

Review



Increasing Prevalence of Allergic Rhinitis in China

Yuan Zhang ^{1,2,3} Luo Zhang ^{1,2,3*}

¹Department of Otolaryngology Head and Neck Surgery, Beijing TongRen Hospital, Capital Medical University, Beijing, China

²Beijing Key Laboratory of Nasal Diseases, Beijing Institute of Otolaryngology, Beijing, China

³Department of Allergy, Beijing TongRen Hospital, Capital Medical University, Beijing, China



Received: Jun 30, 2018

Revised: Aug 16, 2018

Accepted: Aug 28, 2018

Correspondence to

Luo Zhang, MD, PhD

Beijing Key Laboratory of Nasal Diseases,
Beijing Institute of Otolaryngology, No. 17,
Hougou Hutong, Dongcheng District, Beijing
100005, China.

Tel: +86-10-65141136

Fax: +86-10-85115988

E-mail: dr.luozhang@139.com


Copyright © 2019 The Korean Academy of
Asthma, Allergy and Clinical Immunology ·
The Korean Academy of Pediatric Allergy and
Respiratory Disease

This is an Open Access article distributed
under the terms of the Creative Commons
Attribution Non-Commercial License (<https://creativecommons.org/licenses/by-nc/4.0/>)
which permits unrestricted non-commercial
use, distribution, and reproduction in any
medium, provided the original work is properly
cited.

ORCID iDs

Yuan Zhang 

<https://orcid.org/0000-0003-1080-4267>

Luo Zhang 

<https://orcid.org/0000-0002-0910-9884>

Disclosure

There are no financial or other issues that
might lead to conflict of interest.

ABSTRACT

Allergic rhinitis (AR) is a critical public health, medical and economic problem in China. AR is also an important risk factor which will cause many diseases or disorders, especially in children. The trend of AR incidence is still on the rise in recent years and has had a significant effect on the general public. This significant increase is alarming, which highlights an urgent need for better understanding of the prevalence status and characteristics, sensitization patterns, and the associated risk factors of AR in order to improve treatment and develop effective prevention strategies.

Keywords: Allergic rhinitis; prevalence; China; asthma; allergen

INTRODUCTION

Allergic rhinitis (AR) is a hypersensitivity reaction caused when inhaled particles contact the nasal mucosa and induce an immunoglobulin E (IgE)-mediated inflammatory response, which is often accompanied by ocular pruritus, redness and/or lacrimation.¹ AR is one of the most common chronic diseases in all age groups.² Epidemiological studies have revealed that the prevalence of AR has increased progressively in more developed countries, and currently affects 10%–40% of adults and 2%–25% of children^{3–5} worldwide. Likewise, with rapid economic development and more Westernized lifestyles in terms of urbanization and dietary habits of developing nations, a rising trend of AR has also been observed in recent decades,^{6,7} with a widely varying prevalence particularly in China.^{8,9} Furthermore, the potential effect of the considerable increases in air pollutant levels and climate change can also not be ignored with regard to the increased prevalence of respiratory disorders.^{10,11} We have reviewed the prevalence of AR, the incidence of comorbid allergic diseases, and trends and patterns of sensitizing allergens of AR in adults and children in China in 2014.⁸ The aim of this current review was to elaborate changing features of the prevalence of AR in adults as well as children in China and to assess the possible risk factors contributing to such variations according to publications between March 2013 and June 2018 using PubMed. Citations for articles reporting the prevalence of AR in China were identified using *allergic rhinitis* as the primary search term in association with *China*, *prevalence/incidence*, *comorbid allergic disease*, *asthma* and *sensitizing allergens* as the secondary search terms.

CHANGING PREVALENCE OF AR IN ADULTS

It has been widely recognized that the prevalence of AR in many countries has markedly increased over time.^{7,10,12} A nationwide population-based study assessed self-reported AR using validated questionnaire-based telephone interviews in over 38,000 adult subjects from 11 major cities across China between September 2004 and May 2005 demonstrated that the prevalence of AR was 11.1% (8.7%–24.1%).¹³ Recently, a total of 47,216 telephone interviews were conducted and the standardized prevalence of adult AR in 18 major cities was 17.6% in 2011 (9.8%–23%),⁹ suggesting the obviously increasing trend and the instability of the prevalence of AR in China. Comparison of age-adjusted prevalence of self-reported AR between 2005 and 2011 indicated that the standardized prevalence of AR had significantly increased in 8 of 11 cities ($P < 0.01$ for Guangzhou and $P < 0.001$ for Beijing, Changchun, Shenyang, Shanghai, Nanjing, Hangzhou and Xi'an), decreased in 1 city (Urumqi, $P < 0.01$), and not significantly altered in 2 cities (Wuhan and Changsha) as summarized in **Fig. 1**.⁹

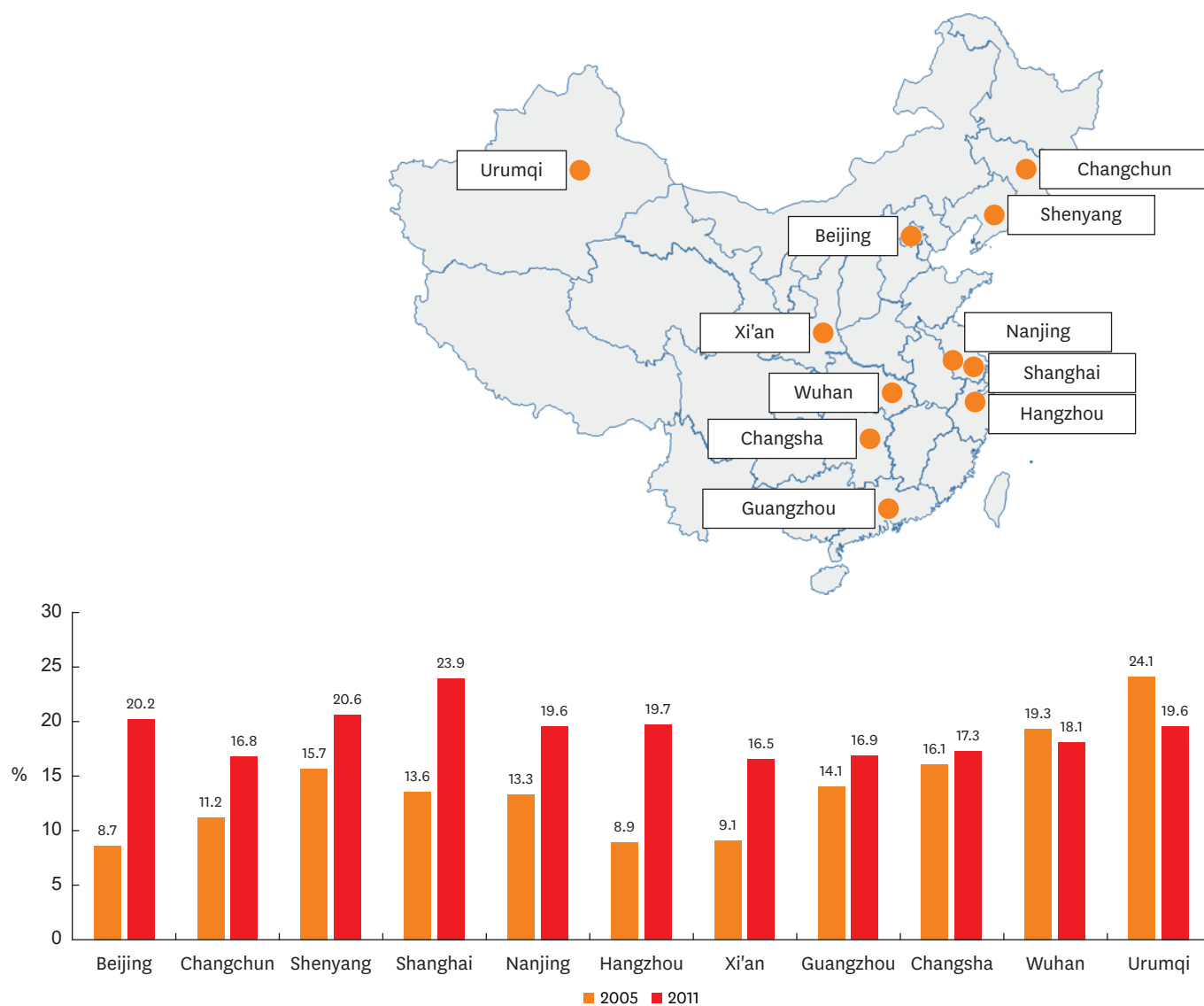


Fig. 1. Comparisons of the prevalence of self-reported AR in 11 major cities in China between 2005 and 2011 according to the study by Wang et al.¹⁴ AR, allergic rhinitis.

Furthermore, a total of 6,043 subjects from the grasslands of Northern China completed a study recently and demonstrated a proportion of 32.4% epidemiologic AR and 18.5% physician-diagnosed pollen-induced AR (PiAR).¹⁴

These nationwide population studies based on telephone interviews revealed the most commonly reported symptoms of AR were sneezing (81.8%), rhinorrhea (60.2%), blocked nose (54.9%), nasal itching (49.6%), ocular itching (42.9%), pharyngolaryngeal symptoms (35.1%), watery eyes (34.5%), ocular swelling (19.1%) and ocular pain (17.6%).⁹ Moreover, another survey performed in Guangzhou via face-to-face interviews demonstrated that the majority (69%) of the patients indicated that the most severe symptom episodes occurred in the early morning and that the symptom severity gradually reduced over time until midnight. Of the total patients, 287 (49%) patients complained that the symptom severity differed according to the season and that the symptom became more severe in most of the patients (75%) in winter (November, December, January and February).¹⁵

The prevalence rate of AR between the urban and rural areas were not always consistent. One cross-sectional investigation performed in 9,899 citizens in Guangzhou from December 2009 to March 2010 using a stratified multistage cluster-sampling method through face-to-face interviews, presented a significantly higher prevalence in the urban area (8.32%) compared with the rural area (3.43%).¹⁵ Another survey via face-to-face interviews using modified validated questionnaires in a community in Beijing (1,499 participants) and a village in Baoding (803 participants) exhibited that the self-reported AR prevalence was significantly lower in the urban area (13.5%) than in the rural area (19.1%).¹⁶ It is still worth mentioning that the confirmable AR prevalence in terms of the positive skin prick test (SPT) results was 7.2% and 6.2% among the urban and rural adults, respectively, reflecting the similar prevalence between urban and rural areas in China.¹⁶ However, a study investigated in the grasslands of Northern China reported that subjects from the urban area showed a higher prevalence of physician-diagnosed PiAR than those from the rural area (23.1% vs. 14.0%, $P < 0.001$), making the comparison of AR prevalence between the urban and rural areas more confusing.¹⁴

CHANGING PREVALENCE OF AR IN CHILDREN

The International Study of Asthma and Allergies in Children (ISAAC) has found a slight worldwide increase in the prevalence of AR among children aged 13–14 years (0.18% per year) and 6–7 years (0.17% per year), as well as a > 20-fold variation in the prevalence of self-reported AR among countries in the same region and between centers in the same country.⁵ Over the 12-year period from 2001 to 2012, the prevalence of current rhinitis and lifetime rhinitis prevalence increased by 5.5% (22.6 vs. 28.1, $P = 0.004$) and 13% (18.6 vs. 31.7, $P < 0.001$) among 2 Isle of Wight birth cohorts.⁶ Comparatively, there was no trend study over time of AR prevalence in children in China; however, the prevalence of asthma in 12,235 children age 0–14 years in Fuzhou, a city in South-east China, was obtained as 1.57% in 1990 and 3.28% in 2000 based on a cross-sectional study with a 2-stage, clustered, stratified random sample design, indicating an increased trend over time.¹⁷ Very recently, the China, Children, Homes, Health cross-sectional study was conducted in Shanghai. Questionnaires were distributed to 17,898 parents or guardians of pre-school children from 72 kindergartens in 5 districts in Shanghai from April 2011 to April 2012. The prevalence of asthma was detected to increase significantly, almost five-fold, from 2.1% in 1990 to 10.2% in 2011.¹⁸

To date, only 1 nationwide survey of the prevalence of AR in children in China has been reported.¹⁹ A total of 23,791 children aged 6–13 years in 8 metropolitan capital cities of provinces in 4 regions were surveyed via standard ISAAC questionnaire between November and December of 2005 using a cluster-stratified sampling method. The study demonstrated that the mean prevalence of childhood AR in the 8 cities was 9.8% (Shanghai 13.1%, Guangzhou 16.8%, Xi'an 3.9%, Wuhan 8.3%, Harbin 4.9%, Chengdu 10.1%, Hohhot 4.5% and Urumqi 10.1%). A smaller cross-sectional survey by Zhao *et al.*²⁰ investigated the self-reported prevalence of childhood AR in 3 major cities in China (Beijing, Chongqing and Guangzhou) using the ISAAC questionnaire between October 2008 and May 2009. A total of 24,290 children aged 0–14 years were interviewed using a multi-stage sampling method; the self-reported prevalence of AR was 14.46%, 20.42% and 7.83% in Beijing, Chongqing and Guangzhou, respectively.²⁰ Actually, besides the questionnaire, the other issues regarding the sampling and investigation were all different between these 2 studies.^{19,20} In this sense, it is hard to attribute the results to 1 reason or clearly explain them. The subjects' age and area source, the sample size and the sampling methods might be all the possible reasons causing the variation, such as the AR prevalence in Guangzhou (16.8% in 2005¹⁹ vs. 7.83% in 2008–2009²⁰). Stratified sampling and cluster sampling surveys were performed among 10–17-year-old students in Changsha city from June 2011 to April 2012 and the prevalence of AR was 15.8%–19.4%.²⁹ A current study analyzed the cohort data of 74,688 junior high school students from a national retrospective birth cohort study in Taiwan, and among subjects mainly at 13–15 years of age, the estimated prevalence of AR was 22.4%.²² In addition, an investigation of clinical features of AR in children in Shanghai demonstrated that 8.6% of AR cases were classified as intermittent mild, 4.2% as persistent mild, 40.5% as intermittent moderate-severe and 46.7% as persistent moderate-severe.²³

There have been numerous reports concerning the prevalence of AR for children compared to adults in the urban versus rural areas. A questionnaire survey was performed on 11,473 children aged 7–12 years in 20 schools from urban Guangzhou and rural Shaoguan, China and showed that the prevalence of self-reported AR (23.2% vs. 5.3%) was significantly higher in Guangzhou compared to Shaoguan.²⁴ Similarly, from a total of 14,884 questionnaires for 3- to 7-year-old children in Shanghai, the prevalence of hay fever in urban children was shown to be higher than that in suburban children as well.¹⁸

CURRENT STATUS OF COMORBID ALLERGIC DISEASES

Attaching great importance to concomitant diseases of AR will contribute to the precise management of AR itself. It has been reported that the incidence of AR accompanied by rhinosinusitis, asthma or atopic dermatitis was significantly higher among individuals having self-reported AR compared to the general population,⁹ while the occurrence of comorbidities such as allergic conjunctivitis, cough and asthma gradually increased from intermittent mild, persistent mild and intermittent moderate-severe to persistent moderate-severe in terms of an AR children study in Shanghai.²³ To be exact, a cross-sectional, hospital-based survey of 1,931 AR patients in two modernized cities (Guangzhou and Zhuhai) in Southern China and the prevalence of concomitant asthma for AR patients is 5.33% (103 of 1,931).²⁵ Comparatively, another cross-sectional questionnaire study surveyed the diagnostic methods and treatment patterns for asthma-AR comorbidity by primary respiratory medicine specialty (PRMs) from 98 hospitals across China and the PRMs reported an estimated prevalence of asthma-AR comorbidity of > 30% at their clinics.²⁶ On the contrary, research conducted

on 20,051 patients with asthma in an outpatient setting covering all territories of China confirmed that AR presented in 69.9% of patients with asthma,²⁷ further suggesting the upper and lower airway relationship between AR and asthma.

PROFILES OF SENSITIZATION TO INHALED ALLERGENS

The changing prevalence of aeroallergen sensitizations may explain different time trends observed in the cohorts suffering from AR as well as the other allergic diseases.⁶ Moreover, a better summary of differences in sensitization patterns within different geographical areas will be beneficial to optimize the optimal panels for allergen tests and efficiently prevent exposure. Due to the effect of geographic, climatic and humanistic factors, the type of allergen inducing AR varies significantly among regions. Zhang *et al.*⁸ summarized the pattern of sensitization to inhalant allergens among AR patients in Mainland China and provided evidence that the prevalence and type of aeroallergens were different in various cities and regions. Dust mites were reported as the most common allergen in many regions for both adults and children. In 2009, Li *et al.*²⁸ surveyed 6,304 patients suffering from asthma and/or rhinitis in 17 cities from four regions of China and showed that *Dermatophagoides farinae* and *Dermatophagoides pteronyssinus* were the predominant aeroallergens in perennial/persistent AR individuals in China. Similar results were gained by the study in single regions such as Guangzhou,²⁵ Xiamen,²⁵ Quanzhou,²⁹ Zhengzhou,³⁰ Qingdao,³¹ Yichang³² and Shanghai.³³ The overall prevalence of specific immunoglobulin E (sIgE)-mediated sensitization to aeroallergens in 5,486 patients with AR who visited the ear, nose, and throat outpatient clinic in Guangzhou, the largest city in Southern China from January 2005 to December 2014 were as follows: 84.4% for house dust mites (HDMS), 23.4% for pet allergens, 21.1% for cockroaches, 9.1% for mold allergens, 7.7% for tree pollen and 6.0% for weed pollen.³⁴ The prevalence of sensitization to aeroallergens decreased with age.³⁴ Such results were also reported in a study of Shanghai children suffering from AR, *i.e.*, with age the prevalence of sIgE against inhalant allergens was significantly increased; however, the opposite trend was observed for food allergens. Additionally, the proportion of children with high levels of sIgE against *Der p* increased with age.²³ Furthermore, the prevalence of sensitization to inhaled allergens was significantly higher in children at > 6 years of age compared to that in those at 3–5 years of age.³¹ As for PiAR patients in Northern China, most of them were sensitized to 2 or more pollens (79.4%) with *Artemisia*, *Chenopodium* and *Humulus scandens* being the most common pollen types.¹⁴ Interestingly, 7,148 subjects with self-reported AR recently completed a standard questionnaire and were assessed for sensitization to relevant allergens by SPT. Eight allergens (*Der f*, mugwort, *Blatella*, hazel, goosefoot, *Penicillium notatum*, animal dander and *Der p*) allowed identification > 96% of sensitized subjects in Central China. Differences in optimal panels were observed between regions, with 5 to 6 allergens being sufficient for North-east (*Der f*, mugwort, *Blatella*, hazel, goosefoot and dandelion), North-west (mugwort, *Der f*, goosefoot, *Blatella* and hazel) and South (*Der f*, *Blatella*, dandelion, ragweed and birch) China,³⁵ providing a cost-effective tool to screen sensitized patients in China. Nevertheless, no significant difference in major allergen positivity was observed in developed and developing regions in Guangdong³⁶ as well as the North-west Hubei Province.³⁷

With respect to the changing pattern of inhaled allergens, a study performed in Guangzhou revealed that the prevalence of sensitization to pet allergens in patients with AR increased at an annual rate of 1.3% (95% confidence interval, 0.85% to 1.67 n%, $P < 0.01$) in the past decade,³⁴ with a similar trend worldwide.³⁸

Last but not least, as regards the fact that correct identification of the culprit allergen is an essential part of diagnosis and treatment in allergic diseases, molecular biology has recently made important advances facilitating such identification and overcoming some of the drawbacks of natural allergen extracts, which consist of a mixture of various proteins that may be allergenic or not, specific for the allergen source or widely distributed.³⁹ New technologies offer the opportunity for a more accurate component-resolved diagnosis (CRD),⁴⁰ especially to polysensitized allergic patients. In China, there is so far a paucity of data about allergen CRD and the prevalence of grass pollen allergen components, in contrast to those from Western countries. As a result, 22.5% of patients with AR and/or asthma were positive for Bermuda-sIgE, 13.6% for Timothy-sIgE and 7.0% for *Humulus scandens*-sIgE. Of the Bermuda-sIgE-positive patients, 53.4% were Cyn d 1-positive and 60.3% were Timothy-sIgE-positive. Of the Timothy-positive patients, 100% were positive for Phl p 4, 17.1% were positive for Phl p 1, and 8.6% tested positive for Phl p 5/6/7/11/12. Patients with high Bermuda-sIgE levels were more likely to be positive for other grasses. In 41.4% of Bermuda grass-positive patients, cross-reactive carbohydrate determinant (CCD)-sIgE was also positive. Sensitization to Phl p 4 was significantly correlated with CCD.⁴¹ In addition, Chan *et al.*⁴² reported that an HDM genome draft produced from genomic, transcriptomic and proteomic experiments revealed allergen genes and a diverse endosymbiotic microbiome, providing a tool to further identify and characterize HDM allergens and to develop diagnostics and immunotherapeutic vaccines.

RISK FACTORS ASSOCIATED WITH AR INCIDENCE

Numerous risk factors have been described for AR in different stages of life, and they can be briefly divided into three aspects: genetic susceptibility, pre- and post-natal events, and environmental exposure as illustrated in **Fig. 2**.

First, genetically predisposed individuals are under increased risk of developing AR when they are exposed to certain environmental and lifestyle factors. In many studies, the most important risk factor for allergic disease development is reported as genetic susceptibility. If there is a family history of atopy in the first and second degrees, this risk increases further. In China, a genetic epidemiologic study involving 23,825 families from Jiangsu Province reported that the average AR heritability of the first, second and third generations was 81.86%.⁴³ A family history of allergies and a patient history of allergies within 6 months of birth were significantly associated with the duration and severity of AR symptoms in a Shanghai children survey.^{15,23}

Secondly, it has also been hypothesized that factors influencing the in utero environment may affect immune system development and later allergic diseases. Based on a cross-sectional survey of 20,803 elementary school students from 8 metropolitan cities in China in 2005, there was a higher likelihood of AR in school-aged children who were not exclusively breast-fed in the first 4 months of their lives (odds ratio [OR], 1.28), children who were born post-term (OR, 1.34), children delivered by Cesarean section (OR, 1.07) or children born to mothers who experienced depressive symptoms during the pre- and post-natal periods (OR, 1.28). AR in school-aged children was found to be associated with pre- and post-natal events. These findings indicate that strategies for reducing exposure to risk factors during pre- and post-natal periods for childhood allergies might be warranted.⁴⁴ A previous study in Shanghai suggested that no siblings, mothers at older ages during pregnancy, shorter breast-feeding period, use of antibiotics in the first year, and home dampness-related exposures, had significant associations

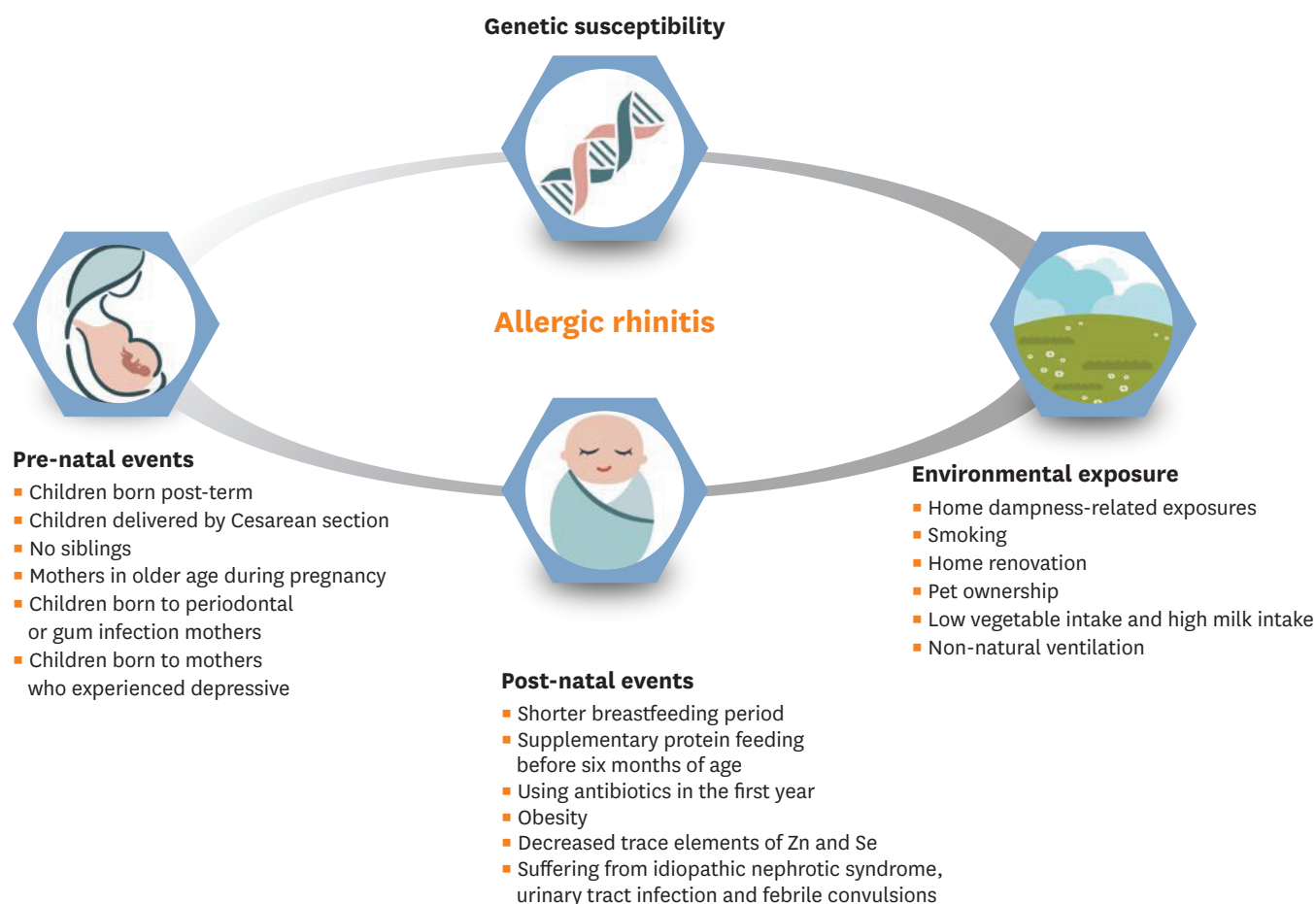


Fig. 2. Profile of risk factors influencing AR incidence according to studies in China. AR, allergic rhinitis.

with increased prevalence of rhinitis.⁴⁵ Another 9-year population-based, retrospective cohort study using Taiwan's National Health Insurance database demonstrated that the overall cumulative incidence and risk of AR in children born to periodontal mothers were significantly higher than those born to non-inflammatory mothers. Gum infection in women during pregnancy is an independent risk factor for allergic diseases in children, and thus its intergenerational consequences should be considered in gestational care.⁴⁶

Moreover, the most hazardous experiences such as disease and drug use, especially during the neonatal period and in childhood are also unavoidable risk factors for AR development. Children with idiopathic nephrotic syndrome (INS) had adjusted hazard ratio (1.71) for AR. Allergic disorders are common in children with INS, especially within the first year after diagnosis.⁴⁷ Urinary tract infection in newborns is significantly associated with the development of AR (OR, 1.41) in childhood and might be a risk factor for subsequent childhood AR.⁴⁸ Risk of AR was found to be 1.21-fold higher in the febrile convulsion group than in the control group, and this risk of AR development is further increased (0.94 vs. 18.9) with frequency of febrile convulsion-related medical visits (1 to 3 visits vs more than 3 visits, $P < 0.0001$). Children with more than 3 febrile convulsion-related medical visits had a significantly higher cumulative incidence of AR.⁴⁹ Another recent study investigated the correlation between AR and trace elements, and reported that the content of Zn and Se

were lower in AR patients serum than in the control group.⁵⁰ In addition, obesity proved to increase the prevalence of AR⁵¹ and excessive weight gain was also confirmed to be an important risk factor developing AR during adolescence, especially among infants born small for their gestational age.²² Self-medication with antibiotics (OR, 6.35),²⁵ especially use of at least 1 course of antibiotics in the first year of life (OR, 7.61) was involved.¹⁷

Thirdly, the current living place, the living place during babyhood, smoking, home renovation and pet ownership were significant risk factors associated with AR prevalence in the population. Change of living environment and lifestyles had strong impacts on the prevalence of AR.^{15,52} Home renovation in the past 2 years was associated with increased likelihood of AR in Chinese women (OR, 1.39).⁵³ Meanwhile, high vegetable intake, low milk intake²⁴ and natural ventilation at home¹⁷ might protect against asthma, while supplementary protein feeding before 6 months of age were harmful.¹⁷

CLIMATE, ENVIRONMENTAL EXPOSURE AND LIFESTYLE IMPACT ON AR PREVALENCE

Over recent decades, AR and allergic diseases have become increasingly common, but the reason for this increased prevalence is still unclear. It has been apparent that genetic variation alone is not sufficient to account for the observed changes; rather, changing climate and environment, together with alterations in lifestyle and eating habits, are likely to have driven the increase in prevalence, and in some cases, the severity of disease.⁵⁴ For example, no difference in PiAR prevalence between the Han and Mongolian populations was observed in a Chinese study, indicating the more important role of high seasonal pollen exposure, but not ethnic-oriented genetic susceptibility.¹⁴

It has been confirmed that the potential effect of the considerable increases in climate change should not be ignored with regard to the increased prevalence of respiratory disorders.^{10,11} Actually, climate change is correlated with allergens for several reasons, especially involving the increase in the amount of pollen produced by each plant.^{55,56} Very recently, Wang *et al.*¹⁴ carried out a sizable population survey in the grasslands of Northern China to study the impact of the intensity and time of pollen exposure on AR prevalence, and they reported that the epidemiologic AR and PiAR was 32.4% and 18.5%, respectively, exhibiting a remarkably higher rate than cities. Furthermore, they found that the PiAR symptoms were positively associated with pollen counts, temperature and precipitation ($P < 0.05$), but negatively with wind speed and pressure ($P < 0.05$). In this sense, PiAR prevalence in the grasslands region being extremely high may be due to high seasonal pollen exposure, which was influenced by local environmental and climate conditions.¹⁴

Concerning environmental exposures from outdoor and indoor, most studies provide evidence that both may contribute to AR development. A study conducted on 2,598 children aged 3–6 years in Changsha demonstrated that early life exposure to traffic-related air pollutants including NO₂ and PM₁₀ during pregnancy and first-year of life may contribute to childhood AR.⁵⁷ Another survey found that pre-conceptional, prenatal, post-natal exposure to outdoor industrial and traffic air pollutants were significantly associated with increases in the risk of childhood asthma, AR and eczema. New furniture was associated with eczema and AR during post-natal exposure, but redecoration was associated with asthma and eczema during prenatal exposure.⁵⁸ A recent study also supported the hypothesis that

childhood allergic diseases originate in fetal life and were triggered by traffic-related air pollution in sensitive trimesters,⁵⁹ *i.e.*, maternal exposure to traffic-related air pollutant NO₂ during pregnancy, especially in specific trimesters, was associated with an increased risk of developing asthma, AR, and eczema in children. There was also evidence proving that the location, type, building area, decoration materials, and construction period of the residence also had significant associations with allergic diseases.⁴⁵ Moreover, a previous study further assessed the impact of residential home characteristics and home environmental risk factors on respiratory diseases in Chinese women, and they reported that living near the main road or near ambient air pollution sources, pet-keeping, humidifier use and home renovation in the past 2 years were all risk factors for asthma, chronic bronchitis and AR.⁵³ Comparatively, a parent-completed questionnaire survey administered in Hong Kong in 2003 assessed the influence of indoor and outdoor environmental factors, concluding that the exposure to moisture and mold in the first year of life increased the risk of asthma (OR, 2.56) and rhinoconjunctivitis (OR, 2.09). Current maternal smoking was also associated with a higher prevalence of asthma (OR, 2.00). No association was observed between outdoor traffic-related air pollutants and the prevalence of allergic conditions. Indoor home environments had a stronger influence on allergy development.⁶⁰

Interestingly, Ouyang *et al.*⁶¹ recently published an article that illuminated the arguments about why air pollutants might increase the sensitization of pollen. Short-term exposure to oak pollen at high concentrations of SO₂ or NO₂ significantly increased their fragility and disruption, leading to subsequent release of pollen cytoplasmic granules into the atmosphere. These results suggested that heightened air pollution during the oak pollen season may possibly increase the incidence of allergic airway disease in sensitized individuals by facilitating the bioavailability of airborne pollen allergens.

PROSPECTS

AR and allergic diseases remain a critical public health, medical and economic problem, and are in fact among the major causes of illness for all ages in China. AR has also proved to be an important risk factor that will cause many diseases or disorders, especially in children. It has been verified that AR exerted significant effects on students' character, memory and sleep quality, emotion, and memory.²¹ Moreover, AR cases showed a greater risk of otitis media with effusion,⁶² Kawasaki disease,⁶³ primary immune thrombocytopenia,⁶⁴ migraine,⁶⁵ attention deficit hyperactivity disorder, and oppositional defiant disorder⁶⁶ in children, sleep disorder-related problems in pregnant women,⁶⁷ and mental disorders in adults.⁶⁸ Unfortunately, the trend of AR incidence is still on the rise and has become significant economic burden on the general public.⁶⁹ In this regard, China is one of the largest countries in the world with a population of around 1.3 billion citizens and with rapidly expanding industrialization, while the growth rate of national AR prevalence from 2006 to 2011 was 6.5%.⁹ The significant increase in health cost is alarming, which highlights an urgent need for better understanding of the prevalence status, etiology and its causative mechanisms in order to improve treatment and to develop effective prevention strategies. For example, optimized instruments based on objective tests have been more adequately employed to assess the prevalence of AR in the past 5 years than before. Meanwhile, the definition of PRMs has been given great attention because PRMs with greater work experience and higher professional titles estimated treating a significantly higher proportion of patients with AR as well as asthma-AR comorbidity.²⁶ Moreover, allergy specialists in China have confirmed that

the incidence of pollen-, dust mites- and mold-sensitive AR can be predicted by employing models based on prevailing meteorological conditions,⁷⁰ providing critical methods for the prevention of environmental exposure. Avoidance behaviors and reverse causation in parental smoking, pet-keeping and dietary habits for childhood rhinitis should be carefully considered. Public health policies should further help patients benefitted by proper diagnosis/treatment as well as help specifically target the local risk factors in order to control AR incidence.⁶⁹

ACKNOWLEDGMENTS

This work was supported by grants from the program for National Key R&D Program of China (2016YFC20160905200), the national natural science foundation of China(81570895, 81420108009, 81400444, 81470678 and 81630023), Changjiang scholars and innovative research team (IRT13082), the special fund of capital health development (2011-1017-06, 2011-1017-02), the special fund of sanitation elite reconstruction of Beijing (2009-2-007), Beijing health bureau program for high level talents (2011-3-043), Beijing Municipal Administration of Hospitals' Mission Plan (SML20150203), Capital citizenry health program (z161100000116062) and Beijing Advanced Innovation Center for Food Nutrition and Human Health, Beijing Technology and Business University (20181045).

REFERENCES

1. Bousquet J, Khaltaev N, Cruz AA, Denburg J, Fokkens WJ, Togias A, et al. Allergic rhinitis and its impact on asthma (ARIA) 2008 update (in collaboration with the World Health Organization, GA(2)LEN and AllerGen). *Allergy* 2008;63 Suppl 86:8-160.
[PUBMED](#) | [CROSSREF](#)
2. Allergic rhinitis: common, costly, and neglected. *Lancet* 2008;371:2057.
[PUBMED](#) | [CROSSREF](#)
3. Meltzer EO, Blaiss MS, Naclerio RM, Stoloff SW, Derebery MJ, Nelson HS, et al. Burden of allergic rhinitis: allergies in America, Latin America, and Asia-Pacific adult surveys. *Allergy Asthma Proc* 2012;33 Suppl 1:S113-41.
[PUBMED](#) | [CROSSREF](#)
4. Bauchau V, Durham SR. Prevalence and rate of diagnosis of allergic rhinitis in Europe. *Eur Respir J* 2004;24:758-64.
[PUBMED](#) | [CROSSREF](#)
5. Asher MI, Montefort S, Björkstén B, Lai CK, Strachan DP, Weiland SK, et al. Worldwide time trends in the prevalence of symptoms of asthma, allergic rhinoconjunctivitis, and eczema in childhood: ISAAC phases one and three repeat multicountry cross-sectional surveys. *Lancet* 2006;368:733-43.
[PUBMED](#) | [CROSSREF](#)
6. Patil VK, Kurukulaaratchy RJ, Venter C, Grundy J, Roberts G, Dean T, et al. Changing prevalence of wheeze, rhinitis and allergic sensitisation in late childhood: findings from 2 Isle of Wight birth cohorts 12 years apart. *Clin Exp Allergy* 2015;45:1430-8.
[PUBMED](#) | [CROSSREF](#)
7. Baena-Cagnani CE, Canonica GW, Zaky Helal M, Gómez RM, Compalati E, Zernotti ME, et al. The international survey on the management of allergic rhinitis by physicians and patients (ISMAR). *World Allergy Organ J* 2015;8:10.
[PUBMED](#) | [CROSSREF](#)
8. Zhang Y, Zhang L. Prevalence of allergic rhinitis in china. *Allergy Asthma Immunol Res* 2014;6:105-13.
[PUBMED](#) | [CROSSREF](#)
9. Wang XD, Zheng M, Lou HF, Wang CS, Zhang Y, Bo MY, et al. An increased prevalence of self-reported allergic rhinitis in major Chinese cities from 2005 to 2011. *Allergy* 2016;71:1170-80.
[PUBMED](#) | [CROSSREF](#)

10. D'Amato G, Holgate ST, Pawankar R, Ledford DK, Cecchi L, Al-Ahmad M, et al. Meteorological conditions, climate change, new emerging factors, and asthma and related allergic disorders. A statement of the World Allergy Organization. *World Allergy Organ J* 2015;8:25.
[PUBMED](#)
11. D'Amato G, Vitale C, Rosario N, Neto HJ, Chong-Silva DC, Mendonça F, et al. Climate change, allergy and asthma, and the role of tropical forests. *World Allergy Organ J* 2017;10:11.
[PUBMED](#) | [CROSSREF](#)
12. Verlato G, Corsico A, Villani S, Cerveri I, Migliore E, Accordini S, et al. Is the prevalence of adult asthma and allergic rhinitis still increasing? results of an Italian study. *J Allergy Clin Immunol* 2003;111:1232-8.
[PUBMED](#) | [CROSSREF](#)
13. Zhang L, Han D, Huang D, Wu Y, Dong Z, Xu G, et al. Prevalence of self-reported allergic rhinitis in eleven major cities in China. *Int Arch Allergy Immunol* 2009;149:47-57.
[PUBMED](#) | [CROSSREF](#)
14. Wang XY, Ma TT, Wang XY, Zhuang Y, Wang XD, Ning HY, et al. Prevalence of pollen-induced allergic rhinitis with high pollen exposure in grasslands of Northern China. *Allergy* 2018;73:1232-43.
[PUBMED](#) | [CROSSREF](#)
15. Li CW, Chen DH, Zhong JT, Lin ZB, Peng H, Lu HG, et al. Epidemiological characterization and risk factors of allergic rhinitis in the general population in Guangzhou city in China. *PLoS One* 2014;9:e114950.
[PUBMED](#) | [CROSSREF](#)
16. Zheng M, Wang X, Bo M, Wang K, Zhao Y, He F, et al. Prevalence of allergic rhinitis among adults in urban and rural areas of china: a population-based cross-sectional survey. *Allergy Asthma Immunol Res* 2015;7:148-57.
[PUBMED](#) | [CROSSREF](#)
17. Tang SP, Liu YL, Wang SB, Weng SF, Chen S, Zhang MJ, et al. Trends in prevalence and risk factors of childhood asthma in Fuzhou, a city in Southeastern China. *J Asthma* 2015;52:10-5.
[PUBMED](#) | [CROSSREF](#)
18. Huang C, Liu W, Hu Y, Zou Z, Zhao Z, Shen L, et al. Updated prevalences of asthma, allergy, and airway symptoms, and a systematic review of trends over time for childhood asthma in Shanghai, China. *PLoS One* 2015;10:e0121577.
[PUBMED](#) | [CROSSREF](#)
19. Li F, Zhou Y, Li S, Jiang F, Jin X, Yan C, et al. Prevalence and risk factors of childhood allergic diseases in eight metropolitan cities in China: a multicenter study. *BMC Public Health* 2011;11:437.
[PUBMED](#) | [CROSSREF](#)
20. Zhao J, Bai J, Shen K, Xiang L, Huang S, Chen A, et al. Self-reported prevalence of childhood allergic diseases in three cities of China: a multicenter study. *BMC Public Health* 2010;10:551.
[PUBMED](#) | [CROSSREF](#)
21. Song Y, Wang M, Xie J, Li W, Zhang X, Wang T, et al. Prevalence of allergic rhinitis among elementary and middle school students in Changsha city and its impact on quality of life. *J Laryngol Otol* 2015;129:1108-14.
[PUBMED](#) | [CROSSREF](#)
22. Lin MH, Hsieh CJ, Caffrey JL, Lin YS, Wang JJ, Ho WC, et al. Fetal growth, obesity, and atopic disorders in adolescence: a retrospective birth cohort study. *Paediatr Perinat Epidemiol* 2015;29:472-9.
[PUBMED](#) | [CROSSREF](#)
23. He S, Li YJ, Chen J. Clinical features of allergic rhinitis in children of Shanghai, China. *Genet Mol Res* 2016;15:1-13.
[PUBMED](#) | [CROSSREF](#)
24. Yang Z, Zheng W, Yung E, Zhong N, Wong GW, Li J. Frequency of food group consumption and risk of allergic disease and sensitization in schoolchildren in urban and rural China. *Clin Exp Allergy* 2015;45:1823-32.
[PUBMED](#) | [CROSSREF](#)
25. Hong H, Yang Q, Zuo K, Chen X, Xia W, Lv M, et al. A hospital-based survey on the prevalence of bronchial asthma in patients with allergic rhinitis in Southern China. *Am J Rhinol Allergy* 2013;27:502-5.
[PUBMED](#) | [CROSSREF](#)
26. Su N, Lin J, Liu G, Yin K, Zhou X, Shen H, et al. Asthma with allergic rhinitis management in China: a nationwide survey of respiratory specialists at tertiary hospitals. *Int Forum Allergy Rhinol* 2015;5:221-32.
[PUBMED](#) | [CROSSREF](#)
27. Lin J, Su N, Liu G, Yin K, Zhou X, Shen H, et al. The impact of concomitant allergic rhinitis on asthma control: a cross-sectional nationwide survey in China. *J Asthma* 2014;51:34-43.
[PUBMED](#) | [CROSSREF](#)

28. Li J, Sun B, Huang Y, Lin X, Zhao D, Tan G, et al. A multicentre study assessing the prevalence of sensitizations in patients with asthma and/or rhinitis in China. *Allergy* 2009;64:1083-92.
[PUBMED](#) | [CROSSREF](#)
29. Que Z, Ran Q, Lin D, Zhang Z, Cai J, Xu Y, et al. Allergen analysis of patients with allergic rhinitis in Quanzhou. *Lin Chung Er Bi Yan Hou Tou Jing Wai Ke Za Zhi* 2013;27:1148-50.
[PUBMED](#)
30. Zhao Y, Li W, Di R, Wang J, Hu P, Ji P. Prevalence of sensitization to allergens in 1 139 patients with allergic rhinitis in Zhengzhou area. *Zhonghua Er Bi Yan Hou Tou Jing Wai Ke Za Zhi* 2014;49:858-60.
[PUBMED](#)
31. Lin H, Lin R, Li N. Sensitization rates for various allergens in children with allergic rhinitis in Qingdao, China. *Int J Environ Res Public Health* 2015;12:10984-94.
[PUBMED](#) | [CROSSREF](#)
32. Zhang L, Han B, Zhang Z, Liu A, Liu G, Du Z, et al. Skin prick test of inhalative allergens for patients with allergic rhinitis in Yichang. *Lin Chung Er Bi Yan Hou Tou Jing Wai Ke Za Zhi* 2014;28:98-101.
[PUBMED](#)
33. Zhou J, Yan P, Zhang D, Ren X, Feng Y, Wang W, et al. Preliminary analysis of SPT tests on 5,843 allergic rhinitis patients from South Shanghai area. *Lin Chung Er Bi Yan Hou Tou Jing Wai Ke Za Zhi* 2014;28:102-7, 112.
34. Wang W, Huang X, Chen Z, Zheng R, Chen Y, Zhang G, et al. Prevalence and trends of sensitisation to aeroallergens in patients with allergic rhinitis in Guangzhou, China: a 10-year retrospective study. *BMJ Open* 2016;6:e011085.
[PUBMED](#) | [CROSSREF](#)
35. Lou H, Ma S, Zhao Y, Cao F, He F, Liu Z, et al. Sensitization patterns and minimum screening panels for aeroallergens in self-reported allergic rhinitis in China. *Sci Rep* 2017;7:9286.
[PUBMED](#) | [CROSSREF](#)
36. Qiu Q, Lu C, Han H, Chen S. Clinical survey and analysis of allergic rhinitis patients' allergens in Guangdong developed and developing regions. *Lin Chung Er Bi Yan Hou Tou Jing Wai Ke Za Zhi* 2013;27:928-31.
[PUBMED](#)
37. Chen X, Chen J, Wang Q, Zhu H, Li G, Kong W. Analysis on allergen of clinical allergic rhinitis patients in North-west mountain area of Hubei Province. *Lin Chung Er Bi Yan Hou Tou Jing Wai Ke Za Zhi* 2015;29:1203-5.
[PUBMED](#)
38. Chan SK, Leung DY. Dog and cat allergies: current state of diagnostic approaches and challenges. *Allergy Asthma Immunol Res* 2018;10:97-105.
[PUBMED](#) | [CROSSREF](#)
39. Douladiris N, Savvatiannos S, Roumpedaki I, Skevaki C, Mitsias D, Papadopoulos NG. A molecular diagnostic algorithm to guide pollen immunotherapy in Southern Europe: towards component-resolved management of allergic diseases. *Int Arch Allergy Immunol* 2013;162:163-72.
[PUBMED](#) | [CROSSREF](#)
40. Valenta R, Lidholm J, Niederberger V, Hayek B, Kraft D, Grönlund H. The recombinant allergen-based concept of component-resolved diagnostics and immunotherapy (CRD and CRIT). *Clin Exp Allergy* 1999;29:896-904.
[PUBMED](#) | [CROSSREF](#)
41. Luo W, Huang H, Zheng P, Wei N, Luo J, Sun B, et al. Major grass pollen allergens and components detected in a Southern Chinese cohort of patients with allergic rhinitis and/or asthma. *Mol Immunol* 2016;78:105-12.
[PUBMED](#) | [CROSSREF](#)
42. Chan TF, Ji KM, Yim AK, Liu XY, Zhou JW, Li RQ, et al. The draft genome, transcriptome, and microbiome of *Dermatophagoides farinae* reveal a broad spectrum of dust mite allergens. *J Allergy Clin Immunol* 2015;135:539-48.
[PUBMED](#) | [CROSSREF](#)
43. Ma L, Chen DL, Zhang RX, Wang XL, Shi YJ, Ji C, et al. Genetic epidemiological study on allergic rhinitis in Nantong region of Jiangsu Province. *Zhonghua Er Bi Yan Hou Tou Jing Wai Ke Za Zhi* 2007;42:643-6.
[PUBMED](#)
44. Li Y, Jiang Y, Li S, Shen X, Liu J, Jiang F. Pre- and postnatal risk factors in relation to allergic rhinitis in school-aged children in China. *PLoS One* 2015;10:e0114022.
[PUBMED](#) | [CROSSREF](#)
45. Wang X, Liu W, Hu Y, Zou Z, Shen L, Huang C. Home environment, lifestyles behaviors, and rhinitis in childhood. *Int J Hyg Environ Health* 2016;219:220-31.
[PUBMED](#) | [CROSSREF](#)

46. Hsieh VC, Liu CC, Hsiao YC, Wu TN. Risk of allergic rhinitis, allergic conjunctivitis, and eczema in children born to mothers with gum inflammation during pregnancy. *PLoS One* 2016;11:e0156185.
[PUBMED](#) | [CROSSREF](#)
47. Wei CC, Lin CL, Shen TC, Sung FC. Occurrence of common allergic diseases in children with idiopathic nephrotic syndrome. *J Epidemiol* 2015;25:370-7.
[PUBMED](#) | [CROSSREF](#)
48. Lin CH, Lin WC, Wang YC, Lin IC, Kao CH. Association between neonatal urinary tract infection and risk of childhood allergic rhinitis. *Medicine (Baltimore)* 2015;94:e1625.
[PUBMED](#) | [CROSSREF](#)
49. Lin WY, Muo CH, Ku YC, Sung FC, Kao CH. Increased association between febrile convulsion and allergic rhinitis in children: a nationwide population-based retrospective cohort study. *Pediatr Neurol* 2014;50:329-33.
[PUBMED](#) | [CROSSREF](#)
50. Ma R, Shen Y, Hou L, Yang Z, Feng N, Yan X, et al. The correlation of allergic rhinitis and trace elements in Ningxia region. *Zhonghua Er Bi Yan Hou Tou Jing Wai Ke Za Zhi* 2014;49:1017-20.
[PUBMED](#)
51. Lei Y, Yang H, Zhen L. Obesity is a risk factor for allergic rhinitis in children of Wuhan (China). *Asia Pac Allergy* 2016;6:101-4.
[PUBMED](#) | [CROSSREF](#)
52. Wang J, Li B, Yu W, Yang Q, Wang H, Huang D, et al. Rhinitis symptoms and asthma among parents of preschool children in relation to the home environment in Chongqing, China. *PLoS One* 2014;9:e94731.
[PUBMED](#) | [CROSSREF](#)
53. Dong GH, Qian ZM, Wang J, Trevathan E, Ma W, Chen W, et al. Residential characteristics and household risk factors and respiratory diseases in Chinese women: the Seven Northeast Cities (SNEC) study. *Sci Total Environ* 2013;463-464:389-94.
[PUBMED](#) | [CROSSREF](#)
54. Huang SK, Zhang Q, Qiu Z, Chung KF. Mechanistic impact of outdoor air pollution on asthma and allergic diseases. *J Thorac Dis* 2015;7:23-33.
[PUBMED](#)
55. D'Amato G, Vitale C, Lanza M, Molino A, D'Amato M. Climate change, air pollution, and allergic respiratory diseases: an update. *Curr Opin Allergy Clin Immunol* 2016;16:434-40.
[PUBMED](#) | [CROSSREF](#)
56. D'Amato G, Cecchi L, D'Amato M, Annesi-Maesano I. Climate change and respiratory diseases. *Eur Respir Rev* 2014;23:161-9.
[PUBMED](#) | [CROSSREF](#)
57. Deng Q, Lu C, Yu Y, Li Y, Sundell J, Norbäck D. Early life exposure to traffic-related air pollution and allergic rhinitis in preschool children. *Respir Med* 2016;121:67-73.
[PUBMED](#) | [CROSSREF](#)
58. Deng Q, Lu C, Ou C, Chen L, Yuan H. Preconceptional, prenatal and postnatal exposure to outdoor and indoor environmental factors on allergic diseases/symptoms in preschool children. *Chemosphere* 2016;152:459-67.
[PUBMED](#) | [CROSSREF](#)
59. Deng Q, Lu C, Li Y, Sundell J, Dan Norbäck. Exposure to outdoor air pollution during trimesters of pregnancy and childhood asthma, allergic rhinitis, and eczema. *Environ Res* 2016;150:119-27.
[PUBMED](#) | [CROSSREF](#)
60. Lam A, Wong GW, Poon CM, Lee SS. A GIS-based assessment of environmental influences on allergy development in children. *Asia Pac J Public Health* 2014;26:575-87.
[PUBMED](#) | [CROSSREF](#)
61. Ouyang Y, Xu Z, Fan E, Li Y, Zhang L. Effect of nitrogen dioxide and sulfur dioxide on viability and morphology of oak pollen. *Int Forum Allergy Rhinol* 2016;6:95-100.
[PUBMED](#) | [CROSSREF](#)
62. Pau BC, Ng DK. Prevalence of otitis media with effusion in children with allergic rhinitis, a cross sectional study. *Int J Pediatr Otorhinolaryngol* 2016;84:156-60.
[PUBMED](#) | [CROSSREF](#)
63. Hwang CY, Hwang YY, Chen YJ, Chen CC, Lin MW, Chen TJ, et al. Atopic diathesis in patients with Kawasaki disease. *J Pediatr* 2013;163:811-5.
[PUBMED](#) | [CROSSREF](#)
64. Chiang MR, Wei CC, Muo CS, Fu LS, Li TC, Kao CH. Association of primary immune thrombocytopenia and common allergic diseases among children. *Pediatr Res* 2015;77:597-601.
[PUBMED](#) | [CROSSREF](#)

65. Wang IC, Tsai JD, Lin CL, Shen TC, Li TC, Wei CC. Allergic rhinitis and associated risk of migraine among children: a nationwide population-based cohort study. *Int Forum Allergy Rhinol* 2016;6:322-7.
[PUBMED](#) | [CROSSREF](#)
66. Lin YT, Chen YC, Gau SS, Yeh TH, Fan HY, Hwang YY, et al. Associations between allergic diseases and attention deficit hyperactivity/oppositional defiant disorders in children. *Pediatr Res* 2016;80:480-5.
[PUBMED](#) | [CROSSREF](#)
67. Cai XH, Xie YP, Li XC, Qu WL, Li T, Wang HX, et al. The prevalence and associated risk factors of sleep disorder-related symptoms in pregnant women in China. *Sleep Breath* 2013;17:951-6.
[PUBMED](#) | [CROSSREF](#)
68. Xi L, Cao F, Zhang Y, Zhang L. Severity of nasal obstruction can predict the anxiety status of patients with allergic rhinitis but not patients with vasomotor rhinitis. *Int Forum Allergy Rhinol* 2016;6:1196-203.
[PUBMED](#) | [CROSSREF](#)
69. Cheng L, Chen J, Fu Q, He S, Li H, Liu Z, et al. Chinese society of allergy guidelines for diagnosis and treatment of allergic rhinitis. *Allergy Asthma Immunol Res* 2018;10:300-53.
[PUBMED](#) | [CROSSREF](#)
70. Ouyang Y, Li J, Zhang D, Fan E, Li Y, Zhang L. A model to predict the incidence of allergic rhinitis based on meteorological factors. *Sci Rep* 2017;7:10006.
[PUBMED](#) | [CROSSREF](#)