

KING SAUD UNIVERSITY COLLEGE OF MEDICINE  $\mathbf{1}^{\text{ST}}$  YEAR,  $\mathbf{3}^{\text{RD}}$  BLOCK

# Treatment of Rhinitis and Cough

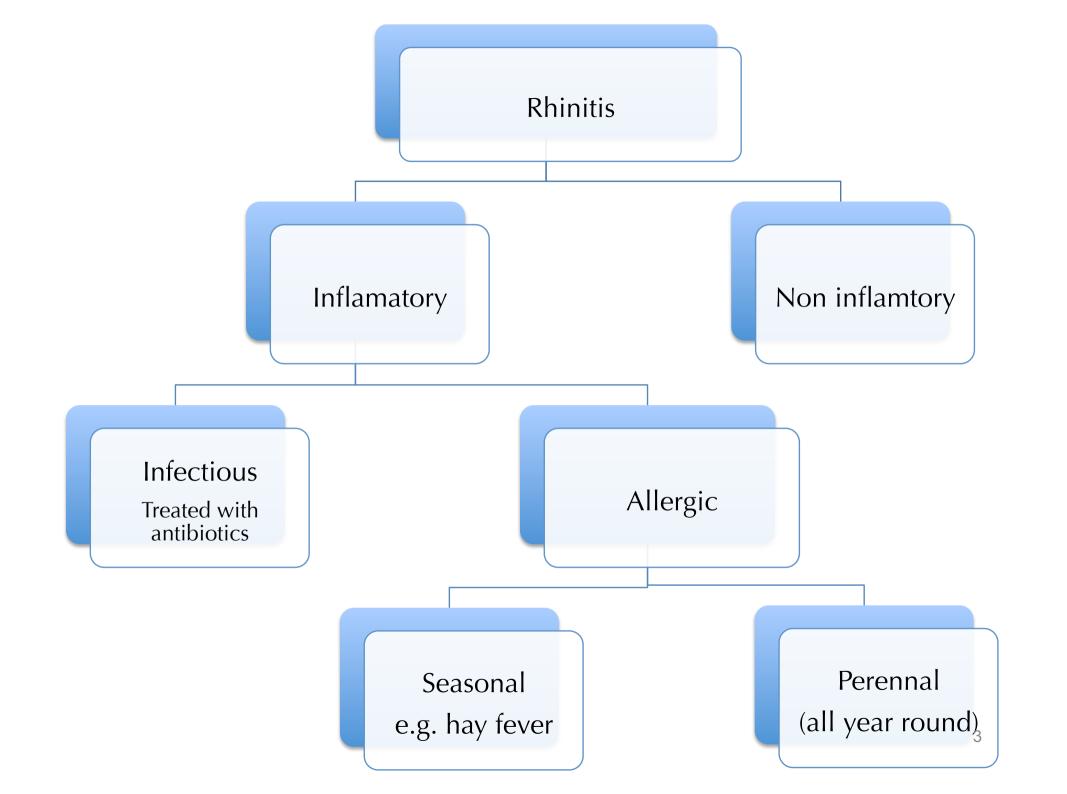




RESPIRATORY BLOCK

### **Objectives:**

- Classify types of rhinitis.
- Specify preventive versus pharmacotherapeutic strategies.
- Expand on the pharmacology of different drug groups used in treatment as antihistamines, anti-allergics, corticosteriods, decongestants and anti-cholinergic.
- Differentiate between productive versus dry irritant cough.
- Compare pharmacology of different expectorants & mucolytics drugs used in treatment of productive cough.
- Contrast between peripherally and centrally acting antitussives.



1- H<sub>1</sub> receptor antagonists

#### **Symptoms:**

Runny nose (rhinorrhea), Stuffy Blocked nose, Sneezing, Nasal congestion, Post-nasal drip, Itching, Catarrh 2-Anti-allergics:

- Mast Cell Stabilizer;
- Leukotriene receptor antagonists

3- Corticosteroids

### **Rhinitis**

Rhin: nose

-itis: inflammation

Pharmacotherapy treatment

4- Decongestants; α-Adrenergic agonists

5- Anticholinergics

6- Antibiotics

7- Mycolytics

### 1 - Anti-Histamines

	1 <sup>st</sup> generation	2 <sup>nd</sup> generation	3 <sup>rd</sup> generation
Example	Diphenhydramine	Loratidine Cetirizine	Desoloratidine Levocetirizine
Duration of action	Short	Longer "better control"	Longer "better control"
Selectivity	Non-selective	Selective	More selective
Crossing BBB	Cross	Poor corssing	No crossing
Drug interactions	with enzyme inhibitors [ macrolides, antifungals, calcium antagonists]	No drug interactions	No drug interactions
Sedating effects	Sedating	Non-sedating	Non-sedating
Side effects	additive pharmacodynamic side effects	Minimal side effects	Minimal side effects

\*Anti-histamines block other receptors than H1 such as: Cholinergic, alpha-Adrenergic, Serotonin But they do not not block Dopamine nor H2

### 1 - Anti-Histamines

Indications linked to H <sub>1</sub> blockers	Indications linked to blocking other receptors than H <sub>1</sub>		
	Drugs	Action	Mechanism
Allergies: -Good control of: Rhinitis, Conjunctivitis, Urticaria, Flu (cough and sneezing) Poor control of Asthma, Otitis, Anaphylaxis, Sinusitis, Atopic dermatitis.	Dimenhydrinate, Diphenhydramine Promethazine	Vertigo & Motion sickness	-
Itching	Promethazine	Anti-emetic	<ul><li>Decrease firing from internal ear to vomiting center</li><li>Decrease firing to vomiting center + Anticholinergic</li></ul>
Others: Insomnia	Chlorpheniramine Dimenhydrinate Promethazine	Anti-parkinsonism	By anticholinergic action that will decrease the extrapyramidal effects.
Sleep aid Vertigo	Cyproheptadine	Increase appetite	By 5-HT modulation ↑ Sedation
Anxiety Cough	Promethazine, Antazoline	Anti-arrhythmic actions	by Na channel blocking action & local anesthetic effects

### 2-ANTI-ALLERGICS

Mast cell stabilizers		Leukotrienes Antagonists	
example	Cromolyn, Nedocromyl	Zafirlukast, Montelukast, Pranlukast	
Mechanism of action	Decrease Histamine release [ by inhibiting Cl channels ] *It does not antagonize released histamine	Block leukotriene actions	
Uses	In children for <u>prophylaxis</u> of perennial allergic rhinitis  (not used a lot in Asthma, work better in upper respiratory tract allergy)	-Prophylaxis of <u>lower respiratory tract</u> allergies e.g. perennial allergen, exercise or aspirininduced asthma  (Work good in Asthma, less actions in upper respiratory tract allergy)	
Side effects	Cough, wheezes, headache, rash.	Elevation of liver enzymes, headache, dyspepsia.	

	3 - CORTICOSTERIODS
example	Beclomethasone, Budesonide, Fluticasone
Mechanism of action	Inhibition of phospholipase A2 ↓ prostaglandin and leukotrienes
Uses	severe intermittent or moderate persistent symptoms
Side effects	Nasal irritation, fungal infection, hoarseness of voice

4. DECONGESTANTS (α-Adrenergic agonists)					
	Systemic Topical				
Example	Pseudoephedrine Phenylethylamines, Imidazoline				
Uses	For treatment of nasal stuffiness				
Side effects	Nervousness, insomnia, tremors, palpitations, hypertension.	Rebound nasal stuffiness			

5. ANTICHOLINERGICS			
Example	Ipratropium		
Uses	<ul> <li>Bronchiodilator in asthma.</li> <li>Control rhinorrhea (excess nasal secretion &amp; discharge).</li> <li>Effective in vasomotor rhinitis (watery hyper-secretion).</li> </ul>		
Side effects	minimal systemic side effects	8	

The respiratoy tract is protected by:

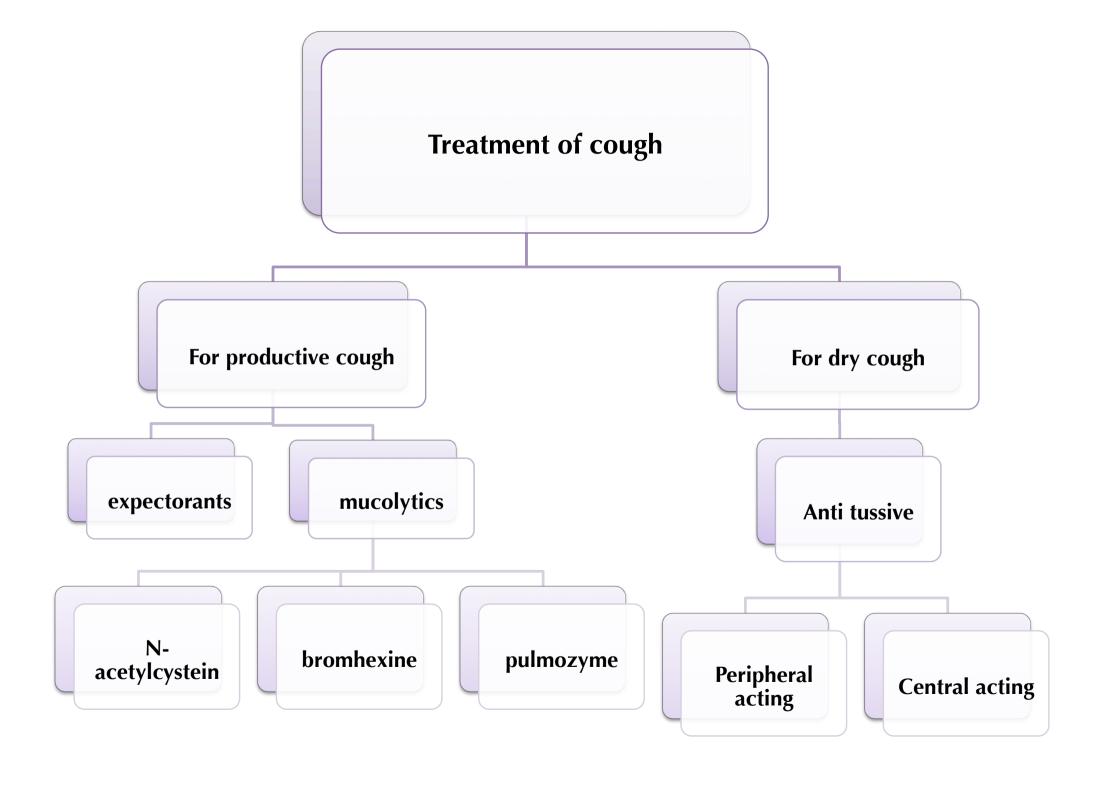
#### **Mucocilliary Clearance:**

Ensures optimum tracheobronchial clearance by forming sputum

(in optimum quantity & viscosity ) exhaled by ciliary movements.

#### **Cough Reflex:**

Exhales sputum out, if not optimally removed by the mucociliary clearance mechanisms.



### **Expectorants**

### Act by removal of mucus through:

	Reflex stimulation	Direct stimulation	
Example	<u>Guaifenesin</u>	Iodinated glycerol, Na or K iodide/acetate , Ammonium chloride, Ipecacuahna	
Mechanism of action	Irritate GIT → stimulate gastro-pulmonary vagal reflex → loosening & thinning of secretions.	Stimulate secretary glands → increase respiratory fluids production.	
Side effects	Dry mouth (Xerostomia) chapped lips risk of kidney stones (†uric acid excretion)	Unpleasant metallic taste hypersensitivity Hypothyroidism swollen of salivary glands (overstimulation of salivary secretion) flare of old TB.	
Indications	Final outcome is that cough is indirectly diminished  Common cold, Bronchitis, Laryngitis, Pharyngitis, Influenza, Measles, Chronic paranasal sinusitis, Pertussis.		

Mucolytics

Act by altering biophysical quality of sputum → becomes easily exhaled by mucociliary clearance or by less intense coughing

	N-Acetylcysteine	Bromhexine, Ambroxol	Pulmozyme (Dornase Alpha or DNAse)	Hypertonic Saline & NaHCO <sub>3</sub>	Steam inhalation
overview	A free radical scavenger used in Acetaminophen overdose. (an antidote for it)	They increase immune defense:	A recombinant of human deoxyribo-nuclease-1 enzyme that is neubilized. Full benefit appears within 3-7 days		
Mechanism of action	Breakdown S-S bonds in Glycoproteins by its reducing SH Gp → less viscid mucous	Synthesize serous mucus (sialomucins of smaller-size) so it is:  Secretolytic + activate ciliary clearance & transport	Cleavage of extracellular bacterial DNA, that contributes to viscosity of sputum in case of infection	Decrease viscoelasticity by increasing thewater content.	Decrease the adhesivness.
Indications	- Most mucolytics → effective as adjuvant therapy in COPD, asthma, bronchitis, (whenever there is excessive and/or thick mucus)  - In bronchiectasis, pneumonia & TB → they are of partial benefit Hardly any benefit in cystic fibrosis & severe infections → Give rhDNAase				
Side effects	Bronchospasm stomatitis, rhinorrhea, rash, nausea & vomiting.	Rhinorrhea, lacrymation, gastric irritation, hypersensitivity	Voice changes, pharyngitis, laryngitis, rhinitis, chest pain, fever, rash		12

### Anti-tussive Agents

Stop or reduce cough by acting either primarily on the peripheral or CNS components of cough reflex.

### PERIPHERALLY ACTING ANTITUSSIVES

#### A. Inhibitors of airway stretch receptors:

- In Pharynx: Use Demulcents → form a protective coating Lozenges & Gargles
- In Larynx: Use Emollients → form a protective coating Menthol & Eucalyptus.
- -In Tracheobronchial Airway→ Use aerosols or inhalational of hot steam

#### **Tincture Benzoin Compound & Eucalyptol**

-During bronchoscopy or bronchography → Use local anaesthetic aerosols

Lidocaine, Benzocaine, and Tetracaine

### **B.** Inhibitors of pulmonary stretch receptors in alveoli:

**Benzonatate:** ★ sensitivity (numbing) of receptors by local anesthetic action.

#### **Side effects:**

drowsiness, dizziness, dysphagia, allergic reactions

- Overdose → mental confusion, hallucination, restlessness & tremors.

### CENTRALLY ACTING ANTITUSSIVES

#### A. OPIOIDS:

activating  $\mu$  opioid receptors

Codeine & Pholcodine

#### **B. NON-OPIODS:**

Antihistaminics (>sedating)

#### **Dextromethorphan:**

- It increase threshold at cough center.
- It has benefits over opiods in being →
- 1. As potent as codeine.
- 2- But no drowsiness.
- 3- Less constipating
- 4- No respiratory depression.
- 5- No inhibition of mucociliary clearance.
- 6- No addiction.

#### **Side effects:**

Nausea, vomiting, dizziness, rash & pruritis in normal doses

In high doses, hallucinations + opiate like side effects on respiration & GIT

### **SUMMARY**

	Mechanism of action	Clinical uses	Side effect
2)anti allergics:- a) Acromolyn & Nedocromyl	a)Mast cell stabilizer	Treatment of rhinitis and act as prophylactic	a)Rash,cough,headache,wheeze
b) Leukotriene receptor antagonists	b) Block leukotriens action		b) Elevation of liver enzymes, headache, dyspepsia.
3)corticosteroids:- Fluticasone & Budesonide	Block phospholipase A2=anti inflammatory drug	Treatment of rhinitis and in allergy cases	Nasal irritation, infection, hoarseness of voice
4)decongestants:- a)topical Imidazoline & Phenylethylamines b)systemic pseudoephedrine	Adrenergic agonists	Treatment of rhinitis and nasal stuffiness	A) Rebound nasal stuffiness  B) Nervousness, insomnia, hypertension
5)anticholinergics:  Ipratropium	Stop over activity of parasympathetic	Treatment of rhinitis and effective in vasomotor rhinitis (watery hyper secretion)	14



- 1- An old man with a productive cough is on one of its medications. But after a while he got Xerostomia and chapped lips. What's the drugs that he's taking?
- A. Antazoline
- B. Guaifenesin
- C. N-Acetylcysteine
- D.Benzonatate
- 2- A medical student who's having an exam after two days, came to the hospital complaining of his runny nose and continuous sneezing and congestions in his nose. What's the best drug to prescribe him with?
- A. Ipecacuahna
- B. Promethazine
- C. Diphenhydramine
- D. Levocetirizine
- 3- A patient with productive cough is on a medication for this cough, but he thinks his voice changes because of this medication. What is it ?
- A. Ipratropium
- B. Zafirlukast,
- C. Dextromethorphan:
- D. Pulmozyme

1-B, 2-D, 3-D

## We hope we made this lecture easier for you Contact us for any questions or comments Good Luck!

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