

RESPIRATORY SYSTEM

**RESPIRATORY FUNCTIONS
MECHANICS OF RESPIRATORY SYSTEM
GAS TRANSPORT**

STEPS IN THE DELIVERY OF O_2 TO THE CELLS

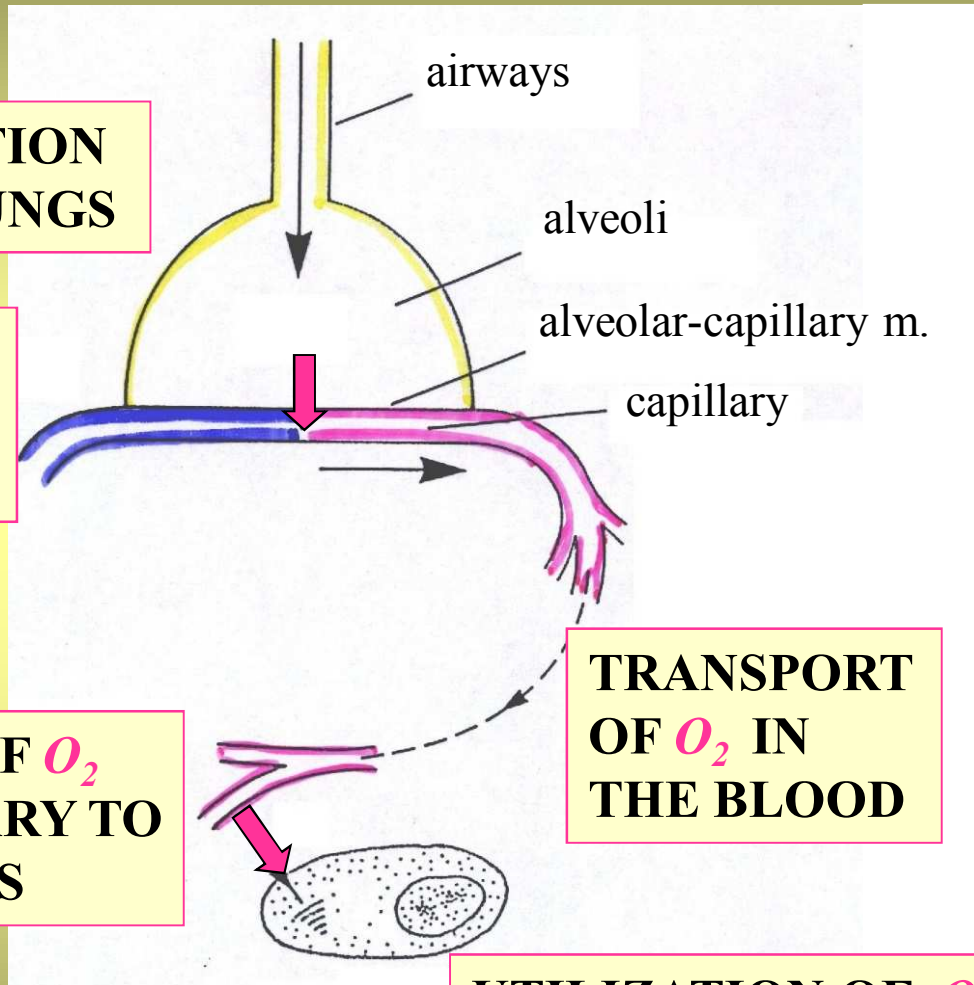
VENTILATION
OF THE LUNGS

DIFFUSION OF O_2 ACROSS
ALVEOLAR-CAPILLARY
MEMBRANE

DIFFUSION OF O_2
FROM CAPILLARY TO
THE CELLS

TRANSPORT
OF O_2 IN
THE BLOOD

UTILIZATION OF O_2
BY MITOCHONDRIA




AT REST

O_2 UPTAKE ~ 300 ml / min

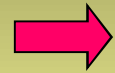
CO_2 OUTPUT ~ 250 ml / min

INTERNAL RESPIRATION

- 
- I PHYSIOLOGY OF AIR PASSAGES**
 - II BASIC MEASURABLE PARAMETERS**
 - III ACTIVE AND PASSIVE FORCES**
 - IV COMPOSITION OF ALVEOLAR AIR**
 - V ALVEOLAR-CAPILLARY MEMBRANE**
 - VI TRANSPORT OF O_2 AND CO_2 IN THE BLOOD**

AIR PASSAGES

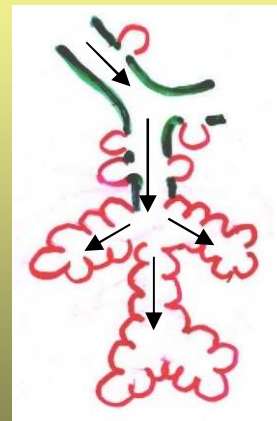
ANATOMICAL DEAD SPACE – **CONDUCTING ZONE**



- **NASAL PASSAGES**
- **PHARYNX**
- **LARYNX**
- **TRACHEA**
- **BRONCHI**
- **BRONCHIOLES**
- **TERMINAL BRONCHIOLES**

Other physiological functions:

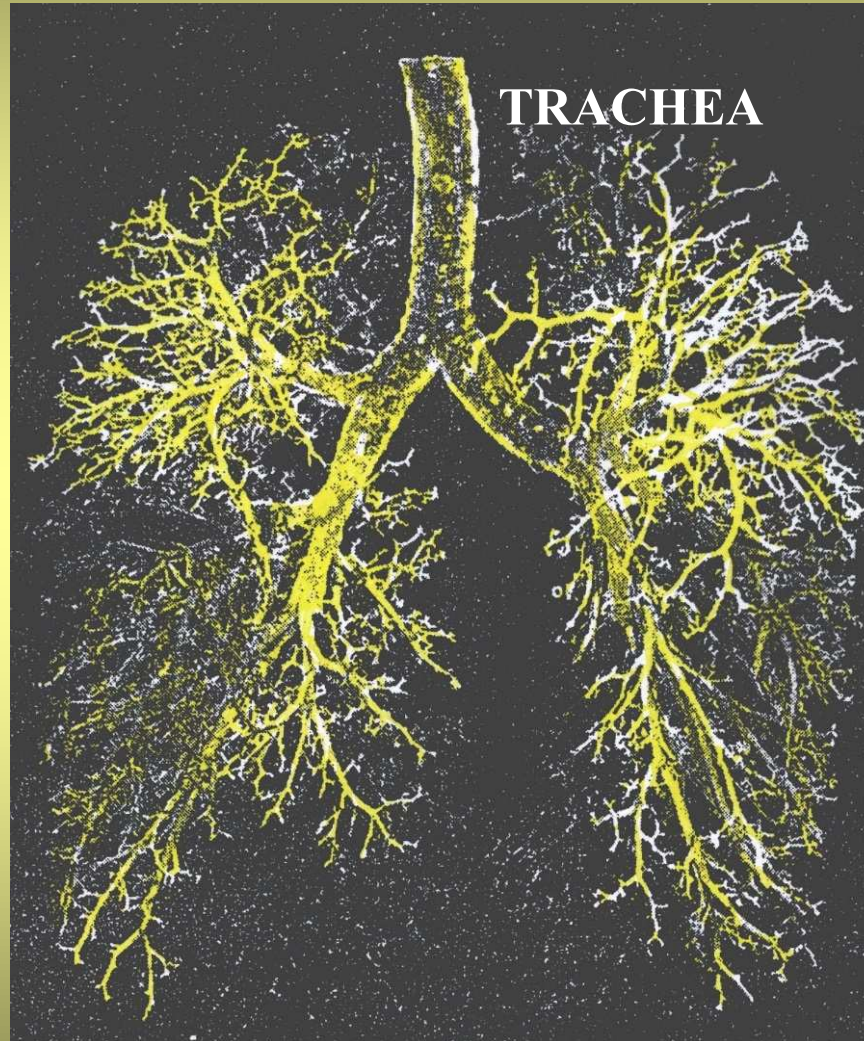
- air is warmed, cleaned and takes up water vapour
- respiratory reflex responses to the irritants
- speech and singing (function of larynx)



RESPIRATORY ZONE (GAS EXCHANGE)

Total alveolar area $\sim 100 \text{ m}^2$

CAST OF HUMAN AIR PASSAGES



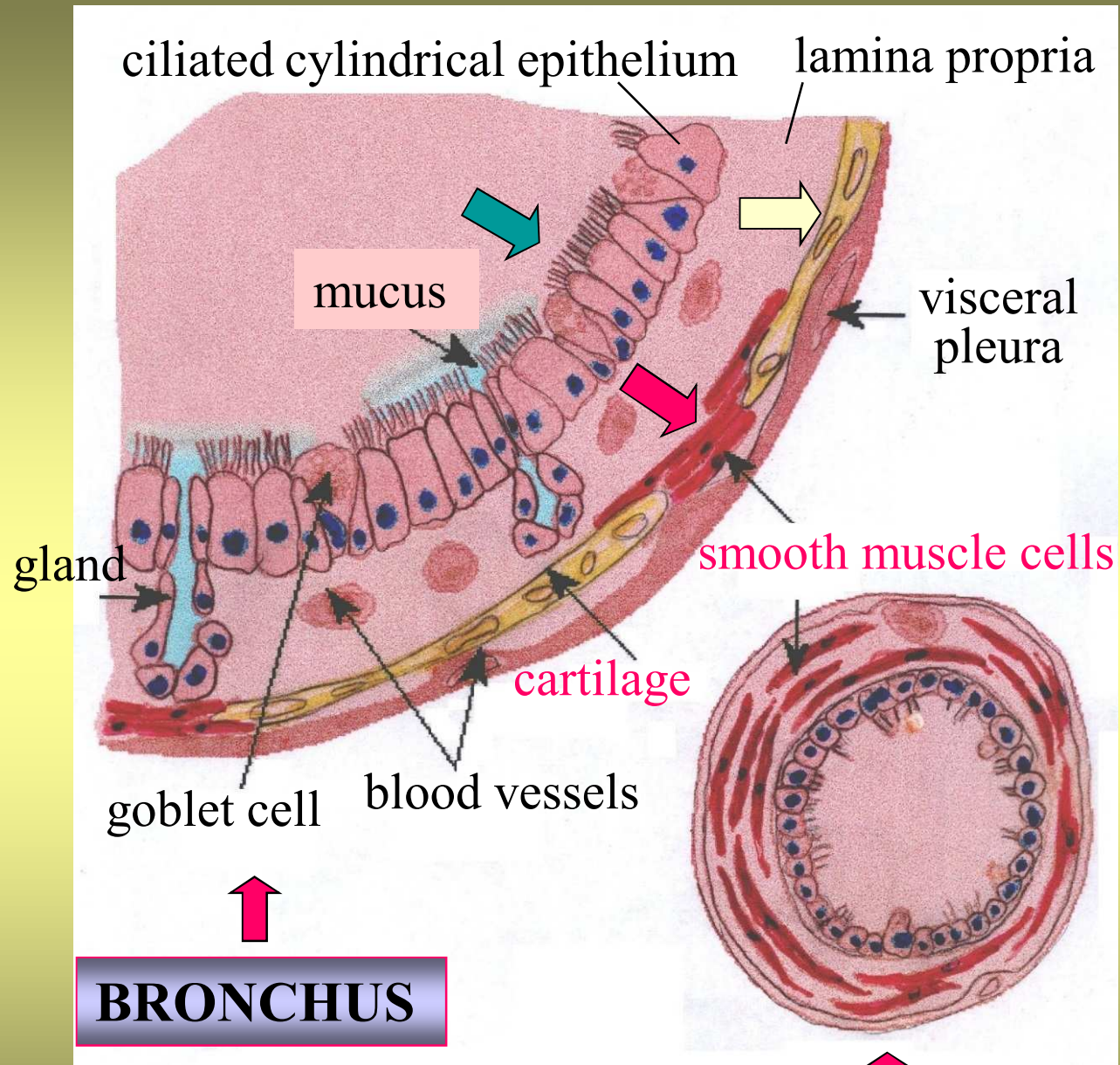
TRACHEA

BRONCHI

BRONCHIOLES

TERMINAL
BRONCHIOLES

AERODYNAMIC RESISTANCE



**AUTONOMIC
INNERVATION of
smooth muscle cells**

Muscarinic receptors:
Acetylcholine activates
bronchoconstriction

β-adrenergic receptors:
Noradrenaline activates
bronchodilatation

TERMINAL BRONCHIOLE

∅ < 1 mm

I AIR PASSAGES

➔ II MEASURABLE PARAMETERS

- DEAD SPACE
- LUNG VOLUMES
- FUNCTIONAL INVESTIGATION
- CHARACTERISTIC PRESSURES

III ACTIVE AND PASSIVE FORCES

- RESPIRATORY MUSCLES
- LUNGS ELASTICITY
- COMPLIANCE
- WORK OF BREATHING

IV COMPOSITION OF ALVEOLAR AIR

V ALVEOLAR-CAPILLARY MEMBRANE

VI TRANSPORT OF GASSES (O_2 and CO_2)

V_T tidal volume ~ 500 ml

$$V_T = V_A + V_D$$

V_A part of tidal volume entering alveoli ~ 350 ml

V_D part of tidal volume remaining in the dead space ~ 150 ml

$f = 12/\text{min}$

$$\dot{V} = V_T \times f$$

**PULMONARY
MINUTE
VENTILATION**

6 l/min

$$\dot{V}_A = V_A \times f$$

ALVEOLAR VENTILATION

4.2 l/min

$$\dot{V}_D = V_D \times f$$

DEAD SPACE VENTILATION

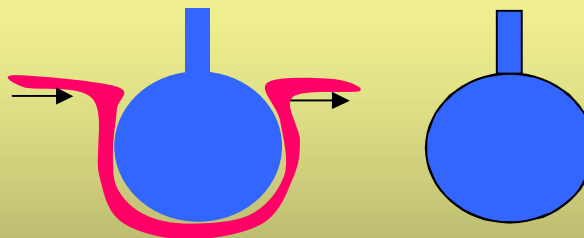
1.8 l/min

DEAD SPACE

**TOTAL GAS VOLUME NOT EQUILIBRATED WITH BLOOD
(without exchange of gasses)**

- **ANATOMICAL** dead space - volume of air passages
- **FUNCTIONAL (total)** dead space

ANATOMICAL dead space + total **VOLUME** of **ALVEOLI** without functional capillary bed



**IN HEALTHY INDIVIDUALS
both spaces are practically identical**

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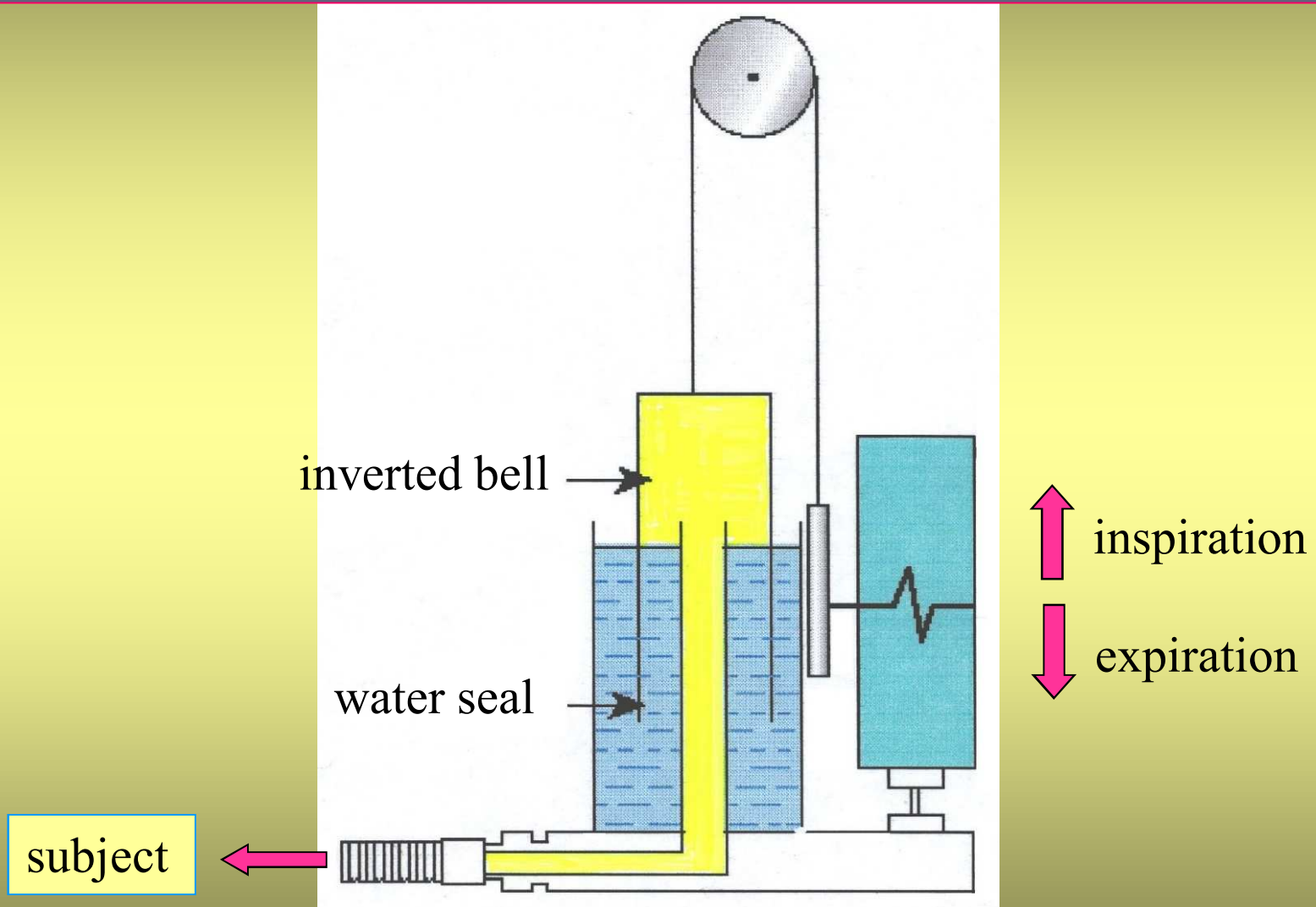
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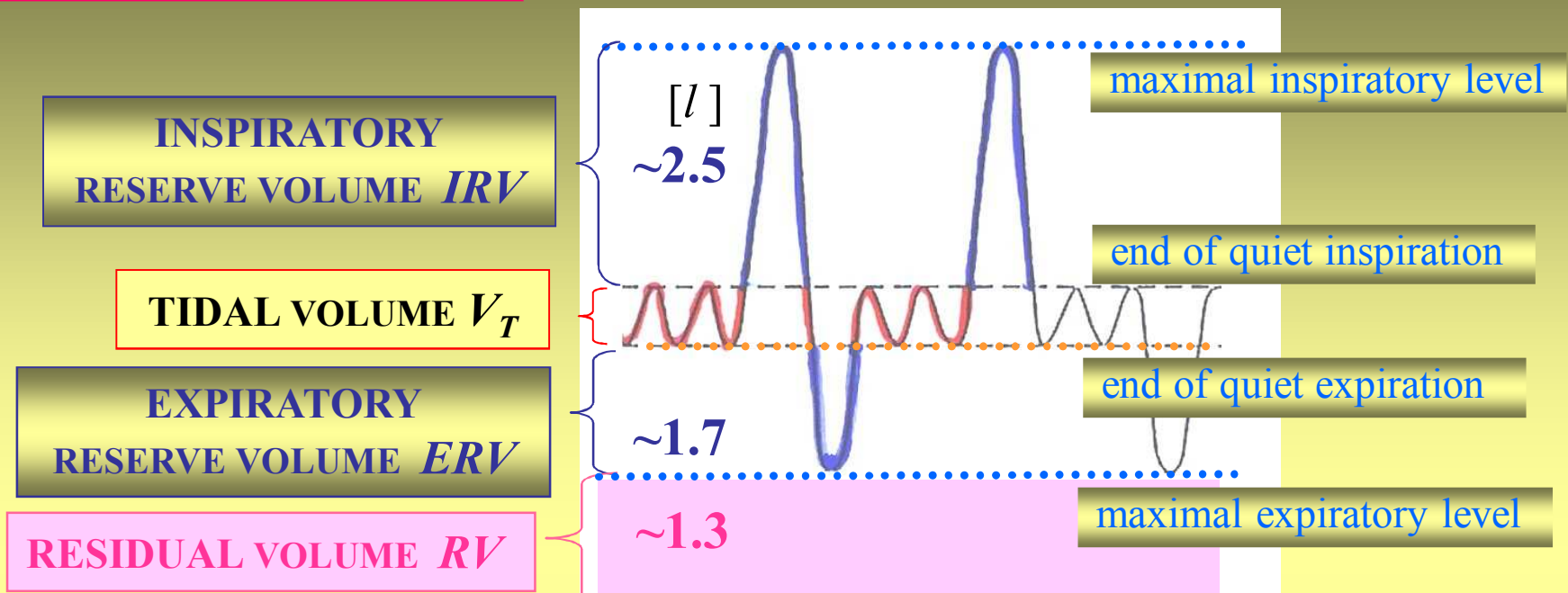
VI TRANSPORT OF GASSES (O_2 and CO_2)

SPIROMETRY

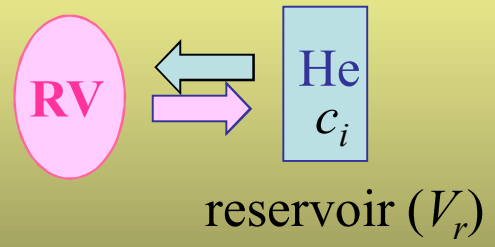
(measurements of lung volumes, capacities, functional investigations, ...)



LUNG VOLUMES



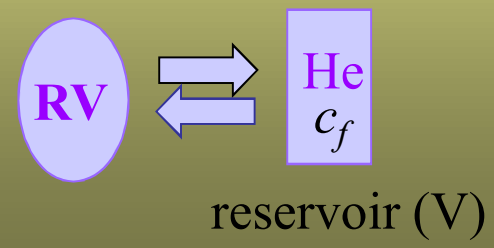
DILUTION METHOD *He*



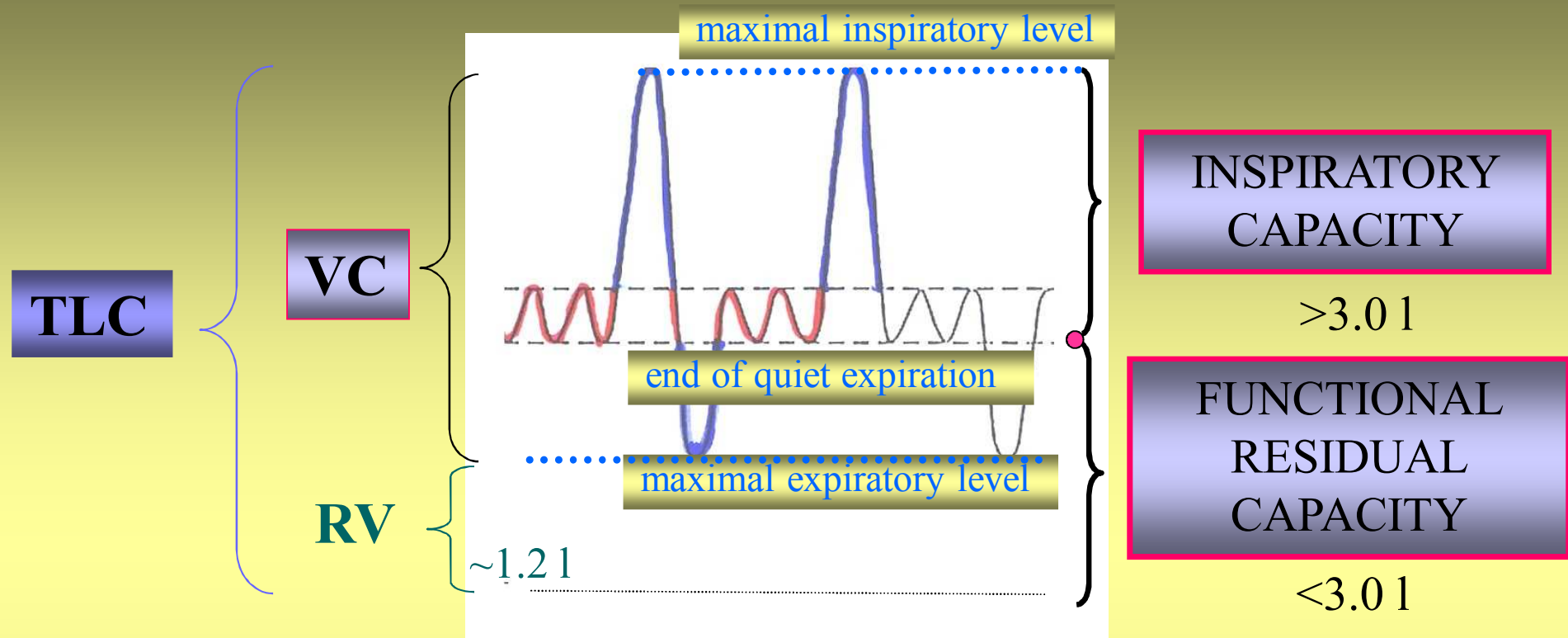
Principle of method: **1** Maximal expiration, **2** Repeated inspiration from and expiration into a reservoir (known volume V_r) with inert gas He (known concentration c_i)

⇒ Equilibration of the air in the residual volume and reservoir

3 Calculation of **residual volume RV** from the initial and final He concentrations in reservoir (c_i, c_f).



$$RV = V_r \frac{c_{iHe} - c_{fHe}}{c_{fHe}}$$



VC **VITAL CAPACITY = $V_T + IRV + ERV$** ~ 4.7 l

VC - the largest amount of air that can be expired after maximal inspiration

TLC **TOTAL LUNG CAPACITY = VC + RV** ~ 6.0 l

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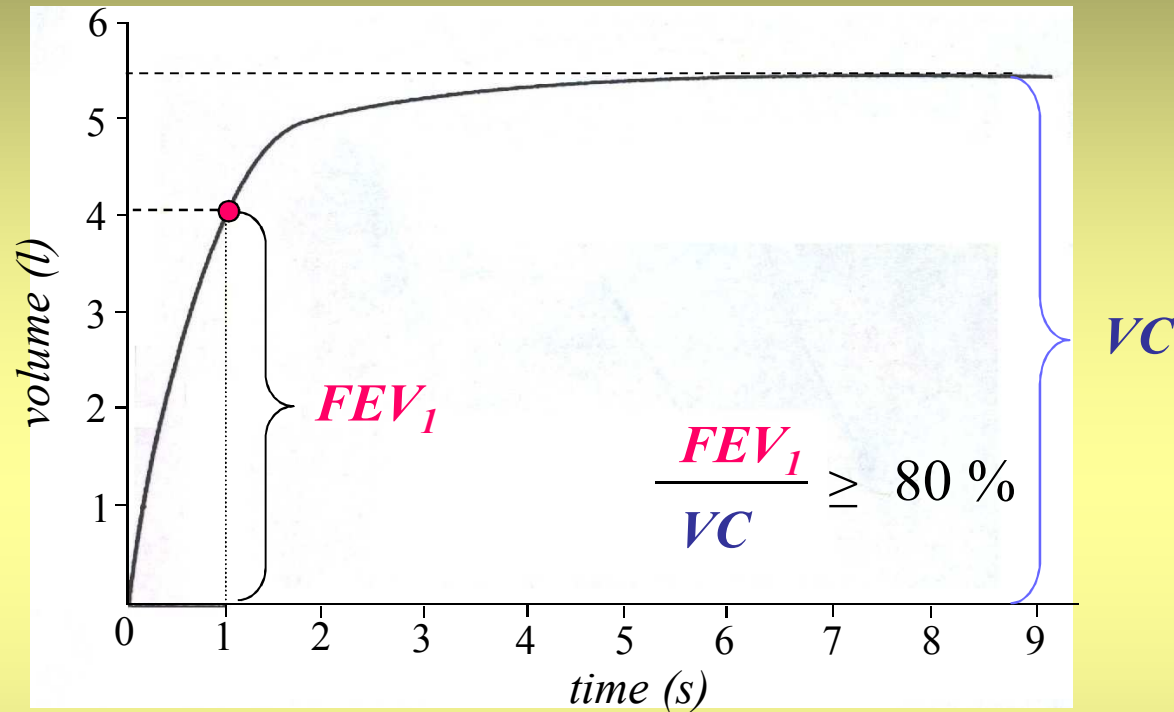
IV COMPOSITION OF ALVEOLAR AIR

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VI TRANSPORT OF GASSES (O_2 and CO_2)

FUNCTIONAL INVESTIGATION OF THE LUNGS

- **TIMED VITAL CAPACITY (FEV_1 - forced expiratory volume per 1 s)**



- **PULMONARY MINUTE VENTILATION RMV (respiratory minute volume) at rest** ($0.5 \text{ l} \times 12 \text{ breathes/min} = 6 \text{ l/min}$)
- **MAXIMAL VOLUNTARY VENTILATION (MVV)** (125-170 l/min)
- **PEAK EXPIRATORY FLOW RATE ($PEFR$)** ($\sim 10 \text{ l/s}$)

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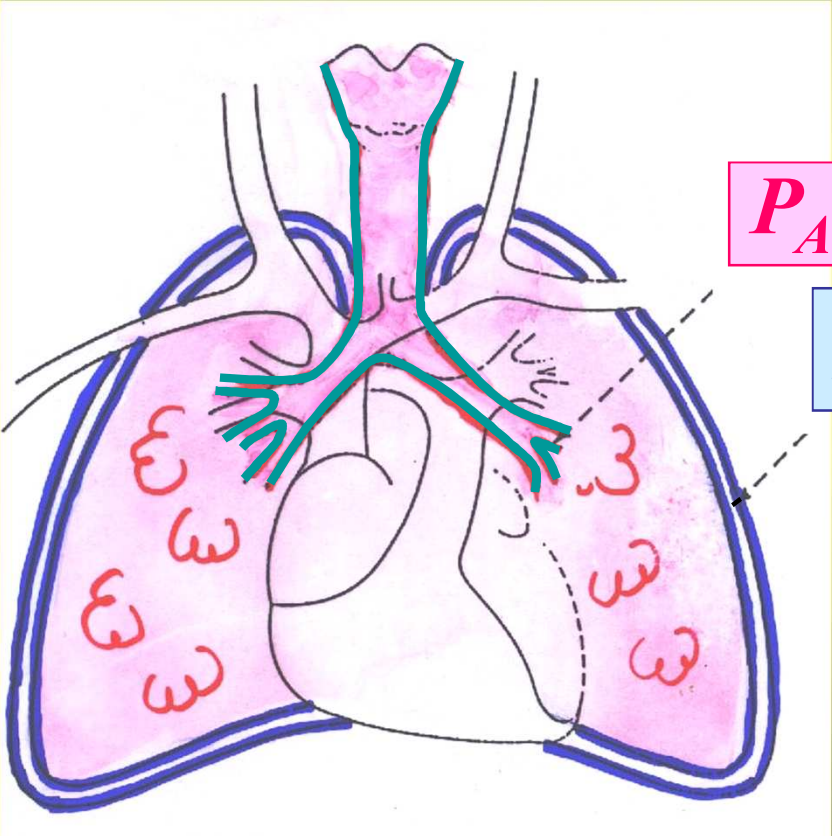
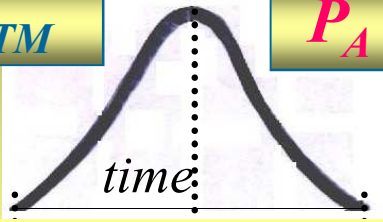
VI TRANSPORT OF GASSES (O_2 and CO_2)

$$P \cdot V = \text{const} \rightarrow P = \frac{\text{const}}{V}$$

TIME COURSE OF PRESSURES AT QUIET RESPIRATION

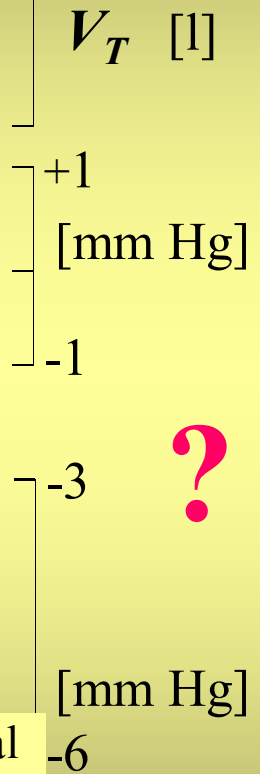
INSPIRATION EXPIRATION

$P_A < P_{ATM}$ $P_A > P_{ATM}$



P_A

P_{PL}



measured curve

theoretical curve

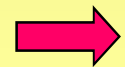
P_A ALVEOLAR (INTRAPULMONARY, LUNG)

P_{PL} INTRAPLEURAL (INTRATHORACIC)

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FORCES PARTICIPATING IN RESPIRATION

- **ACTIVE FORCES** performed by respiratory muscles
- **PASSIVE FORCES** represented by:
 - lungs elasticity
 - chest elasticity

QUIET RESPIRATION

INSPIRATION - active forces of inspiratory muscles prevail

EXPIRATION - only passive (elastic) forces are in action

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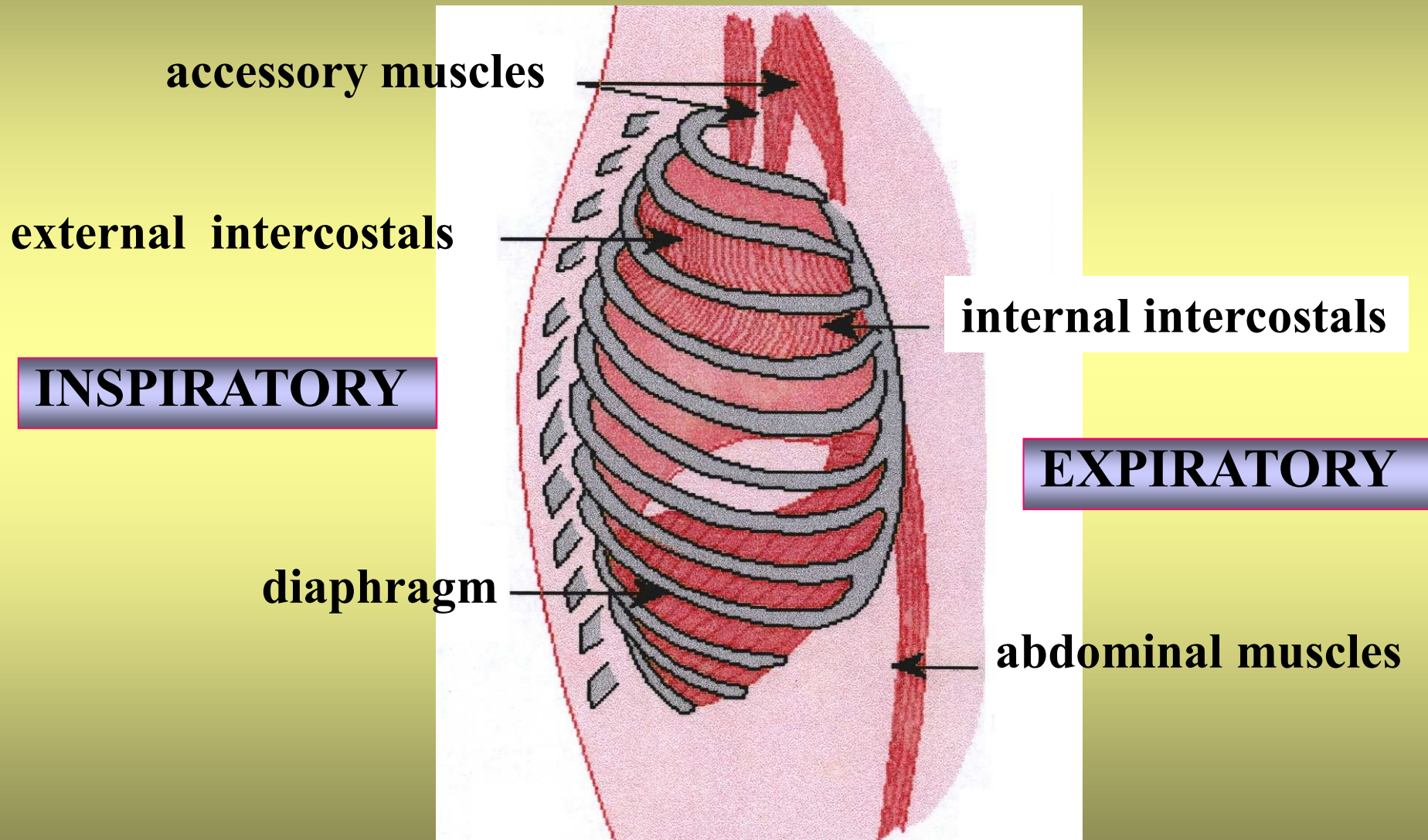
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RESPIRATORY MUSCLES



INSPIRATORY muscles

QUIET breathing

- *diaphragm* ($> 80\%$)
- *external intercostals* ($< 20\%$)

FORCED breathing in addition

- *accessory inspiratory muscles* (mm. scalene)

EXPIRATORY muscles

Only at FORCED breathing

- *internal intercostals*
- *muscles of the anterior abdominal wall*
(abdominal recti, ...)

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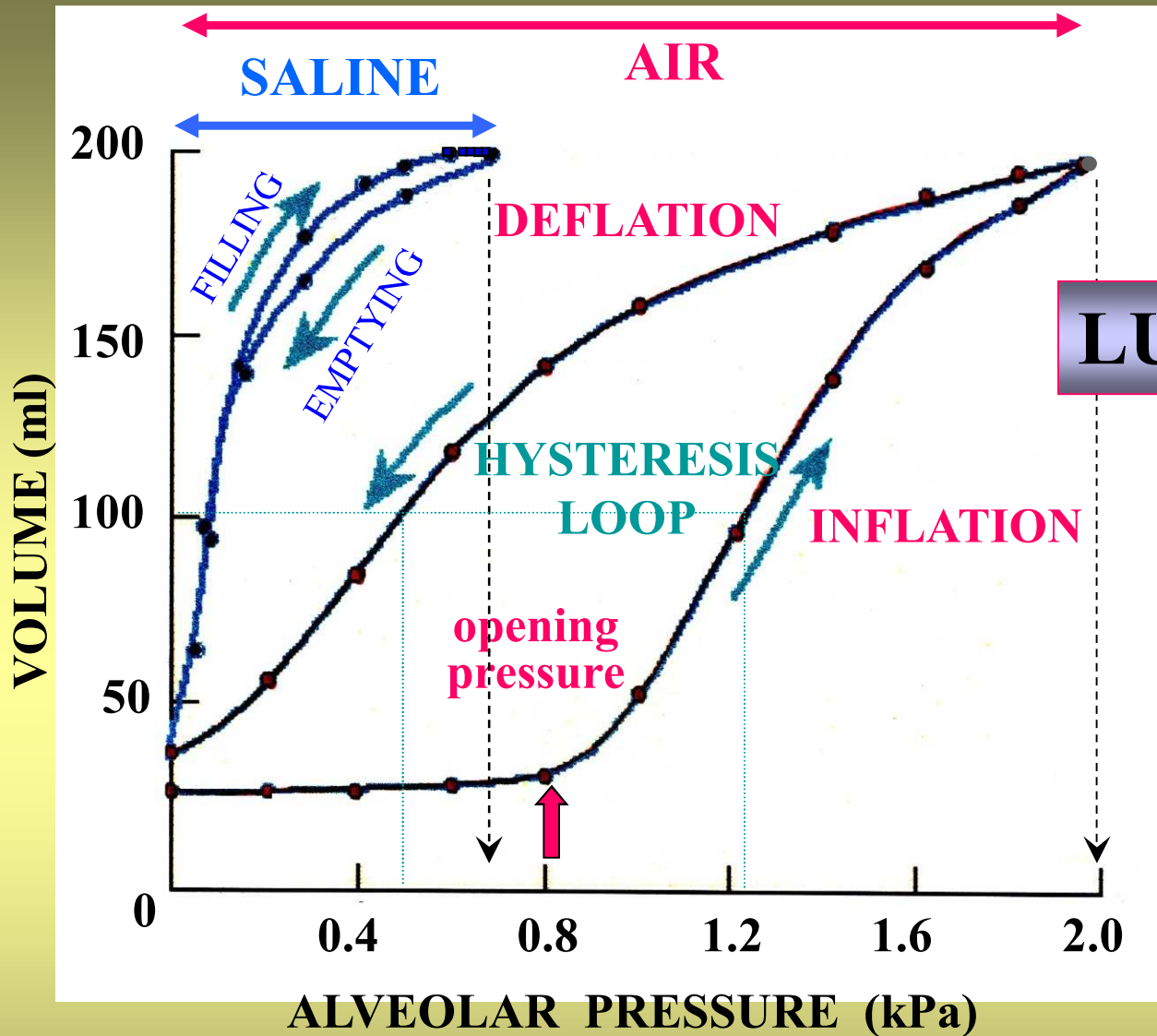
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LUNGS ELASTICITY



1 kPa = 7.5 mm Hg

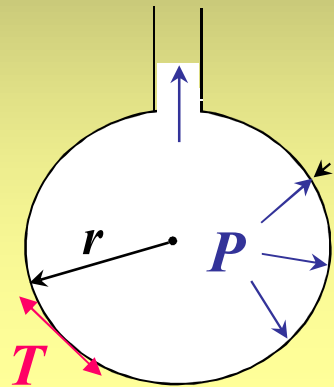
LUNGS ELASTICITY

INHERENT TISSUE ELASTICITY
(elastin and collagen fibres)

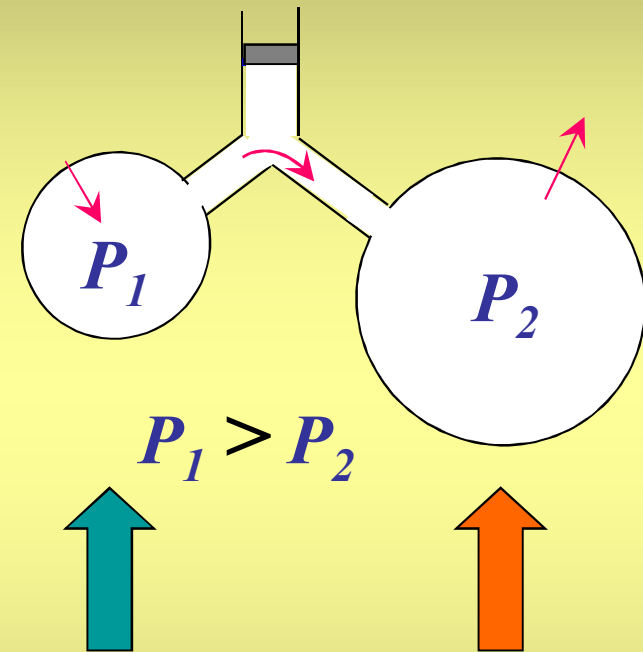
SURFACE TENSION FORCES
air-liquid interface in alveoli

LAW OF LAPLACE

spherical structures



$$P = \frac{2T}{r}$$



P pressure (transmural ΔP)

r radius

T surface tension

PATHOLOGY

- COLLAPSE OF ALVEOLI - ATELECTASIS
- EXPANSION OF ALVEOLI

SURFACTANT

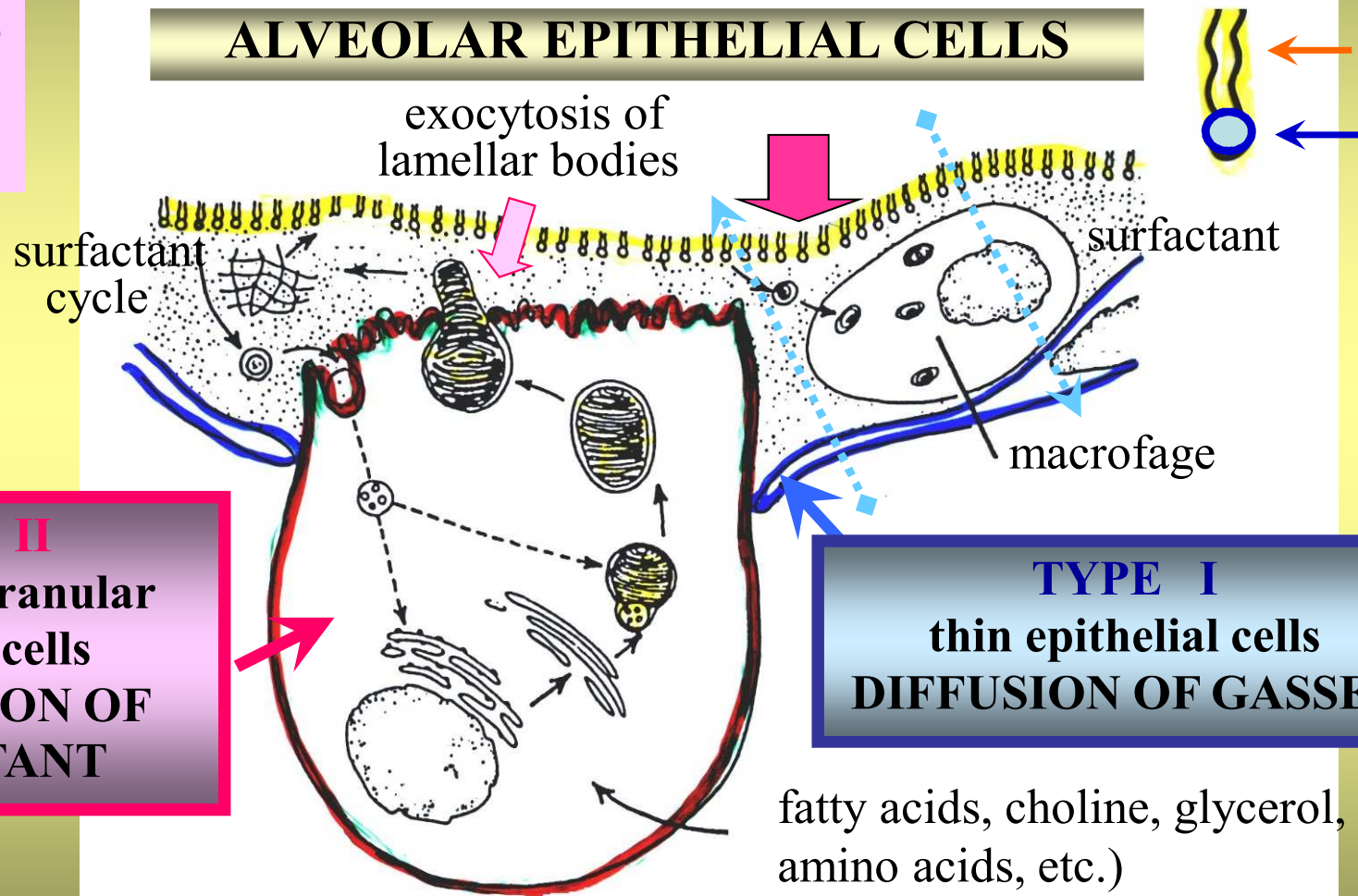
SURFACE TENSION LOWERING AGENT

EFFECT MAINLY IN THE EXPIRED POSITION

PHOSPHOLIPID

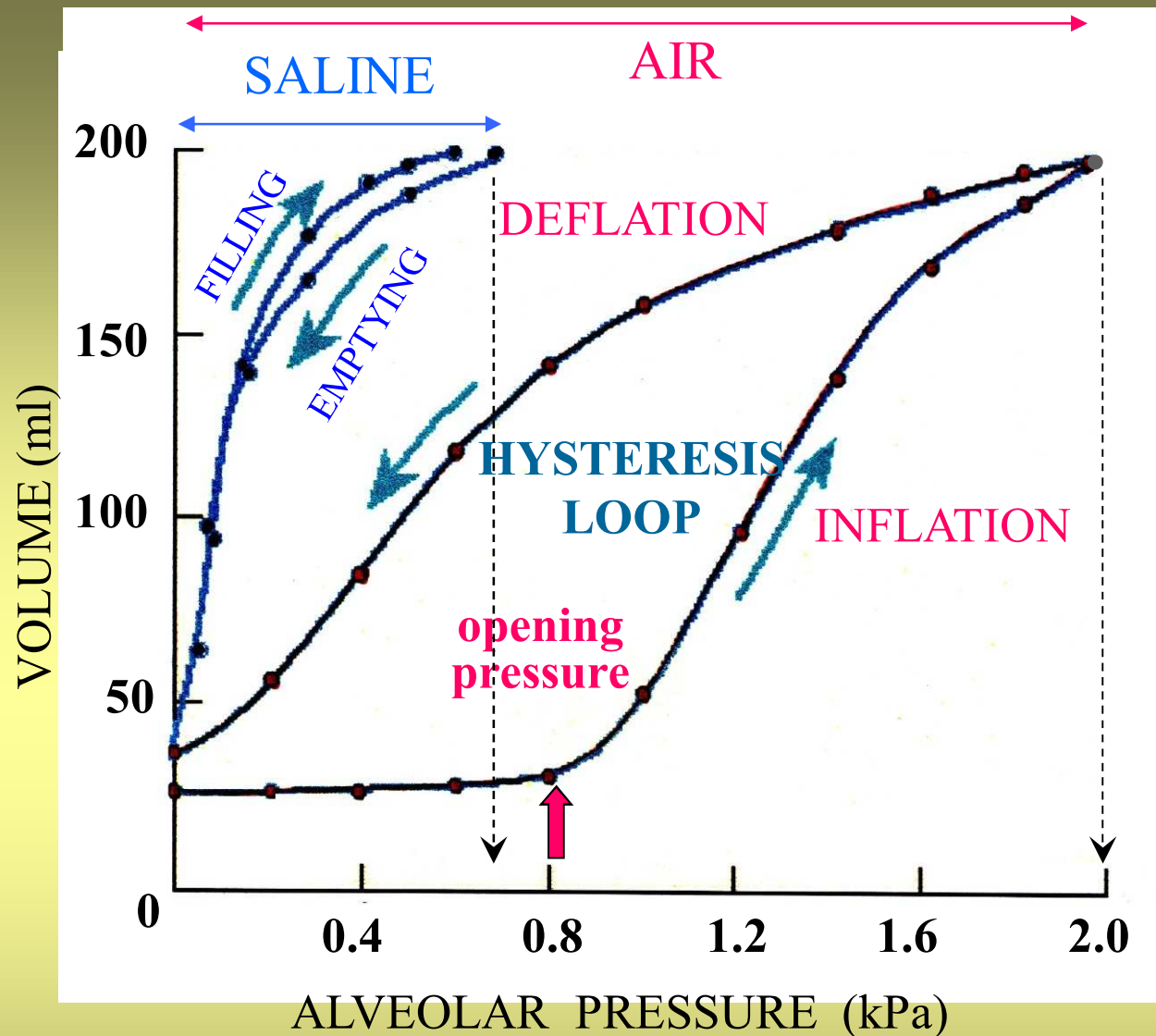
dipalmitoyl
fosfatidyl cholin

ALVEOLAR EPITHELIAL CELLS



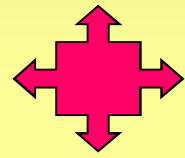
TYPE II
specialized granular
epithelial cells
**PRODUCTION OF
SURFACTANT**

TYPE I
thin epithelial cells
DIFFUSION OF GASSES



**Factors involved in
HYSTERESIS LOOP**

- **LAPLACE LAW** (responsible for high **opening pressure** of alveoli)
- **Dynamic changes in the DENSITY** of surfactant molecules during **INSPIRATION** and **EXPIRATION**



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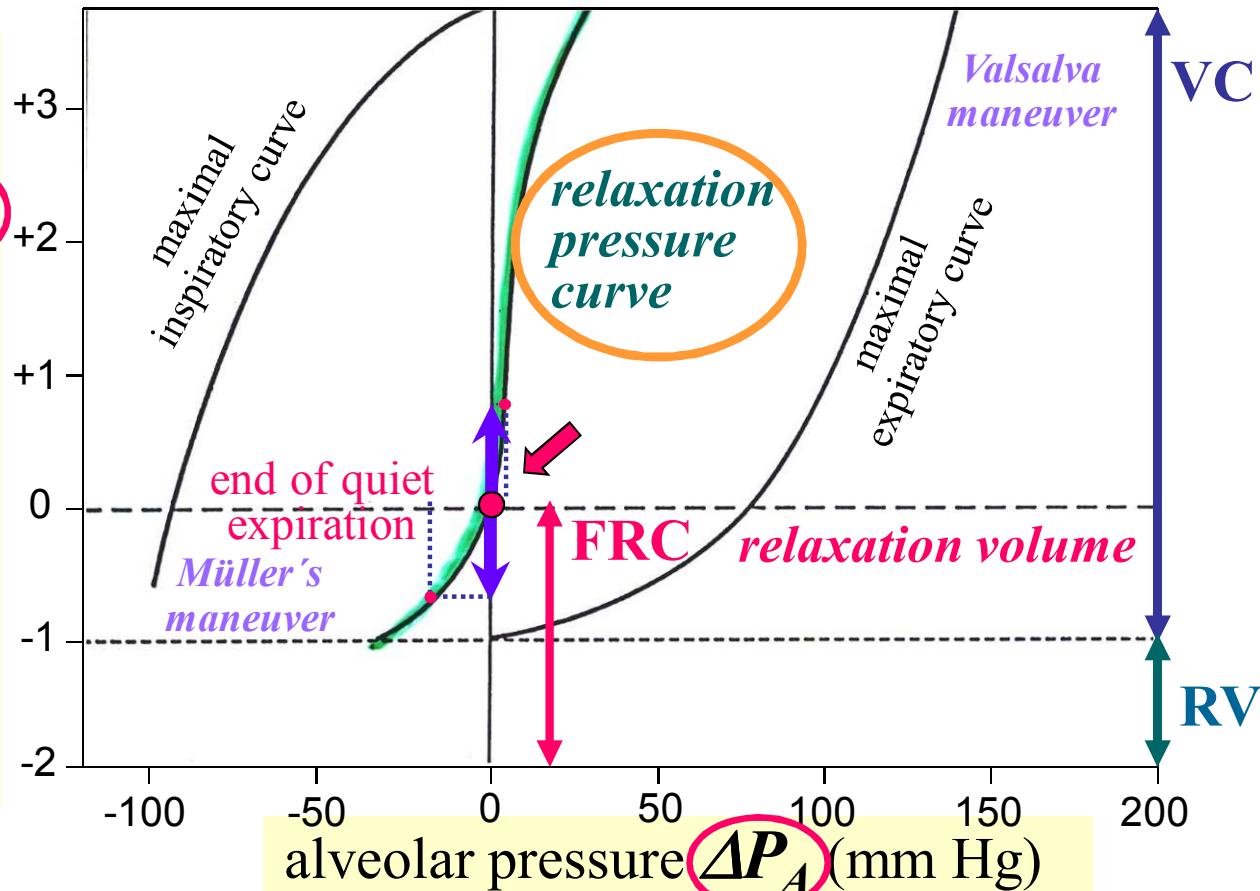
VI TRANSPORT OF GASSES (O_2 and CO_2)

COMPLIANCE (VOLUME STRETCHABILITY)

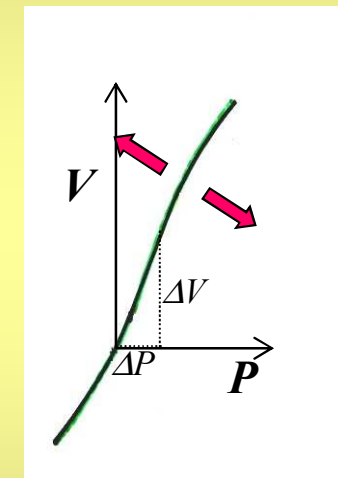
STATIC MEASUREMENT IN CLOSED SYSTEM

$$C = \frac{\Delta V}{\Delta P}$$

change of the volume ΔV (l)



alveolar pressure ΔP_A (mm Hg)



compliance is decreased
 ↑ *stiffness of the tissue*

compliance is increased
 ↓ *stiffness of the tissue*

TOTAL RESPIRATORY SYSTEM
 (lungs and chest)

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VI TRANSPORT OF GASSES (O_2 and CO_2)

TOTAL WORK OF RESPIRATORY MUSCLES AT QUIET BREATHING

ELASTIC (STATIC) WORK (65%)

to overcome the elastic forces of the chest and lungs

DYNAMIC WORK (35%)

- to overcome the resistance of air passages during the air movement – **AERODYNAMIC RESISTANCE** (~ 28%)
- to overcome the friction during mutual movement of inelastic tissues – **VISCOUS RESISTANCE** (~ 7%)

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COMPOSITION OF DRY ATMOSPHERIC AIR

O₂ 20.98 %

N₂ 78.06 %

CO₂ 0.04 %

Other constituents

F_{O₂} ≅ 0.21

F_{N₂} ≅ 0.78

F_{CO₂} = 0.0004

BAROMETRIC (ATMOSPHERIC) PRESSURE AT SEA LEVEL

1 atmosphere = 760 mm Hg

PARTIAL PRESSURES OF GASSES IN DRY AIR AT SEA LEVEL

$$P_{O_2} = 760 \times 0.21 = \sim 160 \text{ mm Hg}$$

$$P_{N_2} = 760 \times 0.78 = \sim 593 \text{ mm Hg}$$

$$P_{CO_2} = 760 \times 0.0004 = \sim 0.3 \text{ mm Hg}$$

$$1 \text{ kPa} = 7.5 \text{ mm Hg (torr)}$$

COMPOSITION OF ALVEOLAR AIR

partial pressures in mm Hg

INSPIRED AIR

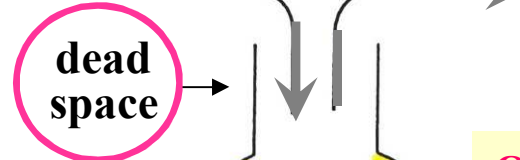
O ₂	158.8
CO ₂	0.3
N ₂	601.0
...	

760 mm Hg

EXPIRED AIR

O ₂	115.0
CO ₂	33.0
H ₂ O	47.0
N ₂	564.0
...	

760 mm Hg



O ₂	100.0
CO ₂	39.0
H ₂ O	47.0
N ₂	...

O ₂	100.0
CO ₂	39.0

right heart

physiological shunts

760 mm Hg

left heart

veins

O ₂	40.0
CO ₂	45.0
H ₂ O	47.0
N ₂	...
...	

O ₂	95.0
CO ₂	41.0
H ₂ O	47.0
N ₂	...
...	

arteries

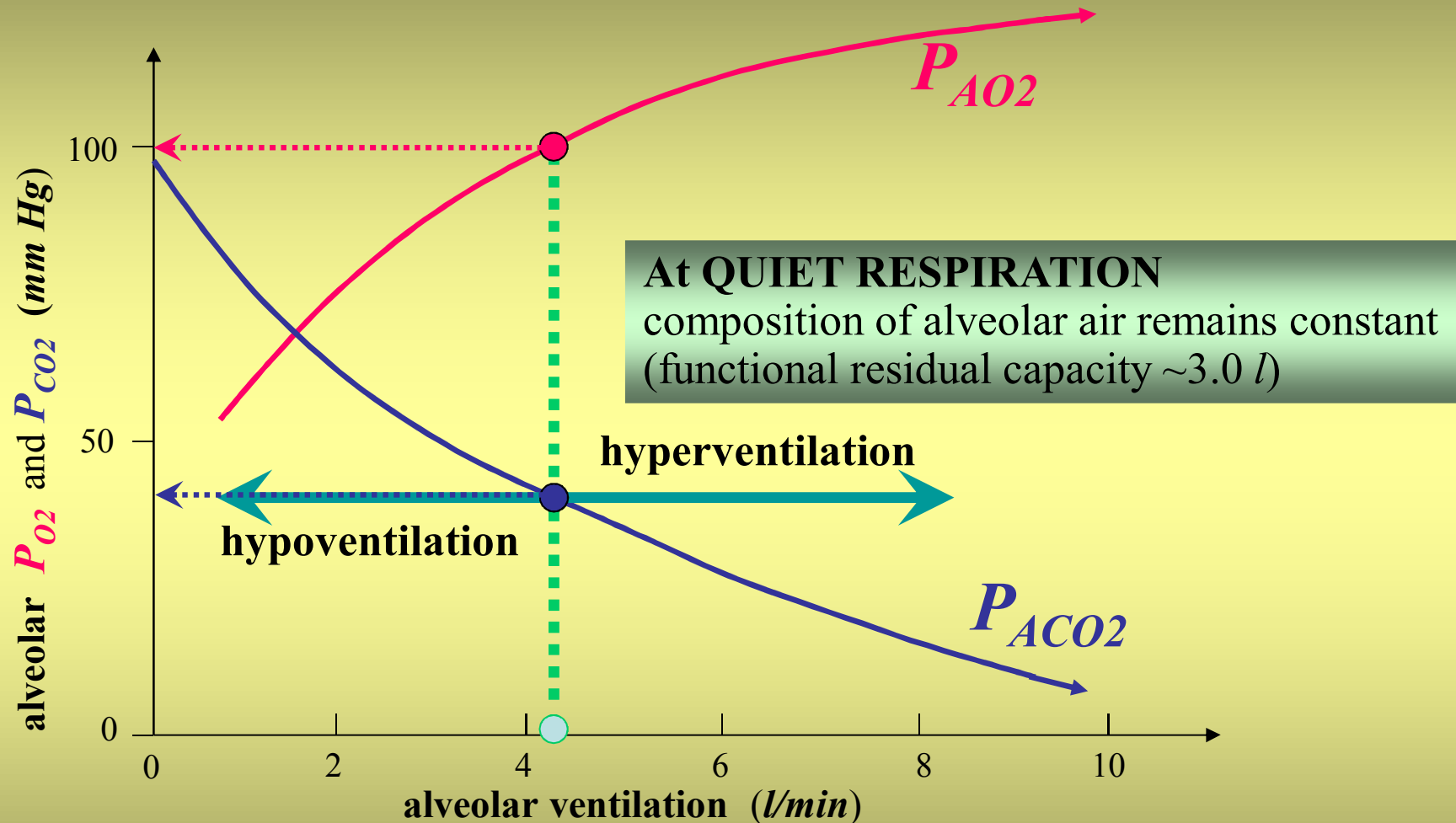
periphery capillaries

O ₂	40.0
CO ₂	45.0
H ₂ O	47.0
N ₂	...
...	

?

?

Alveolar P_{O_2} and P_{CO_2} at voluntary hypo- and hyperventilation



hyperventilation \rightarrow hypocapnia \rightarrow respiratory alkalosis

hypoventilation \rightarrow hypercapnia \rightarrow respiratory acidosis

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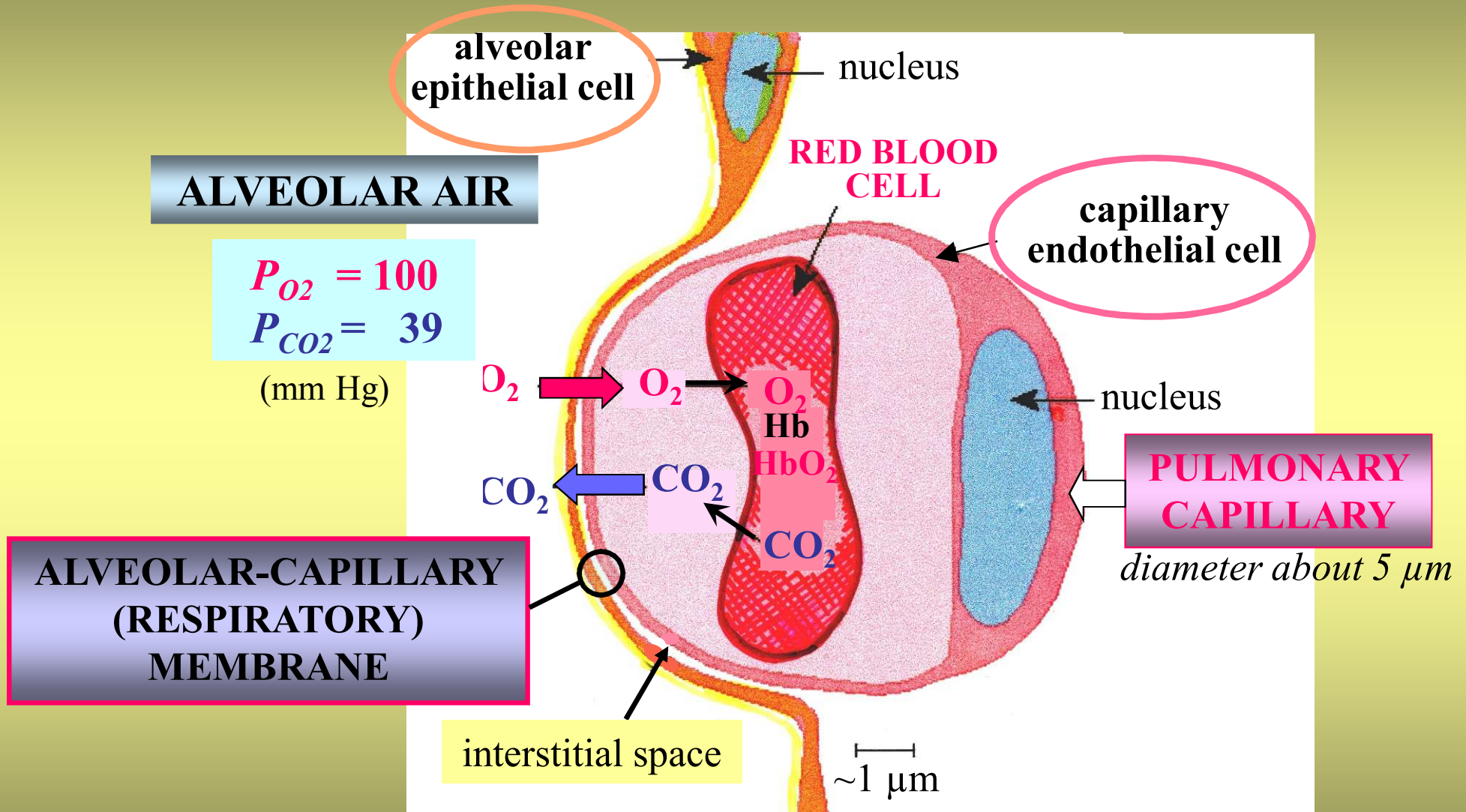
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→ V ALVEOLAR-CAPILLARY MEMBRANE

VI TRANSPORT OF GASSES (O_2 and CO_2)

ALVEOLAR-CAPILLARY (RESPIRATORY) MEMBRANE

DIFFUSION OF GASES



0.75 s

time interval of erythrocyte contact with respiratory membrane at rest

TIME COURSE OF CAPILLARY P_{O_2} AND P_{CO_2} DURING GRADUAL EQUILIBRATION WITH ALVEOLAR AIR

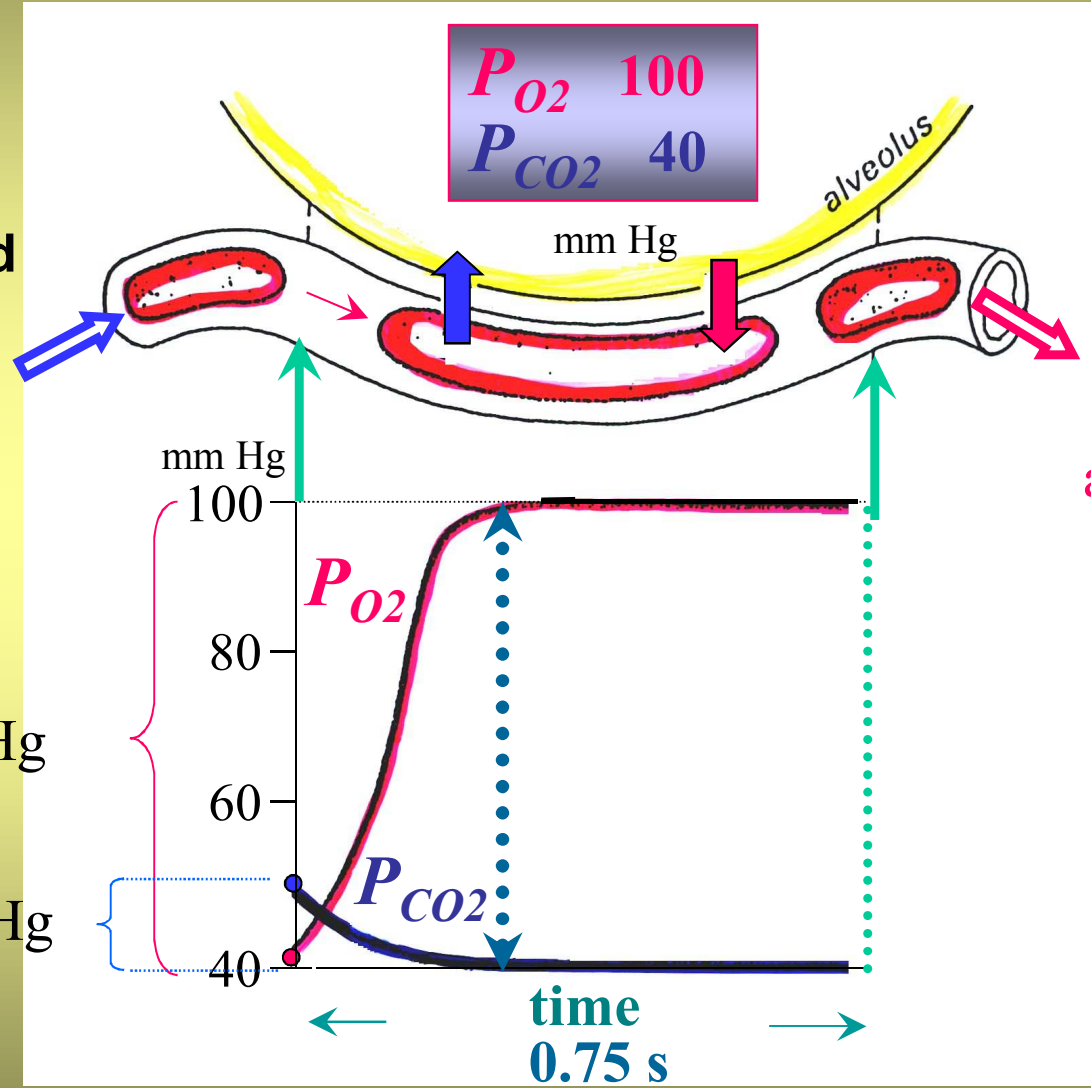
venous blood

P_{O_2} 40
 P_{CO_2} 46

mm Hg

$\Delta P_{O_2} \cong 60$ mm Hg

$\Delta P_{CO_2} \cong 6$ mm Hg



P_{O_2} 100
 P_{CO_2} 40

mm Hg

equalization with alveolar pressures

time interval of contact of erythrocyte with respiratory membrane at rest

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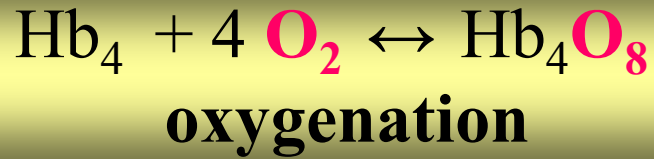
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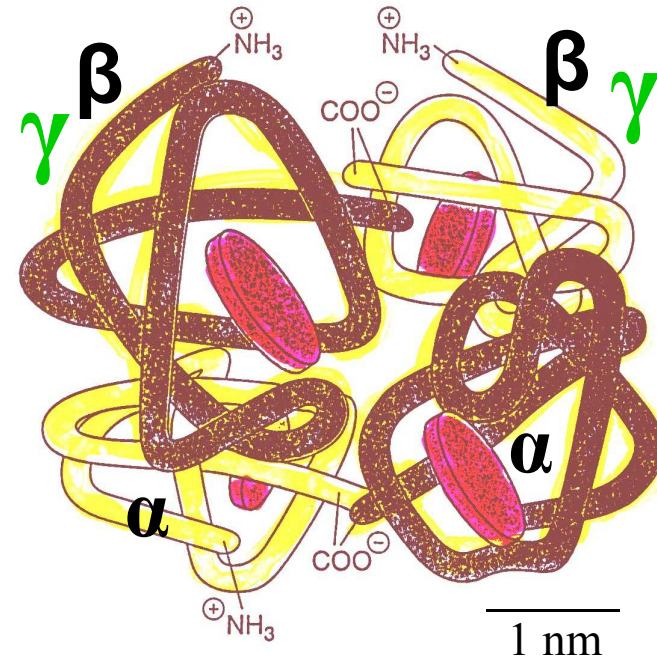
V ALVEOLAR-CAPILLARY MEMBRANE

➔ VI TRANSPORT OF GASSES (O_2 and CO_2)

HAEMOGLOBIN



tetramer

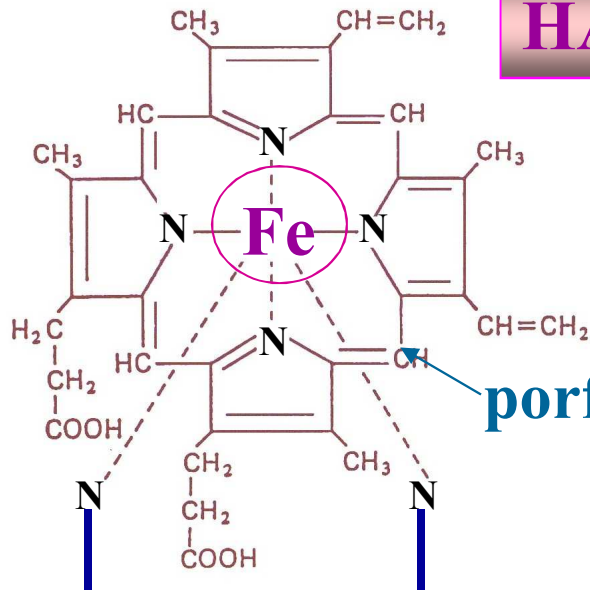


Fe^{2+}

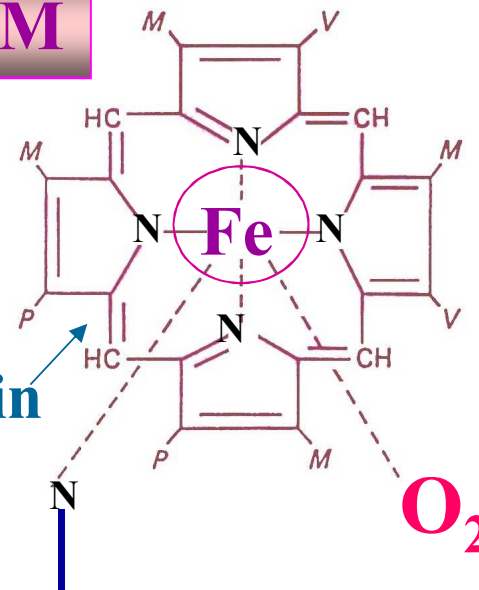
DEOXY

OXY

HAEM



porphyrin



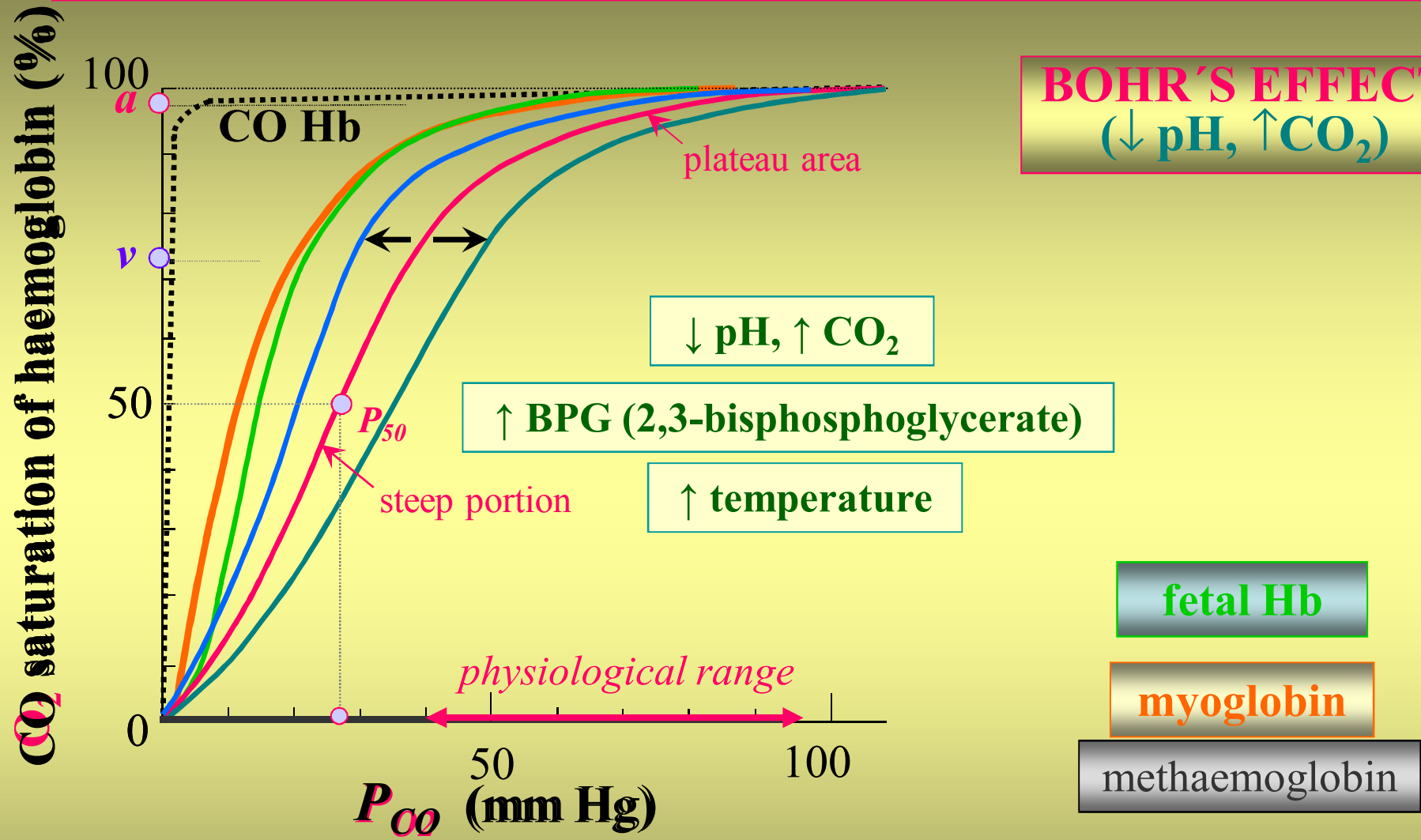
fetal Hb

Fe^{3+} (methaemoglobin)
oxidation

polypeptide chain

polypeptide chain

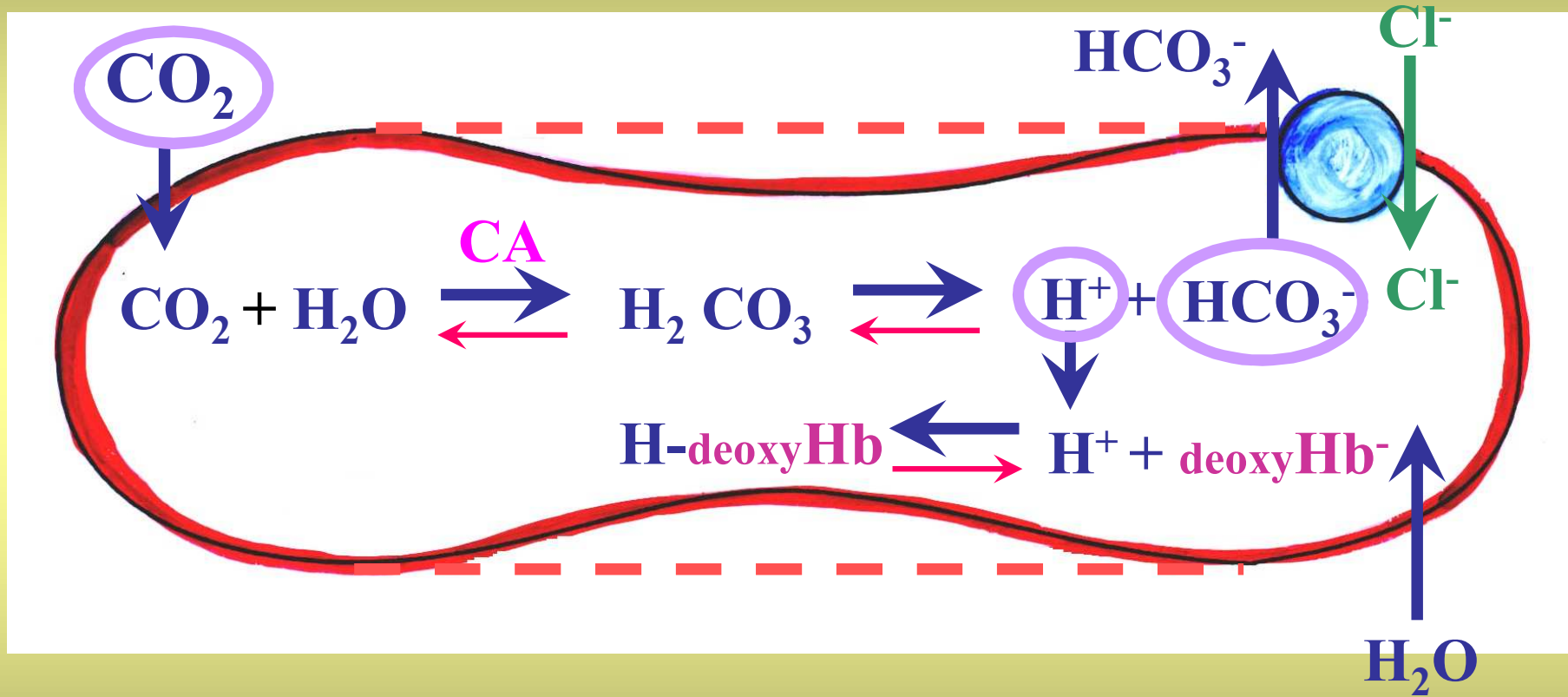
O_2 -HAEMOGLOBIN DISSOCIATION CURVE



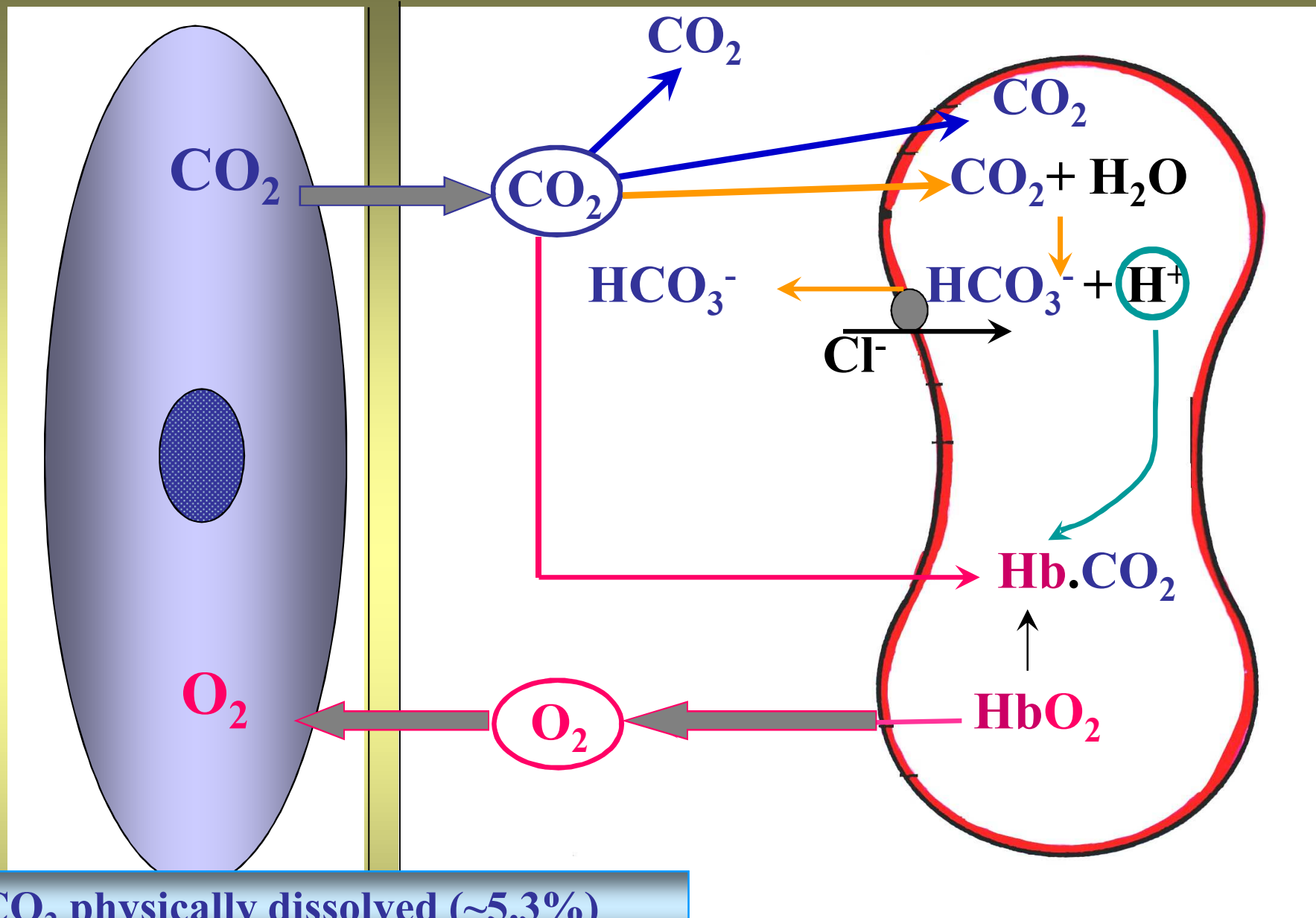
physically dissolved O_2 (1.4%)

TRANSPORT OF CO₂

HAMBURGER CHLORIDE SHIFT



CA – carbonic anhydrase



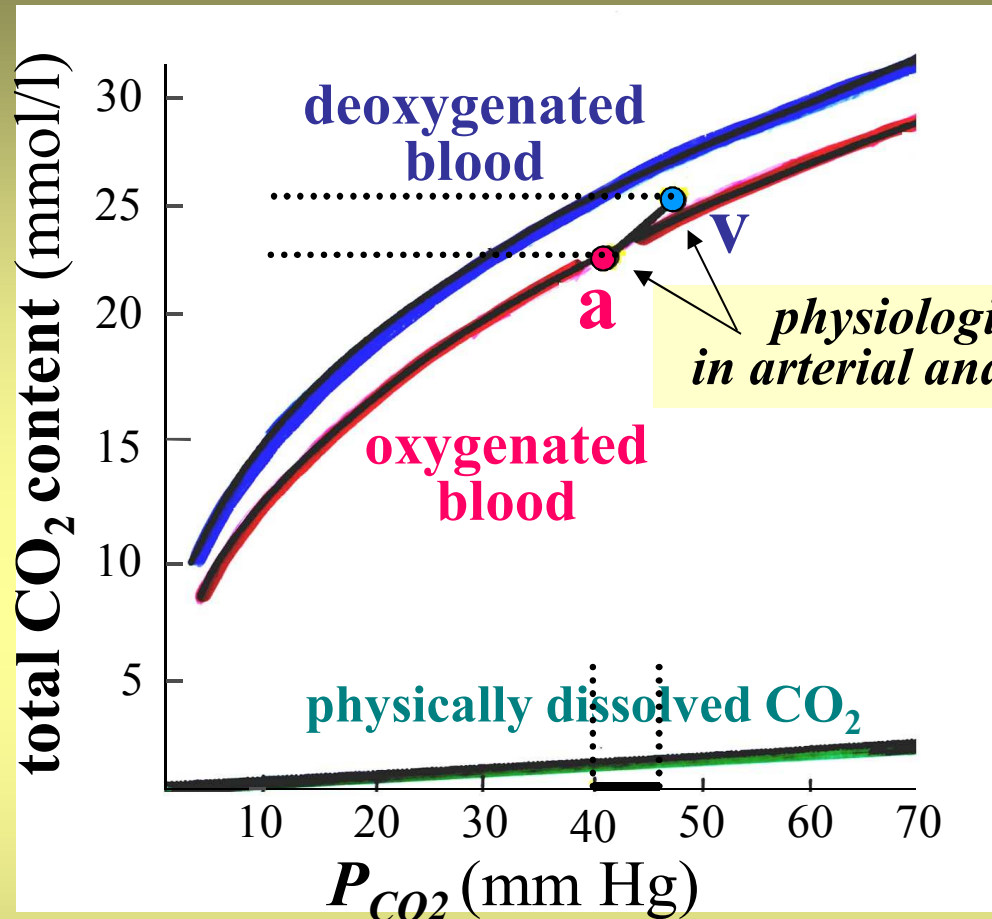
- CO₂ physically dissolved (~5.3%)

- $\text{CO}_2 + \text{Hb-NH}_2 \rightleftharpoons \text{Hb.NH-COO}^-$ (carbamino-Hb) (~5.3%)

- $\text{CO}_2 + \text{H}_2\text{O} \rightleftharpoons \text{HCO}_3^- + \text{H}^+$ (~89%)

60% in plasma, 29% in red blood cell

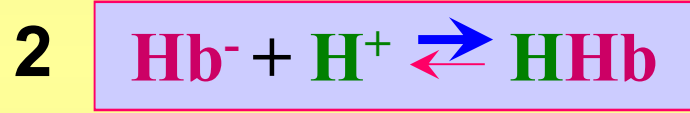
CO₂ DISSOCIATION CURVE



HALDANE EFFECT

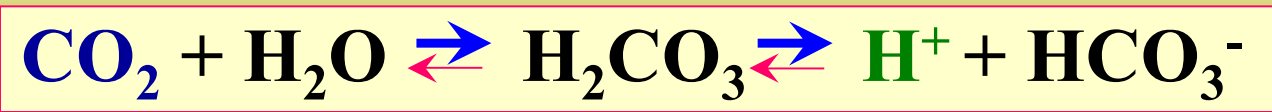
?

DEOXY-Hb

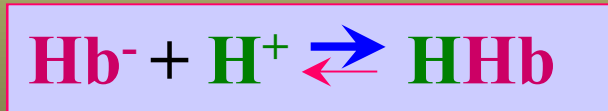


→ deoxygenated blood in peripheral tissues

← oxygenated blood in the lungs

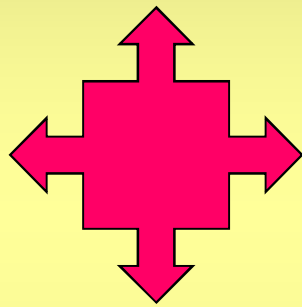


↑
↓



TISSUES: DEOXY-Hb binds H⁺ more readily (weaker acid) ⇒ ↑ amount of chemically bound CO₂

LUNGS: H⁺ is released from OXY-Hb ⇒ ↓ amount of chemically bound CO₂



END