

Asthma in the Elderly

a report by

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Asthma is a common disease in the general population, with an overall prevalence in the US in 2002 of 7% reported by the National Center for Health Statistics.¹ It is caused by a complex interaction of inflammatory cells, mediators, and cytokines, which can induce bronchoconstriction and, as a result, airflow obstruction.² This is a cardinal feature of asthma at any age. There is a common misperception that asthma is predominantly a childhood disease. However, in a survey of elderly persons from four communities in the US, the prevalence of physician-diagnosed asthma was 4%, with another 4% having probable asthma (symptoms of asthma without a diagnosis).³ This is similar to other estimates that 7–9% of individuals over the age of 70 have asthma. New-onset asthma is most common in childhood, but it may occur at any age, even in the eighth and ninth decades of life. It is often under-recognized or misdiagnosed in the elderly. A proper evaluation of asthma symptoms can lead to early diagnosis, proper treatment, and avoidance of unnecessary emergency department visits and hospitalizations.^{4,5}

The Diagnosis of Asthma in the Elderly

Unfortunately, the diagnosis of asthma is frequently overlooked in the geriatric population. Some elderly patients are reluctant to admit their symptoms, or consider them a result of normal aging. Under-reporting of symptoms in the elderly may have many causes, including depression, cognitive impairment, social isolation, denial, and blaming of symptoms

on other comorbid illnesses. Even when symptoms are reported, lung function testing to confirm the diagnosis is often not carried out.⁶

On the other hand, the diagnosis of asthma is frequently confused with chronic obstructive pulmonary disease (COPD),⁷ a disease usually, but not always, associated with cigarette smoking. Both are associated with symptoms of shortness of breath, wheezing, cough, sputum production, and airflow obstruction on pulmonary function testing. In one study of a group of elderly asthmatic patients (mean age of 73 years), only 53% of patients had been correctly identified as having asthma. Furthermore, 19.5% of patients were given the wrong diagnosis of COPD.⁷ These patients were more likely to have their onset of symptoms later in life and have greater degrees of disability.

The presence of airflow obstruction can be confirmed by spirometry showing a reduced forced expiratory volume in one second (FEV₁) and ratio of FEV₁/forced vital capacity (FVC). A ratio of less than 70% increases the probability of asthma in an elderly patient with asthma symptoms. A brisk response to a short-acting bronchodilator may demonstrate the second cardinal feature of asthma: reversible airflow obstruction ('a responder'). When airflow obstruction is found in an elderly patient, attempts should be made to demonstrate reversibility following the inhalation of a short-acting beta adrenergic agent such as albuterol. Age is not a significant predictor of the acute bronchodilator response in asthma. Using American Thoracic Society (ATS) criteria (the post-bronchodilator FEV₁ increases by more than 12% and 200cc), the probability of asthma is significantly increased. Some experts feel this confirms the diagnosis of asthma.⁸ While the response to an inhaled bronchodilator is generally greater with asthma (16 versus 11%), many patients with COPD will also meet the ATS reversibility criteria on any given testing day. However, this response may not persist—52.1% of patients will change 'responder' status between visits.⁹ This makes the test less than reliable to confirm the diagnosis of asthma.^{10,11} If complete reversibility of airflow obstruction is documented, COPD, a disease of fixed airway obstruction, is excluded. A brisk post-bronchodilator response enhances the post-test probability of asthma.

A negative bronchodilator response to a short-acting beta agonist, on the other hand, does not rule out a diagnosis of asthma. Population studies have shown that as many as 30% of patients with fixed airflow obstruction have a past history of asthma.¹² Often, treatment over time will improve lung function and, when documented by spirometry, a diagnosis of asthma can be confirmed. Measurements of the FEV₁ over time may offer the best way of making the diagnosis.



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Fixed Airflow Obstruction in Elderly Asthmatics

In addition to bronchoconstriction, causes of airflow obstruction are mucous plugging, bronchial wall edema, inflammatory cell infiltration, airway smooth muscle hypertrophy, and subepithelial fibrosis. All of these architectural changes are collectively referred to as 'airway remodeling.' While some of these changes may be reversible with treatment, at some point they may become permanent, and this results in fixed airflow obstruction.

There is growing evidence that the airway function of young and middle-aged asthmatics declines at a greater rate than that of normal subjects.¹³ The rate of decline increases with increasing age and in those who smoke cigarettes. These effects are variable since not all individuals show a steeper rate of decline. A longer duration and severity of previous asthma are also important factors.^{14,15} In one random survey of 1,200 elderly asthmatics over the age of 65 years, only one in five patients had normal pulmonary function ($FEV_1 >80\%$ predicted), while a similar number showed moderate to severe airflow obstruction ($FEV_1 <50\%$ predicted) after an inhaled short-acting bronchodilator.¹⁶ Since structural changes of emphysema are minimal in elderly asthmatics, airway remodeling is thought to be the main cause of fixed airflow obstruction.¹⁷

While distinguishing asthma from COPD may be very challenging, there are features seen in elderly asthmatics with fixed airflow obstruction that are distinct from COPD caused by cigarette smoking. When compared with patients with COPD of similar age, they have significantly more eosinophils in the peripheral blood, sputum, and bronchoalveolar lavage (BAL), and on bronchial biopsy have higher numbers of neutrophils in the sputum and BAL, have higher ratios of CD4+/CD8+ T cells infiltrating the airway, and have greater thickness of the airway basement membrane than patients with COPD.¹⁸ Other distinguishing features include higher levels of exhaled nitric oxide, lower emphysema scores on high-resolution

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computed tomography (CT) scans, and higher diffusing capacities. Since many of these tests are impractical, not available for widespread use, or not discriminatory enough for clinical use, the most recent American Thoracic Society (ATS)/European Respiratory Society (ERS) Statement on COPD stated: "Some patients with asthma cannot be distinguished from COPD with the current diagnostic tests. The management of these patients should be similar to that of asthma."¹⁹

While the main diagnostic challenge with asthma in the elderly is distinguishing it from COPD, many current or former smokers have both asthma and COPD. Also, asthma in the elderly may be mimicked by, and often confused with, other diseases, such as congestive heart failure, chronic aspiration, gastroesophageal reflux, and tracheobronchial tumors.²⁰

Treatment of Asthma in the Elderly

There are both short-term and long-term therapeutic objectives for every asthmatic patient, recommended by the National Heart, Lung, and Blood Institute (NHLBI).²¹ Short-term objectives are the control of immediate

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symptoms and improvement in lung function. Long-term objectives are those directed to disease prevention and avoidance of emergency room visits and hospitalizations. In order to meet these therapeutic objectives, four components of asthma care should be addressed.

The first of these is monitoring lung function with peak flow meters and/or office spirometry, which is essential in caring for many elderly asthmatics. Older patients with asthma have been shown to deteriorate for longer periods prior to hospital admission for severe acute asthma than younger patients. For example, twice as many elderly patients will show worsening symptoms for more than 14 days before hospital admission compared with a group of younger patients. One reason for this delay may be the blunted perception of breathlessness that has been found in the elderly compared with younger patients.²²

The second component—treatment of asthma with bronchodilator and anti-inflammatory medication—is initially tailored to the patient's needs, and relies on an international staging system of 'asthma severity,' which is based on symptoms and objective measures of lung function.²¹ Inhaled corticosteroids are the preferred anti-inflammatory agent, and are given to those patients who have asthma symptoms and require a short-acting beta agonist for rescue therapy more than twice a week. Monitoring the response to treatment by an assessment of 'asthma control' will be incorporated into the new NHLBI guidelines, as they are in the new Global Initiative for Asthma (GINA) Guidelines.²³

Third, measures should be taken to avoid respiratory irritants that can cause worsening of symptoms. This third cardinal feature of asthma, bronchial hyper-responsiveness, can be demonstrated as an exaggerated bronchoconstrictive response of the airways to a variety of stimuli, such as aeroallergens, histamine, methacholine, cold air, and environmental irritants. Important provocative factors in the elderly include viral respiratory infections, respiratory irritants, and beta adrenoreceptor antagonists (beta-blockers), which are commonly used in this age group for ischemic heart disease, arrhythmias, and hypertension.

Finally, patient education can be a powerful tool in asthma control. Family members can also be helpful, especially with elderly adults. Active participation by a patient in monitoring lung function, avoiding provocative agents, and making decisions regarding medications provides asthma management skills that give a patient the confidence to control his or her own disease.

Medication for Asthma

The medications used to treat the elderly asthmatic are the same as those used to treat younger patients. Inhaled short-acting beta₂ adrenergic

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agonists are the treatment of choice for the acute exacerbation of asthma symptoms. They can be delivered by metered-dose inhaler (MDI) and compressor-driven nebulizers. Unfortunately, many elderly patients are unable to use the MDI properly, even after proper instruction. Inadequate timing of actuation and inhalation is the most frequent error made. Impaired mental function, weakened or deformed hands, and motor or musculoskeletal diseases are other reasons for inadequate MDI use. Despite the minimal systemic absorption seen with the beta agonists, tachycardia and tremor may be observed. While they have been proved to be safe and effective in all age groups, the relative risks for adverse cardiovascular events is high, and caution should be used with these agents in those patients with underlying cardiac conditions.²⁴

Long-acting beta₂ agonists such as salmeterol and formoterol are helpful for long-term maintenance therapy, and should be used in conjunction with an inhaled corticosteroid to improve asthma control.

The goal of asthma therapy is always to control the disease without systemic steroids. Inhaled corticosteroids are safe and effective treatment for elderly asthmatics. They can reduce airway inflammation after several

months of treatment, but long-term treatment is usually necessary. Long-term use of inhaled corticosteroids has been associated with a good safety profile. They are not associated with an increased risk of fractures at standard doses. High doses of inhaled steroids (>1000mcg per day) are capable of causing hypophyseal-pituitary-adrenal (HPA) axis suppression and systemic complications. Local adverse effects, such as hoarseness, dysphonia, cough, and oral candidiasis, do occur, but can usually be avoided by the use of a spacer or holding chamber.

Prognosis

Longitudinal studies of asthmatic populations have shown that remission from asthma is uncommon in older age groups, occurring in about 20% of patients. Elderly asthmatics with severe symptoms, long-standing disease, reduced pulmonary function, or a concomitant diagnosis of COPD are much less likely to have a remission. The risk for hospitalization in the asthmatic over the age of 65 doubles compared with younger patients, especially for women and non-whites.²⁵ In one study, the risk of re-

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hospitalization was 23% after one year, and 12% of patients died.²⁶ Asthmatics over the age of 65 years account for 60% of asthma-related deaths overall, and black females aged 65 years and older have the highest crude asthma mortality rates.²⁷ Despite the at times severe symptoms and physiological impairment, most elderly patients with asthma can lead active productive lives with the use of appropriate therapy.⁵ ■

1. Asthma Health Care Use and Mortality, 2002 (www.cdc.gov/nchs/products/pubs/pubd/hestat/asthma/asthma.htm).
2. Bousquet J, Jeffery PK, Busse WW, Asthma: From bronchoconstriction to airways inflammation and remodeling, *Am J Respir Crit Care Med*, 2000;161:1720-45.
3. Enright PL, McClelland RL, Newman AB, et al., Underdiagnosis and undertreatment of asthma in the elderly, *Chest*, 1999;116:603-13.
4. NAEP Working Group Report: consideration for diagnosis and managing asthma in the elderly. 1996 National Institute of Health, National Heart, Lung, and Blood Institute, Bethesda, MD: publication No. 96-3662.
5. Braman SS, Asthma in the elderly, *Clin Geriatr Med*, 2003;19:57-75.
6. Bauer BA, Reed CE, Yunginger JW, et al., Incidence and outcomes of asthma in the elderly. A population-based study in Rochester, Minnesota, *Chest*, 1997;111:303-10.
7. Bellia V, Battaglia S, Catalano F, et al., Aging and disability affect misdiagnosis of COPD in elderly asthmatics: the SARA study, *Chest*, 2003;123:1066-72.
8. Enright PL, The diagnosis and management of asthma is much tougher in older patients, *Curr Opin Allergy Clin Immunol*, 2002;2:175-81.
9. Calverly PM, Burge PS, Spencer S, et al., Bronchodilator reversibility testing in chronic obstructive pulmonary disease, *Thorax*, 2003;58(8):659-64.
10. Meslier N, Racineux JL, Six P, et al., Diagnostic value of reversibility of chronic airway obstruction to separate asthma from chronic bronchitis: a statistical approach, *Eur Respir J*, 1989;2:497-505.
11. Kesten S, Rebeck AS, Is the short-term response to inhaled beta-adrenergic agonist sensitive or specific for distinguishing between asthma and COPD?, *Chest*, 1994;105(4):1042-5.
12. Mannino DM, Gagnon RC, Petty TL, et al., Obstructive lung disease and low lung function in adults in the United States: data from the National Health and Nutrition Examination Survey 1988-1994, *Arch Intern Med*, 2000;160:1683-89.
13. Lange P, Parner J, Vestbo J, et al., A 15-year follow-up study of ventilatory function in adults with asthma, *N Engl J Med*, 1998;339(17):1194-1200.
14. Braman SS, Kaemmerlen JT, Davis SM, Asthma in the elderly. A comparison between patients with recently acquired and long-standing disease, *Am Rev Respir Dis*, 1991;143:336-40.
15. Cassino C, Berger KI, Goldring RM, et al., Duration of asthma and physiologic outcomes in elderly nonsmokers, *Am J Respir Crit Care Med*, 2000;162:1423-8.
16. Reed CE, The natural history of asthma in adults: The problem of irreversibility, *J Allergy Clin Immunol*, 2000;103:539-47.
17. Bai TR, Cooper J, Koelmeyer T, et al., The effect of age and duration of disease on airway structure in fatal asthma, *Am J Respir Crit Care Med*, 2000;162:663-9.
18. Fabbri LM, Romagnoli M, Corbetta L, et al., Differences in airway inflammation in patients with fixed airflow obstruction due to asthma or chronic obstructive pulmonary disease, *Am J Respir Crit Care Med*, 2003;167:418-24.
19. Celli BR, MacNee W, Agusti A, et al., Standards for the diagnosis and treatment of patients with COPD: a summary of the ATS/ERS position paper, *Eur Respir J*, 2004;23:932-46.
20. Braman SS, Davis SM, Wheezing in the elderly. Asthma and other causes, *Clin Geriatr Med*, 1986;2:269-83.
21. National Asthma Education and Prevention Program Expert Panel Report 2: Guidelines for the Diagnosis and Management of Asthma (www.nhlbi.nih.gov/guidelines/asthma/asthupdt.htm).
22. Connolly MJ, Crowley JJ, Charan NB, et al., Reduced subjective awareness of bronchoconstriction provoked by methacholine in elderly asthmatic and normal subjects as measured on a simple awareness scale, *Thorax*, 1992;47:410-13.
23. GINA Report, Global Strategy for Asthma Management and Prevention (<http://www.ginasthma.com/GuidelinesResources>).
24. Salpeter SR, Ormiston TM, Salpeter EE, Cardiovascular effects of beta-agonists in patients with asthma and COPD: a meta-analysis, *Chest*, 2004;125:2309-21.
25. Diette GB, Krishnan JA, Dominici F, et al., Asthma in older patients: factors associated with hospitalization, *Arch Intern Med*, 2002;162:1123-32.
26. Hartert TV, Speroff T, Togias A, et al., Risk factors for recurrent hospital visits and death among a population of indigent older adults with asthma, *An Allergy Asthma Immunol*, 2002;89:467-74.
27. Moorman JE, Mannino DM, Increasing US asthma mortality rate: who is really dying?, *J Asthma*, 2001;38:65-70.