

Guideline-directed Asthma Care

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Objectives

- Review asthma prevalence, morbidity and mortality trends
- Describe the four components of the Expert Panel Report-3 (EPR-3)
 - Discuss current standards for determining the diagnosis of asthma, as well as severity and control classifications
 - Highlight asthma triggers (allergens vs. irritants)
 - Compare and contrast the pharmacologic agents recommended by age and severity/control indices
 - Examine the role of patient-provider partnerships in asthma self-management
- Assess the evidence for new therapeutic agents/interventions for asthma from 2017 Global Initiative for Asthma (GINA) and 2018 EPR-4 guidelines
- Identify new risks for asthma related to climate change
- Appraise the current state of asthma adherence

Asthma
prevalence,
morbidity and
mortality
trends

Asthma prevalence: globally

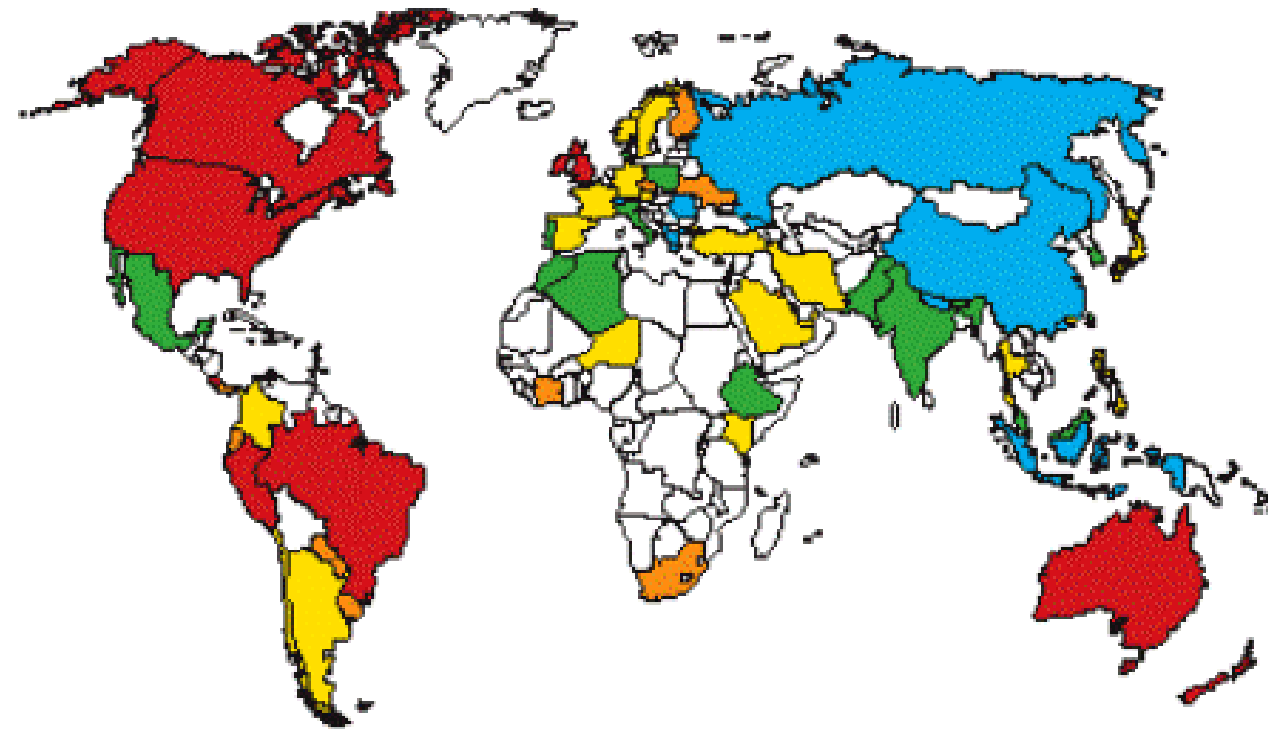


Figure 2. Asthma prevalence, by selected demographic characteristics: United States, average annual 2008–2010

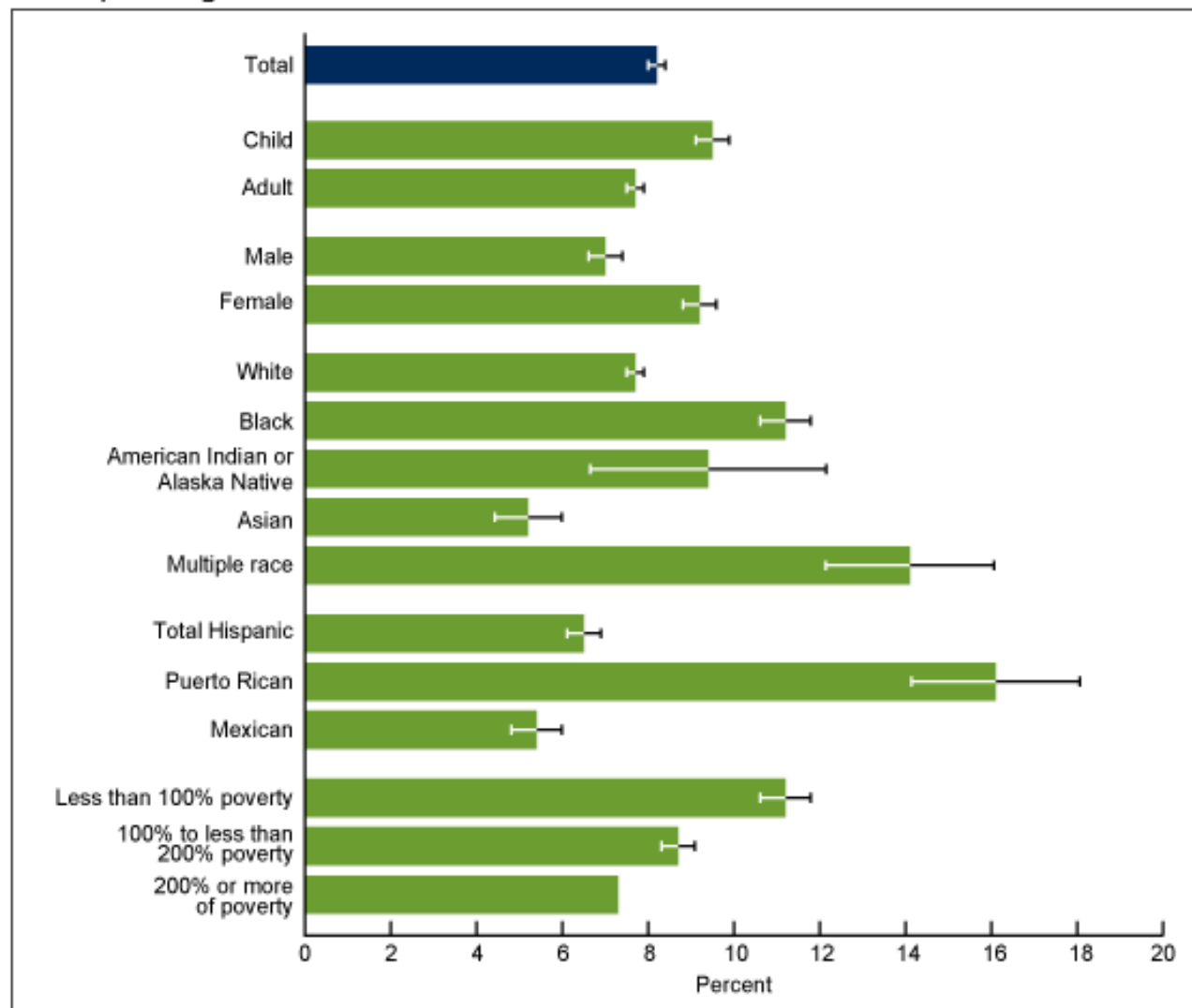


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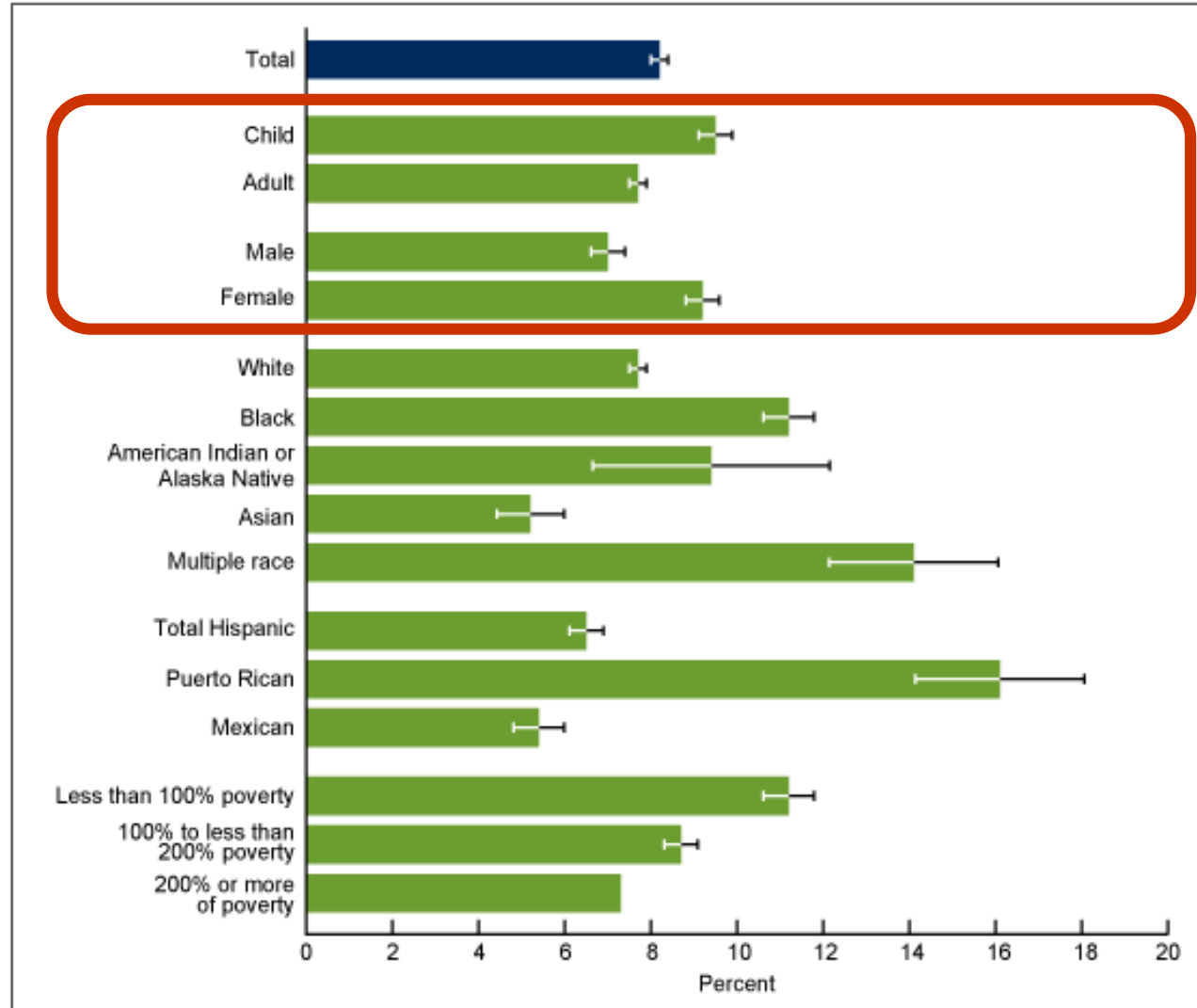


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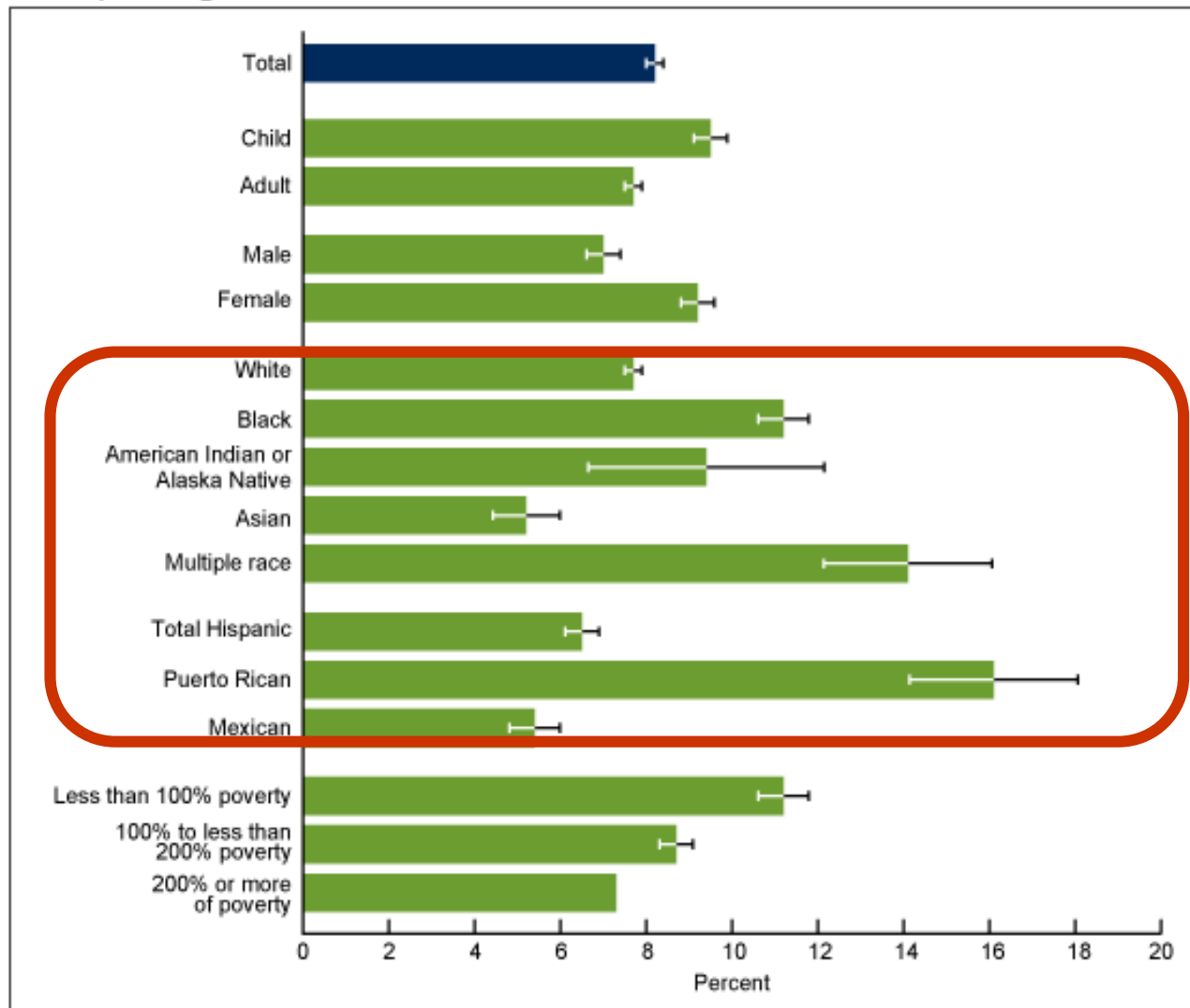
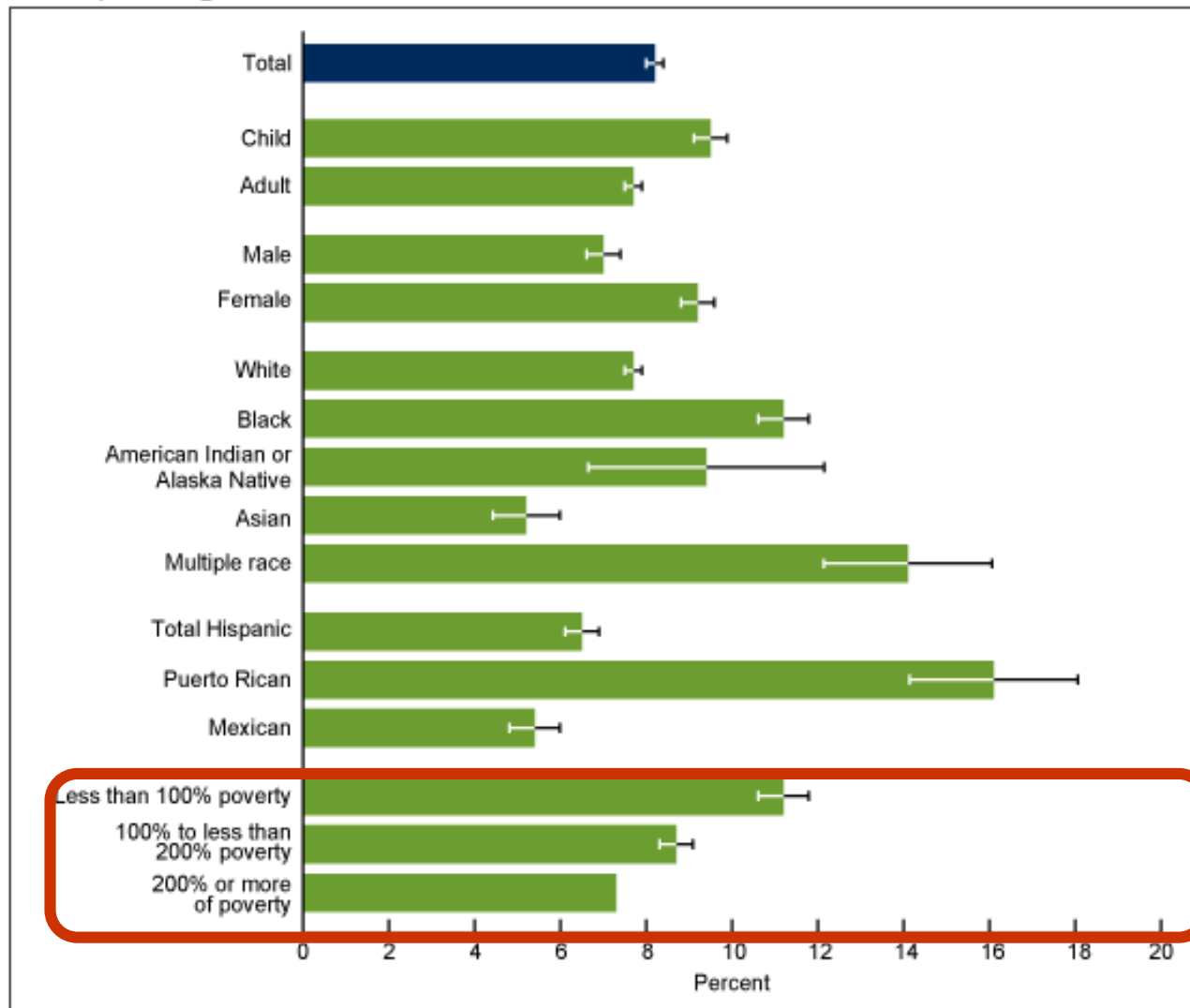


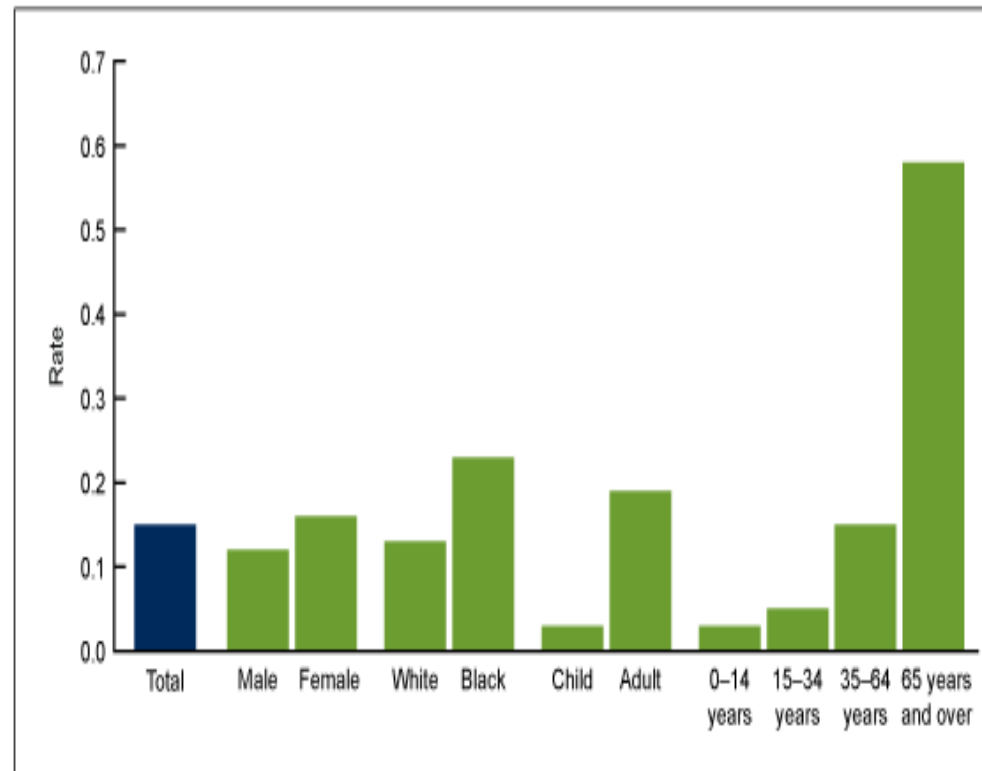
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NCHS Data Brief Number 94, May 2012

Trends in Asthma Mortality in the United States, 2001–2010

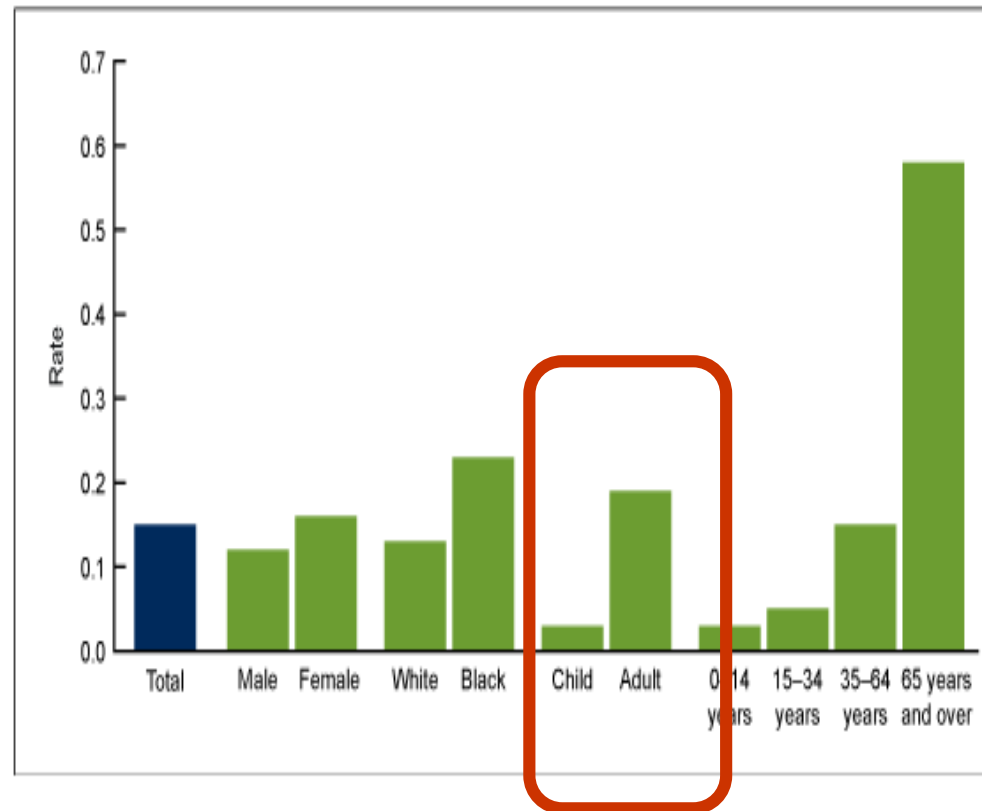
Figure 5. Asthma deaths per 1,000 persons with asthma, by selected demographic characteristics: United States, average annual 2007–2009



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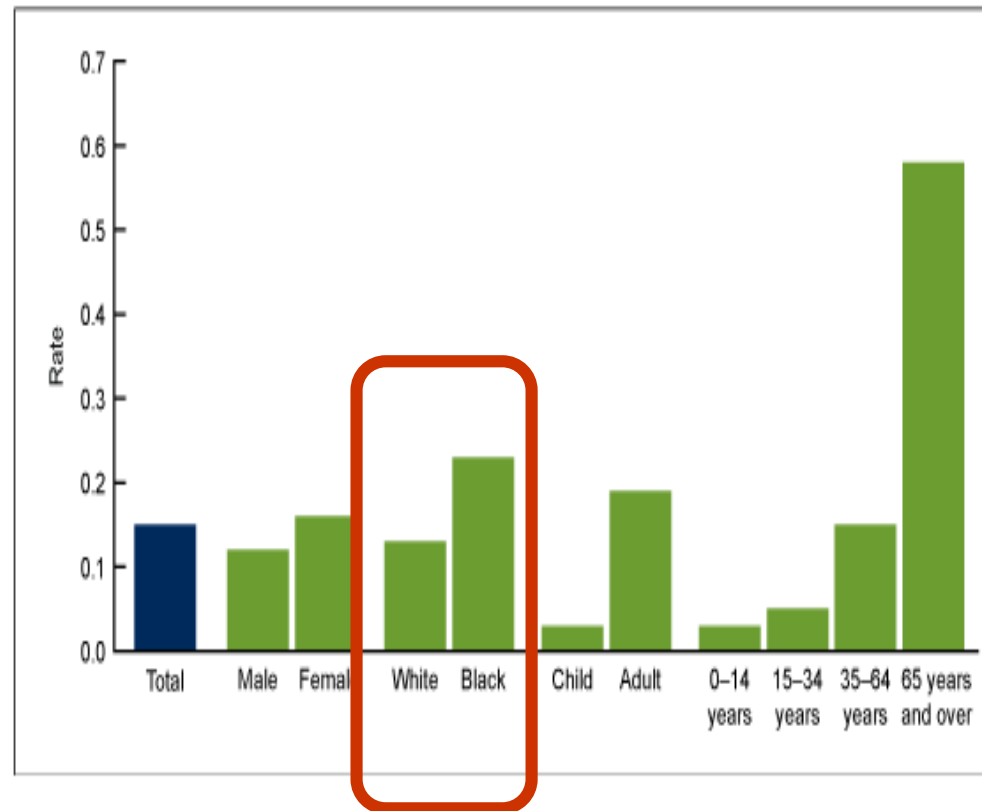
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Describe the
four
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the Expert
Panel Report-3
(EPR-3)

**Component 1- Measures of Asthma
Assessment and Monitoring**

Component 2-Environmental factors

Component 3-Treatment

Component 4-Education for a partnership



Component 1- Measures of Asthma Assessment and Monitoring

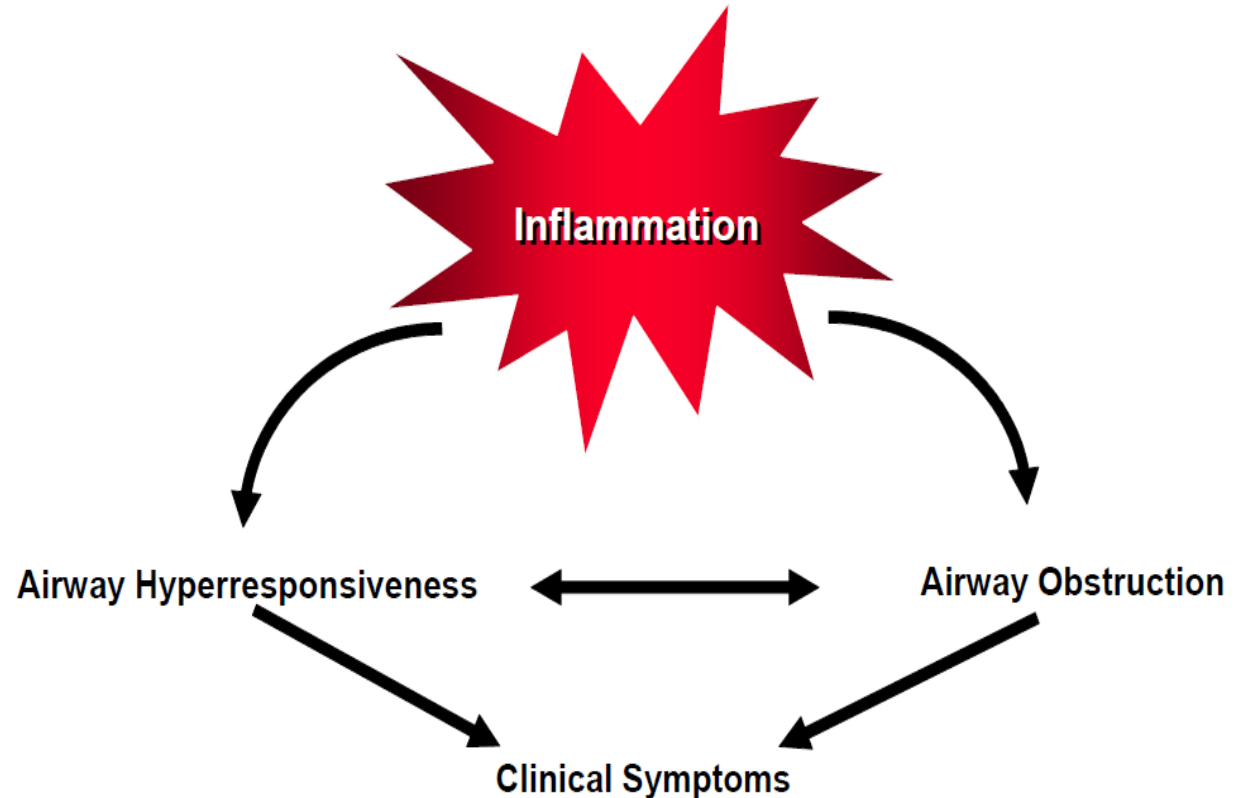
Defining Asthma

Asthma is a heterogeneous disease, usually characterized by chronic airway inflammation.

It is defined by the history of respiratory symptoms such as wheeze, shortness of breath, chest tightness and cough that vary over time and in intensity, together with variable expiratory airflow limitation.

GINA 2017

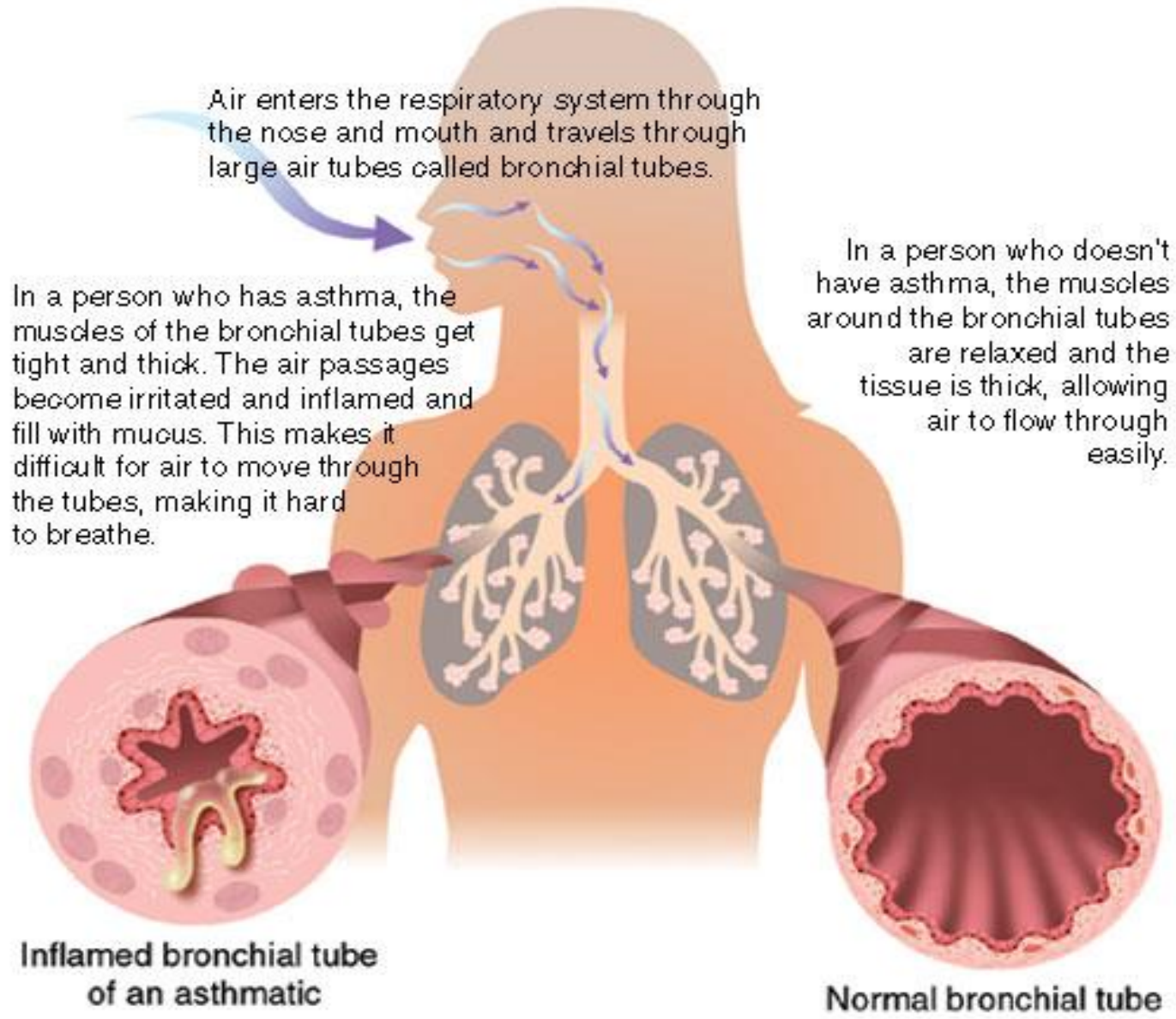
FIGURE 2-1. THE INTERPLAY AND INTERACTION BETWEEN AIRWAY INFLAMMATION AND THE CLINICAL SYMPTOMS AND PATHOPHYSIOLOGY OF ASTHMA



Adapted from: National Heart, Lung, and Blood Institute. *NAEPP Expert Panel Report 3*. Bethesda, MD: National Institutes of Health; 2007.

Asthma is not Reactive Airways Disease

- “Reactive Airway Disease” and “Asthma” are used interchangeably
 - RAD isn’t a specific diagnosis
 - The diagnosis of asthma can-and should be confirmed beginning at age of 5



Source: American Academy of Allergy, Asthma and Immunology

Air enters the respiratory system through the nose and mouth and travels through large air tubes called bronchial tubes.

Albuterol

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In a person who doesn't have asthma, the muscles around the bronchial tubes are relaxed and the tissue is thick, allowing air to flow through easily.

Inflamed bronchial tube of an asthmatic

Normal bronchial tube

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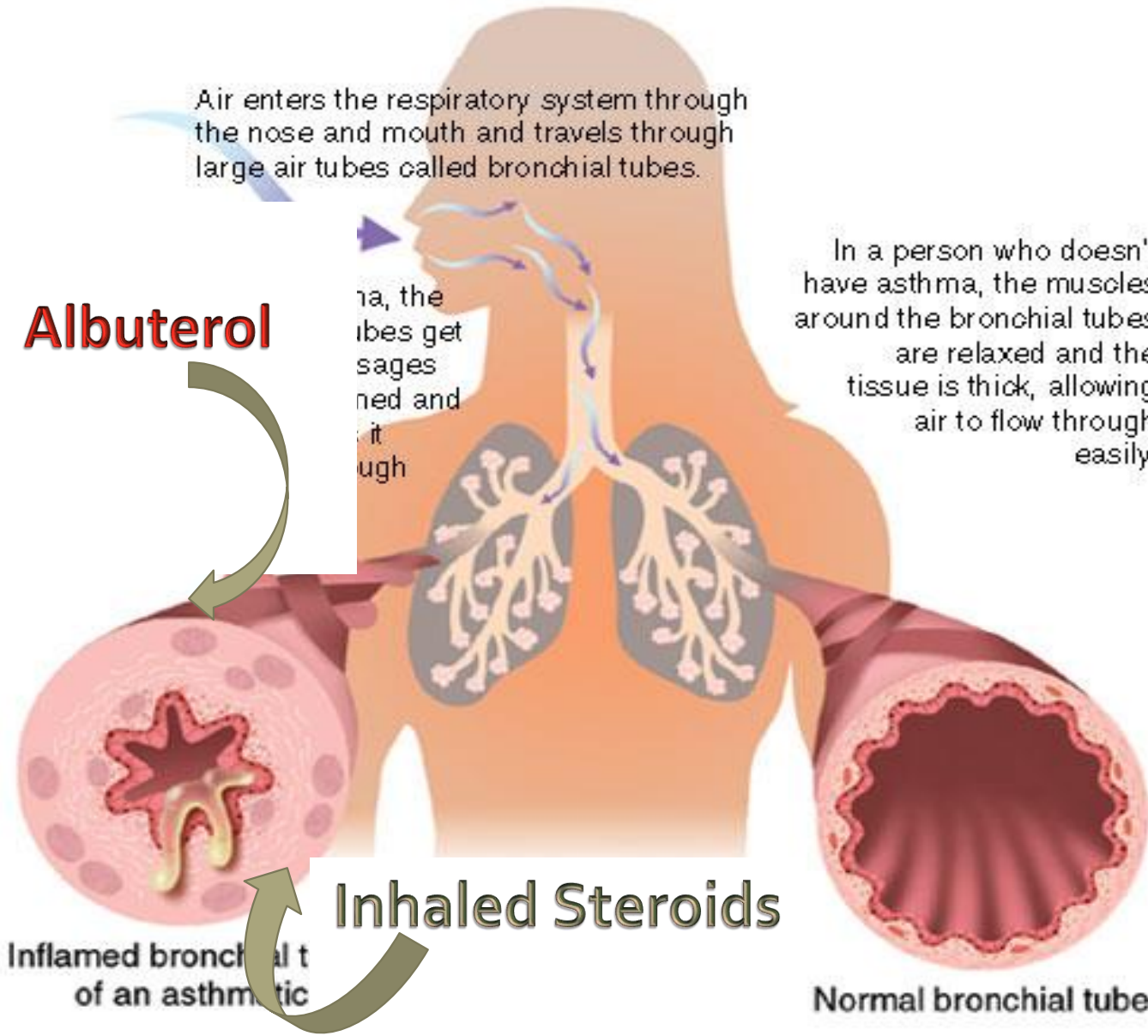
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Inhaled Steroids

Normal bronchial tube

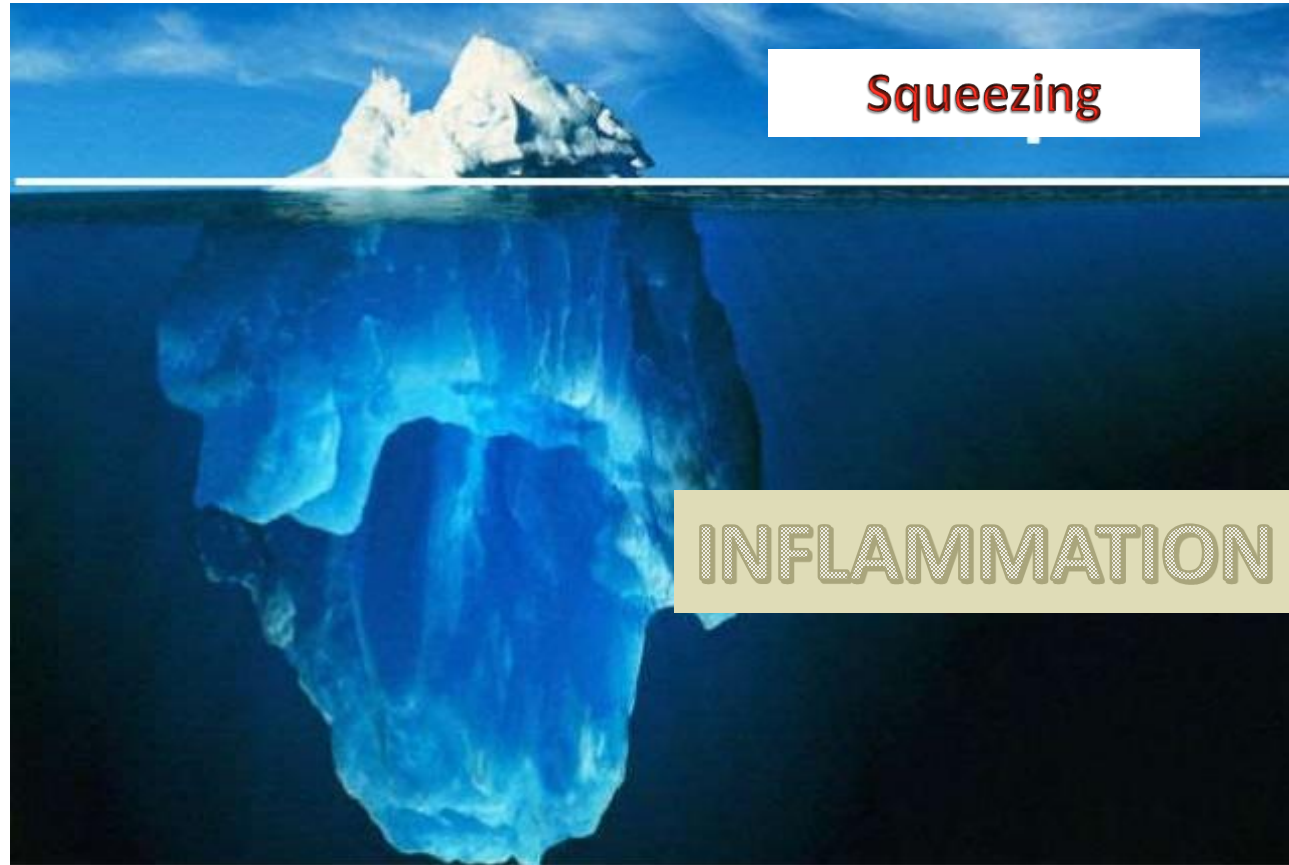
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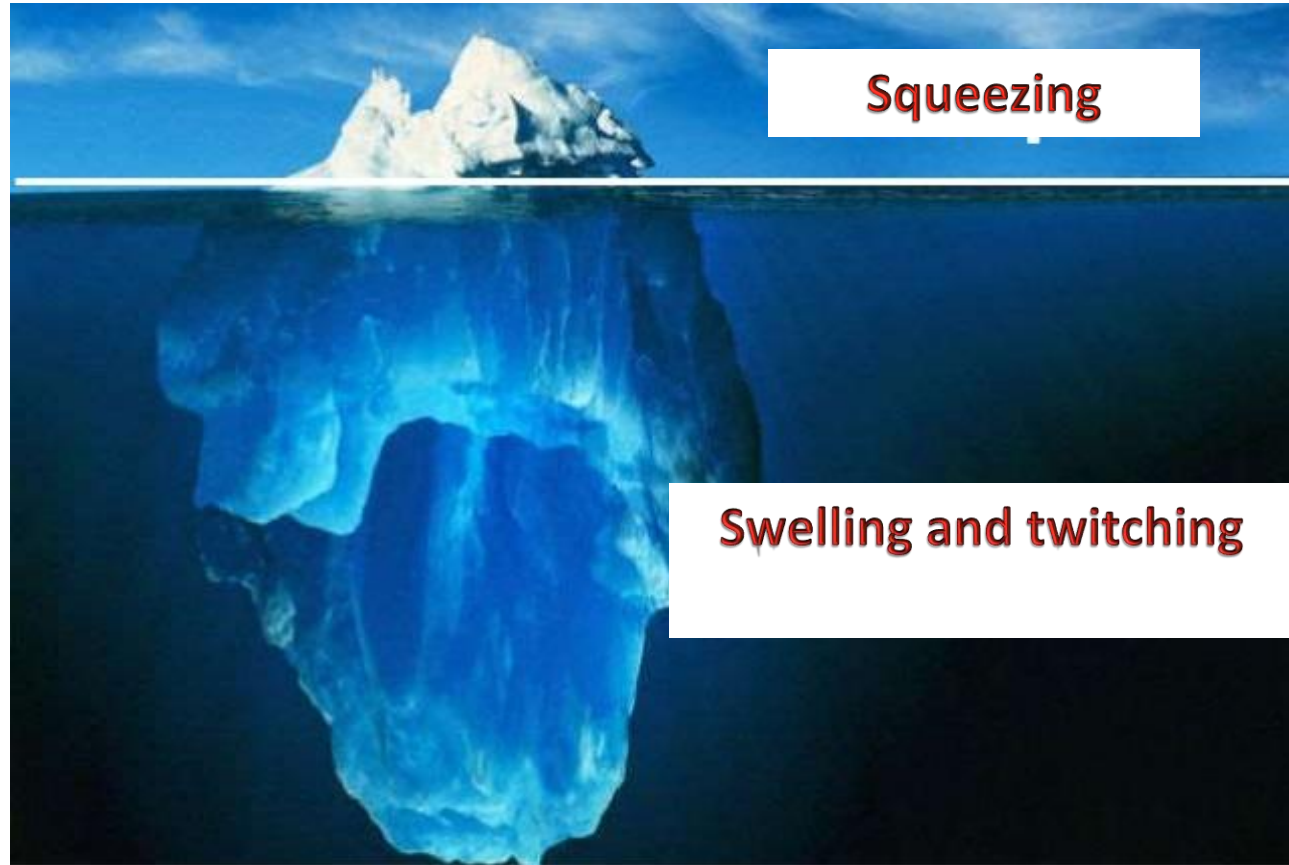
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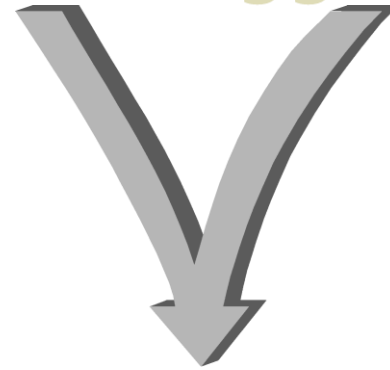


How do you get asthma?

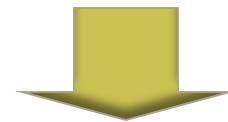
Family (genes)



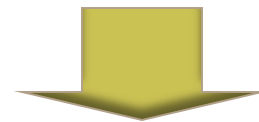
Triggers



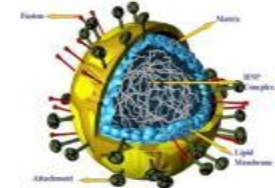
Swelling inside breathing tubes (silent)

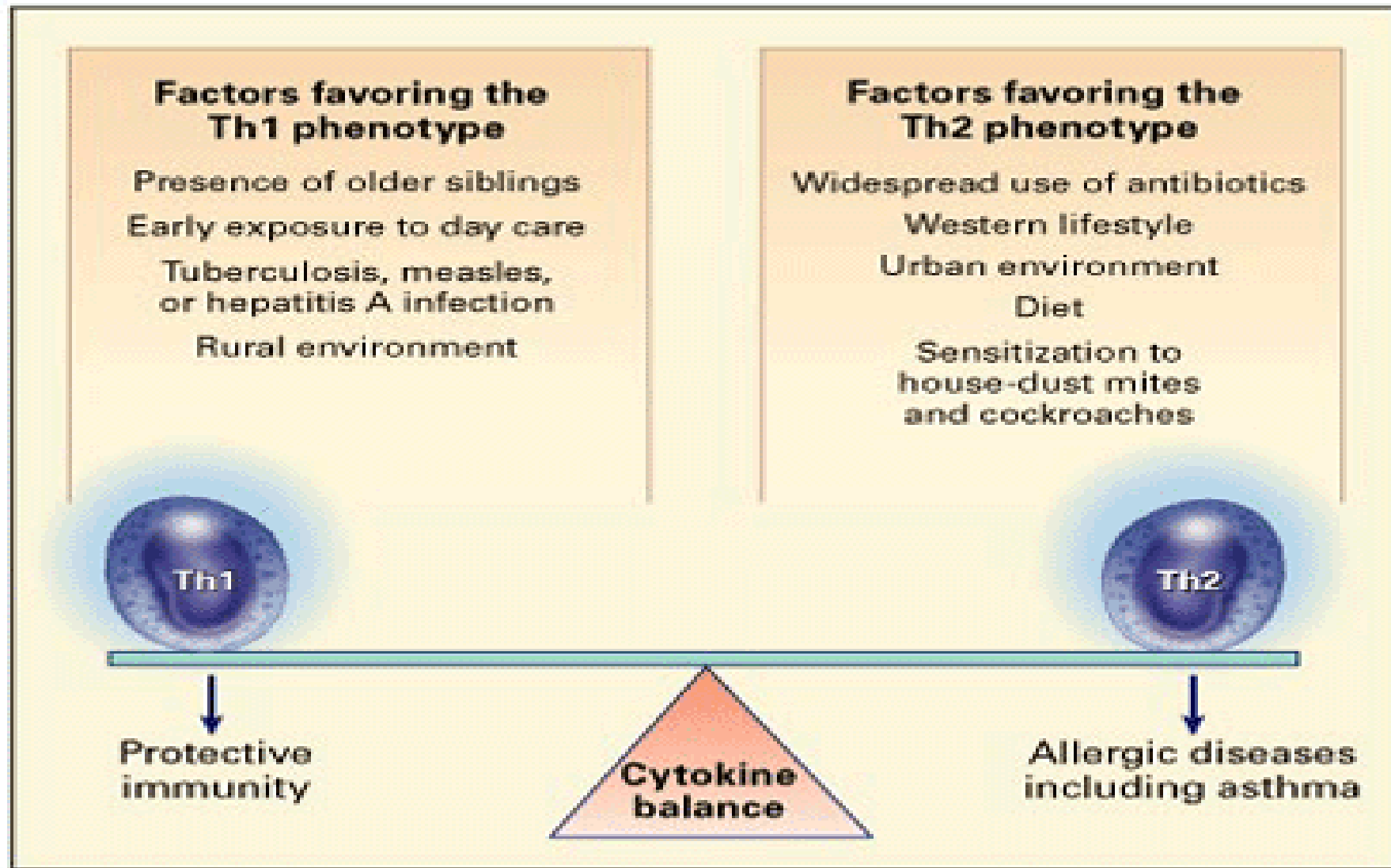


Squeezing of muscle around breathing tubes (noisy)



Wheeze, chest tightness, cough, shortness of breath

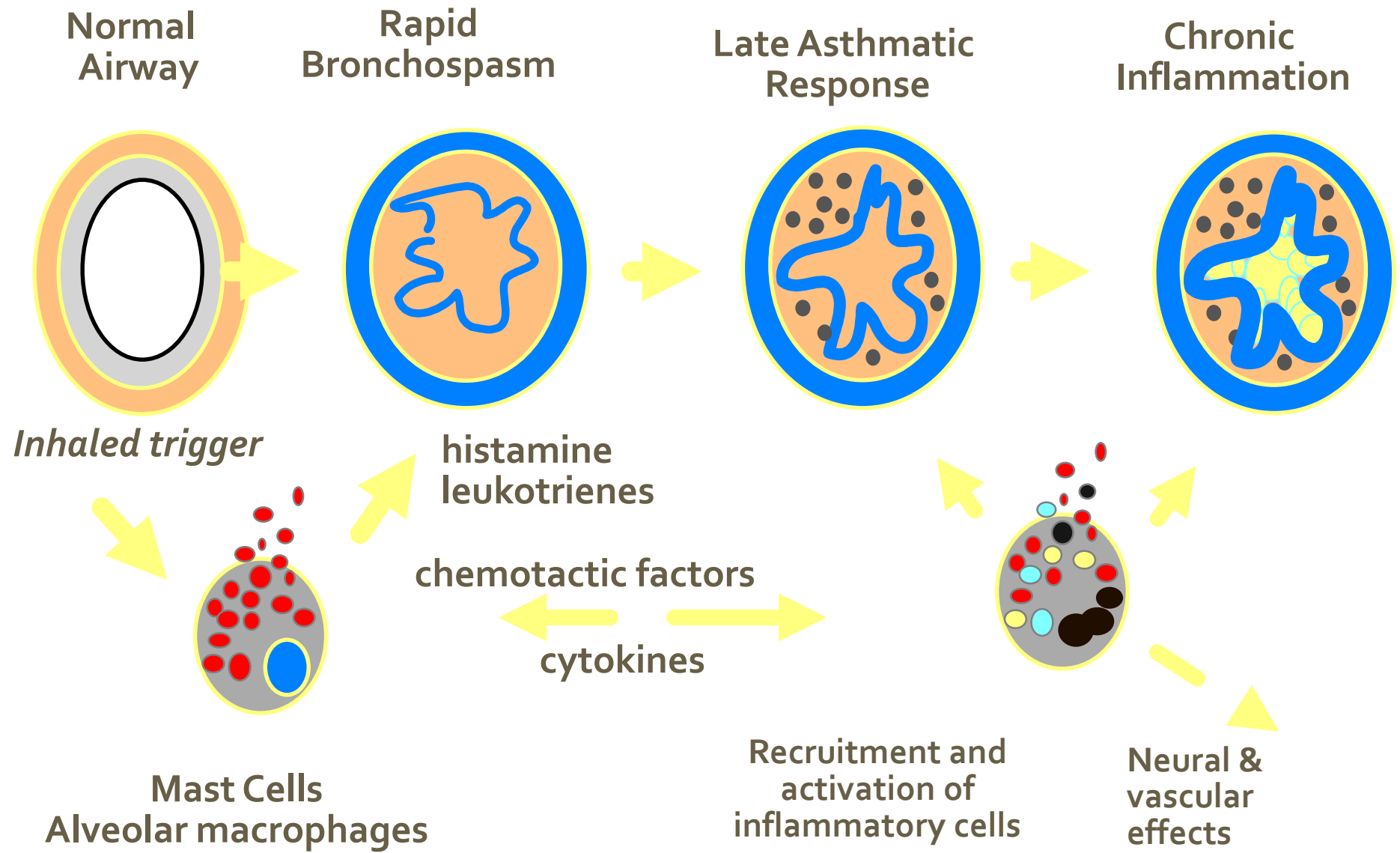




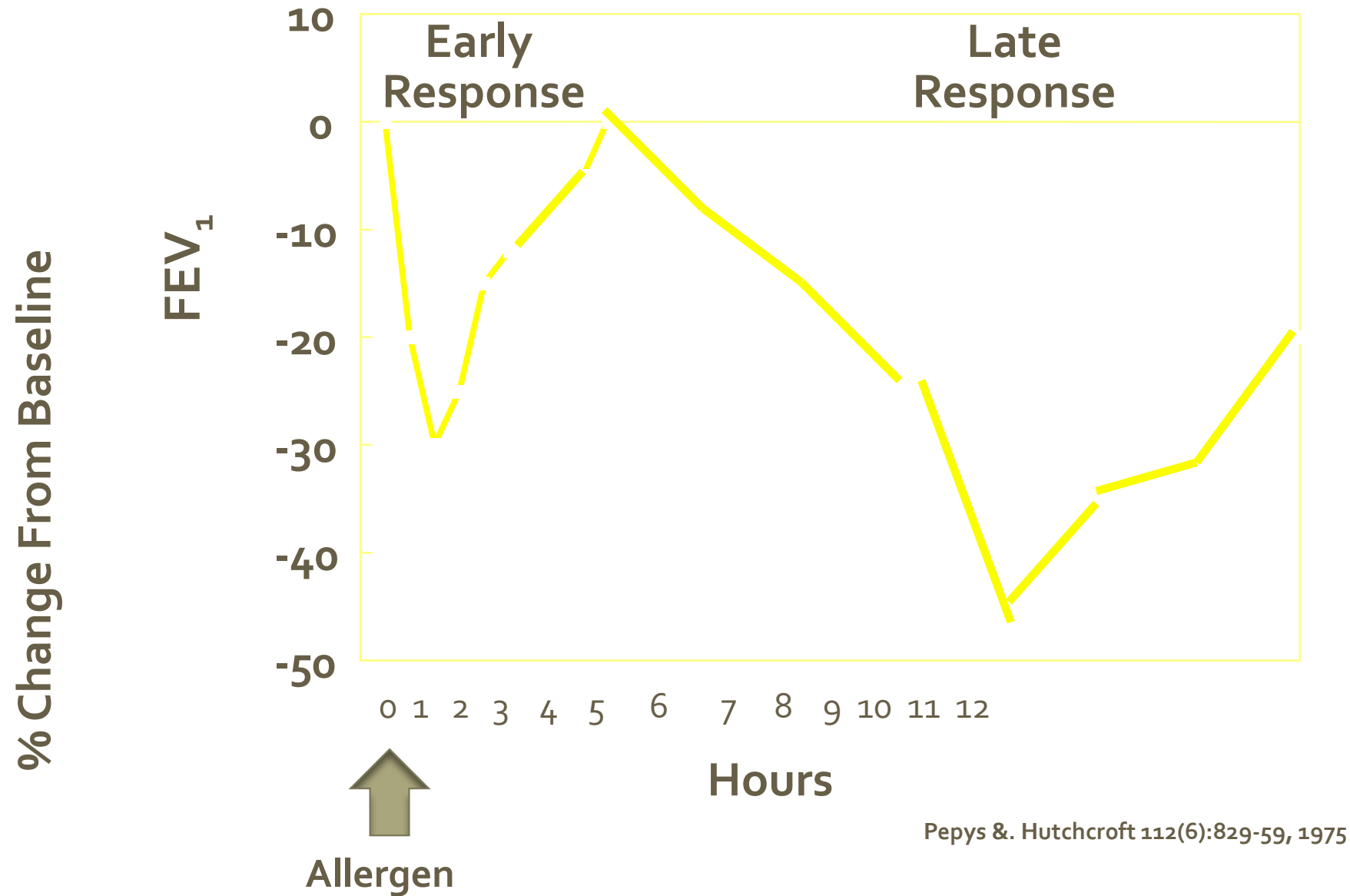
Hygiene Hypothesis

Allergic and non-allergic asthma

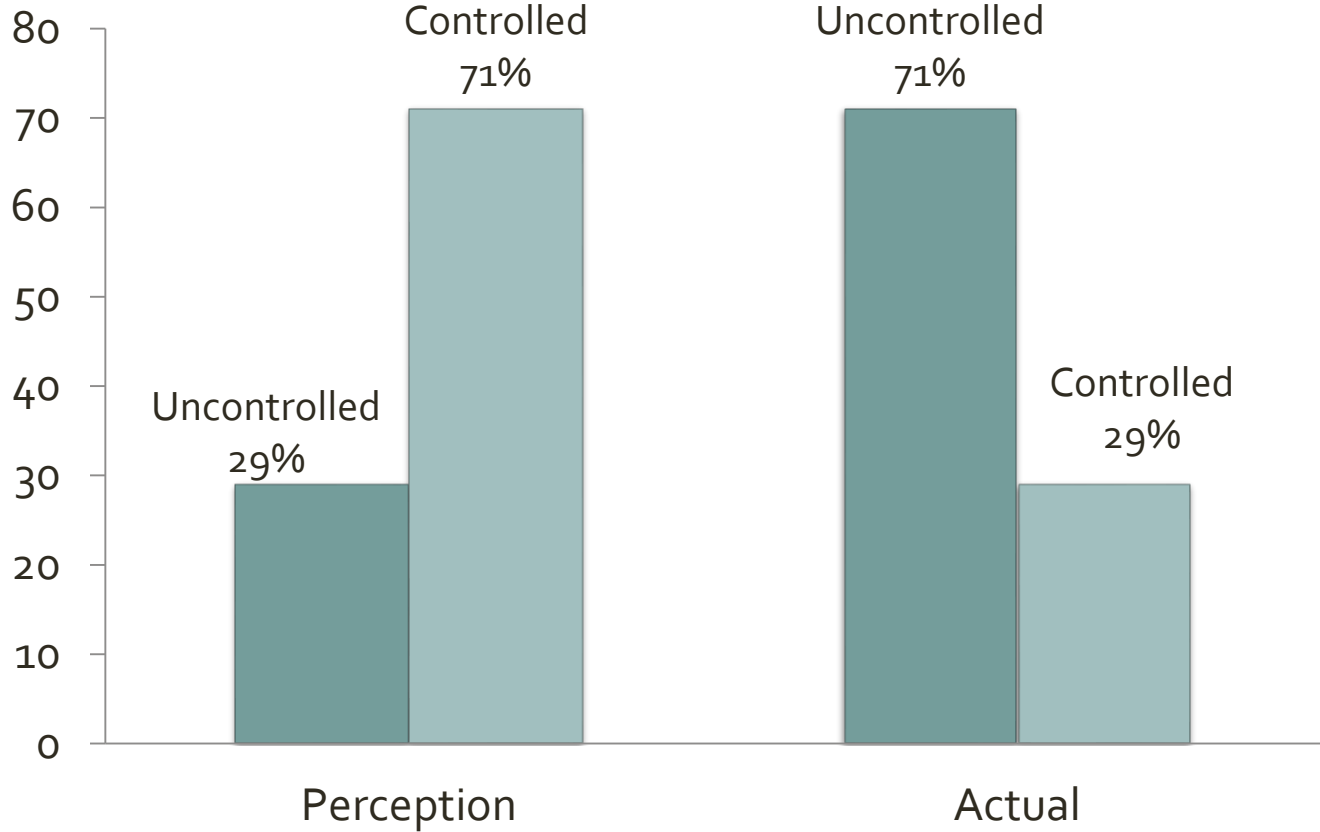
- Allergic asthma, or allergy-induced asthma, is the most common form of the disease
 - Most children have allergic asthma
- Most adults have non-allergic asthma
- Many of the symptoms of allergic and non-allergic asthma are the same but the triggers may differ



Early and Late Asthmatic Response



Asthma control: Perceived and actual

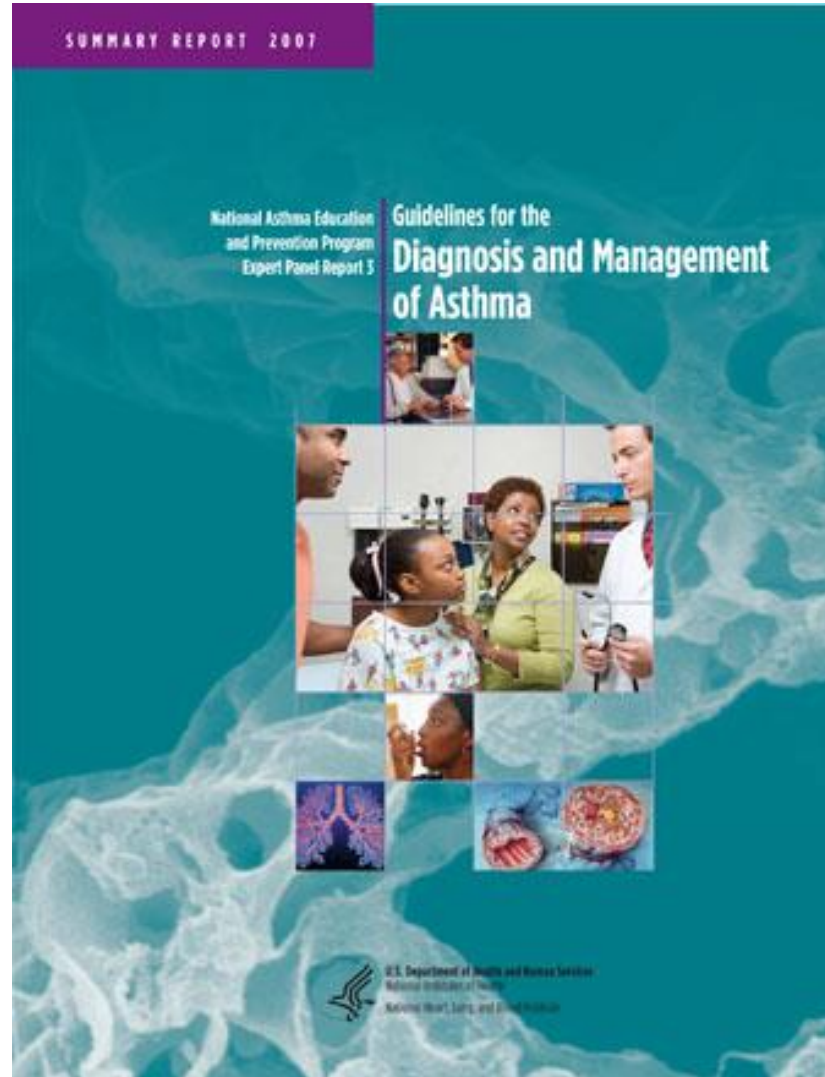


AIM study, 2009

Factors Contributing to Uncontrolled Asthma

- Failure to recognize or respond to signs and symptoms of asthma
- Inadequate treatment for level of severity
- Non-adherence
- Insufficient monitoring
- Failure to avoid or reduce exposure to asthma triggers
- Suboptimal patient-provider communication/partnership

Current standards for determining the diagnosis of asthma, as well as severity and control classifications



2007



Updated yearly

Diagnosis of asthma

- The diagnosis of asthma should be based on:
 - A history of characteristic symptom patterns
 - Evidence of variable airflow limitation, from bronchodilator reversibility testing or other tests

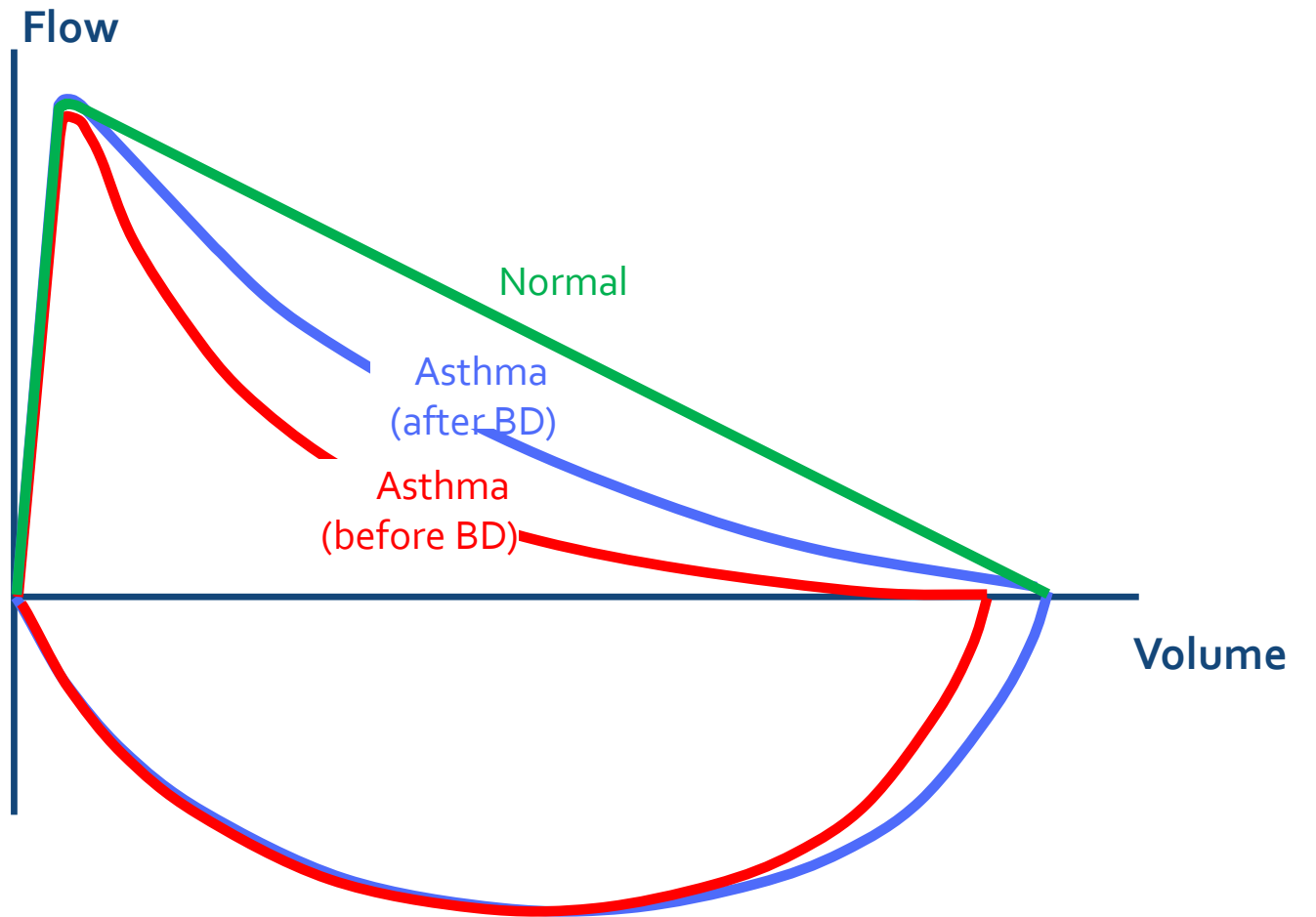
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- Document evidence for the diagnosis in the patient's notes, preferably before starting controller treatment
 - It is often more difficult to confirm the diagnosis after treatment has been started
- Asthma is usually characterized by airway inflammation and airway hyperresponsiveness, but these are not necessary or sufficient to make the diagnosis of asthma.

Typical spirometric tracings



Note: Each FEV₁ represents the highest of three reproducible measurements

Measures of lung function in asthma

Peak flow

- Used only for monitoring
- Can provide falsely high or falsely low readings
- More erroneous measures in children compared to adults
- SYMPTOM BASED ACTION PLAN IS PREFERABLE
- Efforts cannot be quality assured

Spirometry

- Used for diagnosis and monitoring
 - 12% improvement in the FEV₁ pre- post- SABA
 - If – BD response, is asthma ruled out?
 - FEV₁/FVC is better for determining severity
 - FEV₁ is better for predicting an exacerbation
- Efforts can be quality assured

MAPI: Predicting asthma in those 5 years and younger

Modified Asthma Predictive Index (mAPI)

≥4 Wheezing Illnesses *and*

≥1 Major criteria

- Parental asthma
- Atopic dermatitis (MD diagnosed)
- Aeroallergen sensitization

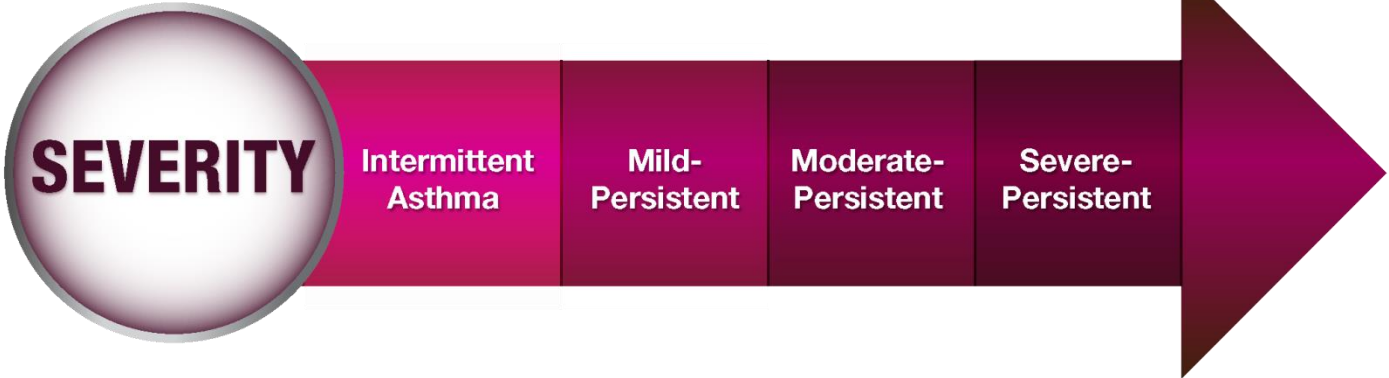
OR

≥2 Minor criteria

- Food sensitization
- Peripheral blood eosinophils ≥4%
- Wheezing apart from colds

Asthma Severity and Control

The intrinsic intensity of the disease process. It is most easily measured in individuals who are not receiving long-term control treatment



The degree to which the manifestations of asthma are minimized and the goals of therapy are met

Asthma
severity

The Pediatrician's Ready Reference Guide

Current NHLBI guidelines on assessing and monitoring asthma control

TABLE
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CLASSIFYING ASTHMA SEVERITY & INITIATION OF TREATMENT BY AGE

COMPONENTS OF SEVERITY		INTERMITTENT			PERSISTENT								
					MILD			MODERATE			SEVERE		
AGE IN YEARS		0-4	5-11	>12	0-4	5-11	>12	0-4	5-11	>12	0-4	5-11	>12
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RISK	Exacerbations requiring systemic corticosteroids	0-1x/year			≥2x/6 months OR >4x/year + risk factors			→			→		
RECOMMENDED STEP FOR INITIATING TREATMENT		STEP 1			STEP 2			STEP 3			STEP 3		STEP 4 OR STEP 5

Abbreviation: FEV, forced expiratory volume; FVC, forced vital capacity; PFT, pulmonary function test; NHLBI, National Heart, Lung, and Blood Institute; SABA, short-acting beta agonist.

From Reddy AP, et al.⁴

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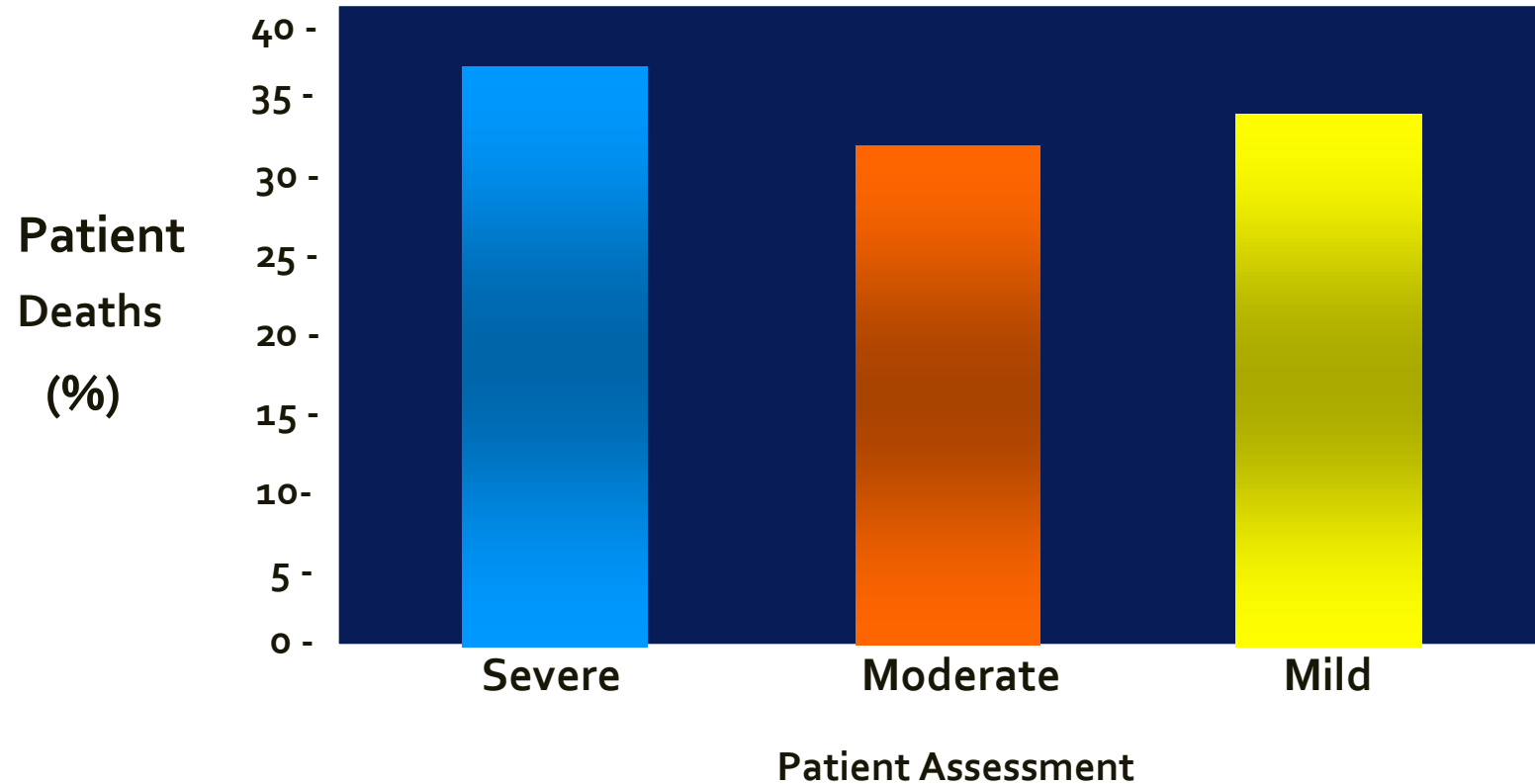
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	SABA use	≤2 days/week			≥2 days/week			Daily			Several times/day		
	Interferes with normal activity	None			Minor			Some			Extremely		
	PFT	FEV ₁	n/a	>80%		n/a	80%		n/a	60%-80%		n/a	<60%
	FEV ₁ /FVC	n/a	>85%	Normal ratio	n/a	>80%	Normal ratio	n/a	75%-80%	Reduced by >5%	n/a	<75%	Reduced by >5%
RISK	Exacerbations requiring systemic corticosteroids	0-1x/year			≥2x/6 months OR >4x/year + risk factors			→			→		
RECOMMENDED STEP FOR INITIATING TREATMENT		STEP 1			STEP 2			STEP 3			STEP 3		STEP 4 OR STEP 5

Abbreviation: FEV, forced expiratory volume; FVC, forced vital capacity; PFT, pulmonary function test; NHLBI, National Heart, Lung, and Blood Institute; SABA, short-acting beta agonist.

From Reddy AP, et al.⁴

Pediatric Patients With Mild Asthma: Mortality Risk



Findings from a cohort study reviewing all pediatric asthma-related deaths (n = 51) in the Australian state of Victoria from 1986 to 1989

Asthma control

Components of Control		Well Controlled			Not Well Controlled			Very Poorly Controlled		
		Ages 0-4 years	Ages 5-11 years	Ages ≥12 years	Ages 0-4 years	Ages 5-11 years	Ages ≥12 years	Ages 0-4 years	Ages 5-11 years	Ages ≥12 years
Impairment	Symptoms	≤2 days/week	≤2 days/week but not more than once on each day	≤2 days/week	>2 days/week	>2 days/week or multiple times on ≤2 days/week	>2 days/week	Throughout the day		
	Nighttime awakenings	≤1x/month		≤2x/month	>1x/month	≥2x/month	1-3x/week	>1x/week	≥2x/week	≥4x/week
	Interference with normal activity	None			Some limitation			Extremely limited		
	SABA* use for symptom control (not to prevent EIB*)	≤2 days/week			>2 days/week			Several times per day		
	Lung function ➔ FEV ₁ * (% predicted) or peak flow (% personal best)	Not applicable	>80%	>80%	Not applicable	60-80%	60-80%	Not applicable	<60%	<60%
	➔ FEV ₁ /FVC*		>80%	Not applicable		75-80%	Not applicable		<75%	Not applicable
	Validated questionnaires [†] ➔ ATAQ* ➔ ACQ* ➔ ACT*	Not applicable	Not applicable	0 ≤0.75 [‡] ≥20	Not applicable	Not applicable	1-2 ≥1.5 16-19	Not applicable	Not applicable	3-4 Not applicable ≤15
Risk	Asthma exacerbations requiring oral systemic corticosteroids [§]	0-1/year			2-3/year	≥2/year		>3/year	≥2/year	
		<i>Consider severity and interval since last asthma exacerbation.</i>								
	Reduction in lung growth/Progressive loss of lung function	Not applicable	Evaluation requires long-term follow-up care.		Not applicable	Evaluation requires long-term follow-up care.		Not applicable	Evaluation requires long-term follow-up care.	
Treatment-related adverse effects	<i>Medication side effects can vary in intensity from none to very troublesome and worrisome. The level of intensity does not correlate to specific levels of control but should be considered in the overall assessment of risk.</i>									

Components of Control		Well Controlled			Not Well Controlled			Very Poorly Controlled		
		Ages 0-4 years	Ages 5-11 years	Ages ≥12 years	Ages 0-4 years	Ages 5-11 years	Ages ≥12 years	Ages 0-4 years	Ages 5-11 years	Ages ≥12 years
Impairment	Symptoms	≤2 days/week	≤2 days/week but not more than once on each day	≤2 days/week	>2 days/week	>2 days/week or multiple times on ≤2 days/week	>2 days/week	Throughout the day		
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Validated questionnaires [†]										
➔ ATAQ*	Not applicable	Not applicable	0	Not applicable	Not applicable	1-2	Not applicable	Not applicable	3-4	
➔ ACQ*			≤0.75 [‡]			≥1.5			Not applicable	
➔ ACT*			≥20			16-19			≤15	
Risk	Asthma exacerbations requiring oral systemic corticosteroids [§]	0-1/year			2-3/year	≥2/year		>3/year	≥2/year	
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Asthma Control Test™ (ACT)

1. In the past 4 weeks, how much of the time did your asthma keep you from getting as much done at work, school or at home?

All of the time	1	Most of the time	2	Some of the time	3	A little of the time	4	None of the time	5
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SCORE

5

2. During the past 4 weeks, how often have you had shortness of breath?

More than once a day	1	Once a day	2	3 to 6 times a week	3	Once or twice a week	4	Not at all	5
----------------------	---	------------	---	---------------------	---	----------------------	---	------------	---

2

3. During the past 4 weeks, how often did your asthma symptoms (wheezing, coughing, shortness of breath, chest tightness or pain) wake you up at night or earlier than usual in the morning?

4 or more nights a week	1	2 or 3 nights a week	2	Once a week	3	Once or twice	4	Not at all	5
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1

4. During the past 4 weeks, how often have you used your rescue inhaler or nebulizer medication (such as albuterol)?

3 or more times per day	1	1 or 2 times per day	2	2 or 3 times per week	3	Once a week or less	4	Not at all	5
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3

5. How would you rate your asthma control during the past 4 weeks?

Not controlled at all	1	Poorly controlled	2	Somewhat controlled	3	Well controlled	4	Completely controlled	5
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5

TOTAL

16

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American Academy
of Pediatrics



Referral to A Specialist

- ✓ When additional diagnostic testing or disease management education is needed
- ✓ When the patient needs an evaluation for immunotherapy
- ✓ When the patient requires an increase in the dose and number of medications as well as frequency of administration
- ✓ If a patient required >2 bursts of oral steroids in a 12-month period, or was hospitalized for asthma
- ✓ Following a life-threatening exacerbation
- ✓ When the patient is not meeting goals of therapy after 3-6 months of treatment
- ✓ When the patient has atypical signs or symptoms
- ✓ When comorbidities are complicating the case, including occupational or environmental exposures

Assessment of asthma summary

1. Asthma control - two domains
 - Assess symptom control
 - Assess risk factors for poor outcomes, including low lung function

Assessment of asthma summary

1. Asthma control - two domains
 - Assess symptom control
 - Assess risk factors for poor outcomes, including low lung function
2. Treatment issues
 - Check inhaler technique and adherence
 - Ask about side-effects
 - Does the patient have a written asthma action plan?
 - What are the patient's attitudes and goals for their asthma?

Assessment of asthma summary

1. Asthma control - two domains
 - Assess symptom control
 - Assess risk factors for poor outcomes, including low lung function
2. Treatment issues
 - Check inhaler technique and adherence
 - Ask about side-effects
 - Does the patient have a written asthma action plan?
 - What are the patient's attitudes and goals for their asthma?
3. Comorbidities
 - Think of rhinosinusitis, GERD, obesity, obstructive sleep apnea, depression, anxiety
 - These may contribute to symptoms and poor quality of life



Component 2: Control of Environmental Factors and Comorbid Conditions that Affect Asthma

Allergens and Irritants

- Allergens
 - Animal dander
 - Cockroach
 - Dust mites
 - Indoor mold
 - Pollen
 - Outdoor Mold

- Irritants
 - Smoke
 - Strong odors
 - Sprays
 - Sulfites in foods

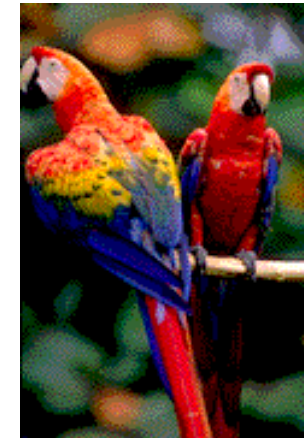
Miscellaneous

- Exercise
- Other medicines
- Weather
- Co-morbid conditions

If allergic to any fur-bearing or feathered animals:



- Remove pets from home
- Keep pets 100% out of bedroom and keep door closed
- Remove carpet and cloth-covered furniture from the home or keep pet out of rooms with these

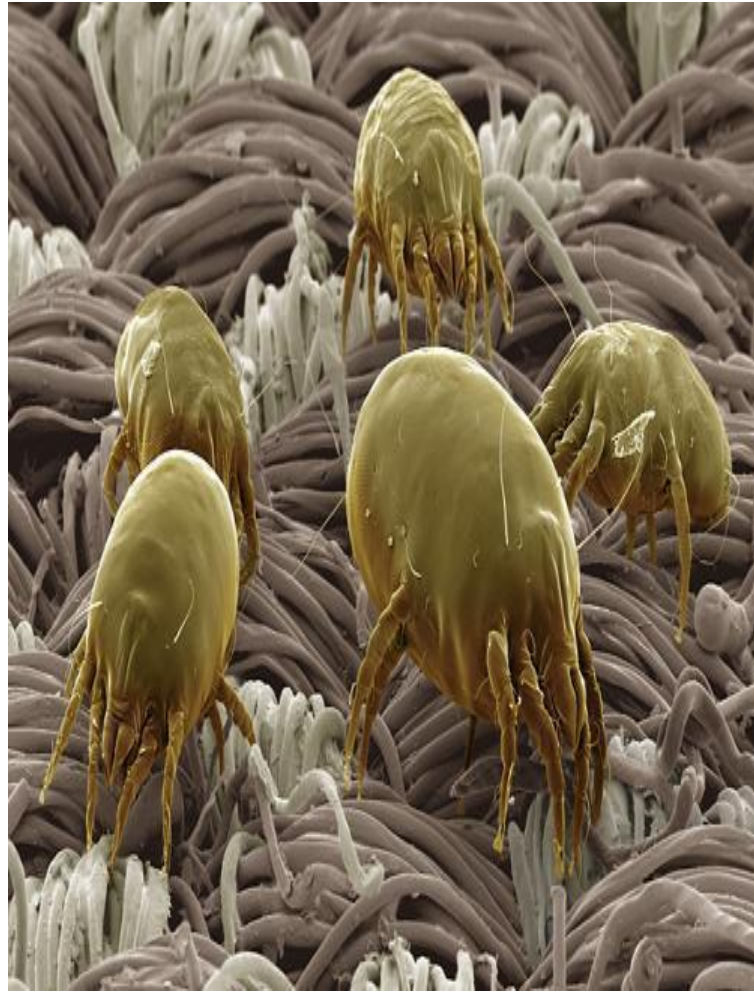


Cockroach Avoidance

- Keep all food out of bedroom
- Store food and garbage in close containers
- Use bait traps
 - If you spray, stay out of home



Dust Mite Control



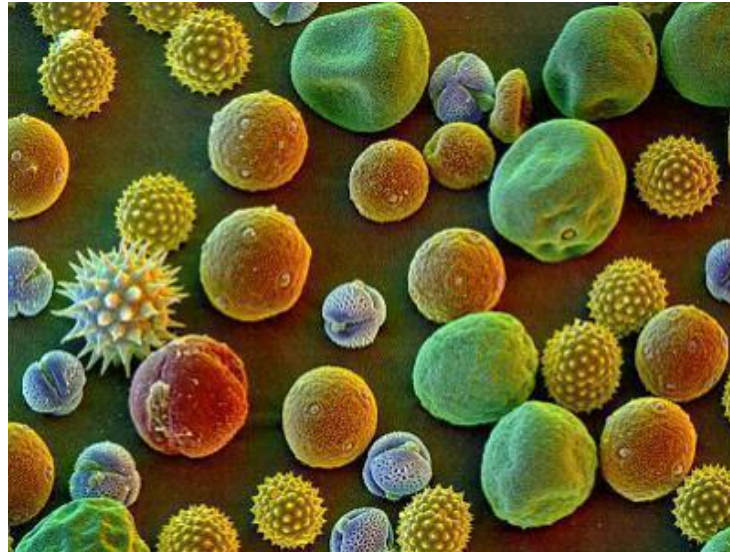
- Encase mattress/ pillow in zippered vinyl cover
- Wash bedding weekly at 130° F
- Reduce clutter
- Have someone vacuum/sweeping
 - Dustmask
 - Central cleaner with outside receptacle
 - Vacuum cleaner with HEPA filter or double-layered bag

Indoor Mold

- Fix source of water
- Clean moldy areas
 - Do not use bleach!
- Remove rugs that have gotten wet
- Dehumidify basements



Pollen and Outdoor Mold



Keep windows closed

Stay indoors from midday-afternoon

Re-evaluate need for additional medicines or intensification of asthma medicines during allergy seasons

Thirdhand smoke



All children are considered “sensitive”

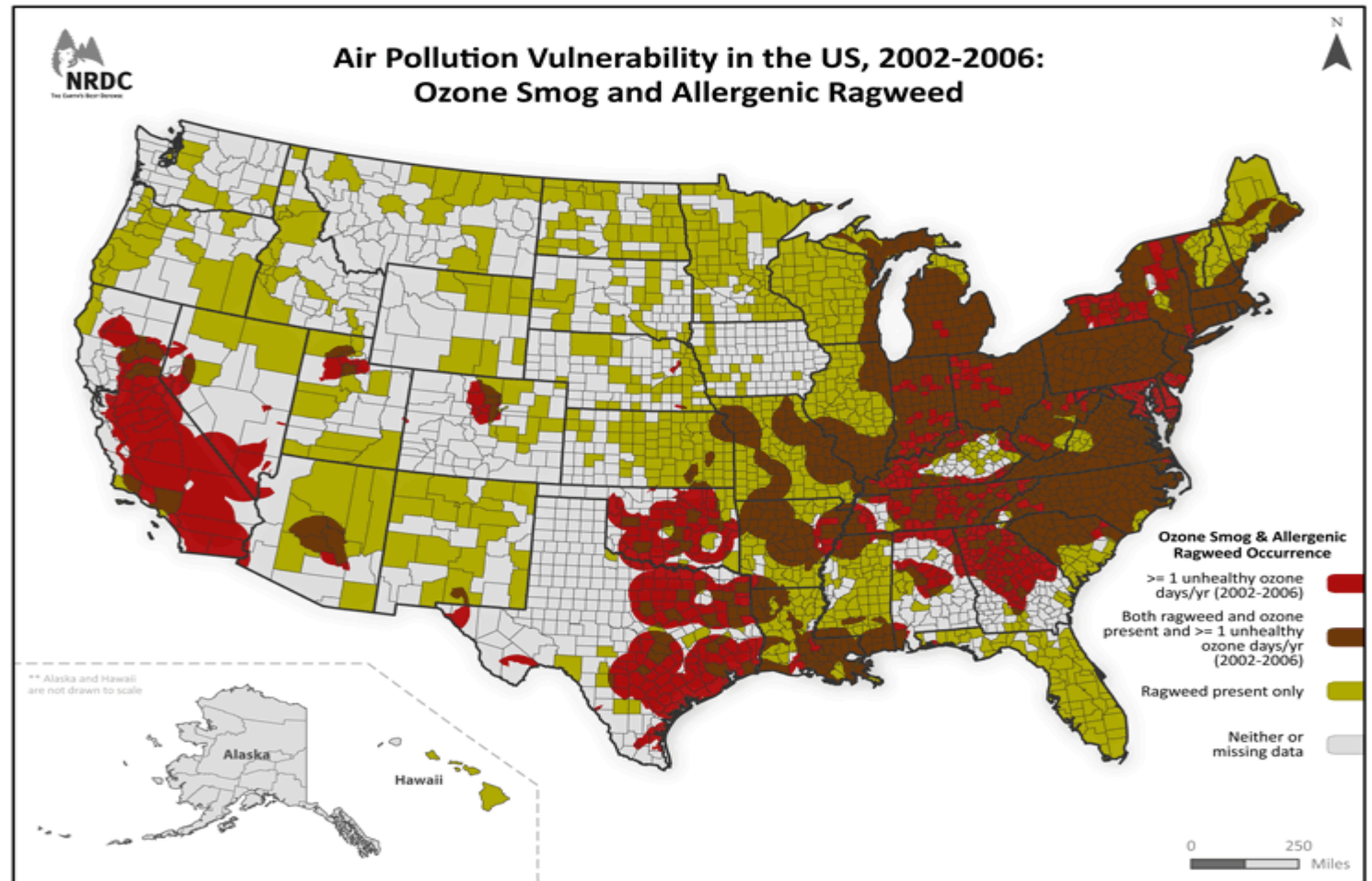
Air Quality Index (AQI) values	Levels of Health Concern	Colors
When the AQI is in this range:	...air quality conditions are:	...as symbolized by this color:
0 to 50	Good	Green
51 to 100	Moderate	Yellow
101 to 150	Unhealthy for Sensitive Groups	Orange
151 to 200	Unhealthy	Red
201 to 300	Very Unhealthy	Purple
301 to 500	Hazardous	Maroon

New risks for
asthma
related to
climate
change

Climate change effects on asthma

- GHGs will drive
 - Higher rates of asthma due to changes in allergen exposure
 - More intense and more frequent storms (“thunderstorm asthma”)
- More wildfires and dust storms will increase transport of PM, bacteria, fungi and influenza
- Excessive heat and/or poor air quality increases symptoms, decrease lung function
- Children are particularly vulnerable
- More flooding will increase mold and microbial growth

Triple threat:
Asthma, ozone
and pollens





Component 3: Pharmacologic + Therapy

Stepping up REMEMBER TO...

- Provide guided self-management education
- Treat modifiable risk factors and comorbidities
- Advise about non-pharmacological therapies and strategies
- Consider stepping up if ... uncontrolled symptoms, exacerbations or risks,
but check diagnosis, inhaler technique and adherence first
- Consider stepping down if ... symptoms controlled for 3 months
+ low risk for exacerbations. Ceasing ICS is not advised.

Stepwise Approach for Managing Asthma Long Term

Step UP if needed (first check inhaler technique, adherence, environmental control, and comorbid conditions) ASSESS CONTROL Step DOWN if possible (and asthma is well controlled for at least 3 months)							
		Step 1	Step 2	Step 3	Step 4	Step 5	Step 6
0-4 Years		Intermittent Asthma	Persistent Asthma: Daily Medication				
	Preferred	SABA as needed	Low-dose ICS	Medium-dose ICS	Medium-dose ICS + LABA or montelukast	High-dose ICS + LABA or montelukast	High-dose ICS + Oral corticosteroids + LABA or montelukast
	Alternative		Cromolyn or montelukast				
	Rescue Medication	Patient education and environmental control at each step. <ul style="list-style-type: none"> • SABA as needed for symptoms. Treatment intensity depends on symptom severity. • With viral respiratory symptoms, SABA every 4-6 hours up to 24 hours (longer with physician consult). • Consider short course of oral corticosteroids if exacerbation is severe or if patient has history of previous severe exacerbations. • Frequent or increasing use of SABA may indicate inadequate control and the need to step up treatment. 					
5-11 Years		Intermittent Asthma	Persistent Asthma: Daily Medication				
	Preferred	SABA as needed	Low-dose ICS	Low-dose ICS + LABA, LTRA, or Theophylline	Medium-dose ICS + LABA	High-dose ICS + LABA	High-dose ICS + LABA + Oral corticosteroids
	Alternative		Cromolyn, LTRA, Nedrocromil, or Theophylline	OR Medium-dose ICS	Medium-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline + Oral corticosteroids
	Rescue Medication	Patient education and environmental control, and management of comorbidities at each step. Step 2-4: Consider subcutaneous allergen immunotherapy for patients who have allergic asthma. <ul style="list-style-type: none"> • SABA as needed for symptoms – up to 3 treatments at 20-minute intervals initially. Treatment intensity depends on symptom severity. • Consider short course of oral corticosteroids. • Increasing use of SABA or use > 2 days/week for symptom relief (not prevention of EIB) generally indicates inadequate control and the need to step up treatment. 					
≥ 12 Years		Intermittent Asthma	Persistent Asthma: Daily Medication				
	Preferred	SABA as needed	Low-dose ICS	Low-dose ICS + LABA OR Medium-dose ICS	Medium-dose ICS + LABA	High-dose ICS + LABA	High-dose ICS + LABA + Oral corticosteroid
	Alternative		Cromolyn, LTRA, Nedrocromil, or Theophylline	Low-dose ICS + LTRA, Theophylline, or Zileuton	Medium-dose ICS + LTRA, Theophylline, or Zileuton	Consider Omalizumab for patients who have allergic asthma	Consider Omalizumab for patients who have allergic asthma
	Rescue Medication	Patient education and environmental control, and management of comorbidities at each step. Step 2-4: Consider subcutaneous allergen immunotherapy for patients who have allergic asthma. <ul style="list-style-type: none"> • SABA as needed for symptoms – up to 3 treatments at 20-minute intervals initially. Treatment intensity depends on symptom severity. • Consider short course of oral corticosteroids. • Increasing use of SABA or use > 2 days/week for symptom relief (not prevention of EIB) generally indicates inadequate control and the need to step treatment. 					
All	Notes	<ul style="list-style-type: none"> • If an alternative treatment is used and response is inadequate, discontinue it and use the preferred treatment before stepping up. • Theophylline requires serum concentration levels monitoring; zileuton requires liver function monitoring. • LABAs are not indicated for acute symptom relief and should be used in combination with an ICS. 					

Stepwise Approach for Managing Asthma Long Term

		Step UP if needed (first check inhaler technique, adherence, environmental control, and comorbid conditions) ASSESS CONTROL Step DOWN if possible (and asthma is well controlled for at least 3 months)					
		Step 1		Step 3		Step 5	
		Step 2		Step 4		Step 6	
0-4 Years		Intermittent Asthma		Persistent Asthma: Daily Medication			
	Preferred	SABA as needed		Consult with asthma specialist if step 3 care or higher is required. Consider consultation at step 2.			
	Alternative	Cromolyn or montelukast		Low-dose ICS		Medium-dose ICS + LABA or montelukast	
	Rescue Medication	Patient education and environmental control at each step. <ul style="list-style-type: none"> SABA as needed for symptoms. Treatment intensity depends on symptom severity. With viral respiratory symptoms, SABA every 4-6 hours up to 24 hours (longer with physician consult). Consider short course of oral corticosteroids if exacerbation is severe or if patient has history of previous severe exacerbations. Frequent or increasing use of SABA may indicate inadequate control and the need to step up treatment. 					
5-11 Years		Intermittent Asthma		Persistent Asthma: Daily Medication			
	Preferred	SABA as needed		Consult with asthma specialist if step 4 care or higher is required. Consider consultation at step 3.			
	Alternative	Cromolyn, LTRA, Nedrocromil, or Theophylline		Low-dose ICS + LABA, LTRA, or Theophylline OR Medium-dose ICS		Medium-dose ICS + LABA	
	Rescue Medication	Patient education and environmental control, and management of comorbidities at each step. Step 2-4: Consider subcutaneous allergen immunotherapy for patients who have allergic asthma. <ul style="list-style-type: none"> SABA as needed for symptoms – up to 3 treatments at 20-minute intervals initially. Treatment intensity depends on symptom severity. Consider short course of oral corticosteroids. Increasing use of SABA or use > 2 days/week for symptom relief (not prevention of EIB) generally indicates inadequate control and the need to step up treatment. 					
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	Alternative	Cromolyn, LTRA, Nedrocromil, or Theophylline		Low-dose ICS + LABA OR Medium-dose ICS		Medium-dose ICS + LABA	
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		Step 1	Step 2	Step 3	Step 4	Step 5	Step 6
0-4 Years		Intermittent Asthma	Persistent Asthma: Daily Medication Consult with asthma specialist if step 3 care or higher is required. Consider consultation at step 2.				
	Preferred	SABA as needed	Low-dose ICS	Medium-dose ICS	Medium-dose ICS + LABA or montelukast	High-dose ICS + LABA or montelukast	High-dose ICS + Oral corticosteroids + LABA or montelukast
	Alternative		Cromolyn or montelukast				
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5-11 Years		Intermittent Asthma	Persistent Asthma: Daily Medication Consult with asthma specialist if step 4 care or higher is required. Consider consultation at step 3.				
	Preferred	SABA as needed	Low-dose ICS	Low-dose ICS + LABA, LTRA, or Theophylline	Medium-dose ICS + LABA	High-dose ICS + LABA	High-dose ICS + LABA + Oral corticosteroids
	Alternative		Cromolyn, LTRA, Nedrocromil, or Theophylline	OR Medium-dose ICS	Medium-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline + Oral corticosteroids
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		Step 1	Step 2	Step 3	Step 4	Step 5	Step 6
0-4 Years		Intermittent Asthma	Persistent Asthma: Daily Medication Consult with asthma specialist if step 3 care or higher is required. Consider consultation at step 2.				
	Preferred	SABA as needed	Low-dose ICS	Medium-dose ICS	Medium-dose ICS + LABA or montelukast	High-dose ICS + LABA or montelukast	High-dose ICS + Oral corticosteroids + LABA or montelukast
	Alternative		Cromolyn or montelukast				
	Rescue Medication	Patient education and environmental control at each step. <ul style="list-style-type: none"> SABA as needed for symptoms. Treatment intensity depends on symptom severity. With viral respiratory symptoms, SABA every 4-6 hours up to 24 hours (longer with physician consult). Consider short course of oral corticosteroids if exacerbation is severe or if patient has history of previous severe exacerbations. Frequent or increasing use of SABA may indicate inadequate control and the need to step up treatment. 					
5-11 Years		Intermittent Asthma	Persistent Asthma: Daily Medication Consult with asthma specialist if step 4 care or higher is required. Consider consultation at step 3.				
	Preferred	SABA as needed	Low-dose ICS	Low-dose ICS + LABA, LTRA, or Theophylline	Medium-dose ICS + LABA	High-dose ICS + LABA	High-dose ICS + LABA + Oral corticosteroids
	Alternative		Cromolyn, LTRA, Nedrocromil, or Theophylline	OR Medium-dose ICS	Medium-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline + Oral corticosteroids
	Rescue Medication	Patient education and environmental control, and management of comorbidities at each step. Step 2-4: Consider subcutaneous allergen immunotherapy for patients who have allergic asthma. <ul style="list-style-type: none"> SABA as needed for symptoms – up to 3 treatments at 20-minute intervals initially. Treatment intensity depends on symptom severity. Consider short course of oral corticosteroids. Increasing use of SABA or use > 2 days/week for symptom relief (not prevention of EIB) generally indicates inadequate control and the need to step up treatment. 					
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	Preferred	SABA as needed	Low-dose ICS	Low-dose ICS + LABA OR Medium-dose ICS	Medium-dose ICS + LABA	High-dose ICS + LABA	High-dose ICS + LABA + Oral corticosteroid
	Alternative		Cromolyn, LTRA, Nedrocromil, or Theophylline	Low-dose ICS + LTRA, Theophylline, or Zileuton	Medium-dose ICS + LTRA, Theophylline, or Zileuton	Consider Omalizumab for patients who have allergic asthma	Consider Omalizumab for patients who have allergic asthma
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Stepwise Approach for Managing Asthma Long Term

		Step UP if needed (first check inhaler technique, adherence, environmental control, and comorbid conditions) ASSESS CONTROL Step DOWN if possible (and asthma is well controlled for at least 3 months)					
		←				→	
		Step 1	Step 2	Step 3	Step 4	Step 5	Step 6
0-4 Years		Intermittent Asthma	Persistent Asthma: Daily Medication				
			Consult with asthma specialist if step 3 care or higher is required. Consider consultation at step 2.				
	Preferred	SABA as needed	Low-dose ICS	Medium-dose ICS	Medium-dose ICS + LABA or montelukast	High-dose ICS + LABA or montelukast	High-dose ICS + Oral corticosteroids + LABA or montelukast
	Alternative		Cromolyn or montelukast				
Rescue Medication	Patient education and environmental control at each step. <ul style="list-style-type: none"> • SABA as needed for symptoms. Treatment intensity depends on symptom severity. • With viral respiratory symptoms, SABA every 4-6 hours up to 24 hours (longer with physician consult). • Consider short course of oral corticosteroids if exacerbation is severe or if patient has history of previous severe exacerbations. • Frequent or increasing use of SABA may indicate inadequate control and the need to step up treatment. 						
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	Preferred	SABA as needed	Low-dose ICS	Low-dose ICS + LABA, LTRA, or Theophylline	Medium-dose ICS + LABA	High-dose ICS + LABA	High-dose ICS + LABA + Oral corticosteroids
	Alternative		Cromolyn, LTRA, Nedrocromil, or Theophylline	OR Medium-dose ICS	Medium-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline + Oral corticosteroids
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0-4 Years		Intermittent Asthma		Persistent Asthma: Daily Medication Consult with asthma specialist if step 3 care or higher is required. Consider consultation at step 2.									
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	Alternative			Cromolyn, LTRA, Nedrocromil, or Theophylline		OR Medium-dose ICS	Medium-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline + Oral corticosteroids				
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	Alternative		Cromolyn, LTRA, Nedrocromil, or Theophylline	OR Medium-dose ICS	Medium-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline	High-dose ICS + LTRA or Theophylline + Oral corticosteroids
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Common treatments for the medical management of Asthma

Quick-relief “rescue”

Short-acting bronchodilators prn
(beta 2 agonists - SABAs - preferred)

“Burst” of systemic corticosteroids

Long-term control

Inhaled corticosteroids (ICS)

Combination therapy (inhaled
corticosteroids and long-acting
beta 2 agonists (ICS/LABA)

Leukotriene modifiers

Biologic agents

*At each treatment step, patient education, environmental control,
and management of comorbidities need to be addressed*

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Rescue SABAs

Quick-relief "rescue"

SABAs preferred: Ventolin, Proventil. Proair
MDI and Respiclick-ALL ALBUTEROL,

Xopenex (LEVALBUTEROL)

SAMAs in the ER: Atrovent (IPRATROPIUM),
Duoneb, Combivent (BOTH ALBUTEROL &
IPRATROPIUM)

OCS: Prednisone, Medrol, Prelone

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Long-term control

ICS:

QVAR (BECLOMETHASONE),
Flovent (FP),

Alvesco (CICLESONIDE),
Aerospan (FLUNISOLIDE),
Asmanex (MOMETASONE),
Pulmicort (BUDESONIDE),
Arnuity (FF),

ArmonAir (FP)

Anti-IL-5 mab:
Nucala
(MEPOLIZUMAB)

SQ q 4 wks

Cinqair
(RESLIZUMAB)

IV q 4 weeks over
20-50 "

Anti-IgE mab: Xolair
(OMALIZUMAB)

SQ q 2-4 wks

Combination therapy:

Advair (FP/SALMETEROL),

Symbicort (BUDESONIDE/FORMOTEROL),

Dulera (MOMETASONE/FORMOTEROL), Breo FF/VILANTEROL,

Airduo (FP and salmeterol)

Leukotriene modifiers: Singulair
(MONTELUKAST), Accolate
(ZAFIRUKAST),

Zyflo (ZILEUTON)

Common treatments for the medical management of Asthma

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Anti-IL-5 mab:
Nucala
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SQ q 4 wks

Cinqair
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IV q 4 weeks over
20-50 "

Anti-IgE mab: Xolair
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Xopenex (LEVALBUTEROL)
SAMAs in the ER: Atrovent (IPRATROPIUM), Duoneb, Combivent (BOTH ALBUTEROL & IPRATROPIUM)

OCS: Prednisone, Medrol, Prelone

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Pulmicort (BUDESONIDE),
Arnuity (FF),
ArmonAir (FP)

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Nucala
(MEPOLIZUMAB)

SQ q 4 wks
Cinqair
(RESLIZUMAB)
IV q 4 weeks over
20-50 "

Anti-IgE mab: Xolair
(OMALIZUMAB)

SQ q 2-4 wks

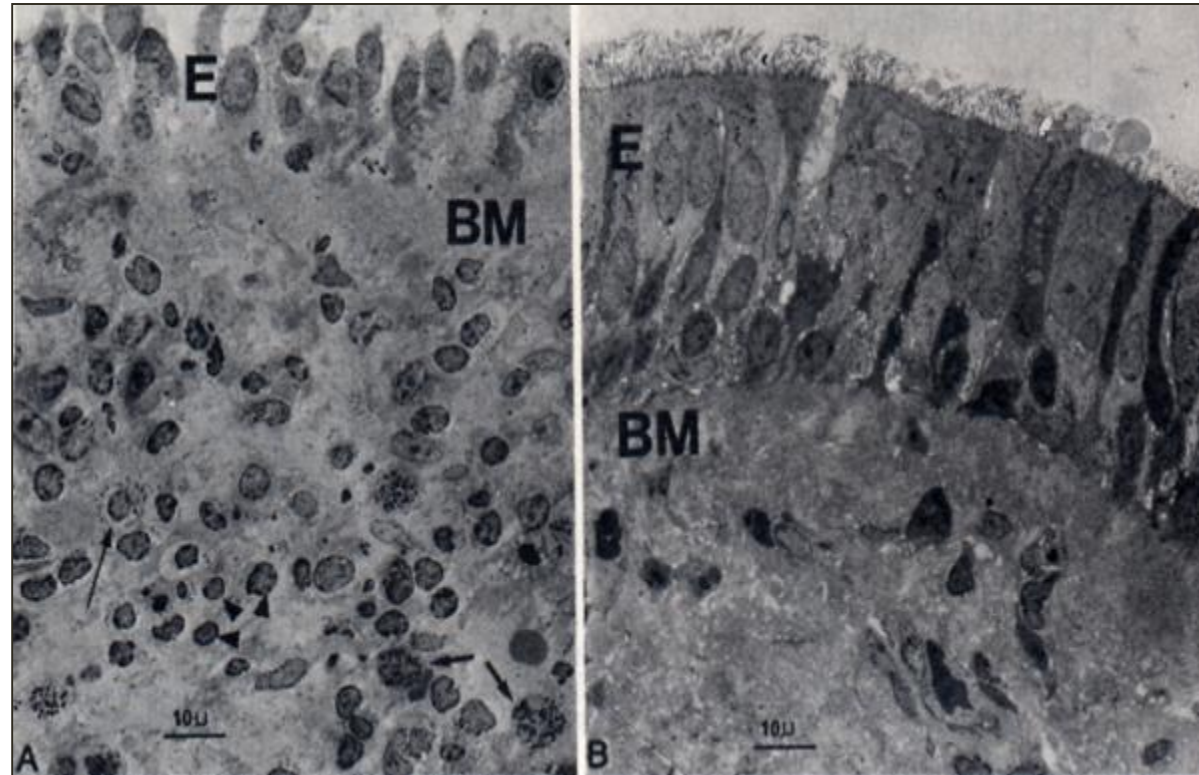
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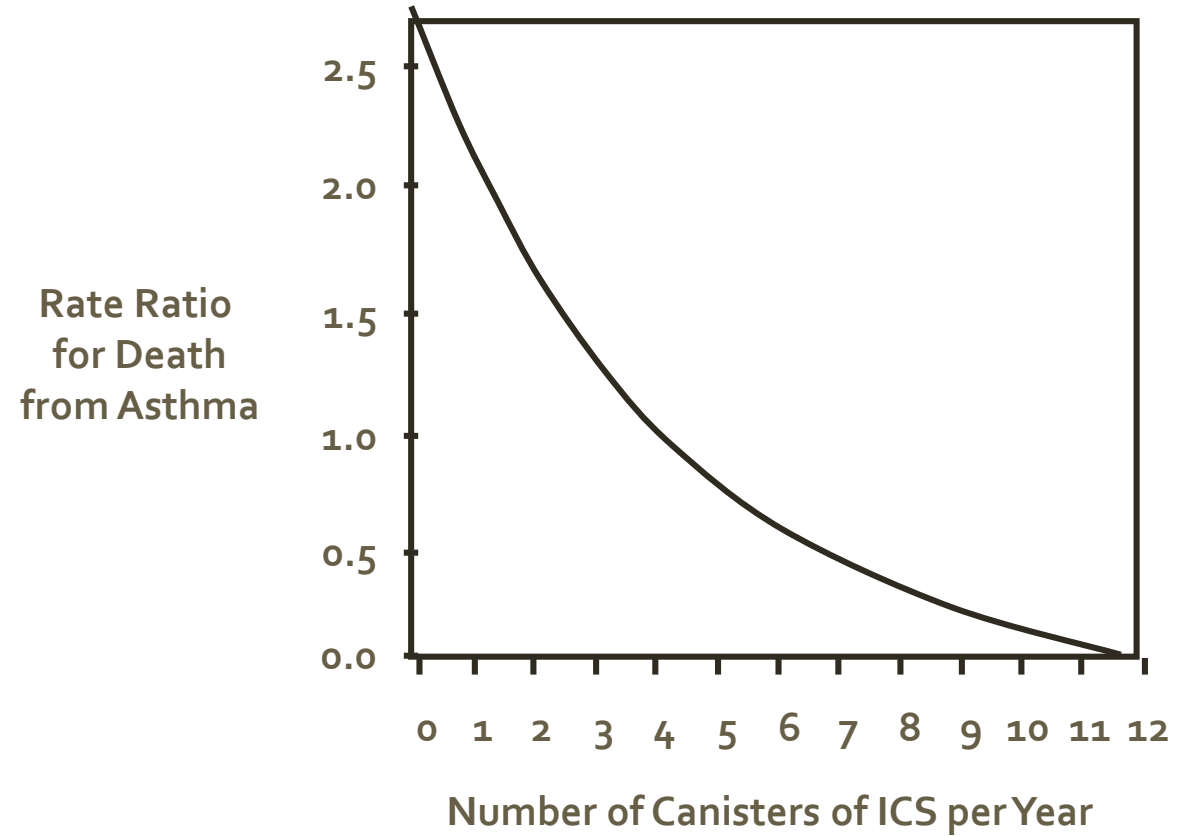
Effects of Inhaled Corticosteroids on Inflammation

E = Epithelium
BM = Basement
Membrane



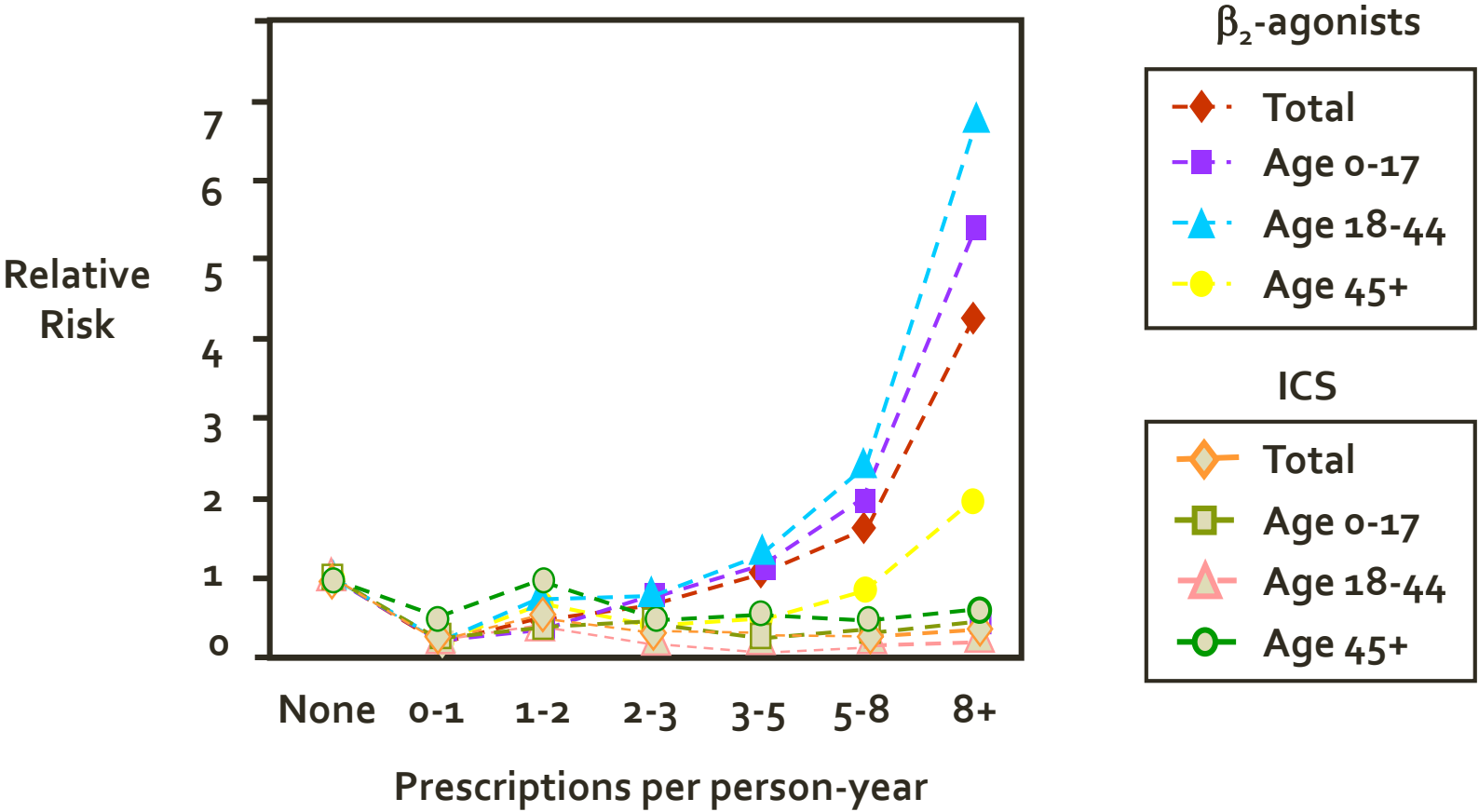
Pre- and post-3-month treatment with budesonide (BUD) 600 mcg b.i.d.

Low-dose ICS and the Prevention of Death From Asthma in Canada



Suissa et al. *N Engl J Med.* 2000;343:332-336.

Relative Risk of Hospitalization in the US



Donahue et al. *JAMA*. 1997;277:887-891.

Characteristics of fluticasone esters (propionate vs. furoate)

- Steroidal backbone (fluticasone)
- Ester substituent (furoate/propionate)
- FF confers higher affinity for both nasal and lung tissue compared with FP
 - Translates to enhanced lung residency and once-daily efficacy in asthma
 - Some evidence that the characteristics of FF may result in superior symptom reduction compared with FP

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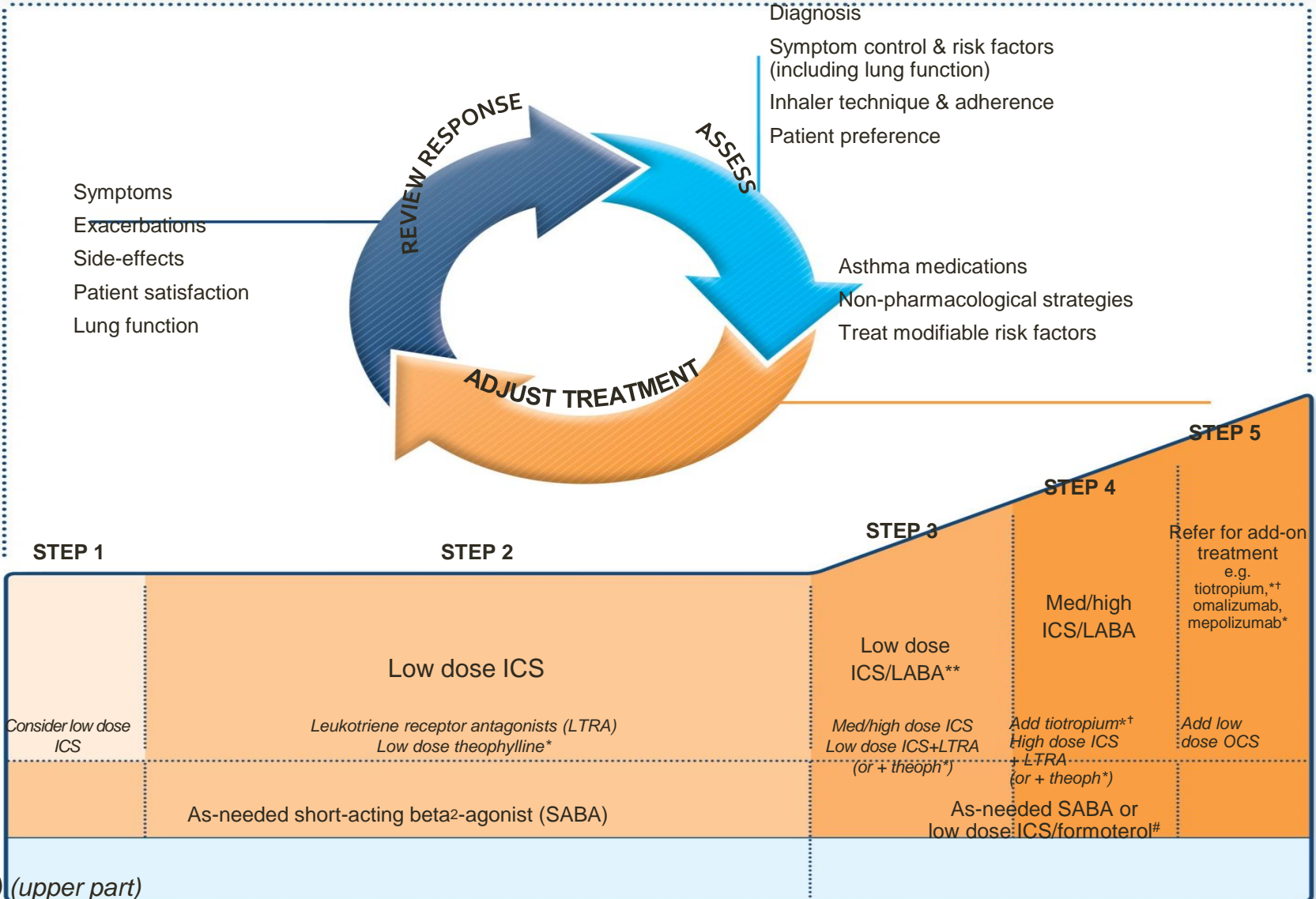
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New therapeutic
agents/interventions
for asthma from
2017 Global Initiative
for Asthma (GINA)
and 2018 EPR-4
guidelines

ICS/LAMAs

ICS/LABA/LAMA?

Stepwise management - pharmacotherapy



*Not for children <12 years

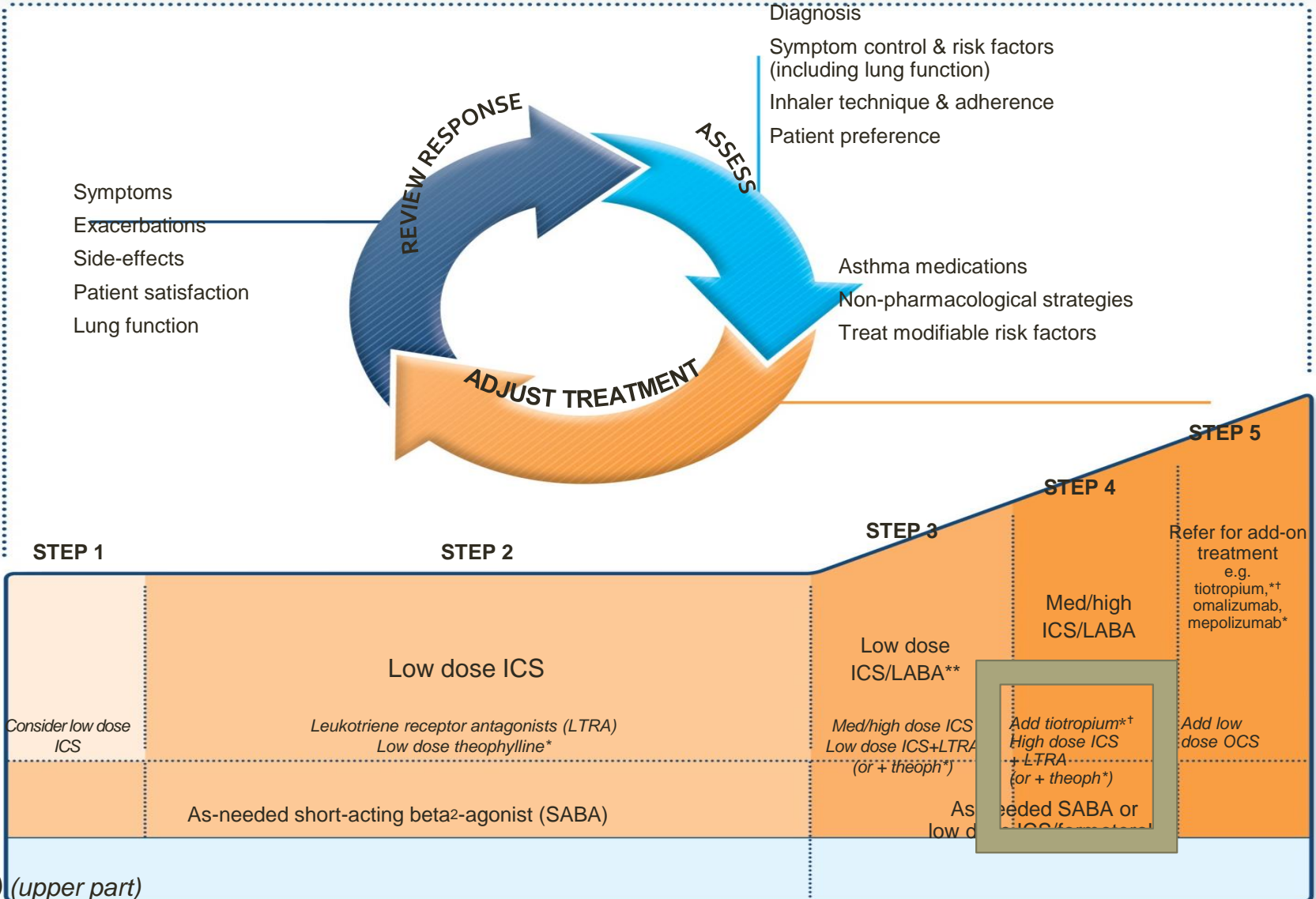
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#For patients prescribed BDP/formoterol or BUD/ formoterol maintenance and reliever therapy

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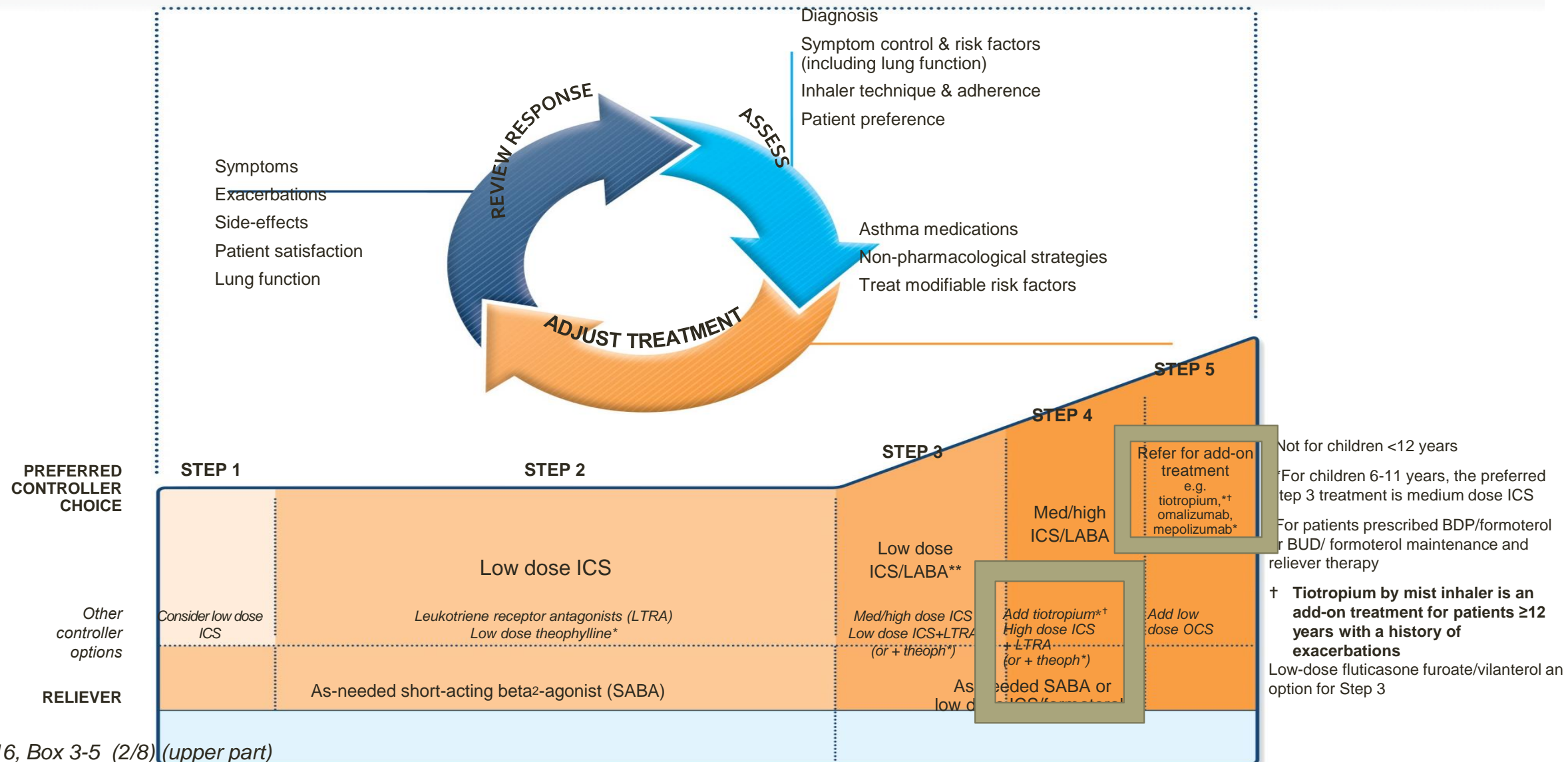
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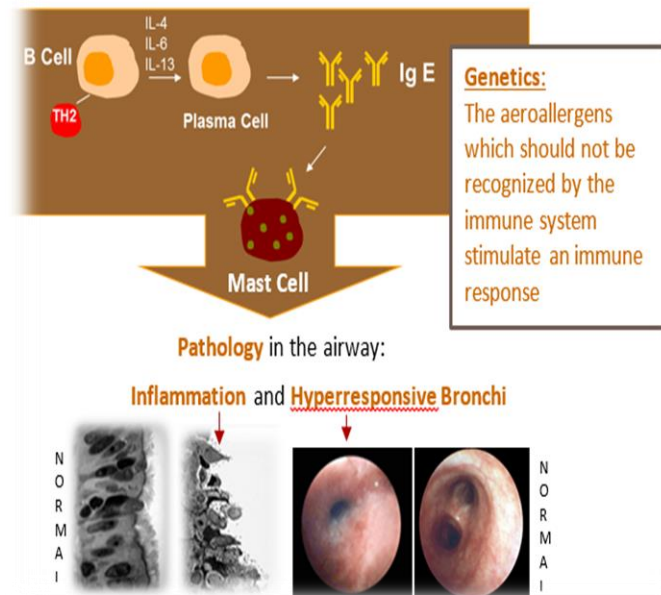
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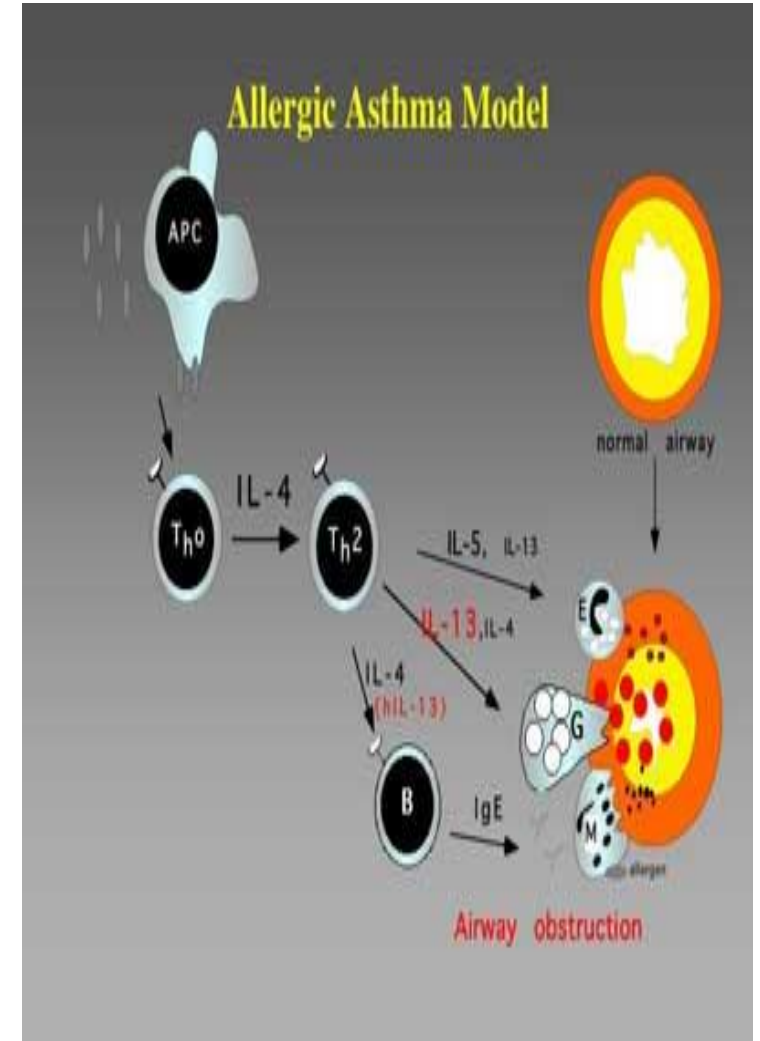
Anti-IgE biologic

- Monoclonal antibody for those
 - 6 and older
 - With severe persistent asthma
 - Which cannot be controlled on conventional therapies
- 1-3 injections every 2 -4 weeks based on IgE levels and weight
- Serum IgE rises
- Black box warning
 - Carry epipen

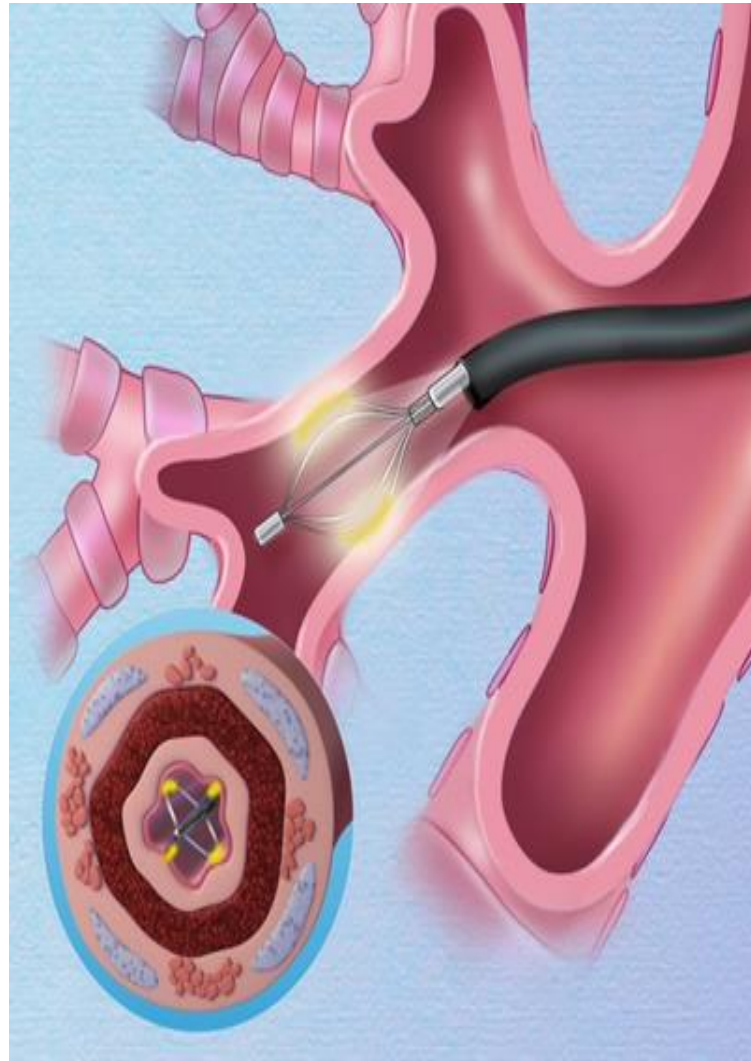


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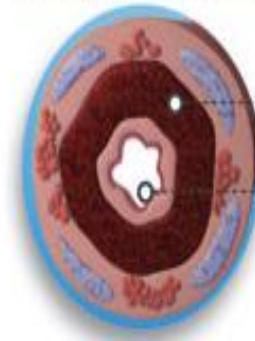
- Monoclonal antibody for those
 - Severe persistent asthma w/ an eosinophilic phenotype
 - Cannot be controlled on conventional therapies
 - Serum eosinophils decrease
- Monoclonal antibody for those
 - 18 and older with severe attacks despite treatment
 - reslizumab (Cinqair) IV every 4 weeks over 20-50 minutes
 - 12 and older
 - mepolizumab (Nucala) 1 injection every 4 weeks



Bronchial thermoplasty



Airway of Person with Severe Asthma



More airway muscle causes airway to narrow

This is the area where Alair applies heat to the airway wall during BT treatment

Airway of Person with Severe Asthma after Treatment



Reduced airway muscle after BT treatment

After BT, the inside airway wall and other tissue heals, but airway muscle is reduced

Apply radio frequency energy to airway wall to decrease smooth muscle
Treat airways > 3 mm during three separate procedures (RLL, LLL, R/LUL)

Bronchial Thermoplasty

Castro M, et al. Am J Respir Crit Care Med. 2010 (n = 288)

- Severe asthmatics randomized to either BT/Sham BT
- 79% vs. 64% improved AQOL scores (≥ 0.5) → large placebo effect
- Reduced/less severe exacerbations and ↓ ED visits & missed days of work
- 8.4% exacerbated with short hospitalization post-BT
- Excluded higher risk group
(> 3 exacerbations/URI or more than 4 steroids pulses in prior year)
- Did not follow placebo group long term
- Five follow-up: Persistent benefit year 1 vs. 5



Component 4- Education for a Partnership in Asthma Care

Pressurized Metered-Dose Inhaler Technique

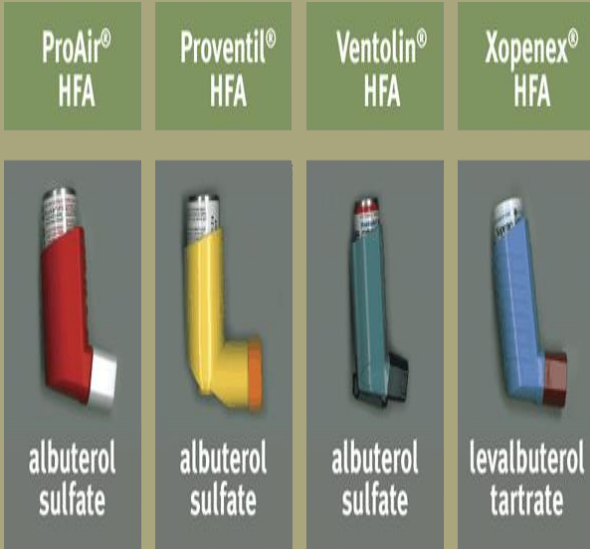


FIGURE 3-14. HOW TO USE YOUR METERED-DOSE INHALER

HOW TO USE YOUR METERED-DOSE INHALER

Using an inhaler seems simple, but most patients do not use it the right way. When you use your inhaler the wrong way, less medicine gets to your lungs.

For the next few days, read these steps aloud as you do them or ask someone to read them to you. Ask your doctor or nurse to check how well you are using your inhaler.

Use your inhaler in one of the three ways pictured below. A or B are best, but C can be used if you have trouble with A and B. Your doctor may give you other types of inhalers.

Steps for Using Your Inhaler

Getting ready

1. Take off the cap and shake the inhaler.
2. Breathe out all the way.
3. Hold your inhaler the way your doctor said (A, B, or C below).

Breathe in slowly

4. As you start breathing in slowly through your mouth, press down on the inhaler one time. (If you use a holding chamber, first press down on the inhaler. Within 5 seconds, begin to breathe in slowly.)

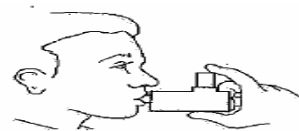
Hold your breath

5. Keep breathing in slowly, as deeply as you can.
6. Hold your breath as you count to 10 slowly, if you can.
7. For inhaled quick-relief medicine (beta₂-agonists), wait about 15–30 seconds between puffs. There is no need to wait between puffs for other medicines.

- A. Hold inhaler 1 to 2 inches in front of your mouth (about the width of two fingers).



- B. Use a spacer/holding chamber. These come in many shapes and can be useful to any patient.



- C. Put the inhaler in your mouth. Do not use for steroids.



Clean your inhaler as needed, and know when to replace your inhaler. For instructions, read the package insert or talk to your doctor, other health care provider, or pharmacist.

Dry powder inhalers

- Rapid inspiration
- Technique is device specific



Choosing an inhaler device for children ≤ 5 years



Age	Preferred device	Alternate device
0–3 years	Pressurized metered dose inhaler plus dedicated spacer with face mask	Nebulizer with face mask
4–5 years	Pressurized metered dose inhaler plus dedicated spacer with mouthpiece	Pressurized metered dose inhaler plus dedicated spacer with face mask, or nebulizer with mouthpiece or face mask

Spacers and VHCs

- Increase medication delivery to the lower airways by reducing oral deposition of particles and by enhancing hand-mouth coordination with activation
- Spacer is a generic term for any open tube placed on the MDI mouthpiece to extend its distance from the mouth
- VHCs are manufactured with a one-way valve that prevents exhalation into the device
- Activate only once into VHC/spacer
- Rinsing with diluted household detergents should prevent static electricity and enhance delivery to lungs (or use anti-static device)

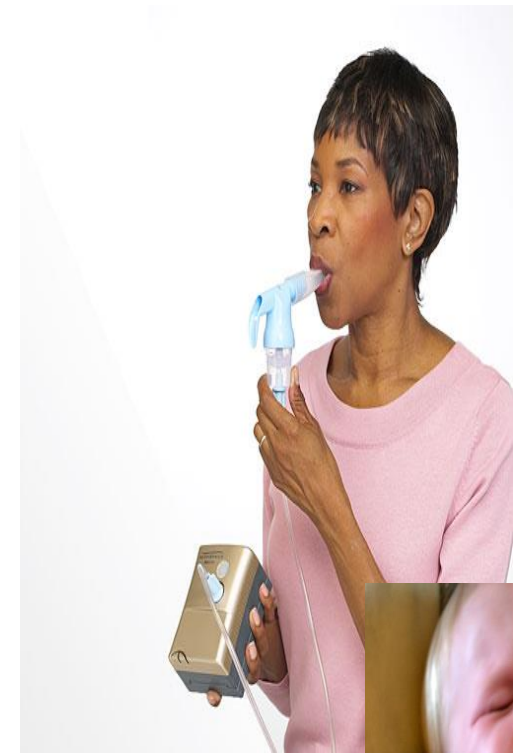


Priming, care

- Priming is necessary to deliver a correct dose into the lungs
- Priming and care is drug/device specific

Nebulizers

- Never use “blow by”
- More expensive and time-consuming than MDIs with VHCs; not any better
- If not cleaned properly, there is a risk of bacterial infections
- After each use, take apart the nebulizer and wash all parts (except tubing and finger valve) in liquid dish soap and water. Rinse with water and shake off any excess. Reattach the nebulizer pieces and tubing to the air compressor and turn on the compressor to dry the nebulizer quickly. Make sure the nebulizer is completely dry before storing







Asthma action plans

- Symptom-based is favored
- “Rules of 2”
- AAP and routine care are the most critical components for good outcomes

ASTHMA ACTION PLAN

Name: _____
Doctor: _____

Green: Feel Good
Orange: Short Wind
Red: Bad Short Wind

Feel Good 	Short Wind 	Bad Short Wind 
<ul style="list-style-type: none">• no short wind• no cough• no whistle breathing	<ul style="list-style-type: none">• tight chest• whistle breathing (wheeze)• short wind when walking or playing	<ul style="list-style-type: none">• short wind all the time• fast breathing• whistle breathing a lot• cannot talk
My medication:  Always use a spacer with your puffer if you have one	My medication:  4 puffs when needed Always carry your blue puffer with you and use it when you have short wind	Short Wind Danger Plan <ul style="list-style-type: none">• sit up• have 4 puffs of blue puffer and wait a short time• send someone to health clinic for help• if you still have bad short wind, take 4 more puffs• keep using the blue puffer until you feel better or the health worker comes
		Dr Comments: _____ _____ _____

Should asthma action plans be based peak flows or symptoms?

- Children assigned to a symptom-based asthma action plan less frequently required an acute care visit for asthma compared to those who received a peak flow based plan
- More children intended to continue using the symptom-based compared to the peak-flow based written action plan

Cochrane systematic reviews Bhogal et al 2009

Symptom monitoring using the Rules of two

Do you....

...Use your rescue inhaler more than 2 times per week?

...Wake up from asthma more than 2 times per month?

...Need Prednisone to treat asthma attacks more than 2 times a year?

If the answer is YES to ANY of ONE question your asthma may not be controlled

Asthma
adherence

Adherence

- Adherence is a process not only an outcome
- Adherence is not dichotomous
 - Chronic underuse
 - Erratic patterns of use
 - Mixed
 - Primary vs. secondary
 - Administration technique
 - Unintentional vs. intentional

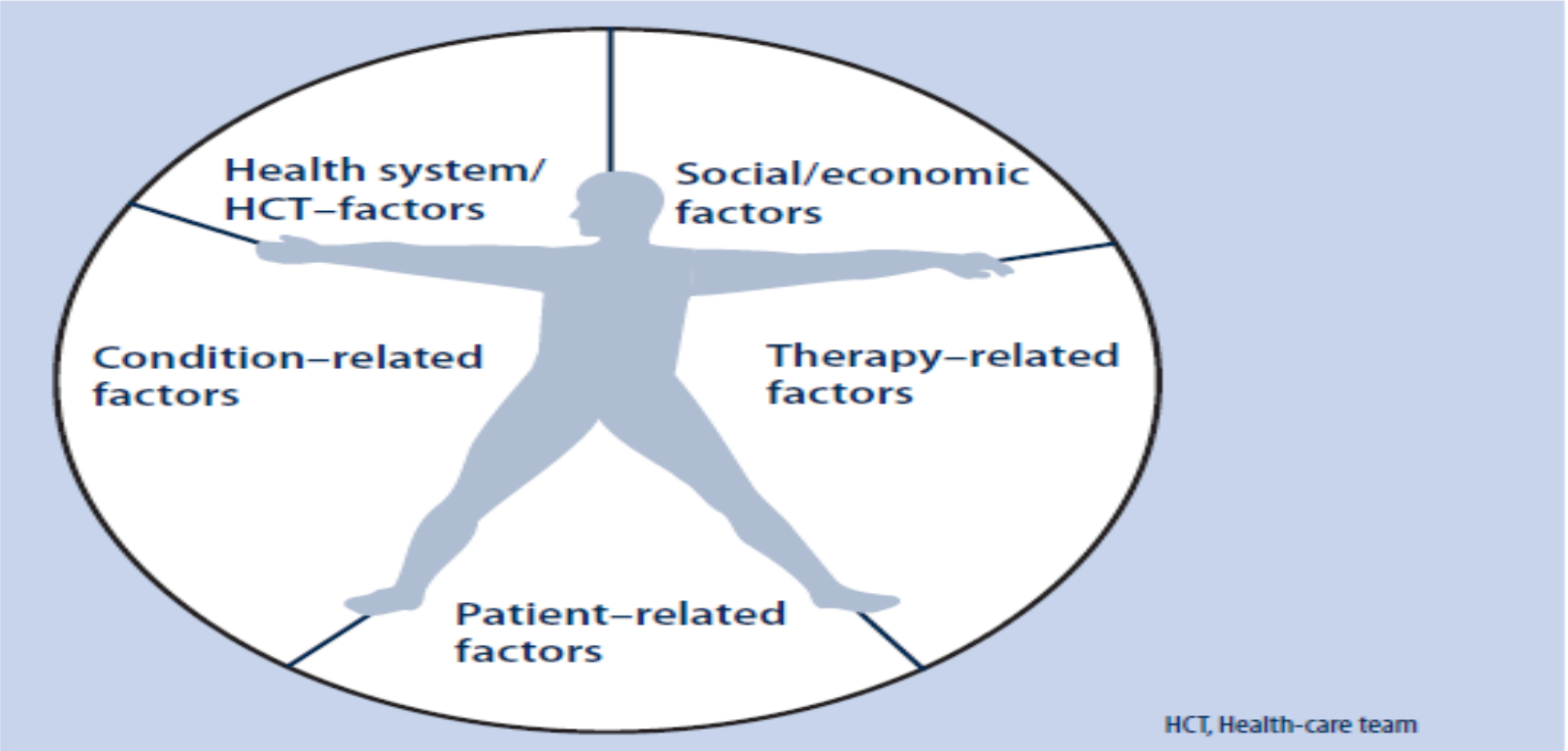
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No factors reliably predict adherence

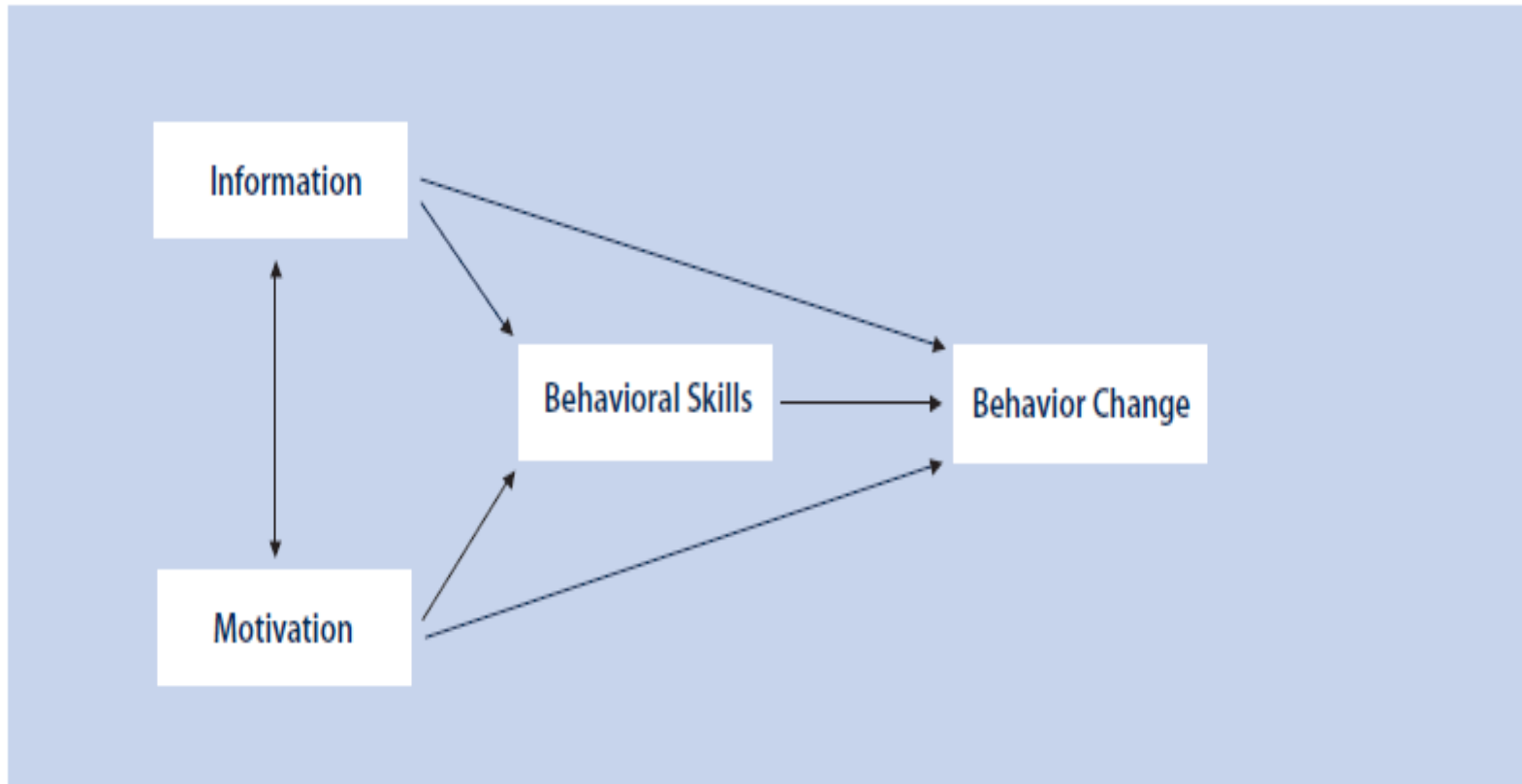
ADHERENCE TO LONG-TERM THERAPIES Evidence for action. WHO, 2003

Figure 3 The five dimensions of adherence



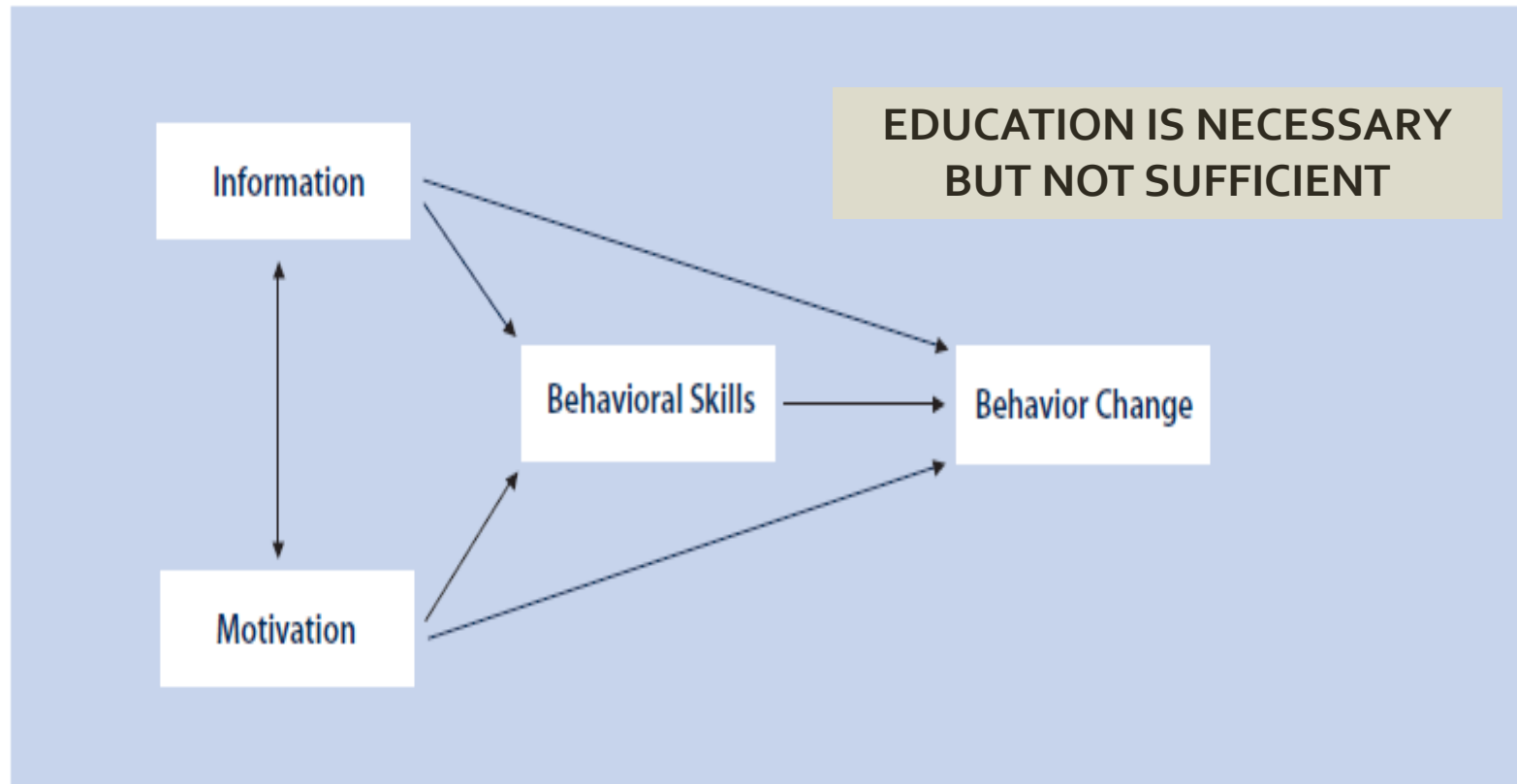
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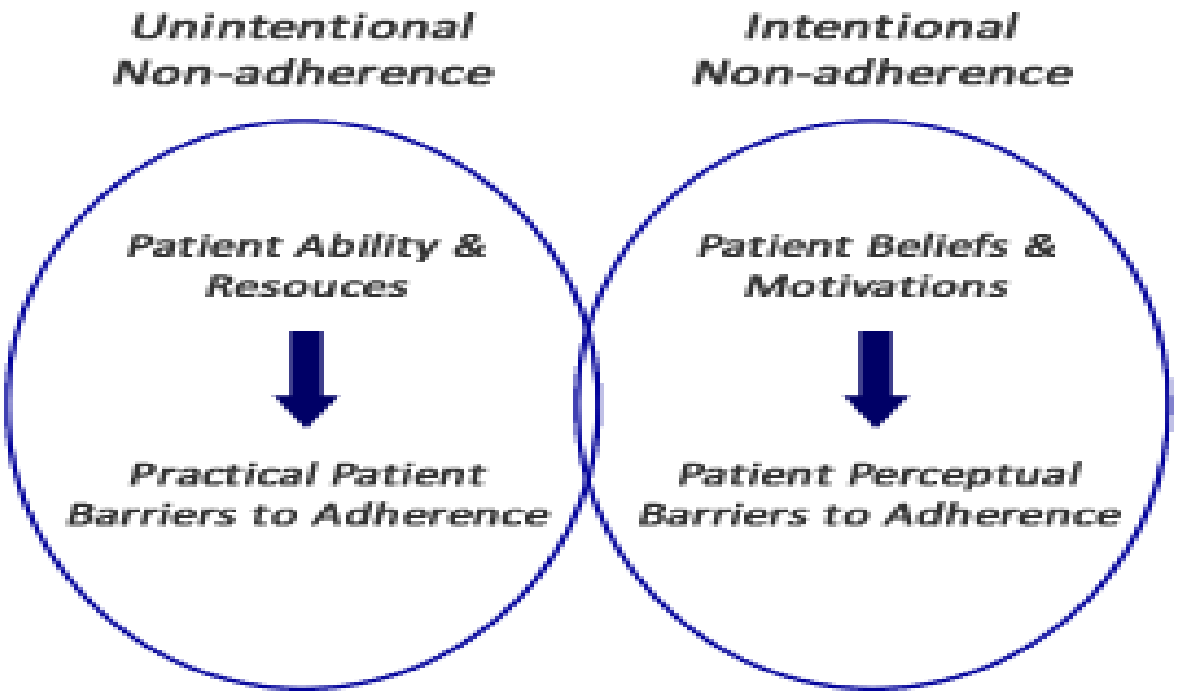


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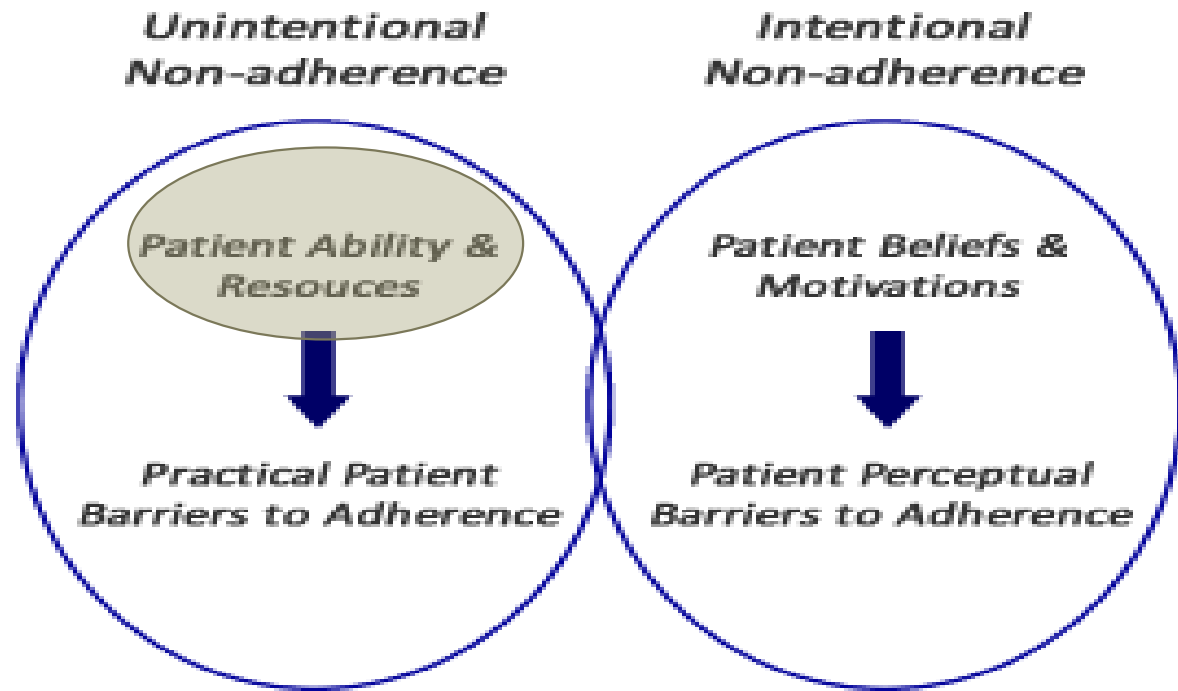


Types of non-Adherence



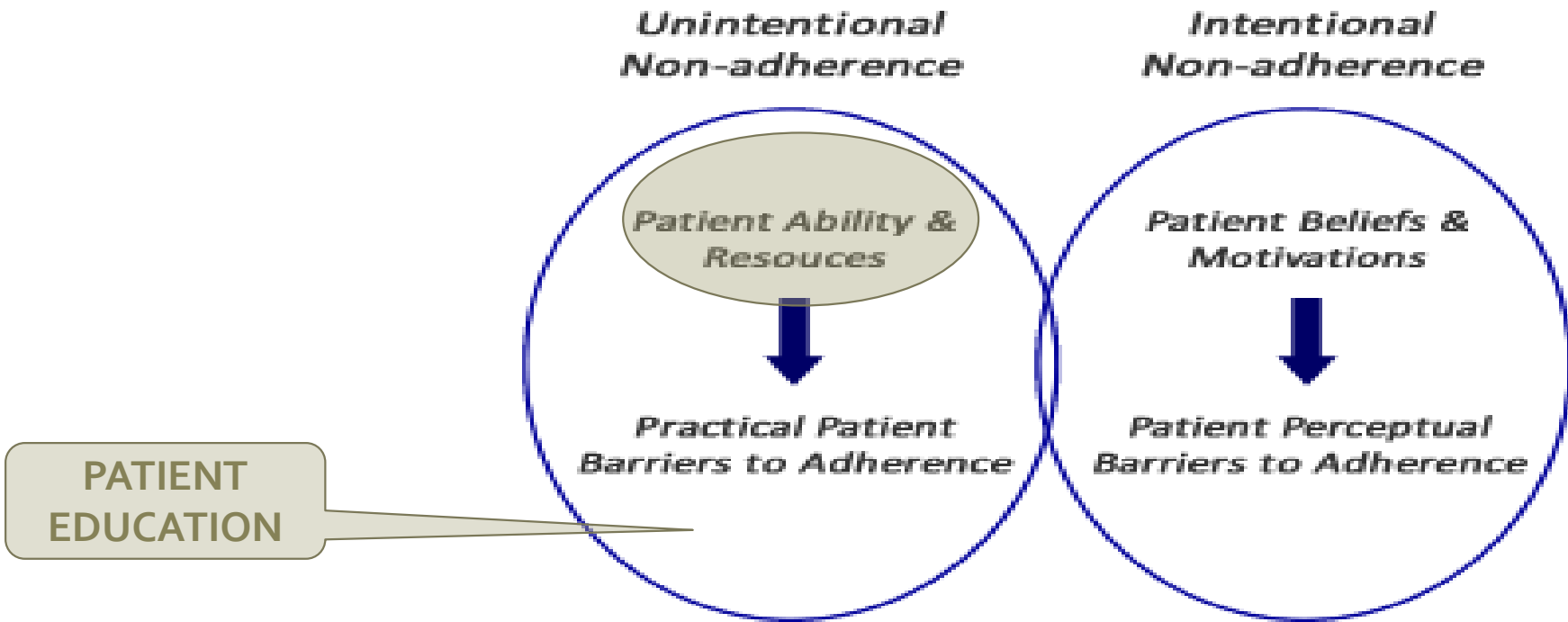
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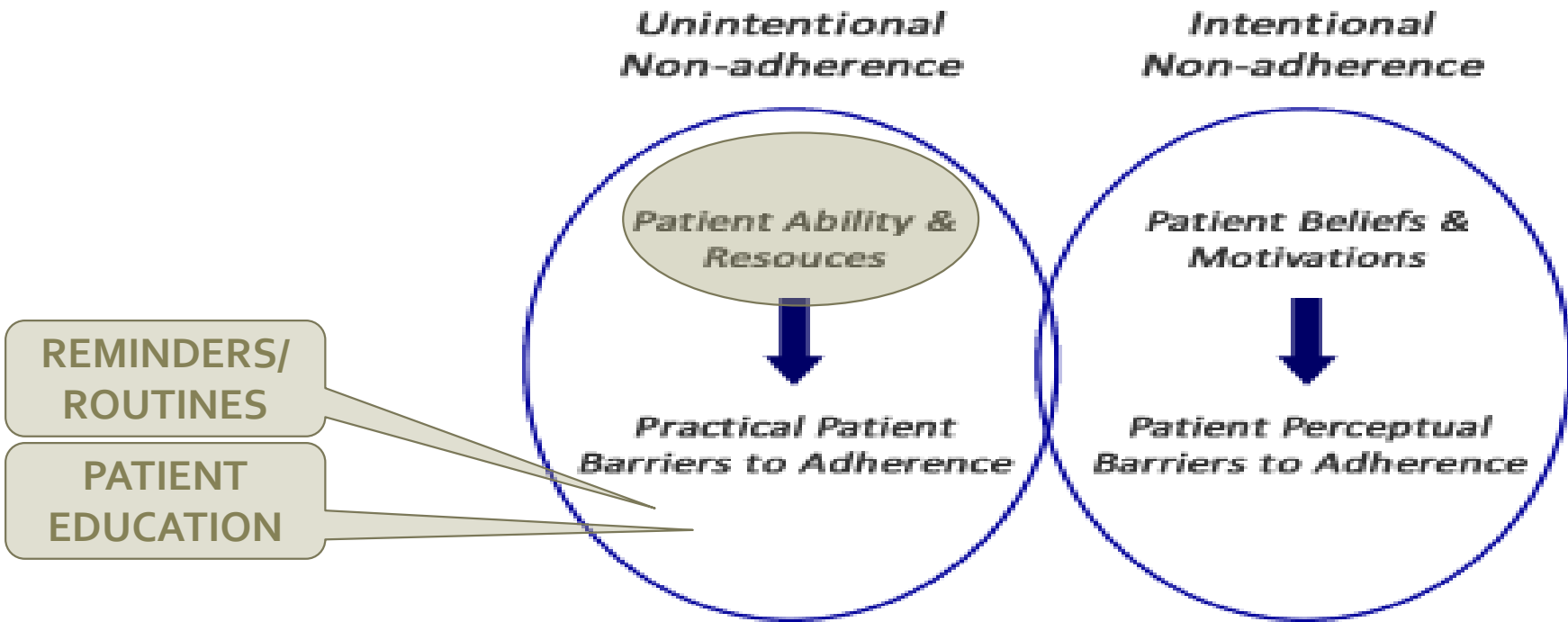
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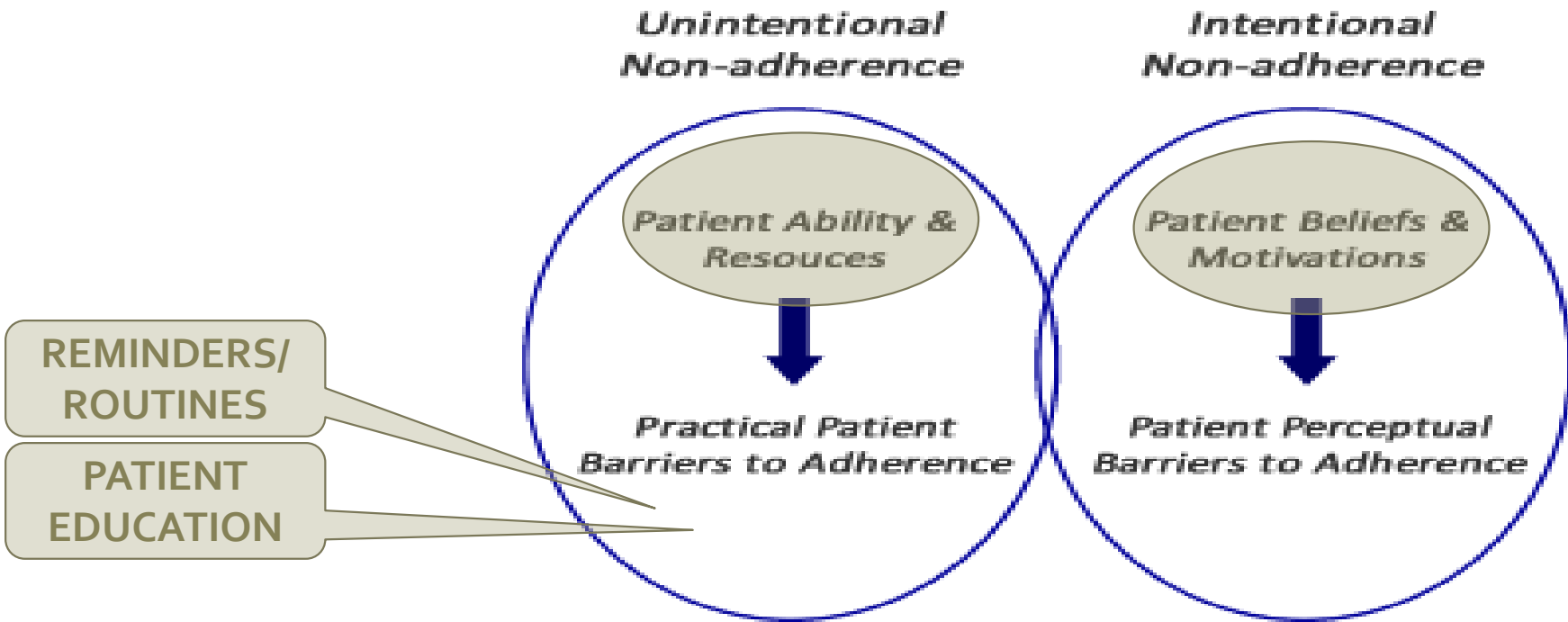
Types of non-Adherence



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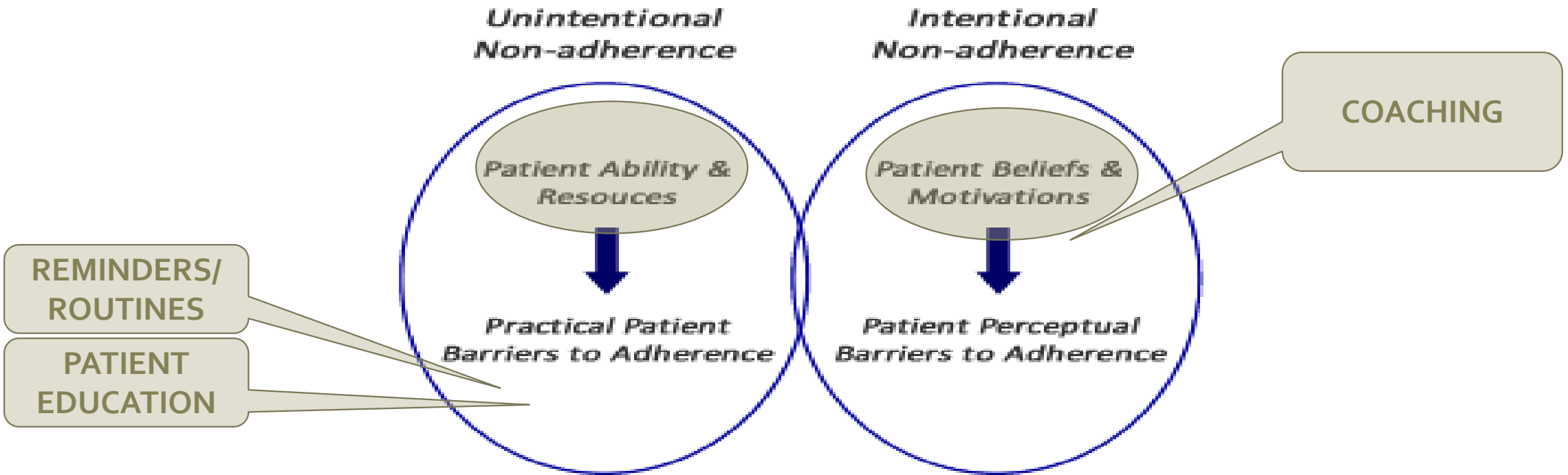
SMART INHALERS

Types of non-Adherence



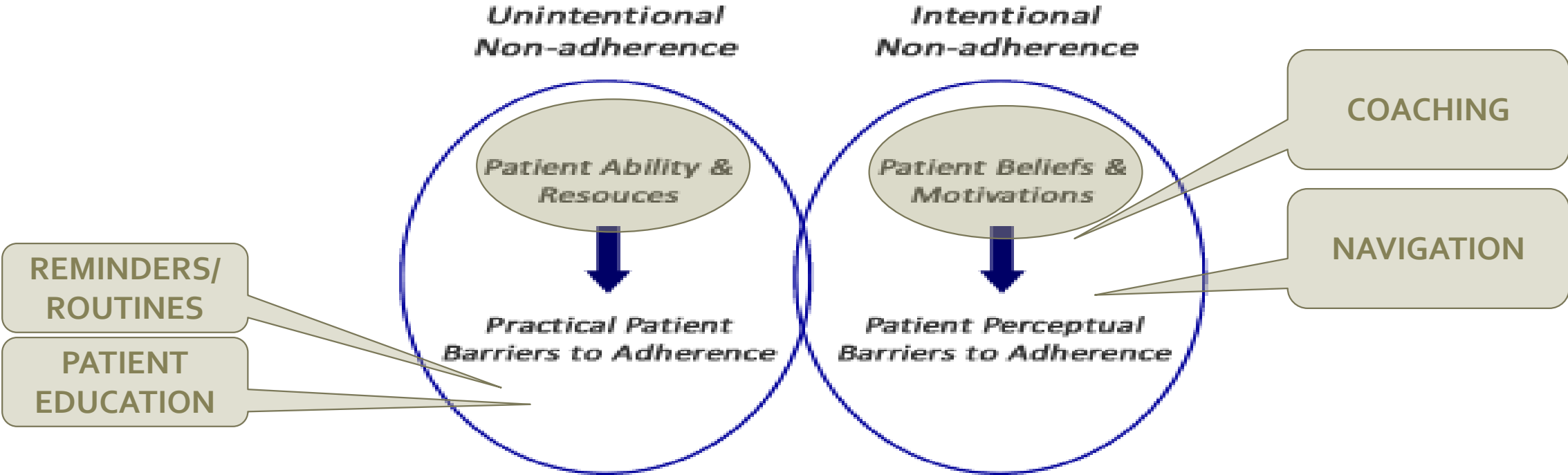
SMART INHALERS

Types of non-Adherence



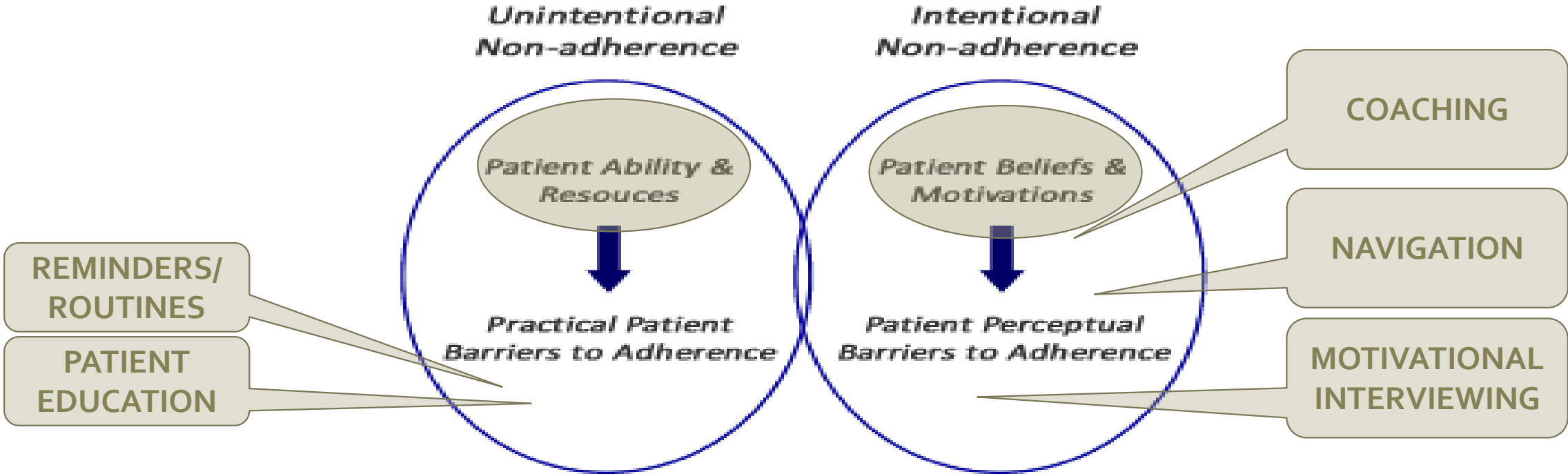
SMART INHALERS

Types of non-Adherence



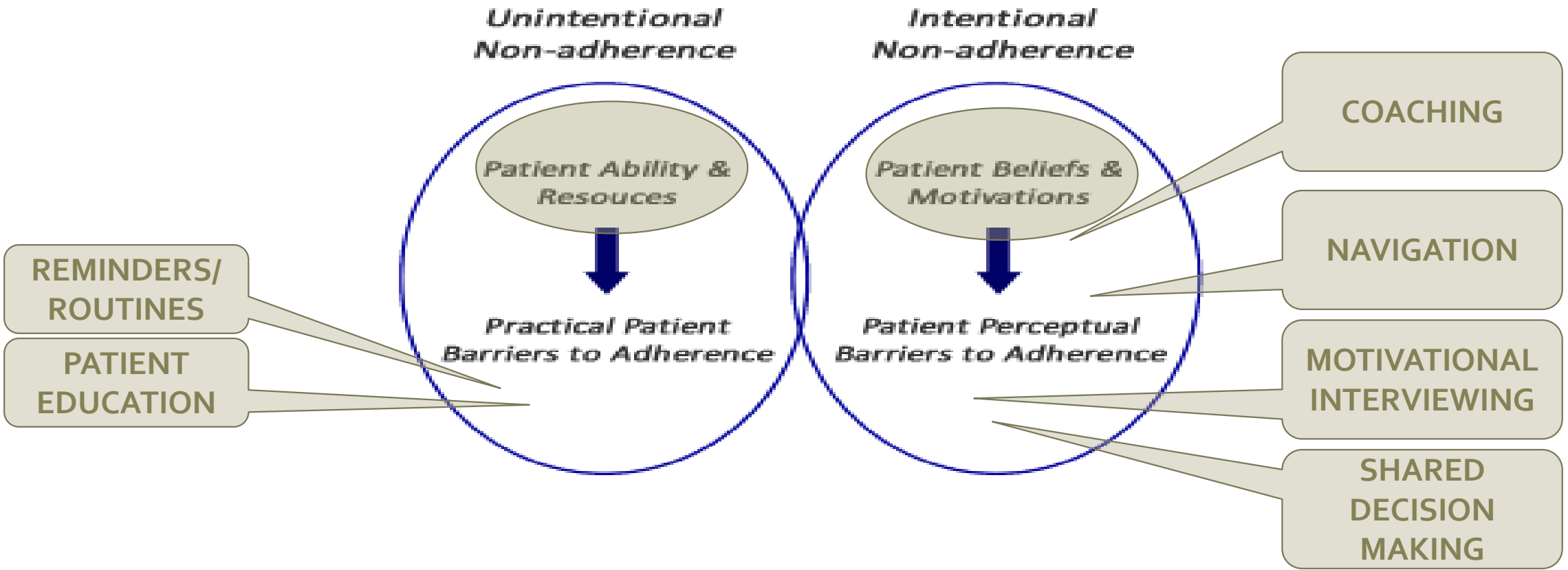
SMART INHALERS

Types of non-Adherence

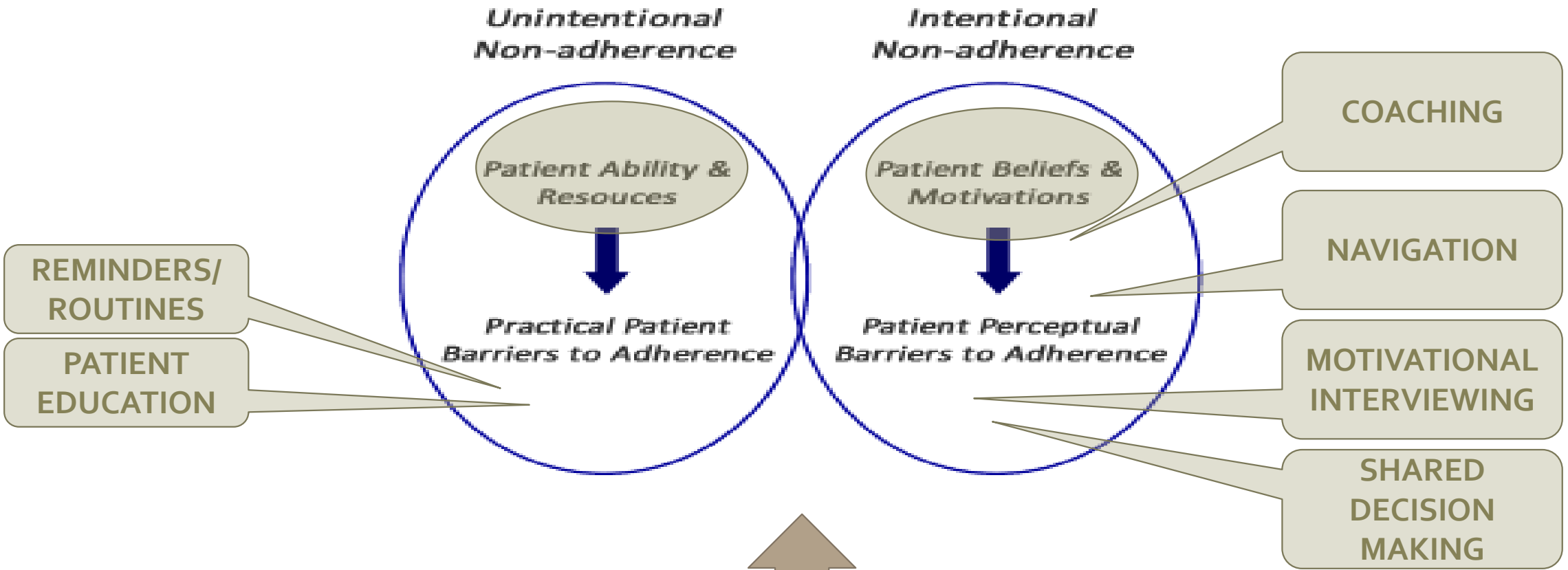


SMART INHALERS

Types of non-Adherence



Types of non-Adherence



SMART INHALERS

Asthma SELF-MANAGEMENT Adherence

Asthma SM Behavior	Adherence	Reference
ICS	47-57% 29 RCTs of 2210 pts	Normansell et al 2017 Cochrane Database of Sys Rev
SABA	N = 53; mean refill 5.1/1.3 canisters	Bollinger et al 2013 AAAI
Environmental Remediation	48%	CDC Vital Signs 2011
Appointment Keeping	29% of pediatric primary care visits (5326/7511 missed)	McGovern et al 2017 J Asthma

Looking ahead to EPR-4

EPR-4 Update in 2018

Asthma Topics

1. Role of Adjustable Medication Dosing in Recurrent Wheezing and Asthma
2. Role of Long Acting Anti-Muscarinic Agents (LAMAs) in Asthma Management as Add-on to ICSs
3. Role of Bronchial Thermoplasty in Adult Severe Asthma
4. Role of Fractional exhaled Nitric Oxide (FeNO) in Diagnosis, Medication Selection, and Monitoring Treatment Response in Asthma
5. Role of Remediation of Indoor Allergens (e.g., House Dust Mites/Animals/Pests) in Asthma Management
6. Role of Immunotherapy in Treatment of Asthma