



Control of ventilation

Sorawit Ongsupankul, Christian John Capirig

Objectives

Introduction

Relevant Anatomy

Phases of Respiration

Inputs to Respiratory Centers

Conclusion

References

Introduction

The regulation of breathing is a complex physiological process that integrates multiple anatomical structures and neural pathways. These components work in harmony to maintain homeostasis by ensuring optimal oxygen delivery and carbon dioxide removal. Understanding this intricate regulation is essential in the context of different respiratory diseases and during mechanical ventilation to provide support that aligns with the patient's natural respiratory drive.

Relevant Anatomy

Medulla Oblongata

The medulla oblongata is the primary control center for breathing and houses critical respiratory groups:

- **Dorsal Respiratory Group (DRG):**
 - **Afferent Inputs:** Receives signals from chemo sensitive areas, peripheral chemoreceptors, baroreceptors, and receptors located in the liver, pancreas, gastrointestinal organs, and lungs.
 - **Function:** Generates the inspiratory ramp signal, which sends action potentials through the corticospinal tract to the phrenic and intercostal nerves, activating the diaphragm and external intercostal muscles to initiate inspiration.
 - **Regulation:** Inhibited by the pneumotaxic center in the pons, signaling the cessation of inspiration.
- **Ventral Respiratory Group (VRG):**
 - **Function:** Fine-tunes both inspiration and expiration, especially during increased respiratory drive, such as during exercise or stress.

Pons

- **Pneumotaxic Center:**
 - **Function:** Inhibits the inspiratory ramp signals from the DRG, regulating the transition between inspiration and expiration and preventing over-inflation of the lungs.

Respiratory Muscles

- **Diaphragm and Intercostal Muscles:** Primary muscles responsible for thoracic expansion and contraction, facilitating the mechanical aspect of breathing.

Peripheral Nervous System

- **Phrenic and Intercostal Nerves:** Transmit neural signals from the central nervous system to the respiratory muscles, initiating contraction and relaxation.

Lung and Airway Receptors

- **Stretch Receptors:** Located in the bronchi and bronchioles, these receptors detect changes in lung volume. When tidal volume exceeds approximately three times the normal value, these receptors send inhibitory signals to the DRG to terminate inspiration, preventing overdistension.

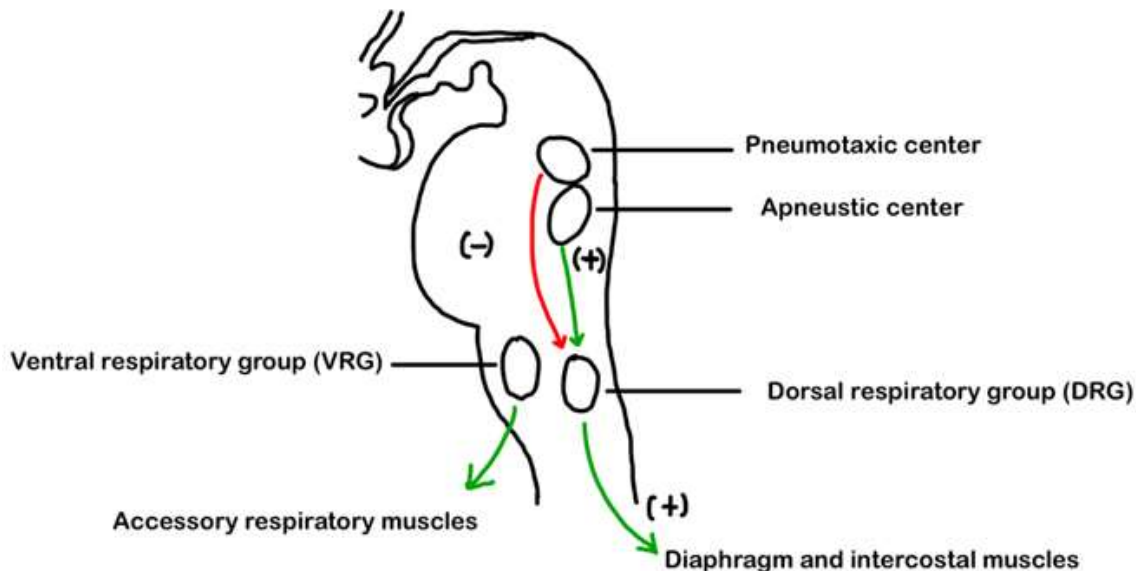


Figure 1 represents the neural control of breathing, highlighting the interaction between key respiratory centers. The pneumotaxic center in the pons inhibits the inspiratory signals from the apneustic center and dorsal respiratory group (DRG), which control the diaphragm and intercostal muscles for inspiration. The ventral respiratory group (VRG) coordinates the accessory respiratory muscles for active breathing, particularly during increased respiratory demand. Positive (+) and inhibitory (-) influences are shown, indicating the regulation of respiratory rhythm and depth.

Phases of Respiration

Inspiration

1. **Initiation:** Neural signals originating from the DRG travel via the phrenic and intercostal nerves to activate the diaphragm and intercostal muscles. This contraction creates negative pleural pressure relative to atmospheric pressure, causing air to rush into the lungs.
2. **Maintenance:** Sustained neural activity keeps the respiratory muscles contracted, ensuring continued air inflow.
3. **Cessation:** The pneumotaxic center sends inhibitory signals to the DRG, halting the inspiratory ramp signals and transitioning to exhalation.

Exhalation

1. **Initiation:** In passive exhalation, the relaxation of inspiratory muscles allows the elastic recoil of the lungs and thoracic cavity to expel air.
2. **Maintenance:** Elastic properties of the lungs facilitate continued air outflow. During active breathing, expiratory muscles (e.g., abdominal muscles) contract to enhance air expulsion.
3. **Cessation:** The chest wall returns to its neutral shape. Airflow ceases when alveolar and airway pressure equalize with atmospheric pressure, resting at the functional residual capacity (FRC).

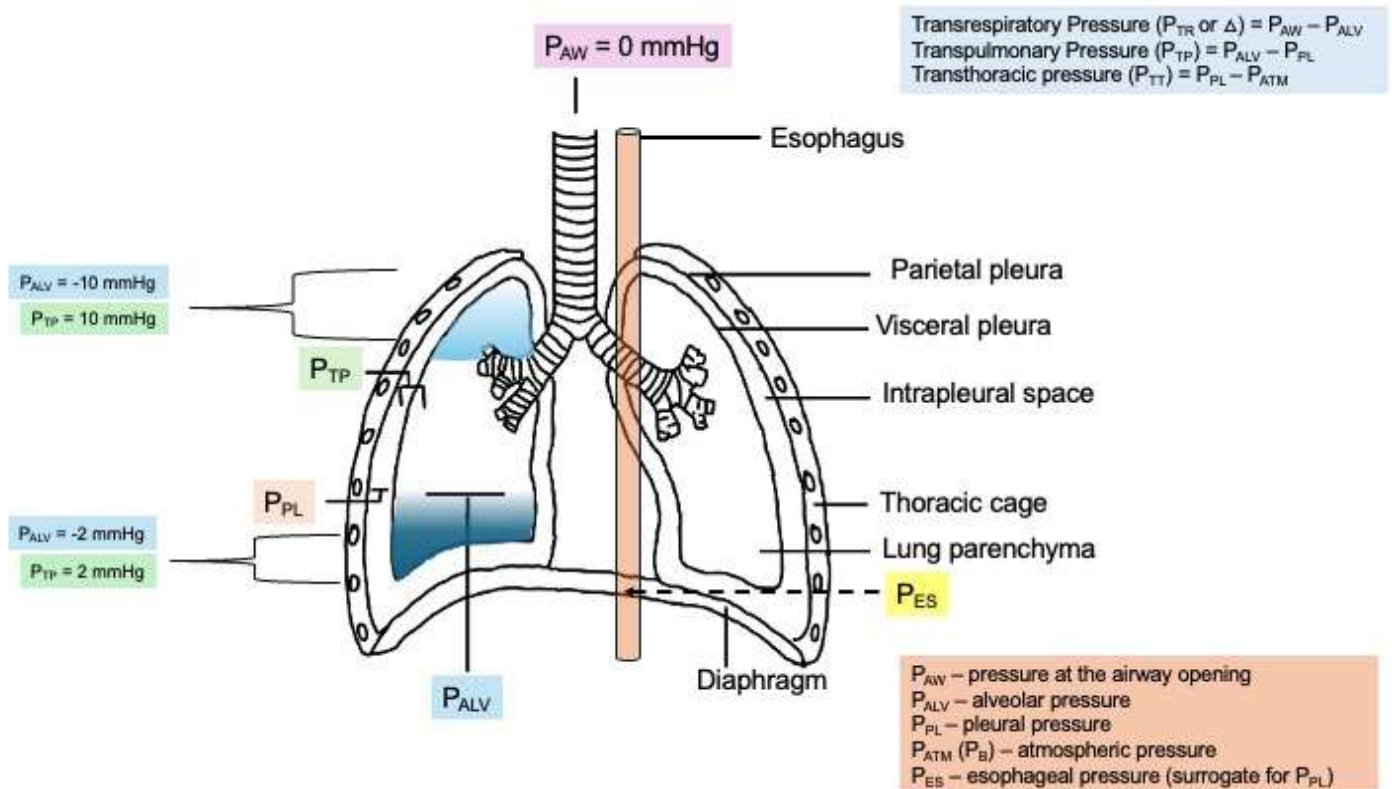


Figure 2 illustrates the pressures involved in normal respiration: P_{ATM} : atmospheric pressure, P_{ALV} : intra-alveolar pressure, P_{PL} : pleural pressure, P_{ES} : esophageal pressure, P_{TP} : trans-pulmonary pressure.

Inputs to Respiratory Centers

Cerebral Cortex

- **Voluntary Control:** Enables conscious modification of breathing patterns, such as during speaking, singing, or holding one's breath.
- **Emotional and Pain Responses:** Emotional states and pain influence respiratory rate and depth through connections in the limbic system.

Chemical Stimuli

- **Partial Pressure of Arterial Carbon Dioxide (P_{aCO_2}):** CO_2 diffuses across the blood-brain barrier and reacts with water to form carbonic acid (H_2CO_3). This dissociates into hydrogen ions (H^+) and bicarbonate (HCO_3^-), increasing H^+ concentration. Central chemoreceptors detect this change, activating the DRG to increase ventilation.

- **pH Levels:** Peripheral chemoreceptors in the carotid and aortic bodies sense changes in blood pH. Acidemia (low pH) increases ventilation to expel CO_2 , while alkalemia (high pH) reduces ventilatory drive.
- **Partial Pressure of Arterial Oxygen (PaO_2):** Peripheral chemoreceptors become significantly activated when PaO_2 falls below 60 mmHg, stimulating increased ventilation to enhance oxygen uptake.

Juxtapulmonary Capillary (J) Receptors

- **Location and Function:** Situated in alveolar walls, these receptors respond to pulmonary capillary engorgement and increased interstitial fluid, as seen in pulmonary edema. Activation results in rapid, shallow breathing and sensations of dyspnea.

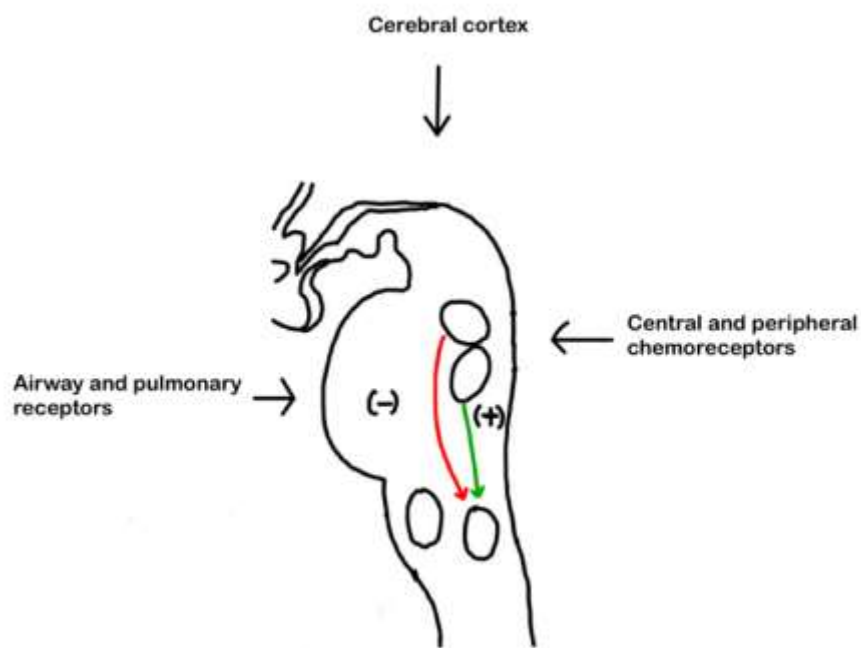


Figure 3 illustrates the integration of inputs in the neural control of breathing. The cerebral cortex provides voluntary control and emotional influence over respiration, while central and peripheral chemoreceptors monitor PaCO_2 , PaO_2 , and pH levels to adjust respiratory drive. Airway and pulmonary receptors detect mechanical changes in the lungs, such as stretch or irritation, influencing respiratory patterns.

Conclusion

The control of ventilation involves intricate neural and mechanical processes evolved to maintain homeostasis. By continuously integrating signals from peripheral and central sensors to adapt ventilation to the body's changing demands. This system not only ensures adequate oxygen delivery and carbon dioxide removal but also maintains acid-base balance and supports metabolic needs. Clinically, it allows us to identify and manage conditions where respiratory control is impaired, and in the context of mechanical ventilation, understanding these physiological mechanisms is essential for optimizing patient care. By aligning ventilator settings with the body's natural respiratory control systems, clinicians can enhance patient outcomes and reduce complications such as ventilator dyssynchrony.

Reference

- Guyton AC, Hall JE. *Textbook of Medical Physiology*. 14th ed. Philadelphia, PA: Elsevier; 2020.
- Boron WF, Boulpaep EL. *Medical Physiology*. 2nd ed. Philadelphia, PA: Elsevier; 2012.
- The process of breathing. In: *Anatomy and Physiology*, Chapter 22.3. Oregon State University. Accessed January 24, 2025. <https://open.oregonstate.education/aandp/chapter/22-3-the-process-of-breathing/>
- Brinkman JE, Toro F, Sharma S. Physiology, Respiratory Drive. In: *StatPearls*. Treasure Island, FL: StatPearls Publishing; 2025. Updated June 5, 2023. Accessed January 24, 2025. <https://www.ncbi.nlm.nih.gov/books/NBK482414/>
- Chourpiliadis C, Bhardwaj A. Physiology, Respiratory Rate. In: *StatPearls*. Treasure Island, FL: StatPearls Publishing; 2025. Updated September 12, 2022. Accessed January 24, 2025. <https://www.ncbi.nlm.nih.gov/books/NBK537306/>
- Ikeda K, Kawakami K, Onimaru H, et al. The respiratory control mechanisms in the brainstem and spinal cord: integrative views of the neuroanatomy and neurophysiology. *J Physiol Sci* 2017; 67(1):45-62. doi:10.1007/s12576-016-0475-y
- Guyenet PG, Stornetta RL, Bayliss DA. Central respiratory chemoreception. *J Comp Neurol* 2010; 518(19):3883-3906. doi:10.1002/cne.22435