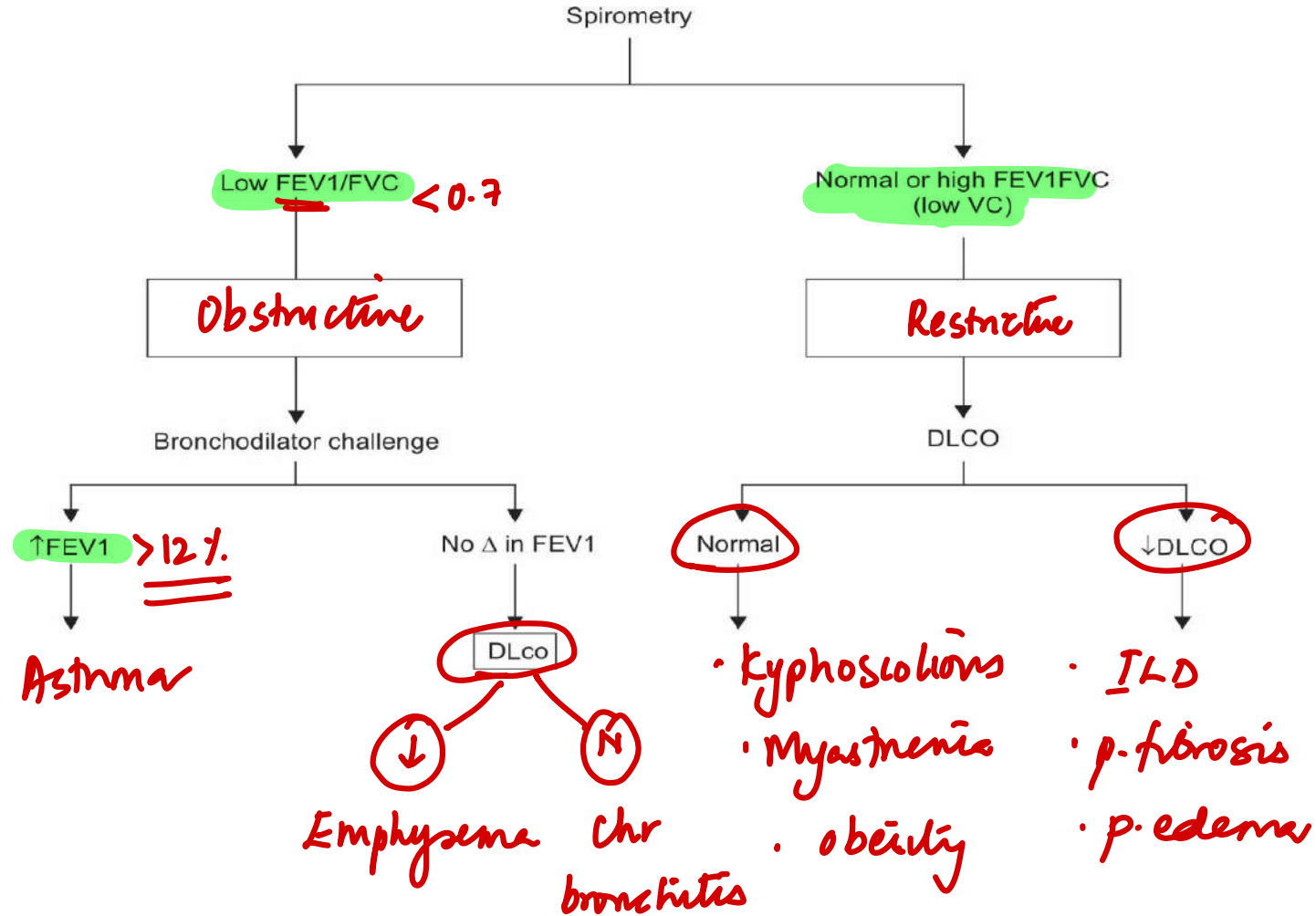
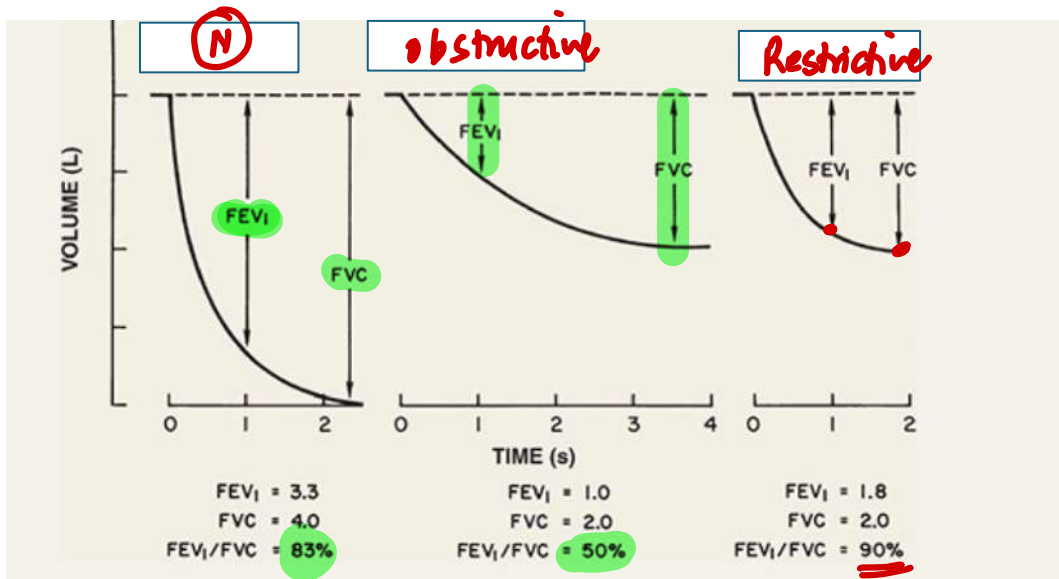
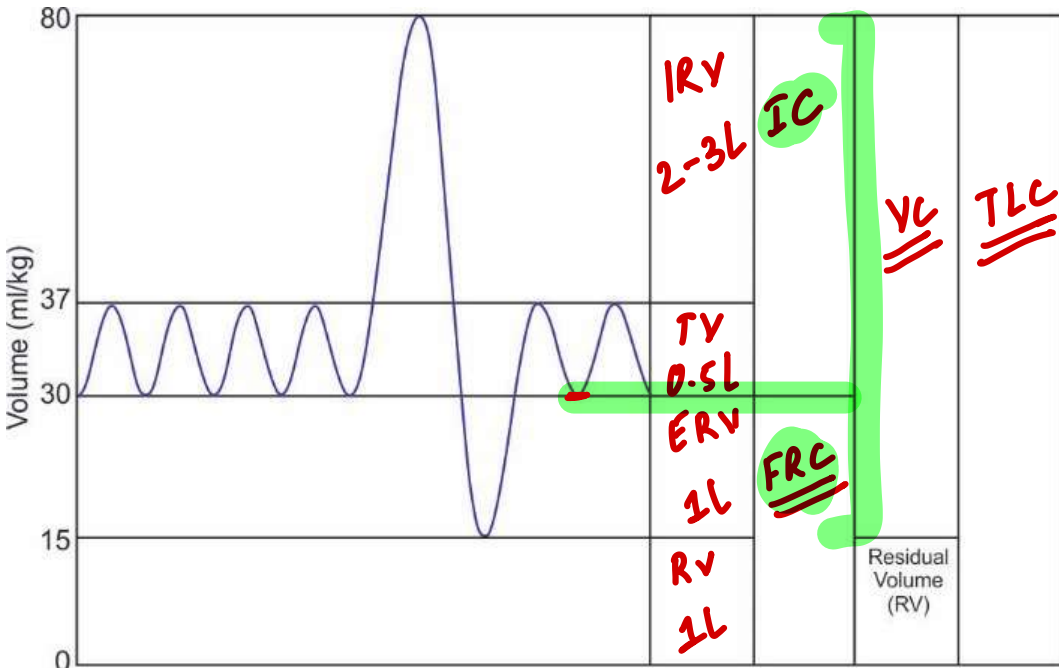


# **INTEGRATED RESPIRATORY SYSTEM**

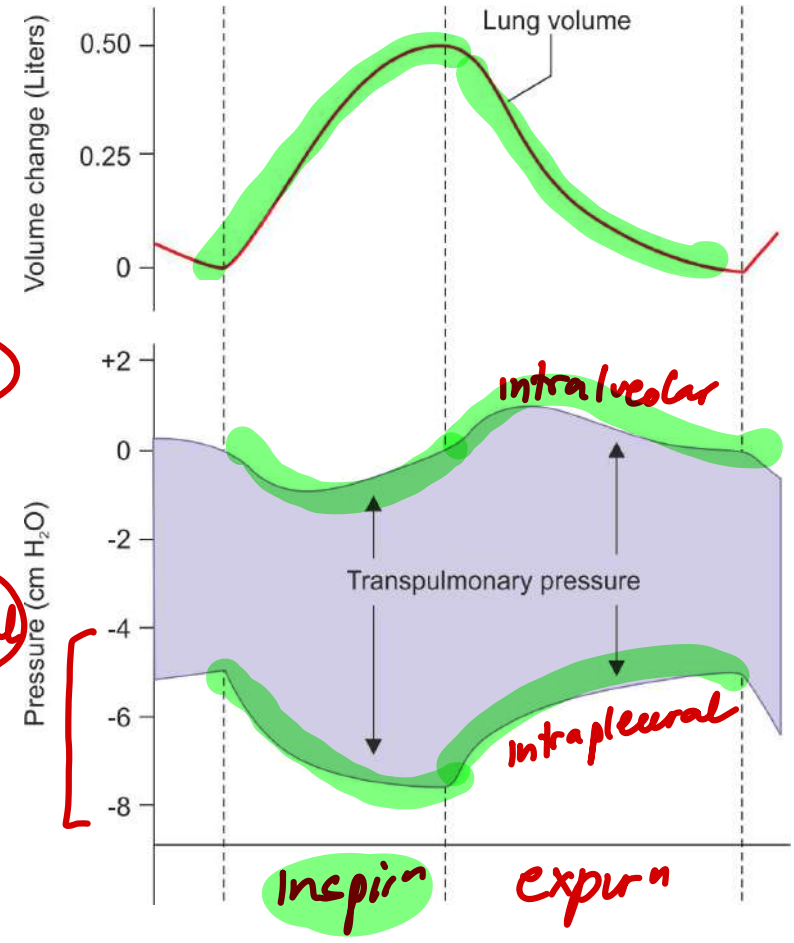
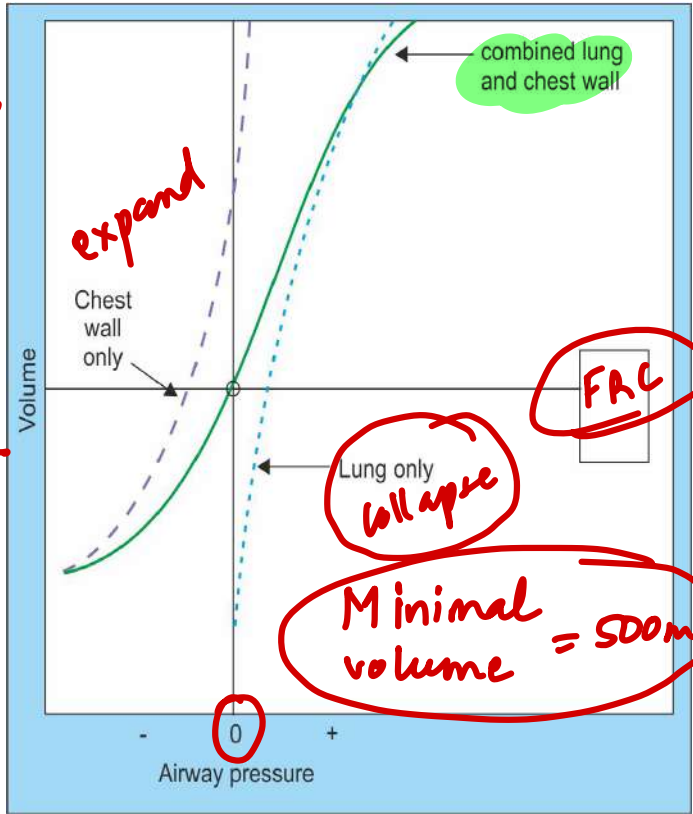
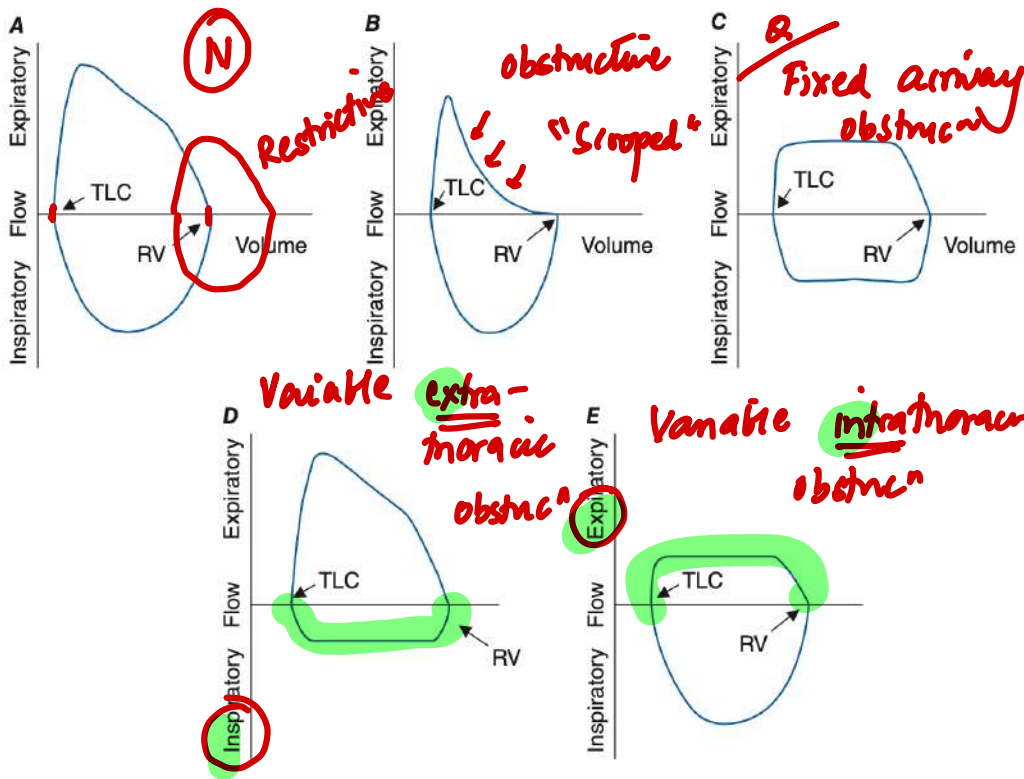
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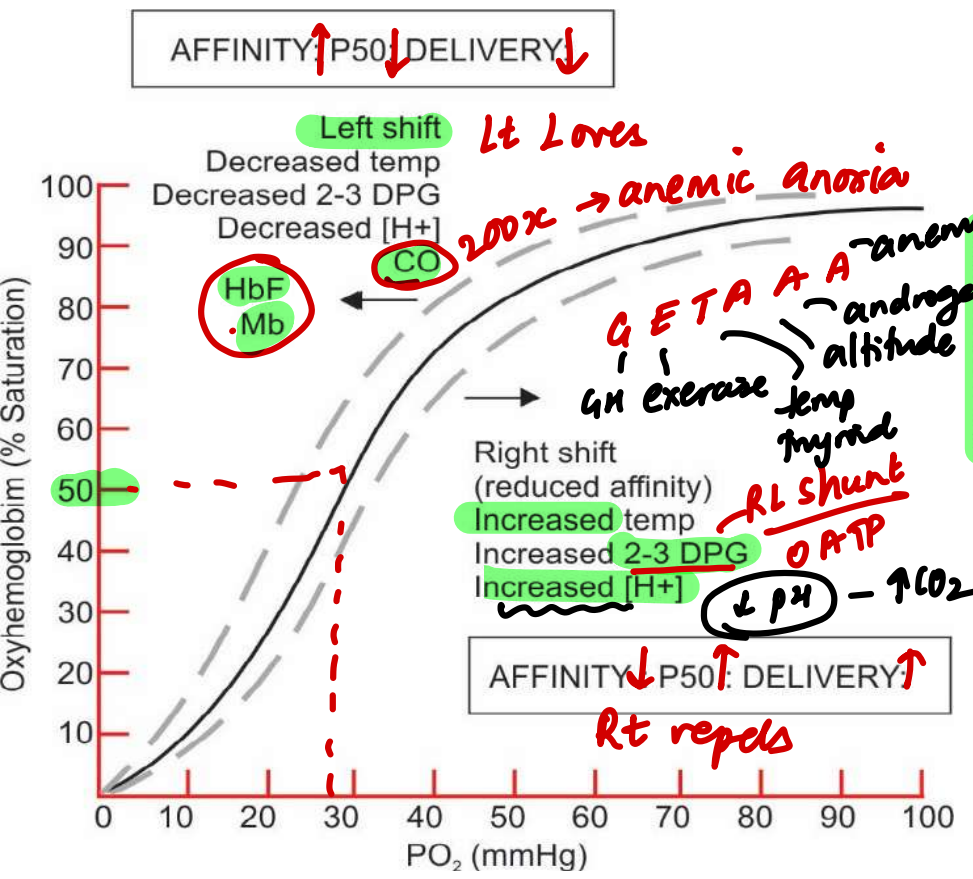
# SPIROMETRY



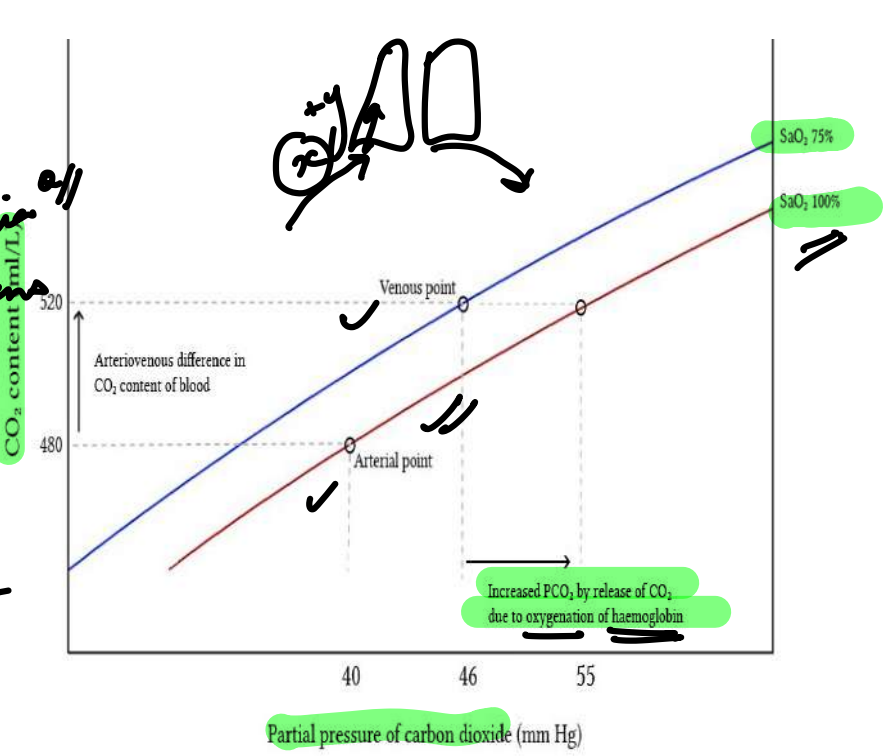
Helium dilution  
Nitrogen washout  
Body plethysmography - best

**RV**

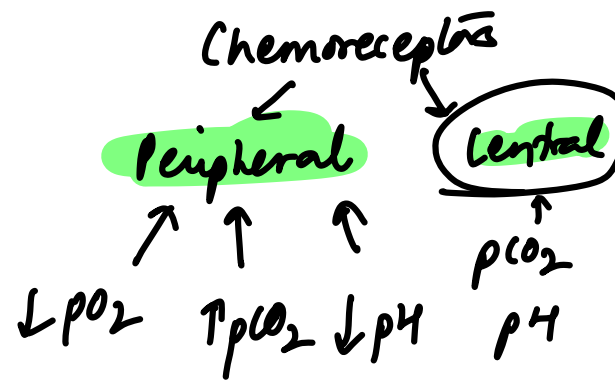
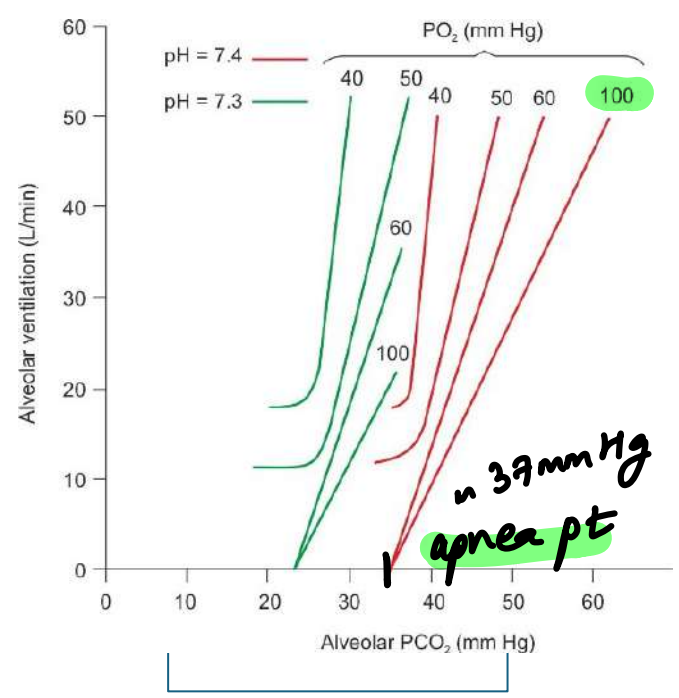




Shape: **Sigmoidal**  
**Positive co-operativity**  
 P50: **27 mm Hg**  
 2,3 DPG binds to: **Globin (B)**

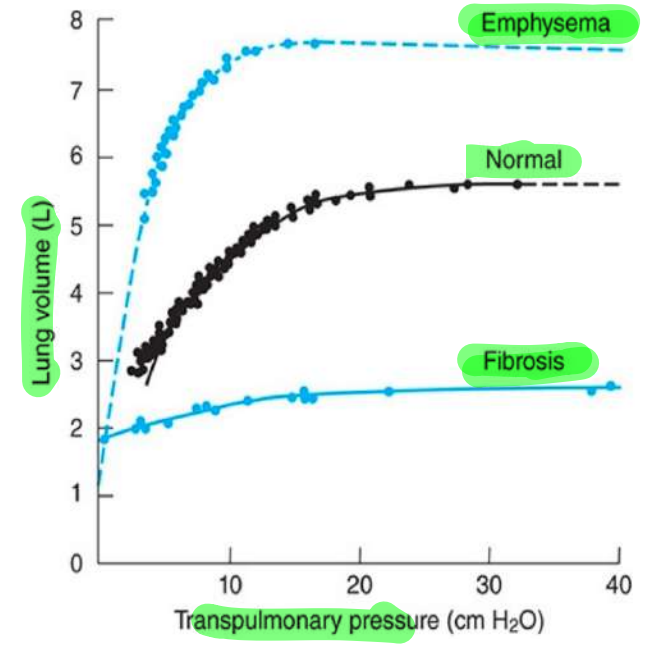
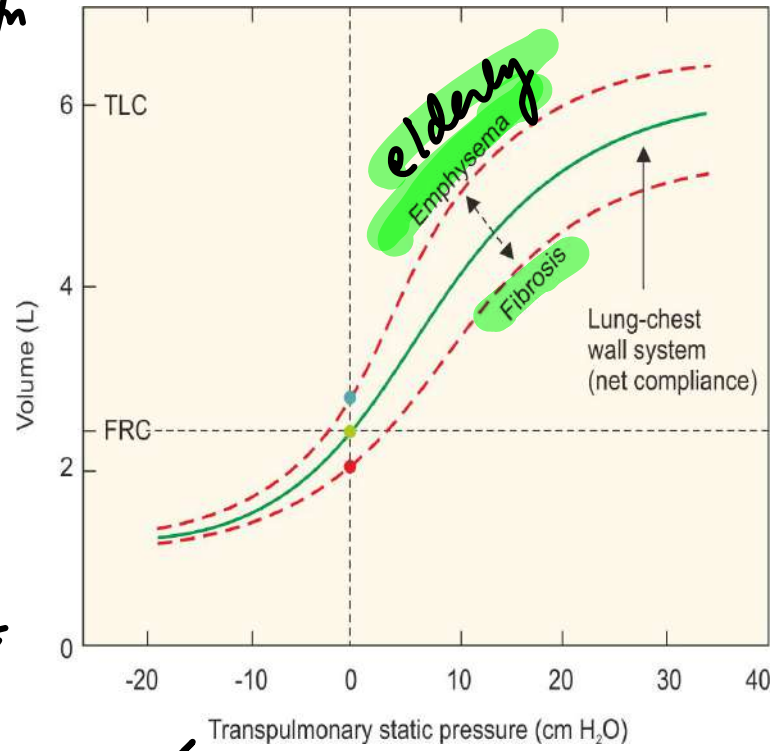
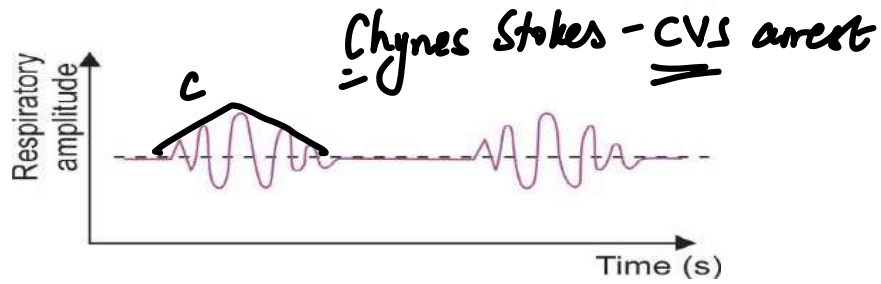
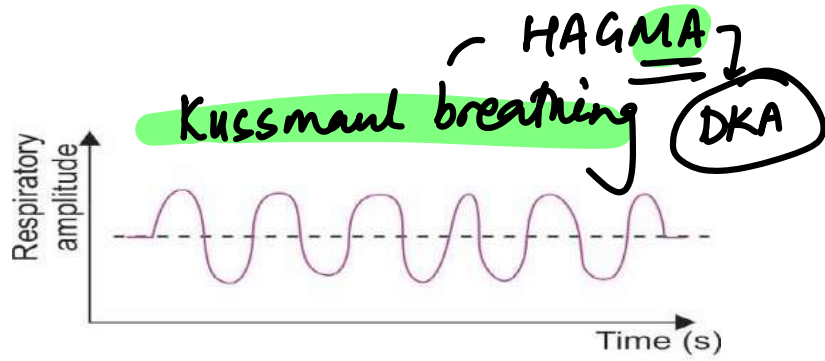
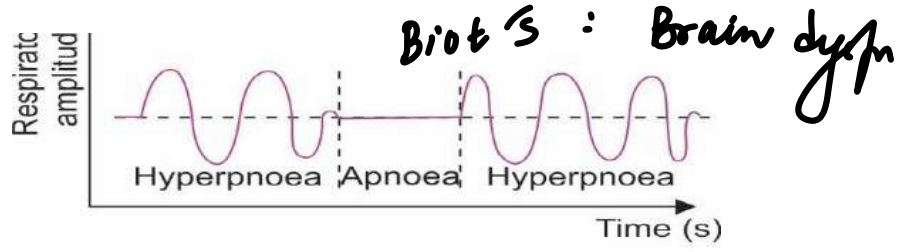


**Haldane effect**  
 $O_2 \uparrow \rightarrow CO_2 \uparrow$



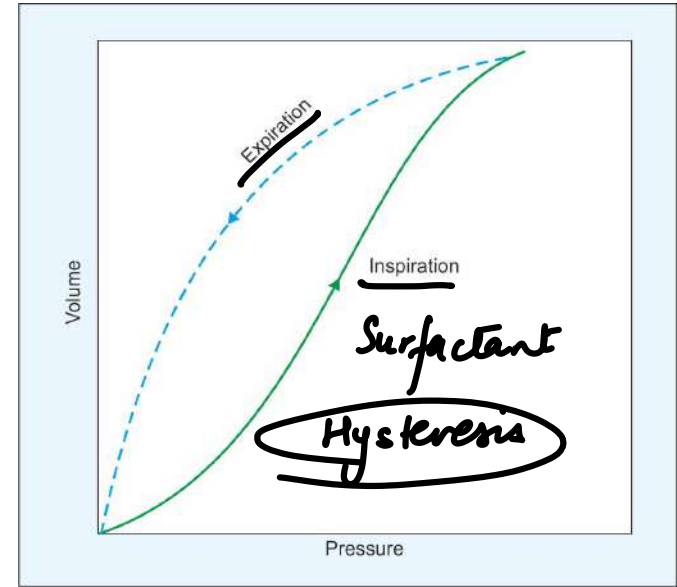
**BOHR effect**

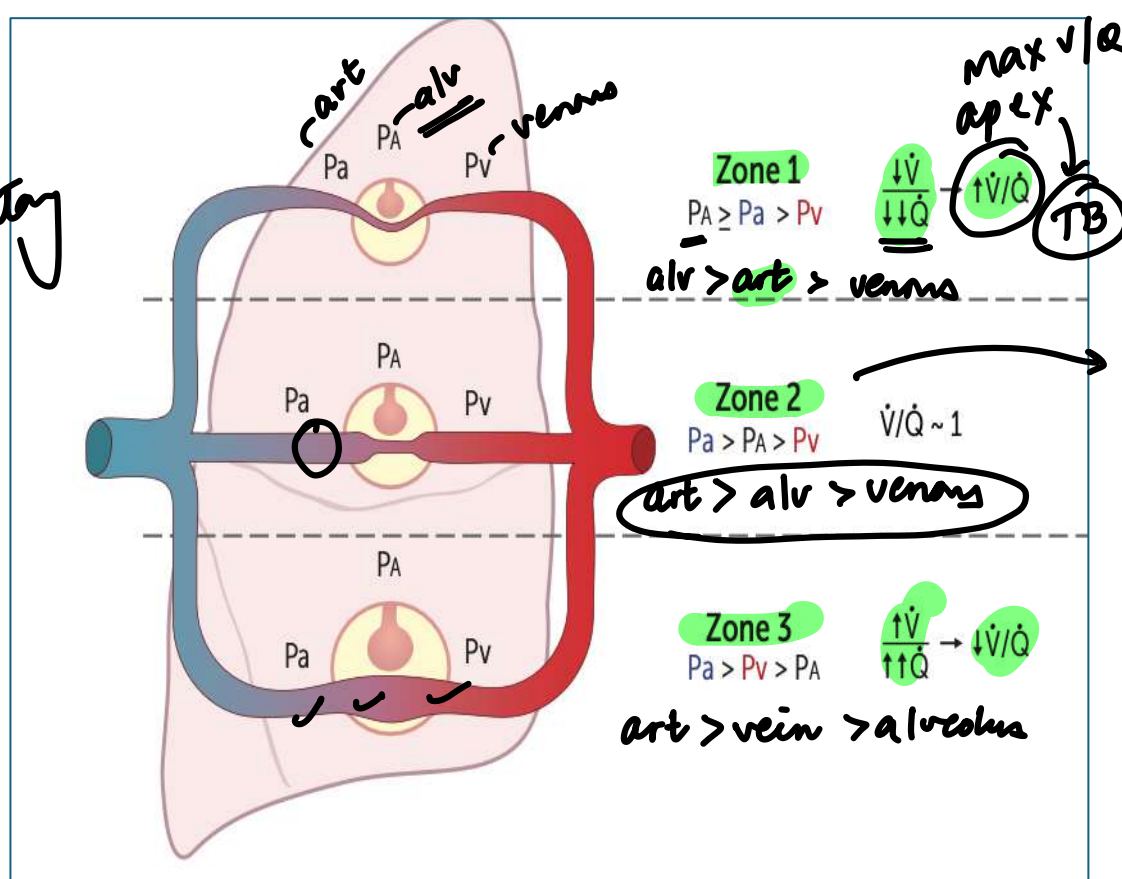
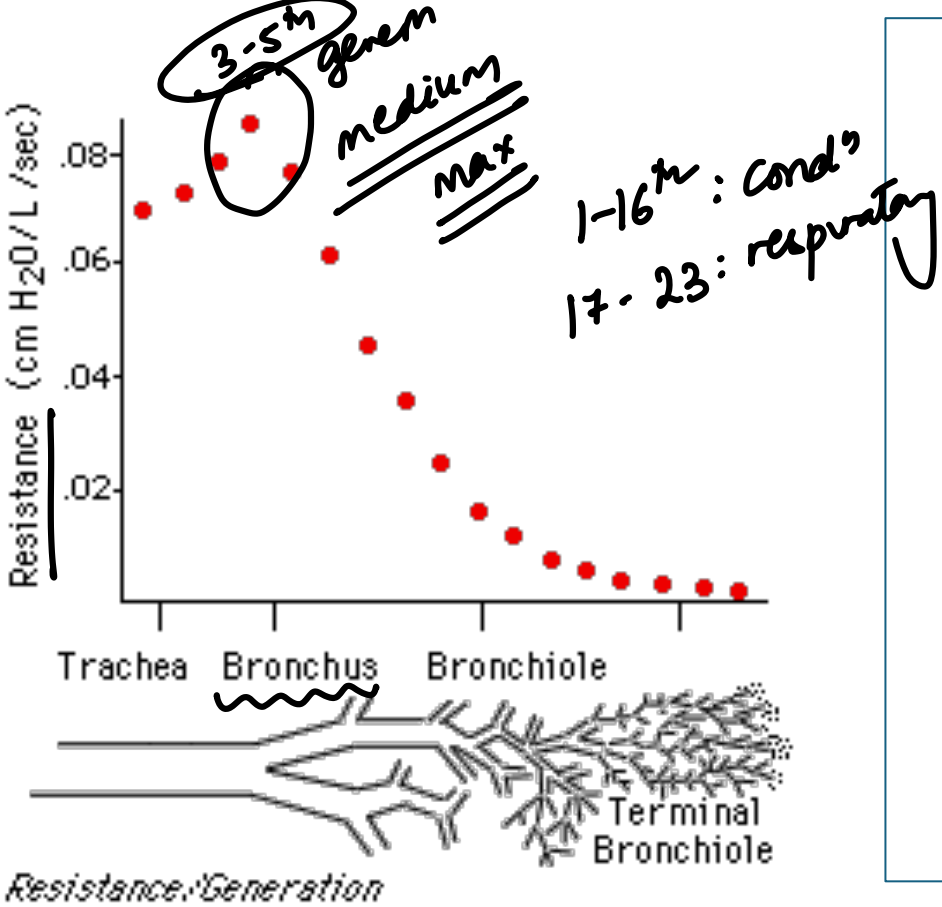
**Partial pressure of carbon dioxide increases - Reduced affinity of  $O_2$**



**Surfactant: Compliance ↑**  
**Surface tension ↓**

**Compliance:  $\Delta V / \Delta P$**





PPV / Hypovolemia:  
 Zone 1

SLUICE/waterfall effect

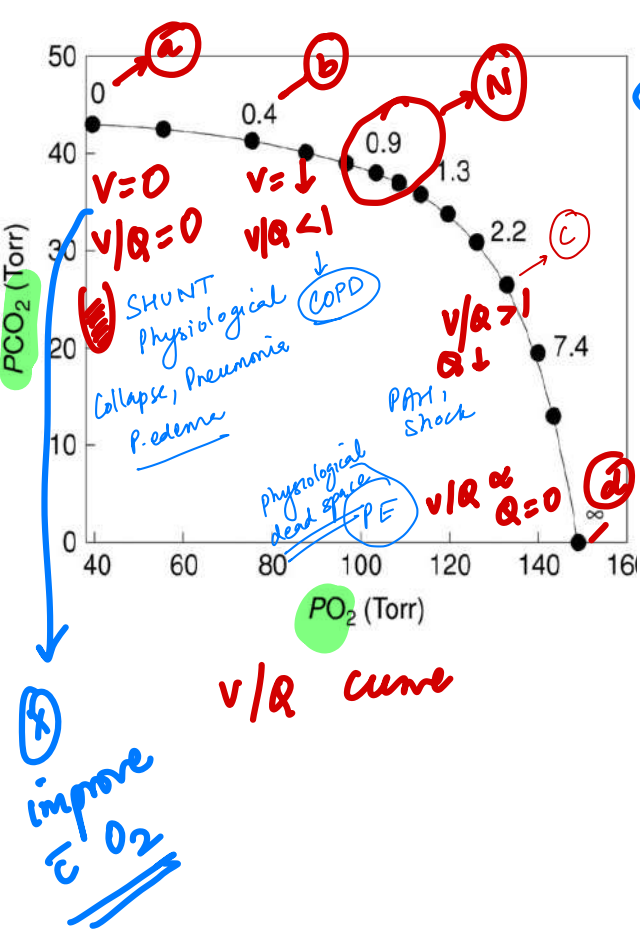
Exercise: Zone 3

Zone 4

$\downarrow$   
 $\uparrow$  interstitial pressure

Supine, exercise: V/Q matching - all lung

$P_{art} > P_{int} > P_{venous}$



# HYPOXIA

**1**  $\downarrow$  inspired oxygen tension ( $P_{iO_2}$ )  
 $P_{iO_2} = F_{iO_2} \times (P_B - P_{H_2O})$ : most commonly due to  $\downarrow P_B$  in high altitude

**2** Hypoventilation (due to  $\uparrow P_{aCO_2}$ )  
 $P_{aO_2} = P_{iO_2} - P_{aCO_2} / RQ$  (eg, CNS depression from opiate overdose, obesity hypoventilation syndrome, neuromuscular weakness)

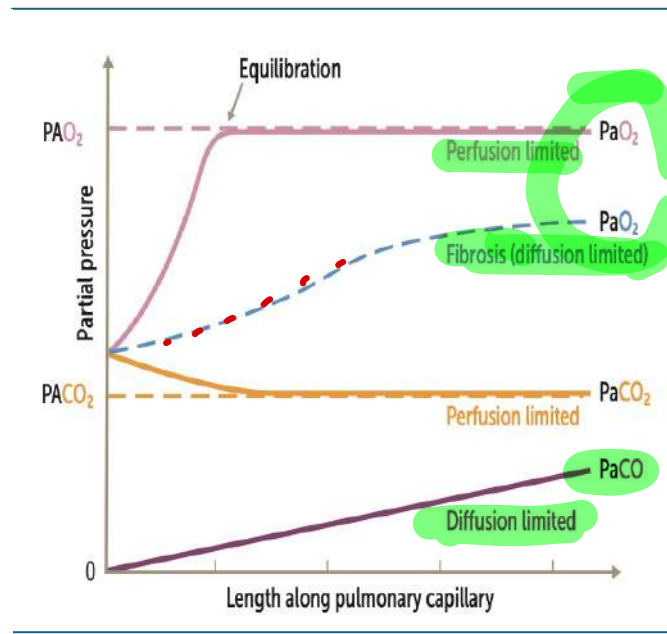
**3** Diffusion limitation (eg, fibrosis)

**4**  $\dot{V}/\dot{Q}$  mismatch  
 Normal perfusion in areas of  $\downarrow$  ventilation (eg, COPD, pulmonary edema, pulmonary embolism)

**5** Right-to-left shunt (the extreme of  $\dot{V}/\dot{Q}$  mismatch)  
 Normal perfusion in areas of no ventilation. Can be anatomic (eg, intracardiac shunt) or physiologic (eg, perfusion of nonventilated alveoli in ARDS)

Alv-arterial

Normal A-a gradient  
 Increased A-a gradient



# FORMULAE

MINUTE VENTILATION

$$RR \times TV$$

ALVEOLAR VENTILATION

$$RR \times (TV - TV_D)$$

OXYGEN CARRYING CAPACITY OF BLOOD

$$C_aO_2 = \text{Hb-bound } O_2 + \text{dissolved } O_2$$

$$C_aO_2 = (O_2 \text{ carrying capacity} \times S_aO_2) + (P_aO_2 \times 0.0031)$$

$$1.34 \times Hb \times SaO_2$$

$$V_D = V_T \times \frac{P_{aCO_2} - P_{eCO_2}}{P_{aCO_2}}$$

*- arterial - expired*

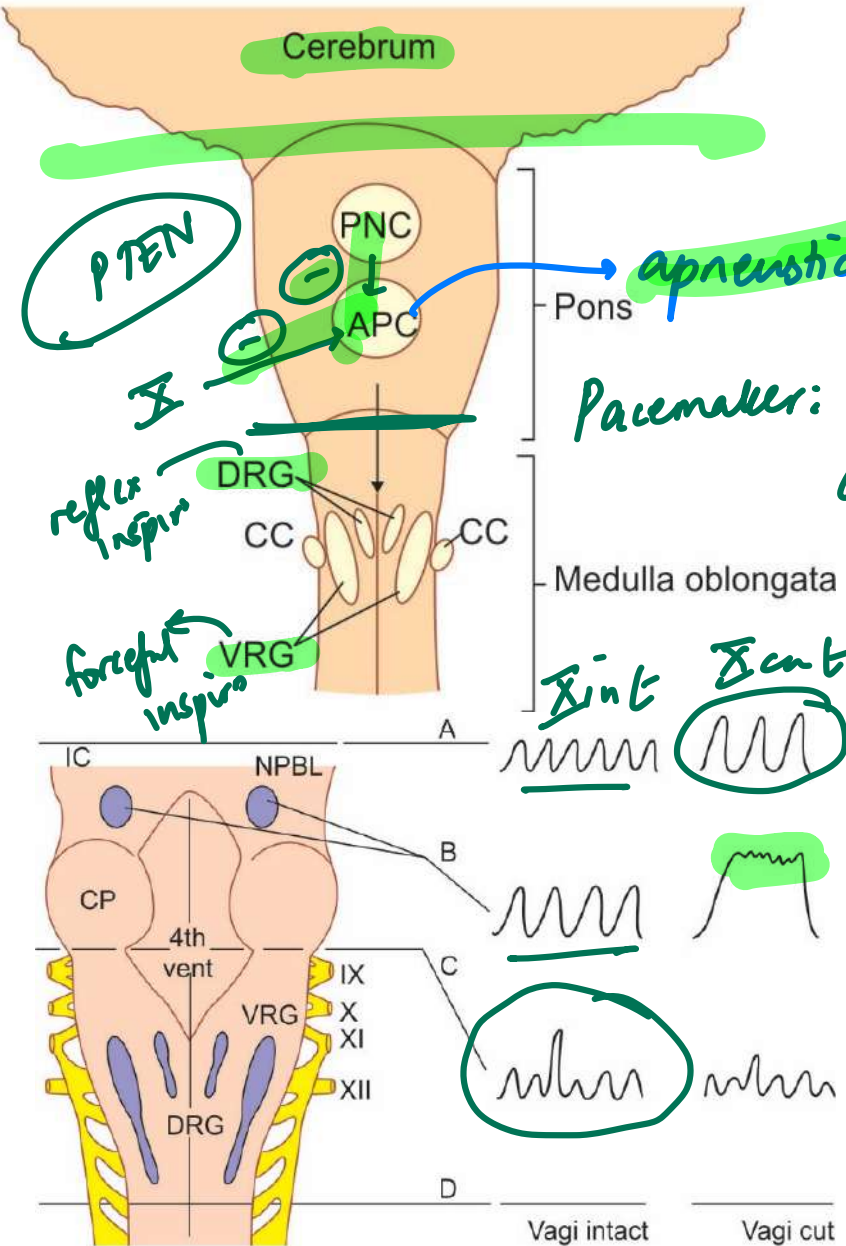
	Hb CONCENTRATION	Sao <sub>2</sub>	Pao <sub>2</sub>	TOTAL O <sub>2</sub> CONTENT
<u>CO poisoning / MethHb</u>	—	↓	—	↓
Anaemia	↓	—	—	↓
Polycythaemia	↑	—	—	↑
Cyanide toxicity	N	N	N	N

*Memic. anoxia*

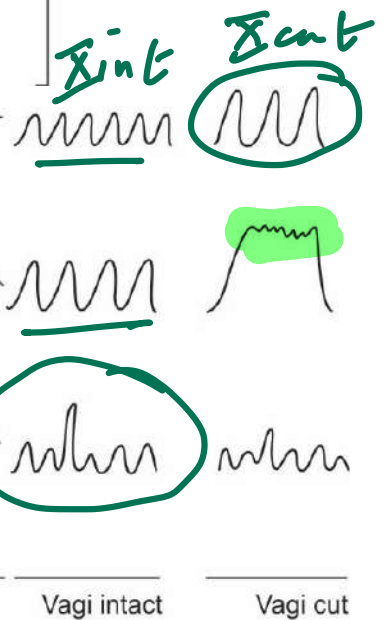
*Histotoxic anoxia*

→ Fe<sup>3+</sup> : Dapsone / Prilocaine<sup>®</sup> / Benzocaine / Nitrites

# RESPIRATORY REGULATION



Transection	Effect
ABOVE PONS	X intact: vol control X cut: ↑ depth
MID-PONS	X intact: ↑ depth X cut: apneusis
PONS-MEDULLA	irregular
BELOW MEDULLA	dead ceases



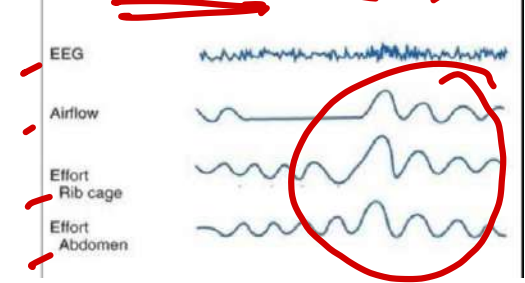
Ondine's curse

≡ Cortical control of breathing

PaO2 normal during day : SLEEP APNEA

Central (CSA)  
Brain dysfunction

Obstructive (OSA)



Obesity BMI > 30  
Hypoventilation

PaCo2 high during day and night

Obesity-Hypoventil<sup>n</sup> = Pickwickian S<sub>x</sub>

# RESPIRATORY FAILURE

TYPE 1  
HYPOXEMIC

$PO_2 \downarrow$

ILD / p. edema

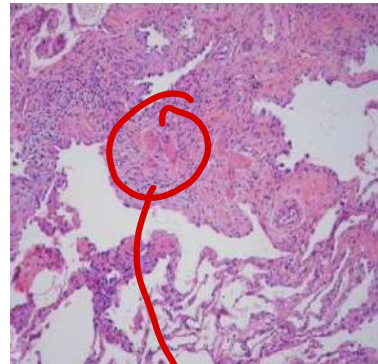
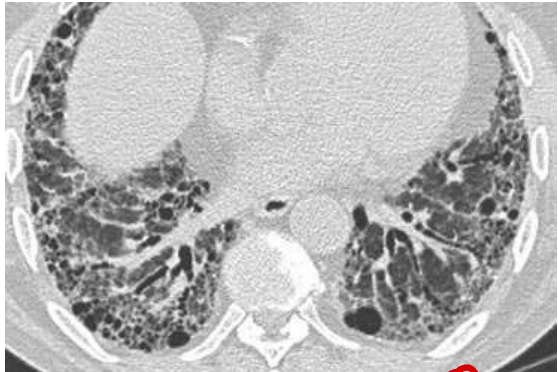
TYPE 2  
HYPERCAPNIC

$PCO_2 \uparrow$

Hypoventil<sup>n</sup>  
OSA / CSA /  
COPD

TYPE 3:  
PERI-OPERATIVE

TYPE 4  
SHOCK WITH  
HYPOPERFUSION



Pirfenidone: anti TGF $\beta$

Nintedanib: anti PDGF

Honey combing (+)

VIP = IPF

Steroids xx

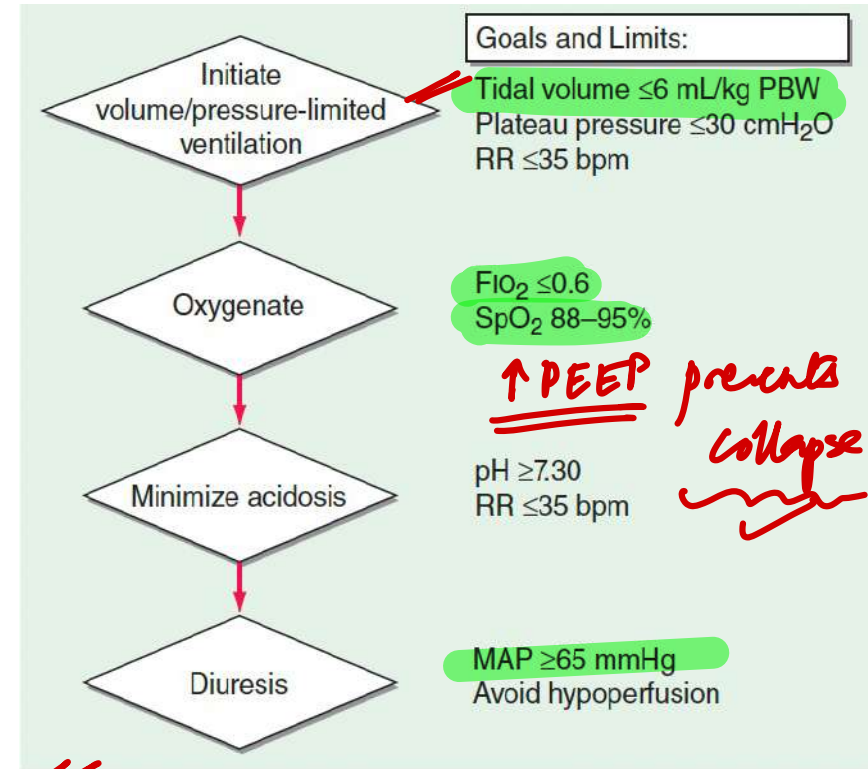
fibroblastic  
foci  
temporal / spatial  
heterogeneity

# ARDS

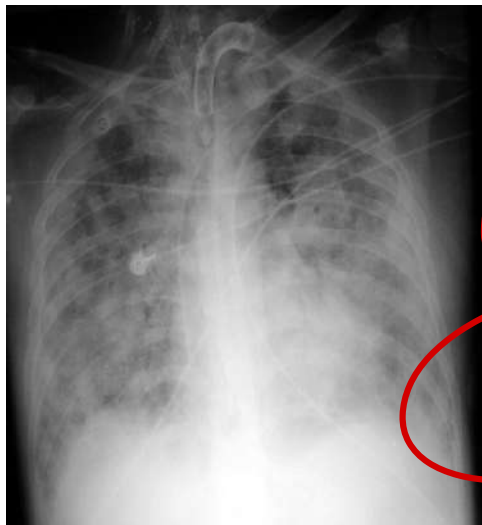
: sepsis / pancreatitis / toxins

Berlin's

DIAGNOSTIC CRITERIA FOR ARDS			
<b>SEVERITY: OXYGENATION</b>	<b>ONSET</b>	<b>CHEST RADIOGRAPH</b>	<b>ABSENCE OF LEFT ARTRIAL HYPERTENSION</b>
PAO2/ FI02 < 300 Mild: 200-300 Moderate: 150-200 Severe: < 100	Acute: within 1 week of clinical insult or new or worsening respiratory symptoms	Bilateral opacities consistent with pulmonary edema not fully explain by effusions, lobar/lung collapse or nodules	PCWP (N)



## PRONE VENTILATION

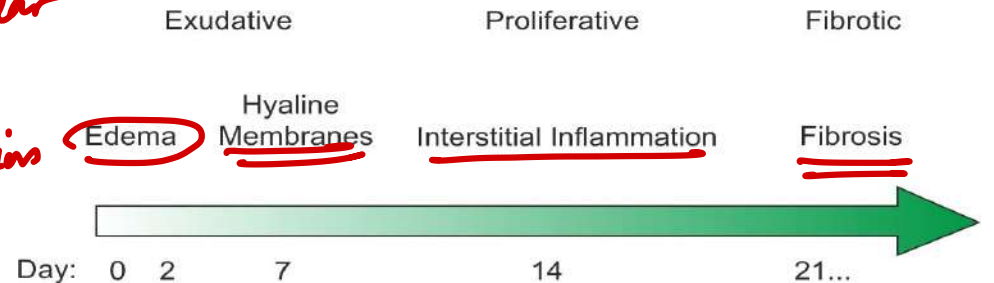


peripheral patchy  
⊗ pl effusions  
NL p. edema  
ARDS & //



bathing (+) & megal  
p. edema  
Kerley lines  
interlobular septal  
pl effusions (+)

Edema  
Hyaline Membranes  
Interstitial Inflammation  
Fibrosis



# PLEURAL EFFUSION

## Light's criteria for pleural effusions

	Transudate	Exudate
Protein (pleural/ serum)	< 0.5	>0.5
LDH (Pleural/serum)	<0.6	>0.6
	Pleural LDH $\leq$ two-thirds upper limit of normal serum LDH	Pleural LDH > two-thirds upper limit of normal serum LDH
Causes	. Cirrhosis / CHF / NS	Inflamm <sup>n</sup> / malignancy

Low glucose: 'REM'

(RA) Empyema Malignancy

High Amylase: Malignancy  
Pancreatitis / Esophageal rupture

Indications of drainage of effusion:

1. Split pleura sign
2. pH < 7.2
3. Glc ↓
4. Gs / culture (+)
5. Loculations



### CHYLOTHORAX

- lymphatic leaks
- Tg > 110mg/dL

Pseudo-chylothorax: RA

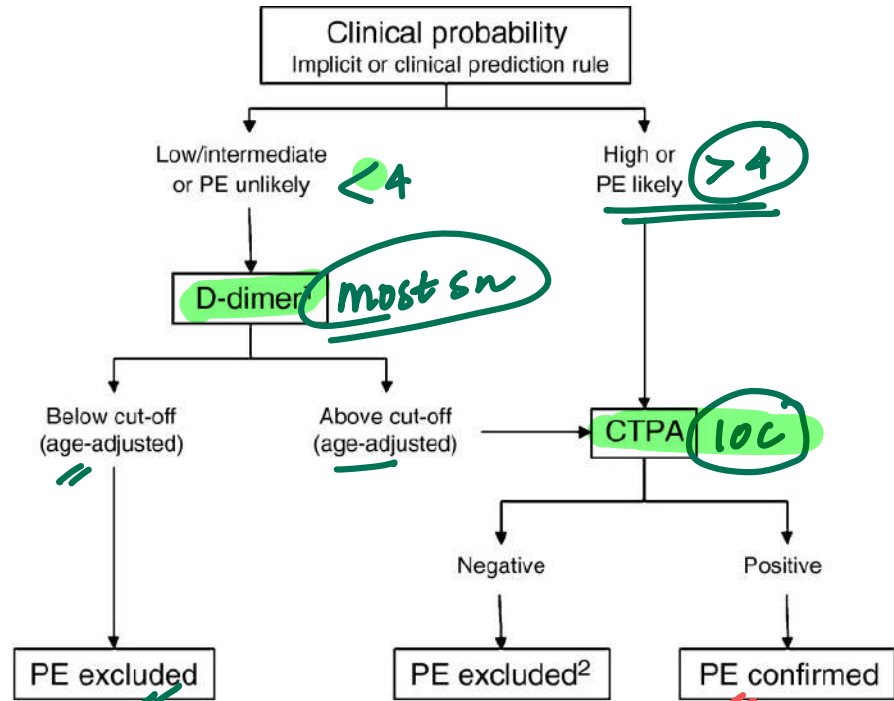
Cholesterol



Empyema  
Pigtail

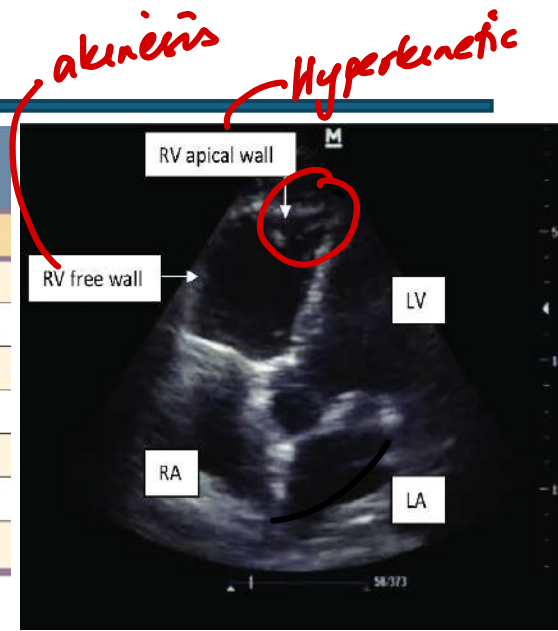
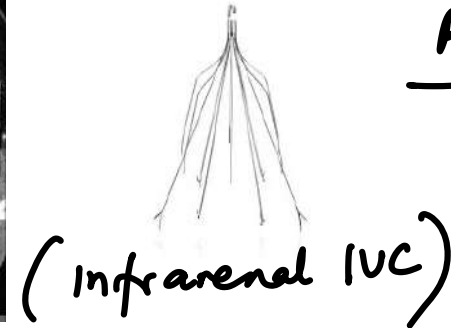
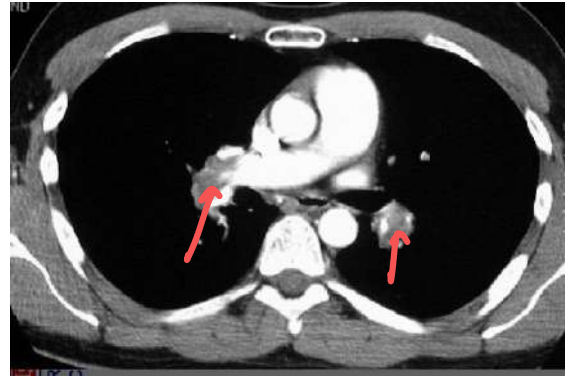
Split pleura sign

# Pulmonary embolism



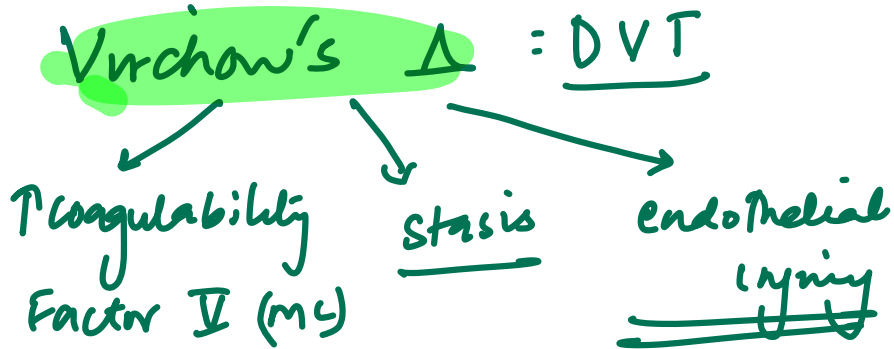
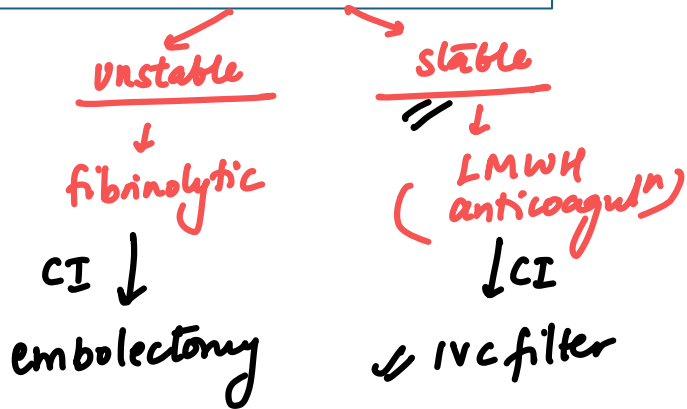
High Clinical Likelihood of Pulmonary Embolism (PE) if Point Score Exceeds 4

CLINICAL VARIABLE	WELL'S SCORE	PE SCORE
Signs and symptoms of DVT		3.0
Alternative diagnosis less likely than PE		3.0
Heart rate >100/min		1.5
Immobilization >3 days; surgery within 4 weeks		1.5
Prior PE or DVT		1.5
Hemoptysis		1.0
Cancer		1.0



RV strain =  
McConnell sign

Treatment algorithm:

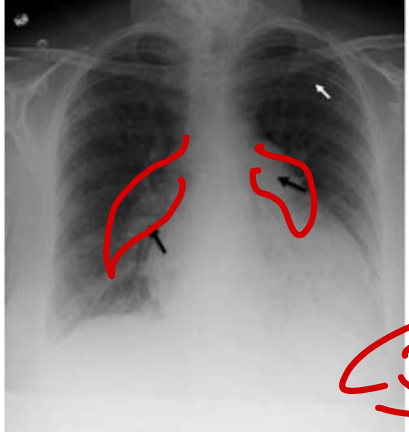
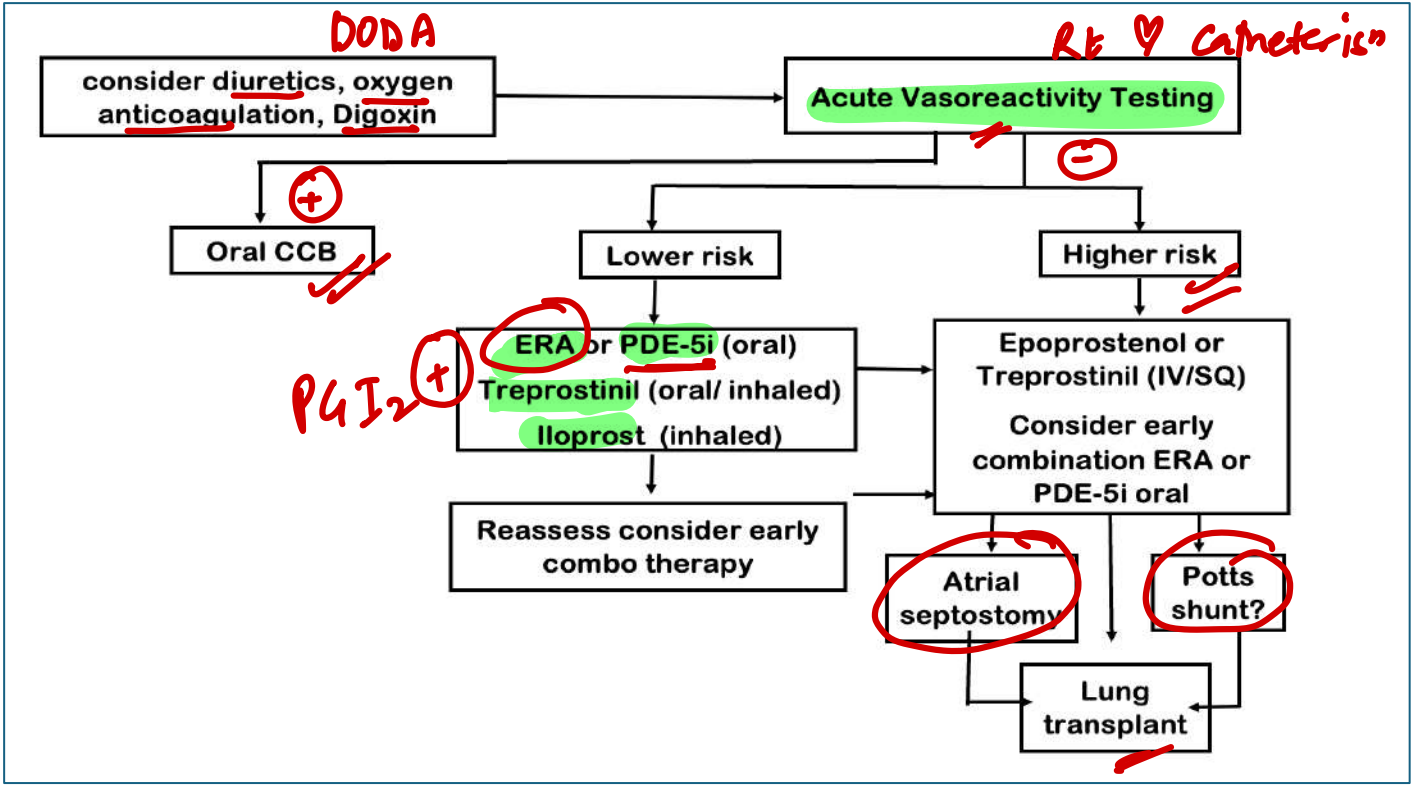


ECG  
MC: sinus tachy  
most sp: S1 QIII TIII  
Xray specific:  
• Westermarck-oligenin  
• Hampton's hump  
• Palla's: PA prominent

# PAH

*Dana Point*

GROUP	DESCRIPTION
1.	<p><b>Idiopathic</b></p> <p><b>Hereditary: BMP2</b></p> <p>Associated with</p> <ul style="list-style-type: none"> <li>Connective tissue diseases</li> <li>HIV infection</li> <li>Portal Hypertension</li> <li>Congenital heart disease</li> <li>Schistosomiasis</li> <li>Chronic haemolytic anaemia</li> </ul>
2.	PAH owing to <b>left heart disease</b>
3.	PAH owing to <b>lung disease and /or hypoxia</b> <i>CREST (mc)</i>
4.	Chronic thromboembolic pulmonary hypertension <i>plexiform arteriopathy</i>
5.	PH with <b>unclear</b> multifactorial mechanism



*Jug handle*

- sle: hepatotoxic*
- ERA: Bosentan, Macitentan, Ambrisentan**
- PDE5i: Sildenafil, Tadalafil** : *(\*) use = nitrates*
- Guanylate cyclase +: Riociguat**

# Chest infections

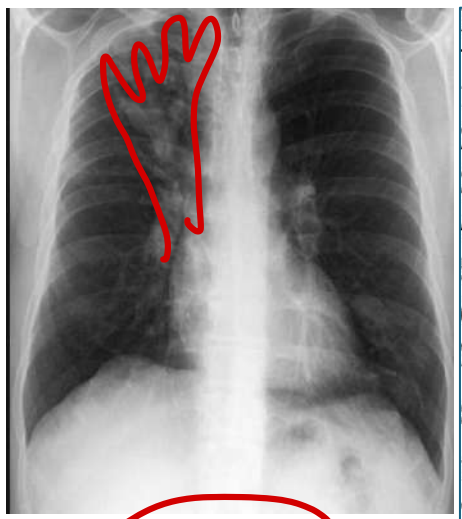
MCC of typical CAP: *S. pneumoniae*  
 MCC of atypical CAP: *Mycoplasma*  
 MCC of HAP (>48hrs of admission): *Gram -ve*  
 MCC of VAP (>48hrs of ventilation):

early: *S. pneumoniae* late: *Pseudomonas*  
*Acinetobacter*

CURB-65 Scoring	
Symptom	Points
Confusion	1
Urea: BUN > 19 mg/dL (> 7 mmol/L)	1
Respiratory rate $\geq$ 30 breaths /min	1
Systolic BP < 90 mm Hg or diastolic BP $\leq$ 60 mm Hg	1
Age $\geq$ 65 years	1

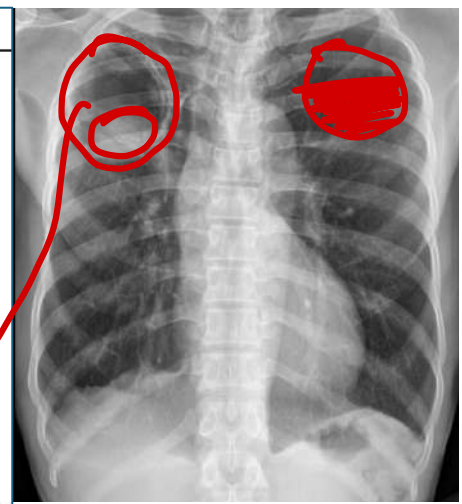
OP: 0-1	Amoxicillin +/- Macrolides / Doxycycline
IP: 2	FQ / BL+ Azithral/Doxy
ICU: 3-5	BL+ FQ/ Azithral

OP: 0  
 IP: 1-2  
 ICU: 3-4



ABPA  
 Hand-in-glove

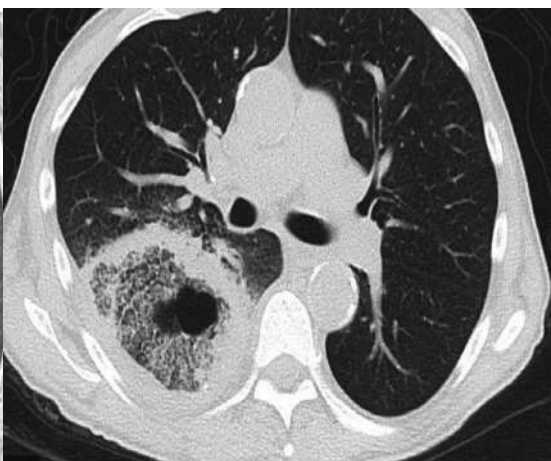
- Primary criteria (1-6 suggestive, +7 definite)
- Episodic bronchial obstruction
  - Peripheral eosinophilia
  - Positive immediate skin test to *Aspergillus*
  - Positive precipitin test to *Aspergillus*
  - Increased total serum IgE
  - History of transient or fixed lung infiltrates
  - Proximal bronchiectasis
- Secondary (supportive) criteria
- Brown plugs/flecks in sputum
  - Positive late (6-12 h/ Arthus) skin test to *Aspergillus*



Fungal ball = Aspergilloma  
 Rx: Symptomatic  
 Monrad

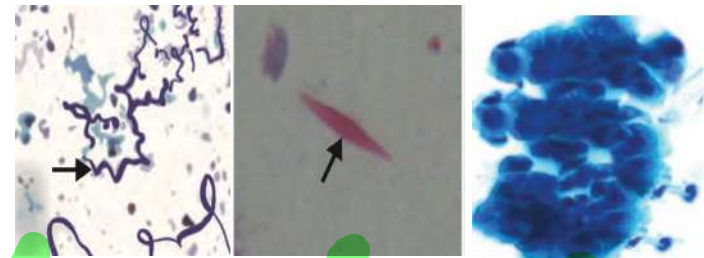


ic / neutropenia  
 Halo sign  
 Rx: VORICONAZOLE



Reverse halo / Bird's nest  
 Rx: MUCOR

# Asthma

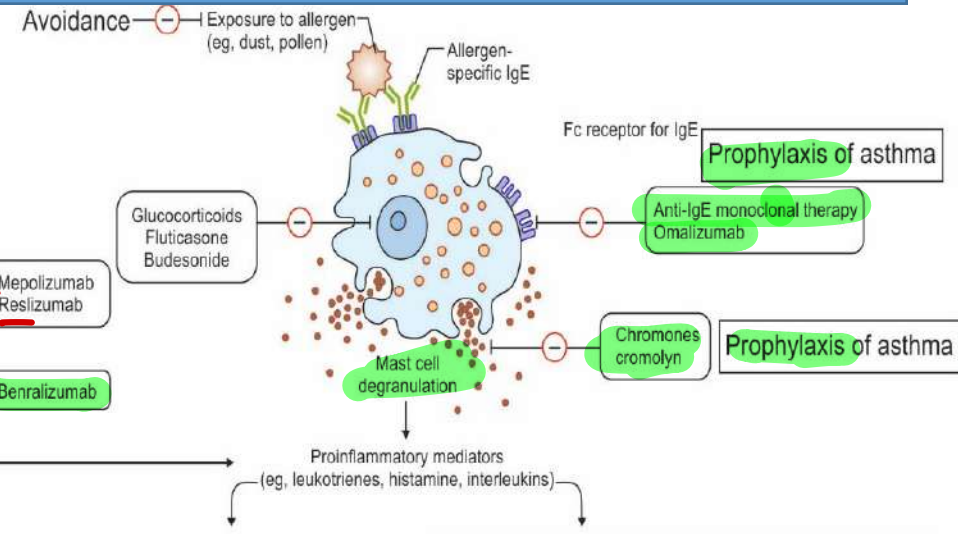


**Craschmann spirals**  
**C-L crystals**  
**Creola bodies**  
 galectin-10 (epithelium)

**GINA**

**IL-4 inhibitor: Dupilumab**  
**IL-13 inhibitor: Tralokinumab, Lebrikizumab**

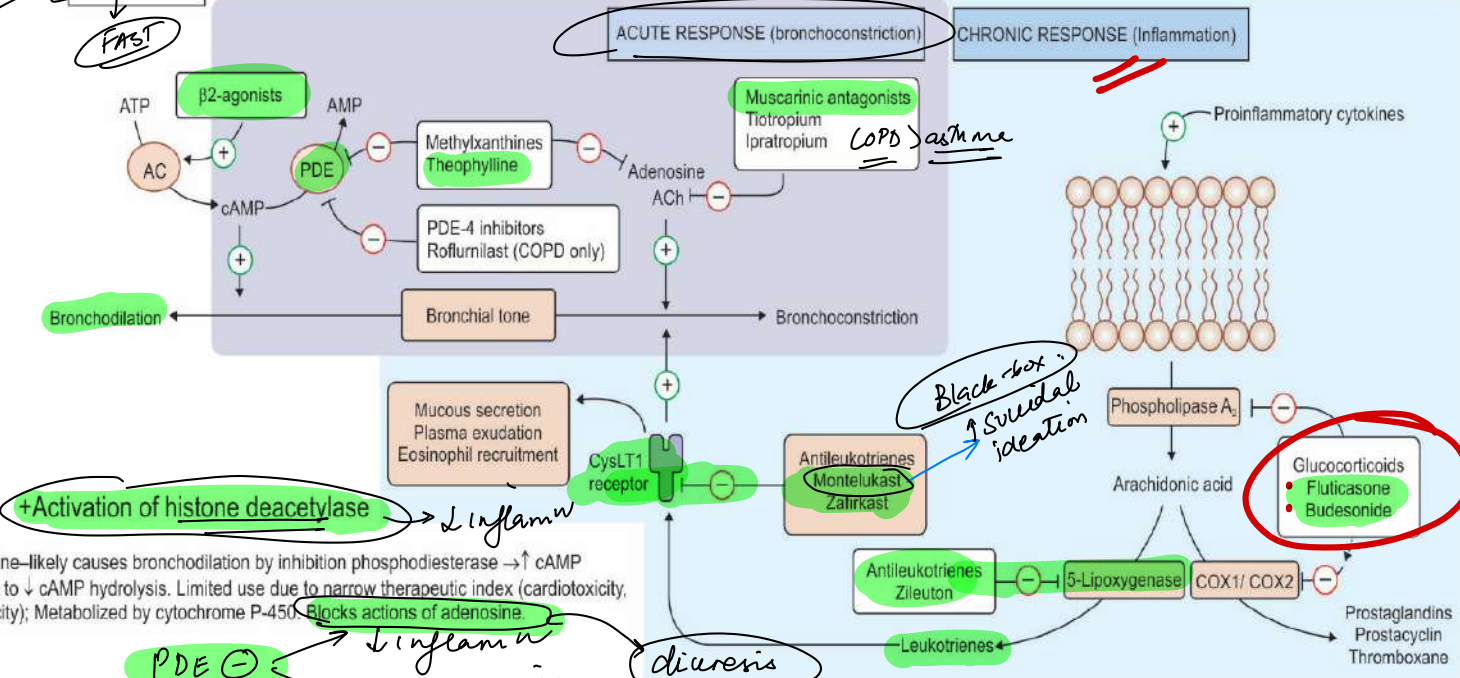
**Th2**  
**IgE**  
**IL-4, 5, 13**



**SABA**  
**LABA**  
**FAST**

Salbutamol  
 Terbutaline  
 Salmeterol  
 Formoterol

**IL-5Ra**  
 Eosinophil



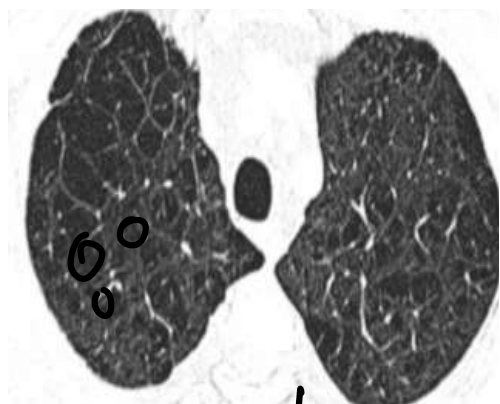
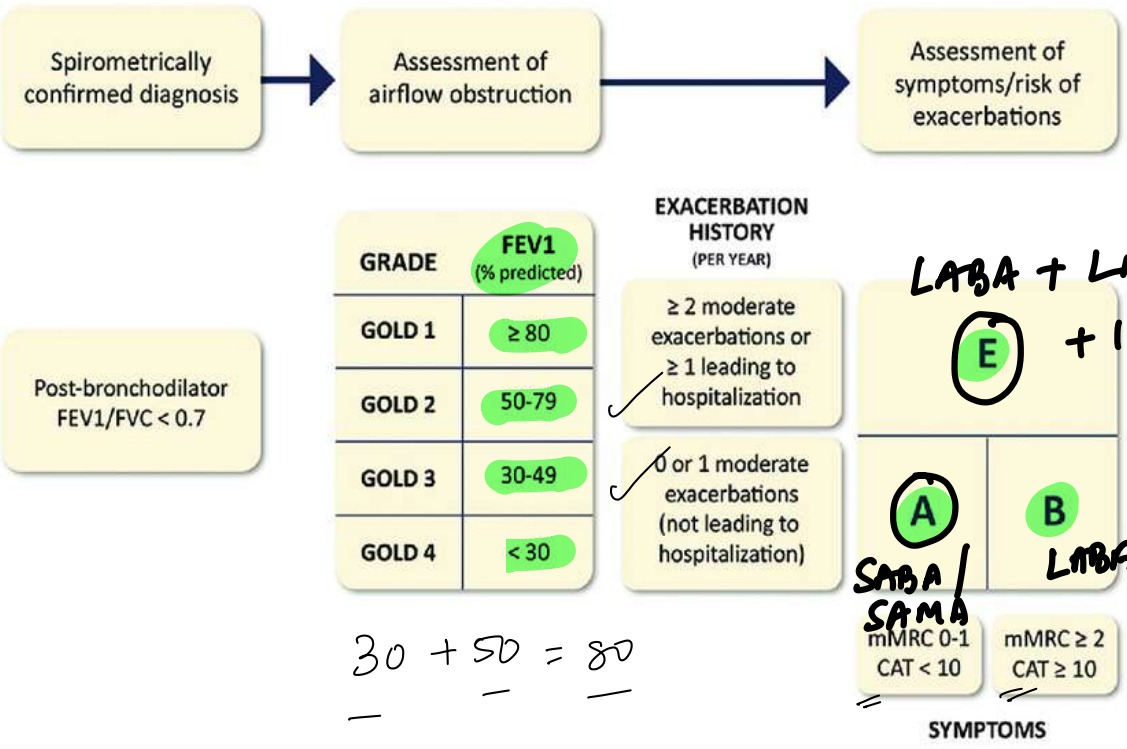
Theophylline—likely causes bronchodilation by inhibition phosphodiesterase → ↑ cAMP levels due to ↓ cAMP hydrolysis. Limited use due to narrow therapeutic index (cardiotoxicity, neurotoxicity); Metabolized by cytochrome P-450. **Blocks actions of adenosine**

**+Activation of histone deacetylase** → ↓ Inflammation  
**PDE ⊖** → ↓ Inflammation  
 ↓ bronchodilation  
 s/e: arrhythmia

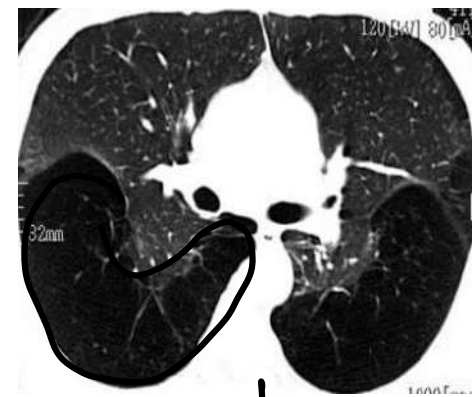
**Step 1,2: Symptoms <5d/week:**  
**LD-ICS-Formoterol as needed**  
**Step 3: Most days/ >1/week nighttime:**  
**LD-ICS-Formoterol**  
**Step 4: Low lung function:**  
**MD-ICS-Formoterol**  
**Step 5: Add LAMA + HD-ICS + Anti-IgE/IL5**

# COPD-GOLD Criteria-2023

## GOLD ABE Assessment Tool



LABA + LAMA + ICS  
**E**  
 - Smokers  
 - v/v  
Centrilobular



α<sub>1</sub>AT def  
**LL**  
Panlobular



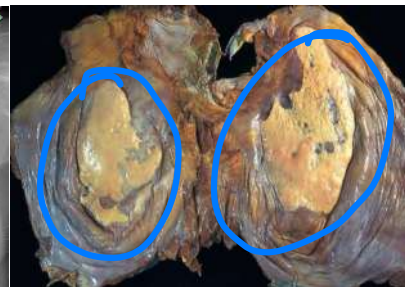
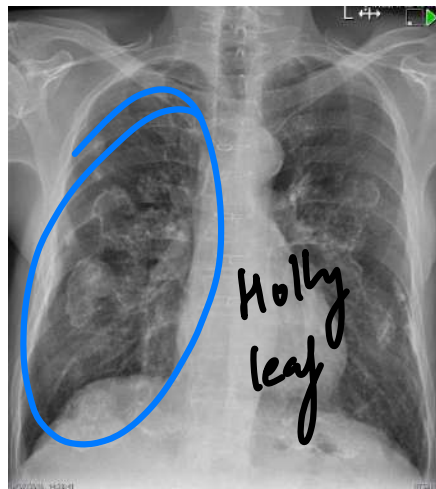
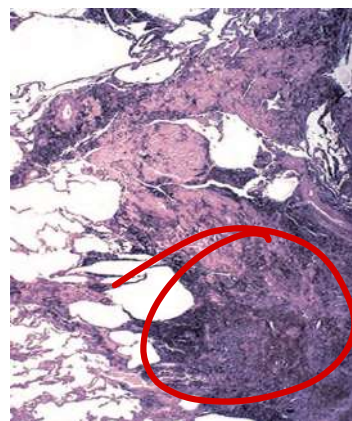
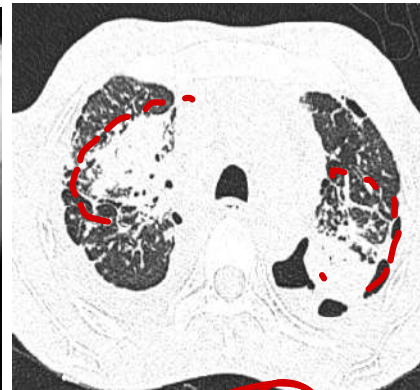
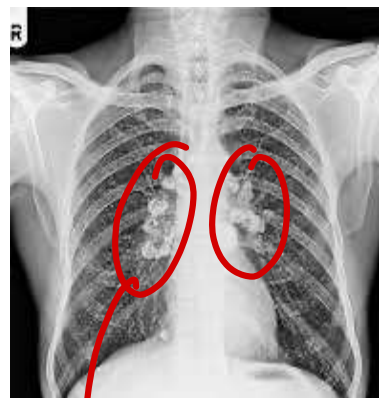
Paraseptal  
mc: Smokers

Reid Index =  $\frac{\text{Mucous gland}}{\text{BM-cartilage}}$

Chronic bronchitis : >40%

# OCCUPATIONAL LUNG DISEASES

V/L exc Asbestosis



**SILICOSIS**

**PMF**

egg shell  $Ca^{2+}$

progressive massive fibrosis

Caplan syndrome

CWP + RA

Erasmus syndrome

CWP + SSc

**CWP**

Pleural plaques  $Ca^{2+}$

Serpentine/ Crysolite  
Amphibole/ Crocidolite

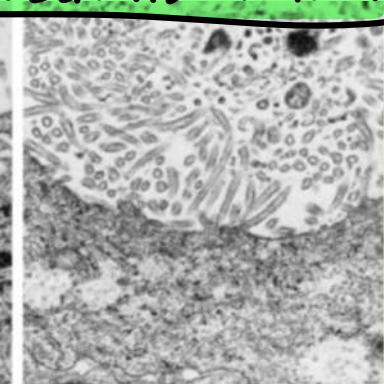
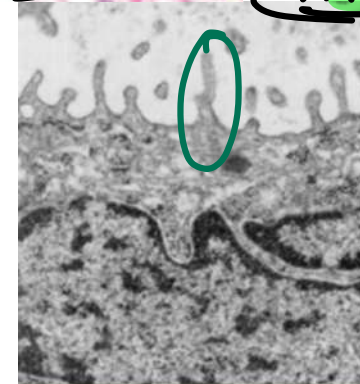


Dumbbell

mc: adeno ca

most sp: **Mesothelioma**

**FROZEN HEMITHORAX**



Short, stubby  
microvilli  
**adca**

TTF1 / Napsin A

long, slender  
microvilli  
**Mesothelioma**

Calretinin / CK5/6

**Berylliosis**

Jewellery / dental alloy

Granulomatosis

**BeLPT**

HLA-DP

mimics  
Sarcoidosis

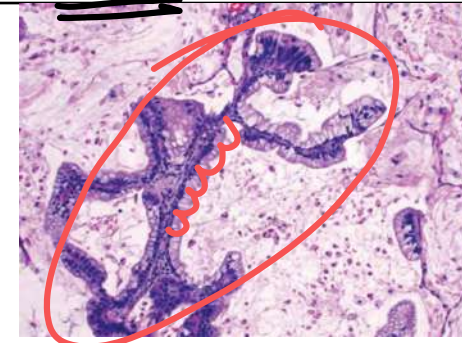
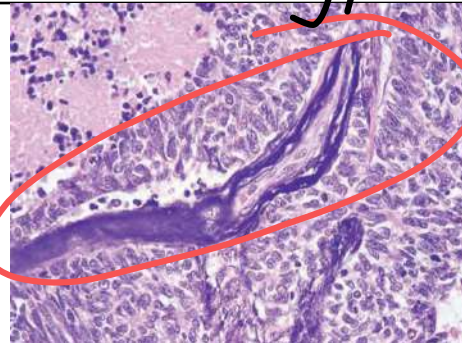
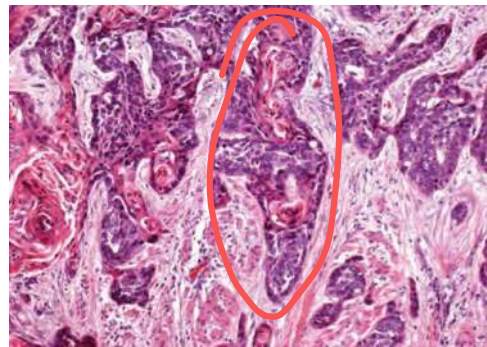
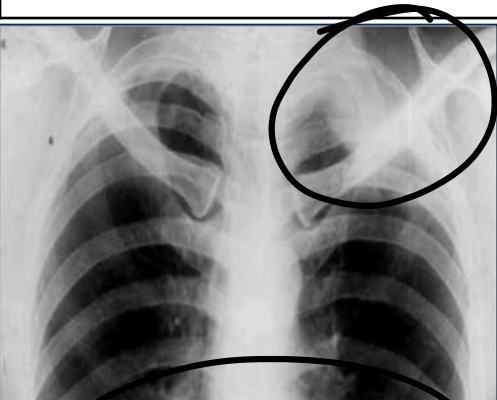
**Cobalt**

Diamond industry

Giant cell interstitial pneumonitis

# CARCINOMA LUNG

	Squamous cell ca	Small cell ca	Adenoca (mc)	Large cell ca
Smoker?	✓	✓	⊖	
Location?	Central	central	peripheral	
GENES	p53	L-myc	EGFR / ALK / KRAS	-
H/P	Keratin pearls	Azzopardi / salt & pepper chromatin	lepidic spread	
IHC	p40 / p63	Synaptophysin / Chromogranin NSE / Bombesin	TTF1 / Napsin A	
Paraneoplastic	↑Ca <sup>2+</sup> : PTHrp	SIADH / Cushing / LEMS	Trousseau Sr / HPOA	Gynecomastia

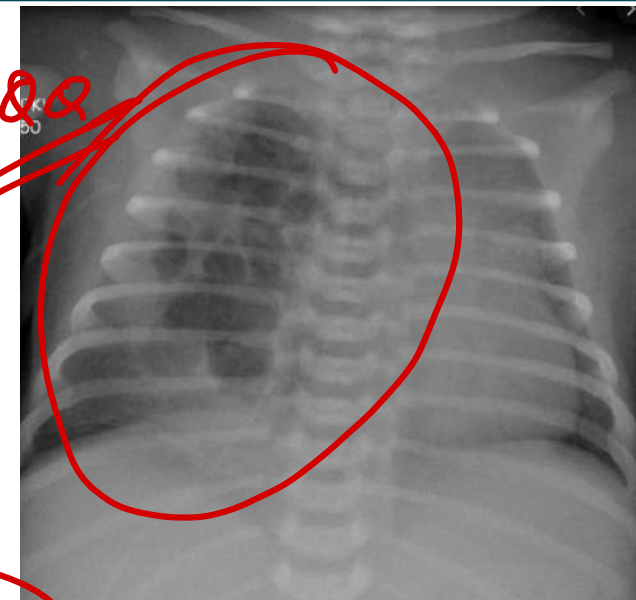
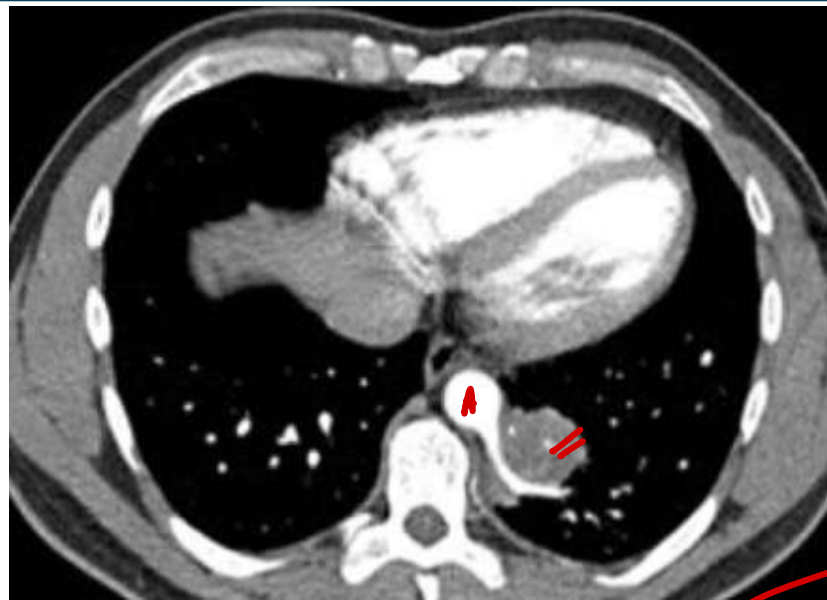
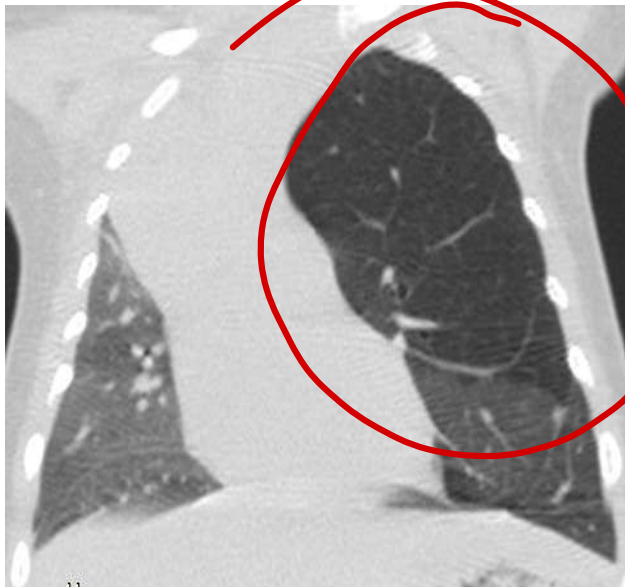


PANCOAST

→ Sq cc > small

IOC: MRI

# Congenital pulmonary malformations



Cong Lobar  
overinflation

CLO

MC: LVL

Sequestration

EXTRALOBAR

Own visceral pleura

Systemic venous drainage

Foregut communication, CDH

Presentation: Perinatal or infancy

INTRALOBAR

Shares pleura with parent lobe

Pulmonary venous drainage

Associated anomalies rare

Adolescence

systemic arterial

CPAM

Cong pulmar  
airway malform<sup>n</sup>