Case study Approach to Asthma Management: Focus on Pharmacology and Guidelines

Wendy L. Wright, DNP, ANP-BC, FNP-BC FAANP, FAAN, FNAP Adult / Family Nurse Practitioner Owner - Wright & Associates Family Healthcare Amherst, New Hampshire Owner – Partners in Healthcare Education, PLLC

Wright, 2024

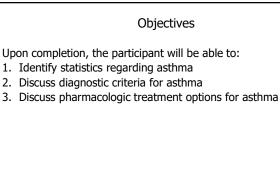
1

# Disclosures

- Speaker Bureau: - Sanofi-Pasteur, Merck, Pfizer, Seqirus, Moderna: Vaccines - AbbVie and Pfizer: Migraines
  - -AstraZeneca: Asthma and COPD
- Consultant:
  - Sanofi-Pasteur, Merck, Pfizer, Moderna, and Seqirus: Vaccines
  - Idorsia: Insomnia
  - -Shield Therapeutics: Iron Deficiency Anemia

Wright, 2024

2



Wright, 2024

# Knowledge Question One Which of the following is true about use of anti-inflammatory

?

5

therapies (corticosteroids) in patients with asthma?

- A. Systemic corticosteroids are an effective, low-risk method for treating most asthma exacerbations
- Anti-inflammatory airway effects of inhaled corticosteroids do not occur until 2-3 days of consecutive use
- C. For patients with mild asthma, SABA-only rescue therapy is preferred to ICScontaining rescue regimens due to improved outcomes and lower corticosteroid exposure
- Adding ICS to a fast-acting bronchodilator for rescue or rescue and maintenance therapy results in fewer exacerbations compared to SABA-only rescue therapy

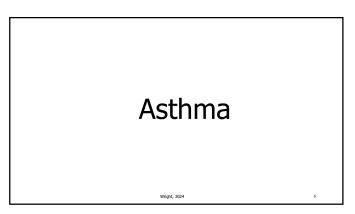
Wright, 2024

4

# ? Knowledge Question Two Which of the following is true about use of SABA in patients with asthma? A. SABA-only rescue therapy is recommended for patients with mild asthma (step 1) in the GINA 2022 Report B. Increasing SABA use is associated with a higher risk of exacerbations C. SABA + ICS rescue therapy is equally as effective for preventing and treating exacerbations as SABA-only rescue therapy D. There are currently several FDA-approved SABA + ICS combination inhalers available in the United States

Wright, 2024

5



### Asthma Definition

#### Definition of asthma:

"Asthma is a heterogenous 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." - 2022 GINA Report

GINA, The Global Initiative for Asthma

al Initiative for Asthma. Global Strategy for Asthma Management and Regentings 2022. Available from: www.ginasthma.org. Fletcher MJ, et al. NPJ Prim Gare Respi

7

#### Prevalence of Asthma

- Asthma affects approximately 339 million people worldwide 1
  - Despite significant advances in guidance and treatments, asthma continues to cause substantial health burden
- Impacts approximately 25.2 million individuals in the United States (18 and older) 2
- Most common chronic disease of childhood affecting 4.2 million children 2

1 GINA, The Global Initiative for Asthma <u>2. https://www.cdc.gov/asthma/asthmadata.htm</u> accessed 01-16-2024 Wright, 2024

8

# Impact of Asthma

er%2010%2C000%

- 9.6 million visits to providers office annually (1)
- 2 million ED visits annually (2)
   500,000 hospitalizations (2)
- 3517 deaths annually (2021)
- Highest rates: adults (5x more likely than children to die)
- Children: boys > girls
- Adults: women > men

https://www.cdc.gov/asthma/asthma\_stasia/asthma-telated-physicianvisits.html:=v-text=chr/0202015%%C/Si200aod%36252/2025%20ch%20af%20physician%20office.children%20and%20278%20p 2025220anog%250ablia.exessed 01-15-2024 2 https://www.cdc.gov/asthma/asthma/data.htm accessed 01-15-2024

Wright, 2024

## Pathophysiology of Asthma

- Likely genetic predisposition with environmental triggers
- Genetic predisposition
- Chromosome: 5Q31-Q33
- Results from repeated exposure to allergens in the individual already equipped with the genetic predisposition
- Upon exposure to an allergen, there is a release of IgE antibodies

Wright, 2024

10

• IgE antibody binds with the antigen

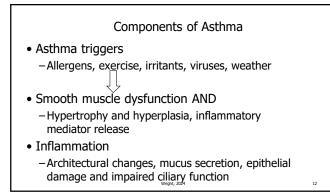
10

Pathophysiology of Asthma

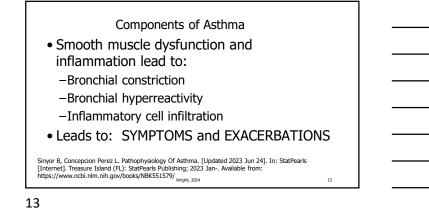
• IgE/allergen complex - then attaches itself to the mast cells on the nasal and bronchial mucosa

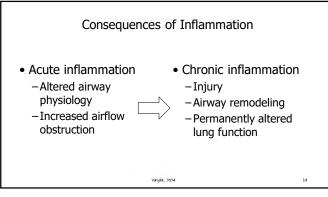
Wright, 2024

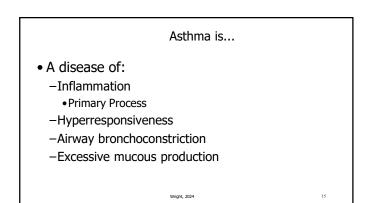
• Release of numerous chemical mediators

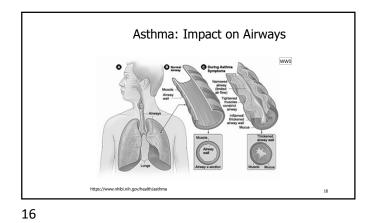




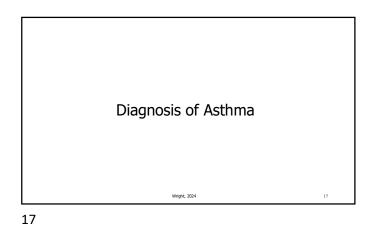








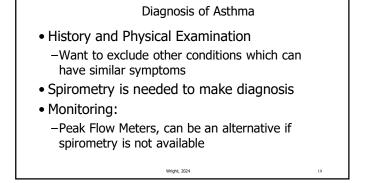




M.E.	
<ul> <li>21-year-old female <ul> <li>-C/o shortness of breath with running; present x months. Accompanied by coughing; occurs 3-5 days per week with exercise only.</li> <li>-Denies CP, audible wheezing, runny nose, dizziness</li> <li>-Has not been previously evaluated</li> <li>-Nonsmoker</li> <li>-Bronchiolitis: infancy</li> </ul> </li> </ul>	
Wright, 2024	18

# WW0 Listed as public domain

Wendy Wright, 2024-01-16T20:40:36.083

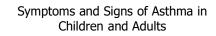


Important:

2% of individuals who present with asthma symptoms have a significant cardiorespiratory condition (other than asthma)

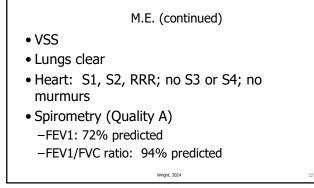
https://www.aafp.org/afp/2020/0615/p762.html accessed 04-01-2021 wight, 2024

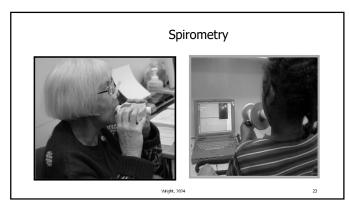
20



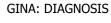
- Coughing, particularly at night or after exercise
- Wheezing
- Chest tightness
- SOB/DOE
- Cough that lingers x weeks after URI

Wright, 2024





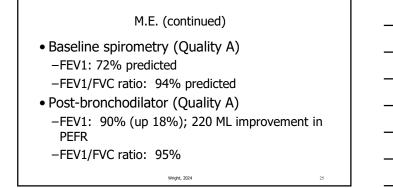
23

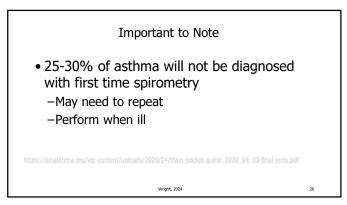


• Various ways to make diagnosis

- -At least once during the diagnostic process (when FEV1 is low), document that FEV1/FVC ratio is also low
- -Improvement of 12% or more in FEV1 and 200 mL from baseline after bronchodilator  $\underline{OR}$
- -Improvement of 12% or more in FEV1 and 200 mL from baseline after 4 weeks of anti-inflammatory  $$_{\rm WHY,\ Z24}$$

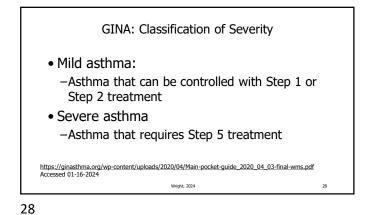
24

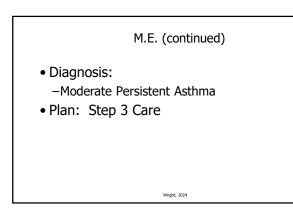


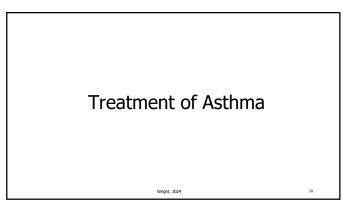


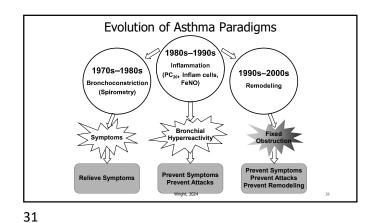
	lassificatio Youths ≥1			,		
Initia	l Diagnosis: De	etermine S	Severity and 1	reatment N	leeded	
Compon	ents of Severity			Persistent		
		Intermittent	Mild	Moderate	Severe	
	Symptoms	≤2 days/week	>2 days/week but not daily	Daily	Throughout the day	
Impairment Normal FEV;/FVC: 8-19 y 85% 20-39 y 80% 40-59 y 75% 60-80 y 70%	Nighttime awakenings	≤2x/month	3-4x/month	>1x/week but not nightly	Often 7x/week	
	(not prevention of EIB)	≤2 days/week	>2 days/week but not >1x/day	Daily	Several times per day	
	Interference with normal activity	None	Minor limitation	Some limitation	Extreme limitation	
	Lung function	Normal FEV <sub>1</sub> between exacerbations				
		FEV <sub>1</sub> >80% predicted	FEV <sub>1</sub> >80% predicted	FEV <sub>1</sub> >60% but <80% predicted	FEV <sub>1</sub> <60% predicted	
		5	FEV <sub>1</sub> /FVC normal	FEV <sub>1</sub> /FVC reduced 5%	FEV <sub>1</sub> /FVC reduced >5%	
Risk	Exacerbations requiring	0-1/year (see note) Consider severity and interval since last exacerbation. Frequency and severity may				
NOA	continenteroide	fluctuate over time for patients in any severity category.				
	controduction dis	Relative annual ri	sk of exacerbations may			
				Step 3	Step 4	
Step 1 Step 2 Recommended Step for initiating Treatment		and consider short course of oral systemic corticosteroids				
ikecommended step		In 2 to 6 weeks, e accordingly,	evaluate level of asthma			
	Wright, 2024					



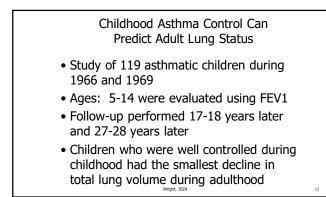




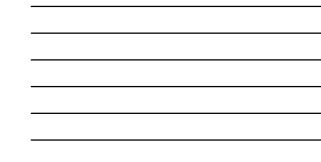


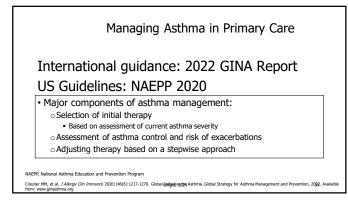






	e 10-14: Cl (Youths≥1:	2 Years	of Age a	nd Adult	s)
Initia	I Diagnosis: De	etermine S	Severity and 1	Freatment N	leeded
Compor	ents of Severity			Persistent	
		Intermittent	Mild	Moderate	Severe
	Symptoms	≤2 days/week	>2 days/week but not daily	Daily	Throughout the da
Impairment Normal FEV <sub>1</sub> /FVC: 8-19 y 85% 20-39 y 80% 40-59 y 75% 60-80 y 70%	Nighttime awakenings	≤2x/month	3-4x/month	>1x/week but not nightly	Often 7x/week
	Short-acting beta <sub>2</sub> -agonist use for symptom control (not prevention of EIB)	≤2 days/week	>2 days/week but not >1x/day	Daily	Several times per day
	Interference with normal activity	None	Minor limitation	Some limitation	Extreme limitation
		Normal FEV <sub>1</sub> between exacerbations			
	Lung function	FEV <sub>1</sub> >80% predicted	FEV <sub>1</sub> >80% predicted	FEV <sub>1</sub> >60% but <80% predicted	FEV <sub>1</sub> < 60% predicted
		FEV <sub>1</sub> /FVC normal	FEV <sub>1</sub> /FVC normal	FEV <sub>1</sub> /FVC reduced 5%	FEV <sub>3</sub> /FVC reduced >5%
	Exacerbations requiring	0-1/year (see note) ≥2/year (see note) Consider severity and interval since last exacerbation. Frequency and severity may			
Risk oral systemic corticosteroids		fluctuate over time for patients in any severity category. Relative annual risk of exacerbations may be related to FEV.			
				Step 3	Step 4
Recommended Ste	ecommended Step for initiating Treatment		Step 2	and consider short of systemic conticoster	hids
		In 2 to 6 weeks, accordingly, Wright,	evaluate level of asthma	control that is achieve	d and adjust therapy



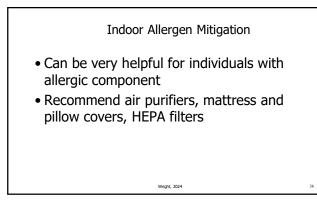


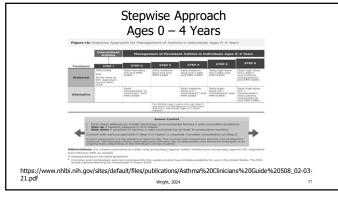


https://www.nhlbi.nih.gov/healthtopics/all-publications-andresources/2020-focused-updates-asthmamanagement-guidelines

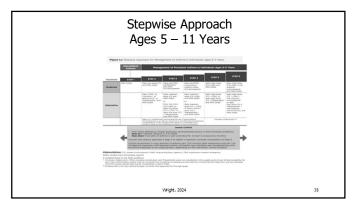
Wright, 2024

35

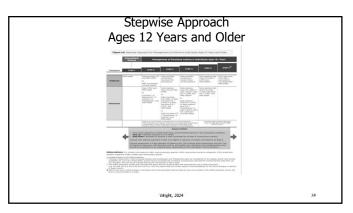




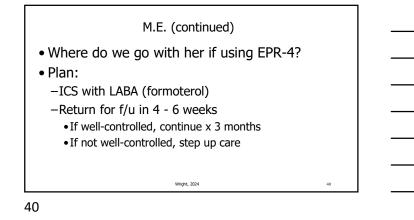














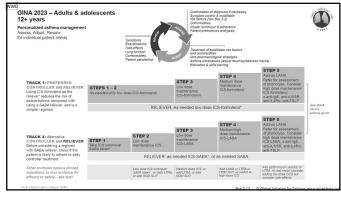




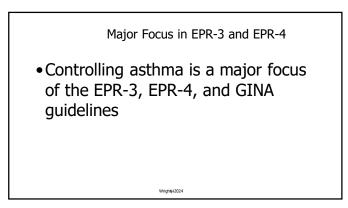
- GINA no longer recommends SABA agent as monotherapy for adolescents and adults with asthma
- All adolescents and adults with asthma should receive an ICS containing product
  - -Symptom driven: ICS/SABA OR ICS/LAMA as needed OR

-Daily: ICS/LABA https://ginastima.org/wp-content/uploads/2020/04/Main-pocket-guide\_2020\_04\_03-final-wms.pdf accessed 01-16-2024









44



ion, 2032. Availai

- Determining asthma control is essential for optimizing therapy and achieving treatment goals
- Validated asthma assessment tools acknowledged in GINA and NAEPP

   GINA and NAEPP each also have a separate set of questions to
   assess control

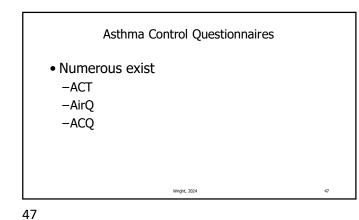
GINA: symptom control should be assessed "at every opportunity"
 NAEPP: periodic assessments at 1–6-month intervals and "ongoing

Cloutier MM, et al. J Allergy Clin Immunol. 2020;146(6):1217-1270. Global Juli 1990 Asthma. Global Strategy for Ast from: www.ginasthma.org

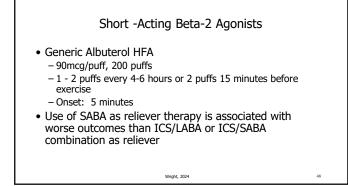
# WW0 I have requested permission to use

Wendy Wright, 2024-01-16T20:58:12.059

(Yo	Assessir uths ≥12 Y	ng Asthma Years of A		
<u> </u>			<u> </u>	atment Needed
Component		Well-controlled	Not Well- controlled	Very Poorly Controlled
	Symptoms	≤2 days/week	>2 days/week	Throughout the day
	Nighttime awakenings	≤2 x/month	1-3x/week	≥4x/week
	Interference with normal activity	None	Some limitation	Extremely limited
Impairment	Short-acting beta <sub>2</sub> - agonist use for symptom control (not prevention of EIB)	≤2 days/week	>2 days/week	Several times per day
	FEV <sub>1</sub> or peak flow	>80% predicted/personal best	60-80% predicted/personal best	<60% predicted/personal best
	Validated Questionnaires ATAQ ACQ ACT	0 ≤0.75* ≥20	1-2 ≥1.5 16-19	3-4 N/A ≤15
	Exacerbations	0-1/year Consider severity and it	≥2/year (see note) d interval since last exacerbation	
	Progressive loss of lung function	Evaluation requires long-term follow-up care		
Risk	Treatment-related adverse effects		upl of intensity door not	m none to very troublesome t correlate to specific levels o assessment of risk.







Short-Acting Beta-2 Agonists

- Usage of these medications more than 2 times/week is indicative of poor control
- 1 inhaler = 200 inhalations
- It is time to begin transitioning patients away from SABA alone

Wright, 2024

49

#### Albuterol and Budesonide Inhaler<sup>7</sup>

- Name: Albuterol and budesonide (Airsupra™)
- Class: SABA/ICS
- Indication: As needed treatment for acute bronchospasm or prevention of bronchospasm and to reduce the risk of acute asthma exacerbations in adults 18 years of age and older
- Dosage: 180 mcg of albuterol and 160 mcg of budesonide
- Dosed: 2 puffs every 4 hours as needed
- Do not exceed 6 doses (12 puffs) in 24 hours.

50

Albuterol and Budesonide In	haler <sup>7</sup> (continued)
<ul> <li>Adverse effects</li> <li>Oral candidiasis: Should instruct to rinse mouth out.</li> <li>Caution: DPI – Acute paradoxical bronchospasm</li> </ul>	<ul> <li>Why a combination?</li> <li>GINA and EPR4 recommend use of ICS whenever SABA is needed to prevent/reduce exacerbations and the need for systemic corticosteroids.</li> <li>Provides ICS/SABA in one inhaler thus decreasing copays</li> </ul>

51

# Albuterol and Budesonide Inhaler<sup>7</sup> (continued)

- Drug interactions
  Budesonide
  - CYP450 3A4 substrate
  - Avoid strong 3A4 inhibitors as they may increase systemic
- exposure of budesonide. • Not currently approved for children

Available December 2023
Canister will have a dosing counter on it to enable patient to see how many doses remain.

52

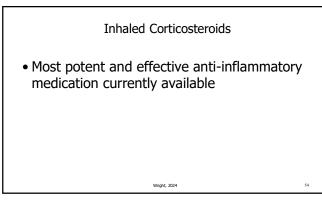
53

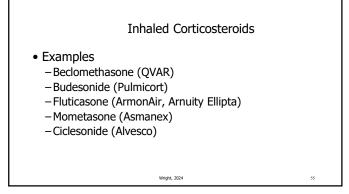
52

**Controller Medications** 

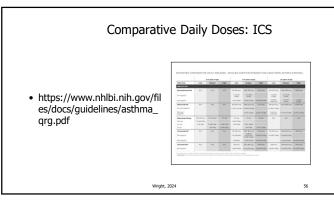
Wright, 2024

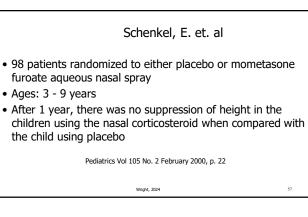
53

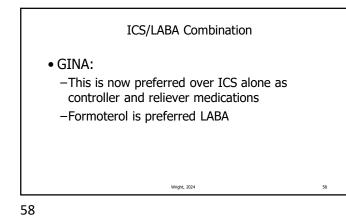


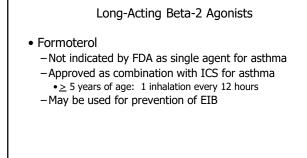








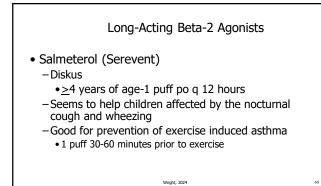


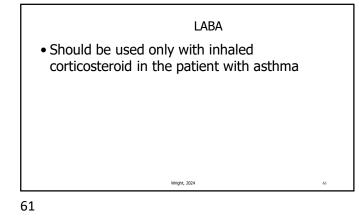


Wright, 2024

59

59



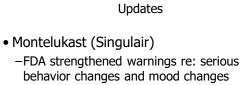


**Combination Products** 

Wright, 2024

- Fluticasone/salmeterol (Advair, AirDuo, Wixela Inhub)
- Budesonide/formoterol (Symbicort)
- Mometasone/formoterol (Dulera)
- Fluticasone/vilanterol (Breo Ellipta)

62



 Per FDA – benefits of treatment may NOT outweigh risks, especially in mild disease or for allergic rhinitis

Wright, 2024

#### Leukotriene Receptor Antagonists

- Cysteinyl leukotriene production in the body has been associated with airway edema, smooth muscle constriction and the inflammatory process
- These medications block the leukotriene receptors which in turn is able to prevent inflammation and bronchoconstriction
- Has been removed from all of the major guidelines due to neuropsychiatric issues

Wright, 2024

64

#### Montelukast

- (Montelukast) Singulair
  - -4 mg Granules once daily: 12 23 months
  - -4 mg tablet for children 2 5 years of age

Wright, 2024

- -5mg qhs for ages 6-14
- -10mg qhs for ages 15 and older

65

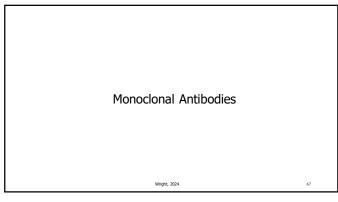
#### Montelukast

• Drug Interactions

- Metabolized through CYP2A6 (minor pathway)
- Phenobarbital: decreases montelukast but no dosage adjustment is required
- Side effects: headache, fatigue, dizziness, Churg-Strauss

Wright, 2024

- Precautions
   Not for an acute exacerbation
- Category: B
- Category. D



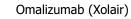
#### Omalizumab (Xolair)

• Indicated for adults and adolescents (6 and older) with moderate to severe persistent asthma who have a positive skin test or *in vitro* reactivity to a perennial aeroallergen

Wright, 2024

- And...whose symptoms are inadequately controlled with inhaled corticosteroids
- SC injection (weight and IGE based)
- Every 2 4 weeks
- Warning: anaphylaxis

68



- Recombinant DNA-derived humanized IgG1 monoclonal antibody that selectively binds to human immunoglobulin E (IgE).
  - IgE antagonist
- Inhibits the binding of IgE to the high-affinity IgE receptor on the surface of mast cells and basophils
- Limits the degree of release of mediators of the allergic response.





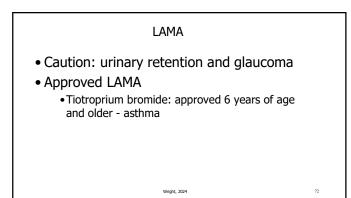
-Long-acting bronchodilator

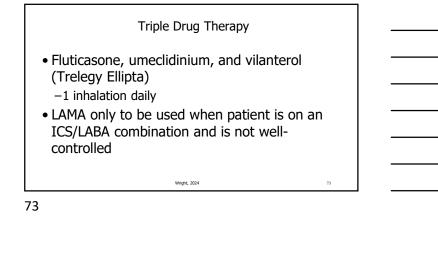
• LAMA

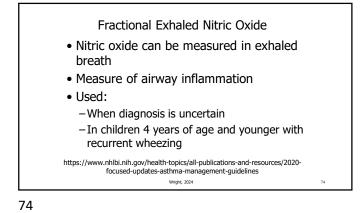
- -Increasing/emerging role in the management of asthma
- -Controller medication
- -LAMA are only added to patient with poorly controlled asthma after LABA/ICS is in place

Wright, 2024

71



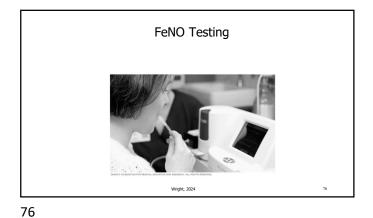




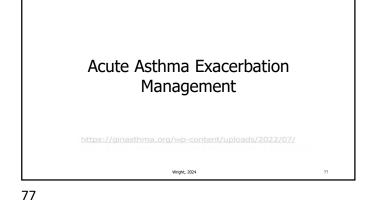


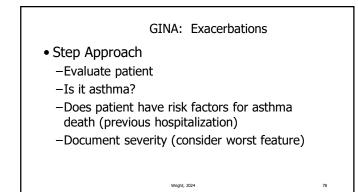
- FeNO > 50 ppb (or > 35 ppb in children ages 5 12 years) are consistent with elevated T2 (Type 2) inflammation and support diagnosis of asthma
- Allergic rhinitis can increase FeNO levels as well; interpret cautiously

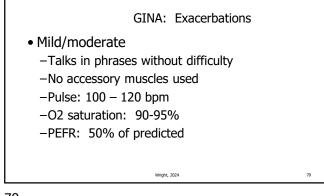
https://www.nhlbi.nih.gov/health-topics/all-publications-and-resources/2020focused-updates-ast<u>hma-management-guidelines</u>



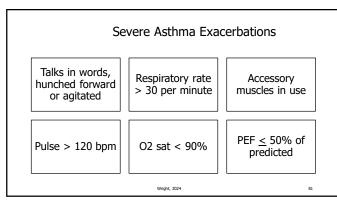








GINA: Exacerbations
Mild/moderate treatment
<ul> <li>–SABA: 4-10 puffs and repeat every 20 minutes for 1 hour</li> </ul>
<ul> <li>Prednisolone: adults: 40-50 mg; children 1-2 mg/kg with max of 40 mg</li> </ul>
<ul> <li>Assess response in 1 hour and if improving: send home with prednisolone prescription and reliever</li> </ul>
Wider 204 80
111git, 221



## GINA: Exacerbations

- Severe
  - -SABA: 4-10 puffs and repeat every 20 minutes for 1 hour
  - -Prednisolone: adults: 40-50 mg; children 1-2 mg/kg with max of 40 mg

Wright, 2024

82

83

?

- -Transfer to an acute care facility
- -Place on oxygen during transfer

82

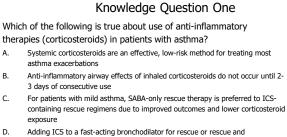
#### Acute Asthma Exacerbation

- Measure FEV1
- Use albuterol: see recommendations (next slide)
- Prednisone, prednisolone, or similar
- What dose and schedule ??
- 5-7 days is recommended
- No taper needed as long as treating for < 14 days

Wright, 2024

- Oral SCS are preferred over IV/IM

83



84

Α.

В.

C.

## Knowledge Question Two

?

85

Which of the following is true about use of SABA in patients with asthma?

- A. SABA-only rescue therapy is recommended for patients with mild asthma (step 1) in the GINA 2022 Report
- B. Increasing SABA use is associated with a higher risk of exacerbationsC. SABA + ICS rescue therapy is equally as effective for preventing and
- treating exacerbations as SABA-only rescue therapy D. There are currently several FDA-approved SABA + ICS combination inhalers available in the United States

Wright, 2024

85

