

# Agaricus Bisporus SAP

Science-based immune support for optimal health

**Agaricus Bisporus SAP** is a hot water-extract from white button mushroom (WBM; *Agaricus bisporus*), the most commonly consumed mushroom in North America and most Western countries. *A. bisporus* constitutes of a significant amount of vitamin D precursor ergosterol, dietary fibres, and antioxidants including vitamins C, B<sub>12</sub>, folates, and polyphenols. In addition, lectins from *A. bisporus* exert various beneficial effects including their role in improving the chemotherapeutic efficacy of cancer treatments, and their ability in inhibiting colon cancer-cell proliferation and enhancing the cellular antioxidant defense mechanisms. As a potent aromatase inhibitor and immunomodulatory agent in increasing natural killer-cell activity and promoting adaptive immune responses, *A. bisporus* extracts have gained recent attention for their potential value in breast and prostate cancer support. Also, studies support the application of *A. bisporus* extracts in type 2 diabetes, metabolic syndrome, cholesterol management, and cardiovascular disease (CVD). Early-stage preclinical evidence suggests the physiological benefits of *A. bisporus* in improving working memory and balance during aging. **Agaricus Bisporus SAP** can be used as an adjunctive support in the treatments of cancers, diabetes, and cholesterol management, as well as in CVD risk alleviation.

## ACTIVE INGREDIENTS

Each capsule contains:

*Agaricus bisporus* 5:1 extract,  
30% polysaccharides providing 15% β-glucans . . . . . 667 mg

**NON-MEDICINAL INGREDIENTS:** Vegetable magnesium stearate and silicon dioxide in a capsule made of vegetable carbohydrate gum and purified water.

**Contains no:** Gluten, soy, wheat, corn, eggs, dairy, yeast, citrus, preservatives, artificial flavour or colour, starch, or sugar.

**This product is non-GMO and vegan friendly.**

**Agaricus Bisporus SAP** contains 90 capsules per bottle.

## DIRECTIONS FOR USE

**Adults:** Take 3 capsules daily or as directed by your healthcare practitioner.

## INDICATIONS

**Agaricus Bisporus SAP** is a source of antioxidants, and can be used as an adjunctive support in breast, prostate, ovary, and colon cancer therapy, and can help:

- Promote healthy inflammatory responses.
- Support healthy immunomodulation.
- With the management of cholesterol and type 2 diabetes, and alleviate CVD risk factors.
- Improve aging-associated loss in working memory and balance.

## CAUTIONS AND WARNINGS

Consult a healthcare practitioner prior to use if you are pregnant or breast-feeding, or if you have diabetes. Do not use if seal is broken. Keep out of reach of children.

## PURITY, CLEANLINESS, AND STABILITY

All ingredients listed for each product lot number of **Agaricus Bisporus SAP** have been tested by an ISO 17025-accredited third-party laboratory for identity, potency, and purity.



Scientific Advisory Panel (SAP):  
adding nutraceutical research  
to achieve optimum health



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## INTRODUCTION

*Agaricus bisporus* is the most commonly consumed mushroom in North America and most Western countries.<sup>[1]</sup> *A. bisporus* constitutes of a significant amount of vitamin D precursor, ergosterol, which usually when exposed to ultraviolet irradiation results in vitamin D<sub>2</sub>. *A. bisporus* contains lectins that have been shown to improve the efficacy of chemotherapeutic treatments by increasing the sensitivity of lung, colon, and glioblastoma cancerous cells to the treatment.<sup>[2][3]</sup> Studies have also demonstrated the ability of lectins from *A. bisporus* in inhibiting colon cancer-cell proliferation and enhancing the cellular antioxidant defense mechanisms.<sup>[2][3]</sup>

## IMMUNOMODULATION

Innate and adaptive immunity are two different yet closely linked components of the immune system. Innate immune response is a fully functional first-line defense system in fighting against invading microorganisms, while the adaptive immunity, i.e. cell-mediated and humoral immunity, ensures a more efficient and sustained response in fighting these invading microorganisms by developing a memory to prevent future infections by the same microorganisms. Antigen-presenting cells (APC) represent an important link between these two components that functions by recognizing, taking up, processing, and eventually presenting a variety of foreign antigens to T cells to initiate the adaptive immune response. The dendritic cells (DC), macrophages, and B cells are three crucial professional APC.<sup>[4][5][6][7][8]</sup>

Dietary supplementation of *A. bisporus* has been shown to increase natural killer-cell activity,<sup>[9][10]</sup> and to promote antigen-presenting function and maturation of DC.<sup>[11]</sup> In one randomized, controlled clinical study with 24 healthy volunteers, participants consuming a normal diet with 100 g of blanched *A. bisporus* showed significant increases in salivary secretory IgA (sIgA) levels compared to those consuming the normal diet without *A. bisporus*.<sup>[12]</sup>

## CANCER

Substantial evidence shows that *A. bisporus* extracts have beneficial effects on various kinds of cancers.

### Breast Cancer

One of the major factors in the development of breast cancer is estrogen. Especially, estrogen production by aromatase/estrogen synthetase in breast cancer is related to increased tumour proliferation.<sup>[13]</sup> *A. bisporus* extracts have been shown to inhibit aromatase activity effectively, particularly in an aromatase-transfected breast cancer-cell line, and suppress breast cancer-cell proliferation.<sup>[13]</sup> Another study in mice showed that *A. bisporus* extracts decreased breast cancer tumour weight and cell proliferation without affecting the rate of apoptosis.<sup>[14]</sup> Conjugated linoleic acid and its derivatives are considered to be the physiologically relevant aromatase inhibitors in *A. bisporus* extracts.<sup>[14]</sup> In addition, microarray analysis of tumours in *A. bisporus* extract-fed mice displayed significant changes in gene expression, especially gene networks that are involved in cell death, growth, and proliferation; lipid metabolism; the tricarboxylic acid cycle; and immune response.<sup>[13][14]</sup> One early-stage clinical study in postmenopausal breast-cancer survivors who were cancer-free after completion of cancer therapy has lent clinical evidence on the breast cancer-preventive role of *A. bisporus* extracts.<sup>[15]</sup> In this study, groups were supplemented with a 12-week course of *A. bisporus* extract at 5, 8, 10, or 13 g doses. The study results showed a modest aromatase inhibition, and *A. bisporus* extract was well-tolerated at all doses.<sup>[15]</sup> The findings of this study suggest the need for a much higher dosage to attain a clinically significant effect. Hence, more clinical studies using *A. bisporus* extracts are warranted to validate the preclinical and early-pilot clinical studies.

### Prostate Cancer

Serum prostate-specific antigen (PSA) is a useful biomarker for the diagnosis of organ-confined prostate cancer.<sup>[16]</sup> A small elevation in PSA level ( $\geq 0.2$  ng/mL) is often considered a sign of persistent disease, usually correlated to subsequent disease progression.<sup>[17]</sup> In a recent phase I clinical study, patients with continuously rising PSA levels were enrolled with the primary objective of testing the treatment effects of *A. bisporus* extracts and associated toxicity.<sup>[1]</sup> In addition, the study evaluated the effect of *A. bisporus* extract on serum PSA/androgen levels, myeloid-derived suppressor cells (MDSCs), and cytokine levels.<sup>[1]</sup> The study results demonstrated an overall PSA response rate of 11%, and two patients receiving 8 and 14 g/d of *A. bisporus* extract showed complete response (PSA declined to undetectable levels that continued for 49 and 30 months). Two more patients who received 8 and 12 g/d experienced partial response. Patients with complete and partial response also displayed higher baseline levels of IL-15 than nonresponders, where a decline in MDSC was observed.<sup>[1]</sup> This study suggests that *A. bisporus* extracts could be a valuable therapeutic option for the treatment of biochemically recurrent prostate cancer.

### Ovarian Cancer

Ovarian cancer, a common gynecological malignancy, is predominantly epithelial in origin.<sup>[18]</sup> In a hospital-based case-control study that investigated the association between mushroom consumption and epithelial ovarian cancer risk in Southern Chinese women, mushroom intake, particularly *A. bisporus* intake, was shown to be inversely related to cancer risk.<sup>[18]</sup>

## DIABETES AND LIPID/CHOLESTEROL METABOLISM

*A. bisporus* contains high levels of dietary fibres and antioxidants, including vitamins C, D, and B<sub>12</sub>; folates; and polyphenols that could be beneficial for cardiovascular diseases and diabetes.<sup>[19]</sup> In an animal study, rats fed *A. bisporus* powder for three weeks showed significant reductions in plasma glucose and triglyceride (TG) levels, liver enzyme activities, and liver weight gain.<sup>[20]</sup> Especially in diet-induced hypercholesterolemic rats, *A. bisporus* supplementation for four weeks resulted in a significant decrease in plasma total cholesterol and low-density lipoprotein cholesterol, hepatic cholesterol, and TG levels, with a simultaneous increase in plasma high-density lipoprotein levels.<sup>[19]</sup> A recent retrospective study determined the impact of *A. bisporus* consumption on type 2 diabetes risk factors in 37 racially diverse adults with metabolic syndrome.<sup>[20]</sup> The subjects studied had participated in a dietary intervention focused on vitamin D bioavailability from *A. bisporus* (100 g/d). The study showed significant beneficial health effects at 16 weeks, including elevation in the antioxidant marker ORAC (oxygen radical absorption capacity) and adiponectin, and significant decreases in serum oxidative stress-inducing factors carboxymethyllysine and methylglyoxal.<sup>[20]</sup> However, more well-controlled studies are required to validate and confirm these research outcomes.

Ergosterol-enriched extracts from *A. bisporus* have been shown to lower hepatic triglyceride and modify the mRNA expression of cholesterol-related genes.<sup>[21]</sup> In another study, researchers observed up to 60% inhibition of HMGCR from methanol-water *A. bisporus* extracts using an in vitro assay.<sup>[22]</sup>

## BALANCE AND WORKING MEMORY

The potential benefits of *A. bisporus* supplementation in aging and balance was explored in a recent animal study.<sup>[23]</sup> This study investigated the effects of dietary mushroom intervention on mobility and memory in aged rats. Rats on the 2% or 5% mushroom-supplemented diet for eight weeks consumed more food, without gaining weight, than rats in the other diet groups. In addition, these rats showed an improvement in balance.<sup>[23]</sup> Rats on the 0.5% mushroom diet showed improved performance in a working-memory version of the Morris water-maze. The study results suggested that the most effective mushroom dose that produced improvements in both balance and working memory was 0.5%, equivalent to about 1.5 ounces of fresh mushrooms for humans. Although human clinical evidence is required to confirm these findings, the results suggest potential beneficial effects on age-related deficits in cognitive and motor function.<sup>[23]</sup>

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