

Asthma transition from childhood into adulthood

Oliver Fuchs*, Thomas Bohmert†, Claus F Rabe, Erik von Mutius

Asthma is the most prevalent chronic respiratory disease both in children and adults and resembles a complex syndrome rather than a single disease. Different methods have been developed to better characterise distinct asthma phenotypes in childhood and adulthood. In studies of adults, most phenotyping relies on biomaterials from the lower airways; however, this information is missing in paediatric studies because of restricted accessibility. Few patients show symptoms throughout childhood, adolescence, and adulthood. Risk factors for this might be genetics, family history of asthma and atopy, infections early in life, allergic diseases, and lung function deficits. In turn, a large proportion of children with asthma lose their symptoms during school age and adolescence. This improved prognosis, which might also reflect a better treatment response, is associated with being male and with milder and less allergic disease. Importantly, whether clinical remission of symptoms equals the disappearance of underlying pathology is unknown. In fact, airway hyper-responsiveness and airway inflammation might remain despite the absence of overt symptoms. Additionally, a new-onset of asthma symptoms is apparent in adulthood, especially in women and in the case of impaired lung function. However, many patients do not remember childhood symptoms, which might reflect relapse rather than true initiation. Both relapse and adult-onset of asthma symptoms have been associated with allergic disease and sensitisation in addition to airway hyper-responsiveness. Thus, asthma symptoms beginning in adults might have originated in childhood. Equivocally, persistence into, relapse, and new-onset of symptoms in adulthood have all been related to active smoking. However, underlying mechanisms for the associations remain unclear, and future asthma research should therefore integrate standardised molecular approaches in identical ways in both paediatric and adult populations and in longitudinal studies.

Introduction

Whooping disorders and asthma are the most prevalent chronic respiratory diseases both in childhood and adulthood, with about 300 million people of all ethnicities affected worldwide across all ages. Despite common pathophysiological factors that lead to characteristic clinical symptoms, individuals affected by asthma vary substantially in their complaints, in their clinical manifestations, including comorbidities, and across age groups.¹

Childhood and adult asthma resemble a complex syndrome rather than a single disease and present with different phenotypes^{2,3} that lead to the recognised state of variable airway obstruction, which at times is not fully reversible or even fixed during the disease course.⁴ Presumably, however, variable airway obstruction is affected through diverse underlying pathophysiological processes adding further to the complexity of our pathophysiological understanding.^{5,6} These diverse underlying mechanisms could be used to further define asthma, which is known as an asthma endotype. This approach also includes complex but refined molecular information in a network approach with the aim to better link biology to clinical presentation.^{7,8} However, symptoms, especially during early life, are rather poor predictors of asthma later in life, and the disease displays a variable course across all ages. These variabilities are possibly the reason why asthma phenotypes have not helped to improve prognosis in the clinical setting for children or for adults, although such predictors and targeted treatment strategies are urgently needed.

Independent of age, several methods for the definition of asthma phenotypes have been developed. These methods are either based on a combination of

clinical and empirical approaches and data obtained from epidemiological studies, including molecular information,^{9,10} or they use objective and unsupervised data-driven techniques to determine endophenotypes and phenotypes from collated data,^{11,12} which remain subjective because of the a-priori choice of data used in the analyses.

However, most endophenotyping and phenotyping have been investigated in children and adults separately, with an inherent bias towards data on underlying pathophysiology in adults and epidemiological data for children. Accordingly, data on the important link between childhood and adult asthma are sparse, despite some prospective studies spanning the age ranges.

Key messages

- For persistent asthma symptoms, epidemiological studies have identified genetic background, family history of asthma and atopy, infections early in life, allergic disease, and lung function deficits as risk factors.
- Data for remission and relapse of asthma symptoms are sparse, but point towards male sex and a milder and less allergic disease being associated with remission of symptoms, and a role of allergic disease and persisting airway hyper-responsiveness for their relapse.
- For new-onset of asthma symptoms in adults, risk is increased for women and in the case of allergic disease, pre-existing airway hyper-responsiveness and lung function impairment.
- Equivocally, persistence into adulthood and relapse and new-onset of asthma symptoms in adulthood have all been related to uptake of active smoking.



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Table of Contents Asthma From Childhood To Adulthood Proefschrift

1. Understanding the eBook Asthma From Childhood To Adulthood Proefschrift
 - The Rise of Digital Reading Asthma From Childhood To Adulthood Proefschrift
 - Advantages of eBooks Over Traditional Books
2. Identifying Asthma From Childhood To Adulthood Proefschrift
 - Exploring Different Genres
 - Considering Fiction vs. Non-Fiction
 - Determining Your Reading Goals
3. Choosing the Right eBook Platform
 - Popular eBook Platforms
 - Features to Look for in an Asthma From Childhood To Adulthood Proefschrift
 - User-Friendly Interface
4. Exploring eBook Recommendations from Asthma From Childhood To Adulthood Proefschrift
 - Personalized Recommendations
 - Asthma From Childhood To Adulthood Proefschrift User Reviews and Ratings
 - Asthma From Childhood To Adulthood Proefschrift and Bestseller Lists
5. Accessing Asthma From Childhood To Adulthood Proefschrift Free and Paid eBooks

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- Asthma From Childhood To Adulthood Proefschrift eBook Subscription Services
- Asthma From Childhood To Adulthood Proefschrift Budget-Friendly Options
- 6. Navigating Asthma From Childhood To Adulthood Proefschrift eBook Formats
 - ePub, PDF, MOBI, and More
 - Asthma From Childhood To Adulthood Proefschrift Compatibility with Devices
 - Asthma From Childhood To Adulthood Proefschrift Enhanced eBook Features
- 7. Enhancing Your Reading Experience
 - Adjustable Fonts and Text Sizes of Asthma From Childhood To Adulthood Proefschrift
 - Highlighting and Note-Taking Asthma From Childhood To Adulthood Proefschrift
 - Interactive Elements Asthma From Childhood To Adulthood Proefschrift
- 8. Staying Engaged with Asthma From Childhood To Adulthood Proefschrift
 - Joining Online Reading Communities
 - Participating in Virtual Book Clubs
 - Following Authors and Publishers Asthma From Childhood To Adulthood Proefschrift
- 9. Balancing eBooks and Physical Books Asthma From Childhood To Adulthood Proefschrift
 - Benefits of a Digital Library
 - Creating a Diverse Reading Collection Asthma From Childhood To Adulthood Proefschrift
- 10. Overcoming Reading Challenges
 - Dealing with Digital Eye Strain
 - Minimizing Distractions
 - Managing Screen Time
- 11. Cultivating a Reading Routine Asthma From Childhood To Adulthood Proefschrift
 - Setting Reading Goals Asthma From Childhood To Adulthood Proefschrift
 - Carving Out Dedicated Reading Time
- 12. Sourcing Reliable Information of Asthma From Childhood To Adulthood Proefschrift
 - Fact-Checking eBook Content of Asthma From Childhood To Adulthood Proefschrift
 - Distinguishing Credible Sources
- 13. Promoting Lifelong Learning
 - Utilizing eBooks for Skill Development

- Exploring Educational eBooks

14. Embracing eBook Trends

- Integration of Multimedia Elements
- Interactive and Gamified eBooks

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