Pediatric Asthma 2011

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Objectives

Upon completion, the participant will be able to:

1. Identify statistics related to incidence/prevalence of asthma

2. Discuss the signs and symptoms of asthma

3. Discuss treatment options for asthma

Asthma

Asthma is...

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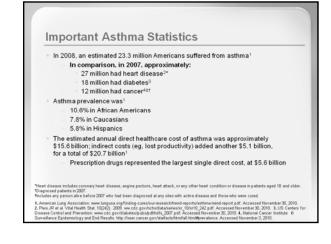
- Derived from the Greek word for panting or breathlessness
- Recurrent airflow obstruction caused by chronic airway inflammation with a superimposed bronchospasm
- Leads to... wheezing, breathlessness and a cough

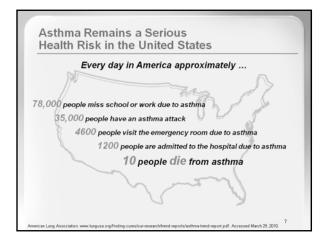
Guidelines for the Diagnosis and Management of Asthma—Update on Selected Topics 2002. NIH, NHLBI. June 2002. NIH publication pp. 02-5075

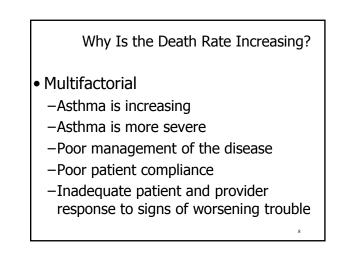
Prevalence of Asthma

- Impacts approximately 21 million individuals in the United States
- Most common chronic disease of childhood affecting 6 million children
- Before adolescence, 2 times more common in boys
- Increasing incidence of this disease
 76% increase in the prevalence of asthma within the past decade

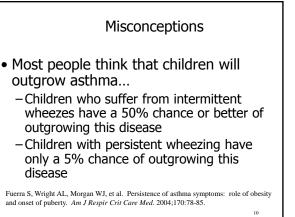
Guidelines for the Diagnosis and Management of Asthma—Update on Selected Topics 2002. NIH, NHLBI. June 2002. NIH publication no. 02-5075.





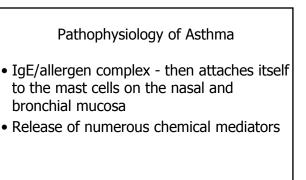




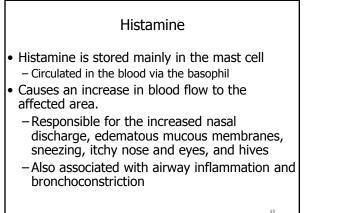


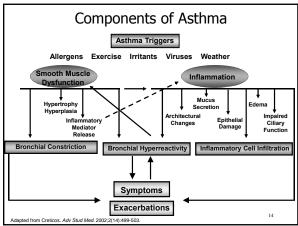
Pathophysiology of Asthma

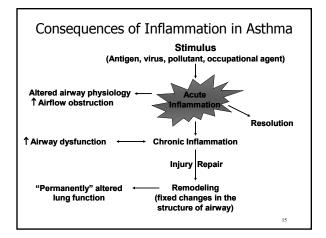
- Genetic predisposition - Chromosome: 5Q31-Q33
- Results from repeated exposure to allergens in the individual already equipped with the genetic predisposition
- Upon exposure to an allergen, there is a release of IgE antibodies
- IgE antibody binds with the antigen

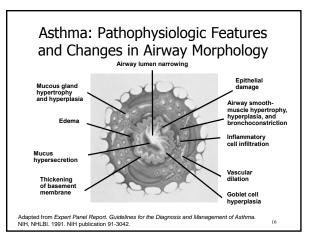


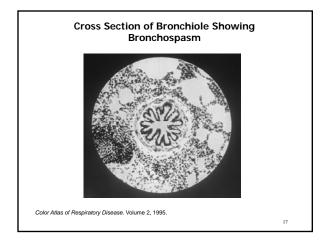
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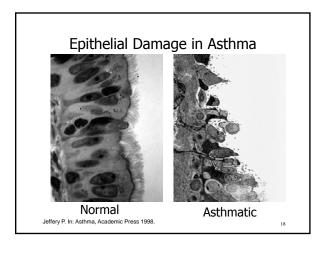


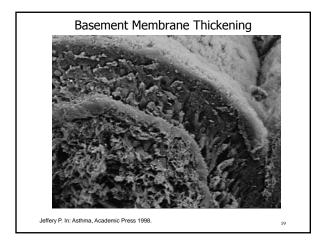


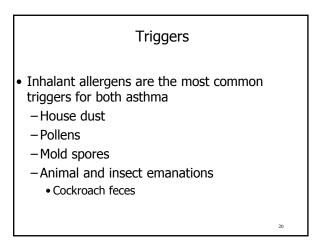


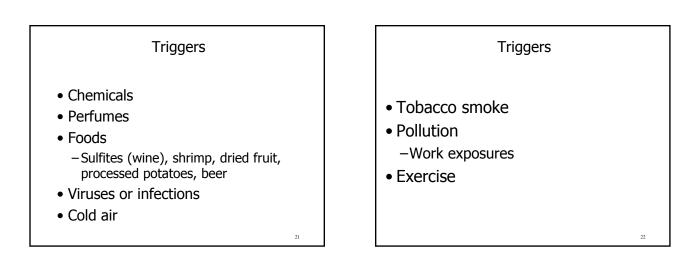






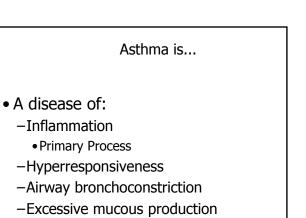


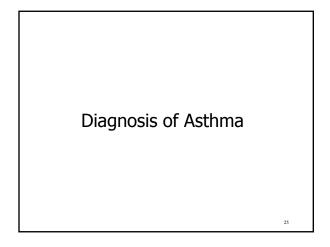


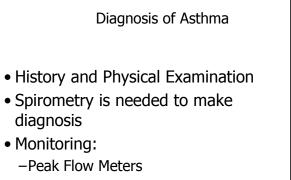


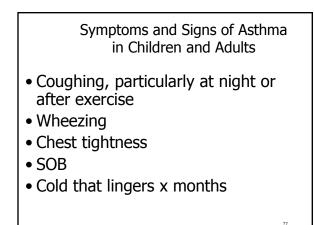
Gastroesophageal Reflux - A Significant Factor in Children

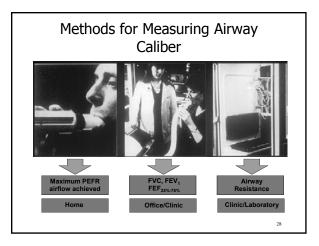
- 84 healthy infants and children referred for an evaluation of daily wheezing
 - -All evaluated for reflux
 - -64% had positive evaluations for reflux
 - After 3 months on anti-reflux treatment,
 64.8% of the infants/children were able to discontinue all daily asthma medications (including nebulized flunisolide)
 - Sheikh S. et. Al. Pediatric Pulmonology. 1999 Sep; 28(3): 181-6

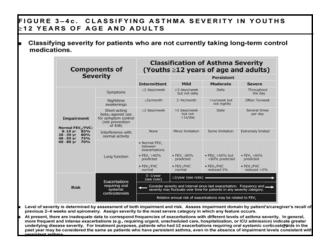


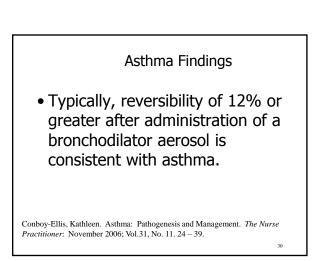


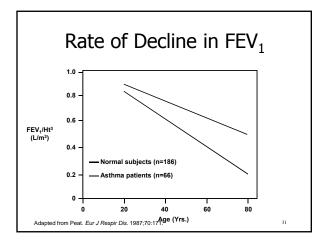


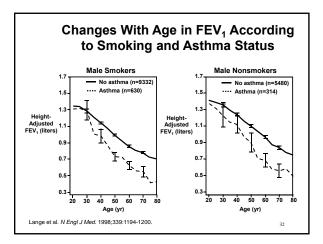




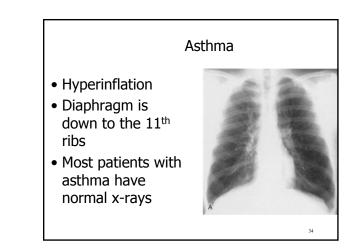








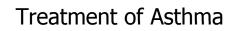
The Biggest Predictor of Sudden Death from Asthma
History of hospitalization with or without intubation
These individuals are at a significant isk for a serious exacerbation again

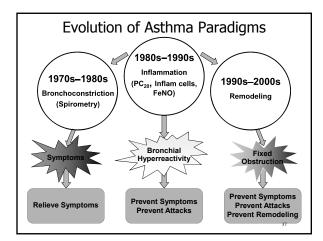


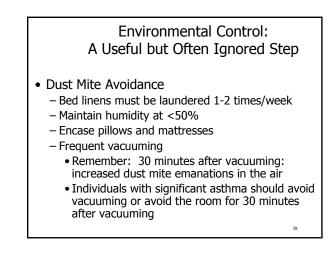
Chronic Asthma Changes

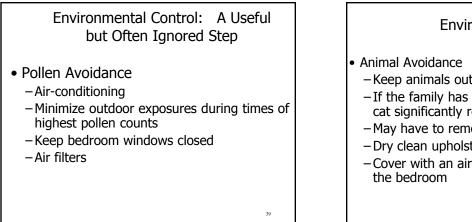
- Increased AP Lateral diameter
- The way you know that AP/Lat diameter is increased is by this clear space between the sternum and the ascending aorta
- Flat diaphragms













- -Keep animals out of the bedroom
- If the family has a cat, weekly washing of the cat significantly reduces the allergen load
- May have to remove animals from home
- Dry clean upholstery and carpets
- -Cover with an air filter any ducts leading into

Environmental Control

- Mold Avoidance
 - -Children/adolescents with allergic rhinitis and/or asthma should not be sleeping in a damp basement
 - -Clean moldy surfaces
 - -Avoid houseplants
 - -Avoid chores that involve damp grass, leaves

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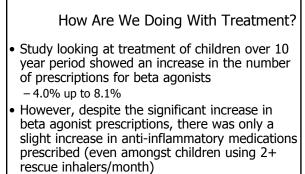
Environmental Control

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- Avoidance of Non-allergic Triggers
 - -Strong emotions
 - -Smoke: No smoking in house or car
 - -Pollution
 - -Cold air
 - -Odors
 - -Exercise

Childhood Asthma Control Can Predict Adult Lung Status

- Study of 119 asthmatic children during 1966 and 1969
- Ages: 5-14 were evaluated using FEV1
- Follow-up performed 17-18 years later and 27-28 years later
- Children who were well controlled during childhood had the smallest decline in total lung volume during adulthood



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- 0.4% up to 2.4% Goodman, DC et. Al. Pediatrics 1999 Aug; 104(2) 187-94

 Stepwise Approach for Managing Asthma in Patients Aged ≥12 Years: NAEPP EPR-3 Guidelines

 Swere Persistent

 Moderate Persistent

 Step 1 Preforred: SABA pm

 Note Persistent

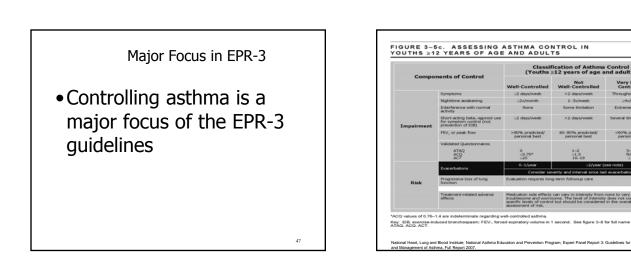
 Step 1 Preforred: SABA pm

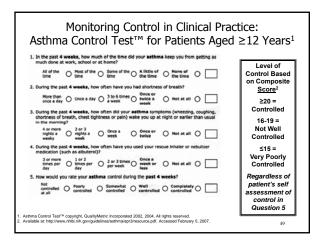
 Note Persistent

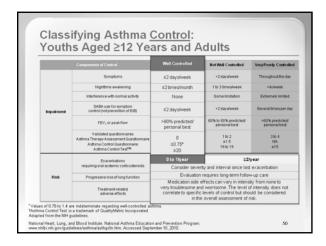
 Step 4 Preforred: Number of the state of the

Step Approach to Therapy
If control is not achieved with therapy, step up the therapy

- Once control is sustained for a minimum of 3 months, can consider stepping down the therapy
- Regardless, therapy should be reviewed q 6 months







Short Acting Inhaled Beta 2 Agonists • Albuterol (Proventil HFA, Ventolin HFA, Pro-Air HFA) – 2 puffs q 4-6 hours or 2 puffs 15 minutes before exercise – Onset: 5 minutes



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 Levalbuterol (Xopenex HFA)
 -1 - 2 inhalations every 4 - 6 hours prn

Short-Acting Beta-2 Agonists

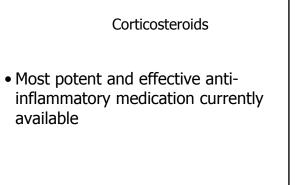
- Usage of these medications more than 2 times/week is indicative of poor control
- Regular, scheduled use of these medications is usually not recommended

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Long-Acting Controller Medications Maintenance or Prevention is the Key

- Good management is the key to preventing exacerbations and hospitalizations
- As with any disease, preventing the problem is always better than treating it



Inhaled Corticosteroids

- Examples
 - -Beclomethasone (Beclovent, Vanceril)
 - -Budesonide (Pulmicort turbuhaler)
 - -Flunisolide (Aerobid)** No longer produced
 - -Fluticasone (Flovent)
 - Triamcinolone Acetonide (Azmacort)*** No longer produced

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- Mometasone (Asmanex)



- Side effects
 - Pharyngitis
 - Dysphonia
 - -Oral Candidiasis
- Precautions
 - High dosages: Increased systemic absorption leading to HPA axis suppression
 - -Not indicated for an acute exacerbation

To Reduce Side Effects of Inhaled Corticosteroids

- Administer with spacers or holding chambers
- Rinse mouth after inhalation
- Use lowest possible dose to maintain control
- Children monitor growth

Schenkel, E. et. al

- 98 patients randomized to either placebo or mometasone furoate aqueous nasal spray
- Ages: 3 9 years
- After 1 year, there was no suppression of height in the children using the nasal corticosteroid when compared with the child using placebo

Pediatrics Vol 105 No. 2 February 2000, p. 22

Remember...

- Poorly controlled asthma often delays growth
- In general, children with asthma tend to have longer periods of reduced growth rates prior to puberty

Mast Cell Stabilizers

- Cromolyn Sodium (Intal)
- No longer being manufactured if not already unavailable, soon to be

Mast Cell Stabilizers

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- Mechanism of Action
 - Reduces the production of histamine and prevents the release from the mast cell
- MDI or Nebulizer Solution
 - -MDI: > 5 years: 2 puffs po qid
 - -Nebulizer Solution: >2 years: 1 ampule qid
 - Begin to work within 15 minutes of inhalation but can take up to 2 weeks to become effective

Mast Cell Stabilizer

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- Side effects
 - -Generally well tolerated
 - Side effects occur in 1:10,000
 - -Cough
 - -Wheezing
 - Rash
 - Nausea
- Category B

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

Leukotriene Receptor Antagonists

- (Zafirlukast) Accolate
 - 10mg bid for ages 5-11
 - 20mg bid for 12 and older
 - Studied in children as young as 5
 - Avoid food 1 hour before and 2 hours after taking: Food decreases the bioavailability of Accolate
 - Metabolism: Metabolized through the CY P450 2C9 and 3A4 pathways
 - Major pathways in the body
 - Numerous other medications use this same pathway

Zafirlukast (Accolate)

- Drug/Drug Interactions
 - Aspirin: Increased zafirlukast levels by 40%
 - Erythromycin: 40% decrease in zafirlukast
 - Theophylline: Postmarketing reports of increased theophylline levels

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-Coumadin: 35% increase in PT/INR

Zafirlukast (Accolate)

- Side effects
 - -Headache (12.9%)
 - Dizziness
 - Nausea
 - Churg Strauss syndrome
- Pregnancy: B
- Precautions
 - -Not for an acute exacerbation

Montelukast (Singulair) • (Montelukast) Singulair –4 mg Granules once daily: 12 – 23 months –4 mg tablet for children 2 - 5 years of age –5mg qhs for ages 6-14 –10mg qhs for ages 15 and older

Montelukast (Singulair)

- Drug Interactions
 - Metabolized through CYP2A6 (minor pathway)
 - Phenobarbital: decreases montelukast but no dosage adjustment is required
- Side effects: headache, fatigue, dizziness, Churg-Strauss
- Precautions
 - Not for an acute exacerbation
- Category: B

Methylxanthines

• Theophylline

- Theo-24, Theo-Dur, Uni-Dur, Slo-Bid
- Bronchodilates and increases the force with which the diaphragm contracts
- 6 years and older
- Difficult to manage and as a result has not really gained wide spread acceptance
- Indicated for individuals with moderate to severe asthma
- Numerous drug interactions

Theophylline Numerous medications, foods and chemicals interact with theophylline All of the following decrease theophylline levels Smoking (cigarettes and marijuana) High protein/low carbohydrate diet

- Phenytoin
- Phenobarbital
- Carbamazepine
- Ketoconazole
- Diuretics

Theophylline

- Theophylline levels (normal 6-15mcg/dL) –15-25: GI upset, N/V, diarrhea, abdominal pain
 - -25-35: Tachycardia, occasional PVC's
 - ->35: Ventricular tachycardia, seizures
- Category: C

Long Acting Inhaled Beta 2 Agonist

- Salmeterol (Serevent)
 - Diskus
 - >4 years of age-1 puff po q 12 hours
 - No role for acute exacerbations
 - Seems to help children affected by the nocturnal cough and wheezing
 - Good for prevention of exercise induced asthma

Long Acting Inhaled Beta 2 Agonist

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- Foradil Aerolizer
 - $-\geq$ 5 years of age: 1 inhalation every 12 hours
 - -Also may be used for prevention of EIB

Combination Products

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- Mometasone/formoterol -Dulera
- Fluticasone/salmeterol –Advair
- Budesonide/formoterol –Symbicort

LABA

- FDA warning regarding increased deaths in patients treated with LABA
 - -Should be used only with inhaled corticosteroid
 - -Should be used for shortest length of time to control symptoms

www.fda.gov/CDER/Drug/infopage/LABA/default.htm accessed 07-20-2010

Omalizumab (Xolair)

- Indicated for adults and adolescents (12 years of age and above) with moderate to severe persistent asthma who have a positive skin test or *in vitro* reactivity to a perennial aeroallergen
- And...whose symptoms are inadequately controlled with inhaled corticosteroids
- SC injection

Omalizumab (Xolair)

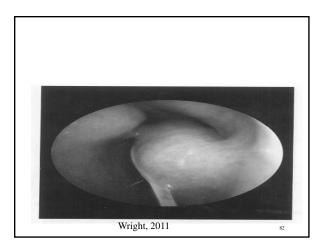
- Recombinant DNA-derived humanized IgG1 monoclonal antibody that selectively binds to human immunoglobulin E (IgE).
- Inhibits the binding of IgE to the high-affinity IgE receptor on the surface of mast cells and basophils
- Limits the degree of release of mediators of the allergic response.

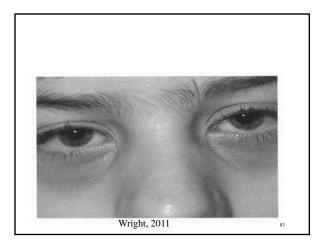
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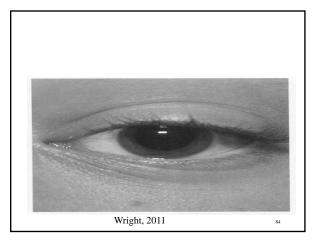
Last....

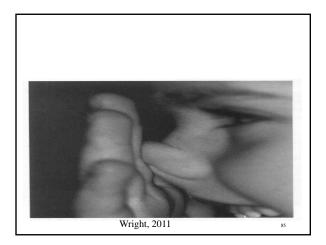
- Don't forget to treat the nose
- 85% of individuals with asthma have concomitant allergic rhinitis

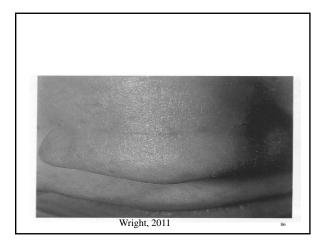


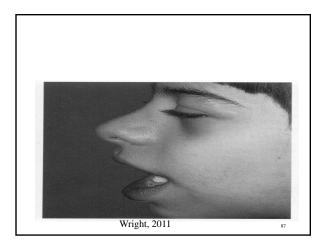


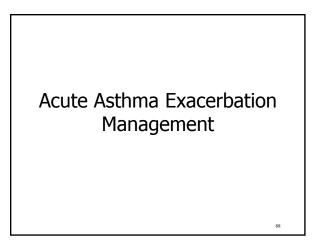












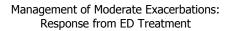
Acute Asthma Exacerbation

- Measure FEV1
- Inhaled short acting beta 2 agonist: Up to three treatments of 2-4 puffs by MDI at 20 minute intervals OR a single nebulizer
- Can repeat x 1 2 provided patient tolerates
- Prednisone – What dose and schedule??

Management of Moderate Exacerbations: Response from ED Treatment

- Good Response
 - -Symptom relief sustained x 1hr; FEV1 or PEF \ge 70%
 - -D/C home
 - -Continue SABA & oral corticosteroid
 - -Consider inhaled corticosteroid (ICS)
 - -Patient education / asthma action plan

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- Incomplete Response
 - Mild-moderate symptoms, FEV1 or PEF 40-69%
 - -SABA, oxygen, oral or IV corticosteroids
 - –Can D/C home
- Poor Response
 - -Marked symptoms, PEF <40%
 - -Repeat SABA immediately
 - -ED / 911; oral corticosteroid

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- Key Differences in the EPR-3 Report
- Point of discharge
 - -FEV1 or PEF \geq 70% predicted
 - Response sustained 60 minutes after last treatment
 - -Normal physical exam
- Continued ED treatment needed -FEV1 or PEF 40-69% predicted
- Consider adjunct therapies -FEV1 or PEF <40% predicted

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Thank you for your time and attention.

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Wright, 2011