

Respiratory distress syndrome



***Eman Farouk Badran
MD. MRCPCH
Professor of Pediatrics
Head division of Neonatology
At Jordan University Hospital***

Third year medical
students

December/2017

Outlines

- **Definition**
- **Physiology**
 - Respiration
 - Surface tension
 - Lung compliance
 - Lung volume
 - surfactant
- **Respiratory distress syndrome**
 - Pathophysiology
 - Incidence
 - Presentation
 - management



Respiration = the series of **exchanges** that leads to the uptake of oxygen by the cells, and the release of carbon dioxide to the lungs

Step 1 = ventilation

- **Inspiration & expiration**

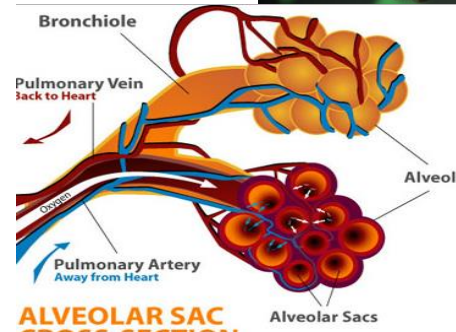
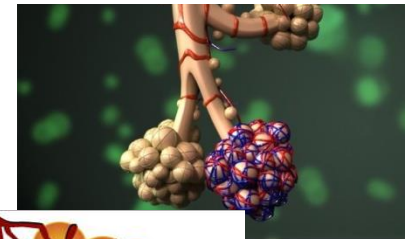
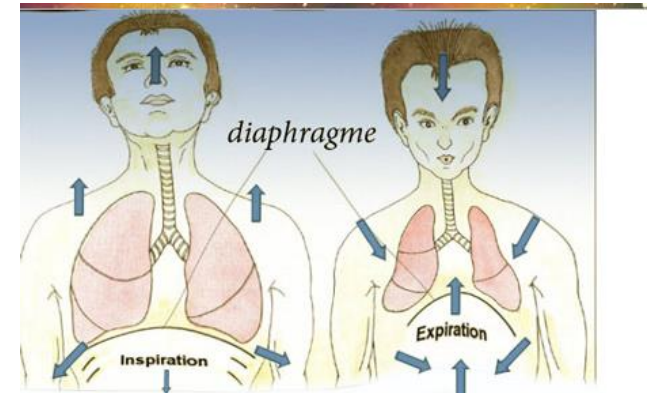
Step 2 = **exchange between alveoli** (lungs) and pulmonary capillaries (blood)

- Referred to as *External Respiration*

Step 3 = **transport of gases** in blood

Step 4 = **exchange between blood and cells**

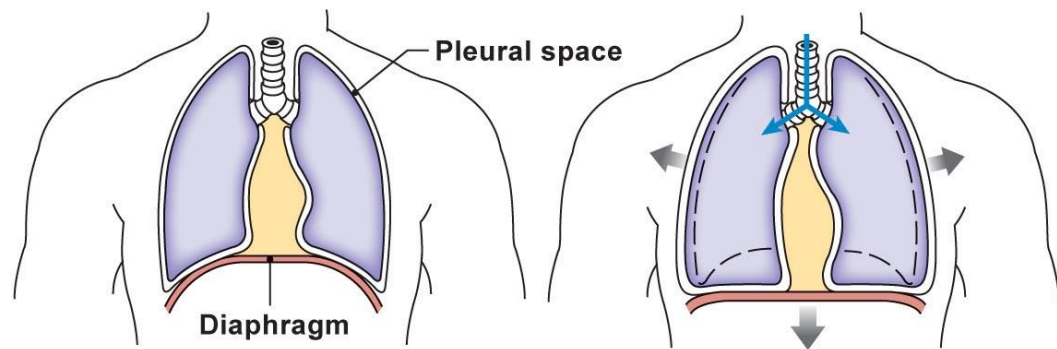
- Referred to as *Internal Respiration*



Ventilation = (inspiration + expiration) responsible muscles

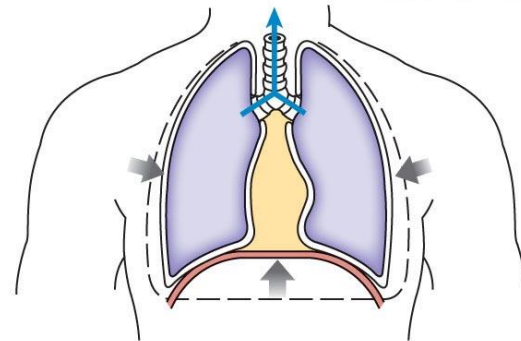
➤ The diaphragm (only creates about 60-75% of the volume change during inspiration)

➤ The muscles of **Inspiration** (external intercostals muscles) & muscles of expiration (internal intercostals muscles)



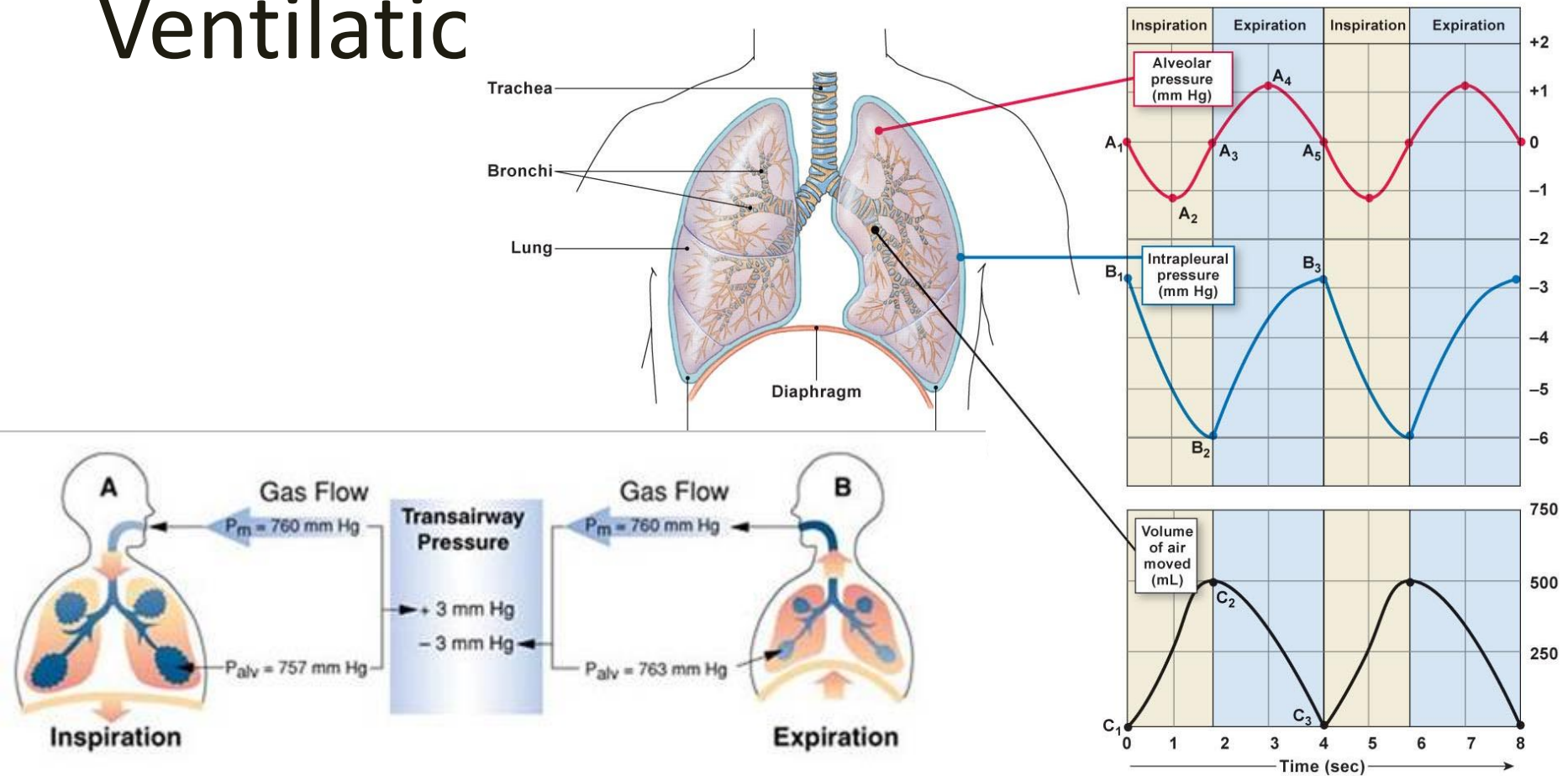
(a) At rest, diaphragm is relaxed.

(b) Diaphragm contracts, thoracic volume increases.



(c) Diaphragm relaxes, thoracic volume decreases.

Ventilatic

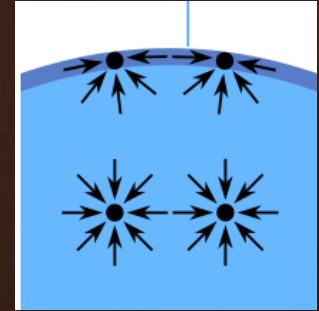


Tidal volume in new born = 4 –6ml / kg

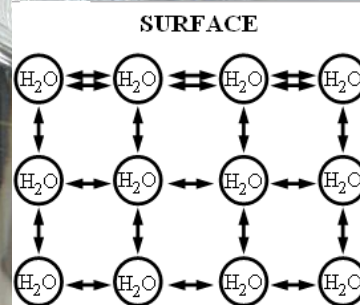
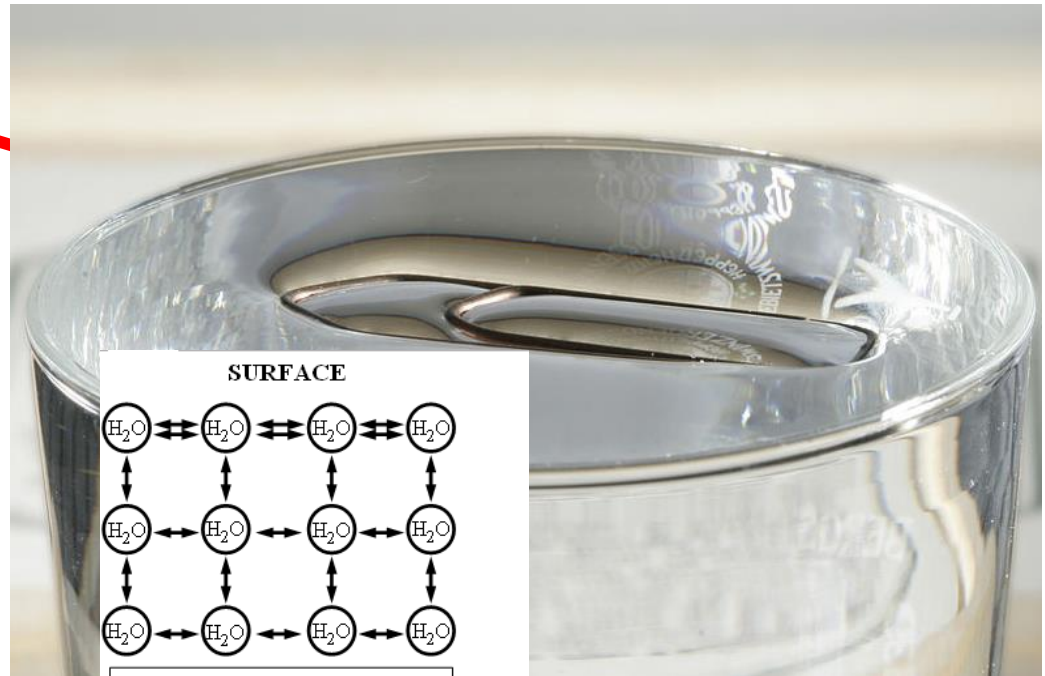
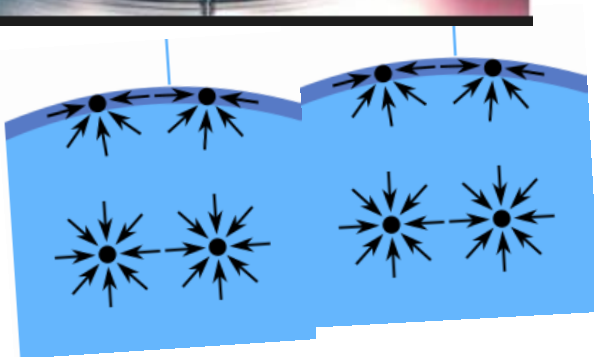
If baby weigh=3kg

TV =12 -18 ml

Surface tension



An air-filled sphere coated with water has a tendency to collapse (reach a minimum volume) due to the pulling force of water surface tension



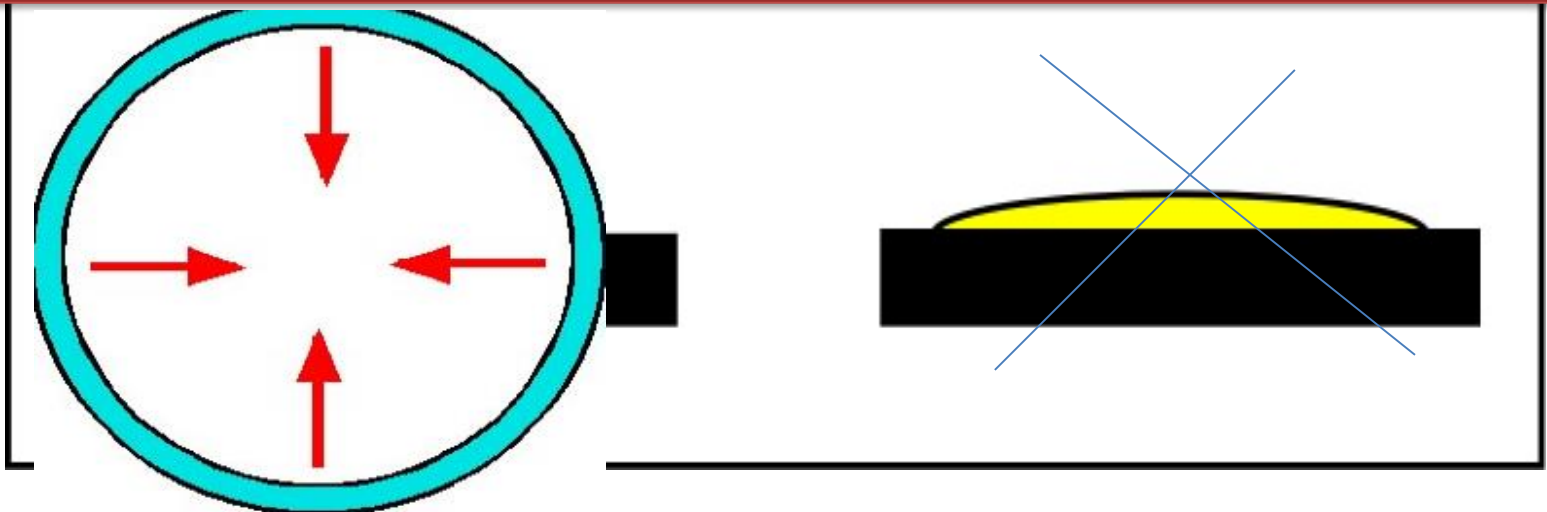
Surface tension—molecules at the surface form stronger bonds

Surface Tension

Water has a VERY HIGH surface tension

Water will attempt to minimize its surface area in contact with air

Surface tension : Attractive forces between molecule at air water interface

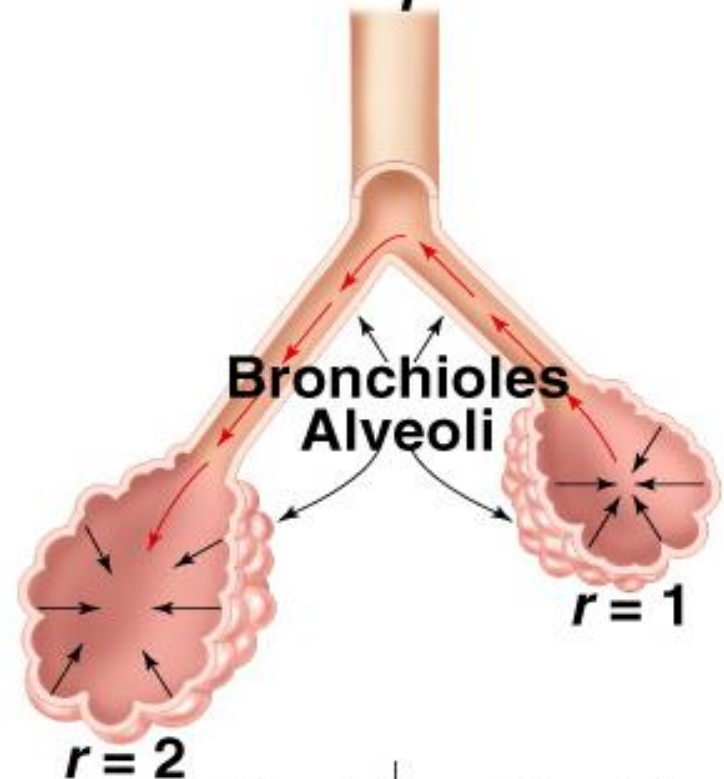


Law of Laplace

Copyright © The McGraw-Hill Companies, Inc. Permission required for reproduction or display.

Law of Laplace

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



$$r = 2$$

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

$$P = T$$

$$P = \frac{2 \times T}{1}$$

$$P = 2T$$

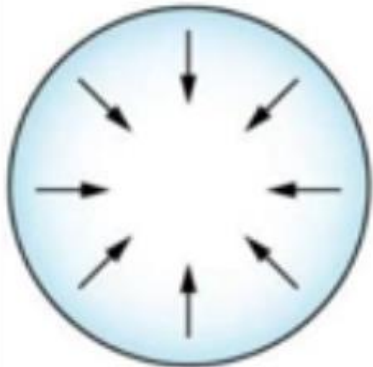
Figure 16.11

- Collapsing Pressure in alveoli is :
 - directly proportional to surface tension
 - and inversely proportional to radius of alveoli
- The smaller the sphere the more surface tension
 - Pressure in smaller alveolus greater

Surface tension

$$P(\text{collapsing Pressure}) = \frac{2 \times T}{r}$$

(a) Pressure is greater in the smaller bubble



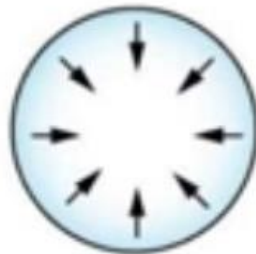
Larger bubble

$$r = 2$$

$$T = 3$$

$$P = (2 \times 3)/2$$

$$P = 3$$



Smaller bubble

$$r = 1$$

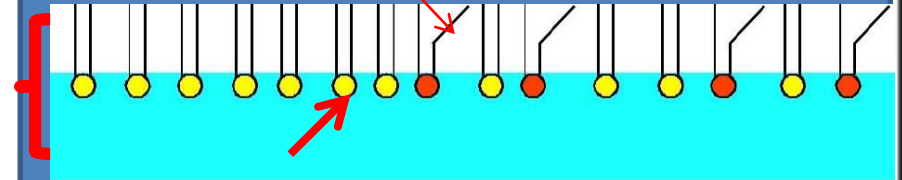
$$T = 3$$

$$P = (2 \times 3)/1$$

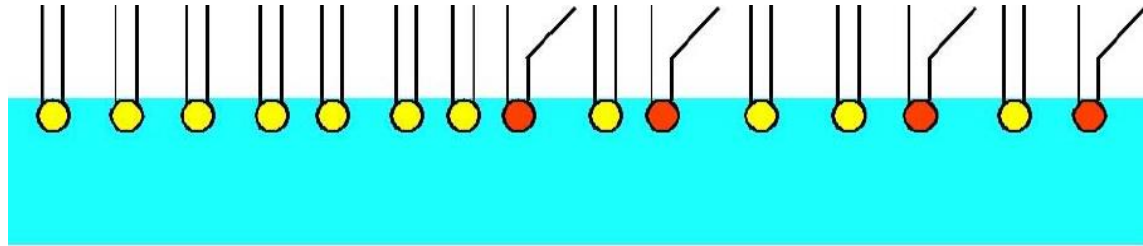
$$P = 6$$

What happen if

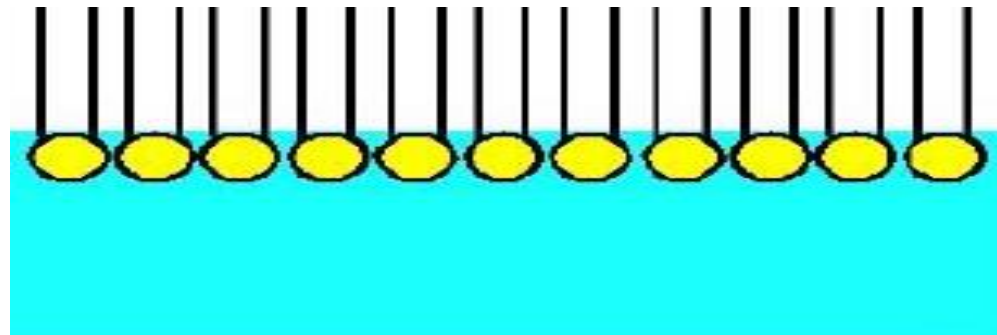
Lipids form a monolayer at the air-water interface



Lipids form a monolayer at the air-water interface



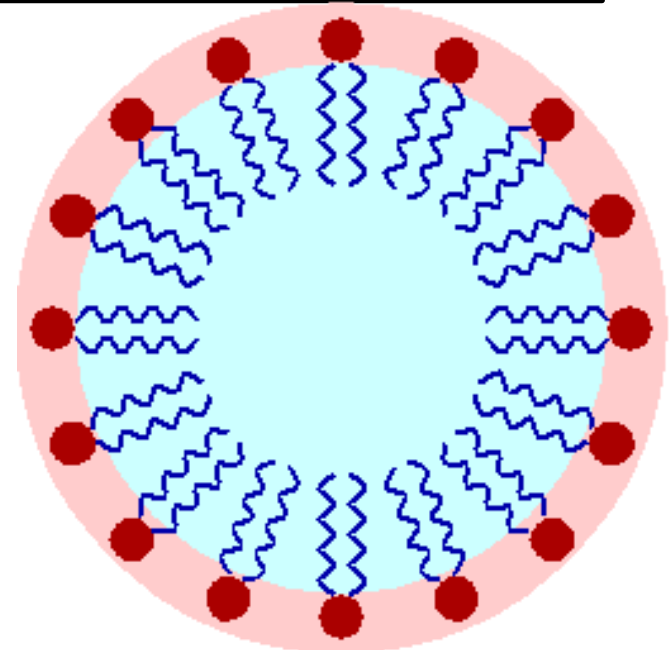
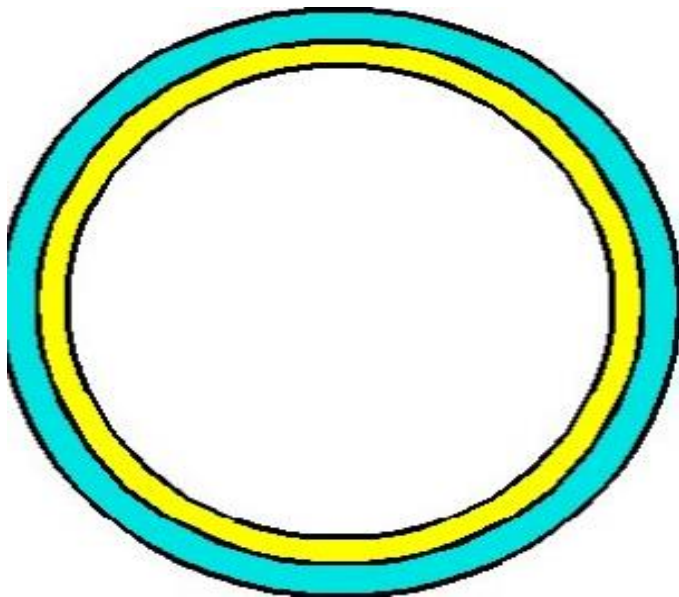
Surface tension decreases as lipid monolayer is compressed



1. Alveoli are coated with lung surfactant in order to reduce the surface tension of water through:

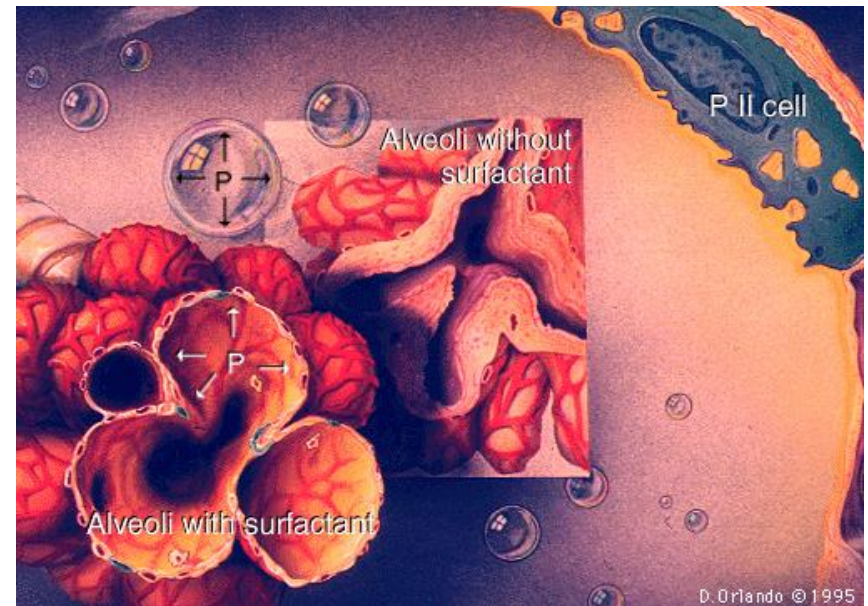
- a) It scatters among the fluid molecule decreasing the attraction between them.
- b) It also spreads over the fluid preventing air-fluid interface.

thus preventing collapse (atelectasis) upon exhalation and decreasing the force necessary to expand the alveoli upon inhalation



Lung Function in respiratory distress syndrome (RDS)

- Reduction in Functional residual capacity (FRC)



Surfactant

Copyright © The McGraw-Hill Companies, Inc. Permission required for reproduction or display.

- produced by alveolar type II cells

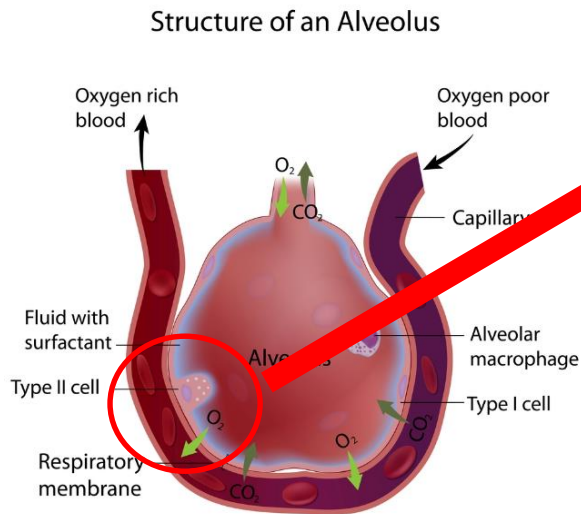
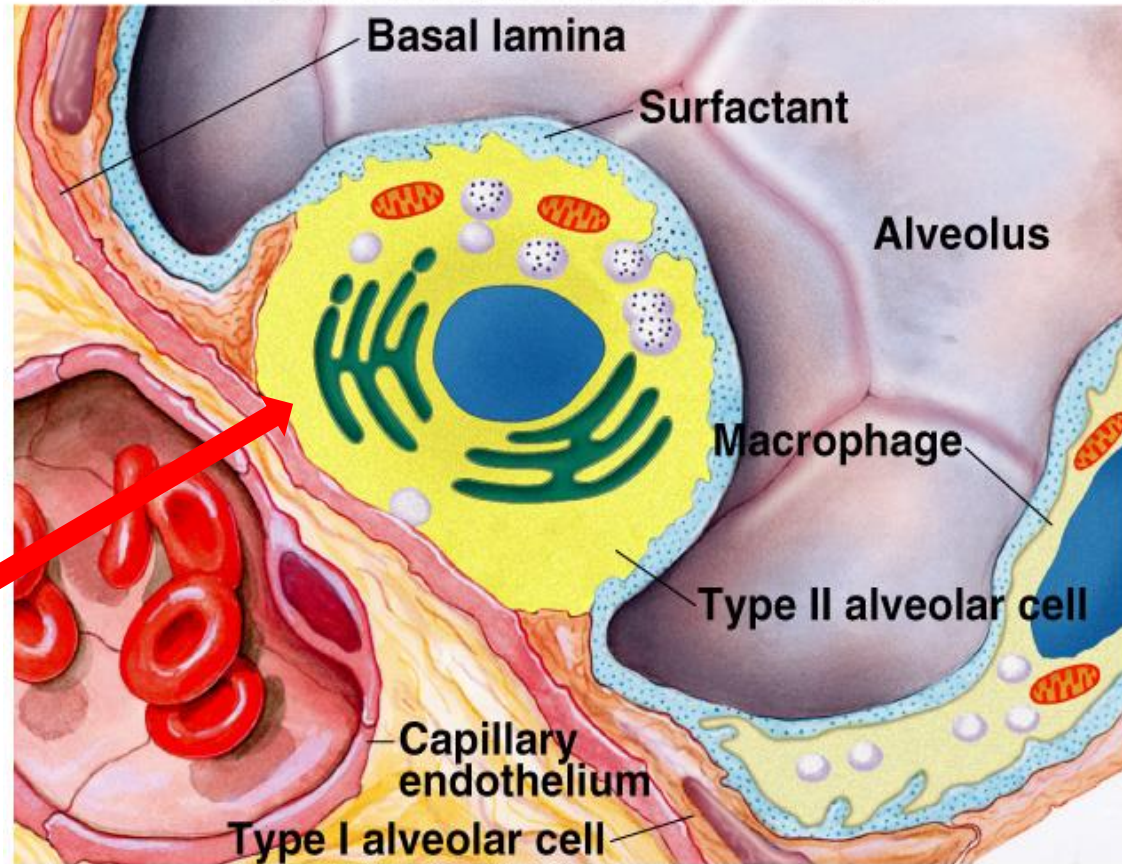


Figure 16.12

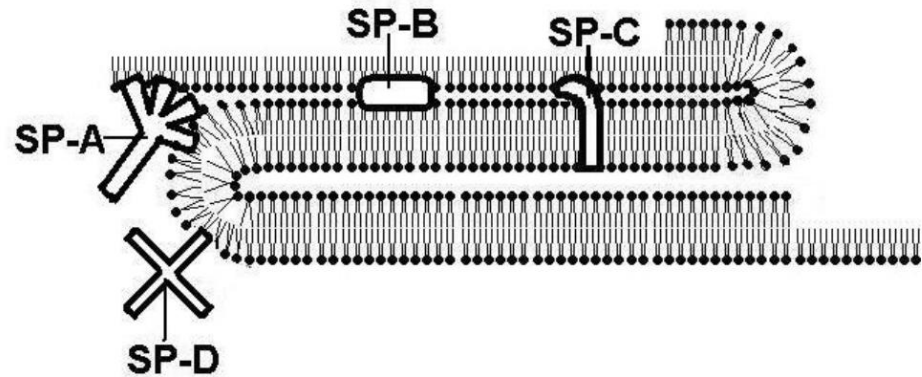
Endogenous Surfactant composition and functions

- **Major Lipids (~90%)**
 - Saturated Phosphatidylcholine **DPPC** (***Lecithin***) 60-80%
 - Unsaturated Phospholipids
 - Phosphatidylglycerol (**PG**) ~10%
- **Proteins (~10%)**
 - SP-A
 - Hydrophilic**, Host defense
 - Surfactant homeostasis
 - SP-B
 - **Hydrophobic**, Spreading, ↓ surface tension
 - SP-C
 - Hydrophilic , Adsorption
 - SP-D: ? Phagocytic function

Surfactant proteins

Surfactant proteins are divided into 2 groups:

- **Large and water-soluble SP-A and SP-D proteins**
- **small, hydrophobic SP-B and SP-C proteins.**



Are of great importance to immune defense mechanisms of the lung

- ability to bind to bacteria, viruses and other pathogens

.....(

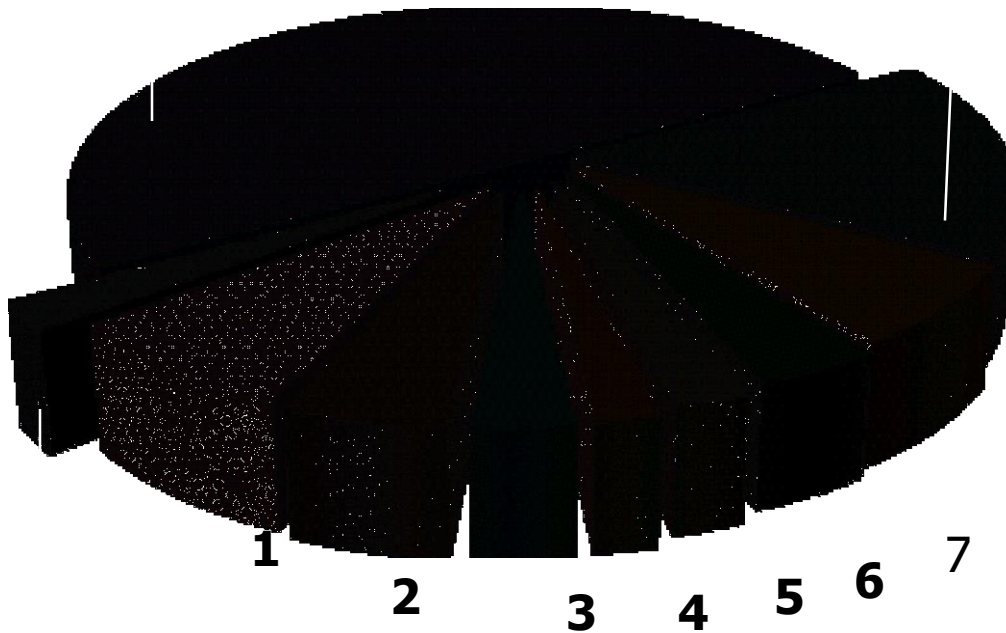
- well as to activate alveolar macrophages

CSurfactant Composition

DPPC - dipalmitoylphosphatidylcholine 60%*

**PG - phosphatidylglycerol
7%***

• *Promotes the spreading of
surfactant throughout the lungs*



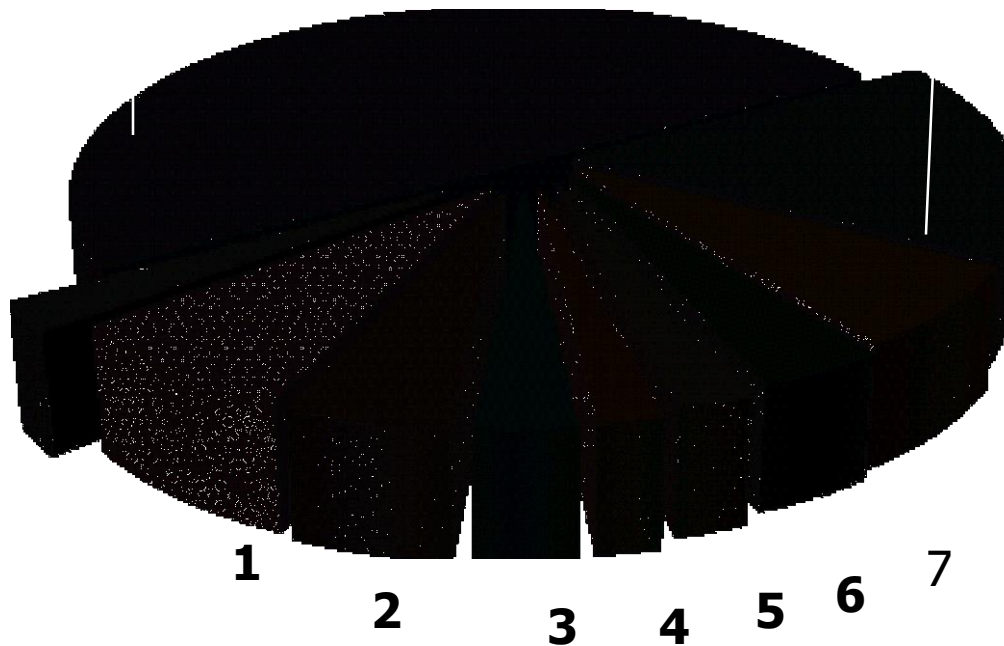
1. Serum proteins 10%
2. Other lipids 5%*
3. Other phospholipids 3%*
4. Phosphatidylinositol 2%*
5. Sphingomyelin 2%*
6. Phosphatidylethanolamine 4%*
7. Unsaturated Phosphatidylcholine 17%*

** By molecular weight*

Prenatal diagnosis

- Lecithin and sphingomyelin ratio in the amniotic fluid, if ratio is more than 2 indicates adequate lung maturity

DPPE - dipalmitoylphosphatidylcholine (lecithin) 60%-80%*



PG - phosphatidylglycerol

7%*

• Promotes the spreading of surfactant throughout the lungs

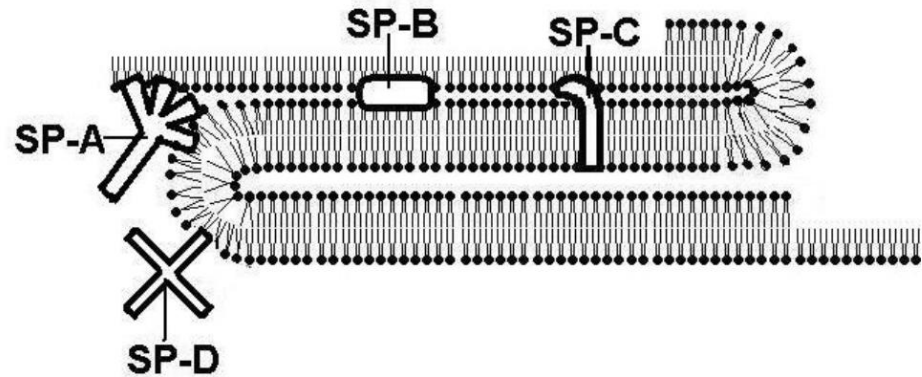
1. Serum proteins 10%
2. Other lipids 5%*
3. Other phospholipids 3%*
4. Phosphatidylinositol 2%*
5. Sphingomyelin 2%*
6. Phosphatidylethanolamine 4%*
7. Unsaturated Phosphatidylcholine 17%*

** By molecular weight*

Surfactant proteins

Surfactant proteins are divided into 2 groups:

- **Large and water-soluble SP-A and SP-D proteins**
- **small, hydrophobic SP-B and SP-C proteins.**



Are of great importance to immune defense mechanisms of the lung

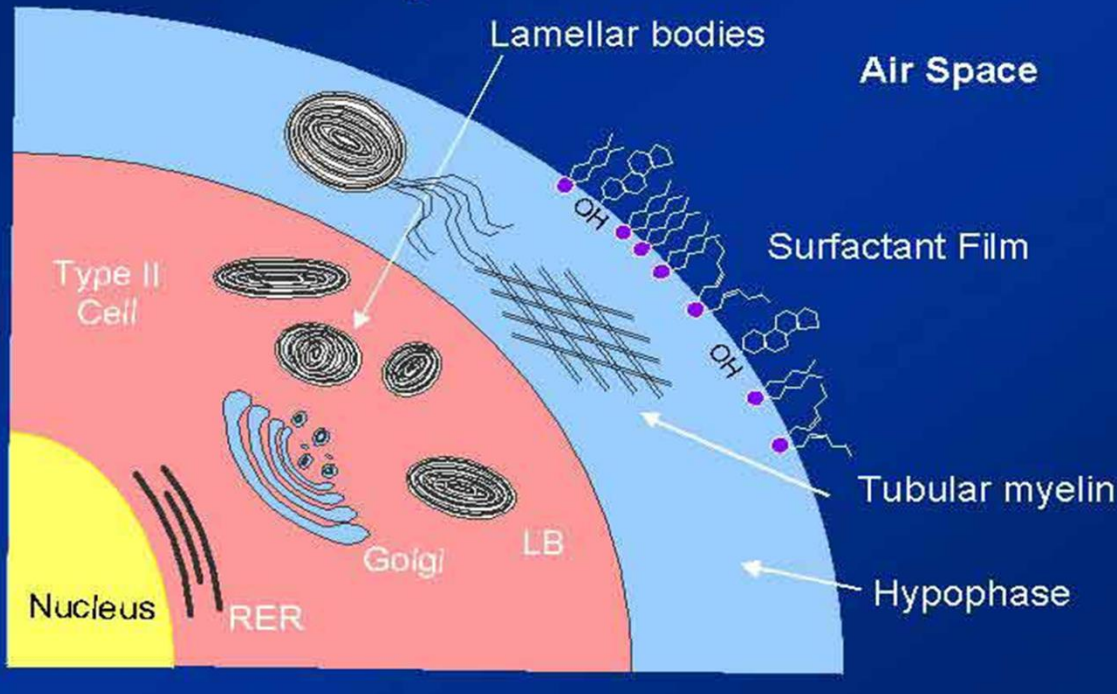
- ability to bind to bacteria, viruses and other pathogens

.....(

- well as to activate alveolar macrophages

Component

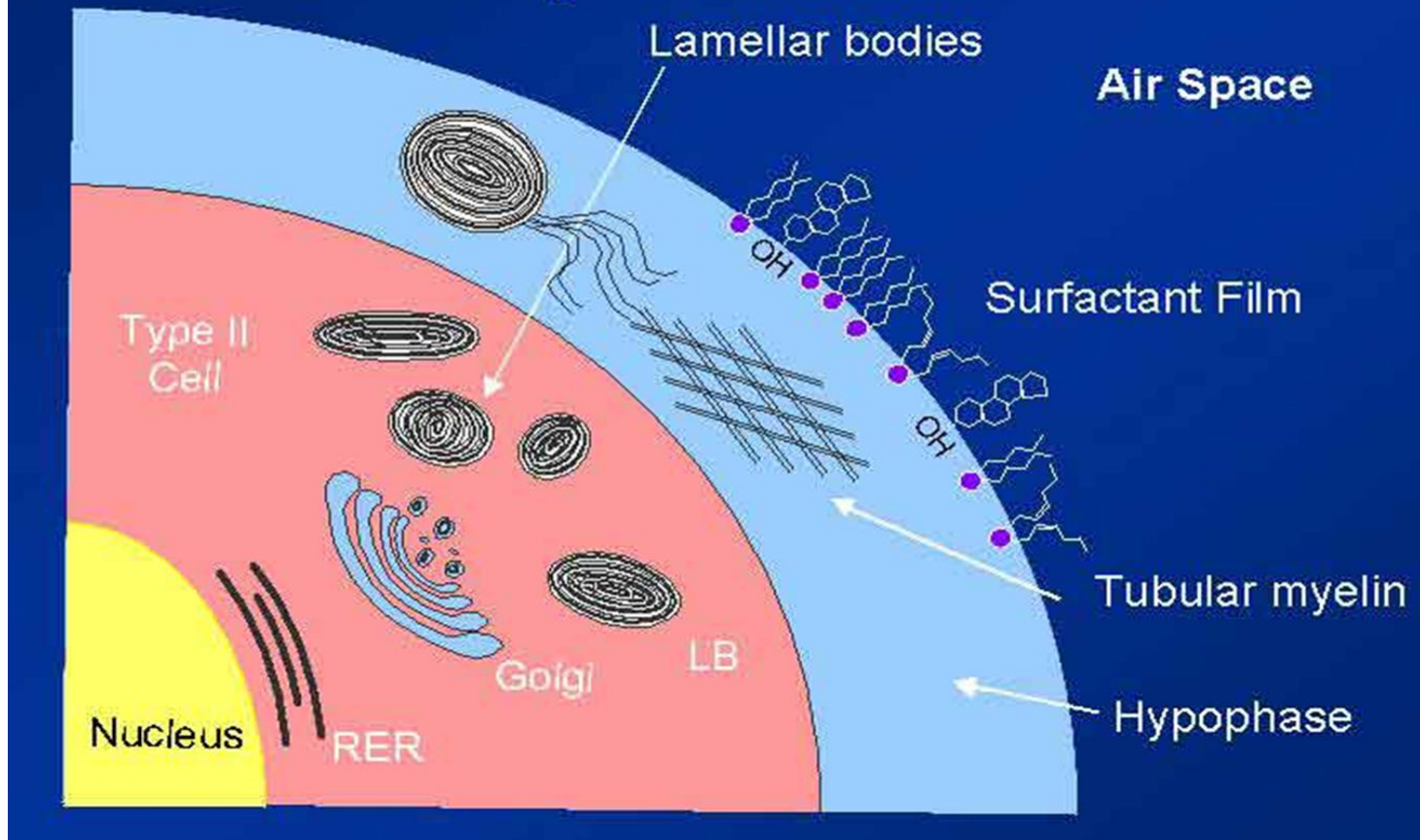
Surfactant Synthesis & Secretion



1-Lipid

- Synthesized in the smooth endoplasmic reticulum moved to Golgi apparatus

Surfactant Synthesis & Secretion

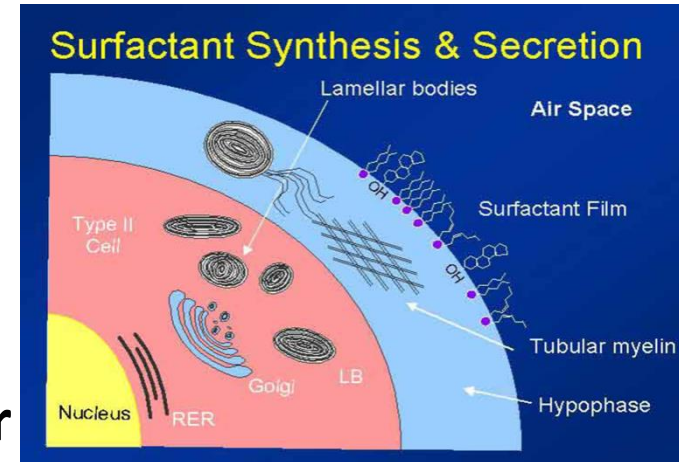


- **Surfactant** is synthesized by ***type II alveolar cells*** from fatty acids that either reach the lung from blood or formed (de novo) inside it. It is stored in organelles know as "***lamellar bodies***".

Component

Lipid

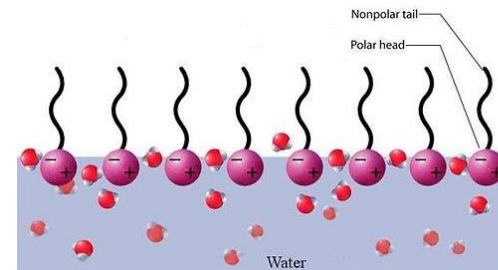
- The main constituent of the monolayer



dipalmitoylphosphatidylcholine (DPPC), which

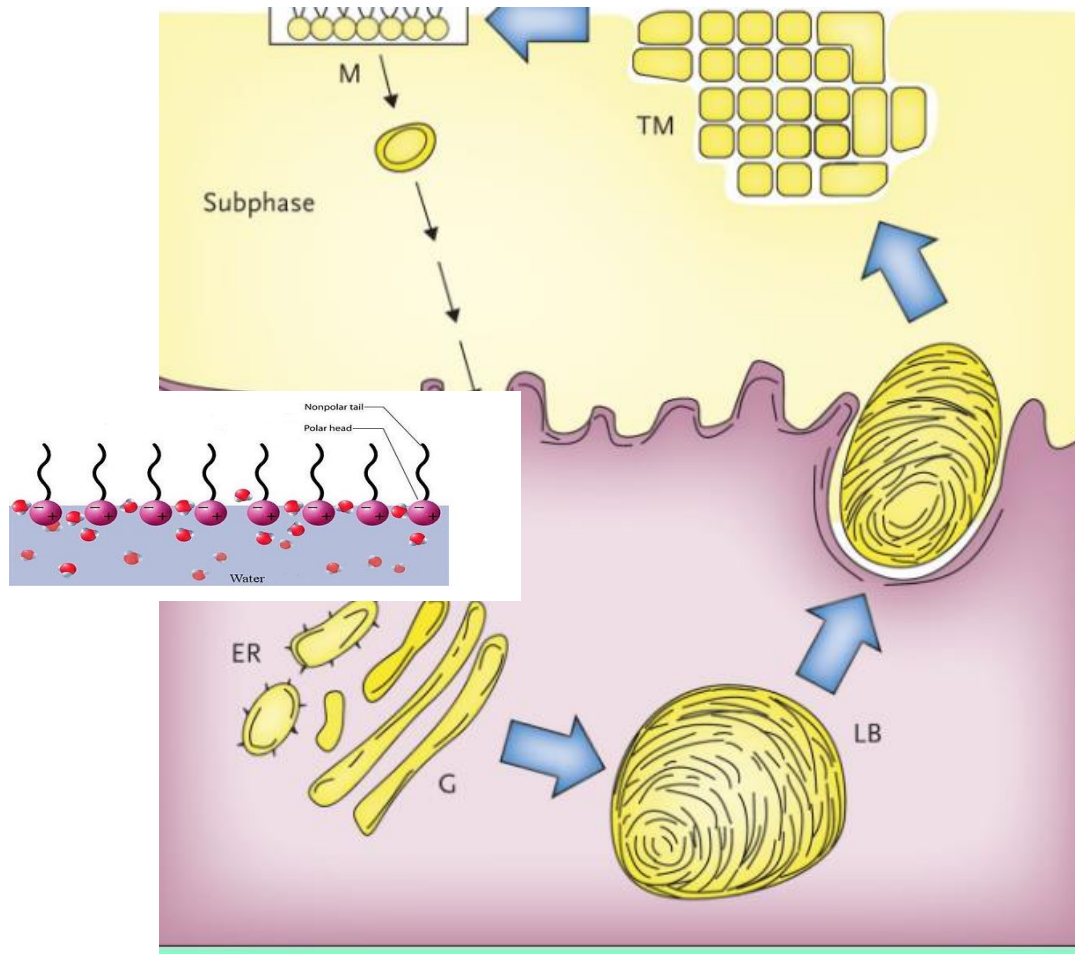
is a bipolar lipid (it has a hydrophilic 'head' and

a lipophilic 'tail')



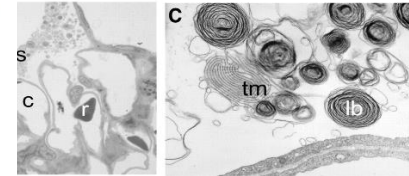
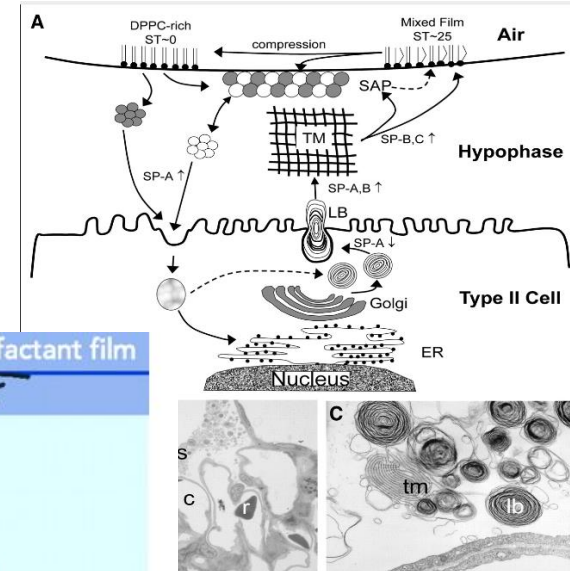
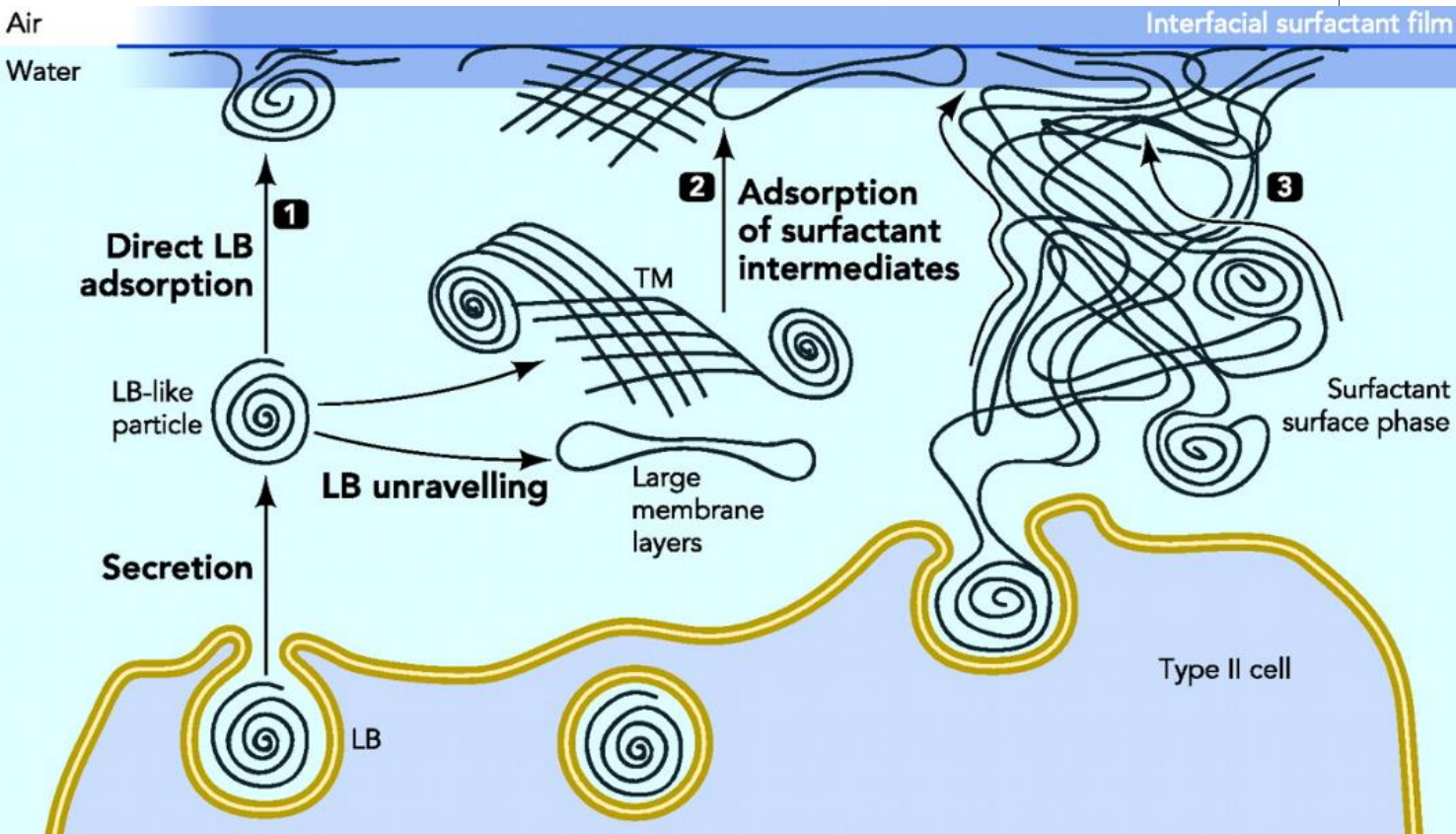
Surfactant synthesis

monomolecular surfactant



Surface Tension

- Surfactant **Lipoprotein complex** that lowered the surface tension synthesized by Type II



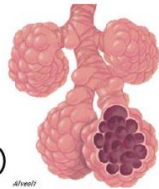
Functions of surfactant:

1-This decreased surface tension:

Roles of Lung surfactant

surfactant decreases surface tension

- ↑ pulmonary compliance
- ↓ alveolar collapse
- Respiratory distress syndrome (RDS)



Fetal lung maturity

- L/S ratio
- phosphatidylglycerol
- foam stability or shake test

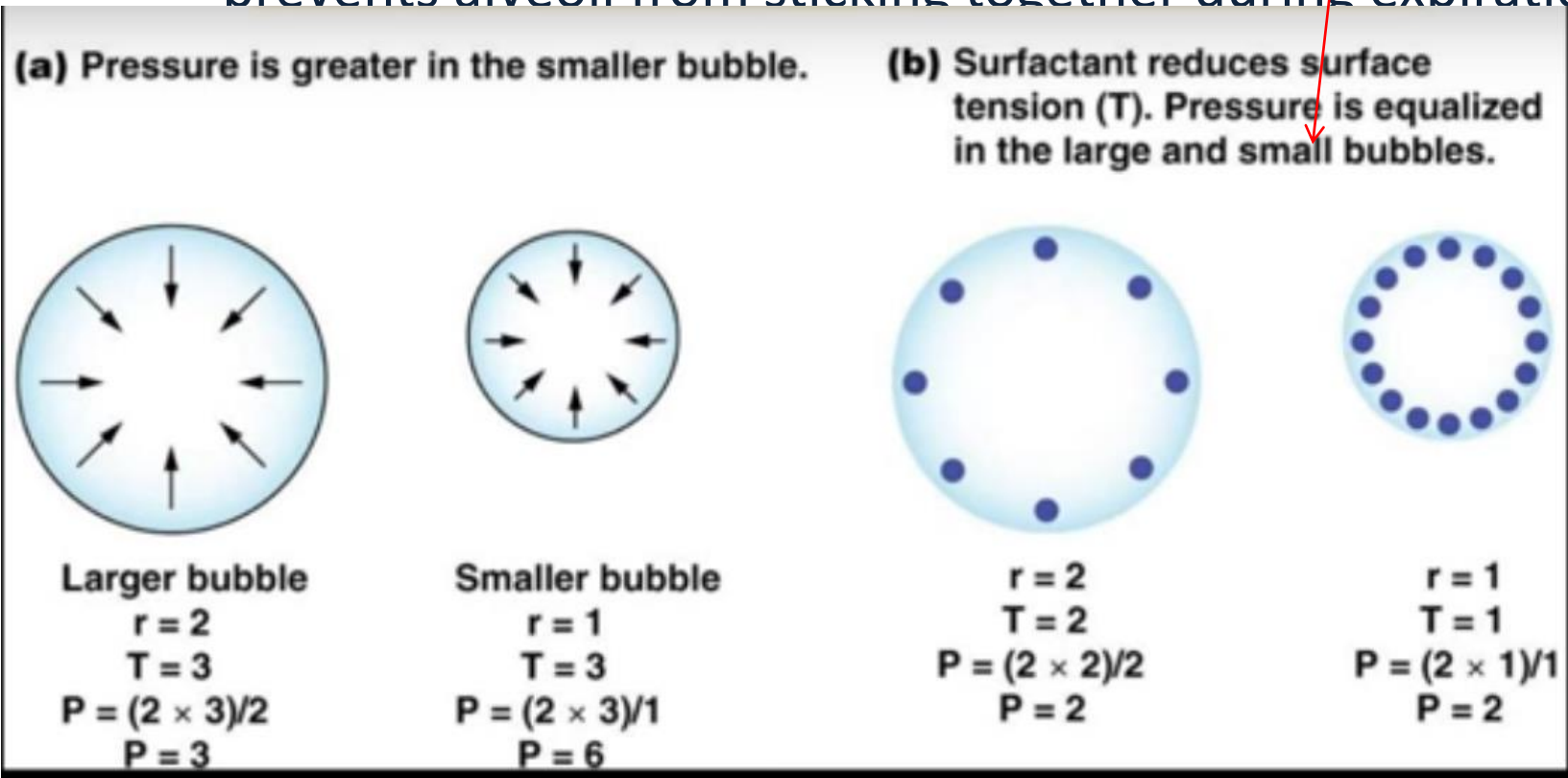
L/S < 1.5	immature
L/S 1.5-1.9	intermediate
L/S ≥ 2	lung maturity

TO TEST

prevention

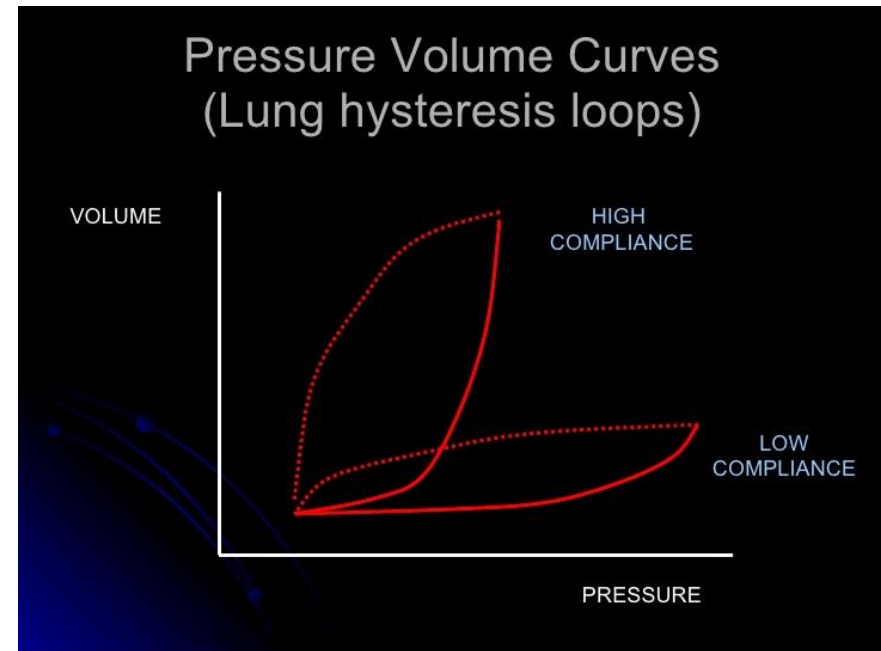
Ventilation in the presence of surfactant

- Disrupts the surface tension & cohesion of water molecules
- Impact?
 - prevents alveoli from sticking together during expiration



Functions of surfactant:

- This decreased surface tension:
 - Increase the lung compliance
 - Helps lung expansion during inspiration



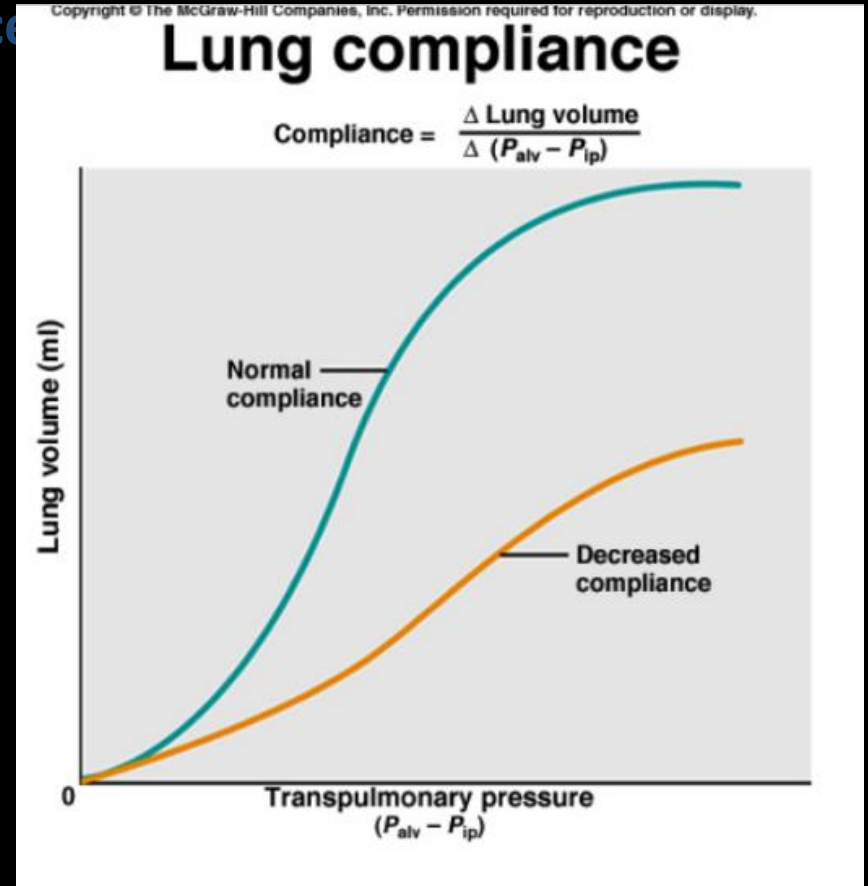
Functions of surfactant:

This decreased surface tension:

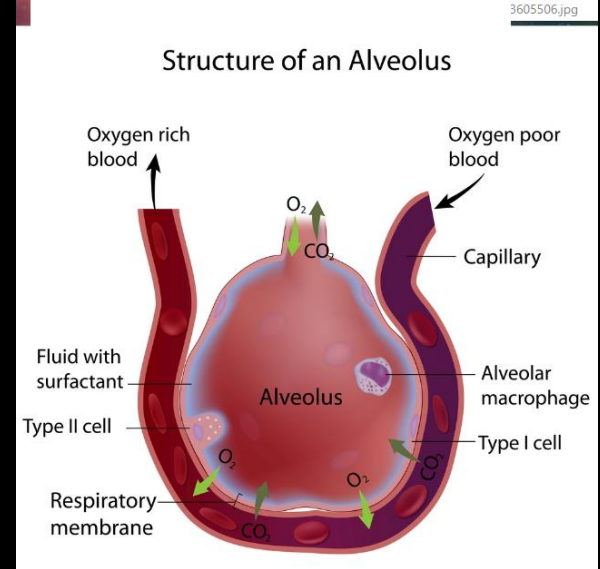
- Increase the lung compliance
 - Helps lung expansion during inspiration
 - stabilize the alveoli :

This protects the alveoli from

- ☐ Collapse during expiration
- ☐ over distention during inspiration
- ☐ Prevent collapse during expiration (atelectasis)



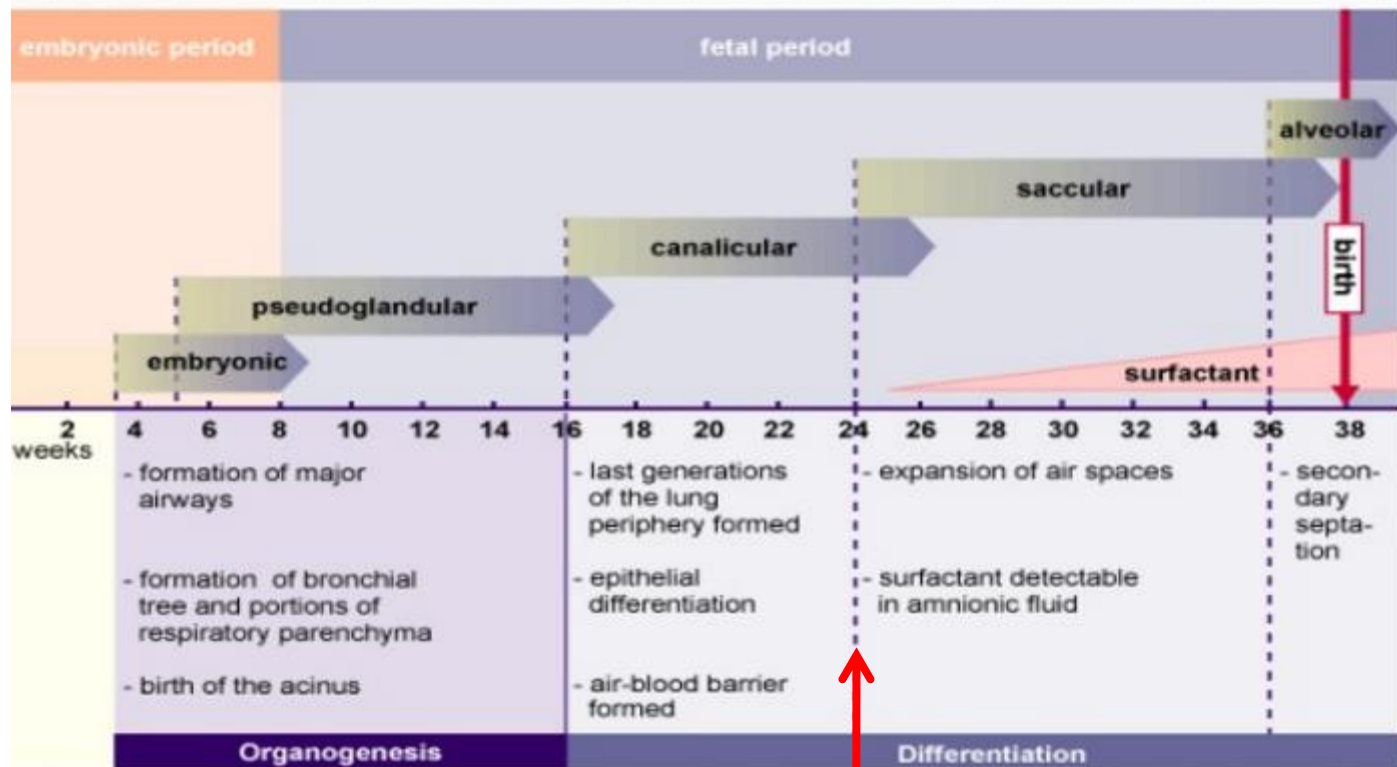
Functions of surfactant:



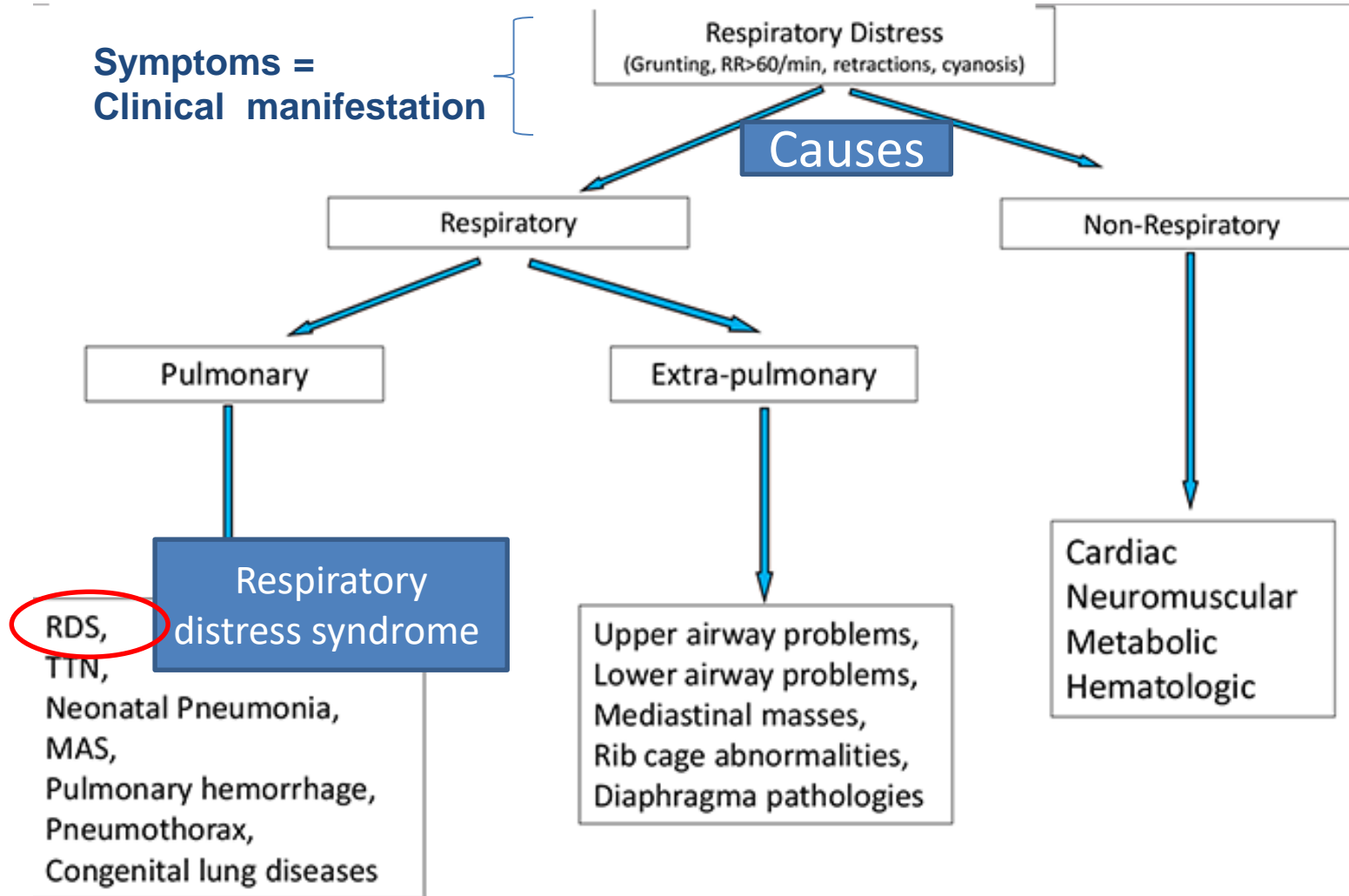
This decreased surface tension:

- **Protects against pulmonary edema as it decreases the filtration forces for the fluid from pulmonary capillaries into alveoli.**

Phases of Lung Development



Symptoms =
Clinical manifestation



Case

Define preterm

Gestation age < 37 weeks
from Last menstrual period

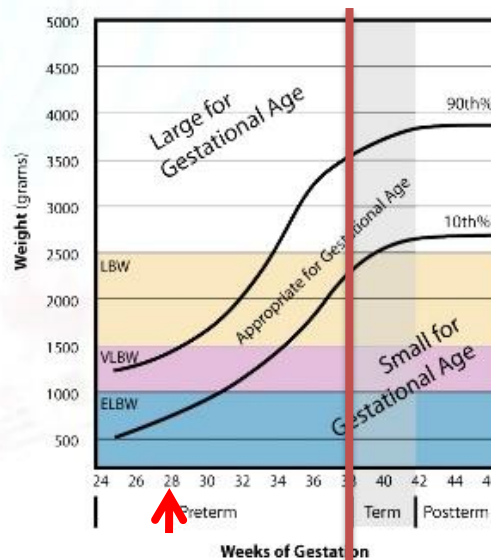
- Baby born preterm at 28 week



Gestational Age

Classification of Size

- LGA** • SGA- small for gestational age-weight below 10th percentile
- AGA** • AGA-weight between 10 and 90th percentiles (between 5lb 12oz (2.5kg) and 8lb 12 oz (4kg).
- SGA** • LGA-weight above 90th percentile
- IUGR-deviation in expected fetal growth pattern, caused by multiple adverse conditions, not all IUGR infants are SGA, may or may not be "head sparing"



What Next ?

Preterm baby
Excepted to
have RDS



CLINICAL MANIFESTATION

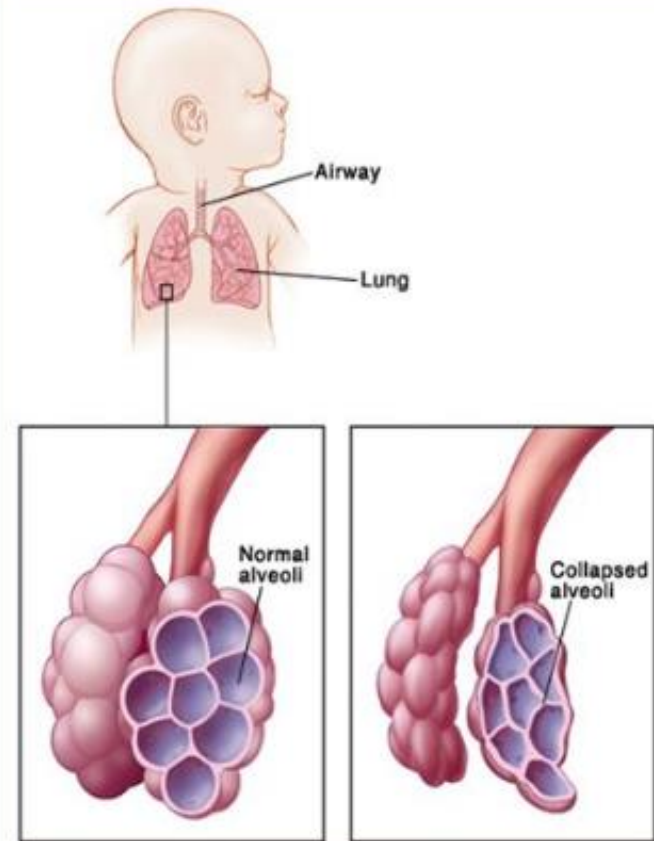
- ▶ Tachypnea
- ▶ Nasal flaring
- ▶ Intercostal, sternal recession
- ▶ Grunting; closure of glottis during expiration
- ▶ Cyanosis





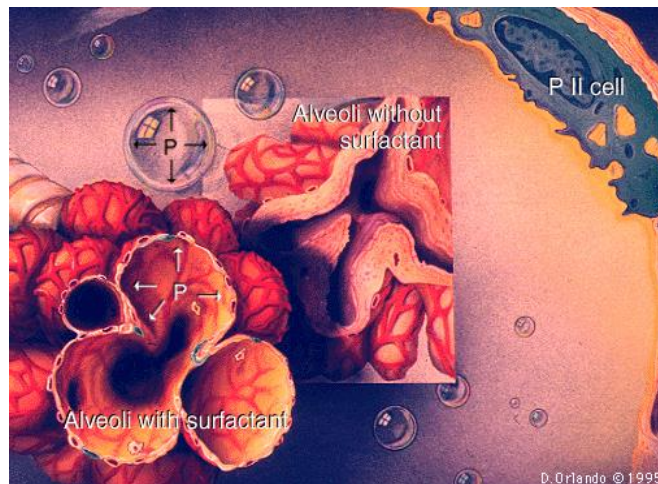
Respiratory Distress Syndrome (RDS)

- Also called hyaline membrane disease.
- Most common cause of respiratory distress in preterm infants.
- Due to structural and functional immaturity of lungs.
 - Underdeveloped parenchyma
 - Surfactant deficiency
 - Type II pneumatocytes
- Results in decreased lung compliance, unstable alveoli



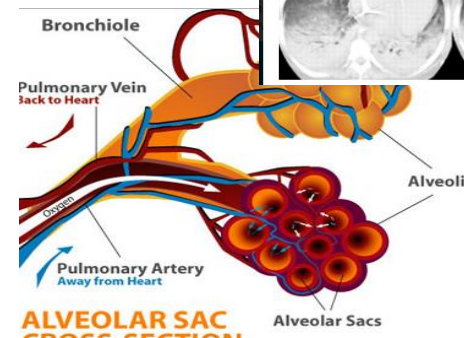
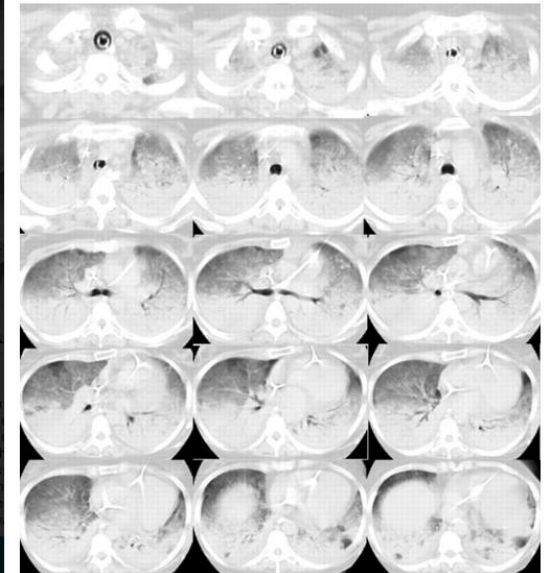
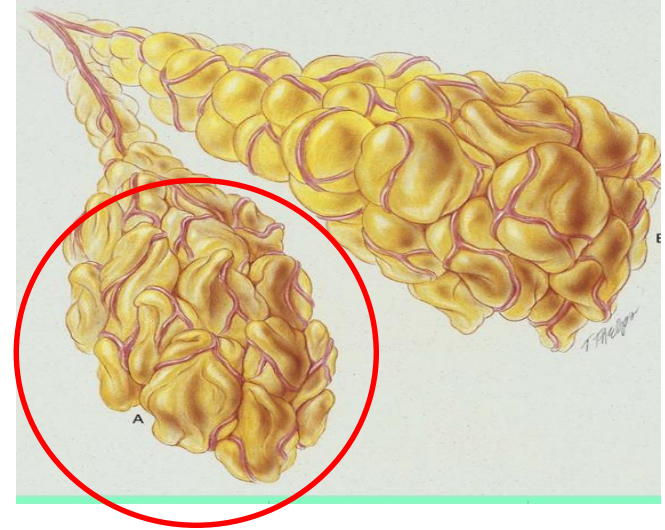
pathophysiology

- Instability of terminal airspaces (difficult to expand during inspiration and atelectasis at expiration) due to elevated surface forces at liquid-gas interfaces (elevated surface tension)



Diminished surfactant :

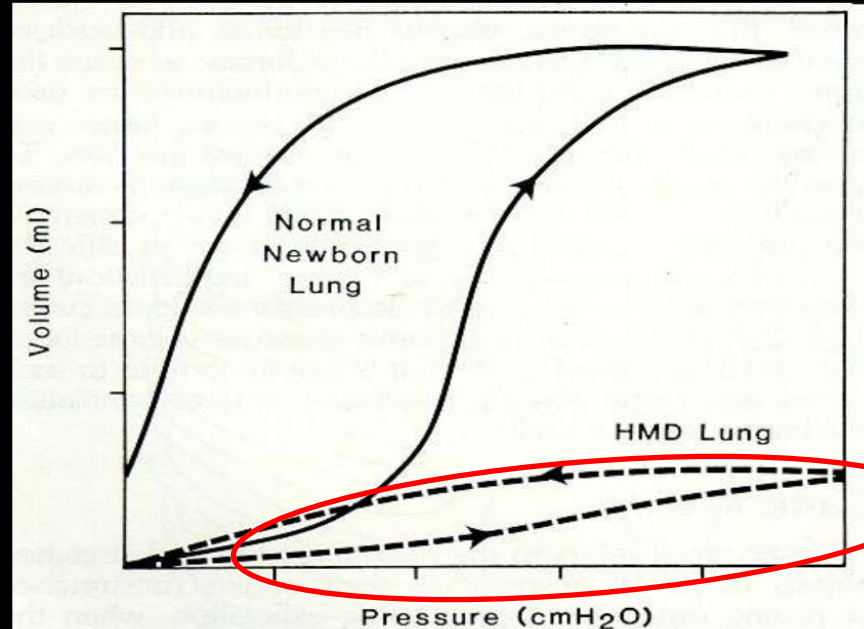
- Progressive Atelectasis
- Loss of functional residual capacity
 - Small lungs and small tidal volume
- Alterations in ventilation perfusion
- Uneven distribution of ventilation



Lung compliance in RDS

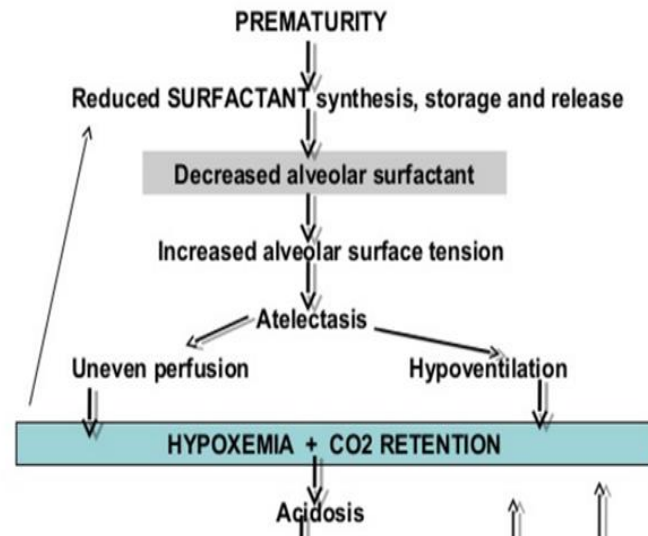
- Lung Compliance is also reduced: from 1-2 to 0.2 - 0.5 ml/cmH₂O/kg

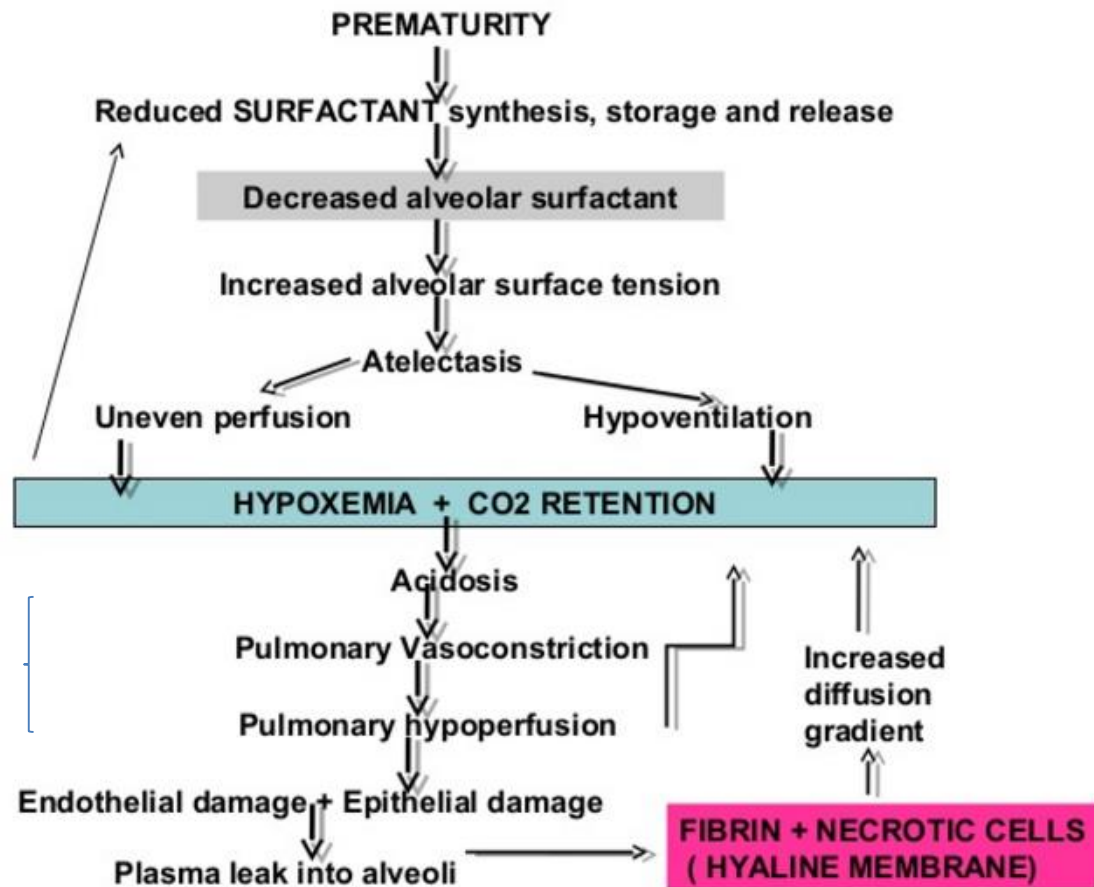
PRESSURE VOLUME LOOP



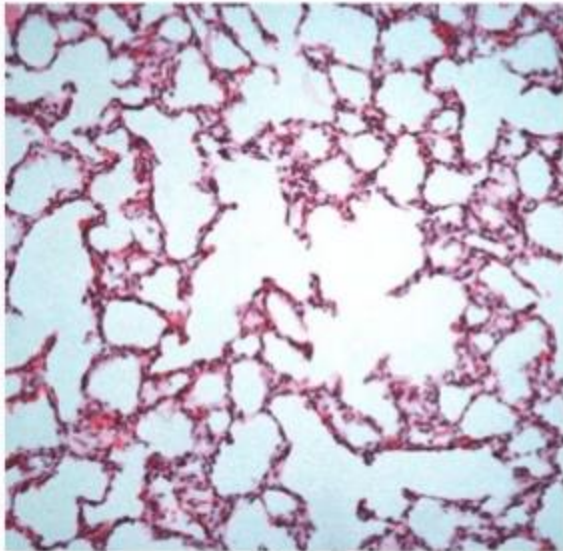
RDS: clinical picture

- At admission of the baby he has
 - Cyanosis
 - ↓ Pulse Oximeter 75% (normal > 95%)
 - Blood gas:
 - ↓ PaO₂ = 45 mmHg (normal 80-108)
 - ↑ Ph = 7.2 (normal 7.35-7.45)
 - CO₂ = 65 mmHg (normal 35-45)

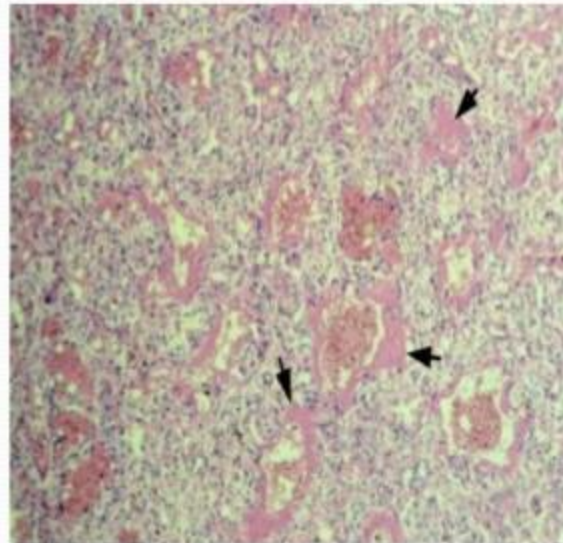




Normal Lung



Hyaline Membranes



➤ Hyaline membrane- combination of sloughed epithelium, protein & edema.

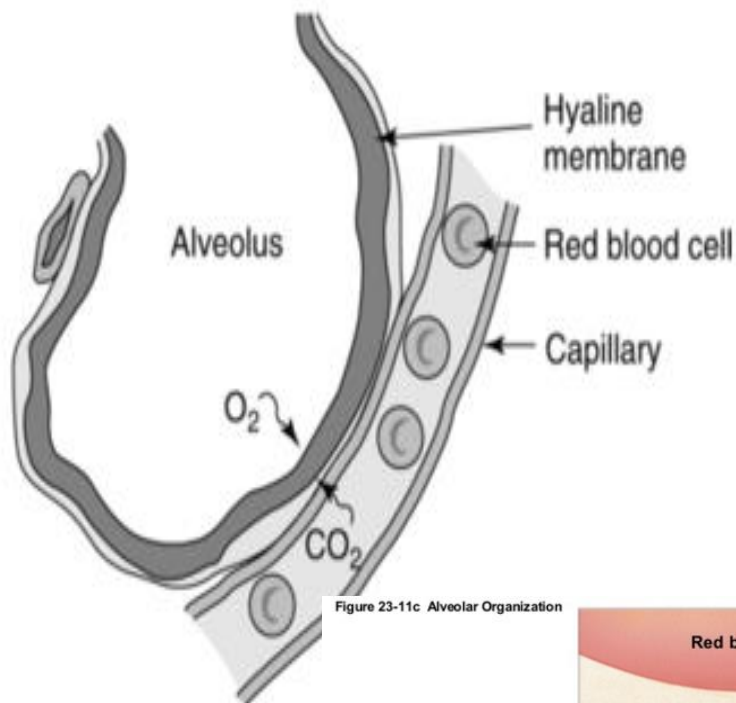
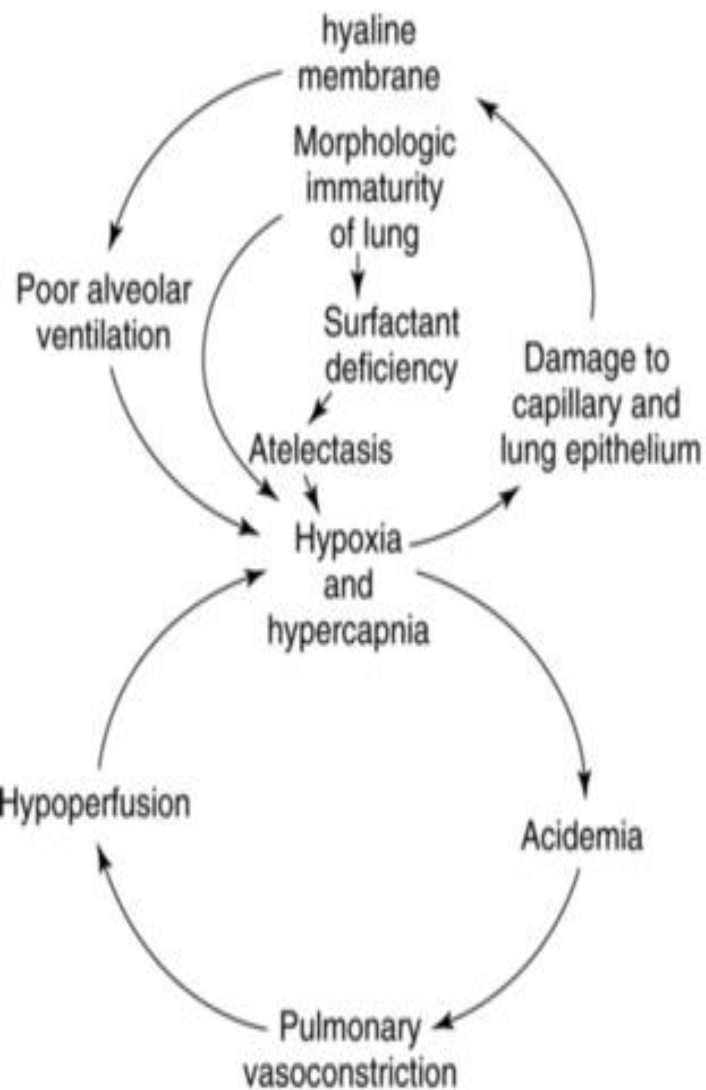
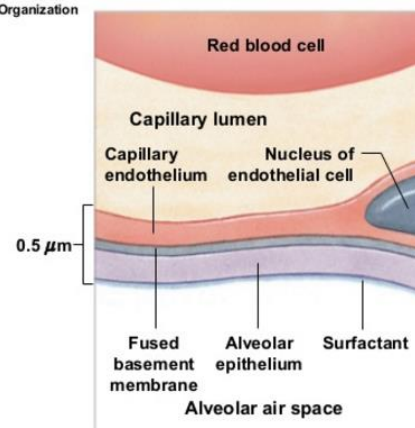
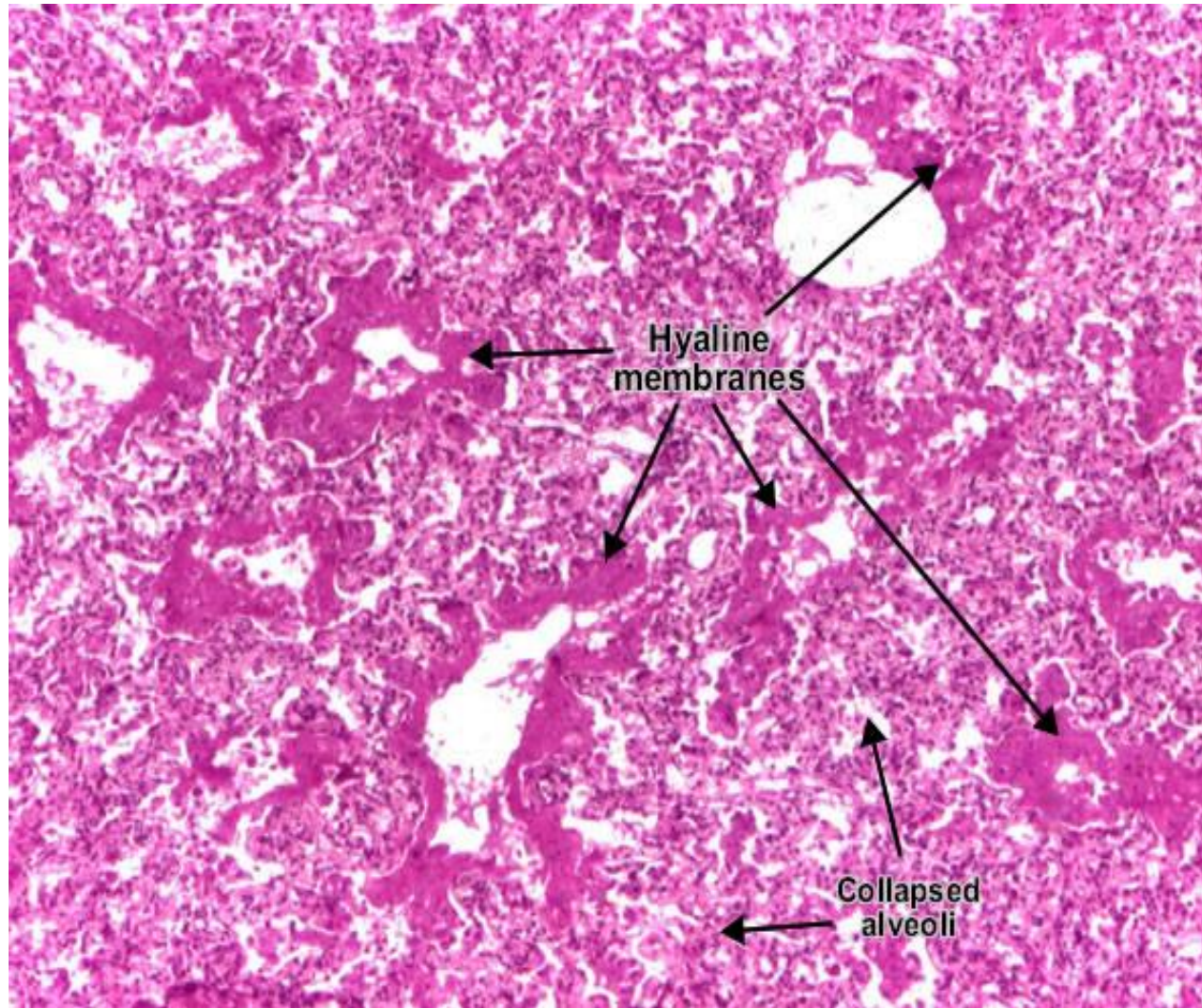


Figure 23-11c Alveolar Organization

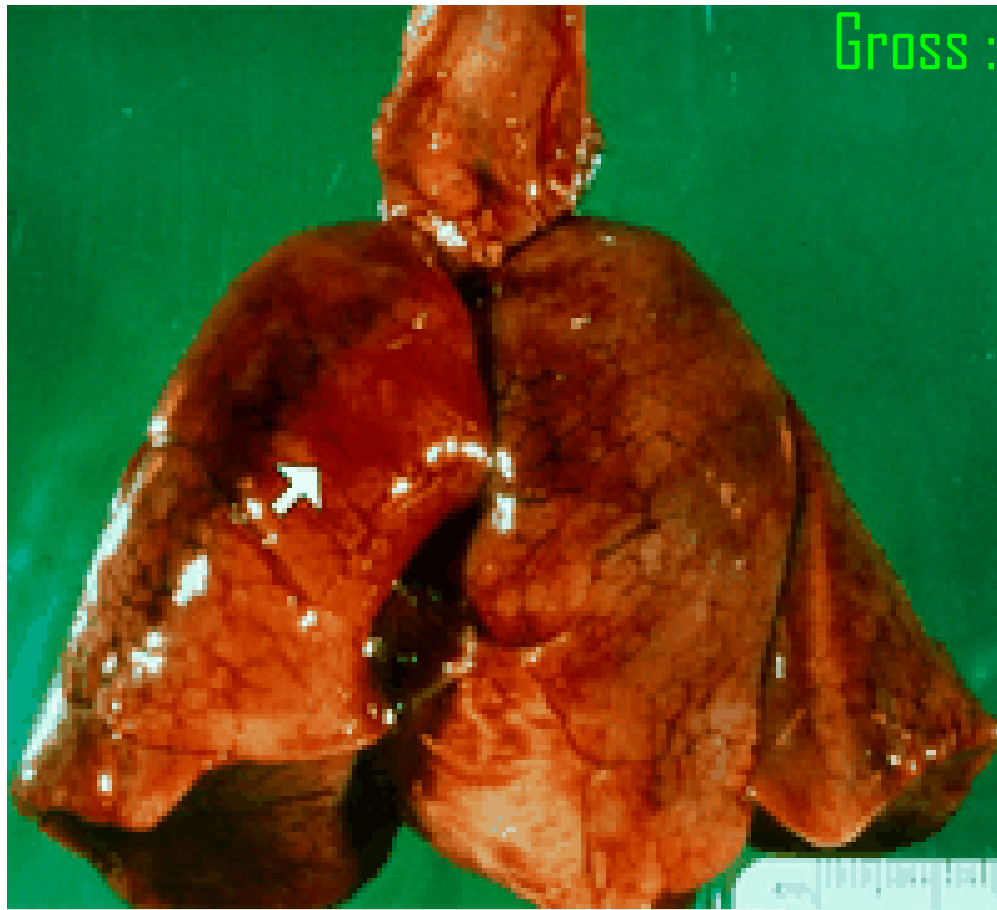


c The respiratory membrane, which consists of an alveolar epithelial cell, a capillary endothelial cell, and their fused basement membranes.

➤ Hyaline membrane- combination of sloughed epithelium, protein & edema.



- Hyaline membrane- combination of sloughed epithelium, protein & edema.



- Photograph of an autopsy specimen demonstrates small atelectatic lungs with focal hemorrhage (arrow) visible on the pleural surface.

Incidence

Respiratory Distress Syndrome (RDS)

- Also known as Hyaline Membrane Disease (HMD)
- Commonest cause of preterm neonatal mortality
- RDS occurs primarily in premature infants; its incidence is inversely related to gestational age and birth weight

Gestational age	Percentages
Less than 28 wks	60-80%
32-36 wks	15-30%
37-39 wk	5%
Term	Rare

Nelson Textbook of Pediatrics, 18th Ed.

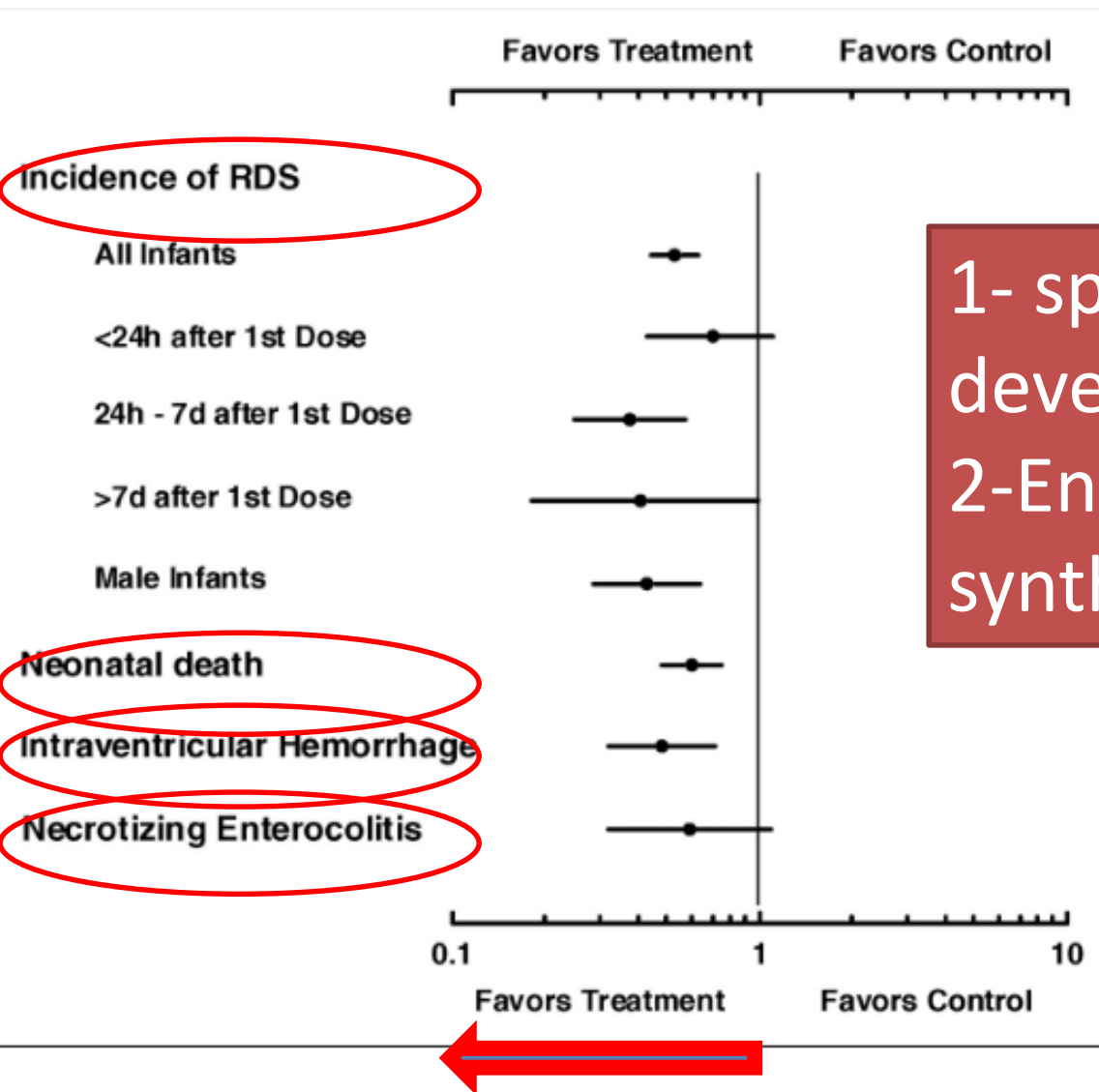
Risk Factors

<u>Increased Risk</u>	<u>Decreased Risk</u>
<ul style="list-style-type: none">• Maternal diabetes• multiple births• cesarean section delivery• perinatal asphyxia• cold stress• history of previously affected infants	<ul style="list-style-type: none">• Chronic or pregnancy-associated hypertension• maternal heroin use• prolonged rupture of membranes• antenatal corticosteroid prophylaxis

Genetic Predisposition to RDS

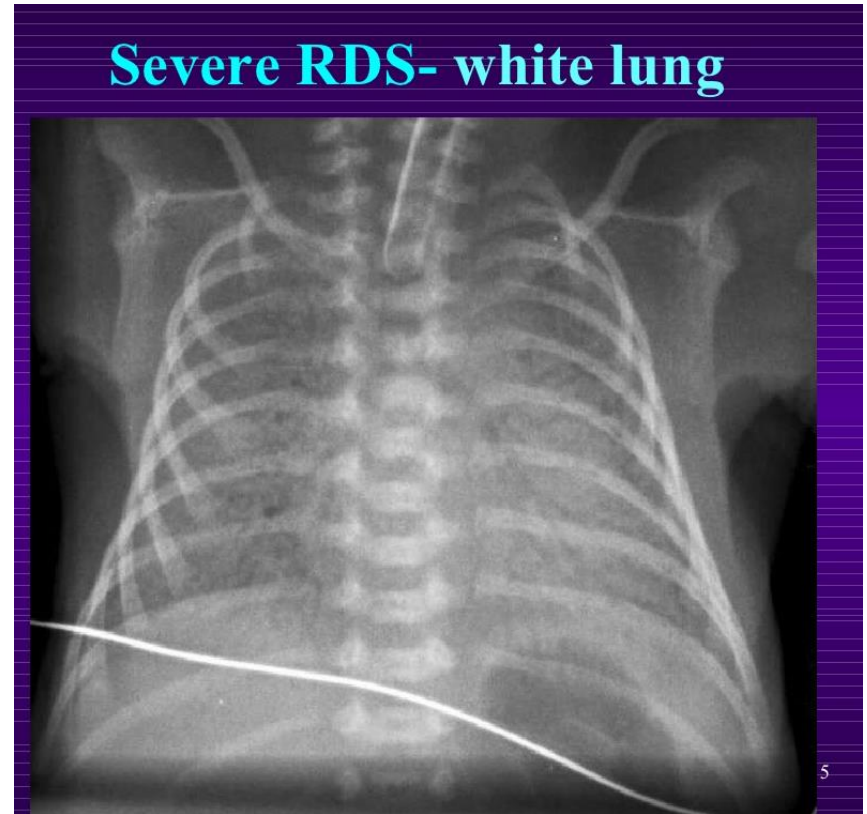
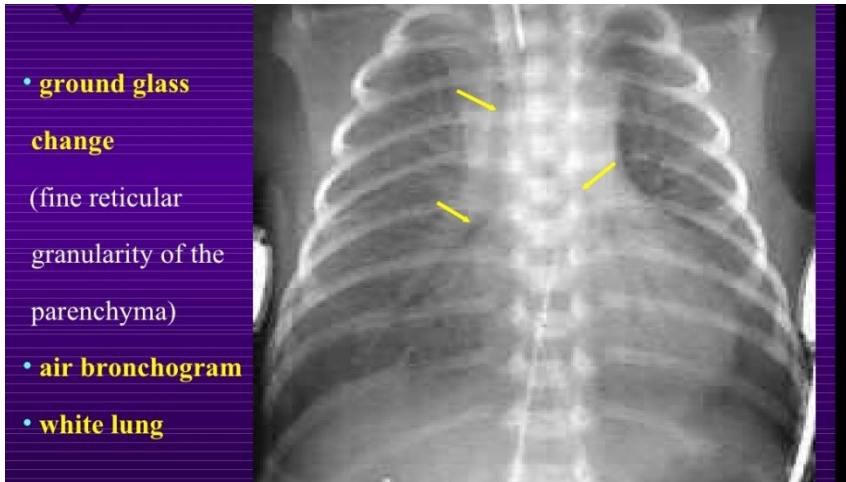
- Susceptibility to RDS is interaction between genetic, environmental and constitutional factors
- ***Very preterm infants***
 - Common alleles predicts RDS: **SP-A** 642, Sp-B121, Sp-C 186 ASN.
- Term Infants: Loss of function mutation of **SP-B, SP-C**, phospholipids transporter ABCA3

Antenatal Corticosteroid Effects



1- speed up lung development
2- Enhance surfactant synthesis

CXR



➤ Chest radiograph: air bronchogram, reticular/ ground-glass appearance after 6-12 hrs to full opacity later on.



Grade 4 - severe case, complete white-out of the lung fields with obscuring of the cardiac border

Prevention

- Prevention of prematurity
- Antenatal corticosteroid therapy

Dexamethasone or betamethasone

↓RDS morbidity and mortality

- PS prophylactic therapy

RDS - Treatment

- Oxygen
 - CPAP
 - Mechanical ventilation
 - Surfactant replacement
 - Supportive Care
-

Respiratory support



Treatment

Oxygen therapy and assist ventilation

1. oxygen therapy

- nasal cannula, mask or headbox oxygen
- keep PaO_2 50-70mmHg, S_aO_2 90-95%

2. CPAP (continuous positive airway pressure)

- Prevent alveolar collapse at end expiration
- Indication: $\text{FiO}_2 > 0.4$, $\text{PaO}_2 < 50$ mmHg or $\text{S}_a\text{O}_2 < 85\%$
- Pressure: 4-6 cmH_2O

CPAP





PS replacement therapy



PS

29

Antenatal steroid and Surfactant goes hand in hand

