



Astragalus

Astragalus membranaceus

Common names

Milkvetch root, Mongolian milkvetch, huang qi

Family

Fabaceae (legume)

Part used

Root

Background and traditional uses

Astragalus, also known as radix astragali or huang qi, the latter translating to ‘yellow leader’, is a perennial plant with a yellow cylindrical root.^{1,2}

Used extensively in traditional Chinese medicine (TCM), astragalus is indigenous to China, Korea, Mongolia and Russia and is considered to have sweet and slightly warm properties.^{1,3,4}

Astragalus root has been used therapeutically for many centuries, with the original report of its use recorded in the Chinese herbal classic, Shen Nong Ben Cao Jing (Divine Husbandman’s Classic of the Materia Medica) in the second century.^{4,5}

Considered to be an important TCM tonic herb for enhancing immunity, astragalus tonifies qi and increases yang energy. It has been used individually and in combination with other herbs to treat qi deficiency and its associated symptoms including fatigue, weakness, poor appetite, diarrhoea and frequent respiratory infections or colds.²⁻⁶

Additionally, astragalus has been used traditionally for its diuretic properties in the treatment of such conditions as spontaneous sweating, oedema, postpartum fever, uterine retention, wound healing, organ prolapse, heavy blood loss and diabetes.^{1,3-5,7}

Actions

Primary:^{3,8}

- Immune enhancing
- Immune modulating
- Adaptogen
- Anti-inflammatory
- Antioxidant

Secondary:^{3,8}

- Antiviral
- Anticarcinogenic
- Antidiabetic
- Cardioprotective
- Hepatoprotective
- Vulnerary
- Diuretic
- Nutritive for bone building

Applications and indications

- Common cold, upper respiratory infections, allergic rhinitis, and to strengthen and regulate the immune system.^{3,9,10}
- Athletic performance and reducing exercise-induced fatigue.^{3,11-13}
- Antiviral activity against Epstein-Barr virus (EBV), hepatitis B and herpes simplex virus (HSV)-1.¹⁴⁻¹⁷
- Conditions requiring diuresis.¹⁸

Active constituents and pharmacodynamics

The primary chemical constituents present in the root of astragalus that contribute to the herbs pharmacological effects are polysaccharides, saponins and flavonoids.^{1,2,19}

The main **polysaccharide** constituents are astragalans I, II and III.^{1,7} These constituents exhibit a range of pharmacological effects including immune stimulating, anti-inflammatory, antioxidant, hepatoprotective, antiasthmatic, antidiabetic²⁴, antitumour and cardioprotective activity.^{1,7,20-28}

Triterpene saponins are also present with the most therapeutically relevant being the astragalosides I-VIII.^{1,2,7} Specifically, astragaloside IV is considered to be a significant active constituent, and is associated with immune stimulating, hepatoprotective, cardioprotective, antiproliferative, antiasthmatic, anti-inflammatory, antioxidant and CNS-protective effects.^{2,19,29,30-32}

The main **flavonoid** constituents are flavones, isoflavones, pterocarpanes and chalcones.^{2,7} Pharmacological activity associated with flavonoids include anti-inflammatory, antiproliferative, immunomodulatory and antioxidant effects.^{2,6,7,21}

Summary of clinical evidence

Immune modulation

Human, animal and *in vitro* evidence has demonstrated the effect of astragalus on immune activity. In a small study, four subjects were sublingually administered three different doses of astragalus on in a three-arm design separated by four week washout periods, with the physiological response to the herb assessed over the subsequent 24-hour period.¹⁰ The doses of astragalus (1:5) were 0.25, 0.75 and 1.5mL per kg, equivalent to a dose of 3.3, 10 and 20g of dried root respectively.

Physiological assessments included white blood cell populations (total WBC, lymphocytes, monocytes, neutrophils, platelets), lymphocyte subpopulations (total T-cells, T-helper cells, T-cytotoxic cells, B-cells and NK cells), and cytokine assay (IL-1 beta, IL-2, sIL-2R, IFN-gamma, IL-5-8, IL-10, IL-12, IL-13 and TNF-alpha). Liver (AST, ALT, bilirubin and ALP) and kidney (sodium creatinine and blood urea nitrogen) panels were also conducted. Subjective assessments included fatigue, malaise, headache and reduced mental focus measures.

Treatment with astragalus resulted in statistically significant, dose-dependent increases in total white blood cell count (monocytes, neutrophils and lymphocytes) ($p=0.042$ for lowest dose and $p=0.011$ for highest dose) after eight to 12 hours compared to baseline levels.

There was also a significant increase from baseline in cytokine levels (interferon-gamma [$p=0.018$], TNF-alpha [$p=0.046$] and soluble IL-2R [$p=0.056$]) and improvement in fatigue and headache symptoms in subjective assessments.

A separate study also observed a beneficial effect on immune cells with oral use of astragalus root extract.⁹ Healthy subjects were given either placebo ($n=2$), a combination of astragalus, *Glycyrrhiza glabra* (licorice) and *Echinacea purpurea* (echinacea) ($n=3$) or one of these herbs individually ($n=4$ each for astragalus and echinacea, $n=3$ for licorice). The tinctures were administered at a dose of 7.5mL twice daily for seven days, providing 1.23g of astragalus, 0.93g echinacea and 0.87g licorice equivalent to dry.

Astragalus administered individually had the strongest effect on immune cell activation compared with the other individual tinctures. Within 24 hours following ingestion, CD69 expression on CD8 and CD4 T-cells were significantly stimulated compared with minimal cell activation by placebo. After seven days, the level of immune cell activation was less than that at 24 hours but was higher than baseline levels. The combination tincture resulted in CD69 immune cell activation levels higher at 24 hours and seven days after ingestion than those observed in the individual tincture groups. These studies demonstrate that astragalus has a stimulating impact on immune cell number and activity.

Physical performance

Human and animal data indicates that astragalus and its constituents may have a beneficial influence in physical performance. Astragalus in combination with *Panax ginseng* (Korean ginseng), *Panax quinquefolius* (American ginseng) and creatine (CrBE) was given to 44 adults aged 55-84 years, three times daily during a 12-week strength training program.¹¹ Compared with placebo or creatine only groups, the CrBE group had more significant improvements in strength, self-reported vigour, and reductions in body fat and cholesterol levels (total, LDL, VLDL, triglycerides, LDL/HDL ratio). There was also a lower incidence of bloating in the CrBE versus creatine-only group.

In a separate study, subjects who had experienced strokes were orally administered either astragalus (equivalent to 2.8g TID) ($n=29$) or placebo ($n=32$) for 28 days, with the primary outcome measures being the Brief Fatigue Index (BFI) and quality of life assessments.³³ At the end of the treatment period the astragalus group had a greater difference in BFI ($p=0.05$), global quality of life, cognitive and social functioning scores compared with placebo ($p=0.05$).

A mice study investigated the effect of astragalus root extract prior to exercise on body composition and physical performance.¹² Animals were orally administered astragalus extract at doses of either 0.615g/kg/day or 3.075g/kg/day or placebo for six weeks (calculated to be equivalent to 3g daily human dosage). Astragalus resulted in improved endurance exercise capacity, hepatic and muscle glycogen content and reduced levels of exercise-induced ammonia and lactate following acute exercise.

A rat study observed that polysaccharides from astragalus given orally at doses of 50, 100 and 200mg/kg for 30 days resulted in a significantly extended length of exhaustive exercise performed and skeletal muscle antioxidant enzyme activity (superoxide dismutase and glutathione peroxidase) compared with placebo.¹³

Antiviral

Preliminary evidence demonstrates that astragalus and its constituents exhibit antiviral effects. Animal and *in vitro* investigations have observed that astragalus inhibited HSV-1 and Cocksackievirus cell concentration and necrotic size.^{14,15}

Constituents of astragalus specifically play a role, with polysaccharides observed to inhibiting EBV expression during the lytic cycle and astragaloside IV reducing replication of human adenovirus *in vitro*.^{16,17}

Diuretic

The traditional application of astragalus as a diuretic has some scientific basis, with evidence indicating a potential beneficial effect.

A double-blind randomised crossover study in 12 healthy males investigated the natriuretic effect of astragalus.¹⁸ Subjects were orally administered a single dose of either astragalus (0.3g/kg aqueous extract) (n=6) or placebo (n=6) and were monitored for the subsequent 24-hour period. Astragalus resulted in an elevated natriuresis rate compared to placebo subsequent to increases in plasma atrial natriuretic peptide and consequent renal response.

Dosage summary

Liquid extract (1:2): 30-60mL weekly³⁵

Dried herb equivalent: 2-30g daily⁸

Immune support: 2-4.5g daily³⁵

Safety information

- Use during pregnancy and lactation is not advised.¹
- Caution is advised in individuals with an autoimmune disorder.³⁶
- Caution is advised regarding concomitant use with immune suppressing medications.⁸
- According to the TCM treatment paradigm, astragalus is contraindicated in individuals with exterior pathogens, excess qi stagnation, excess internal heat, damp obstruction, or with sores and lesions.⁴



References

1. World Health Organization (WHO). *Radix astragali*. Monographs on selected medicinal plants, vol. 1. Geneva: WHO, 1999. Viewed 23 Oct 2017, <http://apps.who.int/medicinedocs/en/d/Js2200e/7.html#Js2200e.7>
2. Auyeung KK, Han QB, Ko JK. *Astragalus membranaceus*: a review of its protection against inflammation and gastrointestinal cancers. *Am J Chin Med* 2016;44(1):1-
3. Astragalus. Natural medicines database. Viewed 23 Oct 2017, <https://naturalmedicines.therapeuticresearch.com/databases/food,-herbs-supplements/professional.aspx?productid=963>
4. Chen J, Chen T. Huang qi (*Radix Astragali*). Chinese medical herbology and pharmacology. California: Art of Medicine Press, 2004.
5. Su G, Chen X, Liu Z, et al. Oral astragalus (huang qi) for preventing frequent episodes of acute respiratory tract infection children. *Cochrane Database Syst Rev*, 2016;12:CD011958.
6. Shahzad M, Shabbir A, Wojcikowski K, et al. The antioxidant effects of radix astragali (*Astragalus membranaceus* and related species) in protecting tissues for injury and disease. *Curr Drug Targets* 2016;17(12):1331-1340.
7. Agyemang K, Han L, Liu E, et al. Recent advances in *Astragalus membranaceus* anti-diabetic research: pharmacological effects of its phytochemical constituents. *Evid Based Complement Altern Med* 2013; 2013:654643.
8. Braun L, Cohen M. Herbs & Natural Supplements: an evidence-based guide, vol. 2, 4th ed. Sydney: Elsevier, 2014.
9. Brush J, Mendenhall E, Guggenheim A, et al. The effect of *E. purpurea*, *A. membranaceus* and *G. glabra* on CD69 expression and immune cell activation in humans. *Phytother Res* 2006;20(8):687-695.
10. Denzler K, Moore J, Harrington H, et al. Characterization of the physiological response following in vivo administration of *Astragalus membranaceus*. *Evidence Based Comp Altern Med* 2016;2016:686107811.
11. Rogers ME, Bohlken RM, Beets MW, et al. Effects of creatine, ginseng and astragalus supplementation on strength, body composition, mood and blood lipids during strength training in older adults. *J Sports Sci Med* 2006;5:60-69.
12. Yeh TS, Chuang HL, Huang WC, et al. *Astragalus membranaceus* improves exercise performance and ameliorates exercise-induced fatigue in trained mice. *Molecules* 2014;19(3):2793-2807.
13. Deng ZH, Hu QL. Effect of *Astragalus membranaceus* polysaccharides on oxidative damage in skeletal muscle of exhaustive exercise rats. *African J Agricultural Res* 2011;6(17):4086-4090.
14. Sun Y, Yang J. Experimental study of the effect of *Astragalus membranaceus* against herpes simplex virus type 1. *Di Yi Jun Yi Da Xue Xue Bao* 2004;24(1):57-58.
15. Peng TQ, Yang YZ, Kandolf R. Effect and mechanism of *Astragalus membranaceus* on coxsackie B3 virus RNA in mice. *Zhongguo Zhong Xi Yi Jie He Za Zhi* 1994;12(11):664-666.
16. Guo Q, Sun X, Zhang Z, et al. The effect of astragalus polysaccharide on the Epstein-Barr virus lytic cycle. *Acta Virol* 2014;58(1):76-80.
17. Shang L, Qi Z, Sun L, et al. Astragaloside IV inhibits adenovirus replication and apoptosis in A549 cells in vitro. *J Pharm Pharmacol* 2011;63(5):688-694.
18. Ai P, Yong G, Dingkun G, et al. Aqueous extract of *Astragali radix* induces human natriuresis through enhancement of renal response to atrial natriuretic peptide. *J Ethnopharmacol* 2008;116(3):413-421.
19. Li L, Hou X, Xu R, et al. Research review on the pharmacological effects of astragaloside IV. *Fundam Clin Pharm* 2017;31(1):17-36.
20. Dun C, Liu J, Qiu F, et al. Effects of astragalus polysaccharides on memory impairment in a diabetic rat model. *Neuropsychiatr Dis Treat* 2016;12:1617-1621.
21. He X, Shu J, Xu L, et al. Inhibitory effect of astragalus polysaccharides on LPS-induced TNF- α and IL-1 β production in THP-1 cells. *Molecules*. 2012;17(3):3155-3164.
22. Pu X, Fan W, Yu S, et al. Polysaccharides from angelica and astragalus exert hepatoprotective effects against carbon-tetrachloride-induced intoxication in mice. *Can J Physiol Pharmacol* 2015;93(1):39-43.
23. Lu Y, Xing QQ, Xu JY, et al. Astragalus polysaccharide modulates ER stress response in an OVA-LPS induced murine model of severe asthma. *Int J Biol Macromol* 2016;93(Pt A):995-1006.
24. Lv J, Zhang Y, Tian Z, et al. Astragalus polysaccharides protect against dextran sulfate sodium-induced colitis by inhibiting NF- κ B activation. *Int J Biol Macromol* 2017;98:723-729.
25. Li Q, Bao JM, Li XL, et al. Inhibiting effect of astragalus polysaccharides on the functions of CD4+CD25 high Treg cells in the tumor microenvironment of human hepatocellular carcinoma. *Chin Med J (Engl)* 2012;125(5):786-793.
26. Dai H, Jia G, Liu X, Liu Z, Wang H. Astragalus polysaccharide inhibits isoprenaline-induced cardiac hypertrophy via suppressing Ca²⁺ mediated calcineurin/NFATc3 and CaMKII β signaling cascades. *Environ Toxicol Pharmacol* 2014 Jul; 38 (1): 263-71.
27. Cao Y, Ruan Y, Shen T, et al. Astragalus polysaccharide suppresses doxorubicin-induced cardiotoxicity by regulating PI3k/Akt and p38MAPK pathways. *Oxid Med Cell Longev* 2014;2014:674219.
28. You Y, Duan Y, Liu SW, et al. Anti-atherosclerotic function of *Astragali Radix* extract: downregulation of adhesion molecules in vitro and in vivo. *BMC Complement Altern Med* 2012;12:54.
29. Ren S, Zhang H, Mu Y, et al. Pharmacological effects of astragaloside IV: a literature review. *J Trad Chin Med* 2013;33(3):413-416.
30. Wang Y, Auyeung KK, Zhang X, et al. Astragalus saponins modulates colon cancer development by regulating calpain-mediated glucose regulated protein expression. *BMC Complement Altern Med* 2014;14:401.
31. Xu C, Tang F, Lu M, et al. Astragaloside IV improves the isoproterenol-induced vascular dysfunction via attenuating eNOS uncoupling-mediated oxidative stress and inhibiting ROS-NF- κ B pathways. *Int Immunopharmacol* 2016;33:119-127.
32. Lu Y, Li S, Wu H, et al. Beneficial effects of astragaloside IV against angiotensin II induced mitochondrial dysfunction in rat vascular smooth muscle cells. *Int J Mol Med* 2015;36(5):1223-1232.
33. Liu CH, Tsai CH, Li TC, et al. Effects of the traditional Chinese herb *Astragalus membranaceus* in patients with post-stroke fatigue: a double-blind, randomised controlled preliminary study. *J Ethnopharmacol* 2016;194:954-962.
34. Mills S, Bone K. Principles and Practice of Phytotherapy Edinburgh: Churchill Livingstone, 2000.
35. Monograph: astragalus. Health Canada 2010. Viewed 24 Oct 2017, <http://webprod.hc-sc.gc.ca/nhp/ndp-bdpsn/monoReq.do?id=>

BioCeuticals®

A division of FIT-BioCeuticals Limited
Unit 1/Level 1, 85 O'Riordan Street
Alexandria NSW 2015
(+61) 2 9080 0900
www.bioceuticals.com.au