* on electrophysiological measures of sensory and cognitive processing in schizophrenia. *Schizophrenia Bulletin*, 43(Suppl 1), S185.

21. Mattson, M. P., Maudsley, S., & Martin, B. (2004 Oct 1). BDNF and 5-HT: A dynamic duo in age-related neuronal plasticity and neurodegenerative disorders. *Trends in Neurosciences*, 27(10), 589-594. <https://doi.org/10.1016/j.tins.2004.08.001>
22. Mirjalili, M., Moyano, E., Bonfill, M., Cusido, R., & Palazón, J. (2009). Steroidal lactones from *Withania somnifera*, an ancient plant for novel medicine. *Molecules*, 14(7), 2373-2393. <https://doi.org/10.3390/molecules14072373>
23. Mishra, L. C., Singh, B. B., & Dagenais, S. (2000 Aug 1). Scientific basis for the therapeutic use of *Withania somnifera* (Ashwagandha): A review. *Alternative Medicine Review*, 5(4), 334-346.
24. Na, K. S., Jung, H. Y., & Kim, Y. K. (2014 Jan 3). The role of pro-inflammatory cytokines in the neuroinflammation and neurogenesis of schizophrenia. *Progress in Neuro-Psychopharmacology & Biological Psychiatry*, 48, 277-286.
25. Ng, Q. X., Koh, S. S., Chan, H. W., & Ho, C. Y. (2017 Jun 1). Clinical use of curcumin in depression: A meta-analysis. *Journal of the American Medical Directors Association*, 18(6), 503-508. <https://doi.org/10.1016/j.jamda.2016.12.071>
26. Patel, S. B., Rao, N. J., & Hingorani, L. L. (2016 Mar 1). Safety assessment of *Withania somnifera* extract standardized for Withaferin A: Acute and sub-acute toxicity study. *Journal of Ayurveda and Integrative Medicine*, 7(1), 30-37. <https://doi.org/10.1016/j.jaim.2015.08.001>
27. Pereira, F. S., Yassuda, M. S., Oliveira, A. M., & Forlenza, O. V. (2008 Dec). Executive dysfunction correlates with impaired functional status in older adults with varying degrees of cognitive impairment. *International Psychogeriatrics*, 20(6), 1104-1115. <https://doi.org/10.1017/S1041610208007631>
28. Perng, C. H., Chang, Y. C., & Tzang, R. F. (2018 May 1). The treatment of cognitive dysfunction in dementia: A multiple treatments meta-analysis. *Psychopharmacology*, 235(5), 1571-1580. <https://doi.org/10.1007/s00213-018-4867-y>
29. Pingali, U., Pilli, R., & Fatima, N. (2014 Jan). Effect of standardized aqueous extract of *Withania somnifera* on tests of cognitive and psychomotor performance in healthy human participants. *Pharmacognosy Research*, 6(1), 12-18. <https://doi.org/10.4103/0974-8490.122912>
30. Rege, N. N., Thatte, U. M., & Dahanukar, S. A. (1999 Jun). Adaptogenic properties of six rasayana herbs used in Ayurvedic medicine. *Phytotherapy Research: An International Journal Devoted to Pharmacological and Toxicological Evaluation of Natural Product Derivatives*, 13(4), 275-291.
31. Rosenberg, P. B., Mielke, M. M., Appleby, B., Oh, E., Leoutsakos, J. M., & Lyketsos, C. G. (2011 Apr). Neuropsychiatric symptoms in MCI subtypes: The importance of executive dysfunction. *International Journal of Geriatric Psychiatry*, 26(4), 364-372. <https://doi.org/10.1002/gps.2535>
32. Shetty, P. (2012 Apr 7). Grey matter: Ageing in developing countries. *The Lancet*, 379(9823), 1285-1287. [https://doi.org/10.1016/s0140-6736\(12\)60541-8](https://doi.org/10.1016/s0140-6736(12)60541-8)
33. Tranter, R., O'Donovan, C., Chandarana, P., & Kennedy, S. (2002 Jul). Prevalence and outcome of partial remission in depression. *Journal of Psychiatry & Neuroscience*, 27(4), 241-247.

34. Van Kesteren, C. F., Gremmels, H., De Witte, L. D., Hol, E. M., Van Gool, A. R., Falkai, P. G., ... Sommer, I. E. (2017 Mar). Immune involvement in the pathogenesis of schizophrenia: A meta-analysis on postmortem brain studies. *Translational Psychiatry*, 7(3), e1075. <https://doi.org/10.1038/tp.2017.4>
35. Wimo, A., Winblad, B., & Jönsson, L. (2010 Mar 1). The worldwide societal costs of dementia: Estimates for 2009. *Alzheimer's & Dementia*, 6(2), 98-103. <https://doi.org/10.1016/j.jalz.2010.01.010>
36. World Health Organisation. Dementia. 2019. Available from: <http://www.who.int/news-room/fact-sheets/detail/dementia> [].
37. Yadav, C. S., Kumar, V., Suke, S. G., Ahmed, R. S., Mediratta, P. K., & Banerjee, B. D. (2010 Aug). Propoxur-induced acetylcholine esterase inhibition and impairment of cognitive function: Attenuation by *Withania somnifera*. *Indian Journal of Biochemistry & Biophysics*, 47, 117-120.