
Virus-specific airway immune response in early life is associated with childhood asthma

M. Elsner Melgaard (Copenhagen, Denmark), B. Lund Krogsgaard Chawes (Copenhagen, Denmark), S. Kjeldgaard Jensen (Copenhagen, Denmark), J. Stokholm (Slagelse, Denmark), A. Malby Schoos (Slagelse, Denmark), S. Brix (Kgs. Lyngby, Denmark), A. Eliassen (Kgs. Lyngby, Denmark), C. Tingskov Pedersen (Copenhagen, Denmark), H. Bisgaard (Copenhagen, Denmark), K. Bønnelykke (Copenhagen, Denmark)

Background: Virus-induced asthma-like symptoms in the first years of life have been associated with an increased risk of later asthma development, but the underlying mechanism is poorly understood.

Objectives: To investigate in situ pathogen-specific differences in upper airway immune mediator levels during episodes with asthma-like symptoms and the association with later development of asthma.

Methods: We included 493 asthma-like episodes in 277 children aged 0-4 years from the COPSAC2010 mother-child cohort. Levels of 18 different immune mediators were assessed in nasal epithelial lining fluid and were compared between children with and without viral PCR-identification (nasopharyngeal sample). Finally, we investigated whether the immune response was associated with the development of asthma.

Results: Viral detection during asthma-like symptoms were associated with upregulation of several Th1 and Treg immune mediators, including IFN- γ , TNF- α , CCL4, CXCL10 and IL10, and downregulation of Th2 and Th17 immune mediators, including CCL13, CCL26 and CXCL8 ($p < 0.05$). Increased levels of CCL4, CXCL8 ($p < 0.05$) and lower IFN- γ levels ($p < 0.06$) during viral episodes were associated with asthma by age six years.

Conclusion: We demonstrated a virus-specific airway immune mediator profile during asthma-like episodes in early childhood. Furthermore, an aberrant immune profile during viral triggered episodes was associated with later asthma development, providing a potential mechanistic clue to the virus-asthma relationship.
