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## Dissociation Curve



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## Dissociation Curve

- Oxy-Haemoglobin Dissociation Curve
- Bohr's Effect
- $\mathrm{CO}_{2}$ Equilibration Curve
- C-D-H effect


## Oxy-Haemoglobin Dissociation Curve

- The loading and unloading of $\mathrm{O}_{2}$ from Hb is described by oxy-haemoglobin dissociation curve.
- If Hb is allowed to equilibrate with various partial pressures of $\mathrm{O}_{2}$ and the values are expressed in a graph plotted between the percentages of Hb on the ordinate against the partial pressure of $\mathrm{O}_{2}$ on the abscissa, the curve obtained is called oxyhaemoglobin dissociation curve.
- For oxy-Hb the curve is " S " shaped



## Dissociation Curve



- For myoglobin (a muscle pigment capable of combining with $\mathrm{O}_{2}$ ) the dissociation curve is rectangular hyperbola
- Under normal conditions, at a $\mathrm{PO}_{2}$ of $100-\mathrm{mm} \mathrm{Hg}$, blood leaving the lungs is $95-98 \%$ saturated with $\mathrm{O}_{2}$. Further increase in $\mathrm{PO}_{2}$ do not increase $\mathrm{O}_{2}$ carrying capacity of blood and the increased $\mathrm{PO}_{2}$ increase the amount of $\mathrm{O}_{2}$ in physical solution according to Henry's law.
- Because the hemoglobin is almost saturated when it leaves the lungs, it is the hemoglobin concentration that determines the amount of $\mathrm{O}_{2}$ transported in blood


## Volume of $\mathrm{O}_{2}$ combined with haemoglobin

- One molecule of Hb can combine with 4 molecules of $\mathrm{O}_{2}$
- One gram of Hb can transport 1.34 ml of $\mathrm{O}_{2}$
- The volume of $\mathrm{O}_{2}$ combined with Hb in each 100 ml blood is = Haemoglobin concentration ( $\mathrm{g} \%$ ) X volume of $\mathrm{O}_{2}$ in each gram of $\mathrm{Hb}(\mathrm{ml} / \mathrm{g}) \times$ Oxygen saturation (decimal) at the partial pressure of measurement.
- If $\mathrm{Hb}=15 \mathrm{~g} \%, \mathrm{O}_{2}$ saturation $97.5 \%$,
- then 100 ml blood can transport 15 X 1.34 X $0.975=19.6 \mathrm{ml} / 100 \mathrm{ml}$ or 19.6 volumes percent.
- 100 ml of blood carries 19.9 ml of $\mathrm{O}_{2}$ i.e. 19.6 ml in combination with haemoglobin and 0.3 ml in physical solution ( $98-99 \mathrm{HO}_{2}$ is transported in combination with haemoglobin).
- If Hb were not present, it would take 66.3 times more blood to transport the same amount of $\mathrm{O}_{2}$.
- At a $\mathrm{PO}_{2}$ of 100 mmHg in arterial blood, Hb is $97.5 \%$ saturated with $\mathrm{O}_{2}$ and can transport 19.6 volumes percent $\mathrm{O}_{2}$, when Hb concentration is $15 \mathrm{~g} / 100 \mathrm{ml}$ blood.
- As the arterial blood reaches the tissue, $\mathrm{O}_{2}$ is unloaded from the blood to tissues, the $\mathrm{O}_{2}$ saturation falls to about $72 \%$ in venous blood.
- At $72 \%$ saturation of $\mathrm{O}_{2}$, the blood will have $14.5 \mathrm{ml} \mathrm{O}_{2}$ ( $\mathrm{Hb} .15 \mathrm{~g} \%$ ). Each $100-\mathrm{ml}$ blood unloads approximately 5 volume percent of $\mathrm{O}_{2}$ and this is called arteriovenous $\mathrm{O}_{2}$ difference.
- $\mathrm{PO}_{2}$ of venous blood is 40 mm Hg .
- $\mathrm{P}_{50}$ of hemoglobin is that $\mathrm{PO}_{2}$ at which haemoglobin is $50 \%$ saturated with $\mathrm{O}_{2}$. In human it is 26.6 mmHg . It indicates affinity of hemoglobin for $\mathrm{O}_{2}$
- $\mathrm{P}_{50}$ is similar for all Hb concentrations.
- Shift of dissociation curve to right results is greater release of $\mathrm{O}_{2}$ from oxy Hb , i.e. a shift to right decrease the affinity of Hb to $\mathrm{O}_{2}$.
- A shift to left increases the affinity of Hb to $\mathrm{O}_{2}$. Hence, $\mathrm{O}_{2}$ released from Hb is decreased.
- The extent of dissociation depends on (i.e.,) the positioning of the oxy haemoglobin dissociation curve is influences by.
$-\mathrm{O}_{2}$ tension
- $\mathrm{CO}_{2}$ tension
- $\mathrm{H}^{+}$ion concentration
- Temperature
- Concentration of BPG (2-3 bisphosphoglycerate) in erythrocytes
- Increase in $\mathrm{H}^{+}$ion concentration and $\mathrm{CO}_{2}$ level shifts the curve to down and right as also increase in temperature and 2-3 BPG.
- The 2-3 BPG is normally present in erythrocytes in higher level than in other cells, it is a by product of glycolytic pathway and its level increases further during chronic hypoxia as during exposure to high altitude, anemia, increased physical exertion etc. When 2-3 BPG binds with hemoglobin, affinity of hemoglobin for $\mathrm{O}_{2}$ is reduced and $\mathrm{O}_{2}$ unloading is increased. Shifting the curve to down and right causes increase in release of $\mathrm{O}_{2}$ from the Hb . Ruminant hemoglobin is unresponsive to 2-3 BPG.
- When metabolic rate of a tissue is increased, both $\mathrm{pCO}_{2}$ and $\mathrm{H}^{+}$ concentration increases; increased metabolic rate also increases heat production and temperature.
- The reduction in pH and rise in temperature reduces the affinity of hemoglobin for $\mathrm{O}_{2}$ shifting the oxy-hemoglobin curve to the right, thereby increasing the unloading of $\mathrm{O}_{2}$ at the tissue level. When the pH is reduced from 7.4 to 7.2 , the hemoglobin saturation is reduced from 72 to $60 \%$ indicating greater unloading of $\mathrm{O}_{2}$.
- The shift of oxy-Hb dissociation curve to down and right by increased $\mathrm{CO}_{2}$ tension and $\mathrm{H}^{+}$ion concentration is termed as Bohr's effect. An increasing concentration of $\mathrm{H}^{+}$and/or $\mathrm{CO}_{2}$ will reduce the affinity of haemoglobin to $\mathrm{O}_{2}$.
- This facilitates more $\mathrm{O}_{2}$ unloading in the tissues especially in tissues with greater demand for $\mathrm{O}_{2}$ like when a tissue's metabolic rate is increased with increase in $\mathrm{CO}_{2}$ production.
- The pH of the tissue decreases, and it promotes the dissociation of $\mathrm{O}_{2}$ from hemoglobin to the tissue, allowing the tissue to obtain enough $\mathrm{O}_{2}$ to meet its demands.
- Conversely, in the lungs, where $\mathrm{O}_{2}$ concentration is high, binding of $\mathrm{O}_{2}$ causes hemoglobin to release $\mathrm{H}^{+}$, which combines with $\mathrm{HCO}_{3}$ to drive off $\mathrm{CO}_{2}$ to alveoli.
- Since these two reactions are closely matched, there is little change in blood pH.


## $\mathrm{CO}_{2}$ Equilibration Curve

- The total quantity of $\mathrm{CO}_{2}$ combined with blood in all forms of transport of $\mathrm{CO}_{2}$ depends on $\mathrm{PCO}_{2}$, which can be expressed through the $\mathrm{CO}_{2}$ equilibration curve.
- Normal blood $\mathrm{PCO}_{2}$ is 40 mm Hg in arterial blood containing 48 volumes percent $\mathrm{CO}_{2}$.
- In venous blood $\mathrm{PCO}_{2}$ is 45 mmHg containing 52 volumes percent $\mathrm{CO}_{2}$ and 4 volumes percent of $\mathrm{CO}_{2}$ is actually exchanged in the process of transporting $\mathrm{CO}_{2}$ from tissues to lungs. Valedictory



## CO2 equilibration curve



- The effect of $\mathrm{O}_{2}$ on $\mathrm{H}^{+}$ion and $\mathrm{CO}_{2}$ loading and unloading from haemoglobin is known as Haldane effect or C-D-H effect. i.e. oxygenation of hemoglobin reduces it's ability to bind with $\mathrm{CO}_{2}$ Deoxygenation of the hemoglobin increases its ability to carry $\mathrm{CO}_{2}$.
- This is a consequence of the fact that reduced (deoxygenated) hemoglobin is a better proton $\left(\mathrm{H}^{+}\right)$acceptor than the oxygenated form.
- The upper part of the curve indicates that for every mm Hg increase in $\mathrm{PCO}_{2}$, a greater volume of $\mathrm{CO}_{2}$ is transported in venous blood than arterial blood which is due to the Haldane effect.
- If Haldane effect were not there, to transport the 4 volumes percent, $\mathrm{PCO}_{2}$ of venous blood would have to be raised to 52 mm Hg .


