### CLINICAL CASE

- A 38-year-old female presented with a worsening shortness of breath, chest tightness, and coughing, especially in the previous 2 weeks.
- She described itchy, watery eyes, congestion and runny nose when around cats and also in the spring and fall.

- Q.1 One of the following treatment options was prescribed to relieve her allergic condition:
- a.Atropine.
- b.Oral aminophylline.
- c.Aggressive therapy with oral steroids for 5 days.
- d.Salbutamol.

- Q.2 Your answer to the last question was based on that this drug will provide the patient with the following advantage:
- a.Its anti-inflammatory effect will dramatically relief her allergic conditions together with her bronchial asthma.
- b.It provides effective bronchodilating activity.
- c. It antagonizes the overactive parasympathetic tone

# BRONCHIAL ASTHMA

By Dr. Rania A. Abdel-Emam

## ANTI-ASTHMATIC DRUGS

- Bronchodilators
- β2-adrenergic agonists
- Anticholinergic drugs
- Methylxanthines
- Antiinflammatory drugs :
- Corticosteroids
- Cromolyn sodium
- Leukotriene antagonists
- Anti-IgE therapy

## **ANTI-INFLAMMATORY DRUGS**

## CORTICOSTEROIDS

#### Mechanism of action:

- HOW DO THEY IMPROVE BRONCHIAL ASTHMA?
- [Potent anti-inflammatory drugs] by inhibition of the formation of inflammatory mediators and suppression of acute and chronic inflammation in airways. This is achieved via:
- 1- inhibition of phospholipase  $A_2$  leading to inhibition of synthesis of inflammatory mediators from arachidonic acid.
- 2- inhibition of COX<sub>2</sub> leading to inhibition of production of PGNs by inflammatory cells.

### CORTICOSTEROIDS

- 3- reduces the levels of circulating eosinophils, lymphocytes, basophils & monocytes leading to decrease bronchial infiltration by chronic inflammatory cells.
- 4- reduces the release of histamine from mast cells leading to decrease capillary permeability, vasoconstriction and tissue edema.

• Potentiate the bronchodilator action of the  $\beta_2$  agonists.

# Uses of corticosteroids in treating bronchial asthma

### According to the route of administration

### INHALATION

- 1<sup>st</sup> line prophylactic therapy
- beclomethasone, triamcinolone, budesonide.
- ADRs:
- 1- Hoarseness of voice.(due to direct effect on vocal cords)
- 2- Oropharyngeal candidiasis. (prevented by rinsing mouth and throat after each use)

### CR SYSTEMICALLY

- Acute exacerbations (short course of oral prednisone for 5 days).
- Status asthmaticus (I.V. methylprednisolone every 6 hours until the attack subsides. Then continue on oral prednisone).
- The dose is gradually tapered over 10-14 days.

### MAST CELL STABILIZERS

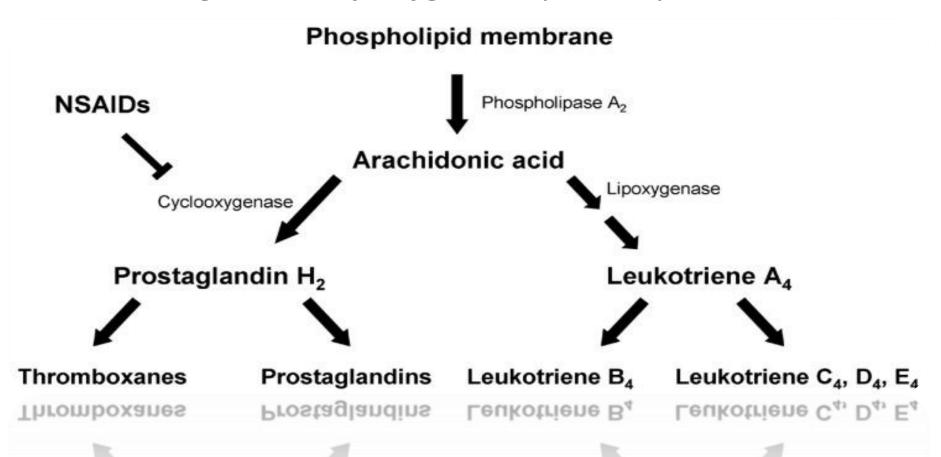
Cromolyn sodium and nedocromil

 Stabilization of mast cell on exposure to allergens preventing degranulation and release of histamine and broncho-constrictor chemical mediators.

- They are not useful in managing an acute asthma attack but only as a prophylactic treatment.
- Given by inhalation.
- Nedocromil more effective than cromolyn and approved for 12 years and older [cromolyn is approved for all ages]

# Leukotriene antagonists

• Leukotrienes are biochemical mediators from arachidonic acid through the **5-lipoxygenase** pathway.

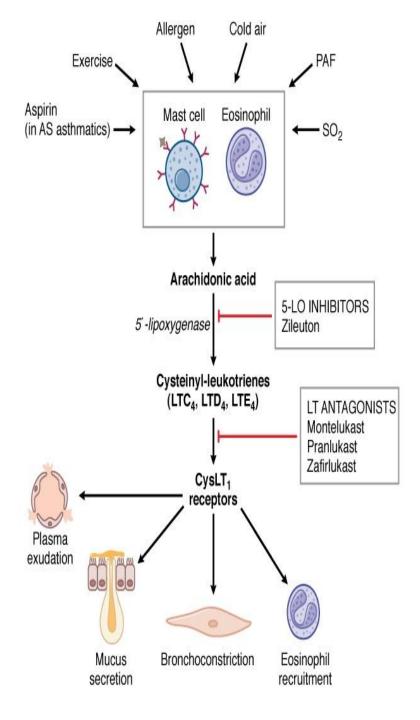


• **Zileuton**: (four times daily)

It is 5-lipoxygenase inhibitor preventing formation of both LTB<sub>4</sub> and the cysteinyl leukotrienes (LTC<sub>4</sub>, LTD<sub>4</sub>, LTE<sub>4</sub>). **HEPATOTOXICITY** 

Zafirlukast & Montelukast:

They are blockers of leukotriene receptors. Prophylactic ttt, Aspirin induced asthma.

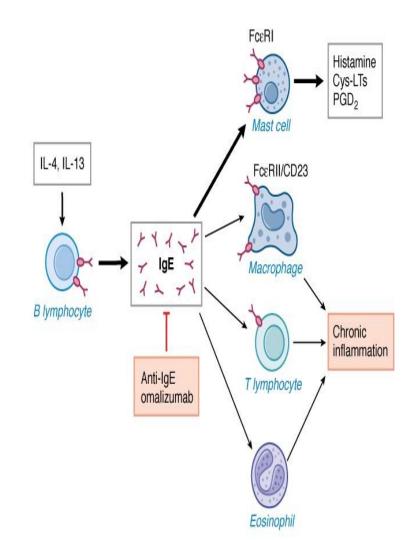


Used as a Prophylactic treatment of mild asthma.

• They are of value as a prophylaxis against aspirininduced asthma where aspirin inhibits cyclooxygenase pathway of arachidonic acid metabolism giving the upper hand to lipoxygenase activity which then will be suppressed by antileukotrienes.

# Anti-IgE therapy Omalizumab

- It is a recombinant monoclonal antibody targeted against free IgE (Anti-antibody).
- IgE bound to omalizumab cannot bind to IgE receptors on mast cells and basophils, preventing the allergic reaction at a very early step in the process.
- Used as a prophylactic treatment



### MANAGEMENT OF ACUTE ASTHMA

#### Mild attacks

Inhalation of a short acting β2 -agonist (salbutamol)

#### **Moderate attacks**

- Inhaled short acting β2 agonist continuously
- Early administration of systemic corticosteroids

#### Severe attacks and status asthmaticus

- Oxygen.
- β 2 agonist (salbutamol), Anticolinergics, Methylxanthines I.V.
- Corticosteroids I.V.
- Mechanical ventilation.

## Long-term management

- Avoidance of exposure to triggering factors
- Treatment of upper respiratory tract infections (e.g. Sinusitis).
- Anti-asthmatic drugs:
  - Long acting B2 agonists (inhaled)
  - Corticosteroids (inhaled)
  - Mast cell stabilizers
  - Leukotriene antagonists
  - Anti-IgE therapy
- Hyposensetization
- Psychotherapy & biofeedback

- Q.3 ONE of the following treatment options was given to her as a maintenance treatment:
- a.lpratropium.
- b.Intravenous aminophylline.
- c.Inhaled corticosteroid and long-acting beta agonist.
- d.Inhaled salbutamol.

- ❖ For avoidance of recurrence of the acute attack of bronchial asthma; one of the following drugs was properly administered:
  - A- Isoprenaline.
  - **B- Epinephrine.**
  - **C-** Cromolyn sodium.
  - D- Ephedrine.

- The drug you selected in the last question is administered through one of the following routes:
- A- Subcutaneously.
- B- Intramuscularly.
- C- By inhalation.
- D- Orally.

