



Allergologia et immunopathologia

Sociedad Española de Inmunología Clínica,
Alergología y Asma Pediátrica

www.elsevier.es/ai



Could imbalances in the composition of gut microbiota be implicated in the origin of atopic dermatitis?



Atopic dermatitis (AD) is one of the most common chronic inflammatory skin diseases with a worldwide prevalence of approximately 20% in children¹. Studies on genetic predisposition to eczema have implicated eczema-related genes; environmental factors have also been shown to contribute to the disease pathogenesis². In recent years, there has been increased interest in the role of intestinal microbiota in the etiopathogenesis of AD, because the immune mechanisms involved in AD are complex and little is known about the role of this microbiota¹.

Research shows that gut microbiota is a key source of immune development and regulation early in life³. In recent decades, some studies have shown that a possible imbalance in the composition of gut microbiota or the relationship of the microbiota with the host may be implicated in the origin of allergic diseases⁴. Industrialisation has brought about improvements in lifestyle but has also produced an impact on the intestinal microbiota that is difficult to assess. Healthy intestinal microbiota usually comes from the maternal microbiota evidenced because identical strains are shared. A Dutch group⁵ has reviewed this transmission of microorganisms from mother to child that can be altered by changes in the maternal microbiota due to diet, health status, exposure to broad-spectrum antibiotics and the mother's genotype. The changes can also be due to a reduction of maternal contact by birth, by caesarean section and the use of artificial lactation. In support of this line of work, Penders et al.³ showed in their study that birth order had a strong effect on the microbiota composition, with increasing number of older siblings, the colonisation rates of lactobacilli ($P < 0.001$) and bacteroides ($P 0.02$) at age five weeks increased, whereas rates of clostridia decreased.

In the present issue of *Allergologia et Immunopathologia*, Melli et al.⁶ present a cross-sectional study about the profile of the gut microbiota in school-age children living in the city of Osasco, in the Metropolitan Region of Sao Paulo, Brazil. They included 81 children: 23 with AD and 58 controls. The analysis showed that a higher prevalence of clostridium difficile, a greater abundance of *Bifidobacterium* spp and a lower

abundance of *Lactobacillus* spp. in the gut microbiota are associated with AD in school-age children. Those results are similar to other studies⁷. Those authors also found that children with AD presented lower total eubacteria count, and a statistically significant difference between groups for the counts of *Lactobacillus* spp, *Escherichia coli*, *Bacterioides fragilis* and *Methanobrevibacter smithii*. Numerous studies in recent years have indeed shown that the microbiota of infants with allergies differs from the microbiota of infants without allergies³.

Some authors speak of the skin-gut axis⁸, referring to changes in the composition and microbial function, called dysbiosis, both in the intestine and in the skin, related to deviations in immune responses that lead to skin diseases such as atopic dermatitis. In a recent and interesting review on the role of Gut Microbiota in Atopic Dermatitis, Petersen et al.¹ summarised that the results were conflicting and the role of the gut microbiota in the development and severity of AD remains unclear. They included 44 studies: 26 were observational and 18 were interventional (use of probiotics) studies. Some studies showed that participants who developed AD had a less diverse gut microbiome than healthy individuals, while others found no significant differences. In addition, observational studies failed to demonstrate overgrowth or lack of specific bacterial species in patients with AD compared with individuals without AD. Nearly half of the interventional studies showed a positive effect of probiotics on the severity of AD, with a concomitant alteration in the gut microbial composition. The remaining studies showed no effect of probiotics on the severity of AD¹.

In conclusion, our understanding on healthy intestinal microbiota and how it is altered in AD is increasing steadily. However, further research would be necessary to know the mechanisms by which intestinal microbes interfere with the immune system and then incorporate this knowledge into therapy. Although there is currently little evidence that probiotics prevent or treat AD, this may become an alternative in the future. We expect many key findings to be made in the next few years.

References

1. Petersen EB, Skov L, Thyssen JP, Jensen P. Role of the Gut Microbiota in Atopic Dermatitis: A systematic Review. *Acta Derm Venereol.* 2019;99:5–11.
2. Kang Y, Cai Y, Pan W. Change in gut microbiota for eczema: Implications for novel therapeutic strategies. *Allergol Immunopathol (Madr).* 2018;46:281–90.
3. Penders J, Gerhold K, Stobberingh E, Thijs C, et al. Establishment of the intestinal microbiota and its role for atopic dermatitis in early childhood. *J. Allergy Clin Immunol.* 2013, 132-: 601-7.
4. Kalliomäki M, Kirjavainen P, Eerola E, Kero P, Salminen S, Isolauri E. Distinct patterns of neonatal gut microflora in infants in whom atopy was and was not developing. *J Allergy Clin Immunol.* 2001;107:129–34.
5. Van Daele E, Knol J, Belzer C. Microbial transmission from mother to child: improving infant intestinal microbiota development by identifying the obstacles. *Rev Microbiol.* 2019;45(5–6):613–48.
6. Melli LC, Carmo-Rodrigues MS, Araújo-Filho HB, Mello CS, Tahan S, Pignatari AC, Solé D, Morais MB. Gut microbiota of children with atopic dermatitis: Controlled study in the metropolitan region of Sao Paulo, Brazil. *Allergol Immunopathol.* 2019;48.
7. Eun Kim J, Sung Kim H. Microbiome of the skin and gut in atopic dermatitis (AD): understanding the pathophysiology and finding novel management strategies. *J. Clin med.* 2019;8:444.
8. Lee SY, Lee E, Park YM, Hong SJ. Microbiome in the Gut-Skin Axis in Atopic Dermatitis. *Allergy Asthma Immunol Res.* 2018 Jul;10:354–62.

Belén García Avilés^a, Nuria Marco Lozano^{b,*},
Francisco Javier Contreras-Porta^c

^a *Department of Pediatrics, San Juan Clinic University Hospital, Alicante, Spain*

^b *Department of Pediatrics, Vega Baja Hospital, Orihuela, Spain*

^c *Allergy Unit. La Paz University Hospital, Madrid, Spain*

* Corresponding author.

E-mail address: numalo27@hotmail.com (N.M. Lozano).

5 February 2020 24 February 2020