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## Infant microbiome and factors determining colonization of the gastrointestinal tract and subsequent health consequences

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## ABSTRACT

Microbiome is a key element in maintaining health, forming all saprophytic, commensal and parasitic microorganisms that invade the human body [1]. The development of microbiota begins already in the prenatal period, providing the way for further studies on the influence of the course and duration of pregnancy, delivery mode, feeding, exposure to environmental factors and gestational age [1,2]. Colonization of the gastrointestinal tract of infants is an important aspect of later health [1]. Probiotics and prebiotics can regulate the composition of bacterial flora of the gastrointestinal tract and thus influence the homeostasis of human body [1].

**KEYWORDS:** microbiome, intestinal microbiota, gastrointestinal colonization

## **INFANT MICROBIOME**

Microbiome represents a collection of all microorganisms living in a specific environment (human organism, host), i.e. bacteria, fungi, viruses and archaea, while the term “microbiome” indicates a collection of their genomes [1,2]. The development of microbiota begins already in the prenatal period, providing the way for further studies on the influence of the course and duration of pregnancy, delivery mode, feeding and exposure to environmental factors [3].

Until recently, it was believed that the gastrointestinal tract of a healthy infant is sterile [4]. However, there are reports that the intrauterine environment is not sterile – bacteria were found in the amniotic fluid, fetal membranes and placenta [5, 6]. Jimenez et al. confirmed the presence of marked DNA of bacteria previously administered to the mother in the neonates of mouse meconium [7].

Therefore, already intrauterine the man has to deal with stimulation of the immune system with antigens from the bacteria originating from the mother [4].

## **KEY ELEMENTS IN THE DEVELOPMENT OF MICROFLORA IN INFANTS**

One of the key factors influencing colonization is the birth type, feeding method, use of drugs, especially antibiotics. [8,9] . After childbirth, the immune system of infants and newborns has to face pathogenic pathogens and gain tolerance to food [4]. This is a crucial time for the proper colonization of microorganisms in the intestine and translates into many aspects, including the development of immune tolerance, angiogenesis and stress response [10,11,12,13]. The delivery method affects the type and number of bacteria acquired by a newborn baby. During natural childbirth a newborn baby comes into contact with the bacterial flora of the vagina and gastrointestinal tract of the mother, the dominant flora are: *Lactobacillus*, *Prevotella*, *Escherichia*, *Enterococcus*, *Bacteroides* and *Bifidobacterium* [4, 10 , 14, 15, 16].

Neonates born by cesarean section are colonized by fewer bacteria [4]. The sterile environment of the operating room, which the child encounters during childbirth by cesarean section, deprives the child of contact with the mother’s natural microflora [4]. This causes colonization with bacteria from the hospital environment and from the mother’s skin [4, 17]. Gastrointestinal microflora resembles the ecosystem of bacteria on the skin, it contains mainly bacteria of genera: *Staphylococcus*, *Corynebacterium*, *Propionibacterium* and *Clostridium* [4]. The basic intestinal flora in infants born by caesarean section may be disturbed for up to 6 months after birth [4,17]. Another equally important factor influencing microorganisms colonizing the gastrointestinal tract of infants is the feeding method [4,17]. The best food for a newborn baby is breast milk, which can be described as a natural synbiotic [4,17].

It contains a number of natural oligosaccharides (prebiotics) that stimulate the growth of healthy lactic acid bacteria in the baby’s intestine [8]. Apart from prebiotics, there are also strains of *Bifidobacterium* and *Lactobacillus* (probiotics), which colonize the gastrointestinal tract of infants and have a beneficial effect on their health [18]. In breastfed children, the number of *Bifidobacterium* increases rapidly, which become the dominant flora and may constitute up to 60-90% of the bacteria in stool [4]. Fermentation products of these bacteria protect the digestive tract against pathogenic strains of *E. coli* and *C. perfringens* [9, 19, 20]. It should also be noted that the use of antibiotics has a catastrophic effect on intestinal microbiota [4]. Treatment with broad spectrum antibiotics contributes to a significant decrease in the number of *Bifidobacterium* and *Bacteroides* with simultaneous competitive Firmicutes increase

[4,17]. The reconstruction of bacterial flora of the gastrointestinal tract after antibiotic treatment may take up to 6-10 months [18].

## **MICROBIOTA AND THE IMMUNE RESPONSE OF THE ORGANISM**

Intestinal microbiota has a significant role in the development and efficient functioning of the immune response of the organism and the nervous system, as well as in the regulation of metabolic

and digestive system functions [19/20]. The proper composition of bacterial flora determines the proper functioning of the whole organism. The intestinal microbiota of a healthy child is formed up to the age of 3 years, in the literature the key role of the first 1000 days of a child's life is emphasized [3].

The most important functions of intestinal microorganisms include: stimulation of the immune system development, modulation of genes expression responsible for strengthening the intestinal barrier and angiogenesis, as well as postnatal development and maturation of the gastrointestinal tract [21/22]. The intestinal microflora fulfils many metabolic tasks; it is responsible for the production of vitamins and digestion of carbohydrates, which are not assimilable to humans, to simple sugars and short-chain fatty acids, which are an additional source of energy for the host cells [4]. It is a source of countless antigens constantly stimulating the intestinal immune system, thus supporting the normal development of the intestinal lymphatic system, development and differentiation of NK cells and plasma cells [4]. Plasma cells, located in the lamina propria of intestinal wall, under the influence of the intestinal flora, produce secretory class A immunoglobulin [3/4]. It is a key element of the humoral immunity of gastrointestinal tract and defends the intestinal mucosa against the intrusion of bacteria from the environment [23]. The maintenance of a balance between the intestinal microbiota and the host is of fundamental importance in order to maintain health [4].

Quantitative and qualitative disturbances in the intestinal microflora composition – intestinal dysbiosis – may increase the risk of many diseases, including hospital infections, necrotizing enterocolitis in premature infants, obesity and autoimmune diseases [4]. They are also associated with more frequent occurrence of inflammatory bowel disease, coeliac disease, depression, cancer and others [24, 25, 26, 27].

Colonization of the gastrointestinal tract by non-physiological microorganisms disturbs and delays the maturation of the immune system and causes imbalance between TH1 and TH2 lymphocytes, which is associated with more frequent occurrence of atopic diseases, including atopic dermatitis and asthma [28, 29].

## **CONCLUSIONS**

Intestinal microbiota affects many aspects of human physiology and the incidence of many diseases, not only in the gastrointestinal tract, but also in the systemic system [4]. Bacterial imbalance of the gastrointestinal tract of infants has long-term health effects, because intestinal microflora has a significant role in the development and maturation of the child's immune system [2,4]. Exposure to microorganisms in the earliest period of life is very important in human immunological programming [2,4,17]. Appropriate microbiota is the basis for proper functioning of the organism.

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