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## COMMENTARY

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# Probiotics and Health

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### Abstract

Probiotics are described as live microbial food ingredients that are beneficial to health of the of host, especially by improving intestinal microbial balance. The major consumption of probiotics is in dairy-based foods form, which is containing intestinal species of *Lactobacillus* and *Bifidobacterium*. A number of potential benefits of probiotics have been proposed, including: adherence to cells; exclusion or reduction of pathogenic adherence; production of acids, hydrogen peroxide, and bacteriocins antagonistic to pathogen growth; safe, noninvasive, noncarcinogenic and nonpathogenic characteristics; and congregation to form a normal balanced flora. The interrelation between probiotics and health are reviewed in this article.

**Key Words:** probiotics - health - dairy food - lactobacilli - bifidobacteria

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### Introduction

Elie Metchnikoff first introduced the probiotic concept in 1908, which observed the long life of Bulgarian peasants who consumed fermented milk foods. He suggested that lactobacilli might counteract the putrefactive effects of gastrointestinal metabolism (Metchnikoff, 1907). In the century which is elapsed since Metchnikoff's work, scientists and consumers have accepted the probiotic concept throughout the world (Fuller, 1997). In the world, the concept of providing functional foods including beneficial components rather than removing potentially harmful components is gaining ground in recent years. It may consider a functional food with the special property of containing live, beneficial microorganisms (Melntosh, 1996). Functional foods and nutraceuticals can prevent and treat diseases. Yogurt and other fermented milks containing probiotics may be considered the first functional foods. More specifically, Fuller defined probiotics as a live microbial feed supplement that beneficially affects the host beyond correcting for traditional nutrient deficiencies by improving the intestinal balance (Fuller, 1992).

The first clinical trials in the 1930s focused on the effect of probiotics on constipation, and research has steadily increased since then. Today probiotics are available in a variety of food products and supplements. Food products containing probiotics are almost dairy products that due to the historical association of lactic acid bacteria with fermented milk. The most frequently used bacteria in these

products include the *Lactobacillus* and *Bifidobacterium* species (Anon., 2000). The probiotic species (bacteria and yeasts) are listed in Table 1.

### Beneficial and nutritional effects of probiotics

The colonic microflora is important to health. The growth and metabolism of the many individual bacterial species

**Table 1. The Probiotic Species (bacteria and yeasts)**

Names of bacteria and yeasts	
<i>Lactobacillus species</i>	<i>Bifidobacterium species</i>
<i>L. acidophilus</i>	<i>B. bifidum</i>
<i>L. casei/L. Paracasei</i>	<i>B. breve</i>
<i>L. fermentum</i>	<i>B. lactis</i>
<i>L. bulgaricus</i>	<i>B. longum</i>
<i>L. cremoris</i>	<i>B. infantis</i>
<i>L. gasseri</i>	<i>B. adolescentis</i>
<i>L. johnsonii</i>	
<i>L. lactis</i>	<i>Streptococcus species</i>
<i>L. plantarum</i>	<i>S. thermophilus</i>
<i>L. reuteri</i>	
<i>L. rhamno sus</i>	<i>Saccharomyces species</i>
<i>L. salivarius</i>	<i>S. boulardii</i>
	<i>S. cerevisiae</i>
<i>Enterococcus species</i>	
<i>E. faecium</i>	

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exerting protective effects in the large bowel depend primarily on the substrates available to them, most of which come from the diet (Oksanen et al., 1990; Siitonen et al., 1990). This has led to attempts to modify the structure and metabolic activities of the community through diet using probiotics. These organisms are non pathogenic and nontoxicogenic, retain viability during storage, and survive passage through the stomach and small bowel. In addition, lactic acid bacteria may improve intestinal mobility and relieve constipation, particularly in seniors.

Probiotics are live microbial food supplements that are widely used in yogurts and other dairy products. Some commercial fermented dairy products which has probiotic properties are given in Table 2. The nutritional effects of probiotics have been shown by the fermentation of food with lactic acid bacteria which increased folic acid content of yogurt, bifidus milk and kefir and to increase niacin and riboflavin levels in yogurt, vitamin B<sub>12</sub> in cottage cheese and vitamin B<sub>6</sub> in Cheddar cheese (Alm, 1982; Shahani and Chandan, 1979). In addition to nutrient synthesis, probiotics may improve the digestibility of some dietary nutrients such as protein and fat (Friend and Shahani, 1984). Short-chain fatty acids such as lactic acid, propionic acid and butyric acid produced by lactic acid bacteria may help maintain an appropriate pH and protect against pathological changes in the colonic mucosa (Anon., 2000).

The claims of beneficial effects of probiotic consumption, which the researchers support, are including intestinal, immune system and the other effects (Dugas et al., 1999; Kaur et al., 2002):

- \* Increased nutritional value (better digestibility, increased bioavailability of minerals and vitamins)
- \* Promotion of intestinal lactose digestion (reducing symptoms of lactose intolerance and malabsorption)
- \* Promotion recovery from diarrhea (rotavirus, travelers)
- \* Positive influence on intestinal flora (antibiotics or radiation induced colitis)
- \* Inhibition of pathogen growth and translocation

- \* Prevention of intestinal tract infections (bacteria or virus induced, *Candida* enteritis, *Helicobacter pylori* ulcus/ neoplasia)
- \* Regulation of gut motility (constipation, irritable bowel syndrome)
- \* Enhancing specific and nonspecific immune response system
- \* Stimulation of gastrointestinal immunity
- \* Improved urogenital health
- \* Prevention of cancer and suppressing tumors
- \* Detoxification of carcinogens
- \* Reduction of catabolic products eliminated by kidney and liver
- \* Prevention of arteriosclerosis (reduction of serum cholesterol)
- \* Prevention of osteoporosis
- \* Better development (growth)
- \* Improved well-being
- \* Synthesized nutrients (folic acid, niacin, riboflavin, vitamins B<sub>6</sub> and B<sub>12</sub>)
- \* Increasing nutrient bioavailability
- \* Decreasing prevalence of allergy in susceptible individuals
- \* Decreasing lower serum cholesterol concentrations
- \* Reduction of blood pressure in hypertensives

### Mechanisms of Probiotic Activity

The mechanism of action of probiotic strains is likely to be multifactor and, from existing evidence, appears to be strain specific. Enhancement of colonization resistance and/or direct inhibitory effects against pathogens is likely to be important in situations in which probiotics have reduced the incidence and duration of gastroenteritis. Probiotic strains have inhibited pathogenic bacteria both *in vitro* and *in vivo* through several different mechanisms. Probiotics exert their effects on the host but the mechanisms are still speculative. They may antagonize pathogens directly through production of antimicrobial and antibacterial compounds such as cytokines and butyric acid (De Vuyst and Vandamme, 1994;

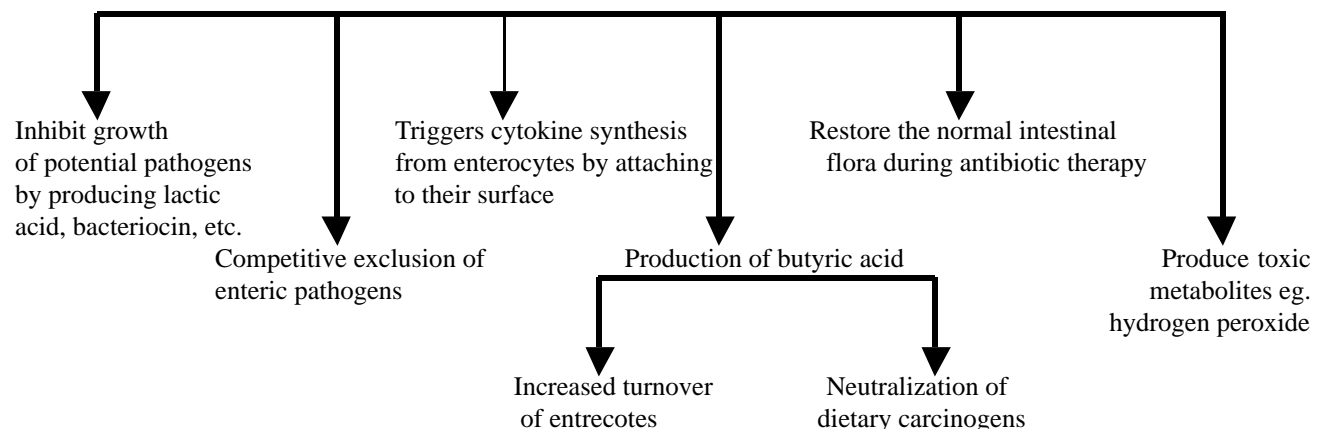
**Table 2. Some Commercial Probiotic Products (Akahn and Ötles, 2002; Miller, 1996; Robinson, 1989; Salji, 1994)**

Product	Microorganism
Acidophilus milk	<i>L. acidophilus</i>
Sweet Acidophilus milk	<i>L. acidophilus</i>
Acidophilus churn out	<i>L. acidophilus</i> , <i>Lactococcus lactis</i> , <i>Lac. Lactis subsp. cremoris</i> , <i>Lac. Lactis biovar diacetylactis</i> , <i>Leuconostoc mesenteroides subsp. cremoris</i>
Acidophilus-yeast milk	<i>L. acidophilus</i> , <i>Saccharomyces fragilis</i> , <i>S. cerevisiae</i>
Acidophilus yogurt	<i>L. acidophilus</i> , <i>L. delbrueckii subsp. bulgaricus</i> , <i>Streptococcus salivarius subsp. thermophilus</i>
Acidophilus-bifidus yogurt	<i>L. acidophilus</i> , <i>Bifidobacterium bifidum</i> , <i>L. delbrueckii subsp. bulgaricus</i> , <i>S. salivarius subsp. thermophilus</i>
Biogarde	<i>L. acidophilus</i> , <i>B. bifidum</i> , <i>S. Salivarius subsp. thermophilus</i>
Bioghurt	<i>L. acidophilus</i> , <i>S. salivarius subsp. thermophilus</i>
Cultura	<i>L. acidophilus</i> , <i>B. bifidum</i> ,
Yakult	<i>L. acidophilus</i> , <i>B. bifidum</i> , <i>B. breve</i> , <i>L. casei</i>

Kailasapathy and Chin, 2000), reduce gut pH by stimulating the lactic acid to produce microflora (Langhendries et al., 1995), compete for binding and receptor sites that pathogens occupy (Fujiwara, 1997; Kailasapathy and Chin, 2000), improve immune function and stimulate immunomodulatory cells (Isolauri et al., 1991 and 1995; Rolfe, 2000), produce lactase which aids in lactose digestion, compete the nutrients and adhere the sites on the gut wall and regulate colonocyte gene expression (e.g. expression of mucin genes) (Fooks and Gibson, 2002; Mack et al. 1999; Steer et al., 2000;). A group of requirements have been identified for a microorganism to be defined as an effective probiotic (Salminen et al., 1996). These are:

1. Adhere to cells;
2. Exclude or reduce pathogenic adherence;
3. Persist and multiply;
4. Produce acids, peroxide, and bacteriocins antagonistic to pathogen growth;
5. To be safe, noninvasive, noncarcinogenic and nonpathogenic;
6. Coaggregate to form a normal balanced flora.

Applying probiotics to stimulate immune function, especially in individuals with underdeveloped or dysregulated immune function, appears to be sound, considering the positive outcomes of feeding studies targeting viral infections, Inflammatory Bowel Disease (IBD) and allergic diseases. It is still unclear which mechanism or, more probably, which spectrum of mechanisms, is used by probiotics within the human gut microbiota to bring about improved health. Further human feeding studies are required to confirm probiotic efficacy in specific disease states such as IBD, colon cancer and gastroenteritis. Probiotic activity is likely to be strain specific and that these disease states are of multifactor etiology (Tuohy et al., 2003). The suggested mechanism of probiotic activity are summarized in Figure 1.



**Figure 1. The Suggested Mechanisms Underlying Probiotic Activity (Kaur et al., 2002)**

## Probiotics and Cancer

Probiotics plays an important role at the intestinal flora in the induction and development of colon cancer by reducing the incidence and number of tumors. The endogenous flora and the immune system take part on the modulation of carcinogenesis. Probiotics may influenced both and this led to trials investigating the role of probiotics in preventing or curing tumors on animals (Wollowski et al., 2001). The probiotic effect that remains the most controversial is the anticancer activity attributed to certain lactic acid bacteria which is reduced cancer risk possibly by counteracting mutagenic and genotoxic effects which are evident by in *vitro* and *in vivo* animal models studies; dietary intervention studies in humans and epidemiological studies correlating cancer to certain dietary regimes. Studies of the effect of probiotic consumption on cancer appear promising.

Several studies showed that some probiotics may decrease the fecal concentrations of enzymes, mutagens, and secondary bile salts that may be involved in colon carcinogenesis (Wollowski et al., 2001). One clinical study showed an increased recurrence-free period in subjects with bladder cancer (Aso and Akazan, 1992). *B. longum* has also been shown to inhibit the incidence of colon, liver, small intestinal and mammary tumors in rats (Reddy and Rivenson, 1993). Another study showed that *Lactobacillus GG* reduces the incidence and number of tumors in animals artificially induced with colon cancer (Goldin et al., 1996). And also one recent study found that dietary supplementation with a strain of *Lactobacillus acidophilus* significantly suppressed the total number of colon cancer cells in rats in a dose-dependent manner (Rao et al., 1999).

## References

- Akalın S and Otles S (2002). Beslenme probiotiklerin önemi. *Gıda*, 7, 70-4.

- Alm L (1982). Effect of fermentation on B-vitamin content of milk in Sweden. *J Dairy Sci*, **65**, 353-9.
- Anon (2000). Probiotics - Friendly bacteria with a host of benefits. Dairy Council of California.
- Aso Y and Akazan H (1992). Prophylactic effect of a Lactobacillus casei preparation on the recurrence of superficial bladder cancer. *Urol Int*, **49**, 125-9.
- De Vuyst L and Vandamme EJ (1994). Antimicrobial potential of lactic acid bacteria. In: De Vuyst L, Vandamme EL, (Eds). Bacteriocins of lactic acid bacteria. Glasgow, United Kingdom: Blackie Academic and Professional, p:91-142.
- Dugas B, Mercenier A, Wijnkoop LL, et al (2000). Immunity and probiotics. *Trends Immunology Today*, **20**, 387-90.
- Fooks LJ and Gibson GR (2002). Probiotics as modulators of the gut flora. *Br J Nutr*, **88**, 39-49.
- Friend BA and Shahani KM (1984). Nutritional and therapeutic aspects of lactobacilli. *J Appl Nutr*, **36**, 125-53.
- Fujiwara S, Hashiba H, Hirota T, Forstner JF (1997). Proteinaceous factor(s) in culture supernatant fluids of bifidobacteria which prevents the binding of enterotoxigenic Escherichia coli to ganglioside GM1. *Appl Environ Microbiol*, **63**, 506-12.
- Fuller R (1992). History and development of probiotics. In Probiotics. The scientific basis. Chapman & Hall Publishers, London.
- Goldin BR, Gualtieri L, Moore RP (1996). The effect of Lactobacillus GG on the initiation and promotion of dimethylhydrazine-induced intestinal tumors in the rat. *Nutr Cancer*, **25**, 197-204.
- Isolauri E, Juntunen M, Rautanen T, Sillanpää P, Koivula T (1991). A human Lactobacillus strain (Lactobacillus casei sp. Strain GG) promotes recovery from acute diarrhea in children. *Pediatrics*, **88**, 90-7.
- Isolauri E, Joensuu J, Suomalainen H, Luomala M, Vesikari T (1995). Improved immunogenicity of oral D 3 RRV reassortant rotavirus vaccine by Lactobacillus casei GG. *Vaccine*, **13**, 310-2.
- Kailasapathy K and Chin J (2000). Survival and therapeutic potential of probiotic organisms with reference to Lactobacillus acidophilus and Bifidobacterium spp. *Immunol Cell Biol*, **78**, 80-8.
- Kopp-Hoolihan (2001). Prophylactic and therapeutic uses of probiotics: a review. *J Am Diet Assoc*, **101**, 229-38.
- Langhendries JP, Detry J, Van Hees J (1995). Effect of a fermented infant formula containing viable bifidobacteria on the fecal flora composition and pH of healthy full-term infants. *J Pediatr Gastroenterol Nutr*, **21**, 177-81.
- Libbey J (1999). Functional Dairy Products. Nutrition and Health Collection.
- Mack DR, et al (1999). Probiotics inhibit enteropathogenic *E. coli* adherence *in vitro* by inducing intestinal mucin gene expression. *Am J Physiol*, **276**, 941-50.
- Metchnikoff E (1907). In The prolongation of life: optimistic studies, (Ed Chalmers Mitchell). London, William Heinemann.
- Miller HJMT (1996). Probiotics-panacea or nostrum? *BNF Nutrition Bulletin*, **21**, 199-202.
- Oksanen PJ, Salminen S, Saxelin M, et al (1990). Prevention of travelers' diarrhea by Lactobacillus GG. *Ann Med*, **22**, 53-6.
- Rao CV, Sanders ME, Indranie C, Simi B, Reddy BS (1999). Prevention of indices of colon carcinogenesis by the probiotic Lactobacillus acidophilus NCFM [TM] in rats. *Int J Oncol*, **14** 939-44.
- Reddy BS and Rivenson A (1993). Inhibitory effect of Bifidobacterium longum on colon, mammary, and liver carcinogenesis induced by 2-amino-3-methylimidazo [4,5] quinoline, a food mutagen. *Canc Res*, **53**, 3914-8.
- Robinson RK (1989). Special yogurts-the potential health benefits. *Dairy Industries International*, **54**, 23-5.
- Rolfe RD (2000). The role of probiotic cultures in the control of gastrointestinal health. *J Nutr*, **130**, 396-402.
- Salminen S, Isolauri E, Salminen E (1996). Clinical uses of probiotics for stabilising gut mucosal barrier: successful strains and future challenges. *Antonie Van Leeuwenhoek*, **70**, 347-58.
- Salji J (1994). Foods for the future. *Food Science and Technology Today*, **8**, 139-43.
- Shahani KM, Chandan RC (1979). Nutritional and healthful aspects of cultured and culture-containing dairy foods. *J Dairy Sci*, **62** 1685-94.
- Siitonen S, Vapaatalo H, Salminen S, (1990). Effect of Lactobacillus GG yoghurt in prevention of antibiotic associated diarrhoea. *Ann Med*, **22**, 57-9.
- Steer T, et al (2000). Perspectives on the role of the human gut microbiota and its modulation by pro- and prebiotics. *Nutr Res Revs*, **13**, 229-54.
- Tuohy KM, Probert HM, Smejkal CW, Gibson GR (2003). Using probiotics and prebiotics to improve gut health. *Reviews Therapeutic Focus*, **8**, 692-700.
- Wollowski I, Rechkemmer G, Pool-Zobel BL (2001). Protective role of probiotics and prebiotics in colon cancer. *Am J Clin Nutr*, **73**, 451-5.