

M.SC Semester III
Core Course XI
Bio-Inorganic Chemistry



TOPIC:-Unit III, Myoglobin

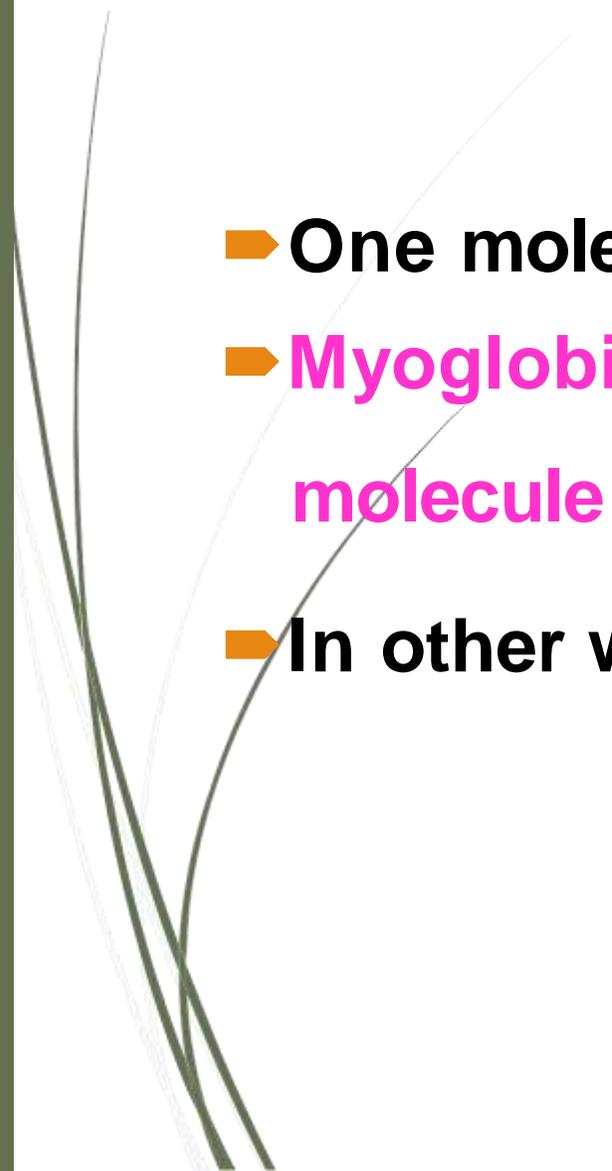
Department of Chemistry
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Myoglobin

- **Myoglobin (Mb) is monomeric O₂ binding hemoprotein**
- **Found in heart and skeletal muscle.**
- **It has single polypeptide (153A.As) chain with heme moiety.**
- **Myoglobin (mol. wt. 17,000) structurally resembles the individual subunits of hemoglobin molecule**
- **Myoglobin functions as a reservoir for oxygen.**
- **It serves as oxygen carrier that promotes the transport of oxygen to the rapidly respiring muscle cells**



Binding of O₂ to haemoglobin

- One molecule of Hb can bind with four molecules of O₂.
 - Myoglobin (with one heme) which can bind with only one molecule of oxygen.
 - In other words, each heme moiety can bind with one O₂.
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Transport of O₂ by haemoglobin

- It can transport large quantities of oxygen
- It can take up and release oxygen at appropriate partial pressures
- It is a powerful buffer.

Oxygen Dissociation Curve (ODC)

- ▶ **The binding ability of hemoglobin with oxygen at physiological pO₂ (partial pressure of oxygen) is shown by the oxygen dissociation curve (ODC)**
- ▶ **At the oxygen tension in the pulmonary alveoli, the Hb is 97% saturated with oxygen.**

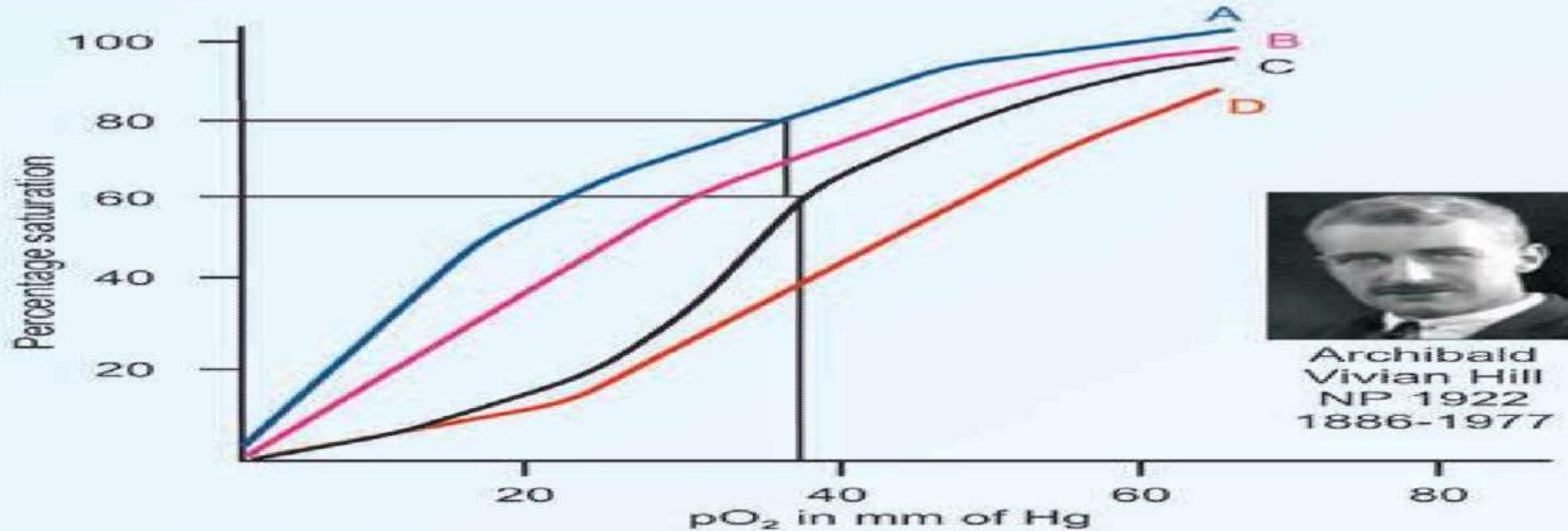
**M.SC Semester III
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TOPIC:-Unit III, Oxygen dissociation curve (ODC)

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Oxygen dissociation curve (ODC)



Archibald
Vivian Hill
NP 1922
1886-1977

A—Theoretical curve as per mass action.
B—Sigmoid curve, due to heme-heme interaction (Hill effect). **C**—Further shift to right due to carbon dioxide (Bohr effect) and BPG. This curve represents the pattern under normal conditions. **D**—Further shift to right when temperature is increased to 42°C.



Christian
Bohr
(1855-1911)

Factors affecting oxygen dissociation curve

- ▶ **Heme-heme Interaction & Cooperativity:**
- ▶ **The oxygen dissociation curve (ODC) is sigmoid shape.**
- ▶ **The binding of O₂ to one heme residue increases the affinity of remaining heme residues for O₂.**
- ▶ **Thus the affinity of Hb for the last O₂ is about 100times greater than the binding of the first O₂ to Hb.**
- ▶ **This is called positive cooperativity**

Release of O₂ from one heme facilitates the release of O₂ from others.

The quaternary structure of **oxy-Hb** is described as **R (relaxed) form**; & **deoxy-Hb** is **T(tight) form**.

$2\alpha + 2\beta$
(Deoxy-Hb – T-form)



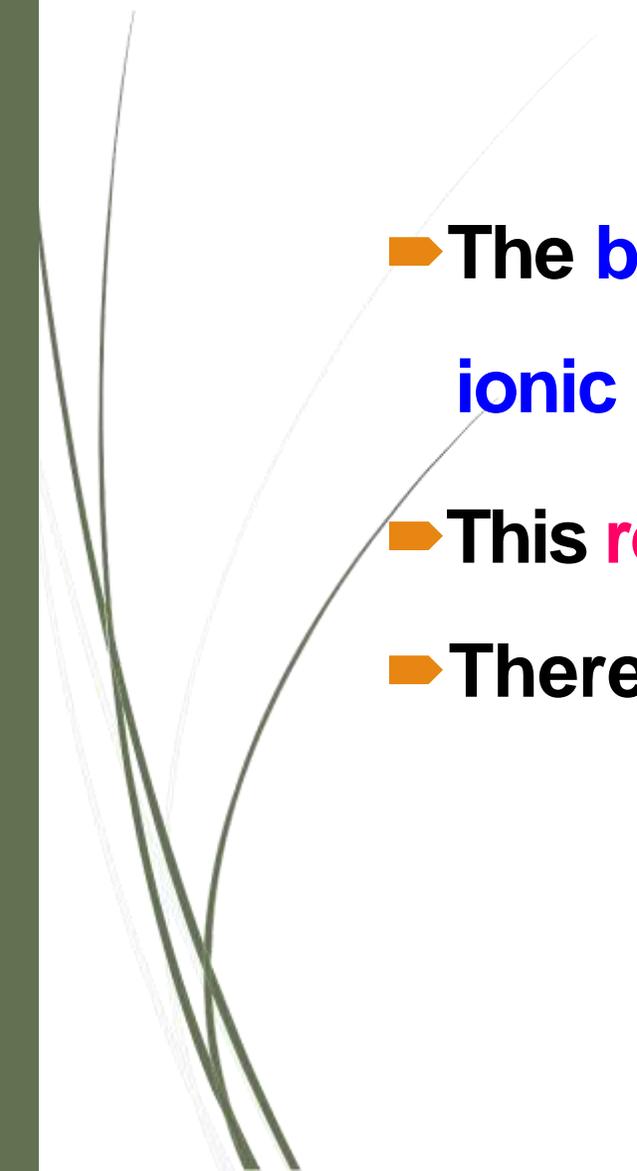
$2\alpha, \beta$
(Oxy-Hb – R-form)

Tand R forms of hemoglobin

- The four subunits ($\alpha_2\beta_2$) of hemoglobin are held together by weak forces.
- The relative position of these subunits is different in oxyhemoglobin compared to deoxyhemoglobin.
- T-form of Hb:
- The deoxy form of Hb exists in T or taut (tense) form.
- The H & ionic bonds limit the movement of monomers.
- The T-form of Hb has low oxygen affinity.

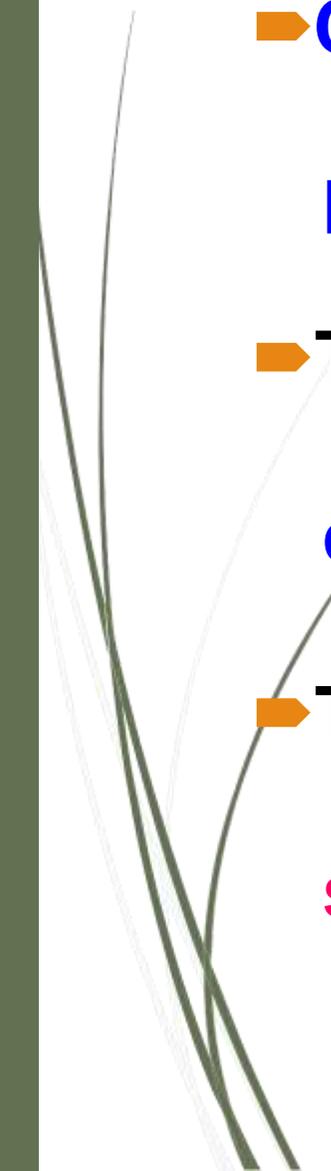


R-form of Hb

- **The binding of O₂ destabilizes some of the hydrogen & ionic bonds particularly between $\alpha\beta$ dimers.**
 - **This results in a relaxed form or R-form of Hb**
 - **Therefore, the R-form has high oxygen affinity.**
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Transport of CO₂ by hemoglobin

- In aerobic metabolism, for every molecule of O₂ utilized, one molecule of CO₂ is liberated.
- Hemoglobin actively participates in the transport of CO₂ from the tissues to the lungs.
- About 15% of CO₂ carried in blood directly binds with Hb.
- The rest of the tissue CO₂ is transported as bicarbonate (HCO₃).

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- ➔ **CO₂ molecules are bound to the uncharged α-amino acids of hemoglobin to form carbamyl hemoglobin.**
 - ➔ **The oxyHb can bind 0.15 moles CO₂/mole heme, whereas deoxyHb can bind 0.40 moles CO₂/mole heme.**
 - ➔ **The binding of CO₂ stabilizes the T(taut) form of hemoglobin structure, resulting in decreased O₂ affinity for Hb.**
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Hemoglobin also helps in the transport of CO₂ as bicarbonate

- ➔ **CO₂ enters the blood from tissues, the enzyme carbonic anhydrase present in erythrocytes catalyses the formation of carbonic acid (H₂CO₃).**
- ➔ **Bicarbonate (HCO₃⁻) & proton (H⁺) are released on dissociation of carbonic acid**
- ➔ **Hb acts as a buffer & immediately binds with protons**

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- **Every 2 protons bound to Hb, 4 oxygen molecules are released to the tissues.**
 - **In the lungs, binding of O₂ to Hb results in the release of protons.**
 - **The bicarbonate & protons combine to form carbonic acid.**
 - **Acted upon by carbonic anhydrase to release CO₂, which is exhaled**