#### TASK:

Each student (or group) receives a card. The smartboard screen will be divided into sections. In your block, draw all the pictures you would need to explain your story. When the class is ready, each student will explain their story with the help of their drawings on the board.

# **Story Event 1: The Journey Begins – Oxygen Loading in the Lungs**

#### Narrative

Meet Alice, a healthy young individual who just took a deep breath. As Alice inhales, air floods her **lungs**, filling tiny, balloon-like structures called **alveoli**. These alveoli are bathed in an environment rich in oxygen, with a **partial pressure of oxygen (pO<sub>2</sub>)** around **100 mmHg**—perfect for oxygen to bind efficiently.

Inside Alice's red blood cells (RBCs), the protein **hemoglobin** eagerly awaits this oxygen. Each hemoglobin molecule is like a sophisticated carrier, comprising **four polypeptide chains**, each embedded with a **heme group** capable of binding one oxygen molecule (O<sub>2</sub>). As oxygen diffuses across the alveolar walls into the blood, it swiftly binds to hemoglobin, forming **oxyhemoglobin** (HbO<sub>2</sub>).

This binding process is highly efficient in the lungs, where hemoglobin becomes almost fully saturated with oxygen—**around 97-100% saturation**—as depicted by the flat upper portion of the **hemoglobin dissociation curve**. This oxygen-rich blood is then pumped by Alice's heart through her **arteries**, embarking on its vital journey to nourish her body's tissues.

#### **Key Facts**

- **Hemoglobin Structure**: Four polypeptide chains, each with a heme group binding one O<sub>2</sub> molecule.
- **Oxygen Saturation in Lungs**: Hemoglobin saturation reaches approximately 97-100% in the high pO<sub>2</sub> environment of the alveoli.
- **Red Blood Cell Structure**: High surface area-to-volume ratio facilitates rapid gas diffusion.
- **Hill Equation Introduction**: Explains cooperative binding of oxygen to hemoglobin, enhancing loading efficiency.

- 1. Lungs and Alveoli Diagram: Highlighting alveoli where gas exchange occurs.
- 2. **Hemoglobin Molecule Illustration**: Showing the four subunits and heme groups binding oxygen.
- 3. **Oxygen-Hemoglobin Dissociation Curve**: Emphasizing the flat upper portion indicating high saturation.
- 4. Red Blood Cell Structure: Illustrating the biconcave shape and high surface area.

## **Story Event 2: The Voyage – Transporting Oxygen Through the Bloodstream**

#### Narrative

With oxygen now securely bound to hemoglobin, Alice's oxygen-rich blood travels through her **arteries**, propelled by the rhythmic beats of her heart. This oxygenated blood embarks on the **systemic circulation**, coursing through her body to reach every organ and tissue—from her bustling brain to her hardworking muscles.

As the blood moves away from the lungs, it encounters tissues with varying **partial pressures of oxygen (pO<sub>2</sub>)**. For instance, **resting tissues** like Alice's liver and kidneys maintain a  $pO_2$  of about 40 mmHg, while active tissues, such as her muscles during exercise, can experience a significantly lower  $pO_2$ .

The **oxygen-hemoglobin dissociation curve**, a beautifully **S-shaped graph**, illustrates how hemoglobin's affinity for oxygen changes with different  $pO_2$  levels. This curve is central to understanding how oxygen is efficiently loaded in the lungs and unloaded in the tissues, ensuring that each part of Alice's body receives the oxygen it needs precisely when it needs it.

#### **Key Facts**

- Systemic Circulation: Pathway of oxygen-rich blood from the lungs to body tissues.
- **Partial Pressure Gradient**: Higher pO<sub>2</sub> in the lungs compared to tissues facilitates oxygen release.
- **Cooperative Binding**: Binding of one O<sub>2</sub> molecule increases hemoglobin's affinity for subsequent O<sub>2</sub> molecules, enhancing loading efficiency.
- Hill Equation Application: Quantifies hemoglobin's cooperative binding, explaining the dissociation curve's shape.

- 1. **Circulatory System Diagram**: Tracing the path of blood from lungs to various body tissues.
- 2. **Hemoglobin Dissociation Curve**: Highlighting different points corresponding to varying pO<sub>2</sub> levels.
- 3. **Blood Flow Illustration**: Showing arteries transporting oxygenated blood through the body.
- 4. **Hill Equation Graphic**: Visual representation of the Hill Equation and its impact on the dissociation curve.

## **Story Event 3: Delivering Oxygen to Resting Tissues**

#### Narrative

As Alice's blood reaches her **resting tissues**, such as her liver and kidneys, it enters an environment with a **moderate pO<sub>2</sub>** of about **40 mmHg**. Here, hemoglobin begins to release some of its bound oxygen. Imagine hemoglobin as a dedicated delivery team—when they arrive in a quiet town (resting tissues), a few team members (oxygen molecules) stay behind to support the community.

At this moderate  $pO_2$ , hemoglobin's saturation drops to approximately 75%—the midportion of the **hemoglobin dissociation curve**. This release is perfectly calibrated to meet the metabolic needs of tissues at rest, ensuring vital organs function smoothly without depleting oxygen reserves.

As hemoglobin releases oxygen, it also binds free hydrogen ions ( $\mathbf{H}^+$ ), helping to buffer the blood's pH and maintain acid-base balance—a crucial role in overall homeostasis.

After delivering oxygen, the blood, now slightly deoxygenated, begins its return journey to the lungs. Since not all oxygen was released in the resting tissues, the returning blood still retains a **relatively high oxygen content**, maintaining efficiency for the next cycle.

#### **Key Facts**

- Moderate pO<sub>2</sub> Environment: Resting tissues operate around 40 mmHg pO<sub>2</sub>.
- Hemoglobin Saturation at Rest: Drops to about 75% as oxygen is released.
- Bohr Effect (Minimal at Rest): Slight influence due to lower CO<sub>2</sub> and stable pH.
- **Buffering Capacity**: Hemoglobin binds free H<sup>+</sup> ions, aiding in pH regulation.

- 1. Muscle Tissue at Rest: Depicting oxygen being delivered to cells.
- 2. Partial Pressure Comparison: Visual comparison of lung pO<sub>2</sub> vs. resting tissue pO<sub>2</sub>.
- 3. Mid-Range Dissociation Curve: Highlighting the 75% saturation point.
- 4. Hemoglobin Buffering H<sup>+</sup> Ions: Illustration of hemoglobin binding hydrogen ions.

## **Story Event 4: Meeting the Challenge – Oxygen Release in Exercising Tissues**

#### Narrative

Now, envision Alice deciding to go for a sprint. Her muscles surge into action, generating more **carbon dioxide (CO<sub>2</sub>)** and producing metabolic byproducts like **lactic acid**, which lowers the **pH** in the muscle tissues. Consequently, the **pO<sub>2</sub>** in these exercising muscles drops dramatically to about **20 mmHg**.

Under these demanding conditions, hemoglobin undergoes a significant transformation known as the **Bohr Effect**. Increased **CO**<sub>2</sub> and lowered **pH** cause the hemoglobin dissociation curve to **shift to the right**, reducing hemoglobin's affinity for oxygen. Additionally, within the red blood cells, molecules like **2,3-Bisphosphoglycerate** (**2,3-BPG**) play a pivotal role by further decreasing hemoglobin's oxygen affinity.

As a result, hemoglobin releases more oxygen, dropping its saturation to about **50-60%**—a crucial adjustment ensuring that Alice's active muscles receive the additional oxygen necessary to sustain intense activity. The presence of **2,3-BPG**, synthesized via the **Rapoport-Luebering shunt**, enhances this oxygen release, especially under conditions like high altitude or anemia where oxygen delivery needs are heightened.

The blood returning from these exercising tissues is now **significantly deoxygenated**, carrying more  $CO_2$  back to the lungs for exhalation. This dynamic interplay ensures that oxygen delivery is finely tuned to the body's varying demands.

#### **Key Facts**

- High Metabolic Activity: Exercise increases CO<sub>2</sub> production and lowers pO<sub>2</sub> to ~20 mmHg.
- **Bohr Effect**: Increased CO<sub>2</sub> and lowered pH shift the dissociation curve right, decreasing oxygen affinity.
- Hemoglobin Saturation During Exercise: Drops to 50-60%, maximizing oxygen delivery.
- **Role of 2,3-BPG**: Enhances oxygen release by reducing hemoglobin's affinity for oxygen.
- **Rapoport-Luebering Shunt**: Pathway in RBCs responsible for synthesizing 2,3-BPG.
- Adaptations: Increased 2,3-BPG levels occur in chronic hypoxia, anemia, and intense exercise.

- 1. Active Muscle Tissue: Illustrating increased oxygen consumption and CO<sub>2</sub> production.
- 2. Rightward Shift of Dissociation Curve: Depicting the Bohr Effect.
- 3. 2,3-BPG Molecules in RBCs: Showing their role in decreasing oxygen affinity.
- 4. Exercising Individual: Demonstrating enhanced blood flow and metabolic activity.

### **Story Event 5: Completing the Cycle – Carbon Dioxide Transport and Release in the Lungs**

#### Narrative

After delivering oxygen to the tissues, Alice's blood, now rich in CO<sub>2</sub>, begins its return journey to the **lungs**. Here, another vital process—the **Haldane Effect**—comes into play. As the blood reaches the lungs, **oxygen binds to hemoglobin**, converting it back to **oxyhemoglobin** (**HbO**<sub>2</sub>). This oxygenation reduces hemoglobin's affinity for CO<sub>2</sub>, facilitating its release.

Inside the red blood cells, the enzyme **carbonic anhydrase** catalyzes the conversion of  $CO_2$  and water into **carbonic acid** (H<sub>2</sub>CO<sub>3</sub>), which then dissociates into **bicarbonate ions** (HCO<sub>3</sub><sup>-</sup>) and hydrogen ions (H<sup>+</sup>). To maintain ionic balance, bicarbonate ions leave the red blood cells in exchange for chloride ions (chloride shift), ensuring the blood remains electrically neutral.

The **Haldane Effect** enhances  $CO_2$  release by decreasing hemoglobin's affinity for  $CO_2$  in the presence of high oxygen levels. As oxygen binds to hemoglobin, it promotes the release of  $CO_2$  into the alveoli. Finally, Alice exhales, expelling the accumulated  $CO_2$  from her body.

#### **Key Facts**

- CO<sub>2</sub> Transport Mechanisms:
  - **Dissolved CO2**: ~5-10% in plasma.
  - **Carbaminohemoglobin (HbCO2)**: ~10-30% bound to hemoglobin.
  - **Bicarbonate Ions (HCO**<sub>3</sub>-): ~60-70% converted in plasma via carbonic anhydrase.
- Haldane Effect: Oxygenation reduces hemoglobin's affinity for CO<sub>2</sub>, facilitating its release.
- **Chloride Shift**: Exchange of bicarbonate ions for chloride ions maintains ionic balance.
- Carbonic Anhydrase: Enzyme that catalyzes the conversion of CO<sub>2</sub> to bicarbonate.
- **Buffering Capacity**: Hemoglobin's ability to bind H<sup>+</sup> ions helps maintain blood pH balance.
- Haldane Effect Interaction with Bohr Effect: Together, they ensure efficient gas exchange tailored to metabolic needs.

- 1. **CO<sub>2</sub> Transport Pathways**: Diagram showing dissolved CO<sub>2</sub>, carbaminohemoglobin, and bicarbonate ions.
- 2. Haldane Effect Illustration: Depicting how oxygen binding reduces CO<sub>2</sub> affinity.
- 3. Chloride Shift Process: Visualizing the exchange of bicarbonate and chloride ions.
- 4. Lungs ExhalationProcess: Showing CO<sub>2</sub> being expelled from the body.

## **Supplementary Story Events (Optional)**

#### Supplementary Event A: Fetal Hemoglobin and Placental Gas Exchange

#### Narrative

During pregnancy, a special form of hemoglobin known as **fetal hemoglobin (HbF)** takes center stage. Unlike adult hemoglobin (**HbA**), HbF consists of two alpha ( $\alpha$ ) and two gamma ( $\gamma$ ) globin chains ( $\alpha_2\gamma_2$ ). HbF has a **higher affinity for oxygen**, ensuring efficient oxygen uptake from the mother's blood across the placenta. This high affinity allows the developing fetus to receive adequate oxygen even when maternal oxygen levels are relatively low.

#### **Key Facts**

- **Fetal Hemoglobin Structure**:  $\alpha_2\gamma_2$  composition.
- **Higher Oxygen Affinity**: Facilitates efficient oxygen transfer from maternal to fetal blood.
- **Physiological Significance**: Ensures the developing fetus receives sufficient oxygen for growth and development.

#### **Suggested Images**

- 1. Maternal-Fetal Placenta Gas Exchange Diagram: Illustrating oxygen transfer.
- 2. HbA vs. HbF Structure Comparison: Highlighting differences in globin chains.
- 3. Fetus Receiving Oxygen: Visual representation of oxygen delivery to fetal tissues.

#### **Supplementary Event B: Clinical Implications of Hemoglobin Function**

#### Narrative

Understanding hemoglobin's role is crucial in medical contexts. For instance, in **Chronic Obstructive Pulmonary Disease (COPD)**, impaired gas exchange leads to increased CO<sub>2</sub> levels, enhancing the **Bohr Shift** and promoting oxygen release despite reduced oxygen uptake. Similarly, **sickle cell anemia** alters hemoglobin's structure, impairing its ability to release oxygen efficiently, leading to symptoms like pain and organ damage.

**Carbon monoxide poisoning** is another critical condition where **CO** binds to hemoglobin with a higher affinity than oxygen, reducing oxygen delivery and exacerbating the Bohr Shift effects by further decreasing hemoglobin's oxygen affinity.

#### **Key Facts**

- **COPD**: Impaired gas exchange increases CO<sub>2</sub> levels, affecting the Bohr Shift.
- Sickle Cell Anemia: Abnormal hemoglobin structure impairs oxygen release.
- **Carbon Monoxide Poisoning**: CO binding reduces oxygen delivery and exacerbates hemoglobin affinity shifts.

#### **Suggested Images**

- 1. **Hemoglobin Mutation in Sickle Cell Anemia**: Showing altered hemoglobin structure.
- 2. COPD Gas Exchange Impairment Diagram: Visualizing reduced oxygen uptake.
- 3. Carbon Monoxide Binding to Hemoglobin: Illustrating competitive binding.

### **Full Story Overview**

#### 1. The Journey Begins – Oxygen Loading in the Lungs

- **Setting**: Alveoli in the lungs.
- Main Characters: Oxygen molecules, hemoglobin.
- **Plot**: Oxygen binds to hemoglobin in high pO<sub>2</sub> environments, saturating hemoglobin for transport.

#### 2. The Voyage – Transporting Oxygen Through the Bloodstream

- **Setting**: Systemic circulation via arteries.
- Main Characters: Oxygen-rich blood, hemoglobin.
- **Plot**: Oxygenated blood travels through the body, maintaining a pO<sub>2</sub> gradient for efficient oxygen delivery.

#### **3. Delivering Oxygen to Resting Tissues**

- **Setting**: Resting tissues (e.g., liver, kidneys).
- Main Characters: Hemoglobin, oxygen molecules.
- **Plot**: Hemoglobin releases oxygen at moderate pO<sub>2</sub>, meeting the needs of resting tissues while also buffering blood pH.

#### 4. Meeting the Challenge – Oxygen Release in Exercising Tissues

- Setting: Active muscles during exercise.
- Main Characters: Hemoglobin, oxygen molecules, CO<sub>2</sub>, 2,3-BPG.
- **Plot**: Under high metabolic demand, the Bohr Effect and 2,3-BPG facilitate greater oxygen release to meet increased needs.

## **5.** Completing the Cycle – Carbon Dioxide Transport and Release in the Lungs

- **Setting**: Returning blood to the lungs.
- Main Characters: CO<sub>2</sub>, hemoglobin, bicarbonate ions.
- **Plot**: The Haldane Effect ensures efficient CO<sub>2</sub> transport and release, completing the gas exchange cycle as blood is oxygenated once again.