# **ProBio SAP**

Science-based probiotics; high dose, multistrain in an enteric capsule

Probiotics are dietary microbial mixtures that beneficially affect the host by improving intestinal microbial balance. Although large numbers of microbes normally inhabit the human intestine, certain strains of bacteria are believed to alleviate the symptoms of lactose intolerance, increase natural resistance to infectious diseases of the intestinal tract, improve digestion, maintain healthy cholesterol levels, reduce risk of cancer, and stimulate gastrointestinal immunity.

### **ACTIVE INGREDIENTS**

#### Each enteric vegetable capsule contains:

Lacticaseibacillus rhamnosus R0011	4.4 billion CFL	J
Lacticaseibacillus rhamnosus R1039	.905 billion CFL	J
Lactobacillus helveticus R0052		
Lactiplantibacillus plantarum R1012	440 million CFL	J
Lacticaseibacillus casei R0215	440 million CFL	J
Bifidobacterium longum R0175	330 million CFL	J
Bifidobacterium infantis R0033		
Bifidobacterium breve R0070		
Streptococcus salivarius ssp. thermophilus R0083	220 million CFL	J
Streptococcus salivarius ssp. thermophilus R0083	55 million CFL	J

Other ingredients: Vegetable magnesium stearate, ascorbic acid, inulin, arabinogalactan, potato starch, saccharose (sucrose), skim milk, maltodextrin, yeast extract (peptone), trehalose and sodium ascorbate, in a capsule composed of hypromellose (vegetable carbohydrate gum), hypromellose acetate succinate, and purified water.

#### This product is non-GMO and vegetarian friendly.

Contains no: Gluten, wheat, eggs, citrus, preservatives, artificial flavour or colour.

Made from 10 specific strains of live cells, chicory root and larch tree (prebiotics).

ProBio SAP (probiotics) contains 90 or 180 enteric capsules per bottle.

Keep refrigerated.

#### **DIRECTIONS FOR USE**

Adults, adolescents, and children ≥ 6 years old: Take 1–2 capsules daily with water or juice or as directed by your healthcare practitioner.

1 capsule provides 11 billion live probiotic cells.

### **INDICATIONS**

Supplementing 1 **ProBio SAP** capsule daily provides doses of live probiotic cells scientifically supported to effectively reduce risk of diarrhea, enhance the immune system, prevent infections, and maintain a good balance of intestinal microorganisms (colon health). **ProBio SAP** may help in management of symptoms of depression and improvement of blood serum lipids.

### **FEATURES**

- ProBio SAP supplies a compatible and complementary blend of 10 probiotic strains to address a
  wide spectrum of gastrointestinal health benefits.
- Enteric-coated vegetable capsule ensures 100% delivery of live microbial cells to the intestine, and is resistant to gastric and bile acids.
- · Fructooligosaccharides and arabinogalactan are prebiotics included in **ProBio SAP** to enhance *Bifidobacterium* and *Lactobacillus* proliferation and implantation, stimulate mineral (calcium and magnesium) absorption, and enhance production of short-chain fatty acids.

## **PURITY, CLEANLINESS, AND STABILITY**

All ingredients listed for all **ProBio SAP** lot numbers have been tested by a third-party laboratory for identity, potency, and purity.



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



351, Rue Joseph-Carrier, Vaudreuil-Dorion, Quebec, J7V 5V5 T 1 866 510 3123 • F 1 866 510 3130 • nfh.ca

# **ProBio SAP**

# Research Monograph

#### WHAT ARE PROBIOTICS?

Probiotics can be defined as cultures of living micro-organisms in mized strain or single strain form, which when administered in certain amount can ameliorate gut microbiota and provide beneficial effects to the host beyond their nutritional potential.[1] Lactobacillus and Bifidobacterium are two main probiotic bacteria which have shown several health benefits when administered in the required amounts.[2]

#### WHAT ARE PREBIOTICS?

Prebiotics are food or supplements that contain indigestible fibres that enhance the growth and proliferation of probiotic bacteria. Examples of prebiotics include fructo-oligosacharides, large polysaccharides such as resistant starches and inulin, lactulose etc. A combination of probiotics and prebiotics is known as synbiotics, which are used for their synergistic effect in providing and proliferating healthy gut microbes. A more beneficial effect has been observed with the use of mixed strain bacteria rather than use of single strains.[1]

#### PROBIOTICS — GENERAL DOSE AND DURATION

Numerous studies that have evaluated the use of probiotics on physiological effects in humans, such as in the treatment of lactose intolerance, diarrhea, and colon cancer biomarkers, recommend ingesting a daily dose of 109-1010 live bacteria. Most ingested probiotics are transient, pass through the intestinal tract in 3-30 days, and do not permanently adhere to the intestinal wall. These probiotics exert their effects as they proliferate and metabolize while in the small intestine and colon. Probiotics must be supplemented regularly to maintain their effectiveness. Different strains of probiotic bacteria exert different effects on human health. Thus, proven effects of one strain or species cannot be transferred to others. [3] As a result, it is best to supplement with a high-dose multistrain probiotic to provide an optimal range of health benefits as well as suit the needs of different individuals.

#### SAFETY OF PROBIOTICS SUPPLEMENTATION

The safety record of probiotics is excellent, with Lactobacilli and Bifidobacteria designated as generally recognized as safe (GRAS).[1] The lack of pathogenicity with probiotic use extends to all age groups and to immunocompromised individuals.[5] While theoretically, as living organisms, probiotics may be responsible for side effects in susceptible individuals including infections, deleterious metabolic activities, excessive immune stimulation and gene transfer, [6] Lactobacilli and Bifidobacteria probiotics are extremely rare causes of infections in humans and have not led to an increase in such opportunistic infections in consumers. On these bases, probiotics are considered safe for human consumption.

#### PROBIOTICS AND NUTRITION RESEARCH

#### Mechanisms of Action[7]

Several studies have characterized the mechanism of action of probiotics. Some of these mechanisms include competitive exclusion of enteric pathogens (via adhesion sites and nutrients), inhibition of pathogen growth by producing lactic acid, butyric acid, bacteriocins, hydrogen peroxide etc., triggering cytokine synthesis from enterocytes by attaching to their surface, restoration of the normal intestinal flora and immunomodulation.[7]

#### **Enhancement of the Immune System**

H Recent meta analysis of clinical trials conducted have shed better light on the role of probiotics in immunomodulation. Systematic meta analysis of 19 clinical trials supports evidence that probiotic supplementation during pregnancy and lactation may reduce risk of eczema and development of immune-mediated diseases in the child.[8] A similar meta analysis of 8 clinical trials indicates probiotic supplementation may reduce the occurrence of respiratory tract diseases, some of which used specific strains such as Lactoacillus casei and strains of Bifidobacterium.[9] Recent evidence also suggests probiotics can help in management of HIV-1 infection by combating bacterial vaginosis in HIV-1 positive women, stabilizing CD4+ T cell counts and enhancing growth in HIV-1 positive children and infants. Probiotics also showed potential with anti-inflammatory and immunomodulatory effects on the gastrointestinal immune system.[10]

#### **Promotion of Gut Health**

The role of probiotics in gut health has been well established, and recent

studies have strengthened this understanding by shedding light on the impact of probiotics on specific gut disorders. Meta analysis of randomized clinical trials revealed 3 clinical trials that showed significant positive association between probiotics and inflammatory bowel disease in children (IBD). The same analysis also revealed an overview of 18 clinical trials showed beneficial effects of probiotic supplementation in patients with ulcerative colitis (UC), specifically combination of Lactobacillus.[11] The far-reaching effects of probiotic supplementation beyond the gut are being explored as well. A summary of clinical studies points to potential benefits of probiotics in the treatment of halitosis, periodontitis, and reduced incidence of dental caries.[12] It has been observed that probiotics through gut microbiota modulation, may show therapeutic potential against chronic liver diseases such as non-alcoholic fatty liver disease (NAFLD), fibrosis progression and liver cirrhosis.[13] Probiotic supplementation could also help alleviate symptoms of lactose intolerance.[14]

#### **Effect on Depression**

Growing evidence points to the importance of probiotic supplementation and their role in management of gut-brain relationship. A recent metaanalysis summarizing the results of clinical randomized clinical trials showed that probiotics significantly reduced the depression scale score in participants.[15]

#### Effect on cysts and tumor growth

Numerous animal and in vitro studies support the use of probiotics as an adjuvant therapy for treatment of tumor growths and different cancers, however further research via clinical trials will be required to strengthen this evidence and understand the mechanism behind it.[16] A randomized, double-blind, placebo controlled trial consucted with polycystic ovary syndrome (PCOS) showed that probiotic supplementation for 12 weeks significantly increased serum sex hormone binding globulin (SHBG) and had beneficial effects on plasma total antioxidant capacity, serum testosterone, C-reactive protein, and plasma malondialdehyde concentrations (n=30).[17] A combination of fiber and probiotics was significantly effective in treatment of diarrhea associated with enteral nutrition in gastric cancer patients.[18]

#### Modulation of serum lipids

A systematic meta-analysis of randomized controlled trials has revealed probiotic supplementation can significantly reduce serum total cholesterol. [19] A meta-analysis of 12 clinical trials looking at the effect of probiotics on type 2 diabetes patients has shown significant effect of probiotics in lowering serum total cholesterol, triglycerides, as well as elevation of HDL-C. The meta-analysis also showed an improvement in glucose control via reduced fasting blood glucose and serum insulin concentration. [20]

- Zheng H.J., et. al. The effect of probiotic and synbiotic supplementation on biomarkers of inflammation and oxidative stress in diabetic patients: A systematic review and meta-analysis of randomized controlled trials. Pharmacol Res, 2019, 142:303-313 Lei W.T., et.al. Effect of probiotic and prebiotics on immune response to influenze vaccination in adults: a
- systematic review and meta-analysis of randomized controlled trials. Nutrients. 2017, 9(11)
- Schrezenmeir, J. and de Vrese M. Probiotics, prebiotics, and synbiotics approaching a definition. Am Journal of Clin Nutr. 2001, 73(2):3615-3645
  Saarela, M., et al. Gut bacteria and health foods — the European perspective. Int J of Food Microbio. 2002,
- 78(1-2):99-117
- Borriello, S.P., et al. Safety of probiotics that contain lactobacilli or bifidobacteria. Clin Infect Diseases. 2003,
- Marteau, P. and F. Shanahan. Basic aspects and pharmacology of probiotics: an overview of pharmacokinetics, mechanisms of action and side-effects. Best Practice & Res in Clin Gastroent. 2003, 17(5):725-740
- Kaur, I.P., et. al. Probiotics: potential pharmaceutical applications. Eur J of Pharmaceut Sci. 2002, 15(1):1-9 Garcia-Larsen V., et. al. Diet during pregnancy and infancy and risk of allergic or autoimmune disease: a systematic review and meta-analysis. PLoS Med. 2018, 15(2):e1002507

- systematic review and meta-analysis. PLoS Med. 2018, 15(2):e1002507
  Meuer S. Probiotics and respiratory tract diseases. Ann Nutr Metab. 2010, 57 Suppl:24-6
  . Cunningham-Rundles S., et. al. Effect of probiotic bacteria on microbial host defense, growth, and immune function in human immunodeficiency virus type-1 infection. Nutrients. 2011, 3(12):1042-70
  . Ganji-Arienaki M., Rafieian-Kopaei M. Probiotics are a good choice in remission of inflammatory bowel diseases: A meta analysis and systematic review. J Cell Physiol. 2018, 233(3):2091-2103
  . Bustamante M., et. al. Probiotics as an Adjunct Therapy for the Treatment of Halitosis, Dental Caries and Periodontitis. Probiotics Antimicrob Proteins. 2019, Feb 7
  . Miloseyic I. et. al. Gutt-110r Axis. Gutt Microbiota and Its Modulation in the Management of Liver Picases.
- 13. Milosevic I., et. al. Gut-Liver Axis, Gut Microbiota, and Its Modulation in the Management of Liver Diseases:
- A Review of the Literature.

  Dhama K., et. al. Probiotics in curing allergic and inflammatory conditions- research progress and futuristic vision. Recent Pat Inflamm Allergy Drug Discov. 2017, 10(2):105-118
- Huang R., et. al. Effect of probiotics on depression: a systematic review and meta-analysis of randomized controlled trials. Nutrients. 2016, 8(8):e483
   So S.S., et. al. Probiotics-mediated suppression of cancer. Curr Opin Oncol. 2017, 29(1):62-72
- 17. Karamali M., et. al. Effects of probiotic supplementation on hormonal profiles, biomarkers of inflammation and oxidative stress in women with polycystic ovary syndrome: a randomized, double-blind, placebo-controlled trial. Arch Iran Med. 2018, 21(1):1-7

  18. Zhao R., et. al. Effects of fiber and probiotics on diarrhea associated with enteral nutrition in gastric cancer
- patients: A prospective randomized and controlled trial. Medicine (Baltimore). 2017, 96(43):e8418
- Wang L., et. al. The effects of probiotics on total cholesteol: a meta-analysis of randomized controlled trials. Medicine (Baltimore). 2018, 97(5):e9679
   Hu Y.M., et. al. Effects of probiotics supplement in patients with type 2 diabetes mellitus: a meta-analysis of
- randomized trials. Med Clin (Barc). 2017, 148(8):362-370