

ADAPTOGENIC MUSHROOM BOOST

CLINICAL APPLICATIONS

Adaptive, Intelligent Support to Modulate and Strengthen the Immune Response
Strengthens the Body, Reduces Fatigue and Improves Stress Resilience

Helps Maintain Normal Inflammatory Balance
Helps Train the Immune System to Eliminate Unwanted Organisms

The use of mushrooms for enhancing physical well-being has a history of human use spanning millennia. This product features a powerful combination of six mushroom extracts with an extensive history of use in supporting human health. This targeted blend of mushrooms contains specialized adaptogenic compounds known as biological response modifiers (BRMs). These unique compounds intelligently modulate immune function and strengthen the immune response to prolonged stress, fatigue and microbial challenges. BRMs actively balance and train the immune system to maintain normal inflammatory balance, respond effectively against environmental toxins, and eliminate unwanted microbes. In addition, this formula is safe and effective for everyday use in a broad range of immune challenges.

Overview

Mushrooms are fungi with spore-producing, umbrella-shaped fruiting bodies and white thread-like structures called mycelia. Mushrooms have a number of biologically active compounds, although the most studied are the various polysaccharides found in the fruiting bodies and cultured mycelia.^{1,2} These polysaccharides are mostly glucans, with glycosidic bonds in either the axial position (α -glucans) or in the equatorial position (β -glucans). In addition to polysaccharides, mushrooms also have other bioactive compounds, such as glycoproteins, terpenoids, sterols and phenolic compounds.³⁻⁵ Mushroom compounds act as biological response modifiers that intelligently modulate immune

cells like macrophages, natural killer (NK) cells and T lymphocytes. Data suggests mushroom polysaccharides and other bioactive compounds can establish Th1 to Th2 immune cell balance, strategically supporting immune function without overstimulation.

Shiitake (Lentinula edodes)

Shiitake is an edible mushroom native to Asia that has been cultivated as both a food and health tonic for many centuries. Both α - and β -glucans derived from shiitake are well-studied bioactive compounds. In Japan, alpha glucans isolated from shiitake are commonly used as supplements.⁶ These alpha glucans have been found to modulate immune function by enhancing the immune responses of both CD4 and CD8 T cells.⁷ Supplementation with shiitake alpha glucans also resulted in a 2.5-fold increase in NK cell activity.⁸ Lentinan is a high molecular weight β -glucan isolated from shiitake mushroom with bioactive properties. Animal studies on the oral administration of lentinan found significantly higher circulating T cell levels in the lentinan group than the control group after four weeks.⁹ Additional studies of shiitake mushroom consumption have shown decreases in C-reactive protein (CRP), a marker of normal inflammatory balance, and increases in secretory IgA (slgA), indicating improved mucosal immunity.¹⁰

Reishi (Ganoderma lucidum)

Reishi mushroom is one of the most widely used mushrooms in the world with a 2000-year-long history of use in traditional

† These statements have not been evaluated by the Food and Drug Administration. This product is not intended to diagnose, treat, cure, or prevent any disease.



8952 E. Desert Cove Ave | Suite E114 Scottsdale, AZ 85260 480-878-0087 @drkristineromine krmdskin.com

Chinese medicine, and it is referred to as ling zhi, or the "mushroom of immortality."^{11,12} Research on mice has found reishi extract maintains normal inflammatory balance.¹³ Reishi mushroom polysaccharide extract supplementation also demonstrated enhanced immune status with increases in NK cell activity and Th1 and Th2 balancing properties.^{14,15} The results of the meta-analysis of five randomized controlled trials showed that patients who had been given reishi mushroom supplements exhibited an increase in the percentage of CD4 and CD8 cells.¹⁶ Several studies have shown that patients in the reishi group had relatively improved quality of life compared to controls. Improved quality of life relating to decreases in plasma concentrations of cytokines $\text{TNF-}\alpha$ and IL-1 was observed in 73.2% of the patients receiving reishi mushroom polysaccharide supplements for 12 weeks.¹⁷ An increase in the counts of CD3, CD4, CD8 and CD56 lymphocytes and NK cell activity was also observed, suggesting a broad immunemodulatory effect of reishi-derived polysaccharides.¹⁸

Chaga (Inonotus obliquus)

Chaga has been used as a folk medicine in Russia and Western Siberia since the 16th century.¹⁹ Chaga grows as a conk (knoblike mass) on the side of birch trees. It contains both betulin and betulinic acid that it metabolizes from the birch tree bark on which it grows. These compounds are effective free radical scavengers that can induce apoptosis in damaged cells.²⁰ Chaga grown on birch displayed activity against a number of human and animal intracellular microbes, which is attributed mainly to betulin, lupeol and sterol content.²¹ Chaga supplementation also resulted in a 54.9% reduction of DNA damage in lymphocytes while also reducing oxidative stress in lymphocytes in vitro.²² In a recent study, chaga extract was found to have α -amylase and α -glucosidase-inhibiting properties. The extract also demonstrated significant free radical scavenging activity and was found to decrease H₂O₂ -induced oxidative damage in hepatic L02 cells.²³

Cordyceps militaris

Cordyceps sinensis grows as a parasitic fungus on caterpillars in the wild. The main bioactive constituent in *Cordyceps* is cordycepin, an adenosine derivative with immunoregulatory properties.²⁴ *Cordyceps militaris* can be more easily masscultivated and has higher levels of cordycepin than *Cordyceps sinensis*.²⁵ Over 200 clinical studies show *Cordyceps* mushrooms improve performance by increasing blood flow, boosting ATP synthesis for more natural energy, and acting as a potent antioxidant to decrease oxidative stress and fatigue.²⁶ A study conducted in Japan with men who were supplementing with *Cordyceps* showed significant increases in the concentrations of creatine as well as catecholamines over the course of two weeks.²⁷ In patients with immune challenges, supplementation with *Cordyceps* mushroom powder helped to balance the activity of the cytokine IL-10.²⁸

Pearl Oyster (Pleurotus ostreatus)

Pearl oyster mushroom is an edible mushroom that is cultivated globally as a vegetarian protein source. Oyster mushrooms are also a source of B vitamins, vitamin C, vitamin D and several minerals.²⁹ They are also a source of glutathione and the fungal antioxidant ergothioneine.³⁰ Pleuran, an insoluble β -glucan isolated from this mushroom, has been shown in athletes to significantly reduce the incidence of respiratory challenges and increase the number of circulating NK cells.³¹ Oyster mushroom has also been found to support turnover of cholesterol in the plasma and liver and help maintain normal cholesterol levels.³² Additionally, a novel ubiquitin-like protein from oyster mushrooms was found to inhibit reverse transcriptase, which is important for immune defense.³³

Turkey Tail (Trametes versicolor)

Turkey tail is one of the best documented mushrooms and the most common polypore mushroom to grow on hardwood. PSK (polysaccharide-K) and PSP (polysaccharopeptide), two protein-bound polysaccharides isolated from turkey tail mushroom mycelia, have been used in clinical trials in Japan since 1970. As a biological response modifier, PSP has been found to positively modulate the immune system and increase T cell proliferation.³⁴ Immune cell production and guality of life scores were also enhanced in both PSK- and PSPsupplemented groups.³⁵ In recent studies, PSP was found to improve pain tolerance by binding to the CB2 receptor and upregulating levels of β -endorphin while reducing levels of IL-1, NO and PGE2.³⁶ Clinical trial results also demonstrated that supplementation with turkey tail mushroom powder increased lymphocyte count and NK cell activity, and CD8+ T cells and CD19+ B cells increased in a dose-dependent fashion.³⁷

Directions

2 capsules per day or as recommended by your health care professional.

Does Not Contain

Gluten, corn, yeast, artificial colors or flavors.

Cautions

If you are pregnant or nursing, consult your physician before taking this product.

Servings Per Container 30	Amount Per Serving	% Daily Value
Calories	5	
Total Carbohydrate	1 g	<1%*
Proprietary Blend	1 g	
Chaga (Inonotus obliquu (Sterile Conk; Canker)	s) Extract (Organic)	**
Reishi <i>(Ganoderma lucio</i> Powder (Fruit Body) (0	<i>lum)</i> Extract Drganic)	**
Cordyceps (Cordycepos (Fruit Body) (Organic)	militaris) Extract	**
Turkey Tail <i>(Trametes ve</i> (Mycelia, Primordia, Fl Compounds) (Organic	e <i>rsicolor)</i> Powder ruit Body, Extracellu)	ular **
Pearl Oyster <i>(Pleurotus o</i> (Mycelia, Primordia, Fl Compounds) (Organic	o <i>streatus)</i> Powder ruit Bodies, Extrace	** Ilular
Shiitake (Lentinula edode (Fruit Body) (Organic)	es) Extract Powder	**

Other Ingredients: Hypromellose (Natural Vegetable Capsules), Microcrystalline Cellulose, Silicon Dioxide and Magnesium Stearate.

References

- 1. Wasser SP. Medicinal mushrooms as a source of antitumor and immunomodulating polysaccharides. *Appl Microbiol Biotechnol*. Nov 2002;60(3):258-274.
- 2. Lull C, Wichers HJ, Savelkoul HF. Anti-inflammatory and immunomodulating properties of fungal metabolites. *Mediators Inflamm*. Jun 9 2005;2005(2):63-80.
- 3. Lindequist U, Niedermeyer TH, Julich WD. The pharmacological potential of mushrooms. *Evid Based Complement Alternat Med*. Sep 2005;2(3):285-299.
- 4. Daba A, Ezeronye O. Anti-cancer effect of polysaccharides isolated from higher basidiomycetes mushrooms. *African Journal of Biotechnology*. 2003;2(12):672-678.
- 5. Ooi VE, Liu F. Immunomodulation and anti-cancer activity of polysaccharide-protein complexes. *Curr Med Chem*. Jul 2000;7(7):715-729.

- 6. Shah SK, Walker PA, Moore-Olufemi SD, Sundaresan A, Kulkarni AD, Andrassy RJ. An evidence- based review of a *Lentinula edodes* mushroom extract as complementary therapy in the surgical oncology patient. *JPEN J Parenter Enteral Nutr.* Jul 2011;35(4):449-458.
- 7. Ritz B. Active hexose correlated compound (AHCC) and immune outcomes in humans: A review. *Natural Medicine Journal*. 2011;3(1):3-7.
- 8. Ghoneum M, Wimbley M, et al. Immunomodulatory and anticancer effects of active hemicellulose compound (AHCC). *Int.J. Immunotherapy*. 1995;11(1):23-28.
- Hanaue H, Tokuda Y, Machimura T, et al. Effects of oral lentinan on T-cell subsets in peripheral venous blood. *Clinical Therapeutics*. 1989 Sep-Oct;11(5):614-622. PMID: 2529966.
- Dai X, Stanilka JM, Rowe CA, et al. Consuming *Lentinula edodes* (Shiitake) Mushrooms Daily Improves Human Immunity: A Randomized Dietary Intervention in Healthy Young Adults. *Journal of the American College of Nutrition*. 34 (6): 478-487. doi:10.1080/07315724.2014.950391.
- 11. Yuen JW, Gohel MD. Anticancer effects of *Ganoderma lucidum*: a review of scientific evidence. *Nutr Cancer*. 2005;53(1):11-17.
- 12. Cheuk W, Chan JK, Nuovo G, Chan MK, Fok M. Regression of gastric large B-Cell lymphoma accompanied by a florid lymphoma-like T-cell reaction: immunomodulatory effect of *Ganoderma lucidum* (Lingzhi). *Int J Surg Pathol.* Apr 2007;15(2):180-186.
- Stavinoha, W.B. Status of *Ganoderma lucidum* in the United States: *Ganoderma lucidum* as an anti-inflammatory agent. Proceedings of the 1st International Symposium on *Ganoderma lucidum* in Japan. Nov 17-18 1991. 99-103. Tokyo.
- 14. Gao Y, Zhou S, Jiang W, Huang M, Dai X. Effects of ganopoly (a *Ganoderma lucidum* polysaccharide extract) on the immune functions in advanced-stage cancer patients. *Immunol Invest*. Aug 2003;32(3):201-215.
- 15. Gao Y, Tang W, Dai X, et al. Effects of water-soluble *Ganoderma lucidum* polysaccharides on the immune functions of patients with advanced lung cancer. *J Med Food*. Summer 2005;8(2):159-168.

- 16. Jin X, Ruiz Beguerie J, Sze DM, Chan GC. *Ganoderma lucidum* (Reishi mushroom) for cancer treatment. *Cochrane Database Syst Rev.* 2012;6:CD007731.
- 17. Oka S, Tanaka S, Yoshida S, et al. A water-soluble extract from culture medium of *Ganoderma lucidum* mycelia suppresses the development of colorectal adenomas. *Hiroshima J Med Sci.* Mar 2010;59(1):1-6.
- 18. Chen X, Hu ZP, Yang XX, et al. Monitoring of immune responses to a herbal immuno-modulator in patients with advanced colorectal cancer. *Int Immunopharmacol*. Mar 2006;6(3):499-508.
- 19. Saar M. Fungi in Khanty folk medicine. *J Ethnopharmacology*. 1991;31(2):175-179.
- 20. Faass N. The healing powers of wild chaga; an interview with Cass Ingram, MD. *Price-Pottenger Journal of Health and Healing*. 2012;35(4):6-11.
- 21. Kahlos K, Lesnau A, et al. Preliminary tests of antiviral activity of two *Inonotus obliquus* strains. *Fitoterpia*. 1996;67(4):344-347.
- 22. Najafzadeh M, et al. Chaga mushroom extract inhibits oxidative DNA damage in lymphocytes of patients with inflammatory bowel disease. *Biofactors*. 2007;31(3-4):191-200.
- 23. Gao X, Santhanam RK, Xue Z, Jia Y, et al. Antioxidant, α-amylase and α-glucosidase activity of various solvent fractions of *l. obliquus* and the preventive role of active fraction against H2O2 induced damage in hepatic L02 cells as fungisome. *J Food Sci.* 2020 Apr;85(4):1060-1069. doi: 10.1111/1750-3841.15084. Epub 2020 Mar 9. PMID: 32147838.
- 24. Zhou X, Luo L, Dressel W, et al. Cordycepin is an immunoregulatory active ingredient of *Cordyceps sinensis*. *Am J Chin Med*. 2008;36(5):967-980.
- 25. Kim H.O., Yun J.W. A comparative study on the production of exopolysaccharides between two entomopathogenic fungi *Cordyceps militaris* and *Cordyceps sinensis* in submerged mycelial cultures. *Journal of Applied Microbiology*. 2005; 99(4):728-38.
- 26. Hrisch K.R., Smith-Ryan A.E., Roelofs E.J., Trexler E.T., Mock M.G. *Cordyceps militaris* improves tolerance to high intensity exercise after acute and chronic supplementation. *Journal of Dietary Supplements*. 2017 Jan 2; 14(1): 42–53.

- 27. Nagata A., Tajima T., Uchida M. Supplemental anti-fatigue effects of *Cordyceps sinensis* (Tochu-Kaso) extract powder during three step wise exercise in human. *Japan Journal of Physical Fitness in Sports. 2006;5:* 145–52
- 28. Ding C, Tian PX, Xue W, et al. Efficacy of *Cordyceps sinensis* in long term treatment of renal transplant patients. *Front Biosci* (Elite Ed). 2011;3:301-307.
- 29. Wani B.A., Bodha R.H., Wani A.H. Nutritional and medicinal importance of mushrooms. *Journal of Medicinal Plants Research* Vol. 4(24), pp. 2598-2604.
- Kalaras MD, Richie JP, Calcagnotto A, Beelman RB. Mushrooms: Arich source of the antioxidants ergothioneine and glutathione. *Food Chem.* 2017 Oct 15;233:429-433. doi: 10.1016/j.foodchem. 2017.04.109. Epub 2017 Apr 20. PMID: 28530594.
- 31. Majtan J. Pleuran (beta-Glucan from *Pleurotus ostreatus*): An Effective Nutritional Supplement against Upper Respiratory Tract Infections. *Med Sport Sci.* 2013;59:57-61.
- 32. Bobek P, Ozdin O, Mikus M. Dietary oyster mushroom (*Pleurotus ostreatus*) accelerates plasma cholesterol turnover in hypercholesterolemic rat. *Physiological Research.* 1995;44 (5): 281-291.
- 33. Wang HX, Ng TB. Isolation of a novel ubiquitin-like protein from *Pleurotus ostreatus* mushroom with anti-human immunodeficiency virus, translation-inhibitory, and ribonuclease activities. *Biochem Biophys Res Commun.* 2000;276(2):587-593.
- 34. Ng TB. A review of research on the protein-bound polysaccharide (polysaccharopeptide, PSP) from the mushroom *Coriolus versicolor* (Basidiomycetes: Polyporaceae). *Gen Pharmacol*. 1998;30(1):1-4. doi:10.1016/s0306-3623(97)00076-1
- 35. Kidd PM. The use of mushroom glucans and proteoglycans in cancer treatment. *Altern Med Rev.* Feb 2000;5(1):4-27.
- 36. Wang K, Wang Z, Cui R, Chu H. Polysaccharopeptide from Trametes versicolor blocks inflammatory osteoarthritis pain-morphine tolerance effects via activating cannabinoid type 2 receptor. *Int J Biol Macromol.* 2019;126:805-810. doi:10.1016/j.ijbiomac.2018.12.212
- 37. Torkelson CJ, Sweet E et al. Phase 1 clinical trial of *Trametes versicolor* in women with breast cancer. *ISRN Oncol.* 2012:10.5402/2012/251632.

+ These statements have not been evaluated by the Food and Drug Administration. This product is not intended to diagnose, treat, cure, or prevent any disease.