



## Quality Improvement Guideline

### Asthma Assessment and Management and Use of FeNO

**Created:**  
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#### **Purpose**

This is a guide for the assessment and management of asthma. It also contains information regarding the use of FeNO in the screening and diagnosis of asthma.

#### **Scope**

The patient population is any patient 5 years or older with a diagnosis of asthma.

#### **Guideline**

##### **Asthma Assessment and Management**

Please refer to the 2018 Global Initiative for Asthma (GINA) Pocket Guide for Asthma Management and Prevention for Adults and Children Older than 5 years. See Appendix for the Pocket Guide.

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The GINA Pocket Guide is also available at <https://ginasthma.org/2018-gina-report-global-strategy-for-asthma-management-and-prevention/>

**NOTE: These guidelines are not to be distributed outside of BSWQA.**

## Use of FeNO

FeNO measurement should be considered in patients to adjust and evaluate the benefits of inhaled corticosteroids and to assess adherence to medication therapy. It should not be used as a diagnostic tool.

### **Additional Resources**

For Baylor Scott & White Epic users, there is an asthma action plan that can be completed and printed for the patient.

**Guideline Developers:** BSWH/BSWQA Asthma/COPD Council

**Approved By:** BSWQA Primary Care Subcommittee, BSWQA Pediatric Subcommittee, BSWQA Quality Improvement Committee, and BSWQA Board of Managers

**BSWQA CY18 Related Focus Measures:** All-Cause 30-day Readmissions, Avoidable ED Visits per 1000, and Avoidable Admits per 1,000

### **References**

Global Initiative for Asthma. Pocket Guide for Asthma Management and Prevention, 2017. Available from: [www.ginasthma.org](http://www.ginasthma.org).

An Official ATS Clinical Practice Guideline: Interpretation of Exhaled Nitric Oxide Levels (FeNO) for Clinical Applications Am J Respir Crit Care Med Vol 184. pp 602–615, 2011.

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# POCKET GUIDE FOR ASTHMA MANAGEMENT AND PREVENTION

(for Adults and Children Older than 5 Years)



*A Pocket Guide for Health Professionals*  
Updated 2018

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**BASED ON THE GLOBAL STRATEGY FOR ASTHMA  
MANAGEMENT AND PREVENTION**

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## **GLOBAL INITIATIVE FOR ASTHMA**

# **POCKET GUIDE FOR HEALTH PROFESSIONALS 2018**

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Names of members of the GINA Committees are listed on page 28.

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# PREFACE

Asthma affects an estimated 300 million individuals worldwide. It is a serious global health problem affecting all age groups, with increasing prevalence in many developing countries, rising treatment costs, and a rising burden for patients and the community. Asthma still imposes an unacceptable burden on health care systems, and on society through loss of productivity in the workplace and, especially for pediatric asthma, disruption to the family, and it still contributes to many deaths worldwide, including amongst young people.

Health care providers managing asthma face different issues around the world, depending on the local context, the health system, and access to resources.

The **Global Initiative for Asthma (GINA)** was established to increase awareness about asthma among health professionals, public health authorities and the community, and to improve prevention and management through a coordinated worldwide effort. GINA prepares scientific reports on asthma, encourages dissemination and implementation of the recommendations, and promotes international collaboration on asthma research.

**The Global Strategy for Asthma Management and Prevention** was extensively revised in 2014 to provide a comprehensive and integrated approach to asthma management that can be adapted for local conditions and for individual patients. It focuses not only on the existing strong evidence base, but also on clarity of language and on providing tools for feasible implementation in clinical practice. The report has been updated each year since then.

The GINA 2018 report and other GINA publications listed on page 28 can be obtained from [www.ginasthma.org](http://www.ginasthma.org).

The reader acknowledges that this **Pocket Guide** is a brief summary of the GINA 2018 report, for primary health care providers. It does NOT contain all of the information required for managing asthma, for example, about safety of treatments, and it should be used in conjunction with the full GINA 2018 report and with the health professional's own clinical judgment. GINA cannot be held liable or responsible for inappropriate healthcare associated with the use of this document, including any use which is not in accordance with applicable local or national regulations or guidelines.

## WHAT IS KNOWN ABOUT ASTHMA?

**Asthma is a common and potentially serious chronic disease** that imposes a substantial burden on patients, their families and the community. It causes respiratory symptoms, limitation of activity, and flare-ups (attacks) that sometimes require urgent health care and may be fatal.

**Fortunately...asthma can be effectively treated**, and most patients can achieve good control of their asthma. When asthma is under good control, patients can:

- ✓ Avoid troublesome symptoms during day and night
- ✓ Need little or no reliever medication
- ✓ Have productive, physically active lives
- ✓ Have normal or near normal lung function
- ✓ Avoid serious asthma flare-ups (exacerbations, or attacks)

**What is asthma?** Asthma causes symptoms such as wheezing, shortness of breath, chest tightness and cough that vary over time in their occurrence, frequency and intensity.

These symptoms are associated with variable expiratory airflow, i.e. difficulty breathing air out of the lungs due to bronchoconstriction (airway narrowing), airway wall thickening, and increased mucus. Some variation in airflow can also occur in people without asthma, but it is greater in asthma. There are different types of asthma, with different underlying disease processes.

**Factors that may trigger or worsen asthma symptoms** include viral infections, domestic or occupational allergens (e.g. house dust mite, pollens, cockroach), tobacco smoke, exercise and stress. These responses are more likely when asthma is uncontrolled. Some drugs can induce or trigger asthma, e.g. beta-blockers, and (in some patients), aspirin or other NSAIDs.

**Asthma flare-ups** (also called exacerbations or attacks) can be fatal. They are more common and more severe when asthma is uncontrolled, or in some high-risk patients. However, flare-ups may occur even in people taking asthma treatment, so all patients should have an asthma action plan.

**A stepwise approach to treatment**, customized to the individual patient, takes into account the effectiveness of available medications, their safety, and their cost to the payer or patient.

**Regular controller treatment**, particularly with inhaled corticosteroid (ICS)-containing medications, markedly reduces the frequency and severity of asthma symptoms and the risk of having a flare-up.

Asthma is a common condition, affecting all levels of society. Olympic athletes, famous leaders and celebrities, and ordinary people live **successful and active lives with asthma**.

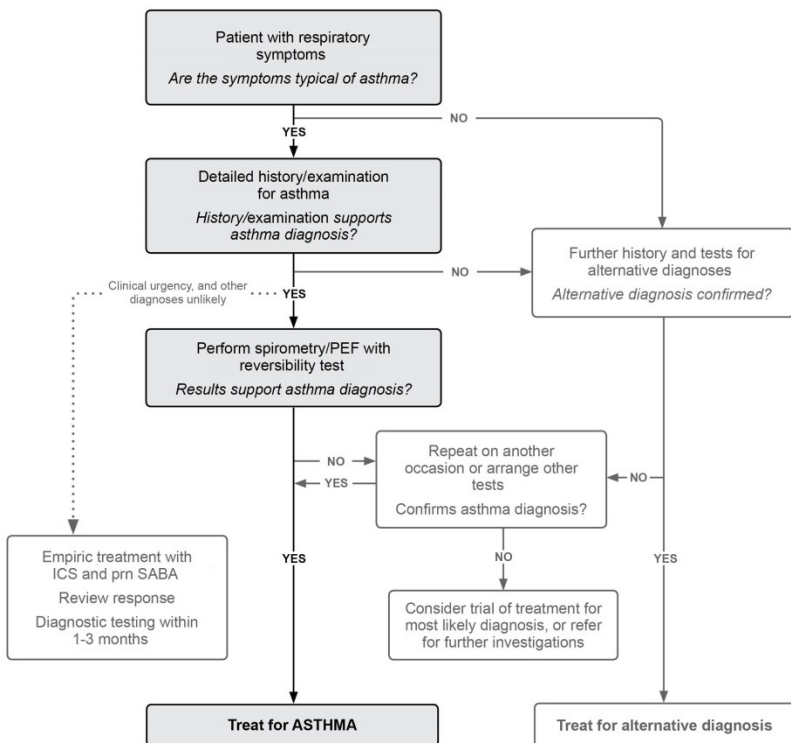
# MAKING THE DIAGNOSIS OF ASTHMA

Asthma is a disease with many variations (heterogeneous), usually characterized by chronic airway inflammation. Asthma has two key defining features:

- a history of respiratory symptoms such as wheeze, shortness of breath, chest tightness and cough that vary over time and in intensity, AND
- variable expiratory airflow limitation.

A flow-chart for making the diagnosis in clinical practice is shown in Box 1, with the specific criteria for diagnosing asthma in Box 2.

## Box 1. Diagnostic flow-chart for asthma in clinical practice



The **diagnosis of asthma** should be confirmed and, for future reference, the evidence documented in the patient's notes. Depending on clinical urgency and access to resources, this should preferably be done before starting controller treatment. Confirming the diagnosis of asthma is more difficult after treatment has been started (see p7).



# CRITERIA FOR MAKING THE DIAGNOSIS OF ASTHMA

## Box 2. Features used in making the diagnosis of asthma

### 1. A history of variable respiratory symptoms

Typical symptoms are wheeze, shortness of breath, chest tightness, cough

- People with asthma generally have more than one of these symptoms
- The symptoms occur variably over time and vary in intensity
- The symptoms often occur or are worse at night or on waking
- Symptoms are often triggered by exercise, laughter, allergens or cold air
- Symptoms often occur with or worsen with viral infections

### 2. Evidence of variable expiratory airflow limitation

- At least once during the diagnostic process, e.g. when FEV<sub>1</sub> is low, document that the FEV<sub>1</sub>/FVC ratio is reduced. The FEV<sub>1</sub>/FVC ratio is normally more than 0.75–0.80 in adults, and more than 0.90 in children.
- Document that variation in lung function is greater than in healthy people. For example:
  - FEV<sub>1</sub> increases by more than 12% and 200mL (in children, >12% of the predicted value) after inhaling a bronchodilator. This is called ‘bronchodilator reversibility’.
  - Average daily diurnal PEF variability\* is >10% (in children, >13%)
  - FEV<sub>1</sub> increases by more than 12% and 200mL from baseline (in children, by >12% of the predicted value) after 4 weeks of anti-inflammatory treatment (outside respiratory infections)
- The greater the variation, or the more times excess variation is seen, the more confident you can be of the diagnosis of asthma.
- Testing may need to be repeated during symptoms, in the early morning, or after withholding bronchodilator medications.
- Bronchodilator reversibility may be absent during severe exacerbations or viral infections. If bronchodilator reversibility is not present when it is first tested, the next step depends on the clinical urgency and availability of other tests.
- For other tests to assist in diagnosis, including bronchial challenge tests, see Chapter 1 of the GINA 2018 report.

\*Calculated from twice daily readings (best of 3 each time), as (the day’s highest PEF minus the day’s lowest PEF) divided by the mean of the day’s highest and lowest PEF, and averaged over 1-2 weeks. If using PEF at home or in the office, use the same PEF meter each time.

**Physical examination** in people with asthma is often normal, but the most frequent finding is wheezing on auscultation, especially on forced expiration.

## **DIAGNOSING ASTHMA IN SPECIAL POPULATIONS**

### **Patients with cough as the only respiratory symptom**

This may be due to chronic upper airway cough syndrome ('post-nasal drip'), chronic sinusitis, gastroesophageal reflux (GERD), vocal cord dysfunction, eosinophilic bronchitis, or cough variant asthma. Cough variant asthma is characterized by cough and airway hyperresponsiveness, and documenting variability in lung function is essential to make this diagnosis. However, lack of variability at the time of testing does not exclude asthma. For other diagnostic tests, see Box 2, and Chapter 1 of the GINA 2018 report, or refer the patient for specialist opinion.

### **Occupational asthma and work-aggravated asthma**

Every patient with adult-onset asthma should be asked about occupational exposures, and whether their asthma is better when they are away from work. It is important to confirm the diagnosis objectively (which often needs specialist referral) and to eliminate exposure as soon as possible.

### **Pregnant women**

Ask all pregnant women and those planning pregnancy about asthma, and advise them about the importance of asthma controller treatment for the health of both mother and baby.

### **The elderly**

Asthma may be under-diagnosed in the elderly, due to poor perception, an assumption that dyspnea is normal in old age, lack of fitness, or reduced activity. Asthma may also be over-diagnosed in the elderly through confusion with shortness of breath due to left ventricular failure or ischemic heart disease. If there is a history of smoking or biomass fuel exposure, COPD or asthma-COPD overlap should be considered (see below).

### **Smokers and ex-smokers**

Asthma and COPD may co-exist or overlap (asthma-COPD overlap), particularly in smokers and the elderly. The history and pattern of symptoms and past records can help to distinguish asthma with fixed airflow limitation from COPD. Uncertainty in diagnosis should prompt early referral, as asthma-COPD overlap has worse outcomes than asthma or COPD alone. Asthma-COPD overlap is not a single disease, but is likely caused by several different mechanisms. There is little good-quality evidence about how to treat these patients, as they are often excluded from clinical trials.

### **Confirming an asthma diagnosis in patients taking controller treatment:**

For many patients (25–35%) with a diagnosis of asthma in primary care, the diagnosis cannot be confirmed. If the basis of the diagnosis has not already been documented, confirmation with objective testing should be sought.

If standard criteria for asthma (Box 2) are not met, consider other investigations. For example, if lung function is normal, repeat reversibility testing after withholding medications for >12 hours. If the patient has frequent symptoms, consider a trial of step-up in controller treatment and repeat lung function testing after 3 months. If the patient has few symptoms, consider stepping down controller treatment; ensure the patient has a written asthma action plan, monitor them carefully, and repeat lung function testing.

## **ASSESSING A PATIENT WITH ASTHMA**

Take every opportunity to assess patients with a diagnosis of asthma, particularly when they are symptomatic or after a recent exacerbation, but also when they ask for a prescription refill. In addition, schedule a routine review at least once a year.

### **Box 3. How to assess a patient with asthma**

<b>1. Asthma control – assess both symptom control and risk factors</b>
<ul style="list-style-type: none"><li>• Assess symptom control over the last 4 weeks (Box 4, p9)</li><li>• Identify any other risk factors for poor outcomes (Box 4)</li><li>• Measure lung function before starting treatment, 3–6 months later, and then periodically, e.g. at least yearly in most patients</li></ul>
<b>2. Treatment issues</b>
<ul style="list-style-type: none"><li>• Record the patient's treatment (Box 7, p14), and ask about side-effects</li><li>• Watch the patient using their inhaler, to check their technique (p18)</li><li>• Have an open empathic discussion about adherence (p18)</li><li>• Check that the patient has a written asthma action plan (p22)</li><li>• Ask the patient about their attitudes and goals for their asthma</li></ul>
<b>3. Are there any comorbidities?</b>
<ul style="list-style-type: none"><li>• These include rhinitis, chronic rhinosinusitis, gastroesophageal reflux (GERD), obesity, obstructive sleep apnea, depression and anxiety.</li><li>• Comorbidities should be identified as they may contribute to respiratory symptoms and poor quality of life. Their treatment may complicate asthma management.</li></ul>

# HOW TO ASSESS ASTHMA CONTROL

**Asthma control** means the extent to which the effects of asthma can be seen in the patient, or have been reduced or removed by treatment. Asthma control has two domains: symptom control and risk factors for future poor outcomes. Questionnaires like Asthma Control Test and Asthma Control Questionnaire assess only symptom control.

**Poor symptom control** is a burden to patients and a risk factor for flare-ups. **Risk factors** are factors that increase the patient’s future risk of having exacerbations (flare-ups), loss of lung function, or medication side-effects.

## Box 4. Assessment of symptom control and future risk

A. Level of asthma symptom control				
In the past 4 weeks, has the patient had:	Well controlled	Partly controlled	Uncontrolled	
Daytime symptoms more than twice/week? Yes <input type="checkbox"/> No <input type="checkbox"/>				
Any night waking due to asthma? Yes <input type="checkbox"/> No <input type="checkbox"/>	None	1–2	3–4	
Reliever needed* more than twice/week? Yes <input type="checkbox"/> No <input type="checkbox"/>	of these	of these	of these	
Any activity limitation due to asthma? Yes <input type="checkbox"/> No <input type="checkbox"/>				
B. Risk factors for poor asthma outcomes				
Assess risk factors at diagnosis and periodically, at least every 1-2 years, particularly for patients experiencing exacerbations.				
Measure FEV <sub>1</sub> at start of treatment, after 3–6 months of controller treatment to record personal best lung function, then periodically for ongoing risk assessment.				
Having uncontrolled asthma symptoms is an important risk factor for exacerbations				
Additional potentially modifiable risk factors for exacerbations, even in patients with few asthma symptoms, include:				
<ul style="list-style-type: none"> <li>• ICS not prescribed; poor ICS adherence; incorrect inhaler technique</li> <li>• High SABA use (with increased mortality if &gt;1x200-dose canister/month)</li> <li>• Low FEV<sub>1</sub>, especially if &lt;60% predicted</li> <li>• Higher bronchodilator reversibility</li> <li>• Major psychological or socioeconomic problems</li> <li>• Exposures: smoking; allergen exposure if sensitized</li> <li>• Comorbidities: obesity; chronic rhinosinusitis; confirmed food allergy</li> <li>• Sputum or blood eosinophilia; elevated FENO in allergic adults taking ICS</li> <li>• Pregnancy</li> </ul>				<p>Having any of these risk factors increases the patient's risk of exacerbations even if they have few asthma symptoms.</p>
Other major independent risk factors for flare-ups (exacerbations) include:				
<ul style="list-style-type: none"> <li>• Ever being intubated or in intensive care for asthma</li> <li>• Having 1 or more severe exacerbations in the last 12 months.</li> </ul>				
Risk factors for developing fixed airflow limitation include preterm birth, low birth weight and greater infant weight gain; lack of ICS treatment; exposure to tobacco smoke, noxious chemicals or occupational exposures; low FEV <sub>1</sub> ; chronic mucus hypersecretion; and sputum or blood eosinophilia				
Risk factors for medication side-effects include:				
<ul style="list-style-type: none"> <li>• <i>Systemic:</i> frequent OCS; long-term, high dose and/or potent ICS; also taking P450 inhibitors</li> <li>• <i>Local:</i> high-dose or potent ICS; poor inhaler technique</li> </ul>				

## What is the role of lung function in monitoring asthma?

Once asthma has been diagnosed, lung function is most useful as an indicator of future risk. It should be recorded at diagnosis, 3–6 months after starting treatment, and periodically thereafter. Most patients should have lung function measured at least every 1-2 years, more often in children and those at higher risk of flare-ups or lung function decline. Patients who have either few or many symptoms relative to their lung function need more investigation.

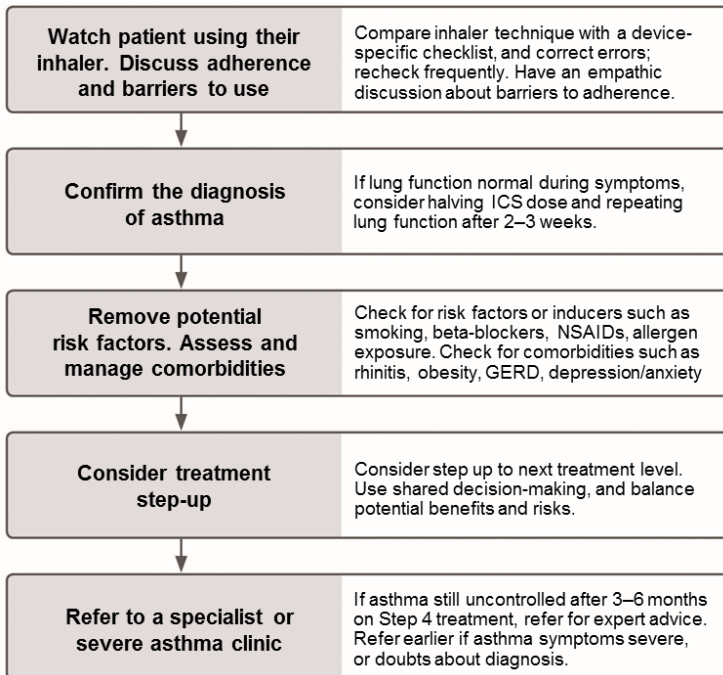
## How is asthma severity assessed?

Asthma severity can be assessed retrospectively from the level of treatment (p14) required to control symptoms and exacerbations. Mild asthma is asthma that can be controlled with Step 1 or 2 treatment. Severe asthma is asthma that requires Step 4 or 5 treatment, to maintain symptom control. It may appear similar to asthma that is uncontrolled due to lack of treatment.

## HOW TO INVESTIGATE UNCONTROLLED ASTHMA

Most patients can achieve good asthma control with regular controller treatment, but some patients do not, and further investigation is needed.

### Box 5. How to investigate uncontrolled asthma in primary care



This flow-chart shows the most common problems first, but the steps can be carried out in a different order, depending on resources and clinical context.

# MANAGEMENT OF ASTHMA

## GENERAL PRINCIPLES

The long-term goals of asthma management are **symptom control** and **risk reduction**. The aim is to reduce the burden to the patient and to reduce their risk of exacerbations, airway damage, and medication side-effects. The patient's own goals regarding their asthma and its treatment should also be identified.

**Population-level recommendations** about 'preferred' asthma treatments represent the best treatment for most patients in a population.

**Patient-level treatment decisions** should take into account any individual characteristics, risk factors, comorbidities or phenotype that predict the patient's likely response to treatment, together with the patient's own preferences, and practical issues such as inhaler technique, adherence, and affordability.

A **partnership** between the patient and their health care providers is important for effective asthma management. Training health care providers in **communication skills** may lead to increased patient satisfaction, better health outcomes, and reduced use of health care resources.

**Health literacy** – that is, the patient's ability to obtain, process and understand basic health information to make appropriate health decisions – should be taken into account in asthma management and education.

## TREATING TO CONTROL SYMPTOMS AND MINIMIZE RISK

Treatment of asthma for symptom control and risk reduction includes:

- Medications. Every patient with asthma should have a reliever medication, and most adults and adolescents with asthma should have a controller medication to reduce the risk of serious exacerbations, even in patients with infrequent symptoms
- Treating modifiable risk factors and comorbidities
- Non-pharmacological therapies and strategies

Importantly, every patient should also be trained in essential skills and guided asthma self-management, including:

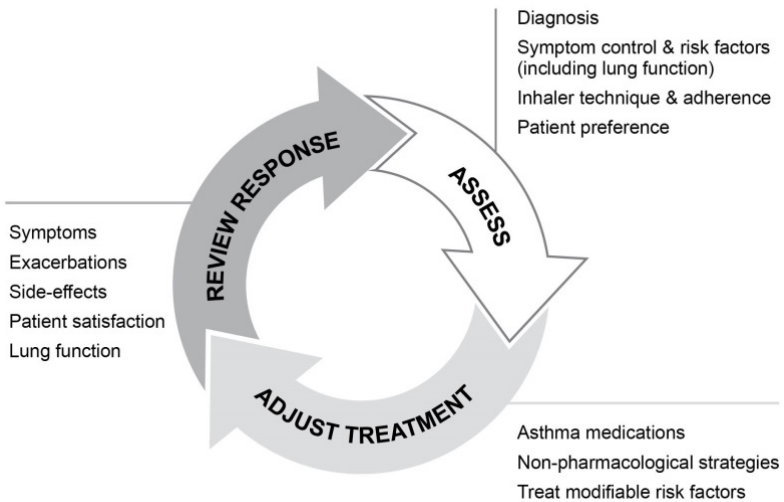
- Asthma information
- Inhaler skills (p18)
- Adherence (p18)
- Written asthma action plan (p22)
- Self-monitoring
- Regular medical review (p8)

## CONTROL-BASED ASTHMA MANAGEMENT

Asthma treatment is adjusted in a continuous cycle to **assess, adjust treatment** and **review response**. The main components of this cycle are shown in Box 6.

The assessment of control includes not only symptom control, but also the patient's individual **risk factors and comorbidities** that can contribute to their burden of disease and risk of poor health outcomes, or that may predict their response to treatment.

### Box 6. The control-based asthma management cycle



## INITIAL CONTROLLER TREATMENT

For the best outcomes, regular daily controller treatment should be initiated as soon as possible after the diagnosis of asthma is made, because:

- Early treatment with low dose ICS leads to better lung function than if symptoms have been present for more than 2–4 years
- Patients not taking ICS who experience a severe exacerbation have lower long-term lung function than those who have started ICS
- In occupational asthma, early removal from exposure and early treatment increase the probability of recovery

**Regular low dose ICS is recommended** for all patients with a diagnosis of asthma and any of the following:

- Asthma symptoms more than twice a month
- Waking due to asthma more than once a month
- Any asthma symptoms plus any risk factor(s) for exacerbations (e.g. needing OCS for asthma within the last 12 months; low FEV<sub>1</sub>; ever in intensive care unit for asthma)

Consider starting at a higher step (e.g. medium/high dose ICS, or ICS/LABA) if the patient has troublesome asthma symptoms on most days; or is waking from asthma once or more a week, especially if there are any risk factors for exacerbations.

If the initial asthma presentation is with severely uncontrolled asthma, or with an acute exacerbation, give a short course of OCS and start regular controller treatment (e.g. high dose ICS, or medium dose ICS/LABA).

Consider stepping down after asthma has been well-controlled for 3 months.

Low, medium and high dose categories for different ICS medications are shown in Box 8 (p14).

### **Before starting initial controller treatment**

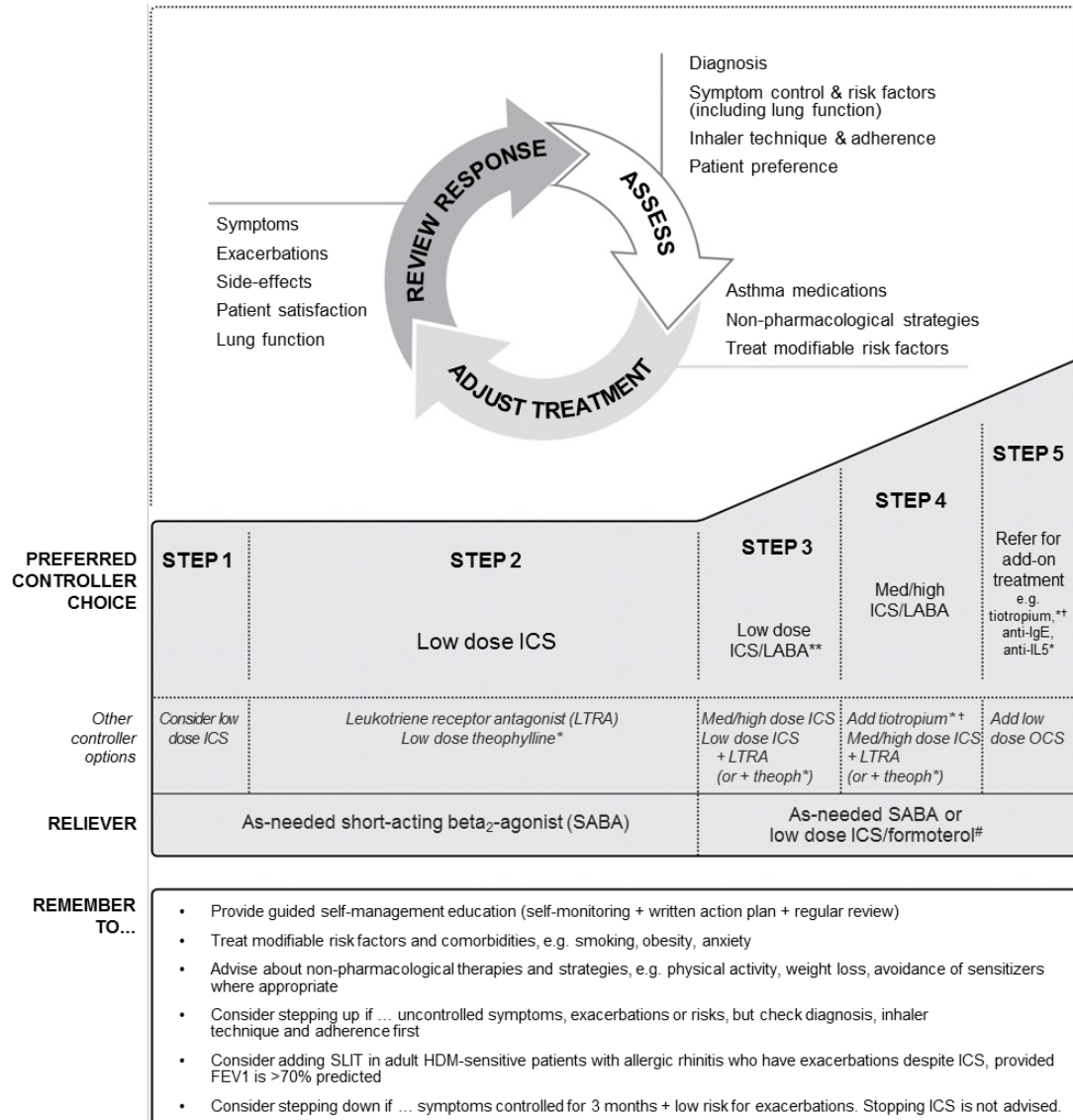
- Record evidence for the diagnosis of asthma, if possible
- Document symptom control and risk factors
- Assess lung function, when possible
- Train the patient to use the inhaler correctly, and check their technique
- Schedule a follow-up visit

### **After starting initial controller treatment**

- Review response after 2–3 months, or according to clinical urgency
- See Box 7 for ongoing treatment and other key management issues
- Consider step down when asthma has been well-controlled for 3 months



## Box 7. Stepwise approach to asthma treatment



\*Not for children <12 years. \*\*For children 6–11 years, the preferred Step 3 treatment is medium dose ICS. # Low dose ICS/formoterol is the reliever medication for patients prescribed low dose budesonide/formoterol or low dose beclometasone/formoterol for maintenance and reliever therapy. †Tiotropium by mist inhaler is an add-on treatment for patients with a history of exacerbations\*.

For medication Glossary, see p25. For details about treatment recommendations, supporting evidence, and clinical advice about implementation in different populations see the full GINA 2018 report ([www.ginasthma.org](http://www.ginasthma.org)).

## Box 8. Low, medium and high daily doses of inhaled corticosteroids (mcg)

Inhaled corticosteroid	Adults and adolescents			Children 6–11 years		
	Low	Medium	High	Low	Medium	High
Beclometasone dipropionate (CFC)*	200–500	>500–1000	>1000	100–200	>200–400	>400
Beclometasone dipropionate (HFA)	100–200	>200–400	>400	50–100	>100–200	>200
Budesonide (DPI)	200–400	>400–800	>800	100–200	>200–400	>400
Budesonide (nebulizer)				250–500	>500–1000	>1000
Ciclesonide (HFA)	80–160	>160–320	>320	80	>80–160	>160
Fluticasone furoate (DPI)	100	n.a.	200	n.a.	n.a.	n.a.
Fluticasone propionate (DPI)	100–250	>250–500	>500	100–200	>200–400	>400
Fluticasone propionate (HFA)	100–250	>250–500	>500	100–200	>200–500	>500
Mometasone furoate	110–220	>220–440	>440	110	≥220–<440	≥440
Triamcinolone acetonide	400–1000	>1000–2000	>2000	400–800	>800–1200	>1200

† CFC: chlorofluorocarbon propellant; DPI: dry powder inhaler; HFA: hydrofluoroalkane propellant. \*Included for comparison with older literature.

## STEPWISE APPROACH FOR ADJUSTING TREATMENT

Once asthma treatment has been started, ongoing decisions are based on a cycle (p.12) to assess, adjust treatment and review response. The preferred treatments at each step are summarized below and in Box 7 (p14); for details, see full GINA 2018 report. See Box 8 (p14) for ICS dose categories.

**STEP 1: As-needed SABA with no controller** This is indicated **only** if symptoms are rare, there is no night waking due to asthma, no exacerbations in the last year, and normal FEV<sub>1</sub>.

*Other options:* regular low dose ICS to reduce risk of severe exacerbations.

**STEP 2: Regular low dose ICS plus as-needed SABA**

*Other options:* LTRA are less effective than ICS; ICS/LABA leads to faster improvement in symptoms and FEV<sub>1</sub> than ICS alone but is more expensive and the exacerbation rate is similar. For purely seasonal allergic asthma, start ICS immediately and cease 4 weeks after end of exposure.

**STEP 3: Low dose ICS/LABA either as maintenance treatment plus as-needed SABA, or as ICS/formoterol maintenance and reliever therapy**

For patients with  $\geq 1$  exacerbation in the last year, low dose BDP/formoterol or BUD/formoterol maintenance and reliever strategy is more effective than maintenance ICS/LABA with as-needed SABA.

*Other options:* Medium dose ICS; for adult patients with rhinitis and allergic to house dust mite (HDM) with exacerbations despite ICS, consider adding sublingual immunotherapy (SLIT), provided FEV<sub>1</sub> is  $>70\%$  predicted.

*Children (6-11 years):* Medium dose ICS. *Other options:* low dose ICS/LABA

**STEP 4: Low dose ICS/formoterol maintenance and reliever therapy, or medium dose ICS/LABA as maintenance plus as-needed SABA**

*Other options:* Add-on tiotropium by mist inhaler for patients  $\geq 12$  years with a history of exacerbations; high dose ICS/LABA, but more side-effects and little extra benefit; extra controller, e.g. add-on LTRA or slow-release theophylline (adults); for adult patients with rhinitis and allergic to HDM with exacerbations despite ICS, consider adding SLIT, provided FEV<sub>1</sub> is  $>70\%$  predicted.

*Children (6-11 years):* Refer for expert assessment and advice.

**STEP 5: Refer for expert investigation and add-on treatment**

Add-on treatments include tiotropium by mist inhaler for patients with a history of exacerbations (age  $\geq 12$  years), anti-IgE (SC omalizumab) for severe allergic asthma  $\geq 6$  years, and anti-IL5 for severe eosinophilic asthma (SC mepolizumab or SC benralizumab for age  $\geq 12$  years or IV reslizumab for age  $\geq 18$  years). Sputum-guided treatment, if available, improves outcomes.

*Other options:* Some patients may benefit from low dose OCS but long-term systemic side-effects commonly occur.

## REVIEWING RESPONSE AND ADJUSTING TREATMENT

### How often should patients with asthma be reviewed?

Patients should preferably be seen 1–3 months after starting treatment and every 3–12 months after that, except in pregnancy when they should be reviewed every 4–6 weeks. After an exacerbation, a review visit within 1 week should be scheduled. The frequency of review depends on the patient's initial level of control, their response to previous treatment, and their ability and willingness to engage in self-management with an action plan.

### Stepping up asthma treatment

Asthma is a variable condition, and periodic adjustment of controller treatment by the clinician and/or patient may be needed.

- *Sustained step-up (for at least 2–3 months)*: if symptoms and/or exacerbations persist despite 2–3 months of controller treatment, assess the following common issues before considering a step-up
  - Incorrect inhaler technique
  - Poor adherence
  - Modifiable risk factors, e.g. smoking
  - Are symptoms due to comorbid conditions, e.g. allergic rhinitis
- *Short-term step-up (for 1–2 weeks)* by clinician or by patient with written asthma action plan (p22), e.g. during viral infection or allergen exposure
- *Day-to-day adjustment by patient* for patients prescribed low dose beclometasone/formoterol or budesonide/formoterol as maintenance and reliever therapy.

### Stepping down treatment when asthma is well-controlled

Consider stepping down treatment once good asthma control has been achieved and maintained for 3 months, to find the lowest treatment that controls both symptoms and exacerbations, and minimizes side-effects.

- Choose an appropriate time for step-down (no respiratory infection, patient not travelling, not pregnant)
- Document baseline status (symptom control and lung function), provide a written asthma action plan, monitor closely, and book a follow-up visit
- Step down through available formulations to reduce the ICS dose by 25–50% at 2–3 month intervals (see Box 3-9 in full GINA 2018 report for details of how to step down different controller treatments)
- Do not completely stop ICS in adults or adolescents with a diagnosis of asthma unless this is needed temporarily to confirm the diagnosis of asthma. Make sure a follow-up appointment is arranged.

## INHALER SKILLS AND ADHERENCE

### Provide skills training for effective use of inhaler devices

Most patients (up to 80%) cannot use their inhaler correctly. This contributes to poor symptom control and exacerbations. To ensure effective inhaler use:

- **Choose** the most appropriate device for the patient before prescribing: consider medication, physical problems e.g. arthritis, patient skills, and cost; for ICS by pressurized metered dose inhaler, prescribe a spacer.
- **Check** inhaler technique at every opportunity. Ask the patient to show you how they use the inhaler. Check their technique against a device-specific checklist.
- **Correct** using a physical demonstration, paying attention to incorrect steps. Check technique again, up to 2–3 times if necessary.
- **Confirm** that you have checklists for each of the inhalers you prescribe, and can demonstrate correct technique on them.

Information about inhaler devices and techniques for their use can be found on the GINA website ([www.ginasthma.org](http://www.ginasthma.org)) and the ADMIT website ([www.admit-inhalers.org](http://www.admit-inhalers.org)).

### Check and improve adherence with asthma medications

At least 50% of adults and children do not take controller medications as prescribed. Poor adherence contributes to poor symptom control and exacerbations. It may be unintentional (e.g. forgetfulness, cost, misunderstandings) and/or non-intentional (e.g. not perceiving the need for treatment, fear of side-effects, cultural issues, cost).

To identify patients with adherence problems:

- Ask an empathic question, e.g. “Most patients don’t take their inhaler exactly as prescribed. In the last 4 weeks, how many days a week have you been taking it? 0 days a week, or 1, or 2 days [etc]?”, or “Do you find it easier to remember your inhaler in the morning or night?”
- Check medication usage, from prescription date, inhaler date/dose counter, dispensing records
- Ask about attitudes and beliefs about asthma and medications

Only a few adherence interventions have been studied closely in asthma and have improved adherence in real-world studies.

- Shared decision-making for medication and dose choice
- Inhaler reminders for missed doses
- Comprehensive asthma education with home visits by asthma nurses
- Clinicians reviewing feedback about their patients’ dispensing records
- An automated voice recognition program with telephone messages triggered when refills were due or overdue

## TREATING MODIFIABLE RISK FACTORS

Exacerbation risk can be minimized by optimizing asthma medications, and by identifying and treating modifiable risk factors. Some examples of risk modifiers with consistent high quality evidence are:

- **Guided self-management:** self-monitoring of symptoms and/or PEF, a written asthma action plan (p22), and regular medical review
- **Use of a regimen that minimizes exacerbations:** prescribe an ICS-containing controller. For patients with 1 or more exacerbations in the last year, consider a low dose ICS/formoterol maintenance and reliever regimen
- **Avoidance of exposure to tobacco smoke**
- **Confirmed food allergy:** appropriate food avoidance; ensure availability of injectable epinephrine for anaphylaxis
- **For patients with severe asthma:** refer to a specialist center, if available, for detailed assessment and consideration of add-on biologic medications and/or sputum-guided treatment.

## NON-PHARMACOLOGICAL STRATEGIES AND INTERVENTIONS

In addition to medications, other therapies and strategies may be considered where relevant, to assist in symptom control and risk reduction. Some examples with consistent high quality evidence are:

- **Smoking cessation advice:** at every visit, strongly encourage smokers to quit. Provide access to counselling and resources. Advise parents and carers to exclude smoking in rooms/cars used by children with asthma
- **Physical activity:** encourage people with asthma to engage in regular physical activity because of its general health benefits. Provide advice about management of exercise-induced bronchoconstriction.
- **Occupational asthma:** ask all patients with adult-onset asthma about their work history. Identify and remove occupational sensitizers as soon as possible. Refer patients for expert advice, if available.
- **NSAIDs including aspirin:** always ask about asthma before prescribing.

Although allergens may contribute to asthma symptoms in sensitized patients, allergen avoidance is not recommended as a general strategy for asthma.

These strategies are often complex and expensive, and there are no validated methods for identifying those who are likely to benefit.

Some common triggers for asthma symptoms (e.g. exercise, laughter) should **not** be avoided, and others (e.g. viral respiratory infections, stress) are difficult to avoid and should be managed when they occur.

## TREATMENT IN SPECIAL POPULATIONS OR CONTEXTS

**Pregnancy:** asthma control often changes during pregnancy. For baby and mother, the advantages of actively treating asthma markedly outweigh any potential risks of usual controller and reliever medications. Down-titration has a low priority in pregnancy. Exacerbations should be treated aggressively.

**Rhinitis and sinusitis** often coexist with asthma. Chronic rhinosinusitis is associated with more severe asthma. Treatment of allergic rhinitis or chronic rhinosinusitis reduces nasal symptoms but does not improve asthma control.

**Obesity:** to avoid over- or under-treatment, it is important to document the diagnosis of asthma in the obese. Asthma is more difficult to control in obesity. Weight reduction should be included in the treatment plan for obese patients with asthma; even 5–10% weight loss can improve asthma control.

**The elderly:** comorbidities and their treatment should be considered and may complicate asthma management. Factors such as arthritis, eyesight, inspiratory flow, and complexity of treatment regimens should be considered when choosing medications and inhaler devices.

**Gastroesophageal reflux (GERD)** is commonly seen in asthma. Symptomatic reflux should be treated for its general health benefits, but there is no benefit from treating asymptomatic reflux in asthma.

**Anxiety and depression:** these are commonly seen in people with asthma, and are associated with worse symptoms and quality of life. Patients should be assisted to distinguish between symptoms of anxiety and of asthma.

**Aspirin-exacerbated respiratory disease (AERD):** a history of exacerbation following ingestion of aspirin or other NSAIDs is highly suggestive. Patients often have severe asthma and nasal polyposis. Confirmation of the diagnosis of AERD may require challenge in a specialized center with cardiopulmonary resuscitation facilities, but avoidance of NSAIDs may be recommended on the basis of a clear history. ICS are the mainstay of treatment, but OCS may be required; LTRA may also be useful. Desensitization under specialist care is sometimes effective.

**Food allergy and anaphylaxis:** food allergy is rarely a trigger for asthma symptoms. It must be assessed with specialist testing. Confirmed food allergy is a risk factor for asthma-related death. Good asthma control is essential; patients should also have an anaphylaxis plan and be trained in appropriate avoidance strategies and use of injectable epinephrine.

**Surgery:** whenever possible, good asthma control should be achieved pre-operatively. Ensure that controller therapy is maintained throughout the peri-operative period. Patients on long-term high dose ICS, or having more than 2 weeks' OCS in the past 6 months, should receive intra-operative hydrocortisone to reduce the risk of adrenal crisis.

## **ASTHMA FLARE-UPS (EXACERBATIONS)**

A flare-up or exacerbation is an acute or sub-acute worsening in symptoms and lung function from the patient's usual status; occasionally it may be the initial presentation of asthma.

For discussion with patients, the word 'flare-up' is preferred. 'Episodes', 'attacks' and 'acute severe asthma' are often used, but they have variable meanings, particularly for patients.

The management of worsening asthma and exacerbations should be considered as a continuum, from self-management by the patient with a written asthma action plan, through to management of more severe symptoms in primary care, the emergency department and in hospital.

### **Identifying patients at risk of asthma-related death**

These patients should be identified, and flagged for more frequent review.

- A history of near-fatal asthma requiring intubation and ventilation
- Hospitalization or emergency care for asthma in last 12 months
- Not currently using ICS, or poor adherence with ICS
- Currently using or recently stopped using OCS (this indicates the severity of recent events)
- Over-use of SABAs, especially more than 1 canister/month
- Lack of a written asthma action plan
- History of psychiatric disease or psychosocial problems
- Confirmed food allergy in a patient with asthma





## MANAGING EXACERBATIONS IN PRIMARY OR ACUTE CARE

**Assess** exacerbation severity while starting SABA and oxygen. Assess dyspnea (e.g. is the patient able to speak sentences, or only words), respiratory rate, pulse rate, oxygen saturation and lung function (e.g. PEF). Check for anaphylaxis.

**Consider alternative causes** of acute breathlessness (e.g. heart failure, upper airway dysfunction, inhaled foreign body or pulmonary embolism).

**Arrange immediate transfer** to an acute care facility if there are signs of severe exacerbation, or to intensive care if the patient is drowsy, confused, or has a silent chest. For these patients, immediately give inhaled SABA, inhaled ipratropium bromide, oxygen and systemic corticosteroids.

**Start treatment** with repeated doses of SABA (usually by pMDI and spacer), early oral corticosteroids, and controlled flow oxygen if available. Check response of symptoms and saturation frequently, and measure lung function after 1 hour. Titrate oxygen to maintain saturation of 93–95% in adults and adolescents (94–98% in children 6–12 years).

**For severe exacerbations**, add ipratropium bromide, and consider giving SABA by nebulizer. In acute care facilities, intravenous magnesium sulfate may be considered if the patient is not responding to intensive initial treatment.

Do not routinely perform chest X-ray or blood gases, or prescribe antibiotics, for asthma exacerbations.

## REVIEWING RESPONSE

**Monitor patients closely and frequently** during treatment, and titrate treatment according to response. Transfer the patient to higher level care if worsening or failing to respond.

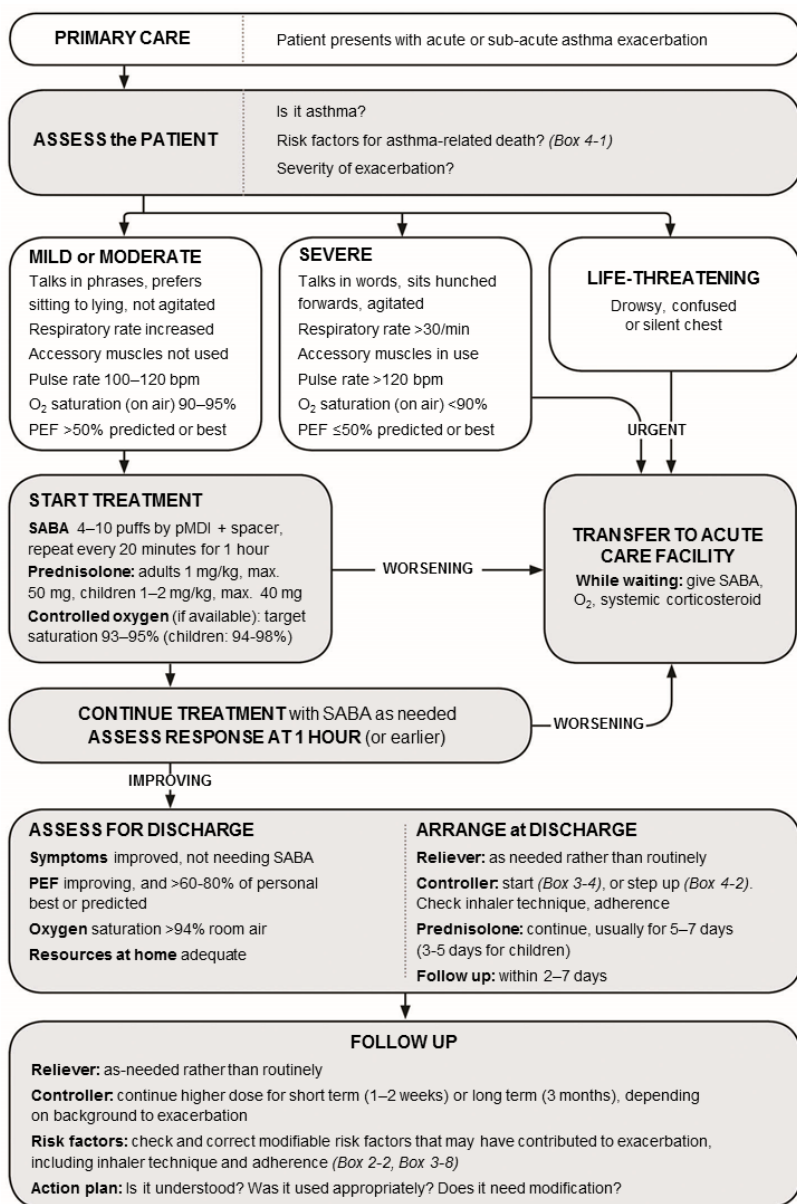
**Decide about need for hospitalization** based on clinical status, symptomatic and lung function, response to treatment, recent and past history of exacerbations, and ability to manage at home.

**Before discharge, arrange ongoing treatment.** For most patients, prescribe regular controller therapy (or increase current dose) to reduce the risk of further exacerbations. Continue increased controller doses for 2–4 weeks, and reduce reliever to as-needed dosing. Check inhaler technique and adherence. Provide an interim written asthma action plan.

**Arrange early follow-up** after any exacerbation, within 2–7 days.

Consider referral for specialist advice for patients with an asthma hospitalization, or repeated emergency department presentations.

## Box 10. Management of asthma exacerbations in primary care



O<sub>2</sub>: oxygen; PEF: peak expiratory flow; SABA: short-acting beta<sub>2</sub>-agonist (doses are for salbutamol)

## FOLLOW-UP AFTER AN EXACERBATION

Exacerbations often represent failures in chronic asthma care, and they provide opportunities to review the patient's asthma management. **All patients must be followed up regularly by a health care provider until symptoms and lung function return to normal.**

Take the opportunity to review:

- The patient's understanding of the cause of the exacerbation
- Modifiable risk factors for exacerbations, e.g. smoking
- Understanding of purposes of medications, and inhaler technique skills
- Review and revise written asthma action plan

Discuss medication use, as adherence with ICS and OCS may fall to 50% within a week after discharge.

Comprehensive post-discharge programs that include optimal controller management, inhaler technique, self-monitoring, written asthma action plan and regular review are cost-effective and are associated with significant improvement in asthma outcomes.

Referral for expert advice should be considered for patients who have been hospitalized for asthma, or who re-present for acute asthma care.

## GLOSSARY OF ASTHMA MEDICATION CLASSES

For more details, see full GINA 2018 report and Appendix ([www.ginasthma.org](http://www.ginasthma.org)) and Product Information from manufacturers.

Medications	Action and use	Adverse effects
<b>CONTROLLER MEDICATIONS</b>		
<b>Inhaled corticosteroids (ICS)</b> (pMDIs or DPIs) e.g. beclometasone, budesonide, ciclesonide, fluticasone propionate, fluticasone furoate, mometasone, triamcinolone	The most effective anti-inflammatory medications for persistent asthma. ICS reduce symptoms, increase lung function, improve quality of life, and reduce the risk of exacerbations and asthma-related hospitalizations or death. ICS differ in their potency and bioavailability, but most of the benefit is seen at low doses (see Box 8 (p14) for low, medium and high doses of different ICS).	Most patients using ICS do not experience side-effects. Local side-effects include oropharyngeal candidiasis and dysphonia. Use of spacer with pMDI, and rinsing with water and spitting out after inhalation, reduce local side effects. High doses increase the risk of systemic side-effects.
<b>ICS and long-acting beta<sub>2</sub> agonist bronchodilator combinations (ICS/LABA)</b> (pMDIs or DPIs) e.g. beclometasone/formoterol, budesonide/formoterol, fluticasone furoate/	When a medium dose of ICS alone fails to achieve good control of asthma, the addition of LABA to ICS improves symptoms, lung function and reduces exacerbations in more patients, more rapidly, than doubling the dose of ICS. Two regimens are available: maintenance ICS/LABA with SABA as	The LABA component may be associated with tachycardia, headache or cramps. Current recommendations are that LABA and ICS are safe for asthma when used in

Medications	Action and use	Adverse effects
vilanterol, fluticasone propionate/formoterol, fluticasone propionate/salmeterol, and mometasone/formoterol.	reliever, and low-dose combination beclometasone or budesonide with formoterol for maintenance and reliever treatment.	combination. LABA should not be used without ICS in asthma due to increased risk of serious adverse outcomes.
<b>Leukotriene modifiers</b> (tablets) e.g. montelukast, pranlukast, zafirlukast, zileuton	Target one part of the inflammatory pathway in asthma. Used as an option for controller therapy, particularly in children. Used alone: less effective than low dose ICS; added to ICS: less effective than ICS/LABA.	Few side-effects in placebo-controlled studies except elevated liver function tests with zileuton and zafirlukast.
<b>Chromones</b> (pMDIs or DPIs) e.g. sodium cromoglycate and nedocromil sodium	Very limited role in long-term treatment of asthma. Weak anti-inflammatory effect, less effective than low-dose ICS. Require meticulous inhaler maintenance.	Side effects are uncommon but include cough upon inhalation and pharyngeal discomfort.
<b>ADD-ON CONTROLLER MEDICATIONS</b>		
<b>Long-acting anticholinergic</b> (tiotropium)	An add-on option at Step 4 or 5 by mist inhaler for patients $\geq 12$ years with a history of exacerbations despite ICS $\pm$ LABA	Side-effects are uncommon but include dry mouth.
<b>Anti-IgE</b> (SC omalizumab)	An add-on option for patients $\geq 6$ years with severe allergic asthma uncontrolled on high dose ICS/LABA.	Reactions at the site of injection are common but minor. Anaphylaxis is rare.
<b>Anti-IL5</b> (SC mepolizumab or benralizumab [ $\geq 12$ y]; IV reslizumab [ $\geq 18$ y])	Add-on options for patients with severe eosinophilic asthma uncontrolled on high dose ICS/LABA	Headache, and reactions at injection site are common but minor.
<b>Systemic corticosteroids</b> (tablets, suspension or intramuscular (IM) or intravenous (IV) injection) e.g. prednisone, prednisolone, methylprednisolone, hydrocortisone	Short-term treatment (usually 5–7 days in adults) is important in the treatment of severe acute exacerbations, with main effects seen after 4–6 hours. Oral corticosteroid (OCS) therapy is preferred and is as effective as IM or IV therapy in preventing relapse. Tapering is required if treatment given for more than 2 weeks. Long-term treatment with OCS may be required for some patients with severe asthma, but side-effects must be taken into account.	Short-term use: some adverse effects e.g. sleep disturbance, reflux, appetite increase, hyperglycaemia, mood changes. Long-term use: limited by the risk of significant systemic adverse effects e.g. cataract, glaucoma, osteoporosis, adrenal suppression. Patients should be assessed for osteoporosis risk and treated appropriately.
<b>RELIEVER MEDICATIONS</b>		
<b>Short-acting inhaled beta<sub>2</sub>-agonist bronchodilators (SABA)</b> (pMDIs, DPIs and, rarely, solution for nebulization or injection) e.g. salbutamol	Inhaled SABAs are medications of choice for quick relief of asthma symptoms and bronchoconstriction including in acute exacerbations, and for pre-treatment of exercise-induced bronchoconstriction. SABAs should be used only as-needed at	Tremor and tachycardia are commonly reported with initial use of SABA, but tolerance to these effects usually develops rapidly. Excess use, or poor

Medications	Action and use	Adverse effects
(albuterol), terbutaline.	the lowest dose and frequency required.	response indicate poor asthma control.
<b>Low-dose ICS/formoterol</b> (beclometasone/formoterol or budesonide/formoterol)	This is the reliever medication for patients prescribed maintenance and reliever treatment. It reduces the risk of exacerbations compared with using prn SABA, with similar symptom control.	As for ICS/LABA above
<b>Short-acting anticholinergics</b> (pMDIs or DPIs) e.g. ipratropium bromide, oxitropium bromide	Long-term use: ipratropium is a less effective reliever medication than SABAs. Short-term use in acute asthma: inhaled ipratropium added to SABA reduces the risk of hospital admission	Dryness of the mouth or a bitter taste.

### Abbreviations used in this pocket guide

BDP	Beclometasone dipropionate	LAMA	Long-acting muscarinic antagonist
BUD	Budesonide	n.a.	Not applicable
DPI	Dry powder inhaler	O <sub>2</sub>	Oxygen
FEV <sub>1</sub>	Forced expiratory volume in 1 second	OCS	Oral corticosteroids
FVC	Forced vital capacity	PEF	Peak expiratory flow
HDM	House dust mite	pMDI	Pressurized metered dose inhaler
ICS	Inhaled corticosteroids	SABA	Short-acting beta <sub>2</sub> -agonists
IV	Intravenous	SC	Subcutaneous
LABA	Long-acting beta <sub>2</sub> -agonists	SLIT	Sublingual immunotherapy

## ACKNOWLEDGEMENTS

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### **GINA Assembly**

The GINA Assembly includes members from 45 countries. Their names are listed on the GINA website, [www.ginasthma.org](http://www.ginasthma.org).

## GINA PUBLICATIONS

- **Global Strategy for Asthma Management and Prevention** (updated 2018). This report provides an integrated approach to asthma that can be adapted for a wide range of health systems. The report was extensively revised in 2014, and has been updated yearly since then. The report has a user-friendly format with many practical summary tables and flow-charts for use in clinical practice.
- **GINA Online Appendix** (updated 2018). Detailed information to support the main GINA report.
- **Pocket Guide for asthma management and prevention for adults and children older than 5 years** (updated 2018). Summary for primary health care providers, to be used in conjunction with the main GINA report.
- **Pocket guide for asthma management and prevention in children 5 years and younger** (updated 2018). A summary of patient care information about pre-schoolers with asthma or wheeze, to be used in conjunction with the main GINA 2018 report.
- **Diagnosis of asthma-COPD overlap** (updated 2018). This is a stand-alone copy of the corresponding chapter in the main GINA report. It is co-published by GINA and GOLD (the Global Initiative for Chronic Obstructive Lung Disease, [www.goldcopd.org](http://www.goldcopd.org)).
- **Clinical practice aids and implementation tools** will be available on the GINA website.

**GINA publications and other resources are available from [www.ginasthma.org](http://www.ginasthma.org)**

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