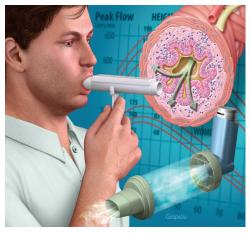
Management of Acute Asthma Exacerbations

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Asthma exacerbations can be classified as mild, moderate, severe, or life threatening. Criteria for exacerbation severity are based on symptoms and physical examination parameters, as well as lung function and oxygen saturation. In patients with a peak expiratory flow of 50 to 79 percent of their personal best, up to two treatments of two to six inhalations of short-acting beta₂ agonists 20 minutes apart followed by a reassessment of peak expiratory flow and symptoms may be safely employed at home. Administration using a hand-held metered-dose inhaler with a spacer device is at least equivalent to nebulized beta₂ agonist therapy in children and adults. In the ambulatory and emer-

gency department settings, the goals of treatment are correction of severe hypoxemia, rapid reversal of airflow obstruction, and reduction of the risk of relapse. Multiple doses of inhaled anticholinergic medication combined with beta₂ agonists improve lung function and decrease hospitalization in school-age children with severe asthma exacerbations. Intravenous magnesium sulfate has been shown to significantly increase lung function and decrease the necessity of hospitalization in children. The administration of systemic corticosteroids within one hour of emergency department presentation decreases the need for hospitalization, with the most pronounced effect in patients with severe exacerbations. Airway inflammation can persist for days to weeks after an acute attack; therefore, more intensive treatment should be continued after discharge until symptoms and peak expiratory flow return to baseline. (*Am Fam Physician*. 2011;84(1):40-47. Copyright © 2011 American Academy of Family Physicians.)



LUSTRATION BY JOHN KARAPELOU

▶ Patient information: A handout on how to treat an asthma attack, written by the authors of this article, is provided on page 49. n 2005, the prevalence of asthma in the United States was nearly 8 percent (close to 9 percent in children younger than 18 years), and approximately 4 percent of Americans (5 percent of children) experienced an asthma attack.^{1,2} There have been many advances in medical therapy to prevent the worsening of asthma symptoms, including an improved understanding of asthma etiology, identification of risk factors for asthma exacerbations, and evidence supporting the benefits of written asthma action plans.

One study of children up to 18 years of age presenting to the emergency department with acute asthma symptoms identified multiple risk factors for a subsequent emergency department visit: age younger than two years, black race or Hispanic ethnicity, persistent asthma, public health insurance, lower asthma quality-of-life scores, and increased use of the health care system during the previous 12 months.³ In adults, variables associated with relapse within eight

weeks of an asthma exacerbation include three or more visits for emergent care in the preceding six months, difficulty performing daily activities because of physical health in the preceding four weeks, and patient selfdischarge from care within 24 hours of hospital admission without achieving 50 percent predicted peak expiratory flow (PEF).4 However, regular monitoring of PEF does not help predict an asthma exacerbation.⁵ Other risk factors for developing an asthma exacerbation include allergen triggers (e.g., pets, seasonal allergens, smoke exposure) and improper use of medications (e.g., not using a spacer, improper use of an inhaler or other delivery device).6

In persons older than two years with asthma, neither the injectable nor the intranasal influenza vaccine increases the likelihood of an asthma exacerbation in the period immediately following vaccination. However, one study of infants found an increase in wheezing and hospital admissions

Clinical recommendation	Evidence rating	References	Comments
Inhaled short-acting beta $_{2}$ agonists are the cornerstones of treatment for acute asthma.	С	14-16	_
An inhaler with a spacer is equivalent to nebulized short-acting beta ₂ agonist therapy in children and adults.	А	17, 18	_
Continuous beta ₂ agonist administration reduces hospital admissions in patients with severe acute asthma.	А	21	_
Inhaled anticholinergic medication improves lung function and decreases hospitalization in school-age children with severe asthma exacerbations.	А	24, 25	When multiple doses are used in combination with short-acting beta ₂ agonist
Intravenous magnesium sulfate increases lung function and decreases hospitalizations in children with an acute asthma exacerbation.	А	29	_
The administration of systemic corticosteroids within one hour of emergency department presentation decreases the need for hospitalization.	А	30	Largest effect noted in patients with severe asthm
Oral and parenteral corticosteroids are equally effective in preventing hospital admission in children.	В	31	_

A = consistent, good-quality patient-oriented evidence; B = inconsistent or limited-quality patient-oriented evidence; C = consensus, disease-oriented evidence, usual practice, expert opinion, or case series. For information about the SORT evidence rating system, go to http://www.aafp.org/afpsort.xml.

Table 1. Classifications	of Severity of an	Asthma Evacarhation
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Degree of severity	Symptoms and signs	Initial PEF (or FEV₁)	Clinical course
Mild	Dyspnea only with activity (assess tachypnea in young children)	PEF ≥ 70 percent of predicted or personal best	Usually treated at home Prompt relief with inhaled short-acting beta ₂ agonist Possible short course of oral systemic corticosteroids
Moderate	Dyspnea interferes with or limits usual activity	PEF 40 to 69 percent of predicted or personal best	Usually requires office or emergency department visit Relief from frequent inhaled short-acting beta₂ agonist Oral systemic corticosteroids; some symptoms last for one to two days after treatment begins
Severe	Dyspnea at rest; interferes with conversation	PEF < 40 percent of predicted or personal best	Usually requires emergency department visit and likely hospitalization Partial relief from frequent inhaled short-acting beta ₂ agonist Oral systemic corticosteroids; some symptoms last for more than three days after treatment begins Adjunctive therapies are helpful
Subset: life threatening	Too dyspneic to speak; perspiration	PEF < 25 percent of predicted or personal best	Requires emergency department visit/hospitalization; possible intensive care unit Minimal or no relief from frequent inhaled short-acting beta ₂ agonist Intravenous corticosteroids Adjunctive therapies are helpful

 FEV_1 = forced expiratory volume in one second; PEF = peak expiratory flow.

Adapted from the National Heart Lung and Blood Institute. National Asthma Education and Prevention Program. Expert panel report 3: Guidelines for the diagnosis and management of asthma; 2007:375. http://www.nhlbi.nih.gov/guidelines/asthma/asthgdln.htm.

after intranasal influenza vaccination.⁷ Seasonal influenza vaccine does not reduce the risk of developing an asthma exacerbation. Influenza vaccination appears to improve asthma-related quality-of-life in children during influenza season.⁷

Diagnosis

Asthma exacerbations can be classified as mild, moderate, severe, or life threatening (*Table 1*).⁶ Criteria for severity are based on symptoms and physical examination parameters, as well as lung function and oxygen

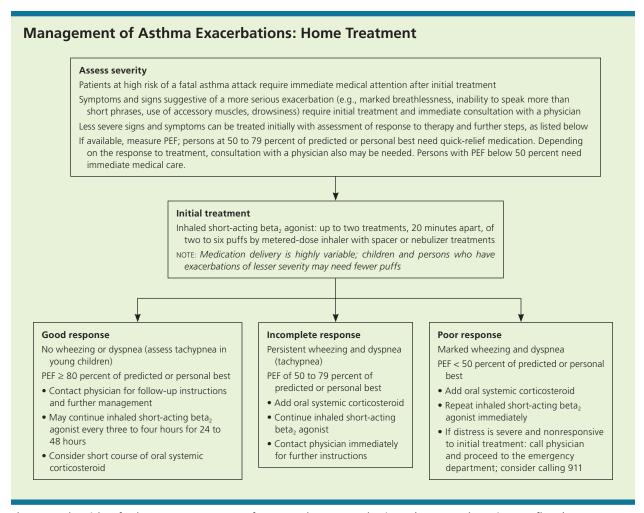


Figure 1. Algorithm for home management of acute asthma exacerbations. (PEF = peak expiratory flow.)

Adapted from the National Heart Lung and Blood Institute. National Asthma Education and Prevention Program. Expert panel report 3: Guidelines for the diagnosis and management of asthma; 2007:382. http://www.nhlbi.nih.gov/guidelines/asthma/asthgdln.htm.

saturation. Although no single parameter has been identified to assess exacerbation severity, lung function is a useful method of assessment, with a PEF of 40 percent or less of predicted function indicating a severe attack in patients five years or older.⁶ The most useful signs for determining the severity of an asthma exacerbation in children younger than five years, or any child unable to perform a PEF, include the use of accessory muscles of respiration, chest wall retractions, tachypnea greater than 60 breaths per minute, cyanosis, and the presence of inspiratory and expiratory wheezing.⁸ For all patients, pulse oximetry on room air is a useful initial assessment. An oxygen saturation of less than 92 to 94 percent one hour after beginning standard treatment is a strong predictor of the need for hospitalization.⁶

Laboratory data are not required for most patients with acute exacerbations. Some tests that may be useful include complete blood count, serum theophylline, and basic chemistries. Chest radiography is not routinely recommended because it has not been shown to alter the care of patients with an uncomplicated asthma

exacerbation.⁹ Measurement of arterial blood gases may be considered if hypoventilation is suspected. Electrocardiography is rarely helpful, unless there is a history or suspicion of cardiac disease.⁶

Management HOME TREATMENT

Early treatment is the most effective strategy for managing asthma exacerbations. It is essential to teach patients how to monitor signs and symptoms, and take appropriate action. Patients who have a written asthma action plan and appropriate medication can often manage mild exacerbations at home (*Figure 1*⁶). Key components of an asthma action plan that have reduced emergency department visits and hospitalization include standard written instructions; criteria based on symptoms or PEF (compared with personal best) to trigger action; two to three action points; and individualized, written instructions on the use of inhaled or oral corticosteroids. Patients at risk of asthma-related death may need more intensive treatment in a monitored setting at the first sign of an

exacerbation (*Table 2*⁶). These patients should have an asthma action plan that emphasizes early communication with their physician.

In children five to 12 years of age with frequent acute exacerbations, a short course of oral prednisolone at the onset of worsening symptoms produced a modest benefit in terms of decreased symptoms, health resource use, and absence from school. Patient- or parent-initiated increases in the dosage of inhaled corticosteroids have been proposed to help with deteriorating asthma symptoms. The data are insufficient to make a recommendation for children; however, a meta-analysis of data from more than 1,200 adults confirms that increasing the dosage does not reduce the risk of a subsequent asthma exacerbation requiring oral corticosteroids.

A randomized controlled trial examined the use of parent-initiated montelukast (Singulair; 4 mg for children two to five years of age and 5 mg for children six to 14 years of age) in children with intermittent asthma, defined as three to six episodes of asthma requiring acute hospital- or office-based care with symptom- and medication-free periods between episodes. When given at the onset of asthma or upper respiratory tract infection symptoms, montelukast therapy resulted in a reduction in unscheduled health care visits and time lost from work and school or childcare.¹³

Inhaled short-acting beta₂ agonists are the cornerstones of treatment for patients with acute asthma.¹⁴⁻¹⁶ In patients with a PEF of 50 to 79 percent of their personal best, up to two treatments of two to six inhalations of a short-acting beta₂ agonist may be safely employed at

Table 2. Risk Factors for Asthma-Related Death

Comorbidities (i.e., cardiovascular disease or other chronic lung disease)

Difficulty perceiving airway obstruction or severity of exacerbation

Illicit drug use

Low socioeconomic status or inner-city residence

Major psychosocial problems or psychiatric disorders

Previous severe exacerbation (e.g., intubation or admission to intensive care unit for asthma)

Two or more hospitalizations or three or more emergency department visits in the past year

Two or more refills of short-acting beta₂ agonists per month

Adapted from the National Heart Lung and Blood Institute. National Asthma Education and Prevention Program. Expert panel report 3: Guidelines for the diagnosis and management of asthma; 2007:377. http://www.nhlbi.nih.gov/guidelines/asthma/asthgdln.htm.

home. Treatments should be 20 minutes apart followed by a reassessment of PEF and symptoms.⁶ Patients who do not achieve a PEF of at least 80 percent of their personal best after two treatments should contact their physician for further instructions. Patients whose PEF declines after treatment should contact their physician and seek emergent care.

Multiple studies have shown that administration using a hand-held metered-dose inhaler with a spacer device is at least equivalent to nebulized short-acting

beta₂ agonist therapy in children older than one year (four puffs per dose) and adults (six puffs per dose).¹⁷ Homemade spacers, such as plastic bottles, foam or paper cups,

A homemade spacer, such as a plastic bottle or paper cup, can be as effective as a commercial spacer.

cardboard tubes, and paper spacers, can be as effective as commercial spacers for the treatment of acute asthma exacerbations.¹⁸ There is no demonstrable difference in terms of safety or effectiveness between levalbuterol (Xopenex) and albuterol.¹⁹

EMERGENCY DEPARTMENT TREATMENT

In the ambulatory and emergency department settings, the goals of treatment are correction of severe hypoxemia, rapid reversal of airflow obstruction, and reduction of the risk of relapse by intensifying therapy and carefully monitoring response (Figure 2).⁶ Correction of hypoxemia and rapid reversal of airflow obstruction are best achieved by oxygen administration and repetitive treatment with short-acting beta₂ agonists. Early use of systemic corticosteroids can reduce the risk of relapse.

The administration of oxygen to maintain saturation of at least 94 percent is recommended in all patients presenting with a moderate to severe asthma exacerbation. Oxygen should be administered as soon as possible, preferably in the prehospital phase in an office setting or in transport by emergency medical services. It has been proposed that the helium and oxygen mixture (heliox), which has a lower density than oxygen, flows more easily through constricted airways and, as a result, improves outcomes in asthma exacerbations. However, there are insufficient data to support the use of heliox in the treatment of acute asthma exacerbations.²⁰

Inhaled short-acting beta₂ agonist treatment is the mainstay of office or emergency department treatment of moderate to severe asthma exacerbations. If the patient can tolerate a measurement of PEF or forced expiratory volume in one second (FEV₁), an initial value should be

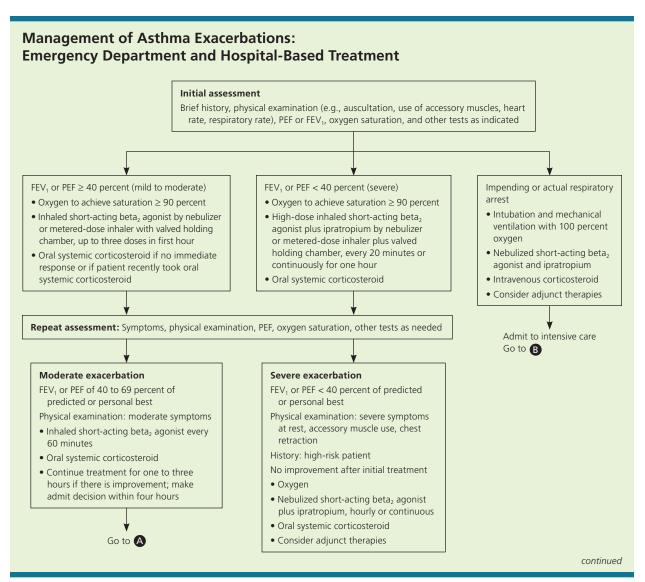


Figure 2. Algorithm for emergency department and inpatient management of acute asthma exacerbations. (FEV₁ = forced expiratory volume in one second; PEF = peak expiratory flow.)

obtained and repeated to monitor treatment response. In patients with severe exacerbations, continuous beta₂ agonist administration has been shown to improve pulmonary function measurements and reduce hospital admission with no notable differences in pulse, blood pressure, or tremor.²¹ The use of high-dose albuterol (7.5 mg via nebulizer every 20 minutes for three doses)²² and intravenous beta₂ agonists does not appear to be beneficial and is not recommended.²³

A meta-analysis of randomized controlled trials compared the combination of inhaled anticholinergics and beta₂ agonists with beta₂ agonists alone in children one to 18 years of age with mild, moderate, or severe exacerbations of asthma. The results showed that adding multiple doses of inhaled anticholinergic medication improves lung function and decreases hospitalizations in school-aged children with severe asthma

exacerbations.^{24,25} The usefulness of inhaled ipratropium for the treatment of asthma exacerbations in adults is less clear, but it does appear to benefit those with a severe exacerbation.^{26,27}

The addition of intravenous magnesium sulfate to standard therapy has been studied in adults and children with divergent results. In adults with severe exacerbations of asthma (PEF of 25 to 30 percent or less of predicted function), intravenous magnesium sulfate therapy resulted in slightly better lung function but no change in rates of hospitalization.²⁸ In children one to 18 years of age, intravenous magnesium sulfate (25 to 100 mg per kg) has been demonstrated to significantly increase lung function and to decrease hospitalizations. Nebulized magnesium sulfate has a weak effect on respiratory function and hospital admission rates in adults, and no effect on either outcome in children.²⁹

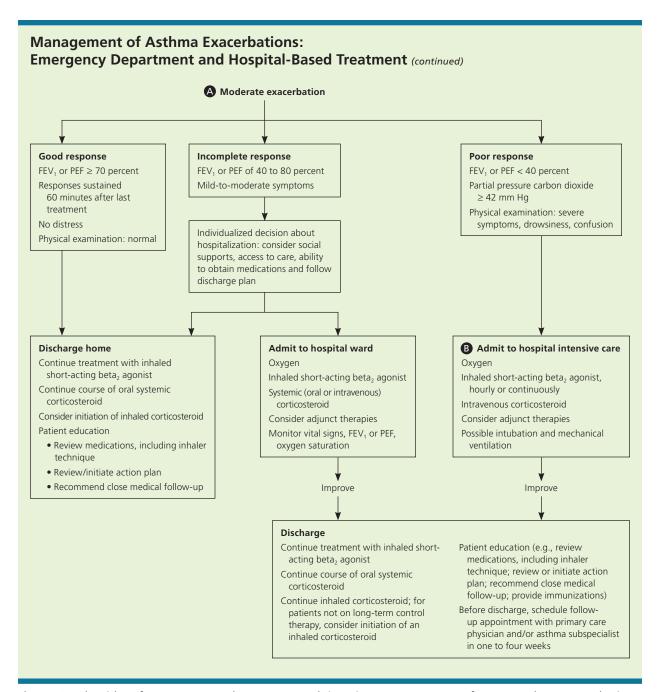


Figure 2. Algorithm for emergency department and inpatient management of acute asthma exacerbations. (FEV $_1$ = forced expiratory volume in one second; PEF = peak expiratory flow.)

Adapted from the National Heart Lung and Blood Institute. National Asthma Education and Prevention Program. Expert panel report 3: Guidelines for the diagnosis and management of asthma; 2007:388. http://www.nhlbi.nih.gov/guidelines/asthma/asthgdln.htm.

The administration of systemic corticosteroids (500 mg hydrocortisone sodium succinate injection [Solu-Cortef] or 125 mg methylprednisolone sodium succinate injection [Solu-Medrol] in adults, or 1 to 2 mg per kg of prednisone or prednisolone in children one to 18 years of age) within one hour of emergency department presentation decreases the need for hospitalization. In a Cochrane review, the most pronounced effect occurred in patients with severe exacerbations.³⁰ Oral

and parenteral corticosteroids are equally effective in preventing hospital admission in children, but only parenteral corticosteroids have been studied in adults.³¹ There is insufficient evidence to recommend the use of inhaled corticosteroids in place of or in conjunction with systemic corticosteroids at the time of discharge from the emergency department. Inhaled corticosteroids do not prevent relapse of symptoms requiring admission or improve quality of life or symptom scores.³²

Asthma Exacerbations

In adults and in hospitalized children one to 16 years of age, corticosteroid use resulted in earlier discharge and fewer symptomatic relapses.³³⁻³⁵ The optimal dosage in children is unknown,34 but in adults, lower dosages (80 mg or less per day of methylprednisolone [Depo-Medrol or 400 mg or less per day of hydrocortisone) are equal to higher dosages in the improvement of lung function, adverse effects, and rates of respiratory failure.35

The addition of intravenous aminophylline to conventional therapy in children and adults has no additional benefit in reducing hospital admissions. It does significantly increase the risk of adverse effects, including vomiting, palpitations, and arrhythmias. 36,37 There are insufficient data to recommend for or against the use of antibiotics in the treatment of acute exacerbations.³⁸ In a Cochrane review, one randomized controlled trial of 30 adults examined the use of noninvasive positive pressure ventilation in the treatment of severe acute exacerbations of asthma as an adjunct to usual care. The intervention showed promising results in objective measure of lung function and reduced rates of hospitalization, but the data are insufficient to make broad recommendations for the use of noninvasive positive pressure ventilation.³⁹ Drinking large amounts of water, high-dose mucolytics, antihistamines, chest physiotherapy, and sedation are all unproven treatments.⁶

POSTDISCHARGE CARE

Patients sent home from the emergency department with systemic corticosteroids (a five- to 10-day nontapering course of 50- to 100-mg prednisone per day in adults) have decreased relapse of asthma symptoms, future hospitalizations, and use of short-acting beta₂ agonists.^{40,41} Although seven to 10 days is the usual treatment duration for oral corticosteroids, three days of therapy (1 mg per kg of prednisone) has been shown to be as effective as five days for the complete resolution of symptoms within one week in children two to 15 years of age.42 There are insufficient data to recommend the initiation of montelukast in place of oral corticosteroids or the use of inhaled corticosteroids in combination with oral corticosteroids at the time of discharge to prevent a relapse of asthma symptoms. 43,44

Allergen avoidance is routinely recommended after emergency department discharge to decrease further acute exacerbations of asthma. Despite multiple trials of allergen control, there are no data showing that pet allergen or dust mite allergen avoidance techniques successfully reduce allergens in the home to levels that improve asthma symptoms. 45,46

Regardless of the therapy chosen in the acute care setting, step-up therapy should be continued for several days to weeks after discharge. Because exacerbations vary in severity, close communication between patients and physicians is required. Symptoms may be controlled quickly, but airway inflammation may persist for two to three weeks.⁴⁷ Scheduled dosing with inhaled beta, agonists should be continued until symptoms and PEF return to baseline.

Data Sources: The National Guidelines Clearinghouse was searched for guidelines on asthma care. The National Asthma Education and Prevention Program's "Expert Panel 3 Report: Guidelines for the Diagnosis and Management of Asthma" section on management of asthma exacerbations was reviewed. Ovid Medline was searched for new information related to the major recommendations of both. PubMed was searched using the key terms asthma + acute + exacerbation. The Cochrane database and Essential Evidence Plus were searched for information pertaining to asthma exacerbations. Search dates: March 2010 and April 2010. Searches on select topics were performed weekly in May and June 2010, with a repeat search in November 2010.

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