Epidemiologic evidence for association between allergic rhinitis and bronchial asthma

Vikas Deep Mishra¹, Jai Krishna Mishra^{2,*}, S.C. Mohapatra³

¹Assistant Professor, Dept. of TB & Chest Diseases, Heritage Institute of Medical Sciences, Varanasi, ²Professor & Head, Dept. of TB & Respiratory Diseases, Institute of Medical Sciences, BHU, Varanasi, ³Dean Academic Affairs, SGT University & Head Community Medicine, FMHS, SGT University, Gurgaon

Corresponding Author:

Email: jkmishra.imsbhu@gmail.com

Abstract

Allergic rhinitis and asthma are often found together, and the characteristics of the two disease are quite similar. Many epidemiologic studies have provided evidence to support the theory that allergic or perennial rhinitis often precedes the development of asthma in same patient, and that, the development of asthma is seen much more frequently in people with both seasonal and perennial rhinitis, than in people with either condition alone. Perennial rhinitis has been associated with increase in non-specific bronchial hyper responsiveness. Rhinitis and asthma were found to be co-morbidities regardless of atopic state. Several studies have demonstrated the frequent co-existence of allergic rhinitis and asthma, that allergi rhinitis usually precedes asthma and that allergic rhinitis is a risk factor for asthma. Further many studies have suggested that early age of onset of atopy may be an important predictive factor of respiratory symptom. However, not all patients with rhinitis present with asthma, and there are some differences between the two condition.

Keywords: Allergic rhinitis, Asthma, Comorbidity.

Introduction

The general pathogenic view of respiratory allergy has deeply changed and evolved during the last two decades. Epidemiologic studies and clinical observation have pointed toward a strong link between allergic rhinitis and asthma. Immunological observation and therapeutic outcomes have further added evidences to the above. Furthermore, an increasing number of evidences have accumulated about (i) the frequent coexistence of rhinitis and asthma; (ii) the possible role of upper respiratory infection in asthma exacerbation; (iii) the presence of rhinitis as a risk factor for developing asthma¹ and, the importance of paranasal sinus infection. These observation taken together lead to the definition of Allergic Rhinobronchitis or United Airway Disease (UAD). Rhinitis is more common than asthma especially in adolescents and young adults. Like asthma its prevalence has increased, (1,2) and it was eventually unknown before the industrial revolution. Pollution and dietary changes have been implicated in this increased incidence.⁽³⁾

Relationship between Rhinitis and Asthma: Epidemiologic studies consistently demonstrate the coexistence of rhinitis and asthma in the same patient. In one study 98.9% of all allergic asthmatic had rhinosinusitis whereas, the prevalence of rhinosinusitis in non-allergic asthmatic subjects was 78.4%.⁽⁴⁾ The author concluded that almost every patient with allergic bronchial asthma had chronic rhinitis. They stated that these findings were consistent with the concept that allergic asthma shall be viewed as a total airway disease and should be treated accordingly. In another study authors found that rhinitis occurs in 78% of patients with allergic asthma and, the focus is often on asthma, so that, the rhinitis may be neglected. In patients with both conditions onset may be with either.⁽⁵⁾ In a study in France, 20,000 households (of which 16,786 i.e. 84% responded) were screened for symptoms of allergic rhinitis (defined as patient having atleast two rhinitis symptom for six months of the year, without any period of remission).⁽⁶⁾ The estimated point prevalence of perennial rhinitis also had history of asthma in comparison to those without perennial rhinitis (3.8%; Odds ratio [OR]=3.26;P \leq 0.001).

In epidemiological studies asthma is present in 20% -50% of patients with rhinitis^(7,8,9) and rhinitis was found to occur in up to 80% of patients with asthma.⁽⁷⁾ In a group of 1245 subjects with documented history of asthma, 6% had perennial allergic rhinitis, 24% had seasonal allergic rhinitis only and 22% had both seasonal and perennial allergic rhinitis.⁽⁷⁾ Leynaert et. al.⁽⁷⁾ analysed data from 34 countries which participated in the European Community Respiratory Health Survey (ECRHS) by taking random sample of 20-44 years old subjects. Asthma was seen in 0.8% of participants without rhinitis, in 6% of participants with seasonal rhinitis only, in 9% of participants with perennial rhinitis only, and in18% of participants with both seasonal and perennial rhinitis. After adjusting for age, sex, family history of asthma, smoking habit, geographical area, and season at the time of examination asthma was found to be strongly associated with rhinitis, not only among atopic subjects [OR= 8.1,95% confidence interval (CI)= 5.4%-12.1%] but also among non atopic subjects [OR,

11.6; 95% CI= 6.2% - 21.9%]. More over the association remained strong when the analysis was confined to non-atopic subjects with IgE level of 80 KU/L or less.

In the Copenhagen Allergy study⁽¹⁰⁾ frequency of rhinitis and asthma related to exposure to pollens, animal dander and mites has been seen. 41% of patients with pollen related rhinitis also had pollen related asthma, whereas, it was 0.1% in those without pollen related rhinitis. For subjects with animal dander allergy, the values were 52% and 0.2% and for subjects with mite related allergy, the values were 41% and 1% respectively. Therefore, in all cases >99% of subjects with allergic asthma also had allergic rhinitis. In a study in Delhi, India investigator found allergic rhinitis cases to be around 10% and asthma to be around 1% in the population during 1964.⁽¹¹⁾ Later studies have reported that in India around 20% to 30% of the population, has allergic rhinitis and around 15% develop asthma,^(12,13)

Aspirin and other non-steroidal anti-inflammatory drugs also induce rhinitis and asthma in certain individual. In a population based study using random samples, aspirin intolerance was found to be more frequent among subjects with allergic rhinitis(8.6%) than those without allergic rhinitis (0.3%; P<0.01).⁽¹⁴⁾ Occupational asthma is a well known entity. But, symptoms of rhinitis are also associated with occupational asthma. In one study 92% of participants with occupational asthma also experienced symptoms of rhinitis.⁽¹⁵⁾ Allergic rhinitis and asthma symptoms can be seen in bakers^(16,17,18) and among people exposed to wood dust.^(19,20,21)

Rhinitis and Non-specific Bronchial Hyperresponsiveness: Patients with allergic rhinitis but with no respiratory symptoms have increased bronchial reactivity to metacholine,^(22,23) although usually less so than asthmatics. Patients with seasonal allergic rhinitis show features of seasonal broncho constriction that is not associated with clinical bronchospasm.⁽²⁴⁾ Patient who are allergic to grass or birch pollen, seasonal increase of carbachol, histamine or methacholine showed bronchial hyper responsiveness and exercise induced bronchoconstriction.^(25,26) In seasonal allergic rhinitis there is slight increase in functional residual capacity and the volume of trapped gas during the hay fever season,⁽²⁷⁾ perhaps reflecting sub-clinical broncho spasm. A mild degree of beta adrenergic hyper- responsiveness in patients with allergic rhinitis could cause a readily reversible increase in resting bronchial tone without asthmatic symptoms. It is frequently possible to obtain bronchoconstriction in patients with allergic rhinitis when allergic challenge is given. However late asthamatic reaction in response to allergic challenge is much more frequent in asthamatic then rhinitis patients.⁽²⁸⁾ Asthma is characterized by hyper reactive airways and epidemiologic studies have established the presence of bronchial hyper responsiveness in patients with allergic rhinitis but no symptoms of clinical

asthma.^(29,30,31) In a more recent study, investigators found that the seasonal rhinitis patients have a reasonably high incidence of non-specific hyperresponsiveness; upto 32% out of season, and upto 48% during the season.⁽³²⁾ Perennial rhinitis patients who are allergic to dust mite or animal dander have the same level of hyper responsiveness as patients of seasonal allergic rhinitis during the season i.e. 48%.

Rhinitis is a Risk Factor for Asthma: Prospective studies have indicated that rhinitis may be a significant risk factor for subsequent development of bronchial asthma. One study, done over a stretch of 23 years reported a three fold increase in the incidence of asthma in patient who previously had allergic rhinitis (without apparent asthma) compared with those who did not had rhinitis (10.5% compared with 3.6%).⁽³³⁾ In another study the investigators showed that clinical, epidemiological and pathophysiological studies suggest a strong functional and immunological relation ship between the nose and the bronchi. Nasal stimulation can induce broncho-constriction by proposed а rhinobronchial reflex and inflammatory mediators originating from nose can trigger bronchial inflammation. Bronchial provocation in sensitized subjects results in nasal inflammation through systemic circulation.⁽³⁴⁾ Allergic rhinitis and extrinsic (or allergic) asthma are part of one atopic (i.e. IgE mediated) condition.^(35,36) Asthma is seen more commonly in patients with perennial rhinitis. In one study it was shown that the presence of physician diagnosed allergic rhinitis in infancy was associated with a doubling of the risk of developing asthma by 11 years of age.⁽³⁷⁾ In an another longitudinal study, the degree to which rhinitis was a risk factor for asthma was studied,⁽³⁸⁾ and even after adjusting for years of follow up, age, sex, atopic status, smoking, presence of COPD, the magnitude of the association was still significantly high (adjusted OR, 3.21%; 95%CI, 2.19-4.71).

The effect of allergic rhinitis was seen in 154 children (aged 3- 17) with the help of a detailed questionnaire which was given to them 8 -11 years later.⁽³⁹⁾ Asthma or wheezing had developed in 19% of subjects and was seen more commonly (P<0.01) in those subjects with perennial allergic rhinitis (34%), than among those with seasonal allergic rhinitis(12.7%).

Conclusion

Epidemiologic studies in general population have provided an evidence that allergic rhinitis and asthma are strongly associated with each other. But, whether allergic rhinitis represents an earlier clinical manifestation of asthma in atopic subjects or if allergic rhinitis itself is a causative factor for asthma is still a matter of debate. More longitudinal studies examining the relationship between these conditions will further provide an important insight into these two co-morbid conditions.

References

- Mygind N. Nasal allergy. 2nd edition. Oxford: *Blackwell* Scientific Publication; 1979: pp219.
- Aberg N. Asthma and allergic rhinitis in Swedish conscripts. *Clinical and Experimental Allergy*, 1989; 19:59-63.
- Ownby DR. Environmental factors versus genetic determinants of childhood inhalant allergies. *Journal of Allergy and Clin Immunol*.1990 Sep;86(3pt1):279-287.
- Kapsali T, Horowitz E, Diemer F, Togias A. Rhinitis is ubiquitous in allergic asthamatics [Abstract]. J Allergy Clin Immunol. 1997;99:s138.
- 5. Pedersen PA, Weeke ER. Asthma and allergic rhinitis in the same patient. *Allergy*.1983 Jan;38(1): 25-9.
- Pariente P, Le-Pen C, Los F, Bousquet J. Quality of life outcomes and the use of antihistamines in a French national population-based sample of patient with perennial rhinitis. *Pharmaco-ecnomics*. 1997, 12:585-95.
- Leynaert B, Bousquet J, Neukirch C, Liard R, Neukirch F. Perennial Rhinitis: an independent risk factor for asthma in non-atopic subjects: results from the European Community Respiratory Health Survey. J Allergy clin immunol. 1999;104:301-304.
- Sibbald B, Rink E. Epidemiology of seasonal and perennial rhinitis: clinical presentation and medical history. *Thorax*. 1991;46:895-901.
- Yawn BP, Yunginger JW, Wollan PC, Reed CE, Silverstein MD, Harris AG. Allergic rhinitis in Rochester Minnesota residents with asthma: frequency and impact on health care charges. *J Allergy Cin Immunol*.1999;103:54-59.
- Linneberg A, Nielsen NH, Madsen F, Frolund F, Dirksen A, Jorgensen T. Secular trends of allergic asthma in Danish adults. The Copenhagen Allergy Study. *Respir Med*.2001;95:258-264.
- 11. Viswanathan R. Defination, incidence, aetiology, and natural history of asthma. *Indian J chest Dis*.1964;6:108-24.
- Anonymous. All India Coordinated Project on Aeroallergens and Human Health Report. Ministry of Environment and Forest, New Delhi;2000.
- Chhabra SK, Gupta CK, Chhabra P, Rajpal S. Prevalence of bronchial asthma in school children in Delhi. *J Asthma*. 1998; 35:291-6.
- 14. Hedman J, Kaprio J, Poussa T, Nieminen MM. Prevalence of asthma aspirin intolerance, nasal polyposis and chronic obstructive pulmonary disease in a population based study. *Int J Epidemiol.* 1999;28:717-722.
- Malo JL, Lemiere C, Desjardins A, Cartier A. Prevalence and intensity of rhinoconjunctivitis in subjects with occupational asthma. *Eur Respir J.* 1997; 10: 1513-1515.
- Brisman J, Jarvholm B. Bakery work, atopy and the incidence of self-reported hay fever and rhinitis. *Eur Respir J*.199913:502-507.
- 17. Baur X. Baker's asthma: causes and prevention. *Int Arch Occup Environ Health*. 1999;72:292-296.
- Musk AW, Venables KM, Crook B, Nunn AJ, Hawkins R, Crook GD et.al. Respiratory symptoms, Lung function, and sensitization to flour in a British bakery. *Br J Ind Med*.1989;46:636-642.
- Fernandez-Rivas M, Perez-Carral C, Senent CJ. OPccupational asthma and rhinitis caused by ash(Fraxinus excelsior) wood dust. *Allery*. 1997; 52:196-199.
- 20. Wilhelmsson B, Jernudd Y, Ripe E, Holmberg K. Nasal hyper sensitivity in wood furniture workers:an allergological and immunological investigation with special reference to mould and wood. *Allergy*. 1984;39:586-595.

- 21. Kanerva L, Vaheri E Occupational allergic rhinitis in Finland. Int Arch Occu Environ Health. 1993, 64:565-568.
- 22. Bakke PS, Baste V, Gulsvik A. Bronchial responsiveness in Norwegian community. American review of respiratory diseases.1991;143:17-22.
- Ramsdal EH, Morris MM, Roberts RS, Hargreave FE. Asymptomatic bronchial hyper responsiveness in rhinitis. *J Allery Clin Immunol.* 1985 may;75(5):573-77.
- 24. Gerblich AA, Schwartz HJ, Chester EH. Seasonal variation of airway function in allergic rhinitis. *J Allergy Clin Immunol.* 1986;77:676-681.
- 25. Karjalainen J, Lindqvist A, Laitinen LA. Seasonal variability of exercise induced asthma especially outdoors: effect of birch pollen allergy. *Clin Exp Allergy*. 1989;19:273-278.
- Madonini E, Briatico-Vangosa G, Pappacoda A, Maccagni G, Cardani A, Saporiti F. Seasonal increase of bronchial reactivity in allergic rhinitis. J Allergy Clin Immunol. 1987;79:358-363.
- Svenonius E, Arborelius M Jr, Kautto R, Lilja B. Lung function studies in children with allergic rhinitis. *Allergy*. 1982 Feb;37(2):87-92.
- Stevens WJ, van Bever HP. Frequency and intensity of late asthmatic reaction after bronchial allergen challenge in asthma and rhinitis. *Allergy*.1989 Sep;44(7):471-476.
- 29. Townley RG, Ryo UX, Kolotkin BM, Kang B. Bronchial sensitivity to methacholine in current and former asthamatic and allergic rhinitis patient and control subject. *J Allergy Clin Immunol.* 1975 Dec;56(6):529-42.
- Fish JE, Rosenthal RR, Batra G, Menkes H, Summer W, Permutt S, Norman P. Airway response to methacholine in allergic and non-allergic subjects. *Am Rev Respir Dis.* 1976;113(5):579-586.
- Braman SS, Burrows AA, Decotus BA, Settipane GA, Corrao WM. Airway hyper responsiveness in allergic rhinitis. A risk factor for asthma. *Chest.* 1987;91:671-674.
- Verdiani P, Di Carlo S, Baronti A. Different prevalence and degree of non specific bronchial hyper reactivity between seasonal and perennial rhinitis. J Allergy Clin Immunol. 1990 Oct;84(4Pt1):576-582.
- 33. Anderson HR, Pottier AC, Strachan DP. Asthma from birth to age 23: incidence and relation to prior and concurrent atopic disease.*Thorax*.1992;47:537-542.
- Palma-Carlos AG, Branco-Ferreira M, Palma-Carlos ML. Allergic rhinitis and asthma: more similarities than differences. *Allergy Immunol(Paris)*. 2001 Jan;33(6):237-41.
- 35. Weiss ST, Sparrow D, O'Connor GT. The inter relationship among allergy, air way responsiveness and asthma. J Asthma. 1993;30(5):329-349.
- Braunstahl GJ. United airway concept: what does it teach us about systemic inflammation in airway disease. Proc Am Thorac Soc.2009 Dec;6(8):652-4.
- Wright AL, Hoberg CJ, Martinez FD, Halonen M, Morgan W, Taussig LM. Epidemiology of physician diagnosed allergic rhinitis in childhood. *Pediatrics*. 1994;94:895-901.
- Guerra S, Sherrill DL, Martinez FD, Barbee RA. Rhinitis as an independent risk factor for adult onset asthma. J Allergy Clin Immunol. 2002;109:419-425.
- Linna O, Kokkonen J, Lukin M. A 10-year prognosis for childhood allergic rhinitis. *Acta Paediatr.* 1992;81:100-102.