

Review Article

Probiotic Era: An immunostimulatory kick to your salivary IgA levels

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ABSTRACT:

Probiotic bacteria or friendly bacteria have created increased awareness within oral health care as a complementary way to prevent diseases caused by the oral microbial flora. They have shown to enhance the production of Salivary IgA levels. SIgA has shown an immunological control over dental caries by preventing adherence of cariogenic microorganisms to hard surfaces and inhibiting the activity of glucosyltransferases.

Key words: Salivary IgA, Probiotics, Glucosyl transferases.

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INTRODUCTION:

Guarner et al in 2005 defined probiotics as living microorganisms which when administered in adequate quantity are beneficial to the host. The term was adopted by INTERNATIONAL SCIENTIFIC ASSOCIATION FOR PROBIOTICS AND PREBIOTIC and proposed by Lilley and Stillwell in 1965.^[1] Elie Metchnikoff in 1900 discovered *Lactobacillus bulgarius* which paved a way to the concept of probiotics and he reported that Bulgarians have a higher life expectancy due to consumption of yoghurt containing lactic acid and bacteria.

The hypothetical mechanism of action for probiotics are that they modify the oral environment by competing with the pathogens for nutrients at the binding sites, act as antioxidants by preventing plaque formation by neutralizing free electron and stimulate non specific immunity, humoral and cellular immune response.^[2,3] The various vehicles used for administration of probiotics are lozenges, tablets, cheese, gelatin, powder, straw, yoghurt, mouth rinse and capsules.^[4]

The most commonly seen probiotic species in saliva belong to *Lactobacillus* amongst which *Lactobacillus acidophilus*, *Lactobacillus casei*, *Lactobacillus fermentum*, *Lactobacillus plantarum*, *Lactobacillus rhamnosus* and *Lactobacillus salivarius*.

DISCUSSION:

Salivary Secretory IgA (SIgA) is the first line of defense produced by local plasma cells (PCs) which boost oral immunity by preventing microbial adherence, neutralizing enzymes, toxins, and viruses; or by acting in synergy with other factors such as lysozyme and lactoferrin, blocking the epithelial receptors and inhibiting the attachment of pathogenic bacteria, especially *Streptococcus mutans* to epithelial cells.^[5-7] Migrating IgA positive B cells are seen in the salivary glands where they differentiate into plasma cells and release SIgA. SIgA has an immunomodulatory effect over caries by preventing the adherence of cariogenic microorganisms to hard surfaces by preventing pellicle formation and by inhibiting the activity of glycosyl transferases.^[8] It has an antibody that blocks the epithelial receptors and inhibits the attachment of pathogenic bacteria, especially *Streptococcus mutans* to epithelial cells.^[9] Ig A represents about 15 to 20% of immunoglobulins in the blood and mucus secreted in the stomach, lungs and intestines that prevents the microbes to bind to epithelial cells of the respiratory and digestive tract. The two forms of IgA : IgA1 (90%) found in serum and produced by bone marrow B cells and IgA2 (10%) produced by B cells located in the mucosae and

secretions of colostrum, maternal milk, tears and saliva differ in the structure. The dimeric molecules of IgA produced by epithelial cells has a polypeptide of 1.5 kD called secretory chain and as shown in Figure 1 is linked by two additional chains.

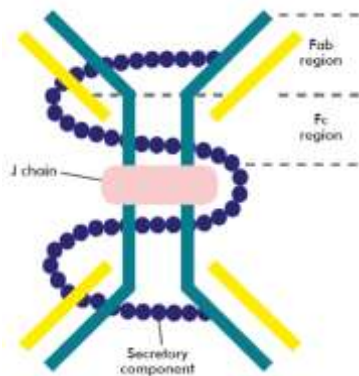


FIGURE 1

Probiotics prevent dental caries by forming a biofilm and thereby preventing proliferation of cariogenic bacteria by antagonizing, competing and leading to decreased acid production. Probiotics have shown to improve the body's immune system through its ability to induce SIgA formation, macrophage activation, modulation of pro-inflammatory cytokines, production of antioxidants like vitamins and lactic acid. The ability of probiotics to modulate the innate and adaptive immune defences has been well documented and probiotics have been seen to release soluble factors which trigger signalling cascades in epithelial cells.^[10]

Lactobacillus is one of the bacteria contained in probiotics which can increase the proliferation of SIgA producing cells e.g. Lactobacillus casei prevents caries by inhibiting the growth of pathogenic bacteria. Streptococcus mutans functions as barrier by strengthening the oral mucosal epithelium by binding to the epitope of the antigen I/II part of Streptococcus mutans, so that streptococcus mutans cannot colonize on the tooth surface.

The most common probiotic strains are Lactobacillus and Bifidobacterium amongst which L.rhamnosus GG, ATCC 53103 (LGG) is the most widely studied probiotic bacterium whose GG was named after Sherwood Gorbach and Barry Goldin on 17th April 1985 who patented it.^[11]

Pahumunto, N et al reported consumption of milk powder with L. paracasei SD1 caused reduction of salivary MS and delayed development of new caries and the strain is safe for use in young children^[12] Chung et al showed L.fermentum significantly inhibited formation of insoluble glucan produced by S.mutans and stopped its adherence onto cuvette walls.^[13]

M.Stensson et al reported that 9 year old children receiving perinatal L.reuteri supplementation from birth to first year of life were caries free and had lower prevalence of approximal caries than those in the control group.^[14] Stecksén-Blicks et al. compared two groups after 21 months for dental caries when L.rhamnosus was supplemented in milk and when fluoride was applied on enamel and noted statistically significant difference between 2 groups with preventive fraction of 75%.^[15]

Comelli EM et al studied 23 dairy bacterial strains for prevention of dental caries and two strains namely S.thermophilus and L.lactis were capable of diminishing the colonization of Streptococcus oralis, Veillonella dispar, Actinomyces naeslundii and Strep.^[16]

Wattanarat et al. found increased levels of SIgA in saliva of children after probiotic administration while reduced salivary IgA levels were seen in athletes.^[17] Ericson et al used a probiotic chewing gum and found no significant impact of bacteria on the SIgA concentration in unstimulated whole saliva but were able to show a post treatment increase in total IgA% per protein after intake of L.reuteri.^[4]

Çaglar et al noted decreased level of S mutans level after 2 week use of L.reuteri enriched yoghurt and results were seen immediately during supplementation and for few days after discontinuation.^[18]

Kotani et al stated oral intake of L.pentosus strain b240 for 12 weeks accelerated salivary IgA levels in elderly. An elevated SIgA secretion rate reached a plateau at week 4, elevating the rate of SIgA secretion during first month and rate remain stable from week 4 to 12 suggesting continuous daily intake of b240 may be efficacious for maintenance of increased salivary SIgA in elderly.^[5] The intake of fermented milk containing L.casei DN-114 001 has shown to induce nonspecific total SIgA in infant saliva by stimulating the gut mucosal immune system to enhance secretion of SIgA in saliva.^[5]

G.S.Pinto et al concluded daily intake of yoghurt with or without B.animalis subsp.lactis for a period of 2 weeks was beneficial in reducing total microbial counts in dental plaque and saliva in orthodontic patients since mouth forms the first part of gastrointestinal tract.^[19]

Probiotics have shown to enhance salivary HNP 1-3 levels however their action to reduce new pit and fissure caries involves microbial interactions. Tao et al demonstrated higher salivary HNP 1-3 levels in caries free children than in those experiencing caries. Increased production of salivary HNP 1-3 has been seen after augmentation of local host immunity after probiotic Lactobacillus paracasei SD1 supplementation. L.paracasei SD1 may exert immunostimulatory effect by enhanced production of HNP1-3 from ductal epithelial cells of submandibular salivary glands.^[17]

Reduction in number of streptococcus mutans after ingestion of probiotics has been observed in number

of studies while no such correlation was seen in other studies. Nase et al enumerated caries preventive role in 594 children aged between 1 to 6 year old by showing significantly reduced dental caries after supplementation with *L.rhamnosus* for 7 months^[20] Significant low dental caries and low counts of *S.mutans* than controls in 3 to 4 year age group was seen.^[20] It has also been well documented that higher titres of salivary IgA levels are seen in people with low dental caries.

Sookkhee and colleagues isolated 3790 strains of lactic acid bacteria and noted *L.paracasei* ssp.*paracasei* and *L.rhamnosus* had high capacity to antagonize important oral pathogens including *Streptococcus mutans* and *Porphyromonas gingivalis*.^[21] Hackioja et al reported that consumption of yoghurt with *L.rhamnosus* probiotic on daily basis hosted this microorganism in the saliva for 3 weeks after stopping the intervention while as Yli-Knuuttila and his colleagues said that the same strain colonized the oral cavity temporarily and consistent consumption is needed for long term beneficial effect.^[22,23]

Oral intake of fermented milk *L.casei* DN-114 001 has been shown to induce non specific total SIgA in infant saliva as seen by Tormo Carnicer R et al because oral intake of b240 stimulates the gut mucosal immune system to enhance secretion of SIgA in saliva.^[5,24]

CONCLUSION:

Probiotics have an immunostimulatory effect which is measurable in saliva. It is seen that probiotic increases salivary IgA levels as long as the intervention is done however no long term effect is seen. Probiotic supplementation can temporarily decrease further caries progression.

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