

COURSE GUIDE

NSC 202 HUMAN ANATOMY III

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Printed 2017

ISBN: 978-058-887-6

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INTRODUCTION

Welcome to the second year course in Human Anatomy, NSC 202 – Human Anatomy III. This is a second year course and a continuation of Human Anatomy I (NSC 102) and Human Anatomy II (NSC 104) where you have increased/improved your knowledge about some body structures and their organisations. You also covered the protective covering of all the body organs as well as the supporting systems. This second part will cover other internal organs that are important to maintenance of life. As indicated in NSC 102, caring always require sound understanding of the normal structure of the body organs as to know what could be wrong and how such manifest. Basic assessments done before planning general and nursing care usually consider the various organs that function within systems and as interrelated systems. You will be required to be able to describe the various organs and discuss the clinical correlates of the body parts. You will enjoy drawing and labelling, as well as seeing some of these organs in real life. You will also see the variations in normal and diseased organs as you are encouraged to participate in all laboratory assignments.

COURSE OVERVIEW

Human Anatomy (III)

Human Anatomy (III) is the second of three courses that covers the major organs that are responsible for life. In this course, we are going to study the special senses, the endocrine system and the nervous system. These are the regulatory systems. The structures and locations of the various organs that make each of the systems will be studied.

COURSE OBJECTIVE

At the completion of this course, you should be able to:

- i. Discuss the anatomy and functions of the special senses
- ii. Describe the various disorders of the special senses
- iii. Discuss the general functions of the endocrine system
- iv. Describe the locations and functions of the glands.
- v. Discuss the management of various endocrine disorders.
- vi. Discuss the formation of the brain and the spinal cord from the most primitive neural tube.
- vii. Discuss the anatomy of the central nervous system.
- viii. Differentiate between structural and functional divisions of the nervous system.
- ix. Discuss the peripheral and autonomic nervous systems

COURSE IMPLEMENTATION DOING THE COURSE

The course will be delivered adopting the blended learning mode, 70% of online but interactive sessions and 30% of face-to-face during laboratory sessions. You are expected to register for this course online before you can have access to all the materials and have access to the class sessions online. You will have hard and soft copies of course materials, you will also have online interactive sessions, face-to-face sessions with instructors during practical sessions in the laboratory. The interactive online activities will be available to you on the course link on the Website of NOUN. There are activities and assignments online for every unit every week. It is important that you visit the course sites weekly and do all assignments to meet deadlines and to contribute to the topical issues that would be raised for everyone's contribution.

You will be expected to read every module along with all assigned readings to prepare you to have meaningful contributions to all sessions and to complete all activities. It is important that you attempt all the Self Assessment Questions (SAQ) at the end of every unit to help your understanding of the contents and to help you prepare for the in-course tests and the final examination

You will also be expected to keep a portfolio where you keep all your completed assignments.

COURSE REQUIREMENTS AND EXPECTATIONS OF YOU

Attendance of 95% of all interactive sessions, submission of all assignments to meet deadlines; participation in all CMA, attendance of all laboratory sessions with evidence as provided in the log book, submission of reports from all laboratory practical sessions and attendance of the final course examination. You are also expected to:

1. Be versatile in basic computer skills
2. Participate in all laboratory practical up to 90% of the time
3. Submit personal reports from laboratory practical sessions on schedule
4. Log in to the class online discussion board at least once a week and contribute to ongoing discussions.
5. Contribute actively to group seminar presentations.

EQUIPMENT AND SOFTWARE NEEDED TO ACCESS COURSE

You will be expected to have the following tools:

1. A computer (laptop or desktop or a tablet)
2. Internet access, preferably broadband rather than dial-up access

3. MS Office software – Word PROCESSOR, Powerpoint, Spreadsheet
4. Browser – Preferably Internet Explorer, Moxilla Firefox
5. Adobe Acrobat Reader

NUMBER AND PLACES OF MEETING (ONLINE, FACE-TO-FACE, LABORATORY PRACTICALS)

The details of these will be provided to you at the time of commencement of this course

DISCUSSION FORUM

There will be an online discussion forum and topics for discussion will be available for your contributions. It is mandatory that you participate in every discussion every week. Your participation link you, your face, your ideas and views to that of every member of the class and earns you some mark.

COURSE EVALUATION

There are two forms of evaluation of the progress you are making in this course. The first are the series of activities, assignments and end of unit, computer or tutor marked assignments, and laboratory practical sessions and report that constitute the continuous assessment that all carry 30% of the total mark. The second is a written examination with multiple choice, short answers and essay questions that take 70% of the total mark that you will do on completion of the course.

Students evaluation: The students will be assessed and evaluated based on the following criteria

○ In-Course Examination:

In line with the university's regulation, in-course examination will come up in the middle of the semester These would come in form of Computer Marked Assignment. This will be in addition to 1 compulsory Tutor Marked Assignment (TMA's) and three Computer marked Assignment that comes after every module.....

○ **Laboratory practical:** Attendance, record of participation and other assignments will be graded and added to the other scores form other forms of examinations.

○ **Final Examination:** The final written examination will come up at the end of the semester comprising essay and objective questions covering all the contents covered in the course. The final examination will amount to 60% of the total grade for the course.

Learner-Facilitator evaluation of the course

This will be done through group review, written assessment of learning (theory and laboratory practical) by you and the facilitators.

GRADING CRITERIA

Grades will be based on the following Percentages

Tutor Marked Individual Assignments		} 10%
Computer marked Assignment	10%	
Group assignment	5%	} 40%
Discussion Topic participation	5%	
Laboratory practical	10%	
End of Course examination	60%	

GRADING SCALE

A = 70-100

B = 60 - 69

C = 50 - 59

F = ≤ 49

SCHEDULE OF ASSIGNMENTS WITH DATES

To be provided for each module by the facilitator in addition to the ones already spelt out in the course materials.

SPECIFIC READING ASSIGNMENTS

To be provided by each module

REFERENCE/FURTHER READING

Hutchinson, M., Mallat, J., Marieb, E.N., Wilhelm P.B. (2007). *A Brief Atlas of the Human Body*. San Francisco: Pearson Education Inc.

Katherine, M. A., Rogers & William, N. S. (2011). *Nurses! Test Yourself in Anatomy and Physiology*

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**MAIN
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MODULE 1 THE SPECIAL SENSES

Unit 1	The Tongue and the Sense of Taste
Unit 2	The Nose and the Sense of Smell
Unit 3	The Ear and the Sense of Hearing
Unit 4	The Eyes and the Sense of Vision

UNIT 1 THE TONGUE AND THE SENSE OF TASTE**CONTENTS**

1.0	Introduction
2.0	Objectives
3.0	Main Content
3.1	Functions of Taste
3.2	Location of Taste Buds
3.3	Histology of Taste Buds
3.4	Taste Preference and Control of Diet
3.5	Clinical Correlates
4.0	Conclusion
5.0	Summary
6.0	Tutor-Marked Assignment
7.0	References/Further Reading

1.0 INTRODUCTION

Taste is mainly a function of the *taste buds* in the mouth, but it is common experience that one's sense of smell also contributes strongly to taste perception. In addition, the texture of food, as detected by tactual senses of the mouth, and the presence of substances in the food that stimulate pain endings, such as pepper, greatly alter the taste experience. The importance of taste lies in the fact that it allows a person to select food in accord with desires and often in accord with the body tissues' metabolic need for specific substances.

2.0 OBJECTIVES

At the end of this unit, you should be able to:

- discuss the anatomy of the tongue and the function of each part.
- describe how taste sensations are created and interpreted
- discuss the five primary taste sensations
- describe the histology and function of a typical taste bud
- explain the neuronal pathways for the sense of taste

3.0 MAIN CONTENT

3.1 Functions of Taste

Substances called **tastants** dissolve in saliva, enter the taste pores, and by various mechanisms cause the taste cells to depolarize. For example, the diffusion of the positively charged ions Na^+ and H^+ into the taste cells causes depolarization. The binding of tastants to receptors on the taste hairs activates G proteins, which results in depolarization by causing Ca^{2+} channels to open or K^+ channels to close. When the taste cells depolarize, they release neurotransmitters that diffuse to sensory neurons associated with the taste cells. The neurotransmitters stimulate action potentials in the sensory neurons, which are conducted to the brain, where the sense of taste is perceived.

Five **primary tastes** have been identified in humans: salty, sour, sweet, bitter, and umami (a Japanese term loosely translated as savoury). The salty taste is stimulated by Na^+ ; sour by acids; sweet by sugars, some other carbohydrates, and some proteins; bitter by alkaloids (bases); and umami by the amino acid glutamate and related compounds. The umami taste sensation is most intense when coupled with the salty taste, hence the popularity of adding salt to tomatoes, ketchup, soy sauce, and the food additive monosodium glutamate (MSG). Even though only five primary tastes have been identified, humans can perceive a fairly large number of different tastes, presumably by combining the five basic taste sensations.

Thresholds for the five primary taste stimuli vary. Sensitivity for bitter substances is the highest. Many alkaloids are poisonous, and the high sensitivity for bitter tastes may be protective because of the avoidance of bitter foods. On the other hand, humans tend to crave sweet, salty, and umami tastes, perhaps in response to the body's need for sugars, carbohydrates, proteins, and minerals.

Many of the sensations thought of as being tastes are strongly influenced by olfactory sensations.

This phenomenon can be demonstrated by pinching one's nose to close the nasal passages while trying to taste something. With olfaction blocked, it is difficult to distinguish between the taste of a piece of apple and the taste of potato. Much of the "taste" is lost by this action. This is one reason that a person with a cold has a reduced sensation of taste. The tongue can detect other stimuli besides taste, such as temperature and texture, and these stimuli can influence the sensation of taste. Food served at the wrong temperature is perceived as distasteful. Stimulation

of hot receptors by capsaicin or black pepper is interpreted as hot or spicy, and stimulation of cold receptors is perceived as fresh or minty.

3.2 Location of Taste Buds

The taste buds are found on three types of papillae of the tongue, as follows:

- (1) A large number of taste buds are on the walls of the troughs that surround the circumvallate papillae, which form a V line on the surface of the posterior tongue.
- (2) Moderate numbers of taste buds are on the fungiform papillae over the flat anterior surface of the tongue.
- (3) Moderate numbers are on the foliate papillae located in the folds along the lateral surfaces of the tongue. Additional taste buds are located on the palate, and a few are found on the tonsillar pillars, on the epiglottis, and even in the proximal oesophagus. Adults have 3000 to 10,000 taste buds, and children have a few more. Beyond the age of 45 years, many taste buds degenerate, causing taste sensitivity to decrease in old age.

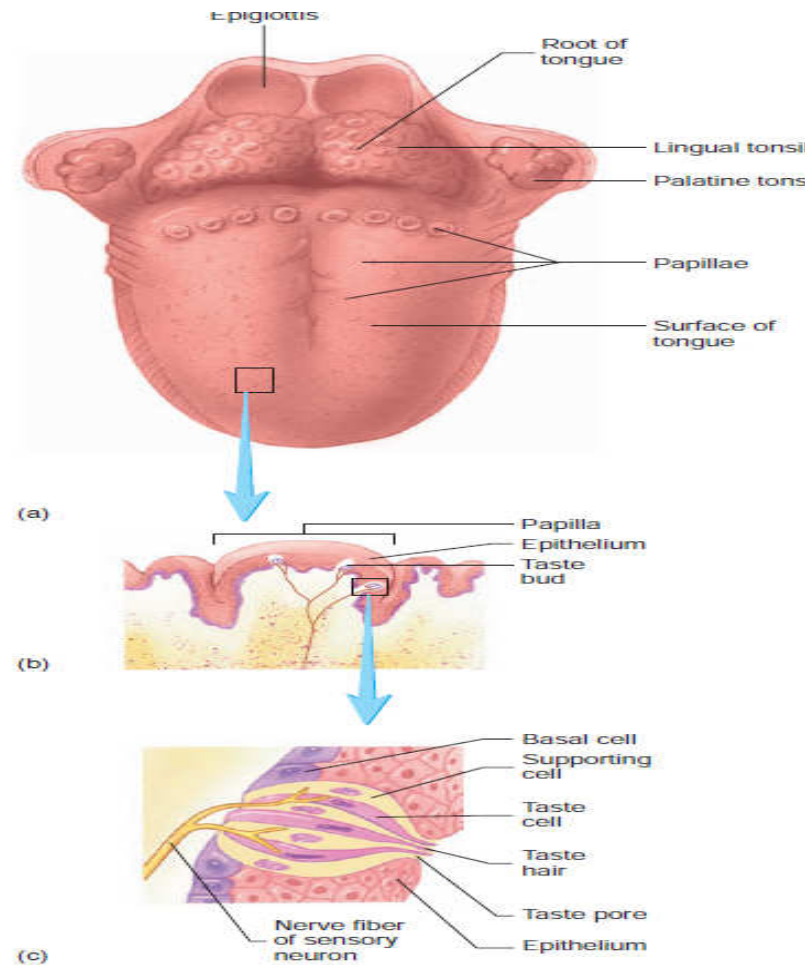


Figure 13.2 Tongue, Papillae, and Taste Buds
 (a) Surface of the tongue. (b) A papilla. (c) A taste bud.

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3.3 Histology of Taste Buds

Each of the 10,000 taste buds on a person's tongue consists of three major types of specialised epithelial cells. The sensory cells of each taste bud consist of about 50 **taste cells**. Each taste cell has several microvilli, called **taste hairs**, extending from its apex into a tiny opening in the epithelium called the **taste pore**. The remaining two nonsensory cell types are **basal cells** and **supporting cells**. Like olfactory cells, the cells of the taste buds are replaced continuously, each having a normal life span of about 10 days.

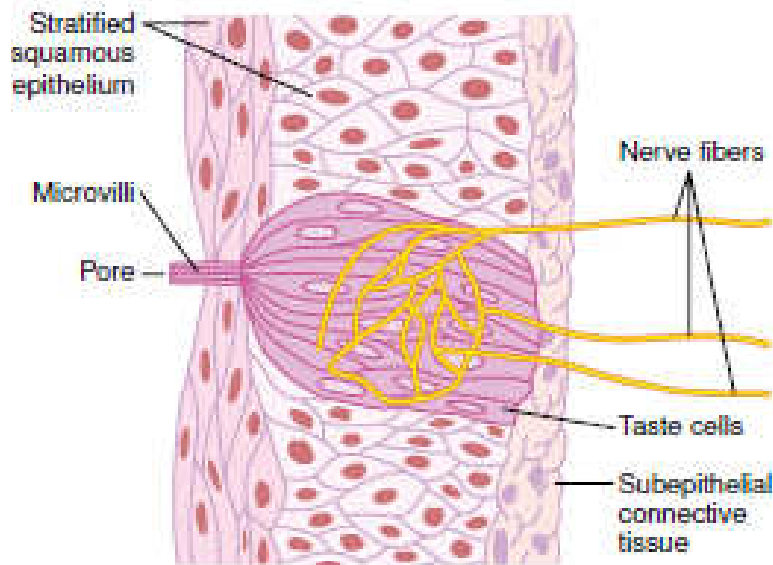


Fig.1.2: Taste Bud

3.4 Taste Preference and Control of Diet

Taste preference simply means that an animal will choose certain types of food in preference to others, and the animal automatically uses this to help control the diet it eats. Furthermore, its taste preferences often change in accord with the body's need for certain specific substances. The following experiments demonstrate this ability of animals to choose food in accord with the needs of their bodies. First, adrenalectomies, salt-depleted animals automatically select drinking water with a high concentration of sodium chloride in preference to pure water, and this is often sufficient to supply the needs of the body and prevent salt-depletion death. Second, an animal given injections of excessive amounts of insulin develops a depleted blood sugar, and the animal automatically chooses the sweetest food from among many samples. Third, calcium-depleted parathyroidectomies animals automatically choose drinking water with a high concentration of calcium chloride.

Also, human beings reject any food that has an unpleasant affective sensation, which in many instances protects our bodies from undesirable substances. The phenomenon of taste preference almost certainly results from some mechanism located in the central nervous system and not from a mechanism in the taste receptors themselves, although the receptors often become sensitised in favour of a needed nutrient. An important reason for believing that taste preference is mainly a central nervous system phenomenon is that previous experience with unpleasant or pleasant tastes plays a major role in determining one's taste preferences. For instance, if a person becomes sick soon after eating a particular type of food, the person generally develops a negative taste preference, or taste aversion, for that particular food thereafter; the same effect can be demonstrated in lower animals.

3.5 Clinical Correlates

Much of what is perceived as a taste defect is truly a primary defect in olfaction resulting in an alteration in taste.

4.0 CONCLUSION

In this unit you learnt main function of the *taste buds* in the mouth, and the common experience that one's sense of smell also contributes strongly to taste perception.

5.0 SUMMARY

- Taste is mainly a function of the *taste buds* in the mouth, but it is common experience that one's sense of smell also contributes strongly to taste perception.
- Receptors on the hairs detect tastants (dissolved substances). Five basic types of taste exist: sour, salty, bitter, sweet, and umami.
- Taste buds usually are associated with papillae – circumvallate, fungiform and foliate papillae.
- Taste buds consist of taste cells, basilar cells, and supporting cells. The taste cells have taste hairs that extend into taste pores.
- Taste preference simply means that an animal will choose certain types of food in preference to others, and the animal automatically uses this to help control the diet it eats. Furthermore, its taste preferences often change in accord with the body's need for certain specific substances.

SELF-ASSESSMENT EXERCISE

At the histology laboratory, examine the histological structure of the papillae with taste buds on the taste bud slide.

6.0 TUTOR-MARKED ASSIGNMENT

1. Taste cells
 - a. are found only on the tongue.
 - b. extend through tiny openings called taste buds.
 - c. have no axons but release neurotransmitters when stimulated.
 - d. have axons that extend directly to the taste area of the cerebral cortex.

2. Which of these is *not* one of the basic tastes?
 - a. spicy
 - b. salty
 - c. bitter
 - d. umami
 - e. sour
3. What are the three kinds of lingual papillae, and where are they located?
4. Do taste receptors undergo adaptation?
5. Describe taste preference in relation to diet control.

7.0 REFERENCES/FURTHER READING

- Hutchinson, M., Mallat, J., Marieb, E.N., Wilhelm P.B. (2007). *A Brief Atlas of the Human Body*. Sna Franscisco: Pearson Education Inc.
- Katherine, M. A., Rogers & William, N. S. (2011). *Nurses! Test Yourself in Anatomy and Physiology*
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- Sadler, T.W. (2004), *Langman's Medical Embryology* (9th ed.).

UNIT 2 THE NOSE AND THE SENSE OF SMELL

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Olfactory epithelium and bulb
 - 3.2 Threshold for detection of odors
 - 3.3 Neuronal pathways for olfaction
 - 3.4 Clinical correlates
- 4.0 Conclusion
- 5.0 Summary
- 6.0 Tutor-Marked Assignment
- 7.0 References/Further Reading

1.0 INTRODUCTION

Smell is the least understood of our senses. This results partly from the fact that the sense of smell is a subjective phenomenon that cannot be studied with ease in lower animals. Another complicating problem is that the sense of smell is poorly developed in human beings in comparison with the sense of smell in many lower animals.

2.0 OBJECTIVES

At the end of this unit, you should be able to:

- describe the histological structure and function of the olfactory epithelium and the olfactory bulb
- explain how the perception of many different odours is possible
- describe the neuronal pathways for the sense of smell.

3.0 MAIN CONTENT

3.1 Olfactory Epithelium and Bulb

A small superior part of the nasal cavity is lined with olfactory epithelium. Ten million olfactory neurons are present within the olfactory epithelium. Olfactory neurons are bipolar neurons with dendrites extending to the epithelial surface of the nasal cavity. The ends of the dendrites are modified into bulbous enlargements with long, specialised cilia, called olfactory hairs, which lie in a thin mucous film on the epithelial surface. The mucus keeps the nasal epithelium moist, traps and dissolves molecules, and facilitates the removal of molecules

and particles from the olfactory epithelium. Airborne odorants become dissolved in the mucus on the surface of the epithelium and bind to molecules on the membranes of the olfactory hairs called **olfactory receptors**. The odorants must first be dissolved in fluid in order to reach the olfactory receptors. When an odorant binds to its receptor, a G protein associated with the receptor is activated. As a result of activation of the G protein, Na^+ and Ca^{2+} channels in the membrane open. The influx of ions into the olfactory hairs results in depolarization and the production of action potentials in the olfactory Neurons. Once an odorant binds to its receptor, however, the receptor accommodates and does not respond to another odorant for some time.

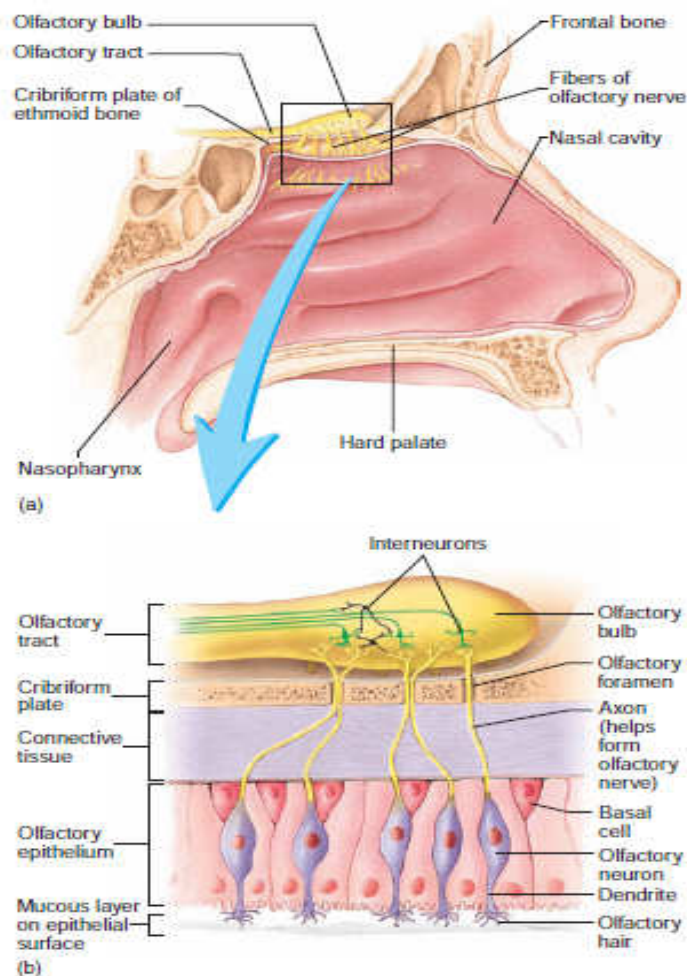


Figure 13.1 Olfactory Region, Epithelium, and Bulb **AP®**

(a) The lateral wall of the nasal cavity (cut in sagittal section), showing the olfactory nerves, olfactory bulb, and olfactory tract. (b) The olfactory cells within the olfactory epithelium, the axons of olfactory neurons passing through the cribriform plate, and the fine structure of the olfactory bulb.

3.2 Threshold for detection of odors

The threshold for the detection of odours is very low, so very few odorants bound to an olfactory neuron can initiate an action potential. Methylmercaptan, which has a nauseating odour similar to that of rotten

cabbage, is added to natural gas at a concentration of about one part per million. A person can detect the odour of about 1/25 billionth of a milligram of the substance and therefore is aware of the presence of the more dangerous but odourless natural gas. There are approximately 1000 different odorant receptors, which can react to odorants of different sizes, shapes, and functional groups. The average person can distinguish among approximately 4000 different smells. It has been proposed that the wide varieties of detectable odors are actually combinations of a smaller number of **primary odours**. The seven primary odours are camphoraceous (e.g., moth balls), musky, floral, peppermint, ethereal (e.g., fresh pears), pungent, and putrid. It is very unlikely, however, that this list is an accurate representation of all primary odours, and some studies point to the possibility of as many as 50 primary odours.

3.3 Neuronal Pathways for Olfaction

Axons from olfactory neurons form the **olfactory nerves (I)**, which pass through the olfactory foramina of the cribriform plate and enter the **olfactory bulb**. There they synapse with interneurons that relay action potentials to the brain through the **olfactory tracts**. Each olfactory tract terminates in the **olfactory cortex** and the **amygdala** in the temporal lobe. It is here that the sensation of smell is perceived. The olfactory cortex is part of the limbic system, projecting to the hypothalamus, amygdala, and hippocampus of the cerebrum. Odours can produce strong emotional reactions, memories, and other responses. For example, the perfume or cologne of a loved one can evoke good feelings, and memories and food odours can cause salivation and hunger. Within the olfactory bulb and olfactory cortex, feedback loops occur that tend to inhibit transmission of action potentials resulting from prolonged exposure to a given odorant. This feedback, plus the temporary decreased in sensitivity at the level of the receptors, results in adaptation to a given odour. For example, if you enter a room that has an odour, you are aware of the odour, but you adapt to the odour and cannot smell it as well after the first few minutes. If you leave the room for some time and then re - enter the room, the odour again seems more intense.

3.4 Clinical Correlates

- i. **Anosmia** - a complete loss of smell
- ii. **Hyposmia** – partial loss of smell or decreased sensation of smell
- iii. **Hyperosmia** – enhanced smell sensitivity
- iv. **Dysosmia** – distortion in odour perception (includes parosmia and phantosmia)

- v. **Parosmia** – distortion of perception of an external stimulus
- vi. **Phantosmia** – smell perception with no external stimulus.

Common causes of smell disorders include:

Aging, sinus and other respiratory infections, smoking, growths in nasal cavities, head injury, hormonal disturbances, dental problems, exposure to certain chemicals, such as insecticides and solvents, radiation for treatment of head and neck cancers.

4.0 CONCLUSION

In this unit you have learnt that the sense of smell is a subjective phenomenon that cannot be studied with ease in lower animals.

5.0 SUMMARY

- Olfaction is the sense of smell.
- Olfactory neurons in the olfactory epithelium are bipolar neurons.

Their distal ends have olfactory hairs. The olfactory hairs have receptors that respond to dissolved substances. There are approximately 1000 different odorant receptors.

- The receptors activate G proteins, which results in ion channels opening and depolarization.
- At least seven (perhaps 50) primary odours exist. The olfactory neurons have a very low threshold and accommodate rapidly.
- Axons from the olfactory neurons extend as olfactory nerves to the olfactory bulb, where they synapse with interneurons. Axons from these cells form the olfactory tracts, which connect to the olfactory cortex. The olfactory bulbs and cortex accommodate to odours.

SELF-ASSESSMENT EXERCISE

At the histology laboratory, examine the olfactory cells slides under the microscope.

At the gross anatomy laboratory, identify the location of the olfactory epithelium and its relationship to the olfactory bulbs and the cribriform plate on the sagittal head model.

6.0 TUTOR-MARKED ASSIGNMENT

1. Olfactory neurons
 - a. have projections called cilia.

- b. have axons that combine to form the olfactory nerves.
 - c. connect to the olfactory bulb.
 - d. have receptors that react with odorants dissolved in fluid.
 - e. all of the above.
2. What are the characteristics of an odorant—a chemical molecule that stimulates a smell receptor?
3. Does adaptation of smell receptors occur?
4. Do all of the volatile chemicals in the nose stimulate smell receptors?
4. Which of the cranial nerves innervate the olfactory mucosa?

7.0 REFERENCES/FURTHER READING

Hutchinson, M., Mallat, J., Marieb, E.N., Wilhelm P.B. (2007). *A Brief Atlas of the Human Body*. San Francisco: Pearson Education Inc.

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UNIT 3 THE EAR AND SENSE OF HEARING AND BALANCE

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- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Auditory Structures
 - 3.2 Auditory Functions
 - 3.3 Neuronal Pathways for Hearing and Balance
 - 3.4 Balance
 - 3.5 Clinical Correlates
- 4.0 Conclusion
- 5.0 Summary
- 6.0 Tutor-Marked Assignment
- 7.0 References/Further Reading

1.0 INTRODUCTION

The organs of hearing and balance are divided into three parts: the external, middle, and inner ear. The **external ear** is the part extending from the outside of the head to the **tympanic membrane**, or **eardrum**. The **middle ear** is an air-filled chamber medial to the tympanic membrane. The **inner ear** is a set of fluid-filled chambers medial to the middle ear. The external and middle ears are involved in hearing only, whereas the inner ear functions in both hearing and balance.

2.0 OBJECTIVES

At the end of this unit, you should be able to:

- describe the anatomy of the ear and the function of each part
- explain how sounds travel through the ear and are interpreted in the brain
- explain the role of the ear in maintaining equilibrium
- describe various disorders of the ear.

3.0 MAIN CONTENT

3.1 Auditory Structures

External Ear/ Auricle

The auricle is the fleshy part of the external ear on the outside of the head. The auricle opens into the external acoustic meatus, a passageway that leads to the tympanic membrane. The auricle collects sound waves

and directs them toward the external acoustic meatus, which transmits them to the tympanic membrane. The external acoustic meatus is lined with hairs and ceruminous glands, which produce cerumen, a modified sebum commonly called earwax. The hairs and cerumen help prevent foreign objects from reaching the delicate tympanic membrane. The tympanic membrane is a thin membrane separating the external ear from the middle ear. It consists of a thin layer of connective tissue sandwiched between two epithelial layers. Sound waves reaching the tympanic membrane cause it to vibrate.

Middle Ear

Medial to the tympanic membrane is the air-filled cavity of the middle ear. Two covered openings, the **oval window** and the **round window** on the medial side of the middle ear, connect the middle ear with the inner ear. The middle ear contains three **auditory ossicles**: the **malleus**, **incus**, and **stapes**. These bones transmit vibrations from the tympanic membrane to the oval window. The malleus is attached to the medial surface of the tympanic membrane. The incus connects the malleus to the stapes. The base of the stapes is seated in the oval window and is surrounded by a flexible ligament. Two small muscles in the middle ear help dampen vibrations of the auditory ossicles caused by loud noises. The **tensor tympani** muscle is attached to the malleus and is innervated by the trigeminal nerve (V). The **stapedius** muscle is attached to the stapes and is innervated by the facial nerve (VII).

Two openings provide air passages from the middle ear. One passage opens into the **mastoid air cells** in the mastoid process of the temporal bone. The other passageway is the **auditory tube**, also called the **pharyngotympanic tube** or the **eustachian tube**. The auditory tube opens into the pharynx and enables air pressure to be equalised between the outside air and the middle ear cavity.

When a person changes altitude, air pressure outside the tympanic membrane changes relative to the air pressure in the middle ear. With an increase in altitude, the pressure outside the tympanic membrane becomes less than the air pressure inside the middle ear and the tympanic membrane is pushed outward. With a decrease in altitude, the air pressure outside the ear becomes greater than in the middle ear and the tympanic membrane is pushed inward. Distortion of the tympanic membrane can make sounds seem muffled and stimulate pain. These symptoms can be relieved by opening the auditory tube to allow air to pass through the auditory tube to equalise air pressure. Swallowing, yawning, chewing, and holding the nose and mouth shut while gently trying to force air out of the lungs are methods used to open the auditory tube.

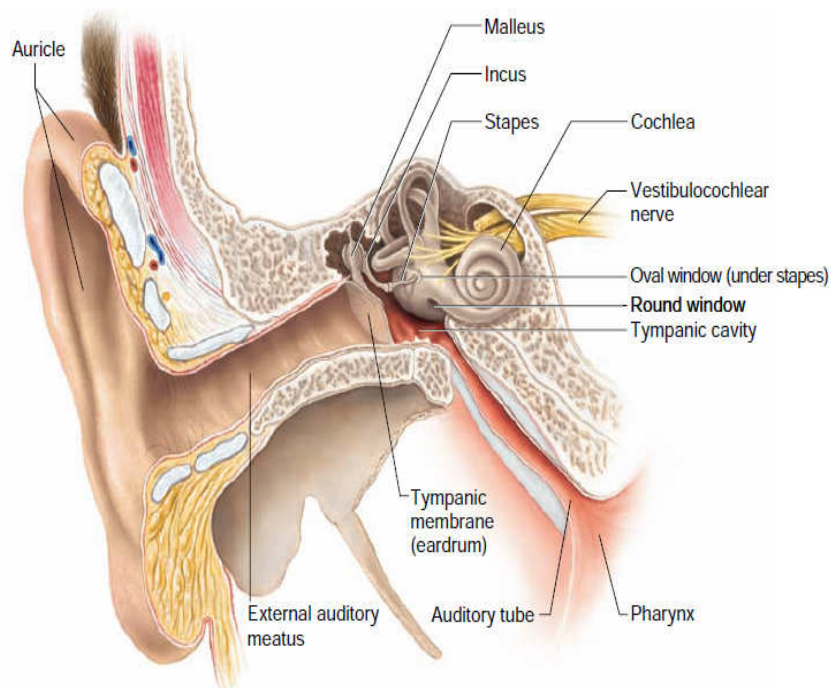


Fig.3.1 Major Parts of the Ear

Inner Ear

The inner ear contains the sensory organs for hearing and balance. It consists of interconnecting, fluid-filled tunnels and chambers within the temporal bone called the bony labyrinth. The bony labyrinth is lined with endosteum. When the inner ear is drawn, it is the endosteum that is depicted. The bony labyrinth is divided into three regions: cochlea, vestibule, and semicircular canals. The vestibule and semicircular canals are involved primarily in balance, and the cochlea is involved in hearing. Inside the bony labyrinth is a similarly shaped but smaller set of membranous tunnels and chambers called the membranous labyrinth. The membranous labyrinth is filled with a clear fluid called endolymph, and the space between the membranous and bony labyrinths is filled with a fluid called perilymph. Perilymph is very similar to cerebrospinal fluid, but endolymph has a high concentration of potassium and a low concentration of sodium, which is opposite from perilymph and cerebrospinal fluid. The membranous labyrinth of the cochlea separates the bony labyrinth into two parts. The cochlea is shaped like a snail shell—that is, a coiled tube.

The base of the cochlea connects to the vestibule and the apex of the cochlea is the end of the coiled tube. The bony core of the cochlea, around which the tube coils, is shaped like a screw with threads called the spiral lamina. A Y-shaped, membranous complex divides the cochlea into three portions. The base of the Y is the spiral lamina. One branch of the Y is the vestibular membrane, and the other branch is the

basilar membrane. The space between these membranes is called the cochlear duct.

3.2 Auditory Functions

Properties of Sound

Vibration of matter, such as air, water, or a solid material, creates sound. No sound occurs in a vacuum. For example, when a tuning fork vibrates, it causes the surrounding air to vibrate. The air vibrations consist of bands of compressed air followed by bands of less compressed air. When these pressure changes are graphed through time, they have a wave form; hence, they are called sound waves. These sound waves are propagated through the air, somewhat as ripples are propagated over the surface of water. The **volume**, or loudness, of sound is a function of wave amplitude, or height, measured in decibels. The greater the amplitude is, the louder the sound. The **pitch** of sound is a function of the wave frequency. It is measured in hertz (Hz), which is the number of waves or cycles per second. The higher the frequency is, the higher the pitch. The normal range of human hearing is 20–20,000 Hz and 0 or more decibels (db). Sounds louder than 125 db are painful to the ear. Timbre is the resonance quality or overtones of a sound. A smooth sigmoid curve is the image of a “pure” sound wave, but such a wave almost never exists in nature. The sounds made by musical instruments and the human voice are not smooth sigmoid curves but, rather, are rough, jagged curves formed by numerous, superimposed curves of various amplitudes and frequencies.

External Ear

The auricle collects sound waves and directs them into the external acoustic meatus. Sound waves travel relatively slowly in air, 332 m/s, and a small time interval may elapse between the time a sound wave reaches one ear and the time it reaches the other. The brain can interpret this interval to determine the direction from which a sound is coming. Sound waves are conducted through the external acoustic meatus and strike the tympanic membrane, causing it to vibrate.

Middle Ear

Vibration of the tympanic membrane causes vibration of the three auditory ossicles of the middle ear, and by this mechanical linkage vibration is transferred to the oval window. The oval window is approximately 20 times smaller than the tympanic membrane. The mechanical force of vibration is amplified about 20-fold as it passes from the tympanic membrane through the auditory ossicles to the oval window because of this size difference. This amplification is necessary

because more force is required to cause vibration in a liquid, such as the perilymph of the inner ear, than is required in air.

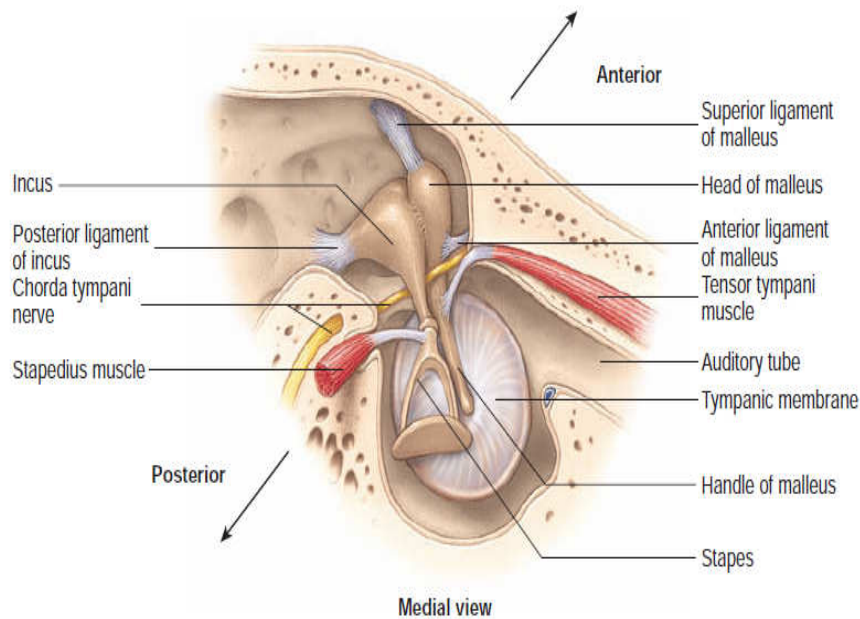


Fig.3.2: Middle Ear

Middle Ear

The tensor tympani and stapedius muscles are attached to auditory ossicles. Excessively loud sounds cause these muscles to reflexively contract and dampen the movement of the auditory ossicles. This **sound attenuation reflex** protects the delicate inner ear structures from damage by loud noises. The sound attenuation reflex responds most effectively to low-frequency sounds and can reduce by a factor of 100 the energy reaching the oval window. The reflex is too slow to prevent damage from a sudden noise, such as a gunshot, and it cannot function effectively for longer than about 10 minutes, in response to prolonged noise.

Inner Ear

As the stapes vibrates, it produces sound waves in the perilymph of the scala vestibuli. Vibrations of the perilymph are transmitted through the vestibular membrane and cause simultaneous vibrations of the endolymph. The mechanical effect is as though the perilymph and endolymph were a single fluid because the vestibular membrane is very thin. Vibration of the endolymph causes distortion of the basilar membrane, which is most important to hearing. As the basilar membrane moves, the hair cells resting on it move relative to the tectorial membrane, which remains stationary. The inner hair cell stereocilia are bent as they move against the tectorial membrane, resulting in the release of neurotransmitter and the production of action potentials in sensory neurons.

The basilar membrane is not uniform throughout its length. The membrane is narrower and denser near the oval window and wider and less dense near the apex of the cochlea. The various regions of the membrane can be compared to the strings in a piano, which has strings of varying length and thickness. As a result of this organisation, sounds with higher pitches cause the basilar membrane nearer the oval window to distort maximally, whereas sounds with lower pitches cause the basilar membrane nearer the apex of the cochlea to distort maximally. Depending on which hair cells are stimulated along the length of the basilar membrane, the brain interprets the pitch and timbre of sounds. Sound volume, or loudness, is a function of sound wave amplitude.

As the volume of sound increases, the vibration of the basilar membrane increases, the stimulation of hair cells increases, and the production of action potentials increases. The brain interprets the higher frequency of action potentials as a louder sound. The outer hair cells are involved in regulating the tension of the basilar membrane, thereby increasing the sensitivity of the inner ear to sounds. Stimulation of the inner hair cells by the nervous system stimulates the contraction of actin filaments within the hair cells, causing them to shorten. This adjustment in the height of the outer hair cells, attached to both the basilar membrane and the tectorial membrane, fine-tunes the tension of the basilar membrane and the distance between the basilar membrane and tectorial membrane. Sound waves in the perilymph of the scala vestibuli are also transmitted the length of the scala vestibuli and through the helicotrema into the perilymph of the scala tympani. This transmission of sound waves is probably of little consequence because the helicotrema is very small. Vibration of the basilar membrane produces most of the sound waves in the perilymph of the scala tympani. Sound waves in the scala tympani perilymph cause vibration of the membrane of the round window. Vibration of the round window membrane is important to hearing because it acts as a mechanical release for sound waves within the scala tympani. If this window were solid, it would reflect the sound waves, which would interfere with and dampen later sound waves. The round window also allows the relief of pressure in the perilymph because fluid is not compressible, thereby preventing compression damage to the spiral organ.

3.3 Neuronal Pathway for Hearing and Balance

The axons of the sensory neurons supplying hair cells form the cochlear nerve. These sensory neurons are bipolar neurons, and their cell bodies are in the cochlear, or spiral, ganglion, located in the bony core of the

cochlea. The cochlear nerve joins the vestibular nerve to become the vestibulocochlear nerve (VIII), which traverses the internal acoustic meatus and enters the cranial cavity. The special senses of hearing and balance are both transmitted by the vestibulocochlear (VIII) nerve.

The term *vestibular* refers to the vestibule of the inner ear, which is involved in balance. The term *cochlear* refers to the cochlea of the inner ear, which is involved in hearing. The vestibulocochlear nerve functions as two separate nerves carrying information from two separate but closely related structures. The auditory pathways within the CNS are very complex, with both crossed and uncrossed tracts. Unilateral CNS damage therefore usually has little effect on hearing. The neurons from the cochlear ganglion synapse with CNS neurons in the cochlear nucleus in the medulla oblongata. These neurons in turn either synapse in or pass through the superior olivary nucleus in the medulla oblongata. Neurons terminating in the superior olivary nucleus may synapse with efferent neurons returning to the cochlea to modulate pitch perception. Nerve fibers from the superior olivary nucleus also project to the trigeminal (V) nucleus, which controls the tensor tympani, and the facial (VII) nucleus, which controls the stapedius muscle. This is part of the sound attenuation reflex pathway.

Ascending neurons from the superior olivary nucleus synapse in the inferior colliculi, and neurons from there project to the thalamus, where they synapse with neurons that project to the cortex. These neurons terminate in the auditory cortex. Neurons from the inferior colliculi also project to the superior colliculi, where reflexes that turn the head and eyes in response to loud sounds are initiated.

Neuronal Pathways for Balance

The axons of the sensory neurons supplying hair cells of the maculae and crista ampullaris form the vestibular nerve. These sensory neurons are bipolar neurons, and their cell bodies are in the vestibular ganglion, a swelling of the vestibulocochlear nerve located in the internal acoustic meatus. The sensory neurons terminate in the vestibular nucleus within the medulla oblongata. Axons run from this nucleus to numerous areas of the CNS, such as the spinal cord, the cerebellum, the cerebral cortex, and the nuclei controlling the extrinsic eye muscles.

Balance is a complex process not simply confined to one type of input. In addition to vestibular sensory input, the vestibular nucleus receives input from proprioceptive neurons throughout the body, as well as from the visual system. People are asked to close their eyes while balance is evaluated in a sobriety test because alcohol affects the proprioceptive and vestibular components of balance (cerebellar function) to a greater extent than it does the visual portion.

Reflex pathways exist between the dynamic balance part of the vestibular system and the nuclei controlling the extrinsic eye muscles (oculomotor, trochlear, and abducent). A reflex pathway allows the maintenance of visual fixation on an object while the head is in motion. This function can be demonstrated by spinning a person around about 10 times in 20 seconds, stopping him or her, and observing eye movements. A slight oscillatory movement of the eyes, called nystagmus, occurs. The eyes track in the direction of motion and return with a rapid recovery movement before repeating the tracking motion.

3.4 Balance

The sense of balance, or equilibrium, has two components: static balance and dynamic balance. Static balance is associated with the vestibule and is involved in evaluating the position of the head relative to gravity. Dynamic balance is associated with the semicircular canals and is involved in evaluating changes in the direction and rate of head movements.

Static balance

Static balance is associated with the utricle and the saccule of the vestibule. It is primarily involved in evaluating the position of the head relative to gravity, although the system also responds to linear acceleration or deceleration, such as when a person is in a car that is increasing or decreasing speed. Most of the utricular and saccular walls consist of simple cuboidal epithelium. The utricle and saccule, however, each contain a specialised patch of epithelium about two to three mm in diameter called the macula. The macula of the utricle is oriented parallel to the base of the skull, and the macula of the saccule is perpendicular to the base of the skull. The maculae resemble the spiral organ and consist of hair cells, sensory neurons, and supporting cells. The “hairs” of the hair cells consist of numerous microvilli, called **stereocilia**, and one cilium, called a **kinocilium**.

The stereocilia and kinocilium are embedded in the **otolithic membrane**, which is a gelatinous mass weighted with crystals of calcium carbonate and protein called **otoliths**. The otolithic membrane moves in response to gravity, bending the hair cells and initiating action potentials in their associated sensory neurons. The stereocilia function much as do the stereocilia of cochlear hair cells, with tip links connected to gated K⁺ channels. Deflection of the hairs toward the kinocilium results in depolarization of the hair cell. The hair cells are constantly being stimulated at a low level by the presence of the otolith-weighted

covering of the macula. When the otolithic membrane moves in response to gravity, the pattern and intensity of hair cell stimulation change. The change in action potentials produced is translated by the brain into specific information about head position or linear acceleration/deceleration. Much of this information is not perceived consciously but is dealt with subconsciously. The body responds by making subtle tone adjustments in the muscles of the back and neck, which are intended to restore the head to its proper neutral, balanced position.

Dynamic Balance

Dynamic balance is associated with the semicircular canals and is involved in evaluating movements of the head. There are three **semicircular canals** placed at nearly right angles to one another. One semicircular canal lies nearly in the transverse plane, one in the frontal plane, and one in the sagittal plane. The arrangement of the semicircular canals enables a person to detect movement in all directions. The base of each semicircular canal is expanded into an **ampulla**. Within each ampulla, the epithelium is specialised to form a **crista ampullaris**. This specialised sensory epithelium is structurally and functionally very similar to the sensory epithelium of the maculae.

Each crista consists of a ridge or crest of epithelium with a curved, gelatinous mass, the **cupula**, suspended over the crest. The hair like processes of the crista hair cells, which are stereocilia similar to those in the maculae and cochlear hair cells, are embedded in the cupula. The cupula contains no otoliths and therefore does not respond to gravitational pull. Instead, the cupula is a float that is displaced by endolymph movements within the semicircular canals. As the head begins to move, the endolymph does not move at the same rate as the semicircular canals, which are part of the temporal one. This difference causes displacement of the cupula in a direction opposite to that of the movement of the head. To appreciate this effect, imagine holding a feather (the cupula) in your hand (the crista ampullaris). If you rapidly move your hand, the feather bends over in the direction opposite the movement as it drags through the air (endolymph). The bending of the stereocilia results in the stimulation of sensory neurons. The brain interprets the direction of head movement based on which hair cells in which crista ampullaris are stimulated. The semicircular canals detect changes in the rate of movement rather than movement alone because displacement of the cupula is most intense when the rate of head movement changes rapidly. As with the static balance, the information the brain obtains regarding dynamic balance is largely subconscious.

If a person continually spins in one direction, the endolymph of the semicircular canals begins to move and catches up with the cupula, and stimulation is stopped. If spinning suddenly stops, the endolymph continues to move because of its momentum, causing displacement of the cupula in the same direction as the head had been moving. The brain interprets this movement of the cupula to mean the head is moving in the opposite direction of the spin, even though the head is no longer moving. This is why a person has a feeling of moving even after he or she has stopped spinning.

3.5 Clinical Correlates

i. Tympanic Membrane Rupture

Rupture of the tympanic membrane may result in hearing impairment. Foreign objects thrust into the ear, pressure, and infections of the middle ear can rupture the tympanic membrane. Sufficient differential pressure between the middle ear and the outside air can also rupture the tympanic membrane. This can occur in flyers, divers, or individuals who are hit on the side of the head by an open hand.

ii. Chorda Tympani

The chorda tympani is a branch of the facial nerve carrying taste impulses from the anterior two-thirds of the tongue. It crosses through the middle ear between the malleus and incus. The chorda tympani have nothing to do with hearing but is just passing through. This nerve can be damaged, however, during ear surgery or by a middle ear infection, resulting in loss of taste sensation from the anterior two-thirds of the tongue on the side innervated by that nerve.

iii. Human Speech and Hearing Impairment

The range of normal human speech is 250–8000 Hz. This is the range that is tested for the possibility of hearing impairment because it is the most important for communication.

iv. Loud Noises and Hearing Loss

Prolonged or frequent exposure to excessively loud noises can cause degeneration of the spiral organ at the base of the cochlea, resulting in high-frequency deafness. The actual amount of damage can vary greatly from person to person. High-frequency loss can cause a person to miss hearing consonants in a noisy setting. Loud music, amplified to 120 db, can impair hearing. The defects may not be detectable on routine diagnosis, but they include decreased sensitivity to sound in specific narrow frequency ranges and a decreased ability to discriminate between two pitches. Loud music, however, is not as harmful as is the sound of a nearby gunshot, which is a sudden sound occurring at 140 db. The sound

is too sudden for the attenuation reflex to protect the inner ear structures, and the intensity is great enough to cause auditory damage. In fact, gunshot noise is the most common recreational cause of serious hearing loss.

v. **Meniere Disease**

Meniere disease is the most common disease involving dizziness from the inner ear. Its cause is unknown but it appears to involve a fluid abnormality in one (usually) or both ears. Symptoms include vertigo, hearing loss, tinnitus, and a feeling of “fullness” in the affected ear. Treatment includes a low-salt diet and diuretics (water pills). Symptoms may also be treated with medications for motion sickness.

4.0 CONCLUSION

In this unit you learnt about the organs of hearing and balance which are divided into three parts: the external, middle, and inner ear.

5.0 SUMMARY

- The osseous labyrinth is a canal system within the temporal bone that contains perilymph and the membranous labyrinth. Endolymph is inside the membranous labyrinth.
- The external ear consists of the auricle and external acoustic meatus. The middle ear connects the external and inner ears. The tympanic membrane is stretched across the external acoustic meatus.
- The malleus, incus, and stapes connect the tympanic membrane to the oval window of the inner ear.
- The auditory tube connects the middle ear to the pharynx and equalises pressure. The middle ear is connected to the mastoid air cells. The inner ear has three parts: the semicircular canals; the vestibule, which contains the utricle and the saccule; and the cochlea.
- The cochlea is a spiral-shaped canal within the temporal bone. The cochlea is divided into three compartments by the vestibular and basilar membranes. The scala vestibuli and scala tympani contain perilymph. The cochlear duct contains endolymph and the spiral organ.
- The spiral organ consists of inner hair cells and outer hair cells, which attach to the tectorial membrane.
- Pitch is determined by the frequency of sound waves and volume by the amplitude of sound waves. Timbre is the resonance quality (overtones) of sound.

- Axons from the vestibulocochlear nerve synapse in the medulla. Neurons from the medulla project axons to the inferior colliculi, where they synapse. Neurons from this point project to the thalamus and synapse. Thalamic neurons extend to the auditory cortex.

SELF-ASSESSMENT EXERCISE

Identify the major anatomical features on the ear models provided.

6.0 TUTOR-MARKED ASSIGNMENT

1. Name the three parts of the ear, and state their functions.
2. Describe the structural components of the *middle ear* and their functions.
3. Which of these structures is found within or is a part of the external ear?
 - a. oval window
 - b. auditory tube
 - c. ossicles
 - d. external acoustic meatus
 - e. cochlear duct
4. Given these auditory ossicles:
 - incus(1)
 - malleus(2)
 - stapes(3)

Choose the arrangement that lists the auditory ossicles in order from the tympanic membrane to the inner ear.

- a. 1,2,3
 - b. 1,3,2
 - c. 2,1,3
 - d. 2,3,1
 - e. 3,2,1
5. The spiral organ is found within the
 - a. cochlear duct.
 - b. scala vestibuli.
 - c. scala tympani.
 - d. vestibule.
 - e. semicircular canals.
 6. An increase in the loudness of sound occurs as a result of an increase in the sound wave.
 - a. frequency
 - b. amplitude
 - c. resonance
 - d. both a and b

7. Interpretation of different sounds is possible because of the ability of it to vibrate at different frequencies and stimulate the:
- vestibular membrane, vestibular nerve
 - vestibular membrane, spiral organ
 - basilar membrane, vestibular nerve
 - basilar membrane, spiral organ

7.0 REFERENCES/FURTHER READING

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UNIT 4 THE EYES AND THE SENSE OF VISION

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Anatomy of the Eye
 - 3.2 Functions of the Eye
 - 3.3 Visual Pathway
 - 3.4 Clinical Correlates
- 4.0 Conclusion
- 5.0 Summary
- 6.0 Tutor-Marked Assignment
- 7.0 References/Further Reading

1.0 INTRODUCTION

The visual system includes the eyes, the accessory structures, and the sensory neurons that project to the cerebral cortex where action potentials conveying visual information are interpreted. The **eye** consists of the **eyeball**, or globe of the eye, and the optic nerve. Much of the information we obtain about the world around us is detected by the visual system. Our education is largely based on visual input and depends on our ability to read words and numbers. Visual input includes information about light and dark, movement, colour and hue.

2.0 OBJECTIVES

At the end of this unit, you should be able to:

- list the accessory structures of the eye, and explain their functions
- describe the anatomy of the eye
- describe the focusing system of the eye and how it adjusts to see distant and near objects
- explain the structure of the retina, and how light entering the eye results in action potentials in the optic nerve
- outline the neuronal pathways for vision.

3.1 Anatomy of the Eye

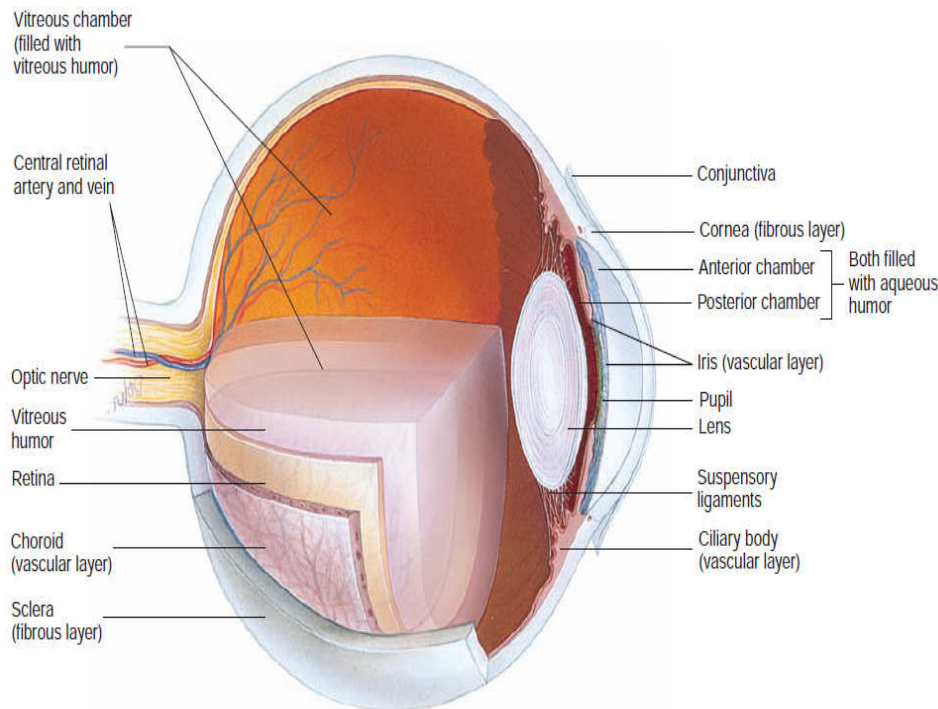


Fig. 4.1: Sagittal Section of the Eye

The eyeball is composed of three layers. From superficial to deep they are the **fibrous layer**, **vascular layer**, and **retina**. The fibrous layer consists of the sclera and cornea; the vascular layer consists of the choroid, ciliary body, and iris.

Fibrous Layer

The **sclera** is the firm, opaque, white, outer layer of the posterior five-sixths of the eyeball. It consists of dense collagenous connective tissue with elastic fibers. The sclera helps maintain the shape of the eyeball, protects its internal structures, and provides an attachment point for the extrinsic eye muscles. A small portion of the sclera can be seen as the “white of the eye”.

The sclera is continuous with the **cornea**, the anterior sixth of the eyeball. The cornea is avascular and transparent, permitting light to enter the eye. The focusing system of the eye refracts, or bends, light and focuses it on the nervous layer (retina). The cornea is responsible for most of the refraction of light entering the eye.

The cornea consists of a connective tissue matrix containing collagen, elastic fibers, and proteoglycans, with a layer of stratified squamous epithelium covering the outer surface and a layer of simple squamous

epithelium on the inner surface. The outer epithelium is continuous with the conjunctiva over the sclera. The transparency of the cornea results from its low water content. In the presence of water, proteoglycans trap water and expand, which scatters light. In the absence of water, the proteoglycans decrease in size and do not interfere with the passage of light through the matrix.

Vascular Layer

The vascular layer of the eye is so named because it contains most of the blood vessels of the eye. The posterior portion of the vascular layer, associated with the sclera, is the **choroid**. This is a very thin structure consisting of a vascular network and many melanin-containing pigment cells so that it appears black in colour. The black colour absorbs light so that it is not reflected inside the eye. If light were reflected inside the eye, the reflection would interfere with vision. The interiors of cameras are black for the same reason.

The choroid is continuous anteriorly with the ciliary body, which consists of an outer ciliary ring and an inner group of ciliary processes. The ciliary ring and the base of the ciliary processes contain smooth muscle called ciliary muscles. Suspensory ligaments attach the ciliary ring and processes to the lens of the eye, and contraction of the ciliary muscles can change the shape of the lens. The ciliary process also produces aqueous humour. The ciliary body is continuous anteriorly with the iris of the eye, which is the “coloured part” of the eye. The iris is a contractile structure, consisting mainly of smooth muscle, surrounding an opening called the pupil. Light enters the eye through the pupil, and the iris regulates the amount of light by controlling the size of the pupil. The iris contains two groups of smooth muscles: a circular group called the sphincter pupillae and a radial group called the dilator pupillae. The sphincter pupillae are innervated by parasympathetic fibers from the oculomotor nerve (III). When they contract, the pupil constricts and less light enters the eye. The dilator pupillae are innervated by sympathetic fibers. When they contract, the pupil dilates and more light enters the eye. The ciliary muscles, sphincter pupillae, and dilator pupillae are sometimes referred to as the intrinsic eye muscles.

The colour of the eye differs from person to person. A large amount of melanin in the iris causes it to appear brown or even black. Less melanin results in light brown, green, or gray irises. Even less melanin causes the eyes to appear blue. If there is no pigment in the iris, as in albinism, the iris is pink because blood vessels in the eye reflect light back to the iris. The genetics of eye colour is quite complex.

Retina

The retina is the inner layer of the eye, covering the inner surface of the eye posterior to the ciliary body. The retina has over 126 million photoreceptor cells, which respond to light. As a result, action potentials are generated and conducted by the optic nerve (II) out of the eye to the cerebral cortex, where the sense of vision takes place. When the posterior region of the retina is examined with an ophthalmoscope, several important features can be observed. Near the center of the posterior retina is a small, yellow spot approximately four (4) mm in diameter, the macula.

In the center of the macula is a small pit, the fovea centralise. The fovea, followed by the macula, has the highest concentration of photoreceptor cells in the retina. Thus, they are the part of the retina most sensitive to light. Just medial to the macula is a white spot, the optic disc, through which the central retinal artery enters and the central retinal vein exits the eyeball. Branches from these vessels spread over the surface of the retina. The optic disc is also the place where axons from the neurons of the retina converge to form the optic nerve, which exits the posterior eye.

The optic disc contains only axons and no photoreceptor cells. Therefore, it does not respond to light and is called the blind spot of the eye.

Chambers of the Eye

The interior of the eye is divided into three chambers: **anterior chamber**, **posterior chamber**, and **vitreous chamber**. The anterior and posterior chambers are located between the cornea and the lens. The iris separates the anterior chamber from the posterior chamber, which are continuous with each other through the pupil. The much larger vitreous chamber is posterior to the lens. A fluid called **aqueous humour** fills the anterior and posterior chambers. Aqueous humour helps maintain intraocular pressure, which keeps the eyeball inflated and is largely responsible for maintaining the shape of the eyeball. The aqueous humour also refracts light and provides nutrition for the structures of the anterior chamber, such as the cornea, which has no blood vessels. Aqueous humour is produced by the ciliary processes as a blood filtrate and is returned to the circulation through a venous ring called the **scleral venous sinus** (canal of Schlemm) (shlem), which is located at the junction of the sclera and cornea. The production and removal of aqueous humour result in the “circulation” of aqueous humour and maintenance of a constant intraocular pressure.

Glaucoma is an abnormal increase in intraocular pressure that results when the rate of production of aqueous humour exceeds its rate of removal. A transparent, jellylike substance called **vitreous humour** fills the vitreous chamber. The vitreous humour helps maintain intraocular pressure and therefore the shape of the eyeball, and it holds the lens and retina in place. It also functions in the refraction of light in the eye. Vitreous humour is not produced as rapidly as is the aqueous humour, and its turnover is extremely slow.

Lens

The **lens** is an avascular, transparent, biconvex disk located behind the pupil. *Biconvex* means that each side of the lens bulges outward. The lens is part of the focusing system of the eye, and light passing through the lens is focused on the retina. The lens is a flexible structure, and changing the shape of the lens is involved with adjusting the focus of light. The lens consists of a layer of cuboidal epithelial cells on its anterior surface and a posterior region of very long, columnar epithelial cells called **lens fibers**. Cells from the anterior epithelium proliferate and give rise to the lens fibers. The lens fibers lose their nuclei and other cellular organelles and accumulate a set of proteins called **crystalline**. A highly elastic, transparent **lens capsule** surrounds the lens. The suspensory ligaments connect the lens capsule to the ciliary body. Through the lens capsule and the suspensory ligaments, the ciliary body can change the shape of the lens.

Accessory Structures

Accessory structures protect, lubricate, and move the eye. They include the eyebrows, eyelids, conjunctiva, lacrimal apparatus, and extrinsic eye muscles.

Eyebrows

The **eyebrows** are curved lines of hair over the orbit. They protect the eyes by preventing perspiration, which can irritate the eyes, from running down the forehead and into them, and they help shade the eyes from direct sunlight.

Eyelids

The **eyelids** are moveable folds covering the anterior surface of the eye when closed. The upper and lower eyelids meet at the **medial** and **lateral angles of the eye**. The medial angle contains a small, reddish-pink mound called the **caruncle**. The caruncle contains some modified sebaceous and sweat glands. The **eyelids** consist of five layers of tissue. From the outer to the inner surface, they are (1) a thin layer of skin on the external surface; (2) a thin layer of loose connective tissue; (3) a layer of skeletal muscle consisting of the orbicularis oculi and levator

palpebrae superioris muscles; (4) a crescent-shaped layer of dense connective tissue called the **tarsal plate**, which helps maintain the shape of the eyelid; and (5) the conjunctiva.

The eyelids, with their associated lashes, protect the eyes from foreign objects. If an object suddenly approaches the eye, the eyelids protect the eye by rapidly closing and then opening, a response called the blink reflex. Blinking, which normally occurs about 25 times per minute, also helps keep the eye lubricated by spreading tears over the surface. Movements of the eyelids are a function of skeletal muscles. The orbicularis oculi muscle closes the lids, and the levator palpebrae superioris elevates the upper lid. The eyelids also help regulate the amount of light entering the eye. **Eyelashes** are attached as a double or triple row of hairs to the free edges of the eyelids. **Ciliary glands** are modified sweat glands that open into the hair follicles of the eyelashes to keep them lubricated. When one of these glands becomes inflamed, it is called a **sty**. **Tarsal glands**, or **meibomian glands**, are sebaceous glands near the inner margins of the eyelids. They produce **sebum**, an oily, semifluid substance that lubricates the lids and restrains tears from flowing over the margin of the eyelids. An infection or a blockage of a tarsal gland is called a **chalazion**, or **meibomian cyst**.

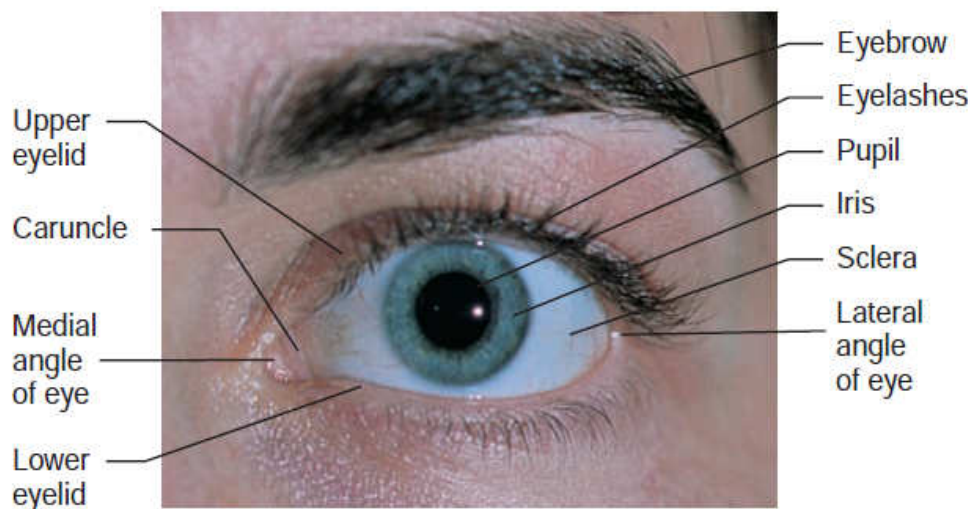


Fig. 4.2: The Left Eye and Accessory Structures.

Conjunctiva

The **conjunctiva** is a thin, transparent mucous membrane. It covers the inner surface of the eyelids and the anterior white surface of the eye. The conjunctiva reduces friction as the eyelids move over the surface of the eye. It also is a barrier to the entry of microorganisms.

Lacrimal Apparatus

The **lacrimal apparatus** consists of a lacrimal gland, lacrimal canaliculi, and a nasolacrimal duct. The lacrimal apparatus produces tears, releases the tears onto the surface of the eye, and removes excess tears from the surface. The **lacrimal gland** is in the superolateral corner of the orbit and is innervated by parasympathetic fibers from the facial nerve (VII). The lacrimal gland produces tears, which leave the gland through several **lacrimal ducts** and pass over the anterior surface of the eyeball. The lacrimal gland produces tears constantly at the rate of about one (1) mL/day to moisten the surface of the eye, lubricate the eyelids, and wash away foreign objects. Tears are mostly water, with some salts, mucus, and lysozyme, an enzyme that kills certain bacteria. Most of the fluid produced by the lacrimal glands evaporates from the surface of the eye, but excess tears are collected in the medial corner of the eye by small tubes called the **lacrimal canaliculi**. One lacrimal canaliculus opens on the inner, medial surface of the upper eyelid, and the other lacrimal canaliculus opens on the inner, medial surface of the lower eyelid. The lacrimal canaliculi connect to the **nasolacrimal duct**, which opens into the inferior meatus of the nasal cavity beneath the inferior nasal concha.

Extrinsic Eye Muscles

The **extrinsic eye muscles** attach to the outside of the eyeball and cause it to move. There are four rectus muscles, the **superior, inferior, medial, and lateral rectus muscles**.

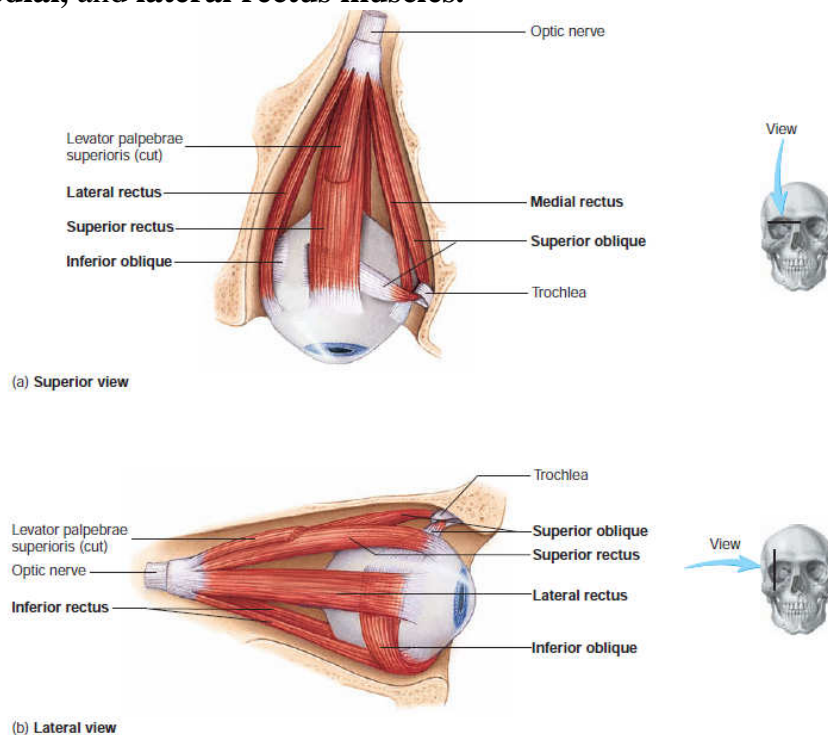


Fig. 4.3: The Extrinsic Muscles of the Right Eye

Rectus means straight, and the fibers of these muscles are nearly straight with the anterior-posterior axis of the eyeball. There are two oblique muscles, the **superior** and **inferior oblique muscles**, so named because they are at an angle to the axis of the eyeball. The movements of the eye can be described graphically by a figure resembling the letter *H*. The clinical test for normal eye movement is therefore called the **H test**. A person's inability to move the eye toward one part of the *H* may indicate dysfunction of an extrinsic eye muscle or the cranial nerve to the muscle. The superior oblique muscle is innervated by the trochlear nerve (IV). The nerve is so named because the superior oblique muscle goes around a little pulley, or trochlea, in the superomedial corner of the orbit. The lateral rectus muscle is innervated by the abducent nerve (VI), so named because the lateral rectus muscle abducts the eye. The other four extrinsic eye muscles are innervated by the oculomotor nerve (III).

3.2 Functions of the Eye

The eye receives light and produces action potentials. When the brain interprets the action potentials, it results in vision.

Properties of Light

The **electromagnetic spectrum** is the entire range of wavelengths, or frequencies, of electromagnetic radiation. Gamma waves have the shortest wavelength and radio waves the longest wavelength. **Visible light**, the portion of the electromagnetic spectrum that can be detected by the human eye, is a small part of the electromagnetic spectrum. Within the visible spectrum, each colour has a different wavelength. An important characteristic of light is that it can be refracted, or bent. As light passes from air to a denser substance, such as glass or water, its speed is reduced. If the surface of that substance is at an angle other than 90 degrees to the direction the light rays are traveling, the rays are bent as a result of variation in the speed of light as it encounters the new medium. This bending of light is called **refraction**. The greater the curvature of the surface, the greater is the refraction of light. If the surface of a lens is concave, with the lens thinnest in the center, the light rays diverge as a result of refraction. If the surface is convex, with the lens thickest in the center, the light rays converge. As light rays converge, they finally reach a point at which they cross. This point is called the **focal point (FP)**, and causing light to converge is called **focusing**.

If light rays strike an object that is not transparent, they bounce off the surface. This phenomenon is called **reflection**. The images we see result from light reflected from objects.

The Eye as a Camera

The eye is optically equivalent to the usual photographic camera. It has a lens system, a variable aperture system (the pupil), and a retina that corresponds to the film. The lens system of the eye is composed of four refractive interfaces: (1) the interface between air and the anterior surface of the cornea, (2) the interface between the posterior surface of the cornea and the aqueous humour, (3) the interface between the aqueous humour and the anterior surface of the lens of the eye, and (4) the interface between the posterior surface of the lens and the vitreous humour. The internal index of air is 1; the cornea, 1.38; the aqueous humour, 1.33; the crystalline (on average), 1.40; and the vitreous humour, 1.34.

Focusing System of the Eye

The light entering the eye passes through the focusing system of the eye to strike the retina. The **focusing system of the eye**, which refracts light, is the cornea, aqueous humour, lens, and vitreous humour. Light passing through the focusing system is refracted, producing a focal point. No image is produced at the focal point. Past the focal point is a place where the image passing through the focusing system can be clearly seen. In a normal eye, the focused image falls on the retina. The image is inverted and reversed right to left because the light rays cross at the focal point.

The cornea and lens are the most important elements of the focusing system of the eye. The cornea is responsible for most of the refraction of light because the greatest contrast in media density is between the air and the cornea. The shape of the cornea and its distance from the retina are fixed, however, so that no adjustment in the location of the focused image can be made by the cornea. Fine adjustments to the location of the focused image are accomplished by changing the shape of the lens. Increasing the curvature of the lens increases the refraction of light, moving the focused image closer to the lens. Decreasing the curvature of the lens decreases the refraction of light, moving the focused image farther from the lens. In cameras, microscopes, and telescopes, focusing is not accomplished by changing lens shape. Instead, focusing is accomplished by moving the lens closer to or farther from the point at which the image will be focused.

Distant and Near Vision

Distant vision occurs when looking at objects 20 feet or more from the eye, whereas **near vision** occurs when looking at objects that are less than 20 feet from the eye. In distant vision, the ciliary muscles in the ciliary body are relaxed. The suspensory ligaments, however, maintain

elastic pressure on the lens, thereby keeping it relatively flat. The condition in which the lens is flattened so that nearly parallel rays from a distant object are focused on the retina is referred to as **emmetropia** and is the normal resting condition of the lens. The point at which the lens does not have to thicken for focusing to occur is called the **far point of vision** and normally is 20 feet or more from the eye.

When an object is brought closer than 20 feet to the eye, the image falling on the retina is no longer in focus. Three events occur to bring the image into focus on the retina: accommodation by the lens, constriction of the pupil, and convergence of the eyes.

1. *Accommodation.* When the eye focuses on a nearby object, the ciliary muscles contract as a result of parasympathetic stimulation from the oculomotor nerve (III). This sphincter like contraction pulls the choroid toward the lens to reduce the tension on the suspensory ligaments. This allows the lens to assume a more spherical form because of its own elastic nature. The more spherical lens has a more convex surface, causing greater refraction of light, which brings the image back into focus on the retina. This process is called **accommodation**.

As an object is brought closer and closer to the eye, accommodation becomes more and more difficult because the lens cannot become any more convex. At some point, the eye no longer can focus the object, and it is seen as a blur. The point at which this blurring occurs is called the **near point of vision**, which is usually about two to three inches from the eye for children, four to six inches for a young adult, 20 inches for a 45-year-old adult, and 60 inches for an 80-year-old adult. This increase in the near point of vision is called **presbyopia**. It occurs because the lens becomes more rigid with increasing age, which is why some older people say they could read with no problem if they only had longer arms.

2. *Pupil constriction.* When we look at a close-up object, the pupil diameter decreases, which increases the depth of focus. The **depth of focus** is the greatest distances through which an object can be moved and still remain in focus on the retina. The main factor affecting depth of focus is the size of the pupil. If the pupillary diameter is small, the depth of focus is greater than if the pupillary diameter is large. With a smaller pupillary opening, an object may therefore be moved slightly nearer or farther from the eye without disturbing its focus. This is particularly important when viewing an object at close range because the interest in detail is much greater, and therefore the acceptable margin for error is smaller. When the pupil is constricted, the light entering the eye tends to pass more nearly through the center of the lens and is more accurately focused than light passing through the edges of the lens. Pupillary diameter also regulates the amount of light entering the eye. The smaller

the pupil diameter, the less light entering the eye. As the pupil constricts during close vision, therefore, more light is required on the object being observed.

3. *Convergence.* Because the light rays entering the eyes from a distant object are nearly parallel, both pupils can pick up the light rays when the eyes are directed more or less straight ahead. As an object moves closer, however, the eyes must be rotated medially so that the object is kept focused on corresponding areas of each retina. Otherwise, the object appears blurry. This medial rotation of the eyes is accomplished by a reflex that stimulates the medial rectus muscle of each eye. This movement of the eyes is called convergence.

Structure and Function of the Retina

The retina consists of an inner, neural layer and an outer, pigmented layer. The **neural layer** contains three layers of neurons: photoreceptor, bipolar, and ganglionic. The photoreceptor layer contains **rods** and **cones**, which are the photoreceptor cells that respond to light. The rods and cones synapse with **bipolar cells**, which in turn synapse with **ganglion cells**. Axons from the ganglion cells pass over the inner surface of the retina, converge at the optic disc (blind spot), and exit the eye as the optic nerve (CN II). The neural layers are separated by the plexiform (like a braid) layers. The outer plexiform layer is where the photoreceptor cells synapse with the bipolar cells and the inner plexiform layer is where the bipolar cells synapse with the ganglion cells.

The **pigmented layer** consists of **retinal pigment epithelium (RPE)**, a single layer of cuboidal epithelial cells filled with melanin. It rests on the **Bruch membrane**, which is the inner layer of the choroid consisting of collagen and elastic fibers. Cells of the RPE phagocytize the spent tips of rods and cones and produce retinal from vitamin A. Along with the choroid, the pigmented layer provides a black-brown matrix that enhances visual acuity by isolating individual photoreceptors and reducing light scattering.

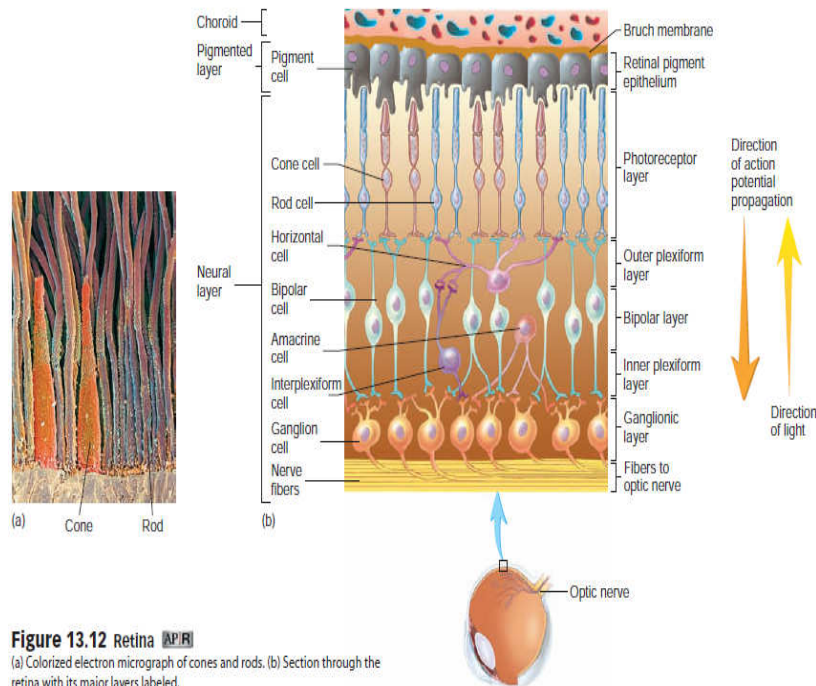


Figure 13.12 Retina (a) Colorized electron micrograph of cones and rods. (b) Section through the retina with its major layers labeled.

Fig. 4.4: Retina

Rods

Rods are responsible for non - colour vision and vision under conditions of reduced light. Even though rods are very sensitive to light, they cannot detect colour, and sensory input reaching the brain from rods is interpreted by the brain as shades of gray. Rods are bipolar neurons with modified, light-sensitive dendrites, which are cylindrical in shape.

Cones

Cones are responsible for colour vision and visual acuity. Colour is a function of the wavelength of light, and each colour results from a certain wavelength of visible light. Cones require relatively bright light to function. As the light decreases, so does the colour of objects that can be seen until, under conditions of very low illumination, the objects appear gray. This occurs because, as the light decreases, the number of cones responding to the light decreases but the number of rods increases. Cones are bipolar photoreceptor cells with a conical, light- sensitive part that tapers slightly from base to apex.

Distribution of Rods and Cones in the Retina

Each eye has approximately 120 million rods and six to seven million cones. The cones are most concentrated in the fovea centralis and the macula. The fovea centralis has approximately 35,000 cones and no rods. The rest of the macula has more cones than rods. Cones are involved in visual acuity, in addition to their role in colour vision. When one is looking at an object directly in front of the eye, the focusing

system of the eye places the image on the macula and fovea centralis. The high concentration of cones makes it possible to see fine details. The rods are 10–20 times more plentiful than cones over most of the retina away from the macula. The high number of rods enables them to “collect” light, and they are more important in low-light conditions.

Inner Layers of the Retina

Within the inner layers of the retina, interneurons modify the signals from the photoreceptor cells before the signal leaves the retina.

Horizontal cells in the outer plexiform layer synapse with photoreceptor cells and bipolar cells. **Amacrine cells** in the inner plexiform layer synapse with bipolar and ganglion cells. **Interplexiform cells** connect cells in the outer and inner plexiform layers, forming feedback loops. The interneurons are either excitatory or inhibitory on the cells with which they synapse. By increasing the signal from some photoreceptors and decreasing the signal from others, these interneurons increase the differences between boundaries, such as the edge of a dark object against a light background.

3.3 Visual Pathway

The optic nerve (II) leaves the eye and exits the orbit through the optic foramen to enter the cranial cavity. Just anterior to the pituitary gland, the optic nerves are connected to each other at the **optic chiasm**. Ganglion cell axons from the nasal (medial) retina cross through the optic chiasm and project to the opposite side of the brain. Ganglion cell axons from the temporal (lateral) retina pass through the optic chiasm and project to the brain on the same side of the body without crossing. Beyond the optic chiasm, the axons form the **optic tracts**. Most of the optic tract axons terminate in the thalamus. Some axons do not terminate in the thalamus but separate from the optic tracts to terminate in the **superior colliculi**, the center for visual reflexes. Neurons from the thalamus form the fibers of the **optic radiations**, which project to the **visual cortex** in the occipital lobe. Neurons of the visual cortex integrate the messages coming from the retina into a single message, translate that message into a mental image, and then transfer the image to other parts of the brain, where it is evaluated and either ignored or acted on.

The image seen by each eye is the **visual field** of that eye. The visual field of each eye can be divided into temporal (lateral) and nasal (medial) parts. The temporal part of a visual field projects onto the nasal retina, which projects to the visual cortex on the opposite side of the brain. The nasal part of a visual field projects onto the temporal retina, which projects to the same side of the brain. The nerve pathways are arranged in such a way that images entering the eye from the right part

of each visual field (right temporal and left nasal) project to the left side of the brain. Conversely, the left part of each visual field (left temporal and right nasal) projects to the right side of the brain.

The visual pathway

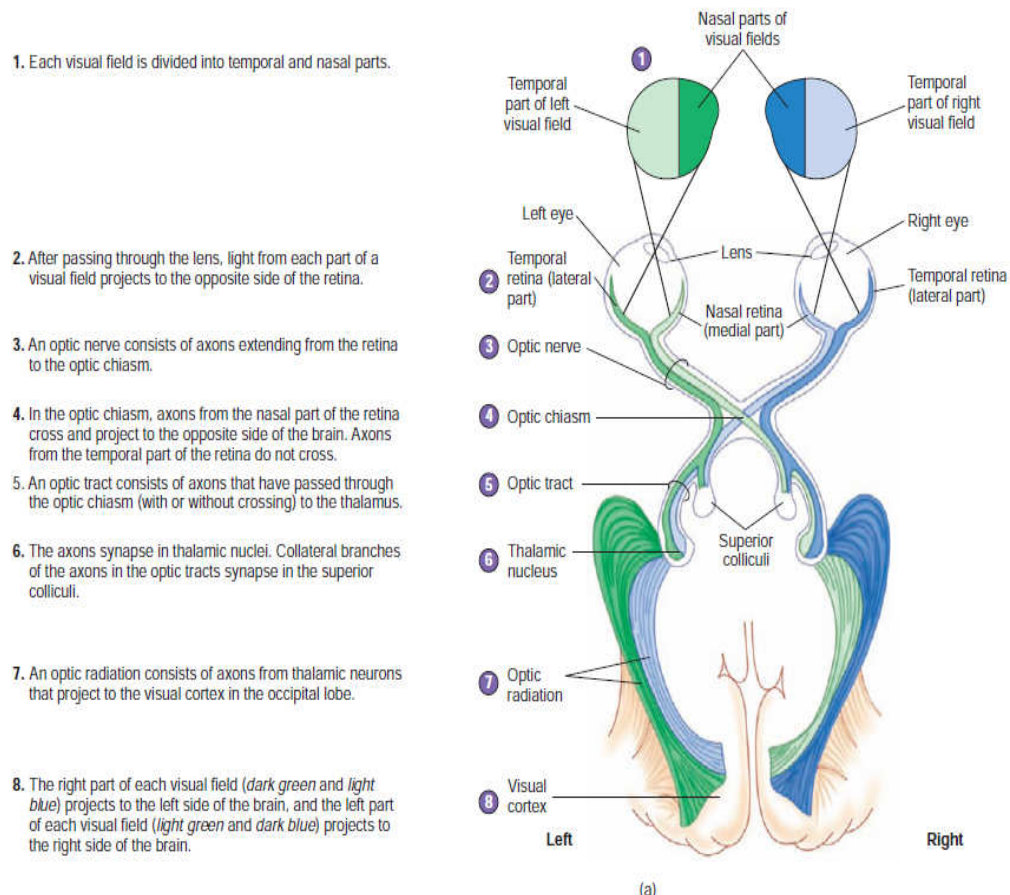


Fig.4.5: The Visual Pathway

3.4 Clinical Correlates

1. Myopia

Myopia or **short - sightedness**, is the ability to see close objects clearly while distant objects appear blurry. Myopia is a defect of the eye in which the focusing system, the cornea and lens, is optically too powerful, or the eyeball is too long. As a result, images are focused in front of the retina. Myopia is corrected by a concave lens that spreads out the light rays coming to the eye so that, when the light is focused by the eye, it is focused on the retina. Such lenses are called “minus” lenses.

2. **Hypermetropia**

Hypermetropia, or long – sightedness is the ability to see distant objects clearly while close objects appear blurry. Hypermetropia is a disorder in which the cornea and lens system is optically too weak or the eyeball is too short. As a result, the focused image is “behind” the retina when looking at a close object. In hypermetropia, the lens must accommodate to bring somewhat distant objects into focus, which would not be necessary for a normal eye. Closer objects cannot be brought into focus because the lens cannot change shape enough to focus the image on the retina. Hypermetropia is corrected by a convex lens that causes light rays to converge as they approach the eye and to focus on the retina. Such lenses are called “plus” lenses.

3. **Presbyopia**

Presbyopia is the normal, presently unavoidable degeneration of the accommodation power of the eye that occurs as a consequence of aging. It occurs because the lens becomes hard and less flexible. The average age for onset of presbyopia is the mid-forties. Avid readers and people engaged in fine, close work may develop the symptoms earlier. Presbyopia can be corrected by the use of “reading glasses” which are worn only for close work and are removed when the person wants to see at a distance. Alternatively, the problem may be corrected by the use of **bifocals**, which have a different lens in the top and the bottom, or by **progressive lenses**, in which the lens is graded.

4. **Astigmatism**

This is a defect in which the cornea or lens is not uniformly curved and the image is not sharply focused. Glasses may be made to adjust for the abnormal curvature as long as the curvature is not too irregular. If the curvature of the cornea or lens is too irregular, the condition is difficult to correct.

5. **Night Blindness**

Everyone sees less clearly in the dark than in the light. A person with night blindness, or nyctalopia, however, may not see well enough in a dimly lit environment to function adequately. Night blindness results from loss of rod function. It can result from general retinal degeneration, such as occurs in retinitis pigmentosa or detached retinas. Temporary night blindness can result from a vitamin A deficiency because vitamin A is necessary to produce retinal.

Patients with night blindness can be helped with electronic optical devices, including monocular pocket scopes and binocular goggles that electronically amplify light.

6. Glaucoma

Glaucoma is a condition involving excessive pressure build-up in the aqueous humour. Glaucoma results from an interference with normal reentry of aqueous humour into the blood or from an overproduction of aqueous humour. The increased pressure within the eye can close off the blood vessels entering the eye and may destroy the retina or optic nerve, resulting in blindness. Everyone older than 40 years should be checked every two to three years for glaucoma; those older than 40 who have relatives with glaucoma should have an annual check up. Glaucoma is usually treated with eye drops, which do not cure the problem but keep it from advancing. In some cases, laser or conventional surgery may be used.

4.0 CONCLUSION

You learnt that the visual system includes the eyes, the accessory structures, and the sensory neurons that project to the cerebral cortex where action potentials conveying visual information are interpreted.

5.0 SUMMARY

- The fibrous layer is the outer layer of the eyeball. It consists of the sclera and cornea. The sclera is the posterior four-fifths of the eyeball. It is white connective tissue that maintains the shape of the eyeball and provides a site for muscle attachment. The cornea is the anterior one-fifth of the eye. It is transparent and refracts light that enters the eye.
- The vascular layer is the middle layer of the eyeball. The black choroid prevents the reflection of light inside the eye. The iris is smooth muscle regulated by the autonomic nervous system. It controls the amount of light entering the pupil. The ciliary muscles control the shape of the lens. The ciliary process produces aqueous humour.
- The retina is the inner layer of the eyeball and contains neurons sensitive to light. The macula (fovea centralis) is the area of greatest sensitivity to light. The optic disc is the location through which nerves exit and blood vessels enter the eye. It has no photosensory cells and is therefore the blind spot of the eye.
- The eyeball has three chambers: anterior, posterior, and vitreous. The anterior and posterior chambers are filled with aqueous humour, which circulates and leaves by way of the scleral venous sinus. The vitreous chamber is filled with vitreous humour.
- Accessory structures of the eye are the eyebrows, the eyelids, the conjunctiva and lacrimal glands.

- Visual pathway comprises ganglion cell axons from the optic nerve, optic chiasm, and optic tracts. They extend to the thalamus, where they synapse. From there, neurons form the optic radiations that project to the visual cortex.

SELF-ASSESSMENT EXERCISE

In the anatomy laboratory, examine the eye model and identify the accessory structures of the eye.

6.0 TUTOR-MARKED ASSIGNMENT

1. Tears
 - a. are released onto the surface of the eye near the medial corner of the eye.
 - b. in excess are removed by the scleral venous sinus.
 - c. in excess can cause a sty.
 - d. can pass through the nasolacrimal duct into the oral cavity.
 - e. contain water, salts, mucus, and lysozyme.
2. The fibrous layer of the eye includes the
 - a. conjunctiva.
 - b. sclera.
 - c. choroid.
 - d. iris.
 - e. retina.
3. Concerning axons in the optic nerve from the right eye,
 - a. they all go to the right occipital lobe.
 - b. they all go to the left occipital lobe.
 - c. they all go to the thalamus.
 - d. they all go to the superior colliculus.
 - e. some go to the right occipital lobe and some go to the left occipital lobe.
4. Contraction of the smooth muscle in the ciliary body causes the
 - a. lens to flatten.
 - b. lens to become more spherical.
 - c. pupil to constrict.
 - d. pupil to dilate.
5. Given these events:
 1. medial rectus contracts
 2. lateral rectus contracts
 3. pupils dilate
 4. pupils constrict
 5. lens of the eye flattens
6. lens of the eye becomes more spherical

Assume you are looking at an object 30 feet away. If you suddenly look at an object that is one (1) foot away, which events occur?

- a. 1,3,6
 - b. 1,4,5
 - c. 1,4,6
 - d. 2,3,6
 - e. 2,4,5
- 6. Why do you get a “runny nose” when you cry?
 - 7. What are the layers of the retina?
 - 8. Which are more numerous, rods or cones?

7.0 REFERENCES/FURTHER READING

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MODULE 2 THE ENDOCRINE SYSTEM

Unit 1	Hormones
Unit 2	Pituitary Gland and Hypothalamus
Unit 3	Thyroid and Parathyroid Glands
Unit 4	Adrenal Glands
Unit 5	Pancreas
Unit 6	Other Endocrine Glands

UNIT 1 HORMONES

CONTENTS

1.0	Introduction
2.0	Objectives
3.0	Main Content
3.1	Functions of the endocrine system
3.2	Transport of hormones in the blood
3.3	Interaction of hormones with their target tissues
3.4	Clinical correlates
4.0	Conclusion
5.0	Summary
6.0	Tutor-Marked Assessment
7.0	References/Further Reading

1.0 INTRODUCTION

The **endocrine system** is composed of **endocrine glands**, which are ductless glands secreting chemical messengers into the circulatory system. In contrast, exocrine glands have ducts that carry their secretions to surfaces. The term *endocrine* is derived from the Greek words *endo*, meaning within, and *krinō*, to separate. The term implies that cells of endocrine glands produce chemical messengers within the glands that influence tissues separated from the glands by some distance. The chemical messengers secreted by endocrine glands are called hormones, a term derived from the Greek word *hormon*, meaning to set into motion. Thus, hormones stimulate responses from cells.

2.0 OBJECTIVES

At the end of this unit, you should be able to:

- describe the functions of the endocrine system
- define the terms endocrine gland and hormone
- explain how the regulation of hormone secretion is achieved

- describe the means by which hormones are transported and excreted.

3.1 Functions of the Endocrine System

The main regulatory functions of the endocrine system are the following:

1. *Metabolism and tissue maturation.* The endocrine system regulates the rate of metabolism and influences the maturation of tissues, such as those of the nervous system.
2. *Ion regulation.* The endocrine system helps regulate blood pH, as well as Na⁺, K⁺, and Ca²⁺ concentrations in the blood.
3. *Water balance.* The endocrine system regulates water balance by controlling the solute concentration of the blood.
4. *Immune system regulation.* The endocrine system helps control the production of immune cells.
5. *Heart rate and blood pressure regulation.* The endocrine system helps regulate the heart rate and blood pressure and helps prepare the body for physical activity.
6. *Control of blood glucose and other nutrients.* The endocrine system regulates blood glucose levels and other nutrient levels in the blood.
7. *Control of reproductive functions.* The endocrine system controls the development and functions of the reproductive systems in males and females.
8. *Uterine contractions and milk release.* The endocrine system regulates uterine contractions during delivery and stimulates milk release from the breasts in lactating females.

3.2 Transport of Hormones in the Blood

Hormones can be defined as chemicals secreted by a cell that affect the functions of other cells. Once released, most hormones enter the bloodstream where they are carried to their target cells. The target cells of a hormone are the cells that contain the receptors for the hormone. A hormone cannot affect a cell unless the cell has receptors for it. Many hormones in the body are derived from steroids. Steroids are soluble in lipids and can therefore cross cell membranes very easily. Once a **steroid hormone** is inside a cell, it binds to its receptor, which is commonly in the nucleus of the cell. The hormone-receptor complex turns a gene on or off. When new genes are turned on or off, the cell begins to carry out new functions, and this is ultimately how steroid hormones affect their target cells. Examples of steroid hormones are **estrogen, progesterone, testosterone, and cortisol.**

Non -steroid hormones are those that are made of amino acids or proteins. Proteins cannot cross the cell membrane easily. Therefore, these hormones bind to receptors on the surface of the cell. The hormone-receptor complex in the membrane usually activates a **G-protein**. The G-protein causes enzymes inside the cell to be turned on. Different chemical reactions then begin inside the cell.

The cell now takes on new functions.

Prostaglandins are local hormones. They are derived from lipid molecules and typically do not travel in the bloodstream to find their target cells. Instead, their target cells are located close by. They have the same effects as other hormones and are produced by many body organs, including the kidneys, stomach, uterus, heart, and brain.

Transport of Hormones in the Blood

Water-soluble hormones (peptides and catecholamines) are dissolved in the plasma and transported from their sites of synthesis to target tissues, where they diffuse out of the capillaries, into the interstitial fluid, and ultimately to target cells.

Steroid and thyroid hormones, in contrast, circulate in the blood mainly bound to plasma proteins. Usually less than 10 percent of steroid or thyroid hormones in the plasma exist free in solution. For example, more than 99 percent of the thyroxine in the blood is bound to plasma proteins. However, protein-bound hormones cannot easily diffuse across the capillaries and gain access to their target cells and are therefore biologically inactive until they dissociate from plasma proteins. The relatively large amounts of hormones bound to proteins serve as reservoirs, replenishing the concentration of free hormones when they are bound to target receptors or lost from the circulation. Binding of hormones to plasma proteins greatly slows their clearance from the plasma.

3.3 Interaction of Hormones with their Target Tissues

Hormones bind to proteins or glycoproteins called receptors. The portion of each protein or glycoprotein molecule where a hormone binds is called a receptor site, or binding site. The shape and chemical characteristics of each receptor site allow only a specific type of chemical messenger to bind to it. The tendency for each type of chemical messenger to bind to a specific type of receptor, and not to others, is called specificity. Insulin therefore binds to insulin receptors but not to receptors for growth hormone. Some hormones, however, can bind to a number of different receptors that are closely related. For example, epinephrine can bind to more than one type of epinephrine

receptor. Hormone receptors have a high affinity for the hormones that bind to them, so only a small concentration of a given hormone results in a significant number of receptors with hormones bound to them.

Hormones are secreted and distributed throughout the body by the circulatory system, but the presence or absence of specific receptor molecules in cells determines which cells will or will not respond to each hormone. For example, there are receptors for thyroid stimulating hormone (TSH) in cells of the thyroid gland, but there are no such receptors in most other cells of the body. Consequently, cells of the thyroid gland produce a response when exposed to TSH, but cells without receptor molecules do not respond to it.

In general, the number of functional receptors affects the amplitude of a cell's response to a hormone. More receptors produce a larger response than fewer receptors. The number of functional receptors can be regulated. In down-regulation, the number of functional receptors is reduced by temporary or permanent removal of receptors from the plasma membrane, inactivation of receptors, or decreased synthesis of replacement receptors. In up-regulation, the number of functional receptors is increased through increased receptor synthesis or availability.

Drugs with structures similar to specific hormones may compete with those hormones for their receptors. A drug that binds to a hormone receptor and activates it is called an agonist for that hormone. A drug that binds to a hormone receptor and inhibits its action is called an antagonist for that hormone. For example, drugs exist that compete with epinephrine for its receptor. Epinephrine agonists activate epinephrine receptors, whereas epinephrine antagonists inhibit them.

3.4 Clinical Correlates

Lipid- and Water-Soluble Hormones in Medicine

Specific hormones are given as treatments for certain illnesses. Hormones that are soluble in lipids, such as steroids, can be taken orally because they can diffuse across the wall of the stomach and intestine into the circulatory system. Examples include the synthetic estrogen and progesterone-like hormones in birth control pills and steroids that reduce the severity of inflammation, such as prednisone. In contrast to lipid-soluble hormones, protein hormones cannot diffuse across the wall of the intestine because they are not lipid-soluble. Furthermore, protein hormones are not transported across the wall of the intestine because they are broken down to individual amino acids by the digestive system. The normal structure of a protein hormone is therefore destroyed, and its physiological activity is lost. Consequently, protein hormones must be

injected rather than taken orally. The most commonly administered protein hormone is insulin, which is prescribed for the treatment of diabetes mellitus.

4.0 SUMMARY

- The main regulatory functions include water balance, uterine contractions and milk release, metabolism and tissue maturation, ion regulation, heart rate and blood pressure regulation, control of blood glucose and other nutrients, immune system regulation, and control of reproductive functions
- Endocrine glands produce hormones that are released into the interstitial fluid and diffuse into the blood. Hormones act on target tissues, producing specific responses. The protein group of hormones includes hormones that are proteins, glycoproteins, polypeptides, and amino acid derivatives. The lipid group of hormones includes hormones that are steroids and fatty acid derivatives.
- Generalizations about the differences between the endocrine and nervous systems include the following: (a) The endocrine system is amplitude-modulated, whereas the nervous system is frequency modulated, and (b) the response of target tissues to hormones is usually slower and of longer duration than their response to neurons.
- Water-soluble hormones, such as proteins, epinephrine, and norepinephrine, are rapidly removed from the blood. These hormones regulate activities that have a rapid onset and a short duration. Lipid-soluble hormones and thyroid hormones are not quickly removed from the blood. They produce a prolonged effect.

SELF – ASSESSMENT EXERCISE

1. When comparing the endocrine system and the nervous system, generally speaking, the endocrine system
 - a. is faster-acting than the nervous system.
 - b. produces effects that are of shorter duration.
 - c. uses amplitude-modulated signals.
 - d. produces more localized effects.
 - e. relies less on chemical messengers.
2. Given this list of molecule types:
 1. nucleic acid derivatives
 2. fatty acid derivatives
 3. polypeptides
 4. proteins

Which could be hormone molecules?

- a. 1,2,3
 - b. 2,3,4
 - c. 1,2,3,4
 - d. 2,3,4,5
 - e. 1,2,3,4,5
3. Which of these regulates the secretion of a hormone from an endocrine tissue?
- a. other hormones
 - b. negative-feedback mechanisms
 - c. nonhormone substance in the blood
 - d. the nervous system
 - e. all of the above
4. Hormones are released into the blood
- a. at relatively constant levels.
 - b. in large amounts in response to a stimulus.
 - c. in a cyclic fashion.
 - d. all of the above.
5. Given these observations:
1. A hormone affects only a specific tissue (not all tissues).
 2. A tissue can respond to more than one hormone.
 3. Some tissues respond rapidly to a hormone, whereas others take many hours to respond.
6. Which of these observations can be explained by the characteristics of hormone receptors?
- a. 1
 - b. 1,2
 - c. 2,3
 - d. 1,3
 - e. 1,2,3
8. A hormone
- a. can function as an enzyme.
 - b. is also a G protein.
 - c. can bind to a receptor.
 - d. is an intracellular mediator.
 - e. all of the above.
7. Given these events:
1. The α subunit of a G protein interacts with Ca^{2+} channels.
 2. Calcium ions diffuse into the cell.
 3. The α subunit of a G protein is activated.

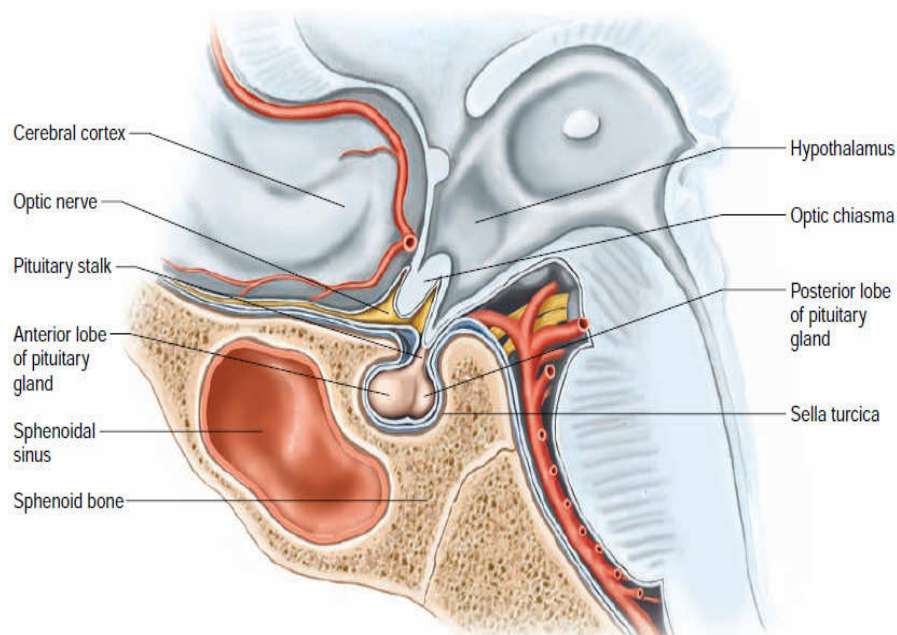
UNIT 2 THE PITUITARY GLAND AND HYPOTHALAMUS

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Structure of the pituitary gland
- 4.0 Summary

1.0 INTRODUCTION

The pituitary gland is located at the base of the brain and is controlled by the hypothalamus. This gland is well protected by a bony structure called the **sella turcica**. Just superior to the gland is the **optic chiasm**, which carries visual information to the brain for interpretation. The pituitary is divided into two lobes—the anterior and the posterior. the location of the pituitary gland



The **pituitary gland**, or **hypophysis**, secretes nine major hormones that regulate numerous body functions and the secretory activity of several other endocrine glands. The **hypothalamus** of the brain and the pituitary gland are major sites where the nervous and endocrine systems interact. The hypothalamus regulates the secretory activity of the pituitary gland. Hormones, sensory information that enters the central nervous system, and emotions, in turn, influence the activity of the hypothalamus.

2.0 OBJECTIVES

At the end of this unit, you should be able to:

- describe the structure of the pituitary gland
- discuss the hormones of the pituitary gland

3.1 Structure of the pituitary gland

The pituitary gland is roughly 1 cm in diameter, weighs 0.5–1.0 g, and rests in the sella turcica of the sphenoid bone. It is located inferior to the hypothalamus and is connected to it by a stalk of tissue called the **infundibulum**. The **posterior pituitary**, or **neurohypophysis** is continuous with the brain. It is formed during embryonic development from an outgrowth of the inferior part of the brain in the area of the hypothalamus. The outgrowth forms the infundibulum, and the distal end of the infundibulum enlarges to form the posterior pituitary.

Relationship of the pituitary gland to the brain

Portal vessels are blood vessels that begin in a primary capillary network, extend some distance, and end in a secondary capillary network. The **hypothalamohypophyseal portal system** is one of two major portal systems. The other is the hepatic portal system. The hypothalamohypophyseal portal system extends from the hypothalamus to the anterior pituitary. The primary capillary network in the hypothalamus is supplied with blood from arteries that deliver blood to the hypothalamus. From the primary capillary network, the hypothalamohypophyseal portal vessels carry blood to a secondary capillary network in the anterior pituitary. Veins from the secondary capillary network eventually merge with the general circulation. Hormones, produced and secreted by neurons of the hypothalamus, enter the primary capillary network and are carried to the secondary capillary network. There the hormones leave the blood and act on cells of the anterior pituitary. They act either as **releasing hormones**, increasing the secretion of anterior pituitary hormones, or as **inhibiting hormones**, decreasing the secretion of anterior pituitary hormones. Each releasing hormone stimulates and each inhibiting hormone inhibits the production and secretion of a specific hormone by the anterior pituitary. In response to the releasing hormones, anterior pituitary cells secrete hormones that enter the secondary capillary network and are carried by the general circulation to their target tissues. Thus, the hypothalamohypophyseal portal system provides a means by which the hypothalamus, using hormones as chemical messengers, regulates the secretory activity of the anterior pituitary..

There is no portal system to carry hypothalamic hormones to the posterior pituitary. Hormones released from the posterior pituitary are produced by neurosecretory cells with their cell bodies located in the hypothalamus. The axons of these cells extend from the hypothalamus through the infundibulum into the posterior pituitary and form a nerve tract called the **hypothalamohypophyseal tract**. Hormones produced in the hypothalamus pass down these axons in tiny vesicles and are stored in secretory vesicles in the enlarged ends of the axons. Action potentials originating in the neuron cell bodies in the hypothalamus are propagated along the axons to the axon terminals in the posterior pituitary. The action potentials cause the release of hormones from the axon terminals, and they enter the circulatory system.

Hormones of the pituitary gland

Posterior Pituitary Hormones

The posterior pituitary stores and secretes two polypeptide hormones called antidiuretic hormone and oxytocin. A separate population of cells secretes each hormone. **Antidiuretic hormone (ADH)** is so named because it prevents the output of large amounts of urine (*diuresis*). ADH binds to membrane-bound receptors and increases water reabsorption by kidney tubules. This results in less water loss from the blood into the urine, and urine volume decreases. ADH can also cause blood vessels to constrict when released in large amounts. Consequently, it is sometimes called **vasopressin**. **Oxytocin** binds to membrane – bound receptors and causes contraction of the smooth muscle cells of the uterus and milk ejection, or milk “let-down” from the breasts in lactating women. Oxytocin plays an important role in the expulsion of the fetus from the uterus during delivery by stimulating uterine smooth muscle contraction. Commercial preparations of oxytocin are given under certain conditions to assist in childbirth and to constrict uterine blood vessels following childbirth. Oxytocin also causes the contraction of uterine smooth muscle in non - pregnant women during menses, which helps expel the uterine epithelium and a small amount of blood. Oxytocin also promotes the movement of sperm cells through the uterus and uterine tubes. Oxytocin has been called the great facilitator of life. In addition to its role in reproduction and lactation, oxytocin produced in the limbic system and other parts of the brain influences a variety of social and non - social behaviours in females and males. In many species, oxytocin promotes pair bonding, sexual behaviour, and parental care. In humans, oxytocin promotes social interactions, feelings of attachment, and maternal behaviour. Oxytocin also inhibits memory, decreases the stress response, reduces feelings of anxiety, suppresses appetite, and raises the pain threshold.

Anterior Pituitary Hormones

Releasing and inhibiting hormones that pass from the hypothalamus through the hypothalamohypophyseal portal system to the anterior pituitary influence anterior pituitary secretions. The hormones secreted are proteins, glycoproteins, or polypeptides. They are transported in the circulatory system and bind to membrane-bound receptor molecules on their target cells. For the most part, each hormone is secreted by a separate cell type. The major hormones of the anterior pituitary, their target tissues, and their effects on target tissues are listed in below. Anterior pituitary hormones include growth hormone, thyroid stimulating hormone, adrenocorticotrophic hormone and related substances, luteinizing hormone, follicle-stimulating hormone, and prolactin.

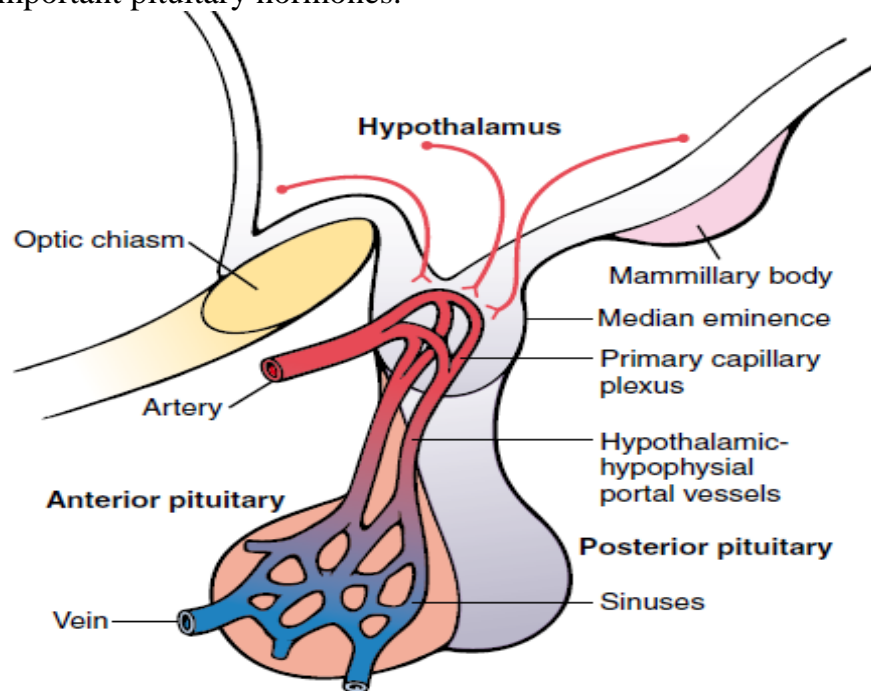
Hormones of the pituitary gland

Hormones	Structure	Target Tissue	Response
Posterior Pituitary			
Antidiuretic hormone (ADH)	Small peptide	Kidney	Increased water reabsorption (less water is lost in the form of urine)
Oxytocin	Small peptide	Uterus; mammary glands	Increased uterine contractions; increased milk expulsion from mammary glands; unclear function in males
Anterior Pituitary			
Growth hormone (GH), or somatotropin	Protein	Most tissues	Increased growth in tissues; increased amino acid uptake and protein synthesis; increased breakdown of lipids and release of fatty acids from cells; increased glycogen synthesis and increased blood glucose levels; increased somatomedin production
Thyroid-stimulating hormone (TSH)	Glycoprotein	Thyroid gland	Increased thyroid hormone secretion
Adrenocorticotrophic hormone (ACTH)	Peptide	Adrenal cortex	Increased glucocorticoid hormone secretion
Melanocyte-stimulating hormone (MSH)	Peptide	Melanocytes in the skin	Increased melanin production in melanocytes to make the skin darker in color
Luteinizing hormone (LH)	Glycoprotein	Ovaries in females; testes in males	Ovulation and progesterone production in ovaries; testosterone synthesis and support for sperm cell production in testes
Follicle-stimulating hormone (FSH)	Glycoprotein	Follicles in ovaries in females; seminiferous tubes in males	Follicle maturation and estrogen secretion in ovaries; sperm cell production in testes
Prolactin	Protein	Ovaries and mammary glands in females	Milk production in lactating women; increased response of follicle to LH and FSH; unclear function in males

Mechanism by which hypothalamus controls pituitary secretion

Almost all secretion by the pituitary is controlled by either hormonal or nervous signals from the hypothalamus. Indeed, when the pituitary gland is removed from its normal position beneath the hypothalamus and transplanted to some other part of the body, its rates of secretion of the different hormones (except for prolactin) fall to very low levels. Secretion from the posterior pituitary is controlled by nerve signals that originate in the hypothalamus and terminate in the posterior pituitary. In contrast, secretion by the anterior pituitary is controlled by hormones called *hypothalamic releasing* and *hypothalamic inhibitory hormones* (or *factors*) secreted within the hypothalamus and then conducted to the

anterior pituitary through minute blood vessels called *hypothalamic-hypophyseal portal vessels*. In the anterior pituitary, these releasing and inhibitory hormones act on the glandular cells to control their secretion. The hypothalamus receives signals from many sources in the nervous system. Thus, when a person is exposed to pain, a portion of the pain signal is transmitted into the hypothalamus. Likewise, when a person experiences some powerful depressing or exciting thought, a portion of the signal is transmitted into the hypothalamus. Olfactory stimuli denoting pleasant or unpleasant smells transmit strong signal components directly and through the amygdaloid nuclei into the hypothalamus. Even the concentrations of nutrients, electrolytes, water, and various hormones in the blood excite or inhibit various portions of the hypothalamus. Thus, the hypothalamus is a collecting center for information concerning the internal well-being of the body, and much of this information is used to control secretions of the many globally important pituitary hormones.



Hypothalamic- hypophyseal system

Growth Hormone

Growth hormone (GH) stimulates the growth of most tissues and, through its effect on the epiphyseal plates of bones, GH plays a role in determining how tall a person becomes. GH promotes the protein synthesis necessary for growth by increasing the movement of amino acids into cells and promoting their incorporation into proteins. It also decreases the breakdown of proteins.

GH plays an important role in regulating blood nutrient levels between meals and during periods of fasting. GH increases lipolysis, the breakdown of lipids. Fatty acids released from fat cells into the blood circulate to other tissues and are used as an energy source. The use of fatty acids as an energy source “spares” the use of blood glucose, helping maintain blood sugar levels. In addition, GH increases glucose synthesis by the liver, which releases glucose into the blood. Thus, through its effects on adipose tissue and the liver,

GH maintains or increases blood sugar levels. GH has indirect effects on some tissues by stimulating the production of polypeptides called **somatomedins**, primarily by the liver but also by skeletal muscle and other tissues. Somatomedins circulate in the blood, stimulating growth in cartilage and bone and increasing the synthesis of protein in skeletal muscles. The best known somatomedins are two polypeptide hormones produced by the liver called **insulin-like growth factor I and II** because of the similarity of their structure to insulin. Two hormones released from the hypothalamus regulate the secretion of GH. Growth hormone–releasing hormone (GHRH) stimulates the secretion of GH, whereas growth hormone–inhibiting hormone (GHIH) inhibits the secretion of GH. Stimuli that influence GH secretion act on the hypothalamus to increase or decrease the secretion of the releasing and inhibiting hormones. Low blood glucose levels and stress stimulate the secretion of GH, and high blood glucose levels inhibit the secretion of GH. An increase in certain amino acids stimulates increased GH secretion. Most people have a rhythm of growth hormone secretion, with daily peak levels occurring during deep sleep. Growth hormone secretion also increases during periods of fasting and prolonged exercise.

Blood growth hormone levels do not become greatly elevated during periods of rapid growth, although children tend to have somewhat higher blood levels of growth hormone than do adults. In addition to growth hormone, genetics, nutrition, and sex hormones influence growth.

Clinical correlates

1. **Gigantism** is a condition of abnormally increased height that usually results from excessive cartilage and bone formation at the epiphyseal plates of long bones. The most common type of gigantism, **pituitary gigantism**, results from excess secretion of GH. The large stature of some individuals, however, can result from genetic factors rather than from abnormal levels of GH.
2. **Acromegaly** is caused by excess GH secretion in adults, and many pituitary giants develop acromegaly later in life. The GH stimulates the growth of connective tissue, including bones.

Bones in adults can increase in diameter and thickness, but not in length because the epiphyseal plates have ossified. The effects of acromegaly are most apparent in the face and hands. Hypersecretion of GH can also cause elevated blood glucose levels and may eventually lead to diabetes mellitus.

3. **Dwarfism**, the condition in which a person is abnormally short, is the opposite of gigantism. Pituitary dwarfism results when abnormally low levels of GH affect the whole body, thus producing a small person who is normally proportioned. **Achondroplasia, or achondroplastic dwarfism**, is the most common type of dwarfism; it produces a person with a nearly normal-sized trunk and head but shorter-than-normal limbs. Achondroplasia is a genetic disorder, not a hormonal disorder. Modern genetic engineering has provided a source of human GH for people who produce inadequate quantities. Human genes for GH have been successfully introduced into bacteria using genetic engineering techniques. The gene in the bacteria causes GH synthesis, and the GH can be extracted from the medium in which the bacteria are grown.

4.0 SUMMARY

- The pituitary gland secretes at least nine hormones that regulate numerous body functions and other endocrine glands. The hypothalamus regulates pituitary gland activity through hormones and action potentials.
- The posterior pituitary develops from the floor of the brain and connects to the hypothalamus by the infundibulum. The anterior pituitary develops from the roof of the mouth.
- The hypothalamohypophyseal portal system connects the hypothalamus and the anterior pituitary. Through the portal system, the hormones inhibit or stimulate hormone production in the anterior pituitary. The hypothalamohypophyseal tract connects the hypothalamus and the posterior pituitary. Hormones are produced in hypothalamic neurons. The hormones move down the axons of the tract and are secreted from the posterior pituitary.
- Antidiuretic hormone (ADH) promotes water retention by the kidneys. Oxytocin promotes uterine contractions during delivery and causes milk ejection in lactating women. Growth hormone (GH) stimulates growth in most tissues and is a regulator of metabolism. GH stimulates the uptake of amino acids and their conversion into proteins and stimulates the breakdown of fats and the synthesis of glucose.

Activity

Examine the histological structure of the pituitary gland and distinguish between the anterior and posterior pituitary glands.

SELF – ASSESSMENT EXERCISE

1. The pituitary gland
 - a. develops from the floor of the brain.
 - b. develops from the roof of the mouth.
 - c. is stimulated by hormones produced in the midbrain.
 - d. secretes only three major hormones.
 - e. both a and b.
2. The hypothalamohypophyseal portal system
 - a. contains one capillary bed.
 - b. carries hormones from the anterior pituitary to the body.
 - c. carries hormones from the posterior pituitary to the body.
 - d. carries hormones from the hypothalamus to the anterior pituitary.
 - e. carries hormones from the hypothalamus to the posterior pituitary.
3. Hormones secreted from the posterior pituitary
 - a. are produced in the anterior pituitary.
 - b. are transported to the posterior pituitary within axons.
 - c. include GH and TSH.
 - d. are steroids.
 - e. all of the above.
4. Oxytocin is responsible for
 - a. preventing milk release from the mammary glands.
 - b. preventing goiter.
 - c. causing contraction of the uterus.
 - d. maintaining normal calcium levels.
 - e. increasing metabolic rate.
5. Growth hormone
 - a. increases the usage of glucose.
 - b. increases the breakdown of lipids.
 - c. decreases the synthesis of proteins.
 - d. decreases the synthesis of glycogen.
 - e. all of the above.

UNIT 3 THE THYROID AND PARATHYROID GLANDS

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Structure of the thyroid gland
- 4.0 Summary

1.0 INTRODUCTION

The **thyroid gland** consists of two lobes connected by a narrow band called the **isthmus**. The lobes are located on each side of the trachea, just inferior to the larynx. The thyroid gland is one of the largest endocrine glands, with a weight of approximately 20 g. It is highly vascular and appears more red than its surrounding tissues.

2.0 OBJECTIVES

At the end of this unit, you will be able to:

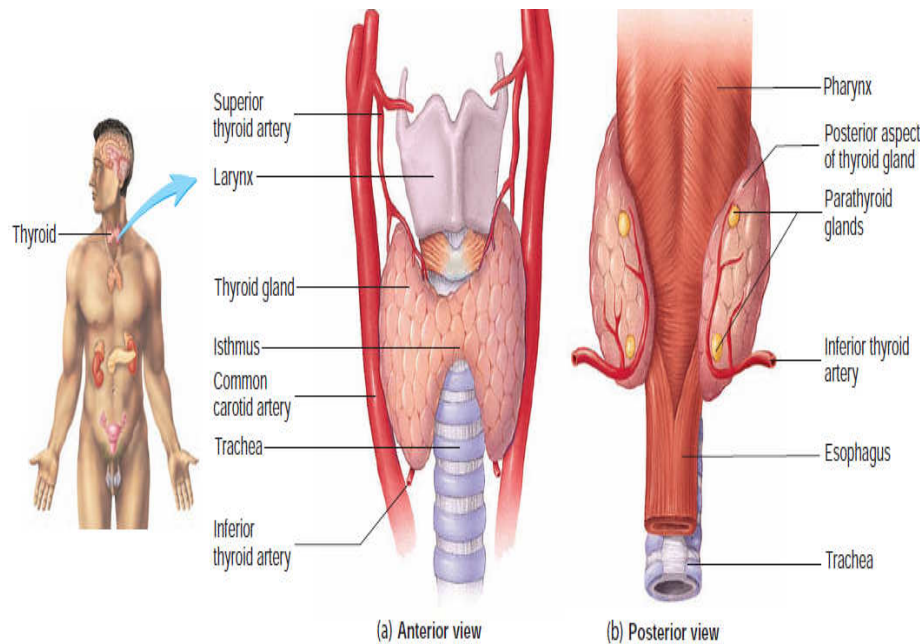
- describe the anatomy of the thyroid and parathyroid gland
- discuss the thyroid hormones and parathyroid hormones.

3.0 MAIN CONTENT

3.1 Structure of the thyroid gland

The thyroid gland contains numerous **follicles**, which are small spheres whose walls are composed of a single layer of cuboidal epithelial cells. Each thyroid follicle is filled with proteins, called **thyroglobulin** which are synthesized and secreted by the cells of the thyroid follicles. Large amounts of the thyroid hormones are stored in the thyroid follicles as part of the thyroglobulin molecules. Between the follicles, a delicate network of loose connective tissue contains scattered **parafollicular cells**, or **C cells**.

Calcitonin is secreted from the parafollicular cells and plays a role in reducing the concentration of Ca^{2+} in the body fluids when Ca^{2+} levels become elevated.



Thyroid and parathyroid glands

Thyroid hormones

The thyroid hormones are **triiodothyronine (T₃)** and **tetraiodothyronine (T₄)**. Another name for T₄ is **thyroxine**. T₃ constitutes 10% of thyroid gland secretions and T₄ 90%. Although calcitonin is secreted by the parafollicular cells of the thyroid gland, T₃ and T₄ are considered to be the thyroid hormones because they are more clinically important and because they are secreted from the thyroid follicles.

T₃ and T₄ Synthesis

Thyroid-stimulating hormone (TSH) from the anterior pituitary stimulates thyroid hormone synthesis and secretion. TSH causes an increase in the synthesis of T₃ and T₄, which are then stored inside the thyroid follicles as part of thyroglobulin. TSH also causes T₃ and T₄ to be released from thyroglobulin and to enter the circulatory system. An adequate amount of iodine in the diet is required for thyroid hormone synthesis because iodine is a component of T₃ and T₄. The following events in the thyroid follicles result in T₃ and T₄ synthesis and secretion:

1. Iodide ions (I⁻) are taken up by thyroid follicle cells by secondary active transport (symport). The movement of the I⁻ is against a concentration gradient of approximately 30-fold in healthy individuals. TSH promotes the uptake of I⁻.
2. Iodide is transported into the follicle lumen and converted into iodine (I₀).

3. Thyroglobulin molecules, which contain numerous tyrosine amino acids, are synthesized, packaged into secretory vesicles, and secreted into the lumen of the follicle.
4. Iodine atoms are bound to a few of the tyrosine amino acids of thyroglobulin, producing monoiodotyrosine, which has one iodine atom, or diiodotyrosine, which has two iodine atoms. After the tyrosines are iodinated, two diiodotyrosine combine to form tetraiodothyronine (T₄), which has four iodine atoms. Also, one monoiodotyrosine and one diiodotyrosine combine to form triiodothyronine (T₃), which has three iodine atoms. A 2–4 months reserve supply of T₃ and T₄ is stored within the thyroid follicles as part of thyroglobulin.
5. Thyroglobulin is taken into the thyroid follicle cells by endocytosis.
6. Lysosomes fuse with the endocytic vesicles, and proteolytic enzymes break down thyroglobulin to release T₃, T₄, and amino acids.
7. T₃ and T₄ are lipid soluble and diffuse through the plasma membranes of the follicle cells into the interstitial fluid and finally into the blood. The remaining amino acids of thyroglobulin are used again to synthesize more thyroglobulin.

Effects of Thyroid Hormones

Thyroid hormones interact with their target tissues in a fashion similar to that of the steroid hormones. They readily diffuse through plasma membranes into the cytoplasm of cells. Within cells, they bind to receptor molecules in the nuclei. Thyroid hormones combined with their receptor molecules interact with DNA in the nuclei to influence genes and initiate new protein synthesis. The newly synthesized proteins within the target cells mediate the cells' response to thyroid hormones. It takes up to a week after the administration of thyroid hormones for a maximal response to develop, and new protein synthesis occupies much of that time. Thyroid hormones affect nearly every tissue in the body, but not all tissues respond identically. Metabolism is primarily affected in some tissues, and growth and maturation are influenced in others.

The normal rate of metabolism for an individual depends on an adequate supply of thyroid hormone, which increases the rate at which glucose, fat, and protein are metabolized. The increased rate of metabolism produces heat. Thyroid hormones increase the activity of Na⁺–K⁺ pumps, which helps increase the body temperature as ATP molecules are broken down. Thyroid hormones also alter the number and activity of mitochondria, resulting in greater ATP synthesis and heat production. The metabolic rate can increase 60%–100% when blood thyroid hormones are elevated. Low levels of thyroid hormones lead to the

opposite effect. Maintaining normal body temperature depends on an adequate amount of thyroid hormones.

Normal growth and maturation of organs also depend on thyroid hormones. For example, bone, hair, teeth, connective tissue, and nervous tissue require thyroid hormones for normal growth and development. Both normal growth and normal maturation of the brain require thyroid hormones.

Regulation of Thyroid Hormone Secretion

Thyroid hormone secretion is regulated by hormones produced in the hypothalamus and anterior pituitary. Thyrotropin – releasing hormone (TRH) is produced in the hypothalamus. Chronic exposure to cold increases TRH secretion, whereas stress, such as starvation, injury, and infections, decreases TRH secretion. TRH stimulates TSH secretion from the anterior pituitary. Small fluctuations in blood levels of TSH occur on a daily basis, with a small nocturnal increase. TSH stimulates the secretion of thyroid hormones from the thyroid gland. TSH also increases the synthesis of thyroid hormones, as well as causing an increase in thyroid gland cell size and number. Decreased blood levels of TSH lead to decreased secretion of thyroid hormones and thyroid gland atrophy. Thyroid hormones have a negative-feedback effect on the hypothalamus and anterior pituitary gland. As thyroid hormone levels increase in the circulatory system, they inhibit TRH and TSH secretion. Also, if the thyroid gland is removed or if the secretion of thyroid hormones declines, TSH levels in the blood increase dramatically.

Calcitonin

In addition to secreting thyroid hormones, the thyroid gland secretes a hormone called **calcitonin** produced by the parafollicular cells. Calcitonin secretion is directly regulated by blood Ca^{2+} levels. As blood Ca^{2+} concentration increases, calcitonin secretion increases, and, as blood Ca^{2+} concentration decreases, calcitonin secretion decreases. Calcitonin binds to membrane-bound receptors of osteoclasts and inhibits them, which reduces the rate of bone matrix breakdown and the release of Ca^{2+} from bone into the blood. Calcitonin may prevent blood Ca^{2+} levels from becoming overly elevated following a meal that contains a high concentration of Ca^{2+} .

The role of calcitonin in humans is unclear. It may be important in slowing bone turnover during periods of rapid growth. Calcitonin helps prevent elevated blood Ca^{2+} levels, but a lack of calcitonin secretion does not result in a prolonged increase in blood Ca^{2+} levels. Other mechanisms controlling blood Ca^{2+} levels, such as parathyroid

hormone and vitamin D, are able to compensate for the lack of calcitonin secretion.

Parathyroid glands

The **parathyroid glands** are usually embedded in the posterior part of each lobe of the thyroid gland. Usually, four parathyroid glands are present, with their cells organized in densely packed masses or cords rather than in follicles. The parathyroid glands secrete a polypeptide hormone called **parathyroid hormone (PTH)**, which is essential for the regulation of blood Ca^{2+} levels. PTH is much more important than calcitonin in regulating blood Ca^{2+} levels. PTH regulates blood Ca^{2+} levels by affecting Ca^{2+} release from bones, Ca^{2+} excretion by the kidneys, and vitamin D formation by the kidneys, which promotes Ca^{2+} absorption by the small intestine. PTH increases the release of Ca^{2+} from bones into blood by increasing the number of osteoclasts in bone, which results in increased bone breakdown. PTH promotes an increase in osteoclast numbers by stimulating stem cells in red bone marrow to differentiate and become osteoclasts. The effect of PTH on osteoclast formation, however, is indirect. PTH binds to its receptors on osteoblasts, stimulating them. The osteoblasts, through surface molecules and released chemicals, stimulate osteoclast stem cells to become osteoclasts. In the kidneys, PTH increases Ca^{2+} reabsorption from the urine into the blood so that less calcium leaves the body in urine. PTH also increases the formation of active vitamin D in the kidneys. The vitamin D is carried by the blood to epithelial cells of the small intestine, where it promotes the synthesis of Ca^{2+} transport proteins. PTH increases blood Ca^{2+} levels by increasing the rate of active vitamin D formation, which in turn increases the rate of Ca^{2+} absorption in the intestine. PTH secretion is directly regulated by blood Ca^{2+} levels. As blood Ca^{2+} concentration increases, PTH secretion decreases; as blood Ca^{2+} concentration decreases, PTH secretion increases. This regulation keeps blood Ca^{2+} levels fluctuating within a normal range of values.

Clinical correlates

1. **Hypothyroidism** is reduced or no secretion of thyroid hormones. It can be caused by inadequate TSH stimulation of the thyroid gland, an inability of the thyroid gland to produce thyroid hormones, or the surgical removal or destruction of the thyroid gland for various reasons. Damage to the pituitary gland can result in decreased TSH secretion. Tumors and inadequate blood delivery to the pituitary because of blood loss during childbirth are causes of pituitary insufficiency. Lack of iodine in the diet can result in decreased thyroid hormone levels because iodine is necessary for the synthesis of thyroid hormones. Damage to the

thyroid gland by drugs, chemicals, or an autoimmune disease (Hashimoto disease) can also reduce thyroid hormone production.

Hyposecretion of thyroid hormones decreases the rate of metabolism. Low body temperature, weight gain, reduced appetite, reduced heart rate, reduced blood pressure, decreased muscle tone, constipation, drowsiness, and apathy are major symptoms.

2. **Hyperthyroidism** is an abnormally increased secretion of thyroid hormones. After diabetes mellitus, the most common endocrine disorder is a type of hyperthyroidism called **Graves disease**. It is an autoimmune disorder that produces a specific immunoglobulin, called **thyroid-stimulating immunoglobulin (TSI)**.
3. **Goiter** is a chronic enlargement of the thyroid gland not due to a tumor. Goiter eventually develops with chronic hypersecretion of thyroid hormones. TSI in Graves disease or elevated TSH produced by pituitary tumors results in continual overstimulation of the thyroid gland. Thyroid hormone synthesis increases and thyroid gland cells increase in size and number, producing goiter.

Hypothyroidism caused by an iodine deficiency in the diet can also cause goiter.

Without adequate iodine to synthesize thyroid hormones, blood levels of thyroid hormones decrease. The reduced negative feedback of thyroid hormones on the anterior pituitary and hypothalamus results in elevated TSH secretion. TSH causes increased thyroid gland cell size and number and increased thyroglobulin synthesis, even though there is not enough iodine to synthesize thyroid hormones. Historically, goiters were common in people from areas where the soil was depleted of iodine. Consequently, plants grown in these areas, called goiter belts, had little iodine in them and caused iodine-deficient diets. Iodized salt has nearly eliminated iodine-deficiency goiters. However, it remains a problem in some developing countries

4.0 SUMMARY

- The thyroid gland is just inferior to the larynx. The thyroid gland is composed of small, hollow balls of cells called follicles, which contain thyroglobulin. Parafollicular cells are scattered throughout the thyroid gland.
- Thyroid hormone (T3 and T4) synthesis occurs in thyroid follicles. Iodide ions are taken into the follicles by secondary active transport (symport), transported to the follicle lumen, and converted to iodine. Thyroglobulin is secreted into the follicle

lumen. Tyrosine molecules with iodine combine to form T3 and T4 within thyroglobulin. Thyroglobulin is taken into follicle cells and is broken down; T3 and T4 diffuse from the follicles to the blood.

- Thyroid hormones are transported in the blood. Thyroid hormones bind to thyroxine-binding globulin and other plasma proteins. The plasma proteins prolong the time that thyroid hormones remain in the blood. T3 and T4 bind with nuclear receptor molecules and initiate new protein synthesis. T3 and T4 affect nearly every tissue in the body. T3 and T4 increase the rate of glucose, fat, and protein metabolism in many tissues, thus increasing body temperature. Normal growth of many tissues is dependent on T3 and T4.
- Thyrotropin-releasing hormone (TRH) and thyroid-stimulating hormone (TSH) regulate T3 and T4 secretion. TRH from the hypothalamus increases TSH secretion. TRH increases as a result of chronic exposure to cold and decreases as a result of food deprivation, injury, and infections. Increased TSH from the anterior pituitary increases T3 and T4 secretion.
- The parafollicular cells secrete calcitonin. An increase in blood calcium levels stimulates calcitonin secretion. Calcitonin decreases blood Ca^{2+} levels by inhibiting osteoclasts.
- The parathyroid glands are embedded in the thyroid gland. Parathyroid hormone (PTH) increases blood Ca^{2+} levels. PTH stimulates an increase in osteoclast numbers, resulting in increased breakdown of bone. PTH promotes Ca^{2+} reabsorption by the kidneys and the formation of active vitamin D by the kidneys. Active vitamin D increases calcium absorption by the intestine. A decrease in blood Ca^{2+} levels stimulates PTH secretion.

Activity

Examine the slides of the thyroid and parathyroid glands under the microscope in the histology laboratory

SELF – ASSESSMENT EXERCISE

1. Describe events in the thyroid follicles that result in the synthesis and secretion of thyroid hormones.
2. Describe the actions of thyroid hormones T3 and T4.
3. What are some common disorders associated with thyroid dysfunction?
4. What two cell types are found in the parathyroid glands?
5. Which of these conditions most likely occurs if a healthy person receives an injection of T3 and T4?

- a. The secretion rate of TSH declines.
 - b. The person develops symptoms of hypothyroidism.
 - c. The person develops hypercalcemia.
 - d. The person secretes more TRH.
6. Which of these occurs as a response to a thyroidectomy (removal of the thyroid gland)?
- a. increased calcitonin secretion c. decreased TRH secretion
 - b. increased T3 and T4 secretion d. increased TSH secretion
7. Calcitonin
- a. is secreted by the parathyroid glands.
 - b. levels increase when blood calcium levels decrease.
 - c. causes blood calcium levels to decrease.
 - d. insufficiency results in weak bones and tetany.
8. If parathyroid hormone levels increase, which of these conditions is expected?
- a. Osteoclast numbers are increased.
 - b. Calcium absorption from the small intestine is inhibited.
 - c. Calcium reabsorption from the urine is inhibited.
 - d. Less active vitamin D is formed in the kidneys.

UNIT 4 ADRENAL GLANDS

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Anatomy of the adrenal glands
- 4.0 Summary

1.0 INTRODUCTION

An adrenal gland sits on top of each kidney. It is divided into two portions—the adrenal medulla and the adrenal cortex. The adrenal medulla is the central portion of the gland and secretes **epinephrine** and **norepinephrine**. These hormones produce the same effects that the sympathetic nervous system produces. They increase heart rate, breathing rate, blood pressure, and all the other actions that prepare the body for stressful situations.

2.0 OBJECTIVES

At the end of this unit, you will be able to:

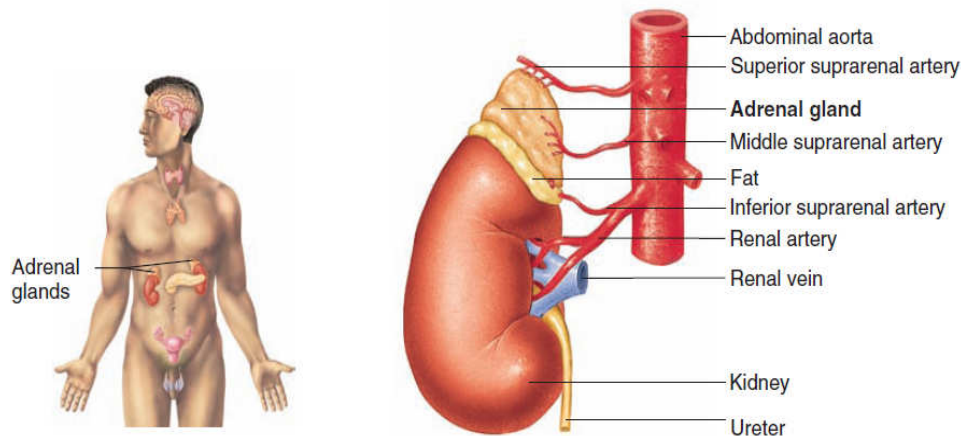
- Describe the anatomy of the adrenal gland
- List the hormones released by the adrenal glands and give the functions of each.

3.0 MAIN CONTENT

3.1 Anatomy of the adrenal glands

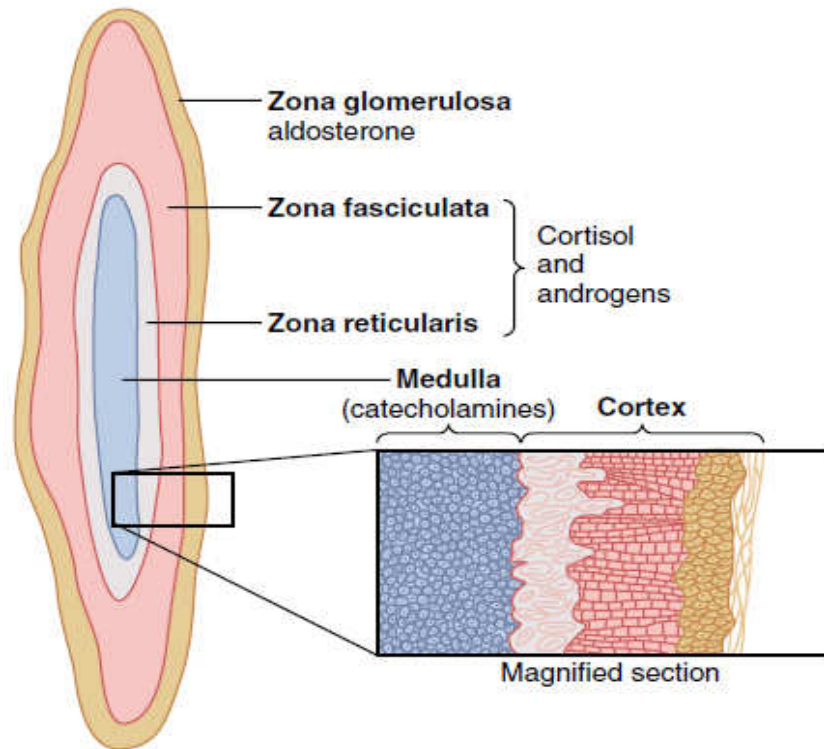
The adrenal glands, or suprarenal (above the kidney) glands, are two small glands that are located superior to each kidney. Each adrenal gland has an inner part, called the adrenal medulla (marrow or middle), and an outer part, called the adrenal cortex (bark or outer). The cortex has three indistinct layers: the zona glomerulosa, the zona fasciculata, and the zona reticularis.

The adrenal glands



The medulla and the three layers of the cortex are structurally and functionally specialized. In many ways, an adrenal gland is four glands in one.

1. The *zona glomerulosa*, a thin layer of cells that lies just underneath the capsule, constitutes about 15 percent of the adrenal cortex. These cells are the only ones in the adrenal gland capable of secreting significant amounts of *aldosterone* because they contain the enzyme *aldosterone synthase*, which is necessary for synthesis of aldosterone. The secretion of these cells is controlled mainly by the extracellular fluid concentrations of *angiotensin II* and *potassium*, both of which stimulate aldosterone secretion.
2. The *zona fasciculata*, the middle and widest layer, constitutes about 75 percent of the adrenal cortex and secretes the glucocorticoids *cortisol* and *corticosterone*, as well as small amounts of *adrenal androgens* and *estrogens*. The secretion of these cells is controlled in large part by the hypothalamic-pituitary axis via *adrenocorticotrophic hormone* (ACTH).
3. The *zona reticularis*, the deep layer of the cortex, secretes the adrenal androgens *dehydroepiandrosterone* (DHEA) and *androstenedione*, as well as small amounts of estrogens and some glucocorticoids. ACTH also regulates secretion of these cells, although other factors such as *cortical androgen-stimulating hormone*, released from the pituitary, may also be involved. The mechanisms for controlling adrenal androgen production, however, are not nearly as well understood as those for glucocorticoids and mineralocorticoids.



Secretion of adrenocortical hormones by the different zones of the adrenal cortex and secretion of catecholamines by the adrenal medulla.

Hormones of the adrenal medulla

Approximately 80% of the hormone released from the adrenal medulla is epinephrine, or adrenaline. The remaining 20% is norepinephrine. The adrenal medulla consists of cells derived from the same cells that give rise to postganglionic sympathetic neurons, which secrete norepinephrine. Epinephrine is derived from norepinephrine. The adrenal medulla and the sympathetic division function together to prepare the body for physical activity, producing the “fight-or-flight” response. Some of the major effects of the hormones released from the adrenal medulla are the following:

1. Increased breakdown of glycogen to glucose in the liver, the release of the glucose into the blood, and the release of fatty acids from fat cells. The glucose and fatty acids are used as energy sources to maintain the body's increased rate of metabolism.
2. Increased heart rate, which increases blood pressure and blood delivery to tissues.
3. Increased vasodilation of blood vessels of the heart and skeletal muscle, which increases blood flow to the organs responsible for increased physical activity. The hormones increase vasoconstriction of blood vessels to the internal organs and skin, which decreases blood flow to organs not directly involved in physical activity.

4. Increased metabolic rate of several tissues, especially skeletal muscle, cardiac muscle, and nervous tissue. The release of adrenal medullary hormones primarily occurs in response to stimulation by sympathetic neurons because the adrenal medulla is a specialized part of the autonomic nervous system.

Several conditions, including exercise, emotional excitement, injury, stress, and low blood glucose levels, lead to the release of adrenal medullary hormones.

Hormones of the adrenal cortex

The adrenal cortex secretes three hormone types: mineralocorticoids, glucocorticoids, and androgens. All are similar in structure in that they are steroids, highly specialized lipids that are derived from cholesterol. Because they are lipid-soluble, they are not stored in the adrenal gland cells but diffuse from the cells as they are synthesized. Adrenal cortical hormones are transported in the blood in combination with specific plasma proteins; they are metabolized in the liver and excreted in the bile and urine. The hormones of the adrenal cortex bind to nuclear receptors and stimulate the synthesis of specific proteins that are responsible for producing the cell's responses.

Mineralocorticoids

The major secretory products of the zona glomerulosa are the mineralocorticoids. The mineralocorticoids are so named because they are steroids, produced by the adrenal cortex, that affect the "minerals" Na^+ , K^+ , and H^+ . **Aldosterone** is produced in the greatest amounts, although other, closely related mineralocorticoids are also secreted. Aldosterone increases the rate of Na^+ reabsorption by the kidneys, thereby increasing blood Na^+ levels. Sodium reabsorption can result in increased water reabsorption by the kidneys and an increase in blood volume, providing ADH is also secreted. Increased blood volume can increase blood pressure.

Aldosterone increases K^+ and H^+ excretion into the urine by the kidneys, thereby decreasing blood levels of K^+ and H^+ . When aldosterone is secreted in high concentrations, it can result in abnormally low blood levels of K^+ and H^+ . The reduction in H^+ can cause alkalosis, an abnormally elevated pH of body fluids.

Glucocorticoids

The zona fasciculata of the adrenal cortex primarily secretes glucocorticoids. The glucocorticoids are so named because they are steroids produced by the adrenal cortex that affect glucose metabolism. The major glucocorticoid is **cortisol**. The target tissues and responses to the glucocorticoids are numerous. The two major types of responses to glucocorticoids are classified as metabolic and anti-inflammatory.

Cortisol increases the breakdown of protein and fat and increases their conversion to forms that can be used as energy sources by the body. For example, cortisol causes proteins in skeletal muscles to be broken down to amino acids, which are then released into the circulatory system.

Cortisol acts on the liver, causing it to convert amino acids to glucose, which is released into the blood or stored as glycogen. Thus, cortisol increases blood sugar levels. Cortisol also acts on adipose tissue, causing fat stored in fat cells to be broken down to fatty acids, which are released into the circulation. The glucose and fatty acids released into the circulatory system are taken up by tissues and used as sources of energy.

Glucocorticoids decrease the intensity of the inflammatory and immune responses by decreasing both the number of white blood cells and the secretion of inflammatory chemicals from tissues. **Cortisone**, a steroid closely related to cortisol, is often given as a medication to reduce inflammation that occurs in response to injuries. It is also given to reduce the immune and inflammatory responses that occur as a result of allergic reactions or diseases resulting from abnormal immune responses, such as rheumatoid arthritis or asthma. In response to stressful conditions, cortisol is secreted in larger than normal amounts. Cortisol aids the body in responding to stressful conditions by providing energy sources for tissues. If stressful conditions are prolonged, however, immunity can be suppressed enough to make the body susceptible to infections. Cortisol secretion is regulated through the hypothalamus and anterior pituitary gland. Stress and low blood glucose levels stimulate increased **corticotropin-releasing hormone (CRH)** from the hypothalamus. CRH stimulates increased **adrenocorticotrophic hormone (ACTH)** secretion from the anterior pituitary gland. ACTH stimulates increased cortisol secretion.

Clinical correlates

I. Adrenal Tumors

The two major disorders of the adrenal medulla are both tumors. Pheochromocytoma is a benign tumor; neuroblastoma is a malignant tumor.

Symptoms, resulting from the release of large amounts of epinephrine and norepinephrine, include hypertension (high blood pressure), sweating, nervousness, pallor, and tachycardia (rapid heart rate). The high blood pressure results from the effect of these hormones on the heart and blood vessels and is correlated with an increased chance of heart disease and stroke.

II. Chronic adrenocortical insufficiency, often called **Addison disease**, results from abnormally low levels of aldosterone and cortisol in the blood. The cause of many cases of chronic adrenocortical insufficiency is unknown, but it frequently results from an autoimmune disease in which the body's defense mechanisms inappropriately destroy the adrenal cortex. Other causes are infections and tumors that damage the adrenal cortex.

I. Aldosteronism is caused by an excess production of aldosterone. Primary aldosteronism results from an adrenal cortex tumor, and secondary aldosteronism occurs when an extraneous factor, such as an overproduction of renin, a substance produced by the kidneys, increases aldosterone secretion. Major symptoms of aldosteronism are hypernatremia (elevated blood Na^+), hypokalemia (decreased K^+), alkalosis (decreased H^+), and high blood pressure due to the retention of water and Na^+ by the kidneys.

IV. Cushing syndrome is a disorder characterized by the hypersecretion of cortisol and androgens and possibly by excess aldosterone production. Most cases are caused by excess ACTH production by nonpituitary tumors, which usually result from a type of lung cancer. Some cases of increased ACTH secretion do result from pituitary tumors. Sometimes adrenal tumors or unidentified causes can be responsible for hypersecretion of the adrenal cortex without increases in ACTH secretion.

4.0 SUMMARY

- The adrenal glands are near the superior poles of the kidneys. The adrenal medulla arises from the same cells that give rise to postganglionic sympathetic neurons.
- The adrenal cortex is divided into three layers: the zona glomerulosa, the zona fasciculata, and the zona reticularis.
- Epinephrine accounts for 80% and norepinephrine for 20% of the adrenal medulla hormones. The adrenal medulla hormones prepare the body for physical activity.
- The zona glomerulosa secretes the mineralocorticoids, especially aldosterone. Aldosterone acts on the kidneys to increase Na^+ and to decrease K^+ and H^+ levels in the blood.
- The zona fasciculata secretes glucocorticoids, especially cortisol. Cortisol increases fat and protein breakdown, increases glucose synthesis from amino acids, decreases the inflammatory response.
- The zona reticularis secretes androgens. In females, androgens stimulate axillary and pubic hair growth and sexual drive

Activity

- I. Examine microscopic slides of the adrenal gland in the histology laboratory
- II. Distinguish between the darker cortex with vertically arranged rows of cells and the lighter medulla.

SELF – ASSESSMENT EXERCISE

1. What controls glucocorticoid secretion?
2. The adrenal medulla
 - a. produces steroids.
 - b. has cortisol as its major secretory product.
 - c. decreases its secretions during exercise.
 - d. is formed from a modified portion of the sympathetic division of the ANS.
 - e. all of the above.
3. If aldosterone secretions increase,
 - a. blood potassium levels increase.
 - b. blood hydrogen levels increase.
 - c. acidosis results.
 - d. blood sodium levels decrease.
 - e. blood volume increases.
4. Glucocorticoids (cortisol)
 - a. increase the breakdown of fats.
 - b. increase the breakdown of proteins.
 - c. increase blood glucose levels.
 - d. decrease inflammation.
 - e. all of the above
5. What controls the secretion of aldosterone (a mineralocorticoid)?

UNIT 5 PANCREAS

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Anatomy of the pancreas
- 4.0 Summary

1.0 INTRODUCTION

The pancreas is located behind the stomach. It is an endocrine gland as well as an exocrine gland. It is considered an exocrine gland because it secretes digestive enzymes into a duct that leads to the small intestine. It is considered an endocrine gland because it contains structures known as islets of Langerhans that secrete hormones into the bloodstream. The islets of Langerhans secrete two hormones—insulin and glucagon.

2.0 OBJECTIVES

At the end of this unit, you will be able to:

- describe the anatomy of the pancreas
- identify the pancreatic hormones and to explain their physiological effects.

3.0 MAIN CONTENT

3.1 Anatomy of the pancreas

The pancreas lies retroperitoneally in roughly the transpyloric plane. For descriptive purposes it is divided into head, neck, body and tail.

The head lies in the C-curve of the duodenum and sends out the *uncinate process* which hooks posteriorly to the superior mesenteric vessels as these travel from behind the pancreas into the root of the mesentery. Posteriorly lie the inferior vena cava, the commencement of the portal vein, aorta, superior mesenteric vessels, the crura of diaphragm, coeliac plexus, the left kidney and suprarenal gland. The tortuous splenic artery runs along the upper border of the pancreas. The splenic vein runs behind the gland, receives the inferior mesenteric vein and joins the superior mesenteric to form the portal vein behind the pancreatic neck.

To complete this list of important posterior relationships, the common bile duct lies either in a groove in the right extremity of the gland or embedded in its substance, as it passes to open into the second part of the duodenum. Anteriorly lies the stomach separated by the lesser sac. To the left, the pancreatic tail lies against the hilum of the spleen. Blood is supplied from the splenic and the pancreaticoduodenal arteries; the corresponding veins drain into the portal system.

Pancreatic hormones

Insulin promotes the uptake of glucose by cells. It therefore reduces glucose concentrations in the bloodstream. It also promotes the transport of amino acids into cells and increases protein synthesis. Glucagon increases glucose concentrations in the bloodstream and slows down protein synthesis.

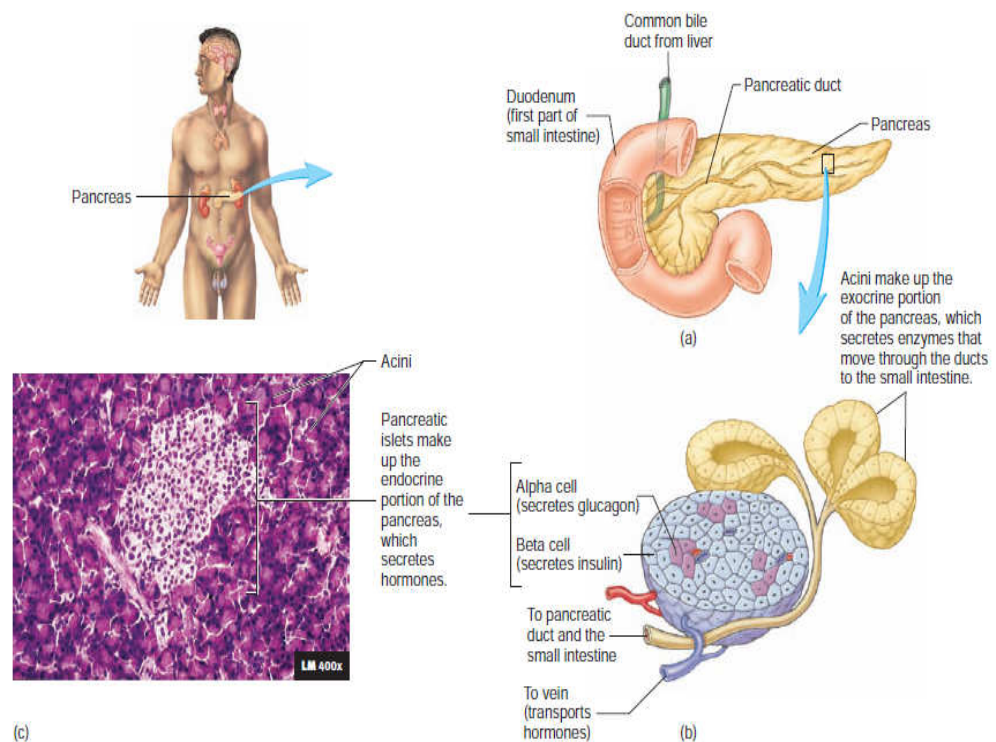


Figure 15.24 Pancreas **APR**

(a) Anterior view of the pancreas. (b) A pancreatic islet consists of clusters of specialized cells among the acini of the exocrine portion of the pancreas. (c) Light micrograph of pancreatic tissue. The stain used for this slide does not distinguish between alpha and beta cells.

Effect of Insulin and Glucagon on Their Target Tissues

Insulin

Insulin increases the uptake of glucose and amino acids by target cells. Once insulin binds to its receptors, the receptors cause specific proteins in the membrane to become phosphorylated. Part of the cell's response to insulin is to increase the number of transport proteins in the membrane of cells for glucose and amino acids. The major target tissues of insulin are the liver, adipose tissue, the skeletal muscles, and the

satiety center within the hypothalamus of the brain. The **satiety center** is a collection of neurons in the hypothalamus that controls appetite. Unlike the satiety center, most of the nervous system does not depend on insulin for the uptake of glucose. Insulin is very important for the normal functioning of the nervous system, however, because insulin regulates blood glucose levels. If blood glucose levels are not maintained within a normal range, the brain malfunctions because glucose is its primary energy source. When insulin levels increase, the movement of glucose and amino acids into cells increases. Glucose molecules that are not immediately used as an energy source are stored as glycogen in the liver, skeletal muscle, and other tissues, or they are converted to fat in adipose tissue. Amino acids are used to synthesize proteins.

When insulin levels decrease, the opposite effects are observed. The movement of glucose and amino acids into tissues slows. Glycogen is broken down to glucose, which is released from the liver, but not from skeletal muscle. Adipose tissue releases fatty acids, and proteins are broken down into amino acids, especially in skeletal muscle. The amino acids are released into the blood, taken up by the liver, and used to synthesize glucose, which is released into the blood.

When insulin levels decrease, the liver uses fatty acids to make **acetoacetic acid**, which is converted to **acetone** and **β -hydroxybutyric acid**. These three substances collectively are referred to as **ketone bodies**. The liver releases the ketone bodies into the blood, from which other tissues take them up and use them as a source of energy. The ketone bodies are smaller, more readily used “packets” of energy than are fatty acids. Ketone bodies, however, are acids that can adversely affect blood pH if too many of them are produced. In addition, when insulin levels are low, the liver releases cholesterol and triglycerides into the blood.

Glucagon

Glucagon increases blood sugar and ketone levels. Glucagon primarily influences the liver, although it has some effect on skeletal muscle and adipose tissue. The pancreas secretes glucagon into the hepatic portal system, which carries blood to the liver from the pancreas and intestines. Glucagon binds to membrane-bound receptors, activates G proteins, and increases cAMP synthesis. In general, glucagon causes the breakdown of glycogen to glucose and increases glucose synthesis from amino acids. The release of glucose from the liver increases blood glucose levels.

Regulation of pancreatic hormones

Insulin

Blood levels of nutrients, neural stimulation, and hormones control the secretion of insulin. Elevated blood levels of glucose directly affect the β cells and stimulate insulin secretion. Low blood levels of glucose directly inhibit insulin secretion. Thus, blood glucose levels play a major role in the regulation of insulin secretion. Certain amino acids also stimulate insulin secretion by acting directly on the β cells. After a meal, when glucose and amino acid levels increase in the circulatory system, insulin secretion increases. During periods of fasting, when blood glucose levels are low, the rate of insulin secretion declines. The autonomic nervous system also controls insulin secretion. The parasympathetic stimulation of digestive system organs is associated with food intake. Parasympathetic stimulation of the pancreas increases its secretion of insulin and digestive enzymes. Sympathetic stimulation inhibits insulin secretion and helps prevent a rapid fall in blood glucose levels during periods of physical activity or excitement. This response is important for maintaining normal functioning of the nervous system. Gastrointestinal hormones involved with the regulation of digestion, such as gastrin, secretin, and cholecystokinin, increase insulin secretion.

Glucagon

Low blood glucose levels stimulate glucagon secretion, and high blood glucose levels inhibit it. Certain amino acids and sympathetic stimulation also increase glucagon secretion. After a high-protein meal, amino acids increase both insulin and glucagon secretion. Insulin causes target tissues to accept the amino acids for protein synthesis, and glucagon increases the process of glucose synthesis from amino acids in the liver.

Clinical correlates

- ❖ Diabetes mellitus results from the inadequate secretion of insulin or the inability of tissues to respond to insulin. As a result, blood sugar levels increase. The two major types of diabetes are type 1 and type 2 diabetes. Type 1 diabetes mellitus, also called insulin-dependent diabetes mellitus (IDDM), results from diminished or absent insulin secretion. It affects approximately 5%–10% of people with diabetes mellitus and most commonly occurs in young people. Type 1 diabetes mellitus develops as a result of autoimmune destruction of the pancreatic islets, and symptoms appear after approximately 90% of the islets have been destroyed. Heredity plays a role in the condition, although the initiation of pancreatic islet destruction may involve a viral infection of the pancreas.

Type 2 diabetes mellitus, also called noninsulin-dependent diabetes mellitus (NIDDM), results from insulin resistance, the inability of tissues to respond normally to insulin. It affects approximately 90%–95% of people who have diabetes mellitus and usually develops in people older than 40–45 years of age, although the age of onset varies considerably. People with type 2 diabetes mellitus have a reduced number of functional receptors for insulin, or one or more of the enzymes activated by the insulin receptor are defective. Heredity influences the likelihood of developing type 2 diabetes, but it is not as important a risk factor as for type 1 diabetes.

- ❖ Three potentially life-threatening conditions are associated with untreated diabetes mellitus: diabetic ketoacidosis, hyperglycaemic hyperosmolar state, and insulin shock.
- ❖ **Diabetic ketoacidosis (DKA)** is the triad of hyperglycemia, ketosis, and acidosis. **Ketosis** is the presence of excess ketone bodies in the blood.
- ❖ **Hyperglycemic hyperosmolar state (HHS)** consists of very elevated blood sugar levels. It is most likely to develop in type 2 diabetics who have enough insulin to prevent ketosis, but not enough insulin to prevent hyperglycemia.
- ❖ **Insulin shock** occurs when there is too much insulin relative to the amount of blood glucose. Too much insulin, too little food intake after an injection of insulin, or increased metabolism of glucose due to excess exercise by a diabetic patient can cause insulin shock. The high levels of insulin cause target tissues to take up glucose at a very high rate. As a result, blood glucose levels rapidly fall to a low level.

4.0 SUMMARY

- The exocrine portion of the pancreas consists of a complex duct system, which ends in small sacs, called acini, that produce pancreatic digestive juices.
- The endocrine portion consists of the pancreatic islets. Each islet is composed of alpha cells, which secrete glucagon, and beta cells, which secrete insulin.
- Insulin's target tissues are the liver, adipose tissue, muscle, and the satiety center in the hypothalamus. The nervous system is not a target tissue, but it does rely on blood glucose levels maintained by insulin.
- Insulin increases the uptake of glucose and amino acids by cells. Glucose is used for energy, stored as glycogen, or converted into fats. Amino acids are used to synthesize proteins. Low levels of insulin promote the formation of ketone bodies by the liver.

- Glucagon's target tissue is mainly the liver. Glucagon causes the breakdown of glycogen to glucose and the synthesis of glucose from amino acids. The liver releases glucose into the blood.
- Insulin secretion increases because of elevated blood glucose levels, an increase in some amino acids, parasympathetic stimulation, and gastrointestinal hormones. Sympathetic stimulation decreases insulin secretion.
- Glucagon secretion is stimulated by low blood glucose levels, certain amino acids, and sympathetic stimulation. Somatostatin inhibits insulin and glucagon secretion.

Activity

- I. Examine the microscopic slides of the pancreatic tissue in the histology laboratory and note the pancreatic islets which are the endocrine portion of the pancreas.
- II. In the gross anatomy laboratory, study the anatomy of the pancreas and its relations.

SELF – ASSESSMENT EXERCISE

1. What are the physiological effects of the pancreatic hormones?
2. What are causes of diabetes mellitus (insulin deficiency)?
3. Within the pancreas, the pancreatic islets produce
 - a. insulin
 - b. glucagon.
 - c. digestive enzymes.
 - d. both a and b.
 - e. all of the above.
4. Insulin increases
 - a. the uptake of glucose by its target tissues.
 - b. the breakdown of protein.
 - c. the breakdown of fats.
 - d. glycogen breakdown in the liver.
 - e. all of the above.
5. Which of these tissues is least affected by insulin?
 - a. adipose tissue
 - b. heart
 - c. skeletal muscle
 - d. brain
 - e. liver

UNIT 6 OTHER ENDOCRINE GLANDS

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Pineal body
- 4.0 Summary

1.0 INTRODUCTION

There are other hormone producing organs which most people are not aware of or consider as parts of the endocrine system. They will be discussed in great details in this chapter, they include the pineal gland, thymus and the hormone – like substances. We will discuss the testis and ovaries as well.

2.0 OBJECTIVES

At the end of this unit, you will be able to:

- discuss the anatomy and functions of other glands with endocrine functions.
- define autocrine and paracrine agents
- discuss the age-related changes that occur in the endocrine system.

3.0 MAIN CONTENT

3.1 Pineal body

The pineal gland is a small, pinecone-shaped structure located superior and posterior to the thalamus of the brain. The pineal gland produces a hormone called melatonin, which inhibits the functions of the reproductive system in some animals. Melatonin helps to regulate your circadian rhythms, which is your biological clock. Your biological clock helps you decide when you should be awake or asleep. Melatonin is also thought to play a role in the onset of puberty. Tumors that destroy the pineal gland correlate with early sexual development, and tumors that result in pineal hormone secretion correlate with retarded development of the reproductive system. It is not clear, however, if the pineal gland controls the onset of puberty.

The amount of light detected by the eyes regulates the rate of melatonin secretion. The axons of some neurons in the retina pass from the optic chiasm to the suprachiasmatic nucleus in the hypothalamus, which influences the pineal gland through sympathetic neurons. Increased light exposure inhibits melatonin

secretion, whereas darkness allows melatonin secretion. Melatonin is sometimes called the “hormone of darkness” because its production increases in the dark. In many animals, longer day length (shorter nights) causes a decrease in melatonin secretion, whereas shorter day length (longer nights) causes an increase in melatonin secretion. For example, in animals that breed in the spring, increased day length results in decreased melatonin secretion. With decreased inhibition of the hypothalamus by melatonin, sex hormone production increases, which promotes the development of reproductive structures and behavior. In the fall, decreased day length results in increased melatonin secretion, decreased sex hormone production, atrophy of reproductive

Thymus gland

The thymus is in the neck and superior to the heart in the thorax; it secretes a hormone called **thymosin**. Both the thymus and thymosin play an important role in the development and maturation of the immune system

Thymosin promotes the production of certain lymphocytes.

Other hormone producing organs include:

Cholecystokinin are released from the gastrointestinal tract. They regulate digestive functions by influencing the activity of the stomach, intestines, liver, and pancreas. The kidneys secrete a hormone in response to reduced oxygen levels in the kidney. The hormone is called erythropoietin. It acts on red bone marrow to increase the production of red blood cells. In pregnant women, the placenta is an important source of hormones that maintain pregnancy and stimulate breast development. These hormones include estrogen, progesterone, and human chorionic gonadotropin, which is similar in structure and function to LH. These hormones are essential to the maintenance of pregnancy

Hormone - like Substances

Autocrine chemical messengers are chemicals released by a cell that affect the cell producing it or affect nearby cells of the same cell type. Examples of autocrine chemical messengers include a group of related chemical mediators called eicosanoids, which are derived from fatty acids. The eicosanoids include prostaglandins, thromboxanes, prostacyclins, and leukotrienes.

Paracrine chemical messengers are chemicals released by a cell that affect nearby cells of a different cell type. Examples of paracrine chemical messengers include growth factors, clotting factors, and histamine. Autocrine and paracrine chemical messengers differ from hormones in that they are not secreted from discrete endocrine glands, they have local effects rather than systemic effects, or they have functions that are not understood adequately to explain their role in the body. The schemes used to classify chemicals on the basis of their functions are useful, but they do not indicate that a specific molecule always performs as the same type of chemical messenger in every place it is found. Some chemical messengers, such as prostaglandins, have both autocrine and paracrine functions. Furthermore, some of these chemicals can also act as hormones. Testosterone produced in the testes has a paracrine effect on the development of sperm cells, but it is released into the blood and has an endocrine effect on skeletal muscle development.

Effects of aging on the endocrine system

Age-related changes to the endocrine system include a gradual decrease in the secretion of some, but not all, endocrine glands. Some of the decreases in secretion may be due to a decrease in physical activity as people age. GH secretion decreases as people age, and the decrease is greatest in people who do not exercise. It may not occur in older people who exercise regularly. Decreasing GH levels may explain some of the gradual decrease in bone and muscle mass and some of the increase in adipose tissue in many elderly people.

Administering GH to slow or prevent the consequences of aging has not been established to be effective, however, and unwanted side effects are possible. A decrease in melatonin secretion may influence age-related changes in sleep patterns. The secretion of thyroid hormones decreases slightly with age. Age-related damage to the thyroid gland by the immune system can occur, and this happens in women more than in men. Approximately 10% of elderly women have some reduction in thyroid hormone secretion.

Parathyroid hormone secretion does not appear to decrease with age. Blood levels of Ca^{2+} may decrease slightly because of reduced dietary calcium intake and vitamin D levels. The greatest risk is a loss of bone matrix as parathyroid hormone increases to maintain blood levels of Ca^{2+} within their normal range.

Reproductive hormone secretion gradually declines in elderly men, and women experience menopause.

There are no age-related decreases in the ability to regulate blood glucose levels. There is an age-related tendency to develop type 2 diabetes mellitus in those who have a familial tendency to do so, and it is correlated with age-related increases in body weight. Thymosin from the thymus decreases with age. Fewer immature lymphocytes are able to mature and become functional, and the immune system becomes less effective in protecting the body. There is an increased susceptibility to infection and to cancer.

4.0 SUMMARY

- The pineal gland produces melatonin, which can inhibit reproductive maturation and may regulate sleep–wake cycles.
- The thymus produces thymosin, which is involved in the development of the immune system.
- The kidneys produce erythropoietin, which stimulates red blood cell production.
- Autocrine chemical messengers are chemicals that locally affect cells producing them or affect cells of the same type.
- There is a gradual decrease in the secretion rate of most, but not all, hormones. Some decreases are secondary to gradual decreases in physical activity.

SELF – ASSESSMENT EXERCISE

1. Melatonin
 - a. is produced by the posterior pituitary.
 - b. production increases as day length increases.
 - c. inhibits the development of the reproductive system.
 - d. increases GnRH secretion from the hypothalamus.
 - e. decreases the tendency to sleep
2. The hormone secretin
 - a. plays an important role in the development and maturation of the immune system.
 - b. is released from the gastrointestinal tract.
 - c. acts on red bone marrow to increase the production of red blood cells.
 - d. is essential to the maintenance of pregnancy.
3. Which of the following statements about autocrine and paracrine agents is true?
 - a. They usually have a local effect, but sometimes can have systemic effects.
 - b. They typically are not produced in discrete endocrine glands.

- c. Paracrine agents affect different cell types from which the paracrine agent is released.
 - d. Eicosanoids and prostaglandins are examples of autocrine agents.
 - e. All of the above.
4. The production of which hormone does not decrease with age?
- a. growth hormone
 - b. melatonin
 - c. thyroid hormones
 - d. parathyroid hormones
 - e. reproductive hormones

MODULE 3 CENTRAL NERVOUS SYSTEM

CONTENTS

Unit 1	Embryology of the central nervous system
Unit 2	Cerebral hemisphere
Unit 3	Brain stem
Unit 4	Diencephalon and Basal ganglia
Unit 5	Ventricles and cerebrospinal fluid
Unit 6	Spinal cord
Unit 7	Blood supply of central nervous system

INTRODUCTION

During a picnic on a sunny spring day, it is easy to concentrate on the delicious food and the pleasant surroundings. The maintenance of homeostasis requires no conscious thought. The autonomic nervous system (ANS) helps keep body temperature at a constant level by controlling the activity of sweat glands and the amount of blood flowing through the skin. The central nervous system (CNS) consists of the brain and spinal cord, the peripheral nervous system (PNS) consists of sensory receptors and nerves outside the CNS. The PNS includes 12 pairs of cranial nerves and 31 pairs of spinal nerves. The CNS receives sensory information, integrates and evaluates that information, stores some information, and initiates reactions..

OBJECTIVES

At the end of this module, you should be able to:

- discuss the formation of the brain and the spinal cord from the most primitive neural tube
- discuss the anatomy of the central nervous system.

UNIT 1 EMBRYOLOGY OF THE CENTRAL NERVOUS SYSTEM

1.0	Introduction
2.0	Objectives
3.0	Main Content
3.1	Anomalies of closure
4.0	Summary

1.0 INTRODUCTION

Brain development occurs at several stages during childhood. The critical period for brain growth appears to be during the first sixteen weeks of life. At birth, a baby's brain weighs approximately 25 per cent of its future adult weight. By the time the child is two years old the brain has increased to 75 per cent, and by six years, 90 per cent of its eventual weight. This, then, indicates phenomenal growth of the central nervous system during the early years. Peripheral nerves continue to become myelinated (grow fatty sheaths to increase nerve transmission rate) and fine physical control appears as the child moves towards adult status. With the unique environment impinging on every waking and sleeping hour, this plastic nervous system constantly matures and changes as demands are put upon it.

2.0 OBJECTIVES

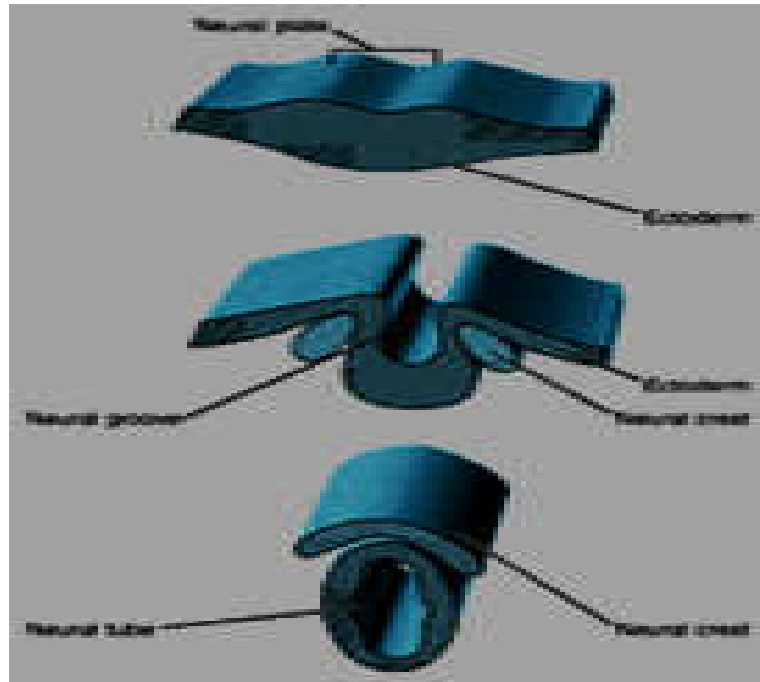
At the end of this unit, you will be able to:

- describe the transformation of the neural tube into the different parts of the definitive brain.
- differentiate between the events that lead to the formation of the brain and the spinal cord.
- discuss the cellular formation, differentiation and migration, and the role of such processes in the formation of the brain and spinal cord.

3.1 Anomalies of closure

The CNS is an ectodermal derivative that begins to differentiate in the 3rd intrauterine week in humans.

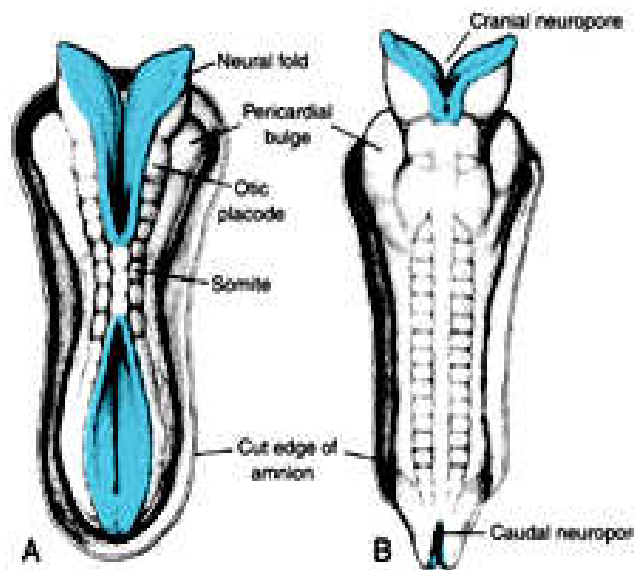
The brain starts its development soon after conception; it is vulnerable to damage through pregnancy. The first indication of the nervous system is within the neural plate, a thickened area of the ectoderm that forms after the ovum has been fertilised in the third week of gestation. Slippers shaped neural plate thickens rostral to the primitive node and midline depression in the neural plate forms the neural groove. Edges of the neural groove rise on both sides of the midline, fusing to convert the groove to a tube. This neural tube is the beginning of the brain and spinal cord. As the neural tube separates from the surface ectoderm cells, the neural folds form the neural crest. Ganglia of the spine, cranial and autonomic nervous system develop from the neural crest. Fusion of the neural groove edges begin in the cervical region and proceed in cephalocaudal directions.



Formation of the neural tube.

A more cranial site of closure is noted in the forebrain to close the cranial neuropore in conjunction with the cervical closure site on the 25th intrauterine day. Caudal neuropore closes in similar fashion 2 days after the cranial counterpart. The neural tube give rise to the brain and spinal cord, the canal of the tube differentiates into the ventricles and the central canal of the spinal cord.

The embryo at twenty-three days shows the hindbrain and midbrain to be formed, and the neural tube closes. In the fourth week the head folds begin to develop as the forebrain grows rapidly. In the fifth week the eye starts to grow, and cerebral hemispheres also develop from this area. The nerves of the branchial arches become the cranial nerves. Peak head breadth growth velocity occurs at thirteen post-menstrual weeks, although a relatively high velocity continues to about thirty weeks. Peak head circumference velocity occurs two to three weeks later, because the cerebellum situated at the back of the skull grows later than the cerebrum. Head volume, representing brain size, has its peak velocity at thirty weeks and growth rapidly slows after this.



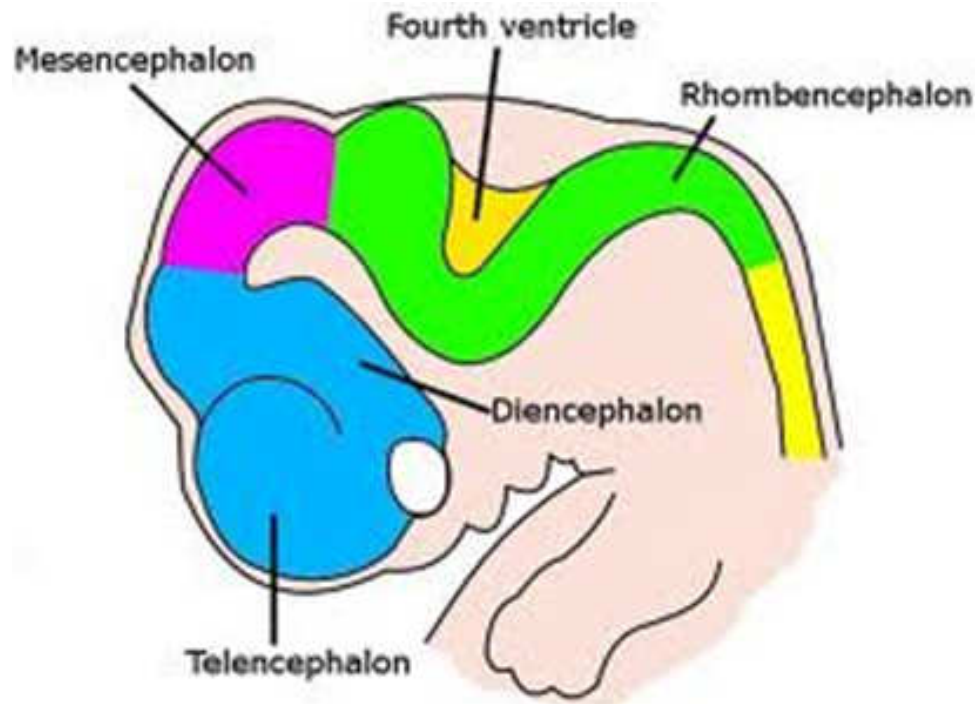
Anomalies of the closure of the neural tube results in several clinical conditions such as anencephaly and spina bifida. Various degrees of these anomalies noted in many regions of the world. The introduction of folic acid supplementation in ante-natal care had drastically reduced the occurrence of such anomalies.



Anencephaly

Spina bifida

3.1.3 BRAIN VESICLE FORMATION

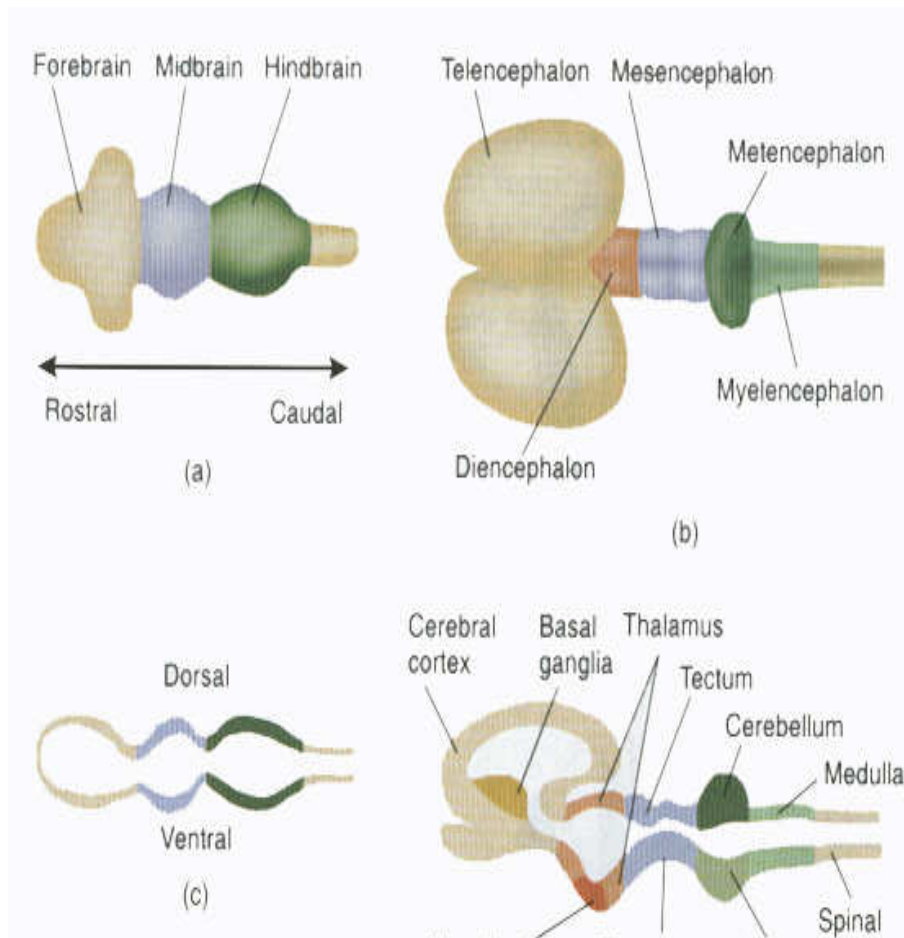


The 4th week witnesses the differentiation of the cranial part of the neural tube into the 3 brain vesicles. Boundary demarcation follows differential growth rate leading to the formation of the prosencephalon, mesencephalon and rhombencephalon cranio-caudally. Same mechanism results in the formation of 2 flexures, cervical and cephalic flexures. Cephalic flexure opens to the ventral surface while the cervical opens to the dorsal surface.

By the 5th week of development, the Prosencephalon is divided into the Telencephalon and Diencephalon by differential growth rate. The Rhombencephalon also divides into the cranial Metencephalon and caudal Myelencephalon roughly by the cephalic (Pontine) flexure. The canal of the neural tube follows the differentiation leading to formation of Lateral, 3rd, 4th ventricles and the Aqueduct of Sylvius.

Telencephalon

The prosencephalon undergoes massive growth in the 5th week. Growth is more in the cranio-lateral portion as against the midline portion. The differential growth rate results in the ballooning of the lateral portion forming the telencephalon while the more quiescent midline portion forms the diencephalon. Ballooning is more in the dorsal part of the telencephalon, the ventral part consist largely of the corpus striatum. The roof plate ependymal layers in addition to surrounding vascular mesenchyme form the choroid plexus.



The choroid plexus bearing portion get “invaginated” due to the lateral expansion of the ballooning lateral wall.

Just beyond the choroid plexus bearing region, there is proliferation of neuroblast resulting in the slightly thickened portion that forms the Hippocampus. The hippocampal portion get “buried” within the enlarging cortex alongside the roof portion. It bulges into the cavity of the lateral ventricle. The enlarging dorsal portion of the telencephalon forms the cerebral cortex. Expansion is in the anterior, posterior and lateral directions forming the frontal, occipital and temporal lobes respectively. The ballooned cerebral cortex envelopes the diencephalon, mesencephalon and upper part of metencephalon.

Axons from the cortical neurons coalesce and pass through the corpus striatum dividing it into the medial caudate nucleus and the lateral lentiform nucleus. The dividing axonal bundle forms the internal capsule.

In the latter quarter of gestation, expansion of the cerebral cortex exceeds the capacity of the cranial vault resulting in convolutions.

The degree of convolutions is directly related to the degree of behavioural and neuronal complexity.

Diencephalon

The portion of the prosencephalon just caudal and medial to the telencephalon. It consists of roof plate and 2 alar plate, basal plate and floor plate are absent. The roof plate consists of ependymal lining which forms the choroid plexus with the surrounding vascular mesenchyme. There is a caudal neuroepithelial thickening on the dorsal surface which evaginates around the 7th week to form the pineal body sitting on the mesencephalon.

The alar plate witness aggregation of cells which is divided into a dorsal thalamus and a ventral hypothalamus by a hypothalamic sulcus. Cells of the thalamus proliferate very rapidly resulting in an inward bulge of the Thalamus obliterating the cavity and forming an adhesion. A ventral thickening of the neuroepithelium results in the formation of the mammillary body which is functionally related to the hypothalamus. An extension of the ventral wall forms the infundibulum. The infundibulum alongside the Rathke's pouch (from the oral cavity) form the pituitary gland sitting in the Sella Turcica. The cephalic flexure topographically makes the connection possible.

Mesencephalon

It has the typical basal and alar portions, the basal (ventral) part mediate motor functions while the alar (dorsal) part mediates sensory function. The sulcus limitans forms the boundary between the 2 sides. The basal plate contains 2 groups of motor cells that aggregate into nuclei viz; **Somatic efferent**; medially placed CN3 and CN4 motor supply to the extraocular muscles.

General visceral efferent; laterally placed nucleus of Edinger-Westphal (sphincter pupillary muscle).

Fibers that connect the spinal cord through the pons from the cerebral cortex pass through the marginal layer over the basal layer. Marginal layer over the basal layer enlarges to accommodate fibers forming the crus cerebri. No particular nucleus is formed in the alar plate of the mesencephalon. The overlying marginal layer of the alar plate had neuroblast wave in it resulting in its enlargement. Initially 2 longitudinal elevation is noticed in the marginal layer of the alar plate, with further development, a transverse furrow separates them. 4 colliculi are thus formed namely the inferior and superior colliculi on both sides of the median sulcus. Serve as synaptic relay centre for auditory and visual impulses respectively.

Metencephalon

It is the cranial portion of the Rhombencephalon. It has well defined basal and alar plate in which neuroblasts form cellular aggregates that form nuclei. The roof plate consists largely of ependymal layer, opened alar plate. It is also the point of formation of the pons and cerebellum.

The fibres from the cerebral cortex to the spinal cord pass through the marginal layer that overlies the basal plate. Passage of the fibres result in the enlargement of the marginal layer forming the Pons. In addition to the fibres, pontine nuclei that are formed from the alar region of the rhombencephalon migrate in the marginal layer into the Pons. The basal plate neuroblast differentiate into neurons that are arranged into 3 motor nuclei mediolaterally;

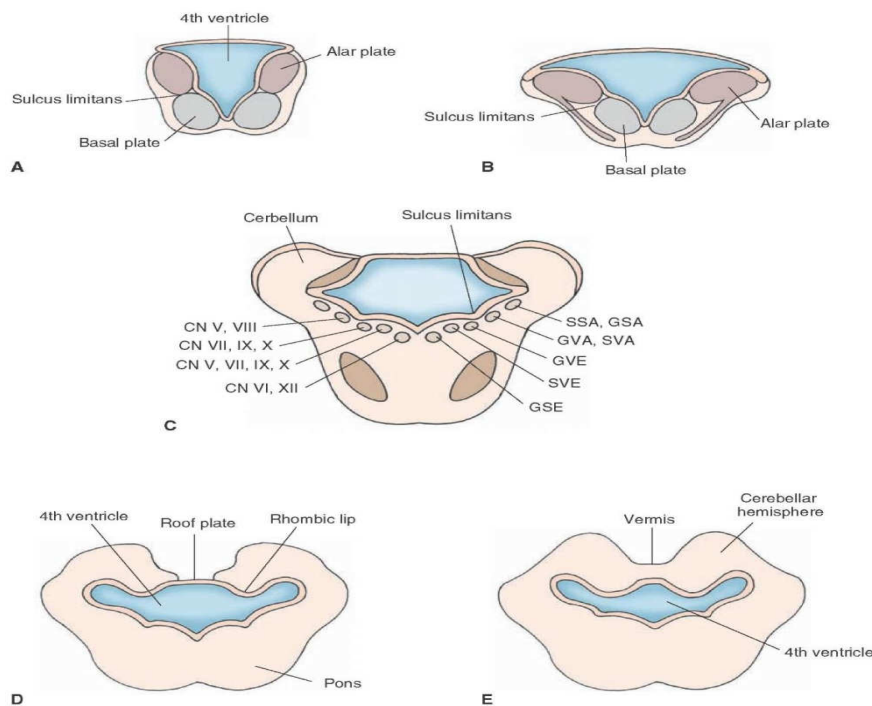
Somatic efferent; CN6 for extraocular mm,

Special visceral efferent; CN5, CN7 for the 1st and 2nd pharyngeal arches.

General visceral efferent; CN9 secretomotor supply submandibular and sublingual glands

The alar plate of the metencephalon contains 3 groups of nuclei that subserve sensory modalities;

Somatic afferent; CN5, CN8 complex



The dorsolateral part of the marginal layer over the alar plate differentiates into the Rhombic lip.

The rhombic lip is wide apart caudally but closes cranially. Pontine flexure ensures the folding of the metencephalon on itself while the rhombic lip differentiates into the cerebellar plates. At 12 weeks, a central vermis and 2 lateral dilated cerebellar hemisphere are identified. A transverse fissure separates the cerebellar plate into an upper and lower parts. The fissure separate the vermis from the Nodule while the hemisphere is separated from the Flocculus. The developing cerebellum consist of 3 layers viz; neuroepithelial, mantle and marginal. The external granular layer differentiates from the neuroepithelial layer. The EGL give rise to granule, purkinje, golgi II neuron cells by the 6th month, while the other cells (basket and stellate) are derived from the marginal layer

Myelencephalon

This is the caudal portion of the rhombencephalon. The dorso-lateral wall is opened up, basal and alar plate are still identifiable separated by the sulcus limitans. 3 set of motor nuclei are identified in the basal plate;

Somatic efferent; forms a motor column continuous cranio-caudally, CN12, (tongue) CN6, CN4, CN3 (eye) at different levels.

Special visceral efferent; continues cranially as a motor column CN11, CN10, CN9 supplying the striated muscles of the soft palate and larynx, pharynx, heart and gut.

General visceral efferent; supply the involuntary muscle of the gut, heart and respiratory system.

The alar plate have similar arrangement;

Somatic afferent; CN8, CN5 for the ear and head surface.

Special visceral afferent; CN9 for the tongue taste bud.

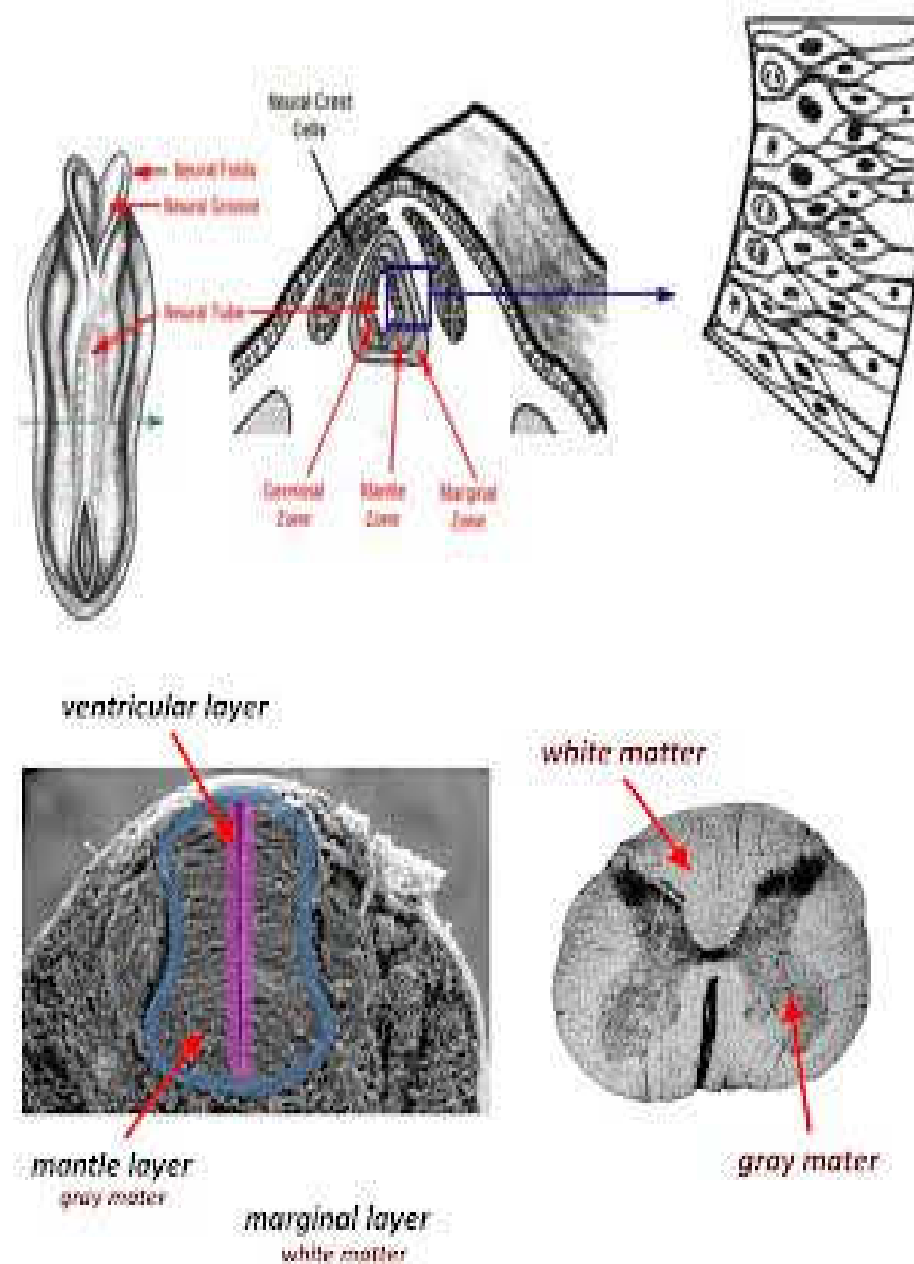
The ependymal lining of the roof plate alongside the vascular mesenchyme form the choroid plexus that produces CSF.

NEUROEPITHELIAL CELLS

The neuroepithelial cells lining the neural tube are of simple columnar type.

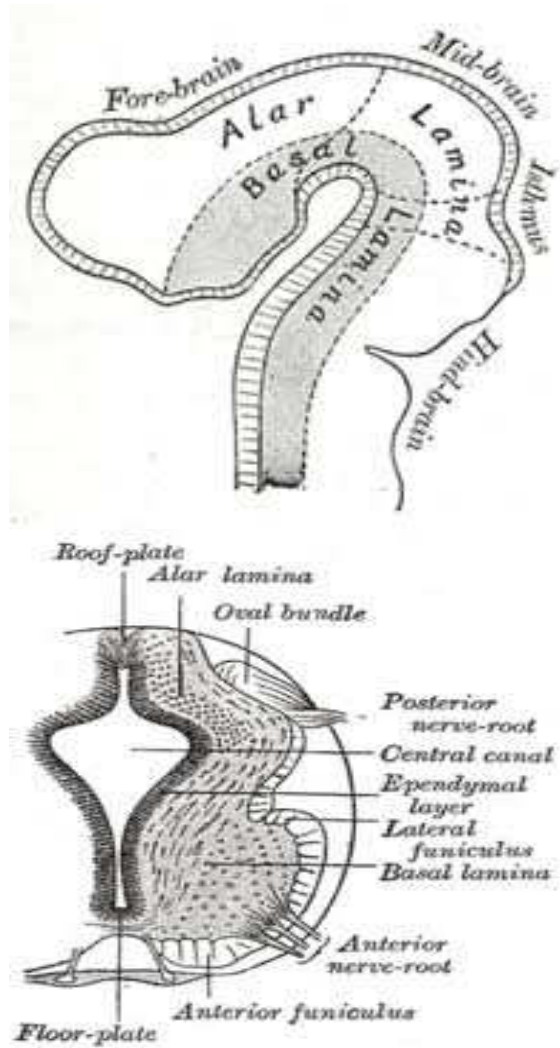
Increase cellular proliferation after closure of tube, giving rise to neuroblast (large deeply stained nucleus and pale cytoplasm). The neuroblast gives rise to the 2nd layer, mantle layer which differentiate

into the gray matter. The axons of the mantle layer neurons coalesce in the 3rd layer (outermost) forming the marginal layer.



The mantle layer differentiates into a basal and alar plate. The roof and floor plate are usually devoid of neuroblast, have only the ependymal layer.

The ependymal layer is the final derivative of the neuroepithelial layer. The dorsal Alar plate houses neurons that subserve sensory modalities, while the ventral basal plate houses neurons that subserve motor modalities.



NEUROHISTOLOGY

Neuroblast; derived from neuroepithelial cell, differentiate through uni, bi to multipolar neurons. Neurons may take different shapes and size depending on function.

Glial cell; produced by the neuroepithelial cells following the production of neurons. Differentiate into astrocyte in the mantle layer and oligodendrocyte in the marginal layer.

Astrocytes are star shaped, mainly protoplasmic in the mantle layer and fibrous in the marginal layer. It is important for the formation of the blood brain barrier.

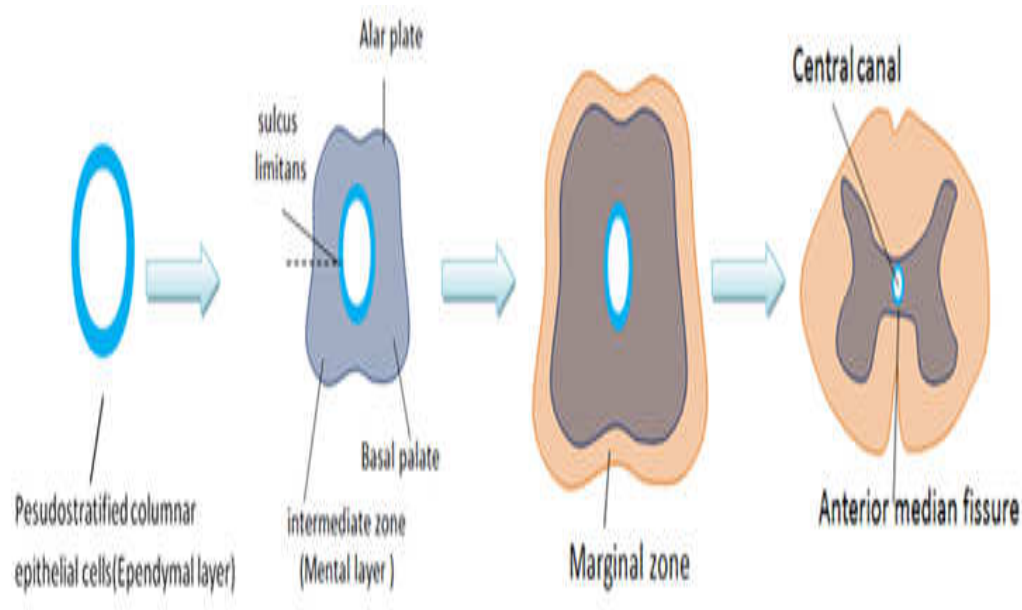
Oligodendrocytes have small round deeply stained nucleus usually with a halo in routine stain, appear in pair or string like formation. It is responsible for myelination in the central nervous system. Myelination of the white matter begins in the 2nd trimester and continues postnatally.

Microglia are of mesenchymal origin, invade the CNS in the latter half of gestation. Usually cigar shaped with fine processes. They are phagocytic cells

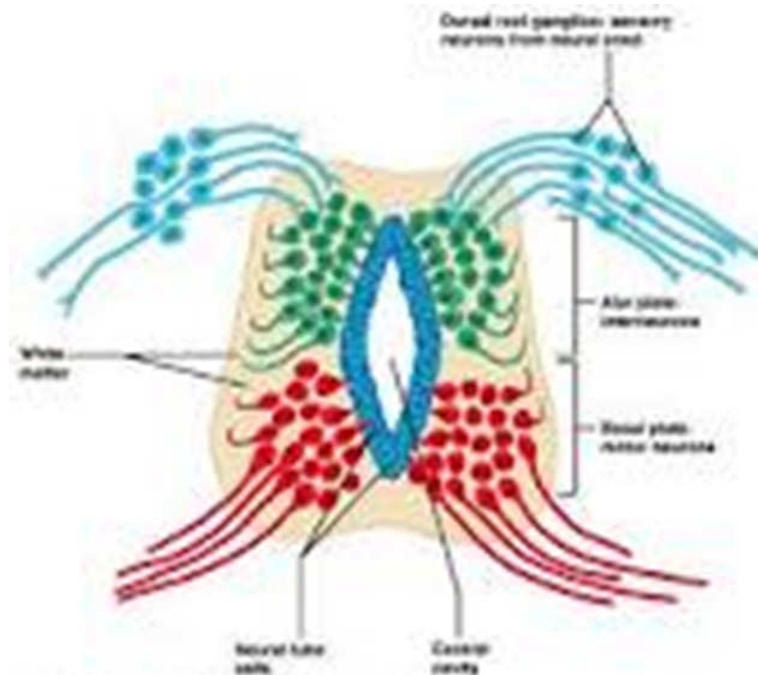
Ependymal cells are the final derivative of neuroepithelial cells. They line the ventricle and central canal. They are cuboidal and low columnar in shape.

SPINAL CORD

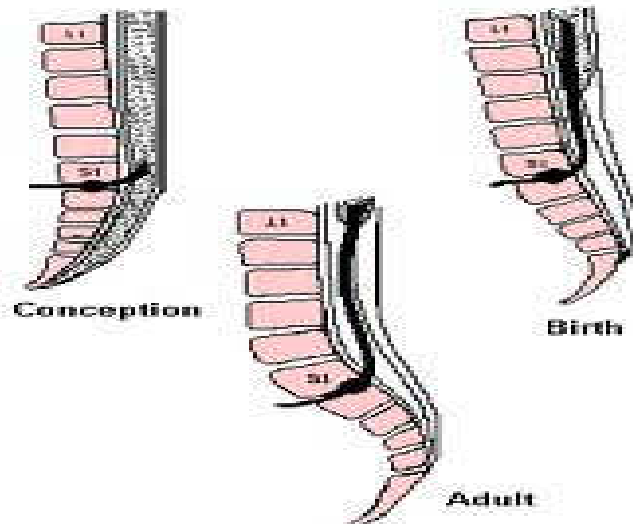
The mantle layer gives rise to the gray matter of the spinal cord, while the marginal layer forms the white matter of the spinal cord. Ventral basal plate subserves motor function while dorsal alar plate subserves sensory functions. Neuroblast aggregation in between the basal and alar plate for the intermediate horn in some part of the developing spinal cord. Intermediate horn neuroblast differentiates into the neurons of the sympathetic portion of the autonomic nervous system. Intermediate horn is limited to the region of T1 to T12 and L2-L3.



The axons of the basal plate neuron coalesce exiting through the marginal layer to form the ventral motor root of the corresponding spinal nerve about the 4th week. Dorsal root ganglia are derivatives of the neural crest cells, its neuroblast differentiate into bipolar neurons. The medial process enters the marginal layer of the alar plate of the SC ending in the dorsal horn or ascending to higher centers. The other process that grows peripherally coalesces with the ventral motor root to form the spinal nerve trunk. Myelination is a function of the Schwann cells in the axons outside the SC while the portion within the SC is myelinated by oligodendrocyte.



The rate of elongation of the spinal cord and the vertebral column is not the same. The vertebral column outpaces the spinal cord resulting in an apparent ascent of the developing spinal cord. At about 3rd month of life, spinal cord extends the entire length of the vertebral column, at birth spinal cord ends at the lower border of L3 while at adulthood it ends at L2/L3 boundary. Implication is the obliquity of spinal nerve forming cauda equine and the formation of filum terminale.



NEURONAL MIGRATION

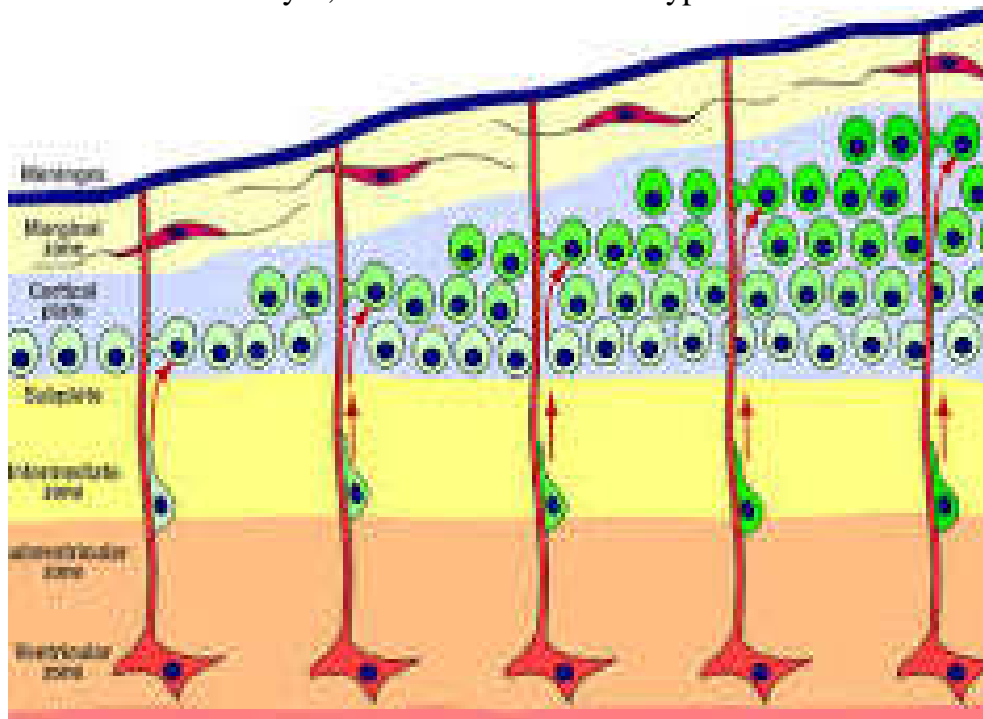
The neuroblast of the cerebral cortex proliferates rapidly around the ventricle, forming neurons and supporting (glia) cells. The glial cells differentiate into the oligodendrocyte and astrocyte.

The neurons that are formed from the neuroblast are largely of 2 types; viz granular (sensory cortex) and pyramidal (motor). Different sizes and

composition of neurons at each point confer a stratified arrangement on the cerebral cortex. Each layer consists of neurons of specified composition and subserves similar functions. Movement of forming neurons from the point of neuroblast proliferation to their definitive position is mediated by internal and external cues.

Mechanism of migration is either radial or tangential means. While radial migration means is mainly employed by the granular and pyramidal neurons, the tangential mean is employed by interneurons. Settling of the neuron in their definitive position is in an inside-out fashion. Migration results in 6-layered cortical arrangement.

- Molecular; few oligodendrocyte almost no neuron.
- External granular; more of round granular neurons.
- External pyramidal; contains small pyramidal neurons.
- Internal granular; contain granular cells that are closely packed.
- Internal pyramidal; contain medium to large pyramidal cells, home to Betz cells.
- Multiform layer; contains different cell type.



Clinical correlates

Anencephaly - this is the absence of a major portion of the brain, skull and scalp that occurs during embryonic r. It is a cephalic disorder that results from a neural tube defect that occurs when the rostral(head) end of the neural tube fails to close, usually between the 23rd and 26th day of conception. With very few exceptions, infants with this disorder do not survive longer than a few hours or possibly days after birth.

Spina bifida - this is a developmental congenital disorder caused by the incomplete closing of the embryonic neural tube. Some vertebrae overlying the spinal cord are not fully formed and remain unfused and open. If the opening is large enough, this allows a portion of the spinal cord to protrude through the opening in the bones. There may or may not be a fluid – filled sac surrounding the spinal cord. Spinal bifida malformations fall into three categories: spina bifida occulta, spina bifida cystic with meningocele, and spina bifida cystic with myelomeningocele. The most common location of the malformations is the lumbar and sacral areas. Spina bifida can be surgically closed after birth, but this does not restore normal function to the affected part of the spinal cord.

4.0 SUMMARY

- The CNS is an ectodermal derivative that begins to differentiate in the 3rd intrauterine week in humans.
- The first indication of the nervous system is within the neural plate, a thickened area of the ectoderm that forms after the ovum has been fertilised in the third week of gestation.
- The neural plate thickens rostral to the primitive node and midline depression in the neural plate forms the neural groove. Edges of the neural groove rise on both sides of the midline, fusing to convert the groove to a tube. The neural tube is the beginning of the brain and spinal cord. As the neural tube separates from the surface ectoderm cells, the neural folds form the neural crest. Ganglia of the spine, cranial and autonomic nervous system develop from the neural crest.
- Anomalies of the closure of the neural tube results in several clinical conditions such as anencephaly and spinal bifida.
- The 4th week witnesses the differentiation of the cranial part of the neural tube into the 3 brain vesicles which leads to the formation of the prosencephalon, mesencephalon and rhombencephalon cranio-caudally. Same mechanism results in the formation of 2 flexures, cervical and cephalic flexures. Cephalic flexure opens to the ventral surface while the cervical opens to the dorsal surface.
- By the 5th week of development, the Prosencephalon is divided into the Telencephalon and Diencephalon by differential growth rate. The Rhombencephalon also divides into the cranial Metencephalon and caudal Myelencephalon roughly by the cephalic (Pontine) flexure.
- The neuroblast of the cerebral cortex proliferates rapidly around the ventricle, forming neurons and supporting (glia) cells. The glial cells differentiate into the oligodendrocyte and astrocyte.

SELF – ASSESSMENT EXERCISE

- i. What is the neural plate?
- ii. What are the differences between the right and left hemispheres of the brain?
- iii. What are the primitive general functional areas of the brain?
- iv. When do the fontanelles of the skull close?

UNIT 2 CEREBRAL HEMISPHERE

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 4.0 Summary

1.0 INTRODUCTION

The cerebrum is the part of the brain that most people think of when the term *brain* is mentioned. The cerebrum accounts for the largest portion of total brain weight, which is about 1200 g in females and 1400 g in males. Brain size is related to body size; larger brains are associated with larger bodies, not with greater intelligence.

The cerebrum is divided into left and right hemispheres by a longitudinal fissure. The most conspicuous features on the surface of each hemisphere are numerous folds called gyri, which greatly increase the surface area of the cortex. The grooves between the gyri are called sulci. The central sulcus, which extends across the lateral surface of the cerebrum from superior to inferior, is located about midway along the length of the brain. The general pattern of the gyri is similar in all normal human brains, but some variation exists between individuals and even between the two hemispheres of the same cerebrum.

Each cerebral hemisphere is divided into lobes, which are named for the skull bones overlying each one.

2.0 OBJECTIVES

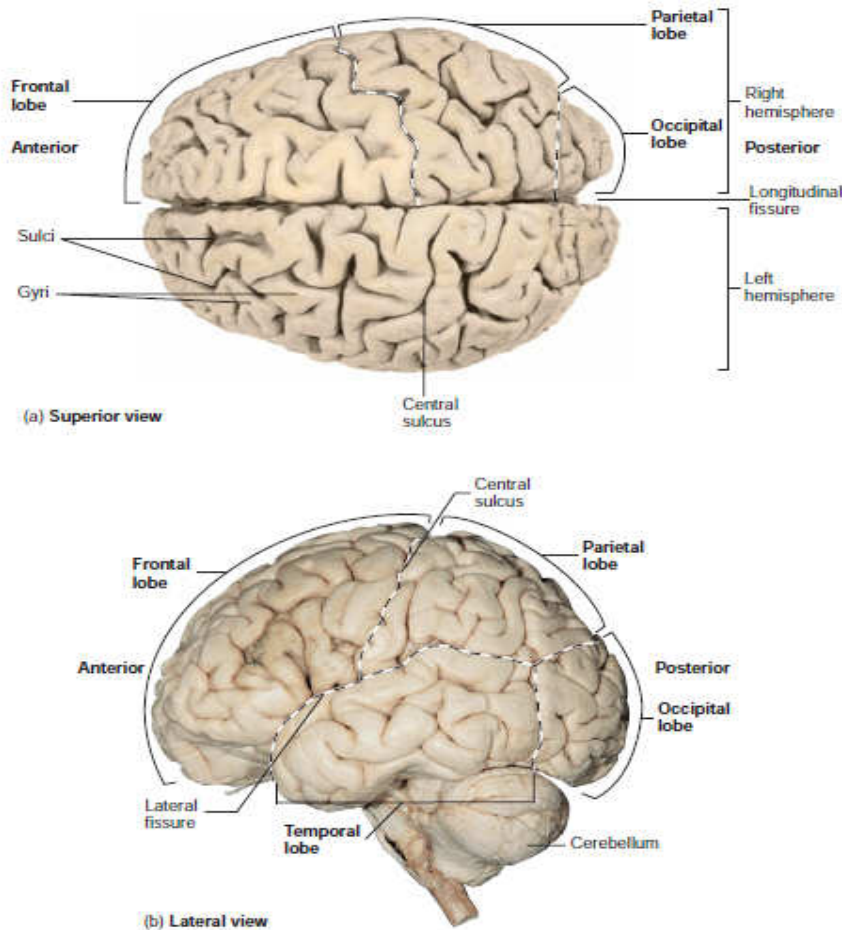
At the end of this unit, you will be able to:

- describe the different regions of the cerebral hemisphere.
- identify the boundaries of the different regions of the cerebral hemisphere.
- identify the different gyri and sulci as well as the functional areas of the cerebrum.

Superolateral surface

Each lobe consists of gyri and sulci that have specific name. 3 main sulci delineate the boundary of the lobes, viz; central, lateral and parieto-occipital. Lateral sulcus (fissure of Sylvius) is the sulcus that separates the frontal from the temporal lobe. The anterior end of the sulcus penetrates (anterior and ascending rami) the inferior frontal gyrus

dividing it into the Orbital, Triangularis and Operculum parts. Central sulcus; (fissure of Rolando) runs on the superolateral surface of the cerebrum just caudal to the antero-posterior midpoint. It runs downward and forward stopping short of the lateral sulcus. It separates the frontal from the parietal lobe. The precentral gyrus is just anterior to it, while the postcentral gyrus is just behind it.



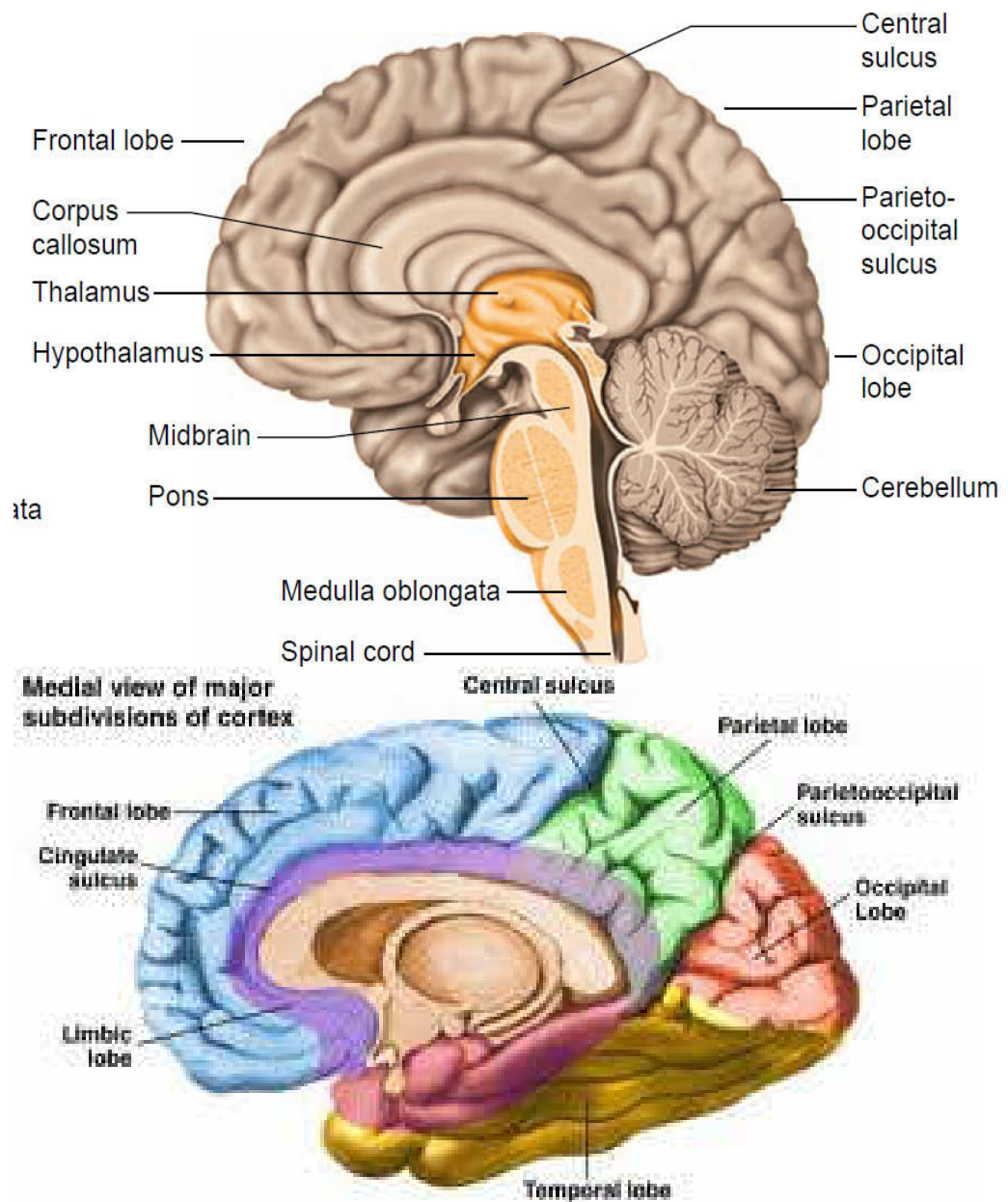
Medial surface

On the medial surface of the cerebral hemisphere, the corpus callosum stands out as a collection of white matter that connects the 2 hemisphere. The cingulate gyrus lies above the corpus callosum while the medial and superior frontal gyri lie above it separated by the cingulate sulcus.

The paracentral lobule caps the medial end of the central sulcus. The calcarine and parieto-occipital sulci form a Y shaped furrow lying on its side facing posteriorly. The precuneus gyrus lies above the upper arm and limb of the 'Y'. The cuneus gyrus lies in between the 2 arms of the 'Y'. The lingual gyrus lies below the 'Y' at the inferio-medial border of the cerebrum. The inferior surface revealed the rectus gyrus, olfactory tract/bulb and orbital gyrus and sulcus from medial to lateral.

Parahippocampal gyrus lies medially on the inferior surface of temporal lobe terminate in the uncus

The brain stem is the other major structure on the inferior surface of an intact brain.



Cortical area and structure

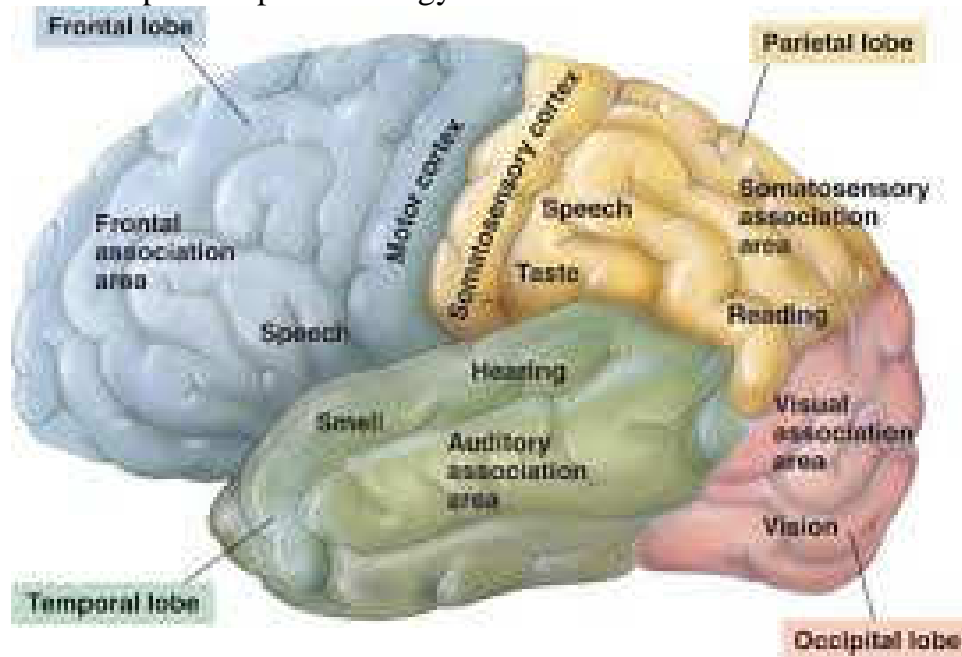
The primary motor sensory area corresponds with the precentral gyrus and frontal lobe (Area 4 and 6). Afferent fibers exit via the corticonuclear and pyramidal bundles, modulated by the Thalamus and Cerebellum. The supplementary motor area corresponds with part of the medial surface of the frontal lobe (Area 6 and 8). The primary sensorimotor area corresponds with the postcentral gyrus and the

adjoining medial surface of the parietal lobe (Area 3, 2 and 1). Subserve touch, kinaesthetic and vibration senses.

The inferior portion of postcentral gyrus corresponds with the supplementary sensory area (Area 40 and 43), subserve pain and temperature modalities. Anterior Motor speech area (Broca) corresponds to the inferior orbital gyrus of the dominant hemisphere and the pars triangularis (Area 44 and 45). Posterior speech area (Wernicke) is the posterior end of the superior and middle temporal gyri and the lower part of the parietal lobe.

Frontal eye field (Area 6, 8 and 9) is the centre of the middle frontal gyrus, it subserves the voluntary eye movement and accommodation functions.

Olfactory area corresponds with the Uncus and adjoining area. Visual area (Area 17) is the medial surface of occipital lobe, posterior lip of calcarine sulcus as far as the occipital pole. The striate cortex is the association visual area (Area 18 and 19). Auditory area (Area 41 and 42) is the floor of the lateral sulcus and the superior temporal gyrus, the surrounding Area 22 is the association area. Gustatory area is the inferior lip of the postcentral gyrus.



CORTICAL STRUCTURE

The cerebral cortex consists of aggregates of cells in definite pattern.

The assembly of cells is along morphological and functional lines.

Over 90% of the cortex has a 6 layered arrangement.

Layer I; plexiform or molecular layer, few cells mostly glia.

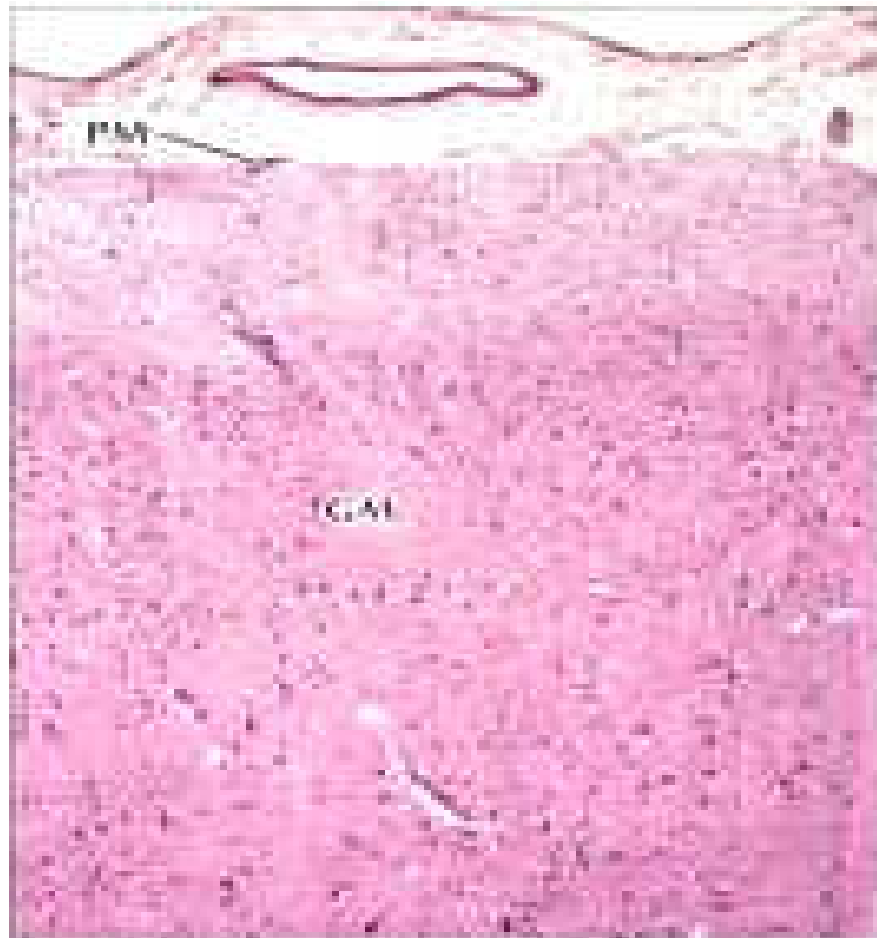
Layer II; external granular layer, cells predominantly small granular type
Layer III; external pyramidal, predominantly pyramidal cells.

Layer IV; internal granular, consist of medium to large size granular cells.

Layer V; internal pyramidal, pyramidal cells are medium to large size, giant pyramidal cells (Betz) characterise this layer

Layer VI; multiform layer, different cell types are found in the layer.

The ratio of granular to pyramidal cells in any layer is dependent on the modality it subserves



Cerebral white matter

The white matter of the cerebrum is made up of fibres that can be grouped into 3 viz;

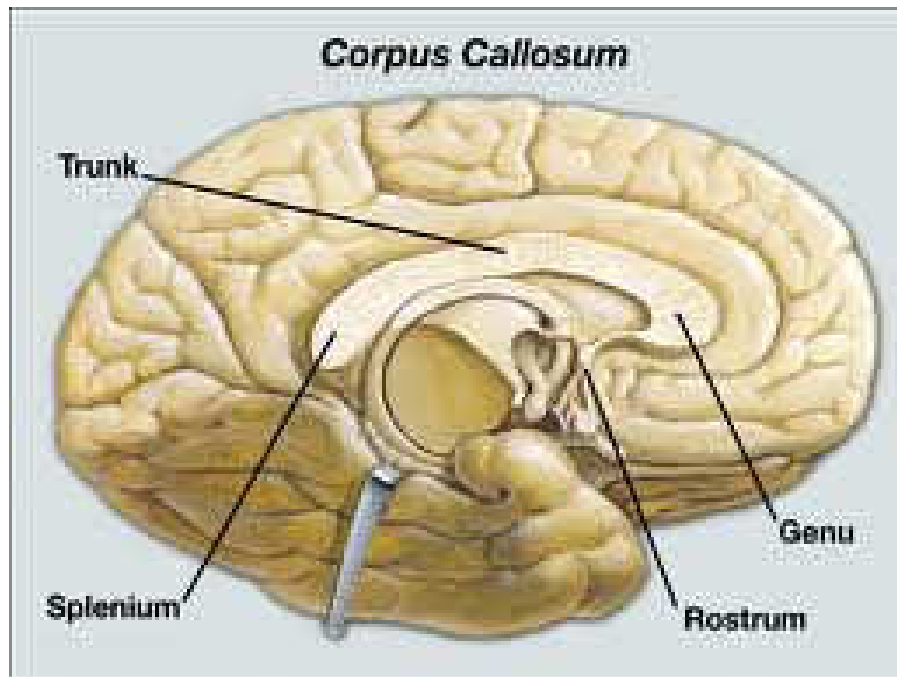
Commissural fibres; these fibres connect the 2 cerebral hemispheres in a symmetrical fashion. They coalesce to form the corpus callosum, anterior and posterior commissures.

Association fibres; they connect different parts of the same cerebral hemisphere.

Projection fibres; they connect the grey matter of the cerebral hemisphere to subcortical nuclei in the brain stem, spinal cord etc.

Commissural fibre; The corpus callosum is the most prominent commissural fibres collection.

It provides asymmetrical connection between the hemispheres. From its commencement at the anterior commissure it thickens posteriorly. It has 4 parts viz, Rostrum, Genu, Body, Splenium.



Forceps minor is formed by the laterally extending fibers of the genu, while that of the splenium form the Forceps major. Forceps minor connects the frontal cortex while Forceps major connects the occipital cortex. The laterally extending fibers of the remaining parts of the corpus callosum form a sheet of white matter that form the roof and lateral walls of the different parts of the lateral ventricles

Projection fibres

The Internal capsule is the most prominent projection fibre. Consist of afferent fibres from thalamic nuclei and efferent fibres that travel through the cerebral peduncle of midbrain.

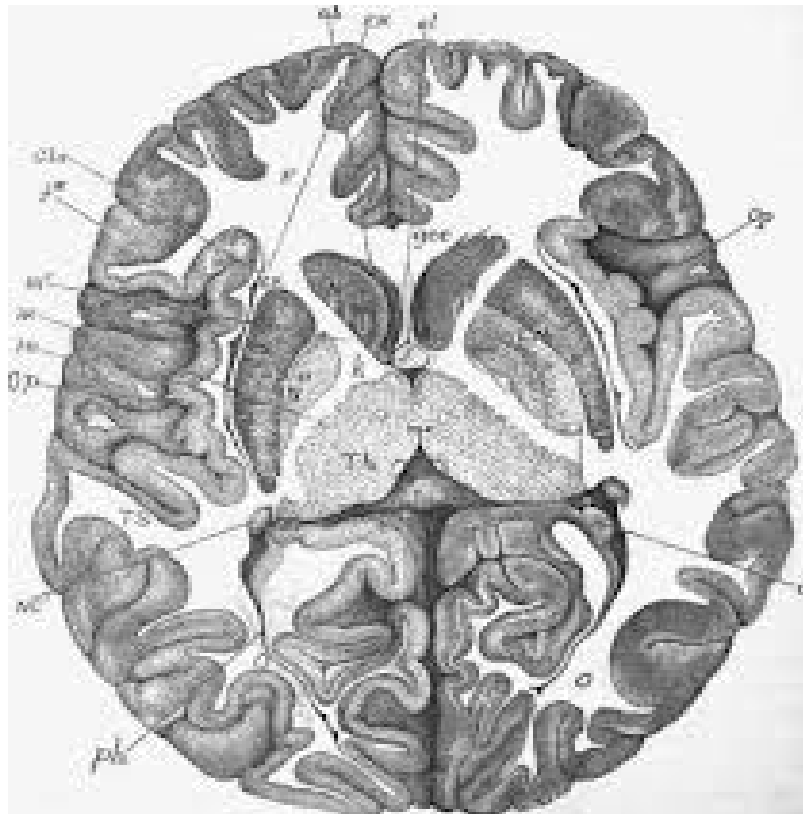
It is separated from the cavity of the lateral ventricle by the caudate nucleus. The globus pallidus of the lentiform nucleus indents its lateral edge conferring a lateral concavity on it. It has an anterior limb, a posterior limb connected by the genu. Retrolentiform and sublentiform parts are posteriorly located.

The Anterior Limb lies between the head of caudate (medially) and the lentiform (laterally) nuclei. Contain frontopontine fibres that travel from cell bodies in the frontal part of the cortex to the pontine nuclei on which they arborize. Passing through the thalamus they occupy the medial part of the base of the cerebral peduncle. Fibres from the frontal eye field to the oculomotor nucleus are suspected to pass through it.

Genu is the bent portion of the internal capsule. Contain corticonuclear fibres from the cerebral cortex to the motor nuclei of cranial nerves in the brain stem.

Posterior limb is located between the Thalamus (medially) and the lentiform nucleus (laterally).

The anterior 2/3rd contains corticospinal fibers that travel from the cortical cell bodies to the anterior horn cells of the spinal cord. The fibers pass through the brainstem to the lower part of the medulla where most of them decussate to form the lateral corticospinal tract. Haemorrhage or thrombosis of the striate artery around the posterior fiber results in paralysis of the opposite skeletal muscles. Fibers of the Broca's speech are affected on the left side. Posterior 1/3rd of the posterior limb contain Thalamocortical fibers from thalamus to sensory cortex.



Retrolentiform part; located in the posterior end of lentiform nucleus, contains the corticopontine fibres which occupy the lateral part of the cerebral peduncle as well as optic radiations.

Sublentiform part is located below the posterior end of lentiform nucleus, contains auditory radiations from MGB to the auditory area of cortex.

Clinical correlates

- ❖ Head injuries are classified as open or closed. In an open injury the cranial cavity contents are exposed to the outside, whereas in a closed injury the cranial cavity remains intact. Closed injuries, which are more common than open injuries, involve the head striking a hard surface or an object striking the head. Such injuries may result in brain trauma.
- ❖ The most common type of traumatic brain injury (75%–90%) is a concussion, which is an immediate, but transient, impairment of neural function, such as a loss of consciousness or blurred vision. In some cases, postconcussion syndrome occurs a short time after the injury. The syndrome includes increased muscle tension or migraine headaches, reduced alcohol tolerance, difficulty in learning new things, reduction in creativity, changes in motivation, fatigue, and personality changes. The symptoms may be gone in a month or may persist for as long as a year.
- ❖ A hematoma is a localized mass of blood released from blood vessels but confined within an organ or a space. An accumulated mass of blood can apply pressure to the brain, causing damage to brain tissue. Blood is toxic to brain tissue. Hematomas of the brain are classified according to where the bleeding occurs. Epidural hematomas result from an accumulation of blood in the epidural space and occur in about 1%–2% of major head injuries. Subdural hematomas result from an accumulation of blood in the subdural space and occur in 10%–20% of major head injuries.

4.0 SUMMARY

- The cortex of the cerebrum is folded into ridges called gyri and grooves called sulci, or fissures. The longitudinal fissure divides the cerebrum into left and right hemispheres. Each hemisphere has five lobes.
- The frontal lobes are involved in voluntary motor function, motivation, aggression, the sense of smell, and mood. The parietal lobes contain the major sensory areas receiving sensory input, such as touch, pain, temperature, balance, and taste. The

occipital lobes contain the visual centers. The temporal lobes evaluate smell and hearing input and are involved in memory, abstract thought, and judgment.

- Gray matter forms the cortex and nuclei of the cerebrum. White matter forms the cerebral medulla, which consists of three types of tracts (Association, Commissural and Projection fibers) connect cerebrum to other parts of the brain and the spinal cord.

Activity

In the gross anatomy laboratory, locate and identify the following anatomical features on the human brain models:

- Cerebral hemispheres, gyri, sulci, lobes, olfactory bulbs, optic nerves, optic chiasma and corpus callosum

SELF –ASSESSMENT EXERCISE

1. Which of these cerebral lobes is important in voluntary motor function, motivation, aggression, sense of smell, and mood?
 - a. frontal
 - b. insula
 - c. occipital
 - d. parietal
 - e. temporal
2. Fibers that connect areas of the cerebral cortex within the same hemisphere are:
 - a. projection fibers.
 - b. commissural fibers.
 - c. association fibers.
 - d. all of the above.
3. The basal nuclei are located in the
 - a. inferior cerebrum.
 - b. diencephalon.
 - c. midbrain.
 - d. all of the above.
4. Describe the two layers of the cerebrum. Why is the outer layer convoluted?
5. Distinguish between a sulcus and a fissure.

UNIT 3 BRAIN STEM

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Cranial nerve nuclei
- 4.0 Summary

1.0 INTRODUCTION

The Brain Stem is a structural connection between the diencephalon and the spinal cord. The **brainstem** consists of the medulla oblongata, pons, and midbrain. It connects the spinal cord and cerebellum to the remainder of the brain, and 10 of the 12 pairs of cranial nerves arise from it. In general, the posterior part of the brainstem contains ascending tracts from the spinal cord, cerebellum, and cranial nerves, whereas the anterior part of the brainstem contains descending tracts involved with motor control. The brainstem contains several nuclei involved in vital body functions, such as the control of heart rate, blood pressure, and breathing.

The caudal portion of the brain stem transit into the spinal cord at the level of the foramen magnum. It contains ascending and descending fibers interspersed with cellular aggregate known as the nuclei. The corticospinal (motor), posterior column (fine touch, vibration sense and proprioception) and spinothalamic (crude touch, temperature and pain).

2.0 OBJECTIVES

At the end of this unit, you will be able to:

- to describe the anatomy of the different parts of the brain stem.
- to identify the characteristic features of the midbrain, pons and medulla oblongata.

3.0 MAIN CONTENT

3.1 Cranial nerve nuclei

The cranial nerve nuclei 3-12 are located in the brain stem in addition to other anatomically distinct nuclei.

The midbrain contains CN 3, 4

The Pons contains CN 5, 6, 7

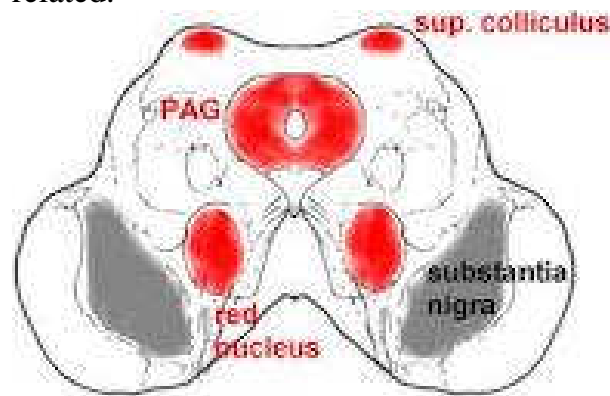
The Medulla contains CN 9, 10, 11, 12

The CN 8 is located in the pontomedullary junction. Diffuse cellular arrangements are also located in the brain stem forming the Reticular system. The reticular formation is related to several vital centers. The sensory part of the CN 5 spans the whole of the brain stem as far as the upper spinal cord. All cranial nerves emerge from the ventral surface of the brain stem except the Trochlear.

Mid-brain

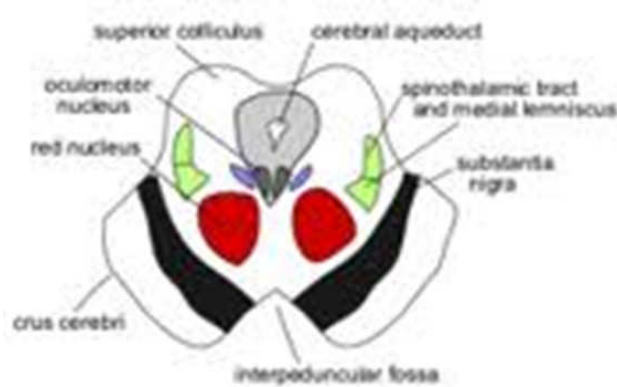
It is the shortest part of the brainstem measuring about 2cm. Ventrally it is characterized by a V shaped crura that converge on the superior surface of the pons. Dorsally, the superior and inferior colliculi form 2 pairs of round elevation. The superior cerebellar peduncle connects the cerebellum to the dorsal surface of the midbrain caudal to the inferior colliculus.

The medial geniculate body and lateral geniculate body are lateral to the superior colliculus, while the pineal gland and pulvinar are superiorly related.



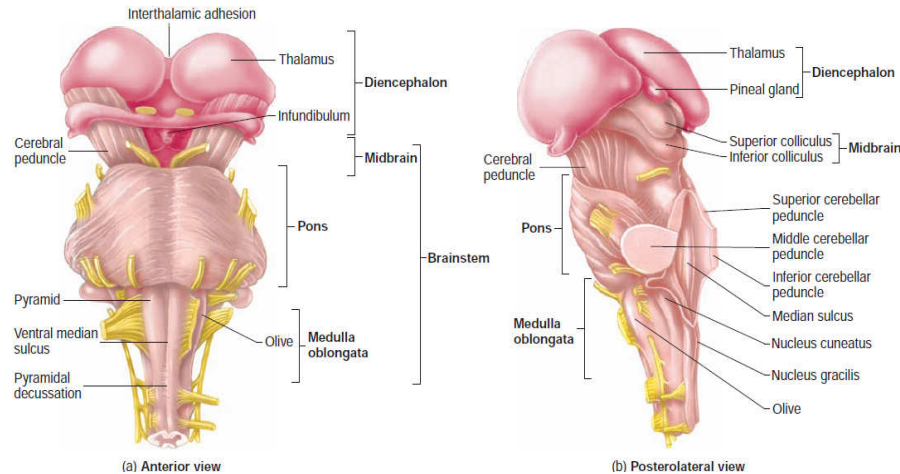
- ❖ The CN 3 emerges from the ventral surface on the medial side of the cerebral peduncle, while the CN 4 courses ventrally from its point of emergence on the dorsal surface inferior to the inferior colliculus.
- ❖ The 2 CN run between the superior cerebellar and posterior cerebral arteries which provide blood supply for the midbrain.
- ❖ It is ventro-dorsally divisible into the base, tegmentum and tectum.
- ❖ Histologically the superior and inferior colliculi provide a logical reference point.
- ❖ Superior colliculus level; the oculomotor nucleus is close to midline with the Edinger-Westphal nucleus cranial to it. The red nucleus is lateral and ventral to CN 3, the axons of the CN pass through it. It receives input from the contralateral cerebellar hemisphere (dentate nucleus) via the superior cerebellar peduncle.

- ❖ Inferior colliculus level; the nucleus of CN 4 lie ventral to the aqueduct, its fibers run dorsally decussating dorsal to the aqueduct.



- ❖ The Substantia Nigra forms the boundary between the crus and the tegmentum. Its cells contain melatonin giving rise to nigrostriatal fibers.
- ❖ Fibers project to the other basal ganglia nuclei. Loss of dopamine producing cells explains Parkinsonism. Mesencephalic nucleus of CN 5 lies in the central gray matter, while spinal and medial lemnisci lie in the lateral region of tegmentum.
- ❖ The reticular formation is located between the medial lemniscus and the central gray matter.
- ❖ The crura contain corticospinal and corticonuclear fibers in the middle 2/3rd portion while frontopontine and temporopontine fibers occupy the medial and lateral 1/6th.
- ❖ The tectum is occupied by the superior and inferior colliculi with inputs from the retina and cochlea respectively.
- ❖ Efferent fibers travel in the tectospinal and tectobulbar tracts for general light and sound reflexes
- ❖ The pretectal nucleus is located at the junction between the midbrain and the diencephalon subserving the pupillary light reflexes.

3.3.4 Pons



The parts of the brainstem

The pons forms a bridge between the midbrain, medulla and the cerebellum. It contains the respiratory center. Its transversely running fibers coalesce to form the middle cerebellar peduncle. The motor and sensory part of the CN 5 emerges from its ventral surface just lateral to the midline.

A midline groove houses the basilar artery, while the superior cerebellar artery curves around its boundary with the midbrain. At the pontomedullary junction CN 6 emerges at the midline while CN 7 and 8 emerge more laterally. The cerebellum covers the dorsal surface of the pons with the pontine part of the 4th ventricle in between them. The central aqueduct opens directly into the 4th ventricle. The superior medullary velum (anchored on SCP) intervenes as the roof between the ventricle and the cerebellum. The upper part and lower part of the pons are distinguishable by the presence or absence of opened 4th ventricle. The ventral part of the pons has pontine nuclei with corticopontine fibers passing through them.

Upper pons; the motor nucleus of CN 5 is located on the lateral part of the floor of 4th ventricle, while the main sensory nucleus (touch) is lateral to it. The sensory nucleus is continuous with the mesencephalic nucleus (proprioception) in the midbrain.

Lower pons; the nucleus of CN 6 lie on the floor of the 4th ventricle near midline while that of CN 7 lies ventral and lateral to it. Fibers of CN 7 course dorsal to the nucleus of CN 6 raising a facial colliculus on the floor of the 4th ventricle. The nucleus of CN 8 lie on the floor of the 4th ventricle lateral to the CN 7 extending to the medulla, the vestibular portion is more medial.

Vestibular part receives fibers from the internal acoustic meatus, while the cochlea part receives fibers from the spiral ganglia of cochlea. Both travel in the inferior cerebellar peduncle.

Blood supply is from the basilar artery superior cerebellar and anterior inferior cerebellar arteries.

Medulla oblongata

This is the portion of the brain stem that connects the pons to the spinal cord. It is continuous with the spinal cord at the level of foramen magnum. Dorsally the upper part is covered by the cerebellum with the caudal part of the 4th ventricle intervening. This portion of the 4th ventricle is roofed by a layer of ependyma and pia matter.

The caudal part of the dorsal surface presents 2 elevations namely the gracile and cuneate tubercle secondary to underlying nuclei medio-laterally. Ventrally, the upper part is grooved in the midline, the pyramid and olive are elevations lateral to the midline. The pyramid is secondary to the corticospinal fibers while the olive is secondary to the inferior olivary nucleus. The most lateral elevation forms the lateral boundary, the inferior cerebellar peduncle.

The CN 6, 7 and 8 emerge from the pontomedullary junction medio-laterally. The CN 9, 10 and 11 emerge as rootlets lateral to olive and CN 12 between pyramid and olive. 3 levels of medulla are identifiable, the open part, the upper closed part and the lower closed part.

Open part; the floor of the 4th ventricle is the site of most cranial nerve nuclei. The hypoglossal nucleus is located adjacent to the midline beneath the hypoglossal trigone. The nucleus of the vagus is located lateral to the hypoglossal beneath the vagal trigone. It controls the cardiac and visceral muscle and glandular secretions. The nucleus of tractus solitarius lie lateral to that of vagal, its cranial part receives fibers from chorda tympani, glossopharyngeal and internal laryngeal branch of vagus.

The nucleus ambiguus lies lateral containing motor cell bodies for larynx, soft palate, pharynx and upper oesophagus skeletal muscles. The inferior olivary nucleus is a crenated C-shaped strip of gray matter with its open end facing the opposite cerebellar peduncle.

Upper closed part; it shows the central decussation of the fibers that form the medial lemniscus (fibers of gracile and cuneate nuclei).

Lower closed part; it is characterized by the pyramidal decussation. Centre for the control of reflex activities such as coughing, swallowing and vomiting. Centers for the control of respiratory, cardiac and motor function are also located in the medulla.

Blood supply of the medulla: anterior spinal branch of vertebral artery (medial medullary syndrome).Posterior inferior cerebellar artery (lateral medullary syndrome).

Clinical correlates

Parkinsonism: this is a neurological syndrome characterized by tremor, hypokinesia, rigidity and postural instability. The neurodegenerative condition Parkinson's disease is the most common cause of parkinsonism. The motor symptoms of this disease result from the death of dopamine – generating cells in the substantia nigra. Modern treatments are effective at managing the early motor symptoms of the disease, these include levodopa and dopamine agonists.

Brainstem stroke syndromes: there are a large number of well defined brainstem stroke syndromes, most of which involve ischemia in the distribution of the basilar or vertebral arteries. Magnetic resonance imaging(MRI) is frequently needed to make a specific diagnosis.

4.0 SUMMARY

- The midbrain is superior to the pons. The corpora quadrigemina consists of four colliculi. The two inferior colliculi are involved in hearing and the two superior colliculi in visual reflexes.
- The substantia nigra and red nucleus help regulate body movements. The cerebral peduncles are the major descending motor pathway
- The pons is superior to the medulla. Ascending and descending tracts pass through the pons. The pons connects the cerebrum and the cerebellum. Pontine nuclei regulate breathing, swallowing, balance, chewing, and salivation.
- The medulla oblongata is continuous with the spinal cord and contains ascending and descending tracts. Medullary nuclei regulate the heart, blood vessels, breathing, swallowing, vomiting, coughing, sneezing, hiccuping, balance, and coordination. The pyramids are tracts controlling voluntary muscle movement.

Activity

In the gross anatomy laboratory, identify the components of the brain stem in the human brain model provided.

SELF – ASSESSMENT EXERCISE

1. Important centers for heart rate, blood pressure, breathing, swallowing, coughing, and vomiting are located in the
 - a. cerebrum.
 - b. medulla oblongata.
 - c. midbrain.
 - d. pons.
 - e. cerebellum.
2. In which of these parts of the brain does decussation of descending tracts involved in the conscious control of skeletal muscles occur?
 - a. cerebrum
 - b. diencephalon
 - c. midbrain
 - d. pons
 - e. medulla oblongata
3. Important respiratory centers are located in the
 - a. cerebrum.
 - b. cerebellum.
 - c. pons and medulla oblongata.
 - d. midbrain.
 - e. limbic system
4. The cerebral peduncles are a major descending motor pathway found in the
 - a. cerebrum.
 - b. cerebellum.
 - c. pons.
 - d. midbrain.
 - e. medulla oblongata.
5. The superior colliculi are involved in, whereas the inferior colliculi are involved in
 - a. hearing, visual reflexes
 - b. visual reflexes, hearing
 - c. balance, motor pathways
 - d. motor pathways, balance
 - e. respiration, sleep

UNIT 4 DIENCEPHALON AND BASAL GANGLIA

CONTENTS

- 1.0 Introduction
- 2.0 objectives
- 3.0 Main Content
 - 3.1 Structure of the diencephalon
- 4.0 Summary

1.0 INTRODUCTION

The forebrain consists largely of white matter with buried grey matter aggregates.

Largest grey matter aggregate is the thalamus. Other grey matter aggregate lies lateral to the thalamus known as the basal ganglia/nuclei. The unexpanded portion of the forebrain that is in the midline is known as the diencephalon. The grey matter aggregates are lateral relations of the diencephalon, the thalamus more medial than the basal ganglia.

3.0 OBJECTIVES

At the end of this unit, you will be able to:

- describe the diencephalon and basal ganglia
- discuss the anatomy of the thalamus and its surrounding structures
- discuss the anatomical basis of Parkinson's disease.

3.1 Structure of the diencephalon

The 3rd ventricle is the cavity of the diencephalon. It is a slit-like space which lies in the sagittal plane roofed by the fornix and the corpus callosum. The lateral wall of the 3rd ventricle is largely formed by the thalamus. The thalamus is separated from the hypothalamus on the lateral wall of the 3rd ventricle by the hypothalamic sulcus. The sulcus runs from the interventricular foramen towards the aqueduct of the midbrain. The lower part of the lateral wall and the floor of the 3rd ventricle is formed by the hypothalamus. The optic chiasma forms the anterior end of the floor of the 3rd ventricle. The infundibulum is just behind the optic chiasma and projects downward forming the pituitary stalk. The pituitary stalk provides a path of communication between the hypothalamus and the posterior lobe of pituitary gland. Other features

on the external surface of the floor are mammillary bodies and posterior perforated substance.

The median eminence is the portion of the floor just behind the infundibulum. It contains cells that control the anterior pituitary gland. The anterior wall of the 3rd ventricle is the lamina terminalis extending from the anterior commissure to the optic chiasma. The body of the fornix separates the anterior commissure from the interventricular foramen. Between the rostrum, genu and body of the corpus callosum and the body of the fornix is a 2 ply thin partition known as the septum pellucidum. The roof is formed by pia matter which covers the superior surface of the thalamus. The roof is invaginated by the pair of choroid plexus.

The posterior wall tapers towards the midbrain. The pineal gland projects posteriorly from the posterior wall, above the superior colliculus. This small conical body secretes melatonin, its stalk is attached to the posterior commissure. Calcification of the pineal gland is common after 40 years and is thus useful for radiologist as an indicator of brain symmetry.

Thalamus

It is the largest of the cellular aggregate of the forebrain. It forms the lateral wall of the diencephalon. The medial surfaces of the thalamus lie parallel to each other slightly bulging into the 3rd ventricle. An interthalamic adhesion joins the 2 thalami in about 60-80% of cases, there is however no exchange of substance via this connection. The pulvinar is the posterior end of the medial surface of the thalamus. The opening of the interventricular foramen forms the anterior limit of the thalamus, while the posterior commissure is the posterior limit. It extends from the 3rd ventricle to the medial side of the posterior limb of the internal capsule.

The superior surface of the thalamus tapers anteriorly from the pulvinar to the small anterior pole. The pia matter covering of the superior surface extends to the pulvinar as far backward as the pineal gland. The lateral edge of the superior surface is covered by ependyma and forms part of the floor of the lateral ventricle. The substance of the thalamus is divided into lateral, medial and anterior portions by the sheet of white matter known as the internal medullary lamina.

The lateral surface is closely related to the internal capsule. Lateral boundary corresponds with the position of the stria terminalis and the terminal vein. The inferior surface is medially continuous with the hypothalamus separated by the hypothalamic sulcus.

The thalamic nuclei are grouped, following structural delineation by the internal medullary lamina and function.

Anterior nuclei group; the anterior end of the thalamus separated from the other part by the limbs of the internal medullary lamina. Connected to the mammillary body and projects to the cingulate gyrus.

Medial nuclei group; the gray substance medial to the internal medullary lamina, it projects to the frontal cortex.

Lateral nuclei group; mass of cells anterior to the pulvinar on the lateral side of the thalamus, project to different parts of the brain including motor cortex, postcentral gyrus and parietal lobe. It receives fibres from the medial lemniscus, spinothalamic and trigeminal tracts.

Posterior nuclei group; this consists of the pulvinar and the medial and lateral geniculate bodies. The pulvinar is the large posterior nuclei that projects to the temporal and parietal lobes.

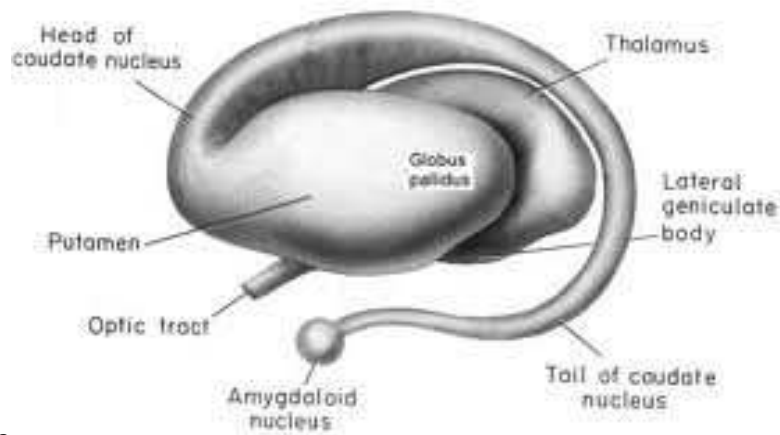
The medial geniculate body is ventral to the pulvinar and lateral to the midbrain, receives fibres from the lateral lemniscus and inferior colliculus, projects into the temporal lobe.

The lateral geniculate body is an oval elevation on the posterior end of the pulvinar, receives fibres from the optic tract and projects to the visual cortex.

Basal ganglia

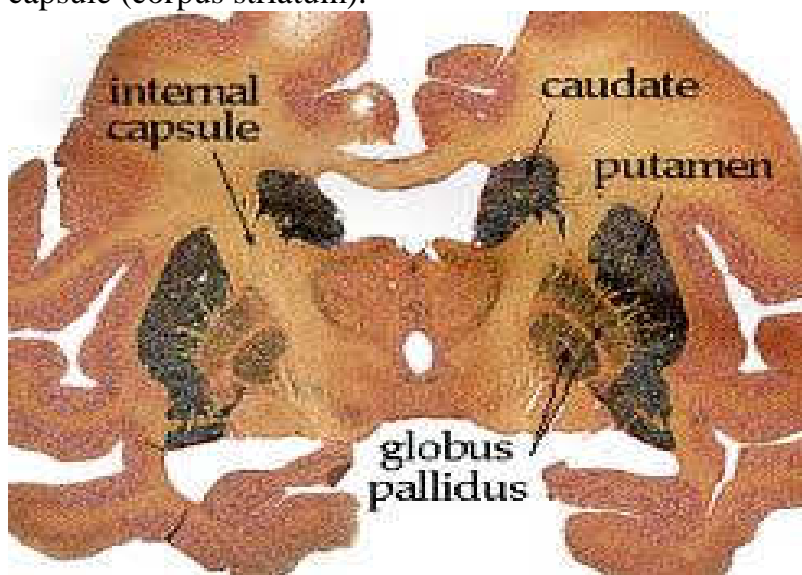
This consist of the caudate nucleus, lentiform nucleus, amygdaloid body and claustrum. The lentiform nucleus consists of an outer putamen and an inner globus pallidus. The caudate and putamen form the striatum. It is the larger part of the basal ganglia and its afferent part.

Caudate nucleus: It is a comma shaped structure with a demonstrable bulbous head, body and tail attached to the amygdaloid body.



The ganglia basal

The convexity of the caudate nucleus projects into the ventricle. The internal capsule fibres run in the concavity of the nucleus. The lentiform nucleus; consist of the putamen and globus pallidus. It is biconvex in shape with the putamen lateral and smaller medial globus pallidus. The caudate nucleus and the putamen are joined by many interconnecting fibre running through the anterior part of the internal capsule (corpus striatum).



Amygdaloid body: a group of grey matter at the tip of the tail of the caudate nucleus. It lies on the roof of the inferior horn of the lateral ventricle. It has numerous neurons that connect it to the frontal, temporal and olfactory lobes. Its efferent bundle (stria terminalis) runs parallel to the concavity of the caudate nucleus. The claustrum is a saucer shaped thin lamina of grey matter, lateral to the putamen.

Functional anatomy

- The corpus striatum is the afferent centre of the basal ganglia receiving fibres from the cortex, thalamus and substantia nigra.
- The globus pallidus is the efferent pathway of the basal ganglia sending fibres to the thalamus, subthalamic nucleus, substantia nigra etc
- The basal ganglia offer supraspinal control over skeletal muscle movement, modulating the rate, range and coordination.
- Several neurotransmitter are involved in the different pathways such as the acetylcholine, dopamine, glutamate, serotonin etc
- Parkinsonism is the most common anomaly of the basal nuclei. It involves a decrease in dopamine in the nigrostriatal pathway following destruction of dopaminergic cells of the substantia nigra. Huntington's disease is as a result of destruction of the neurons in the striatum.

Clinical correlates

Huntington's disease - this is a neurodegenerative genetic disorder that affects muscle coordination and leads to cognitive decline and behavioural symptoms. The disease can affect both male and female and presents mostly in mid – adult life although physical symptoms can begin at any age. It is much more common in Western European people.

Parkinsonism - as discussed in the earlier unit.

4.0 SUMMARY

- The diencephalon is located between the brainstem and the cerebrum. It consists of the thalamus, subthalamus, epithalamus, and hypothalamus.
- The thalamus consists of two lobes connected by the interthalamic adhesion. The thalamus functions as an integration center. All sensory input that reaches the cerebrum, except for the sense of smell, synapses in the thalamus. Pain is registered in the thalamus.
- The thalamus interacts with other parts of the brain to control motor activity. The thalamus is involved with emotions and pain perception. The subthalamus is inferior to the thalamus and is involved in motor function.
- The epithalamus is superior and posterior to the thalamus and contains the habenula, which influence emotions through the sense of smell. The pineal gland may play a role in the onset of puberty and the sleep–wake cycle.
- The hypothalamus is inferior to the thalamus. The hypothalamus is a major integrating center for the ANS, helping control heart rate, blood vessel diameter, urine release from the urinary

bladder, and the movement of food through the digestive tract. The hypothalamus regulates body temperature, hunger, satiety, thirst, and swallowing and interacts with the reticular activating system.

- The hypothalamus is involved with sensations (sexual pleasure, good feelings, rage, and fear). The mammillary bodies are reflex centers for olfaction.

SELF – ASSESSMENT EXERCISE

1. The major relay station for sensory input that projects to the cerebral cortex is the
 - a. hypothalamus.
 - b. thalamus.
 - c. pons.
 - d. cerebellum.
 - e. midbrain.
2. Which part of the brain is involved with olfactory reflexes and emotional responses to odours?
 - a. inferior colliculi
 - b. superior colliculi
 - c. mammillary bodies
 - d. pineal gland
 - e. pituitary gland
3. Which of the following is a function of the hypothalamus?
 - a. regulates autonomic nervous system functions
 - b. regulates the release of hormones from the posterior pituitary
 - c. regulates body temperature
 - d. regulates food intake (hunger) and water intake (thirst)
 - e. all of the above.

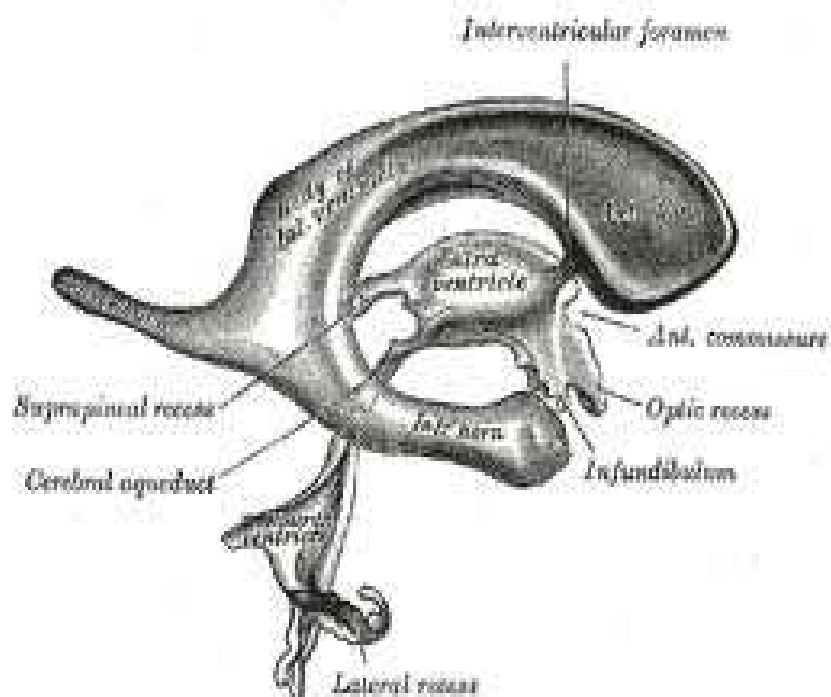
UNIT 5 THE VENTRICLES AND CEREBROSPINAL FLUID

CONTENTS

- 1.0 Introduction
- 2.0 Objective
- 3.0 Main Content
 - 3.1 Third ventricle
- 4.0 Summary

1.0 INTRODUCTION

The ventricles are cavities within the brain for the production and drainage of Cerebrospinal fluid (CSF). It consists of the lateral ventricles, the 3rd ventricle and the 4th ventricle. The ventricular system of the brain is continuous with the central canal of the spinal cord. The ventricles contain CSF that cushions the brain and spinal cord within its bony container as well as provide nourishment. They are modified derivative of the neural canal.



3.1.1 Learning objective

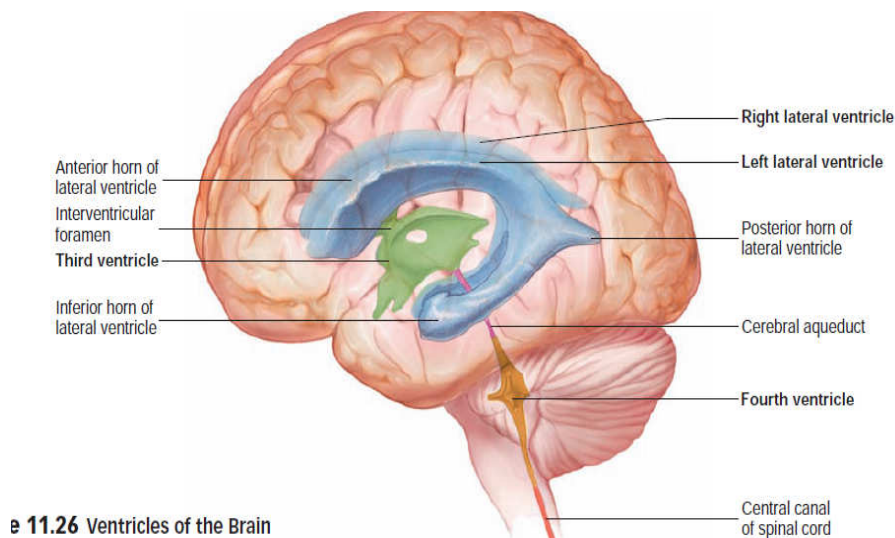
At the end of this unit, you will be able to:

- I. Differentiate the parts of the ventricular system and relate them to the different regions of the brain.

- II. Describe the walls of the different parts of the ventricular system.
- III. Discuss the production and flow of the cerebrospinal fluid.

3.1.2 Lateral ventricle

It is a C-shaped continuous cavity within the cerebral hemisphere. It consists of the body, anterior, posterior and inferior horns. Its medial wall is marked by the slit-like opening on the medial surface of the hemisphere. This marks the point of invagination of the pia mater and ependyma for the formation of the choroid plexus which produces the CSF.



The choroid plexus of the lateral ventricle is continuous with that of the 3rd ventricle. The lateral ventricle produces most of the CSF.

Anterior Horn; it is triangular in outline with the apex pointing laterally. The fornix minor forms the roof, while the floor is indented by the head of the caudate nucleus.

Laterally the roof and the floor converge while they are kept apart by the septum pellucidum on the medial side. The lateral ventricle communicates with the 3rd ventricle via the interventricular foramen of Monro located just behind the anterior column of fornix. It is also the point of continuation of the Choroid plexus of the body and 3rd ventricle.

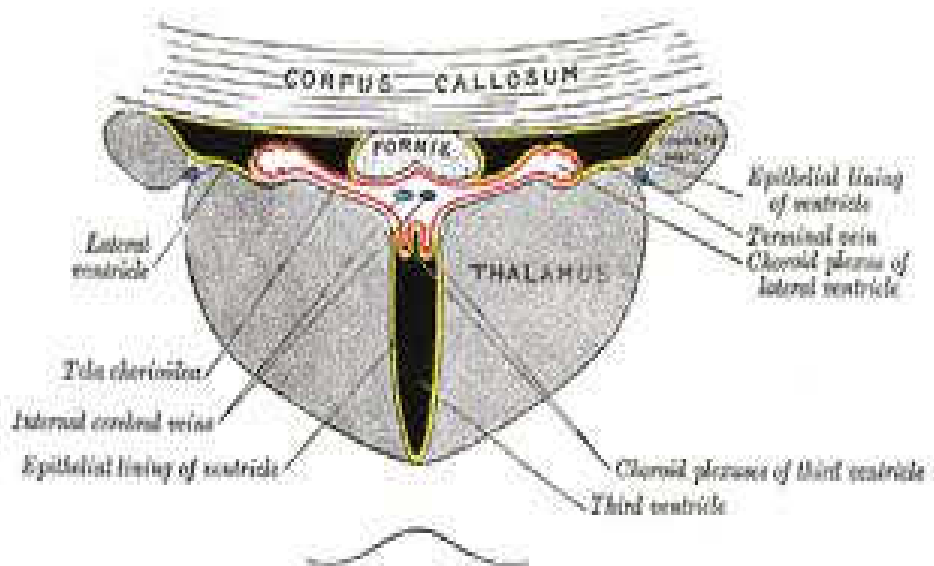
Body; the interventricular foramen of Monro is the boundary between the anterior horn and body. The Thalamus and the body of the caudate nucleus bulge into its floor. Thalamostriate vessels run between them. The roof is formed by the corpus callosum and the body of fornix. Invagination of the choroid plexus is via the medial wall (septum

pellucidum). Posterior Horn; the existence is variable. The collateral eminence bulges on its floor. 2 bulges are on the medial wall, upper bulb of posterior horn and lower calcar avis. The 2 can obliterate the horn. The tapetum forms the roof and lateral wall. The optic radiation is related to the lateral wall.

Inferior Horn; it is the largest part of the ventricles. The Hippocampus (medial) and the collateral eminence (lateral) form the floor of the horn. The tail of the caudate nucleus forms the roof ending in a convexities formed by the amygdaloid body and pes hippocampi. Lateral wall formed by the tapetum.

Third ventricle

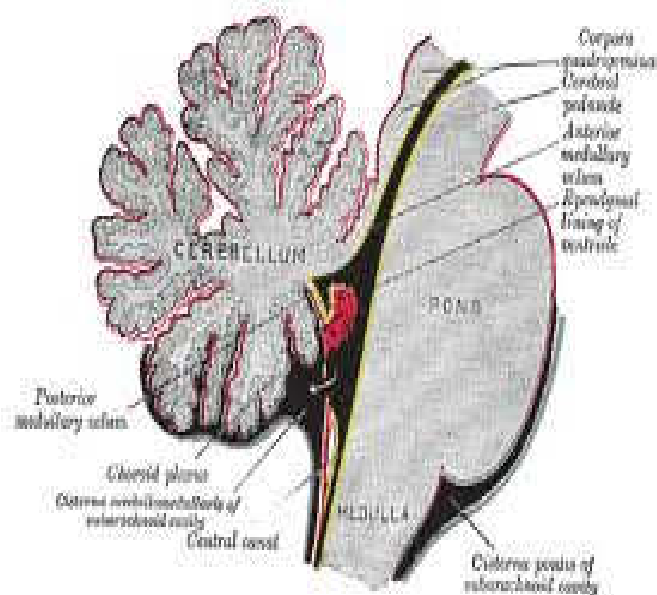
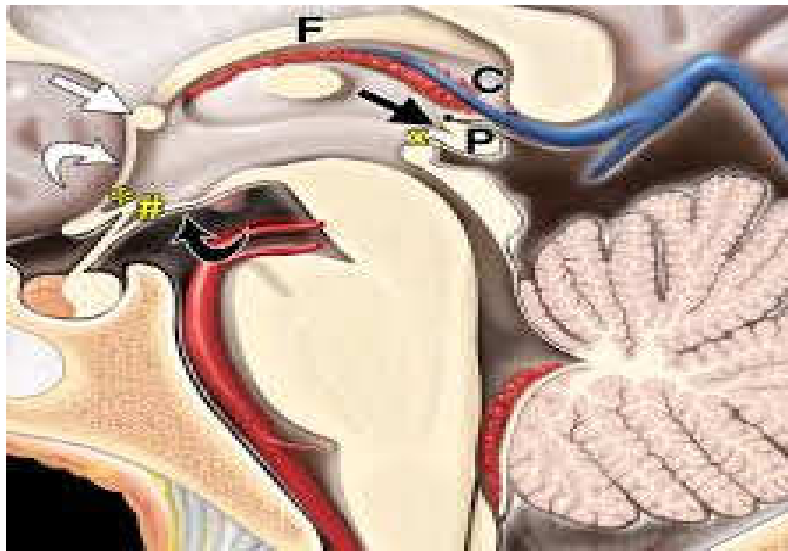
It is a slit-like space beneath the fornix and corpus callosum. Bounded anteriorly by the anterior column of fornix and the interventricular foramen of Monro. The lateral wall is largely occupied by the Thalamic bulge. The hypothalamic sulcus on the lateral wall connects the IVFM to the aqueduct of the midbrain.



Fourth ventricle

It communicates with the 3rd ventricle via the aqueduct of the midbrain. The pons and the upper medulla are ventral relations while the cerebellum is largely a supero-dorsal relation. The floor is diamond shaped with a tent like roof. It is triangular in sagittal section. Caudally the lateral boundary of the floor is formed by the gracile and cuneate tubercles with the inferior cerebellar peduncle superimposed on them. The lateral boundary of the floor cranially is formed by the superior cerebellar peduncle.

A thin stretch of white matter (superior medullary velum) connects the 2 superior cerebellar peduncle forming the cranial part of the roof. The caudal part of the roof is formed by a combination of pia and ependyma. The foramen of Magendie is a central opening of it. The cerebellomedullary cistern communicates via the foramen.



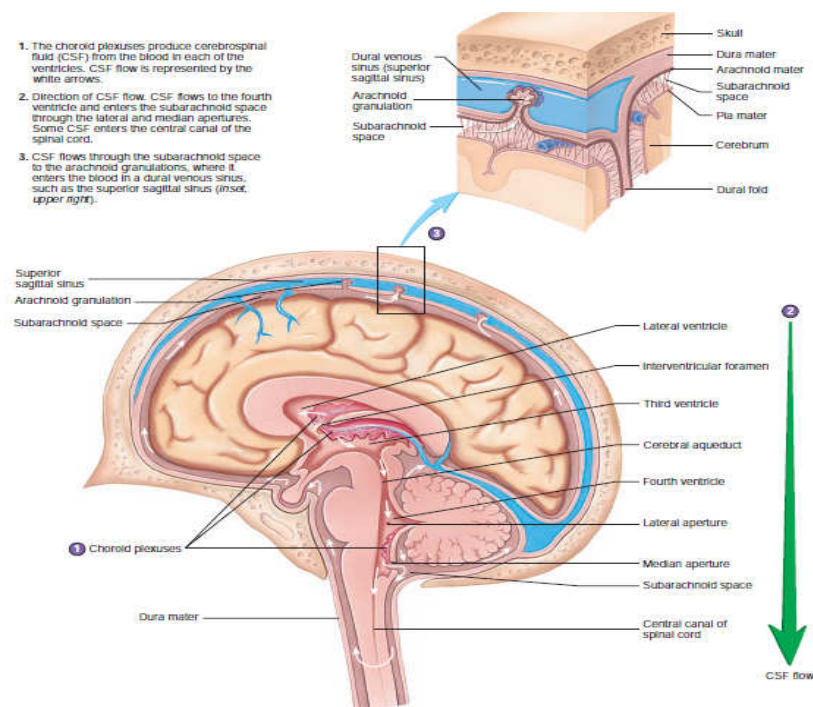
The narrow lateral recess has the foramen of Luschka that communicates with the pontine cistern. The T-shaped choroid plexus indents the cranial part of the roof. The floor (Rhomboid fossa) has a midline groove (median sulcus) which connects the aqueduct of Sylvius to the central canal of the spinal cord.

The cranial part of the floor is divided into the medial pontine part and lateral vestibular part. The pontine part of the floor has a bulge (facial colliculus) adjacent to the median sulcus. Laterally extending white

matter (medullary striae) is found at the widest part of the floor. The vestibular area overlies the vestibular nuclei. The floor of the medullary part is divided into 2 by a faint groove that extends laterally from median sulcus towards the lateral recess. The medial small triangle (hypoglossal trigone) overlying the hypoglossal nucleus, while the lateral (vagal trigone) overlies the dorsal nucleus of vagus.

Cerebrospinal fluid

It is produced by a modified ependymal cells and choroid plexus which is found in most part of the ventricular system. 70% of the CSF is produced from the ventricles while the remaining 30% is produced from brain capillaries. Total CSF volume is about 130ml, about 30ml is found in the ventricles while the subarachnoid space contains about 100ml. The body of the lateral ventricle and the 3rd ventricle are major site of production. The fluid drains through the IVFM to the 3rd ventricle, through the Aqueduct of Sylvius to the 4th ventricle. It exits via the Foramina of Magendie and Luschka into the subarachnoid space or flow straight into the central canal of spinal cord. The fluid flows around the superior sagittal sinus and is reabsorbed via the arachnoid granulation into the venous system.



Flow of CSF

Clinical correlates

- I. **Lumbar puncture** is a sterile procedure done to withdraw C.S.F. from the spinal subarachnoid space and must be performed well clear of the termination of the cord. A line joining the iliac crests passes through the 4th lumbar vertebra and therefore the

intervertebral spaces immediately above or below this landmark can be used with safety. The spine must be fully flexed (with the patient either on his side or seated) so that the vertebral interspinous spaces are opened to their maximum extent. The needle is passed inwards and somewhat cranially exactly in the midline and at right angles to the spine; the supraspinous and interspinous ligaments are traversed and then the dura is penetrated, the latter with a distinct 'give'. Occasionally root pain is experienced if a root of the cauda equina is impinged upon, but usually these float clear of the needle. At spinal puncture C.S.F. can be obtained for examination; antibiotics, radio-opaque contrast medium or anaesthetics may be injected into the subarachnoid space, and the C.S.F. pressure can be estimated (normal, when lying on the side, 80–180mm C.S.F.).

II. Hydrocephalus

CSF is produced at a rate of approximately 0.4 mL/min. If CSF returns to the blood at the same rate, the volume of CSF in the ventricles and subarachnoid space is constant. If the apertures of the fourth ventricle or the cerebral aqueduct are blocked, CSF can accumulate within the ventricles. This condition is called **internal hydrocephalus**, or **noncommunicating hydrocephalus**, and it results in increased CSF pressure. The production of CSF continues, even when the passages that normally allow it to exit the brain are blocked. Consequently, fluid builds inside the brain, causing pressure, which compresses the nervous tissue and dilates the ventricles. Compression of the nervous tissue usually results in irreversible brain damage. If the skull bones are not completely ossified when the hydrocephalus occurs, the pressure may also severely enlarge the head. The cerebral aqueduct may be blocked at the time of birth or may become blocked later in life because of a tumor growing in the brainstem. Internal hydrocephalus can be successfully treated by placing a drainage tube (shunt) between the brain ventricles and abdominal cavity to eliminate the high internal pressures. There is some risk of infection being introduced into the brain through these shunts, however, and the shunts must be replaced as the person grows. A subarachnoid hemorrhage may block the return of CSF to the circulation. If CSF accumulates in the subarachnoid space, the condition is called **external hydrocephalus**, or **communicating hydrocephalus**. In this condition, pressure is applied to the brain externally compressing neural tissues and causing brain damage.

4.0 SUMMARY

- The lateral ventricles in the cerebrum are connected to the third ventricle in the diencephalon by the interventricular foramina.

The third ventricle is connected to the fourth ventricle in the pons by the cerebral aqueduct. The central canal of the spinal cord is connected to the fourth ventricle.

- The fourth ventricle is connected to the subarachnoid space by the median and lateral apertures.
- CSF is produced from the blood in the choroid plexus of each ventricle by ependymal cells. CSF moves from the lateral to the third and then to the fourth ventricle. From the fourth ventricle, CSF enters the subarachnoid space through three apertures.
- CSF leaves the subarachnoid space through arachnoid granulations and returns to the blood in the dural venous sinuses.

Activity

Locate and identify the ventricles, canals and capillary beds associated with the circulation of cerebrospinal fluid on the brain models provided or preserved animal brains in the anatomy laboratory/Museum.

SELF – ASSESSMENT EXERCISE

1. Cerebrospinal fluid is produced by the _____, circulates through the ventricles, and enters the subarachnoid space. The cerebrospinal fluid leaves the subarachnoid space through the
 - a. choroid plexuses, arachnoid granulations
 - b. arachnoid granulations, choroid plexuses
 - c. dural venous sinuses, dura mater
 - d. dura mater, dural venous sinuses
2. Describe the ventricles of the brain.
3. What are the physical characteristics of CSF?

UNIT 6 THE SPINAL CORD

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 General structure of the spinal cord
- 4.0 Summary

1.0 INTRODUCTION

The **spinal cord** is extremely important to the overall function of the nervous system. It is the major communication link between the brain and the PNS (spinal nerves) inferior to the head. The spinal cord participates in the integration of incoming information and produces responses through reflex mechanisms.

2.0 OBJECTIVES

At the end of this unit, you will be able to:

- discuss the parts of the spinal cord
- describe the three meningeal layers surrounding the spinal cord and brain
- describe the spinal cord in cross section and the origin of spinal nerves
- describe the blood supply of the spinal cord

3.0 MAIN CONTENT

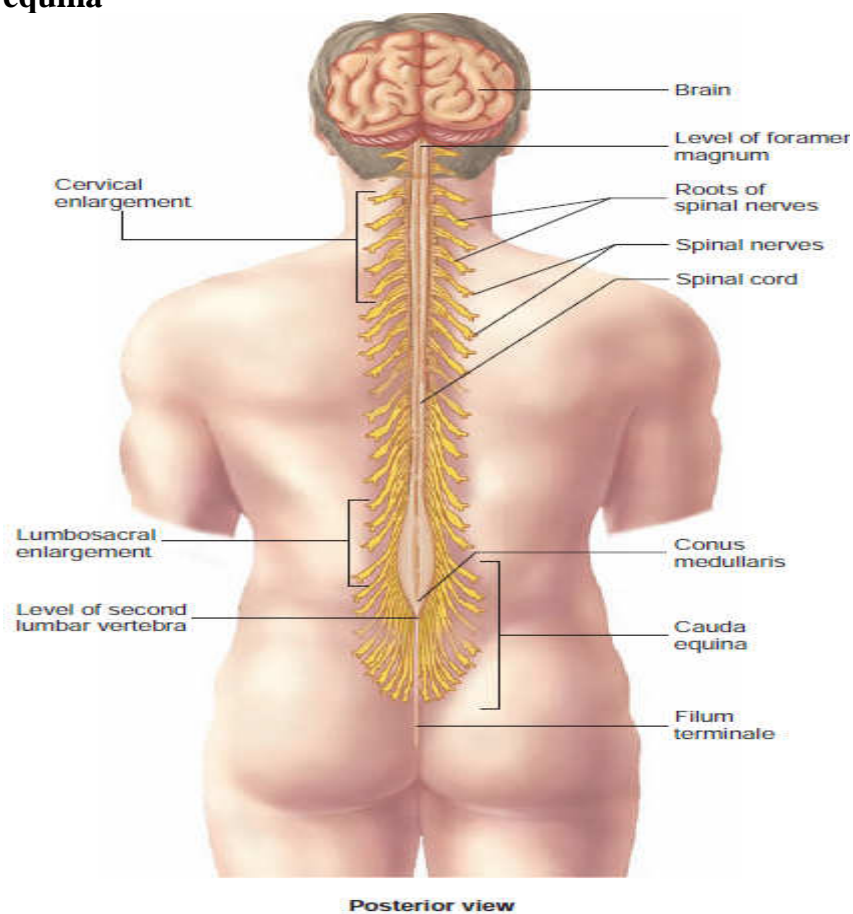
3.1 General structure of the spinal cord

The spinal cord connects to the brain at the level of the foramen magnum and extends inferiorly in the vertebral canal to level L1–L2 of the vertebral column. It is considerably shorter than the vertebral column because it does not grow as rapidly as the vertebral column during development. The spinal cord gives rise to 31 pairs of spinal nerves, which exit the vertebral column through intervertebral and sacral foramina.

Develops from the caudal portion of the neural tube, it is a region of less differentiation. It consists of 31 segments conferred by the emergence of the spinal nerves; cervical -8, thoracic- 12, lumbar-5, sacral-5, coccygeal- 1. Measuring about 42-45cm in adult.

The spinal cord is not uniform in diameter throughout its length. The **cervical enlargement** in the inferior cervical region is where spinal nerves supplying the upper limbs originate. The **lumbosacral enlargement** in the inferior thoracic, lumbar, and superior sacral regions is the site where spinal nerves supplying the lower limbs originate.

Immediately inferior to the lumbosacral enlargement, the spinal cord tapers to form a conelike region called the **conus medullaris**. Nerves arising from the inferior lumbosacral enlargement and the conus medullaris extend inferiorly through the vertebral canal before exiting the vertebral column. The numerous roots (origins) of these nerves resemble the hairs in a horse's tail and are therefore called the **cauda equina**



the spinal cord and segments

Meninges of the spinal cord

The spinal cord is a long cylindrical structure with flattened anterior-posterior surfaces. The spinal cord and brain are surrounded by connective tissue membranes called **meninges**. The most superficial and thickest membrane is the **dura mater**. The dura mater forms a sac, often called the **thecal sac**, which surrounds the spinal cord. The thecal sac attaches to the rim of the foramen magnum and ends at the level of the

second sacral vertebra. The spinal dura mater is continuous with the dura mater surrounding the brain and the connective tissue surrounding the spinal nerves. The dura mater around the spinal cord is separated from the periosteum of the vertebral canal by the **epidural space**, which contains spinal nerves.

Internal structure of the spinal cord

The internal structure of the spinal cord consists of the white and gray matter.

White matter is a collection of myelinated fibres, while cellular aggregates are in the gray matter.

H-shaped gray matter located within the cylindrical white matter, bridge of the h incorporating the central canal. Dorsal limbs of the h form the posterior horn which extends close to surface.

The anterior horns are farther away from the surface. The medial anterior horn cell innervate trunk muscles, lateral part supplies the limbs. Ventral part of lateral horn supplies the proximal muscles while dorsal part supplies distal muscles of limbs. The white matter is bounded by the limbs of the H-shaped gray matter are grouped as the anterior, posterior and lateral white column.

A narrow white commissure separates the anterior median fissure from the grey commissure.

Spinal segments

Segmental differences exist with respect to the size of the grey horn and the white column.

Cervical segment; large oval shaped structure, relatively larger white matter. Posterior horn is smaller than the anterior. Posterior intermediate septum divides the posterior column into graciles and cuneatus. The posterior horn is enlarged from the C5 level downwards. Reticular process is at the base of the posterior horn.

Thoracic segments; the gray matter is relatively smaller than the white matter. Gracilis and cuneatus extend as far as mid thoracic segments, while gracilis continues caudally. T1 is more of a cervical in outline and gray and white matter proportion. Lateral horn at the base of the anterior horn, give rise to the preganglionic sympathetic efferent. Dorsal nucleus of Clarke located on the medial side of the base of the posterior horn, most obvious in T10-T12.

Lumbar segment; almost circular in transverse section. White matter proportionally less than gray matter. Gracilis has thinned out. Blunt lateral process on the lateral side of the base of stout anterior horn. Dorsal nucleus of Clarke in L1-L2.

Sacral segment; small cylindrical structure. Gray matter relatively more than the white matter.

Thickened gray commissure, posterior and anterior horn. Small lateral horn for the preganglionic parasympathetic cells.

Clinical correlates

Introduction of Needles into the Subarachnoid Space

Several clinical procedures involve the insertion of a needle into the subarachnoid space inferior to the level of the second lumbar vertebra. The needle is introduced into either the L3/L4 or the L4/L5 intervertebral space. The needle does not contact the spinal cord because the spinal cord extends only approximately to the second lumbar vertebra of the vertebral column. The needle enters the subarachnoid space, which extends to level S2 of the vertebral column. The needle does not damage the nerve roots of the cauda equine located in the subarachnoid space because the needle quite easily pushes them aside.

In **spinal anesthesia**, or spinal block, drugs that block action potential transmission are introduced into the subarachnoid space to prevent pain sensations in the lower half of the body.

A **spinal tap** is the removal of CSF from the subarachnoid space. A spinal tap may be performed to examine the CSF for infectious agents (meningitis), for the presence of blood (hemorrhage), or for the measurement of CSF pressure. A radiopaque substance may also be injected into this area, and a **myelogram** (radiograph of the spinal cord) may be taken to visualize spinal cord defects or damage.

Damage to the spinal cord can disrupt ascending tracts from the spinal cord to the brain, resulting in the loss of sensation, and/or descending tracts from the brain to motor neurons in the spinal cord, resulting in the loss of motor functions. Automobile and motorcycle accidents are leading causes, followed by gunshot wounds, falls, and swimming accidents. Spinal cord injury is classified according to the vertebral level at which the injury occurred, whether the entire cord is damaged at that level or only a portion of the cord, and the mechanism of injury. Most spinal cord injuries occur in the cervical region or at the thoracolumbar junction and are incomplete.

4.0 SUMMARY

- The spinal cord gives rise to 31 pairs of spinal nerves. The spinal cord has cervical and lumbosacral enlargements from which spinal nerves of the limbs originate.
- The spinal cord is shorter than the vertebral column. Nerves from the end of the spinal cord form the cauda equina.
- Three meningeal layers surround the spinal cord. From superficial to deep they are the dura mater, arachnoid mater, and pia mater.
- The epidural space is between the periosteum of the vertebral canal and the dura mater, the subdural space is between the dura mater and the arachnoid mater, and the subarachnoid space is between the arachnoid mater and the pia mater.
- Gray matter is divided into horns. The dorsal horns contain sensory axons that synapse with interneurons. The ventral horns contain the neuron cell bodies of somatic motor neurons, and the lateral horns contain the neuron cell bodies of autonomic motor neurons.
- The dorsal root contains sensory axons, the ventral root has motor axons, and spinal nerves have sensory and motor axons.

Activity

I. In the anatomy museum, identify the major features on the models and slide of a cross – section of the spinal cord; also note the three layers of the meninges on the cross- section models.

SELF – ASSESSMENT EXERCISE

1. The spinal cord extends from the
 - a. medulla oblongata to the coccyx.
 - b. level of the third cervical vertebra to the coccyx.
 - c. level of the axis to the lowest lumbar vertebra.
 - d. level of the foramen magnum to level L1–L2 of the vertebral column.
 - e. axis to the sacral hiatus.
2. Axons of sensory neurons synapse with the cell bodies of interneurons in the of spinal cord gray matter.
 - a. anterior horn
 - b. lateral horn
 - c. posterior horn
 - d. gray commissure
 - e. lateral funiculi

3. Cell bodies for spinal sensory neurons are located in the
 - a. anterior horn of spinal cord gray matter.
 - b. lateral horn of spinal cord gray matter.
 - c. posterior horn of spinal cord gray matter.
 - d. dorsal root ganglia.
 - e. posterior columns.
4. Contrast the epidural and the subarachnoid spaces.
5. Describe the appearance of the spinal cord in cross section.
6. Discuss the anatomy of the terminal portion of the spinal cord.

UNIT 7 BLOOD SUPPLY OF THE CENTRAL NERVOUS SYSTEM

CONTENTS

- 1.0 Introduction
- 2.0 Objective
- 3.0 Main Content
 - 3.1 Internal carotid system
- 4.0 Summary

1.0 INTRODUCTION

The brain requires a tremendous amount of blood to maintain its normal functions. The brain has a very high metabolic rate, and brain cells are not capable of storing high-energy molecules for any length of time. In addition, brain cells depend almost entirely on glucose as their energy source. Thus, the brain requires a constant blood supply to meet the demands of brain cells for both glucose and oxygen. Even though the brain accounts for only about 2% of the total weight of the body, it receives approximately 15%–20% of the blood pumped by the heart. Interruption of the brain's blood supply for only seconds can cause unconsciousness, and interruption of the blood supply for minutes can cause irreversible brain damage.

2.0 OBJECTIVE

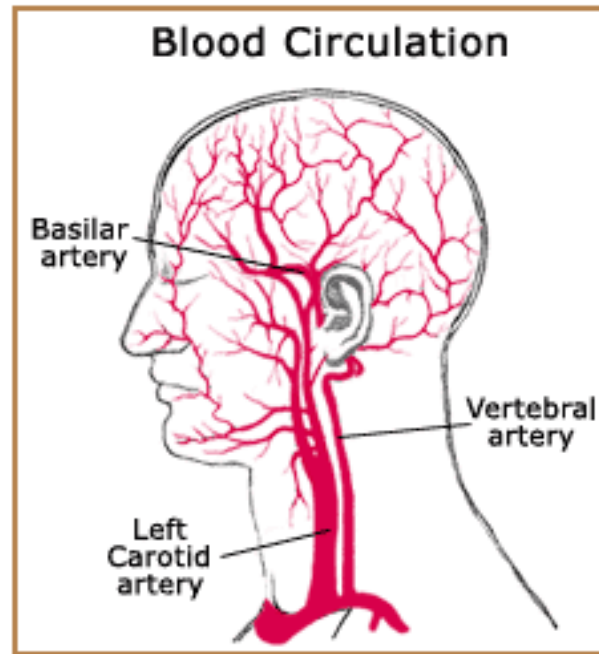
At the end of this unit, you will be able to:

- describe the arterial supply of the brain and the spinal cord.
- describe the venous drainage of the brain and the spinal cord.
- define the blood-brain barrier and its effect on the movement of materials into and out of the brain.

3.0 MAIN CONTENT

3.1 Internal carotid system

The main blood supply to the central nervous system include the internal carotid, vertebral and basilar systems of arteries.



The internal carotid system consists of the anterior and middle cerebral arteries, anterior choroidal and posterior communicating arteries. The internal carotid artery proceeds superiorly alongside the optic chiasma, bifurcates into the middle cerebral artery and anterior cerebral artery.

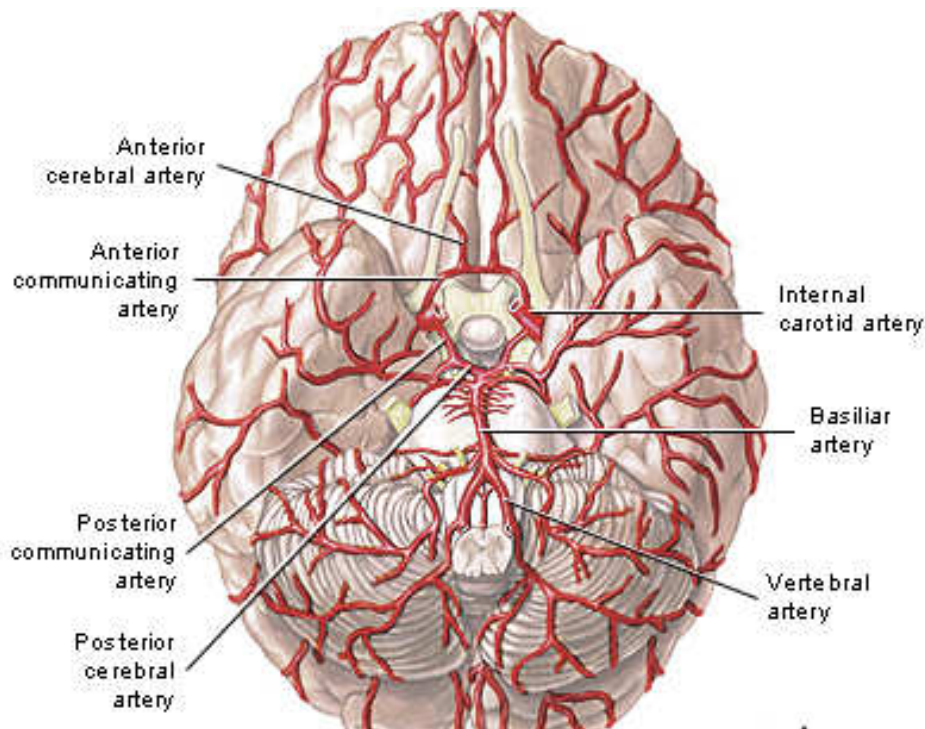
It contributes to the formation of the choroid plexus of the lateral ventricle (especially inferior horn). It gives 2 smaller branches before bifurcating viz; anterior choroidal artery and posterior communicating artery.

Anterior Choroidal Artery; It runs posterior to the optic tract, clinically significant, long and thin artery. It is frequently involved in CVA. It supplies:

Optic tract, Lateral geniculate body, Optic radiation, Uncus, Amygdala, Hippocampus, Internal capsule, Lentiform nucleus (globus pallidus and putamen) and Thalamus

Posterior Communicating Artery; It runs posterior inferiorly to the optic tract joining the posterior cerebral artery. It participates in the formation of the circle of Willis.

Anterior Cerebral Artery; It runs superiorly and medially to the optic nerve to enter the longitudinal fissure.



The two are interconnected by the anterior communicating artery just before entering the longitudinal fissure. It runs posteriorly following the corpus callosum supplying the medial surfaces of frontal and parietal lobes rostral to the parieto-occipital sulcus.

It divides into 2 prominent branches:

Peri-callosal artery; adjacent to the corpus callosum.

Calloso-marginal artery; which follows the cingulate sulcus.

It supplies the paracentral lobule (medial aspects of the precentral and postcentral gyri), and then its occlusion will cause restricted contralateral motor and somatosensory deficits.

Middle Cerebral Artery; runs laterally into the lateral sulcus, supply the insula and temporal pole. Numerous branches spread out from the sulcus to supply the lateral surface of the cerebrum. Its occlusion causes major motor and somatosensory deficits. Also, if the left hemisphere is involved, language deficits are almost invariably found. Deep structures of diencephalon and telencephalon are supplied by its numerous small branches (lenticulostriate arteries).

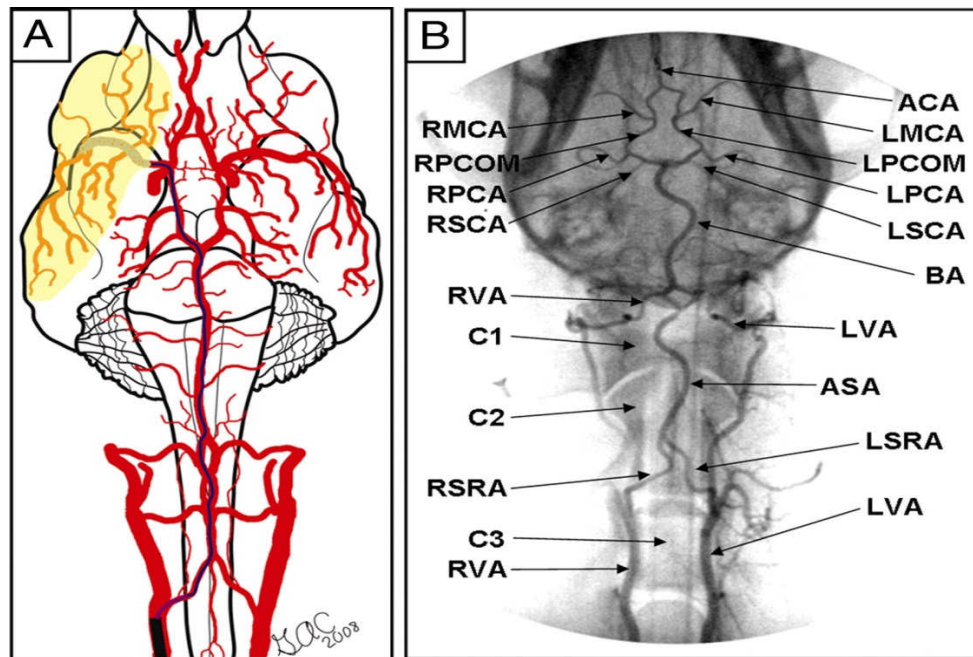
Vertebral system

Vertebral Arteries; they run alongside the medulla, fuse at the pontomedullary junction to form the basilar artery. Before the union it gives 3 branches;

The posterior spinal artery; runs along the dorsolateral aspect of SC supplying the dorsal 1/3rd.

The anterior spinal artery; join together to form a single anterior spinal artery, run on the antero-medial surface of SC to supply ventral 2/3rd.

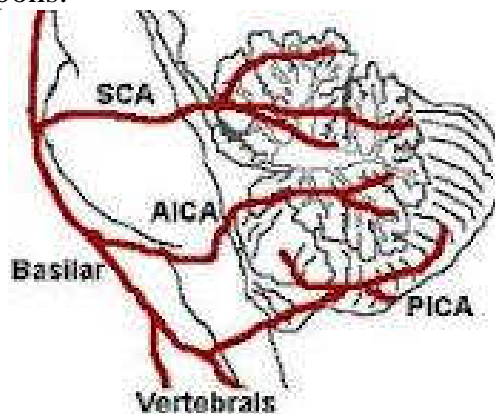
Posterior Inferior Cerebellar Artery (PICA); approaches the cerebellum to supply its inferior surface. As it curves around the brainstem, the artery supplies the choroid plexus of the 4th ventricle and much of the lateral medulla.



Basilar system

The basilar artery runs in the pontine groove, bifurcates at midbrain level into 2 posterior cerebral arteries. Also give rise to the following;

Anterior inferior cerebellar artery; formed rostral to formation of basilar artery, supplies the anterior part of the inferior cerebellar surface and the lower part of pons.



Superior cerebellar artery; it is formed just before the bifurcation of basilar artery, supplies the upper part of pons, superior surface of cerebellum and caudal part of midbrain.

The internal auditory artery; It supplies the middle ear, occlusion causes vertigo and deafness. Often comes from AICA. Numerous unnamed branches; e.g pontine arteries.

Posterior Cerebral Artery; it runs around the midbrain passing through the superior cistern.

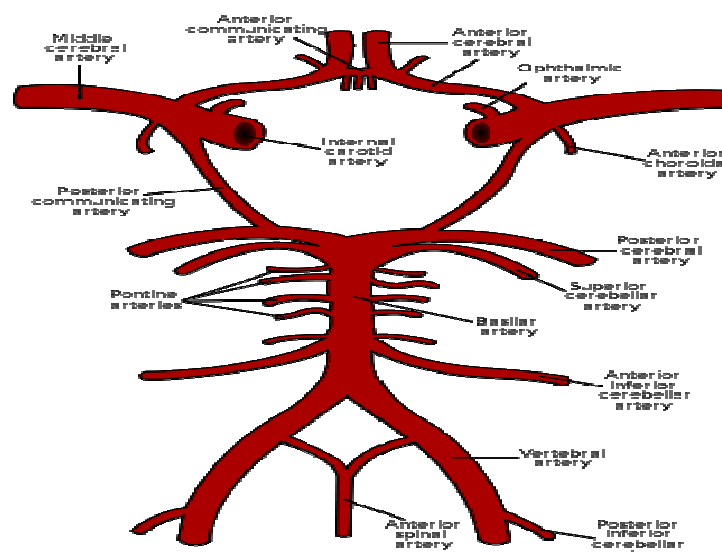
It supplies the medial surfaces of the occipital and temporal lobes. Also send branches to the upper part of midbrain and lower part of diencephalon. It also gives rise to several posterior choroidal arteries. Since the primary visual cortex is located in the occipital lobe, occlusion of the posterior cerebral artery could lead to visual loss among other symptoms

Posterior Choroidal Arteries; form the choroid plexus of 3rd and that of body of 4th ventricles.

The Circle of Willis

The posterior cerebral artery is connected to the internal carotid artery by the posterior communicating artery. This completes an arterial polygon called the circle of Willis which allows for anastomotic blood flow in case of occlusion. Several small arteries arise from the circle which are grouped thus;

The antero-medial; supplies the head of caudate, anterior limb of internal capsule and anterior part of hypothalamus.



The circle of willis

The anterolateral; also known as lenticulostriate, supplies the internal capsule, globus pallidus, putamen, head of caudate, claustrum and external capsule.

The posteromedial; supplies medial part of midbrain, anteriomedial part of thalamus, subthalamic region, middle and posterior part of hypothalamus.

The posterolateral; also known as thalamogeniculate artery, supply the posterior part of thalamus and lateral part of midbrain.

Venous drainage

The principal route of venous drainage of the brain is through a system of cerebral veins. These empty into the dura venous sinuses and ultimately into the internal jugular vein. There is also a collection of emissary veins.

These connect the extracranial veins with the dural sinuses. There is a basilar venous plexus around the base of the brain that communicates with the epidural venous plexus of the spinal cord. Cerebral veins are divided into superficial and deep groups.

Generally, the superficial veins lie on the surface of the cerebral hemispheres and empty into the superior sagittal sinus. The deep veins drain internal structures and eventually drain into the straight sinus. Cerebral veins are valve-less. They are also interconnected by numerous functional anastomoses both within a group and between the superficial and deep groups.

Clinical correlates

Blood Brain Barrier

- i. Branches of the arteries supplying the brain are located in the subarachnoid space. Smaller branches from the arteries in the subarachnoid space extend into the brain and quickly divide into capillaries. The epithelial cells of these capillaries are surrounded by the foot processes of astrocytes. The astrocytes promote the formation of tight junctions between the epithelial cells. The epithelial cells with their tight junctions form the blood–brain barrier, which regulates the movement of materials from the blood into the brain.

Materials that would enter many tissues by passing between the epithelial cells of capillaries cannot pass through the blood–brain barrier because of the tight junctions. Most materials that enter the brain pass through the epithelial cells. Lipid-soluble substances, such as nicotine, ethanol, and heroin, can diffuse through the plasma membranes of the

epithelial cells and enter the brain. Water-soluble molecules, such as amino acids and glucose, move across the plasma membranes of the epithelial cells by mediated transport.

- ii. The permeability characteristics of the blood–brain barrier must be considered when developing drugs designed to affect the CNS. For example, Parkinson disease is caused by a lack of the neurotransmitter dopamine, which normally is produced by certain neurons of the brain. A lack of dopamine results in decreased muscle control and shaking movements. Administering dopamine is not helpful because dopamine cannot cross the blood–brain barrier. Levodopa (L-dopa), a precursor to dopamine, is administered instead because it can cross the blood–brain barrier. CNS neurons then convert levodopa to dopamine, which helps reduce the symptoms of Parkinson disease.

4.0 SUMMARY

- The brain requires tremendous amounts of blood to function normally.
- The blood–brain barrier is formed by the endothelial cells of the capillaries in the brain. It limits what substances enter brain tissue.
- The principal route of venous drainage of the brain is through a system of cerebral veins
- The main blood supply to the central nervous system includes the internal carotid, vertebral and basilar systems of arteries.
- The internal carotid system consists of the anterior and middle cerebral arteries, anterior choroidal and posterior communicating arteries
- Vertebral arteries run alongside the medulla, fuse at the pontomedullary junction to form the basilar artery. It has three divisions: anterior spinal, posterior spinal and posterior inferior cerebellar artery.
- The basilar system with several branches.

Activity

In the anatomy museum, identify the arterial supplies and venous drainage of the central nervous system.

SELF – ASSESSMENT EXERCISE

1. Which of the following statements is true?
 - a. The brain requires constant delivery of glucose and oxygen to function.
 - b. The foot processes of astrocytes surround capillaries in the brain.
 - c. Tight junctions between capillary epithelial cells form the blood – brain - barrier.
 - d. Glucose passes through capillary epithelial cells by mediated transport.
 - e. all of the above .
2. Discuss the importance of the *blood–brain barrier* in maintaining homeostasis within the brain.
3. *True or false:* Alcohol passes readily through the BBB because it is a lipid-soluble compound.
4. Discuss the circle of willis.
5. Discuss the venous drainage of the central nervous system.

MODULE 4 – THE PERIPHERAL AND AUTONOMIC NERVOUS SYSTEM

INTRODUCTION

Anatomically, the nervous system is divided into the **central nervous system (CNS)** and the **peripheral nervous system (PNS)**. The CNS includes the *brain* and the *spinal cord* (see previous module). The PNS (described in this module) includes the *cranial nerves*, arising from the inferior aspect of the brain, and the *spinal nerves*, arising from the spinal cord.

The **autonomic nervous system (ANS)** is a functional division of the nervous system. It consists of components within the CNS and specific nerves. The ANS is subdivided into *sympathetic* and *parasympathetic divisions* that provide innervation to smooth and cardiac muscles, as well as glands. The ANS functions autonomically to maintain homeostasis and carry out many involuntary functions in the body. The peripheral nervous system consists of nerves that branch off the CNS. These nerves are called peripheral nerves and are classified in two types—**cranial nerves** and **spinal nerves**.

OBJECTIVE

At the end of this module, you should be able to:

- i. Differentiate between structural and functional divisions of the nervous system.
- ii. Discuss the peripheral and autonomic nervous systems

CONTENT

Unit 1	Cranial nerves
Unit 2	Spinal nerves
Unit 3	Autonomic nervous system

UNIT 1 CRANIAL NERVES

CONTENT

- 1.0 Introduction
- 2.0 objectives
- 3.0 Main content
 - 3.1 Names and function
- 4.0 Summary

1.0 INTRODUCTION

Cranial nerves are peripheral nerves that originate from the brain. Roman numerals and names designate the twelve different cranial nerves. The cranial nerves also have names in numeric order, this mnemonic can help you remember their names:

On Occasion Our Trusty Truck Acts Funny; Very Good Vehicle AnyHow. The mnemonic corresponds to the **O**lfactory (I), **O**ptic (II), **O**culomotor (III), **T**rochlear (IV), **T**rigeminal (V), **A**bducent (VI), **F**acial (VII), **V**estibulocochlear (VIII), **G**lossopharyngeal (IX), **V**agus (X), **A**ccessory (XI), and **H**ypoglossal (XII) nerves

2.0 OBJECTIVES

At the end of this unit, you will be able to:

- Discuss the distribution and function of each of the cranial nerves
- Discuss the brain stem reflexes

3.1 Names and function

- I. *Olfactory nerves* carry smell information to the brain for interpretation.
- II. *Optic nerves* carry visual information to the brain for interpretation.
- III. *Oculomotor nerves* are found within the muscles that move the eyeball, eyelid, and iris.
- IV. *Trochlear nerves* act in the muscles that move the eyeball.
- V. *Trigeminal nerves* carry sensory information from the surface of the eye, the scalp, facial skin, the lining of the gums, and the palate to the brain for interpretation. They also are found within the muscles needed for chewing.
- VI. *Abducens nerves* act in the muscles that move the eyeball.
- VII. *Facial nerves* are found in the muscles of facial expression as well as in the salivary and tear glands. These nerves also carry sensory information from the tongue.
- VIII. *Vestibulocochlear nerves* carry hearing and equilibrium information from the inner ear to the brain for interpretation.
- IX. *Glossopharyngeal nerves* carry sensory information from the throat and tongue to the brain for interpretation. They also act in the muscles of the throat.
- X. *Vagus nerves* carry sensory information from the thoracic and abdominal organs to the brain for interpretation. These nerves are also found within the muscles in the throat, stomach, intestines, and heart.

- XI. *Accessory nerves* are found within the muscles of the throat, neck, back, and voice box.
- XII. *Hypoglossal nerves* are found within the muscles of the tongue.

Trigeminal nerve and dental anesthesia

Trigeminal means three twins, and the sensory distribution of the trigeminal nerve in the face is divided into three regions, each supplied by a branch of the nerve. The three branches— **ophthalmic**, **maxillary**, and **mandibular**—arise directly from the trigeminal ganglion, which serves the same function as the dorsal root ganglia of the spinal nerves. Only the mandibular branch contains motor axons, which bypass the trigeminal ganglion, much as the ventral root of a spinal nerve bypasses a dorsal root ganglion.

The maxillary and mandibular branches are important in dentistry. The maxillary nerve supplies sensory innervation to the maxillary teeth, palate, and gingiva. The mandibular branch supplies sensory innervation to the mandibular teeth, tongue, and gingiva. The various nerves innervating the teeth are referred to as **alveolar nerves**. The **superior alveolar nerves** to the maxillary teeth are derived from the maxillary branch of the trigeminal nerve, and the **inferior alveolar nerves** to the mandibular teeth are derived from the mandibular branch of the trigeminal nerve.

Dentists inject anesthetic to block sensory transmission by the alveolar nerves. The superior alveolar nerves are not usually anesthetized directly because they are difficult to approach with a needle. For this reason, the maxillary teeth are usually anesthetized locally by inserting the needle beneath the oral mucosa surrounding the teeth. The inferior alveolar nerve probably is anesthetized more often than any other nerve in the body. To anesthetize this nerve, the dentist inserts the needle somewhat posterior to the patient's last molar and extends the needle near where the mandibular branch of the trigeminal nerve enters the mandibular foramen. Several nondental nerves are usually anesthetized during an inferior alveolar block. The mental nerve, which supplies cutaneous innervation to the anterior lip and chin, is a distal branch of the inferior alveolar nerve. When the inferior alveolar nerve is blocked, the mental nerve is blocked also, resulting in a numb lip and chin. Nerves lying near the point where the inferior alveolar nerve enters the mandible often are also anesthetized during inferior alveolar anesthesia. For example, the lingual nerve can be anesthetized to produce a numb tongue. The facial nerve lies some distance from the inferior alveolar nerve, but in rare cases anesthetic can diffuse far enough posteriorly to anesthetize that nerve. The result is a temporary facial palsy, with the injected side of the face drooping because of flaccid muscles, which

disappears when the anesthesia wears off. If the facial nerve is cut by an improperly inserted needle, permanent facial palsy may occur.

Brain stem reflexes

Many of the brainstem reflexes are associated with cranial nerve function. The circuitry of most of these reflexes is too complex for our discussions. In general, these reflexes involve sensory input from the cranial nerves or spinal cord, as well as the motor output of the cranial nerves. Turning of the eyes toward a flash of light, a sudden noise, and a touch on the skin are examples of brainstem reflexes. Moving the eyes to track a moving object is another complex brainstem reflex. Some of the sensory neurons from cranial nerve VIII (vestibulocochlear) form a reflex arc with neurons of cranial nerves V (trigeminal) and VII (facial), which send axons to muscles of the middle ear and dampen the effects of very loud, sustained noises on delicate inner ear structures. Reflexes that occur during the process of chewing allow the jaws to react to foods of various hardness and protect the teeth from breakage. Both the sensory and motor components of the reflex arc are carried by cranial nerve V (trigeminal). Reflexes involving input through cranial nerve V (trigeminal) and output through cranial nerve XII (hypoglossal) move the tongue about to position food between the teeth for chewing and then move the tongue out of the way so that it is not bitten.

Brainstem reflexes are used to determine if the brainstem is functioning. Brainstem reflexes are mediated by cranial nerve sensory nuclei, cranial nerve motor nuclei, and reticular formation nuclei. The presence of a reflex indicates that sensory input reaches the brainstem and the nuclei are functioning to produce a response. The absence of a reflex is taken to indicate damage to the nuclei involved in the reflex.

Clinical correlates

Cranial Nerve Disorders

Trigeminal neuralgia, also called tic douloureux, involves the trigeminal nerve and consists of sharp bursts of pain in the face. This disorder often has a trigger point in or around the mouth, which, when touched, elicits the pain response in some other part of the face. The cause of trigeminal neuralgia is unknown. Facial palsy (called Bell palsy) is a unilateral paralysis of the facial muscles. The affected side of the face droops because of the absence of muscle tone. Although the cause of facial palsy is often unknown, it can result from inflammation of the facial nerve or a stroke or tumor in the cerebral cortex or brainstem. The facial nerve passes from deep to superficial through the parotid gland, and temporary facial palsy can result from inflammation of the parotid gland. Temporary facial palsy can even result from extreme cold in the face, where the superficial branches of the facial nerve are located.

Disorders of the cranial nerves can be determined using the following tests:

- The olfactory nerves (I) are tested by asking a patient to smell various substances.
 - Cranial nerves III, IV, and VI are tested by asking a patient to track the movement of the physician's finger. If a patient cannot move her eyeballs properly, there may be damage to one of these nerves. Recall that these nerves control the muscles that move the eyeballs.
 - Cranial nerve V controls the muscles needed for chewing. To assess this nerve, a patient is asked to clench his teeth. The physician then feels the jaw muscles. If they feel limp or weak, this nerve may be damaged.
 - If a person can no longer make facial expressions, then cranial nerve VII may be damaged. This nerve controls the muscles needed to make facial expressions.
- I. If a patient cannot extend his tongue and move it from side to side, cranial nerve XII may be damaged. This nerve controls tongue movement.

5.0 SUMMARY

- Cranial nerves are designated by Roman numerals (I–XII) or specific names.
- The two types of general functions are sensory and motor. Sensory includes special senses and general senses. Motor includes somatic motor and parasympathetic.
- The trigeminal nerve (V) has three branches—ophthalmic, maxillary, and mandibular. The superior alveolar nerve (branch of the maxillary) and the inferior alveolar nerve (branch of the mandibular) are used for dental anesthesia.
- Many reflexes are mediated through the brainstem. The brainstem is considered to be non-functional when reflexes at all levels of the brainstem are non-functional and there is no spontaneous breathing, which is mediated through the medulla oblongata

SELF – ASSESSMENT EXERCISE

1. The cranial nerve involved in chewing food is the
 - a. trochlear (IV).
 - b. trigeminal (V).
 - c. abducent (VI).
 - d. facial (VII).
 - e. vestibulocochlear (VIII).

2. The cranial nerve involved in moving the tongue is the
 - a. trigeminal (V).
 - b. facial (VII).
 - c. glossopharyngeal (IX).
 - d. accessory (XI).
 - e. hypoglossal (XII).
3. The cranial nerve involved in feeling a toothache is the
 - a. trochlear (IV).
 - b. trigeminal (V).
 - c. abducent (VI).
 - d. facial (VII).
 - e. vestibulocochlear (VIII).
4. From this list of cranial nerves:
 1. optic (II)
 2. oculomotor (III)
 3. trochlear (IV)
 4. trigeminal (V)
 5. abducent (VI)

Select the nerves that are involved in moving the eyes.

- a. 1,2,3 b. 1,2,4 c. 2,3,4 d. 2,4,5 e. 2,3,5

5. From this list of cranial nerves:
 1. trigeminal (V)
 2. facial (VII)
 3. glossopharyngeal (IX)
 4. vagus (X)
 5. hypoglossal (XII)

Select the nerves that innervate the salivary glands.

- a. 1,2 b. 2,3 c. 3,4 d. 4,5 e. 3,5

6. From this list of cranial nerves:
 1. oculomotor (III)
 2. trigeminal (V)
 3. facial (VII)
 4. vestibulocochlear (VIII)
 5. glossopharyngeal (IX)
 6. vagus (X)
7. Where do the cranial nerves attach to the brain?
8. Of all the cranial nerves, which is most important to a dentist?

UNIT 2 SPINAL NERVES

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Structure of nerves
- 4.0 Summary

1.0 INTRODUCTION

Spinal nerves are peripheral nerves that originate from the spinal cord. There are 31 pairs of spinal nerves: 8 pairs of cervical nerves (numbered C1 through C8), 12 pairs of thoracic nerves (numbered T1 through T12), 5 pairs of lumbar nerves (numbered L1 through L5), 5 pairs of sacral nerves (numbered S1 through S5), and one pair of coccygeal nerves (Co).

2.0 OBJECTIVES

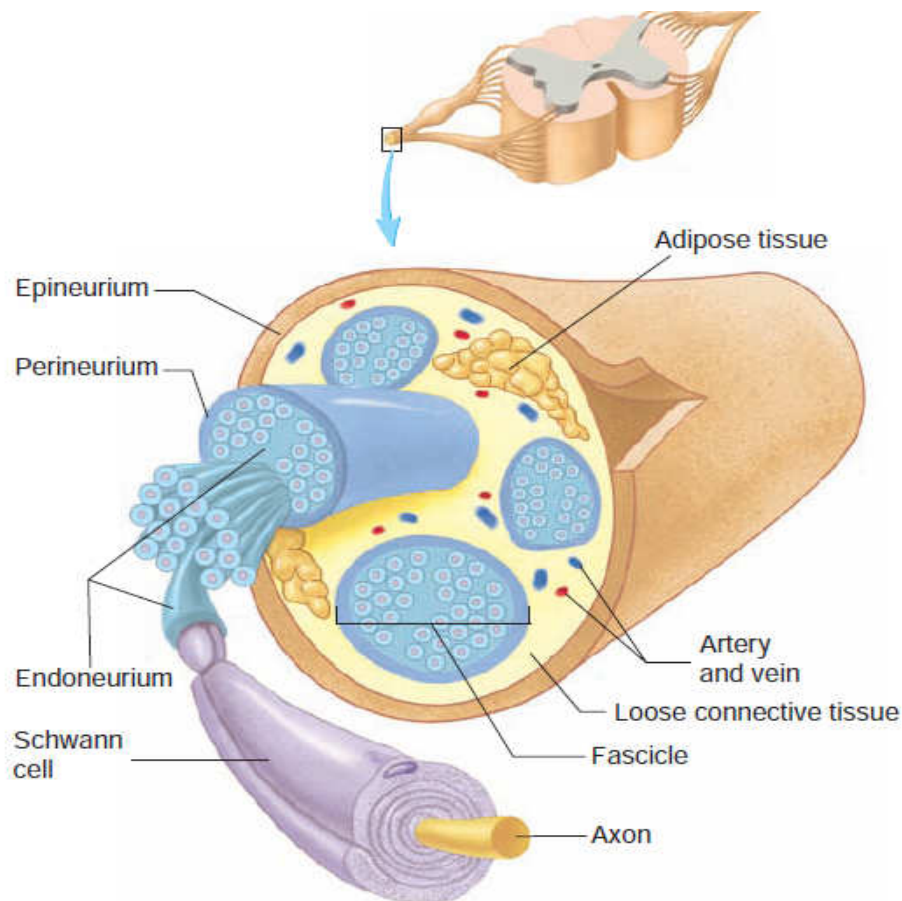
At the end of this unit, you should be able to:

- Locate and describe the spinal nerves.
- Discuss the spinal *reflex arc*.

3.0 MAIN CONTENT

3.1 Structure of nerves

Nerves consist of collections of axons, Schwann cells, and connective tissue in the PNS. Each axon, or nerve fiber, and its Schwann cell sheath are surrounded by a delicate connective tissue layer, the endoneurium. A heavier connective tissue layer, the perineurium, surrounds groups of axons to form nerve fascicles. A third layer of dense connective tissue, the epineurium, binds the nerve fascicles together to form a nerve. The connective tissue of the epineurium is continuous with the dura mater surrounding the CNS. An analogy of this relationship is a coat (the dura) and its sleeve (epineurium). The connective tissue layers of nerves make them tougher than the nerve tracts in the CNS.



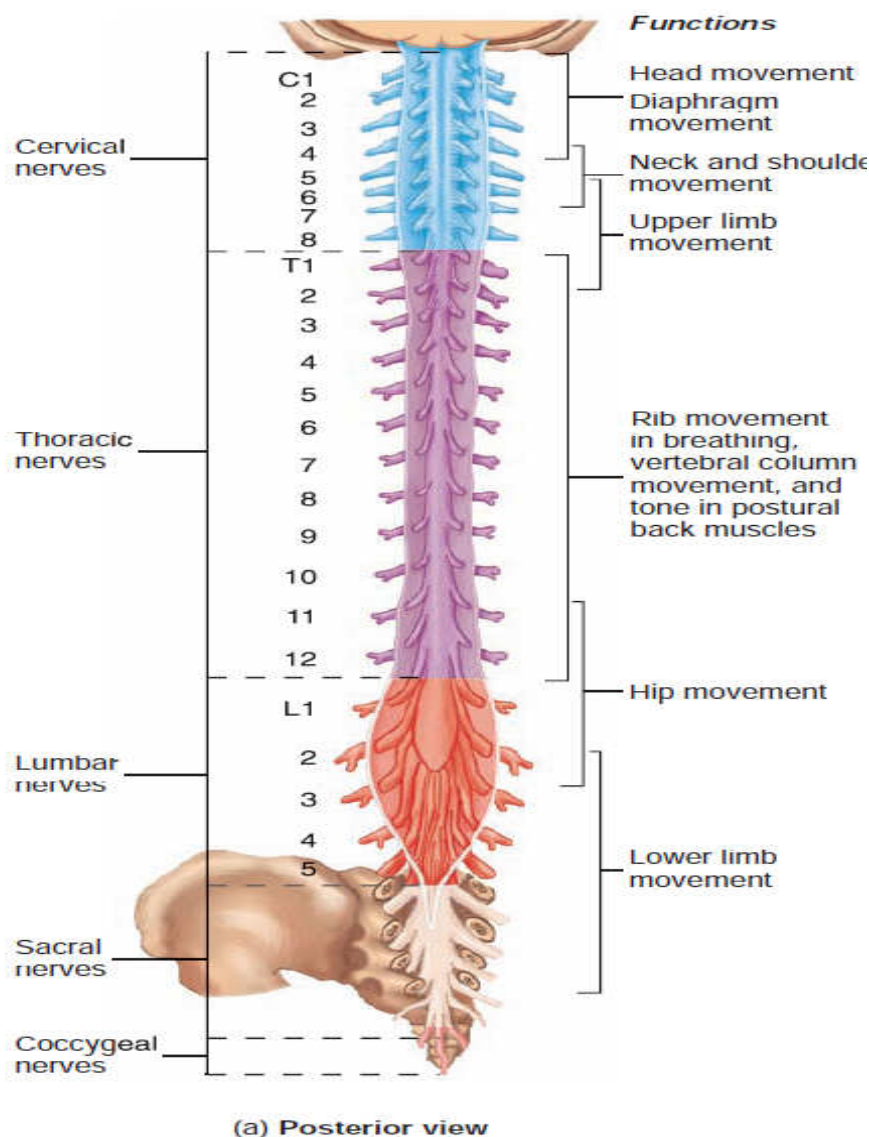
Structure of a nerve

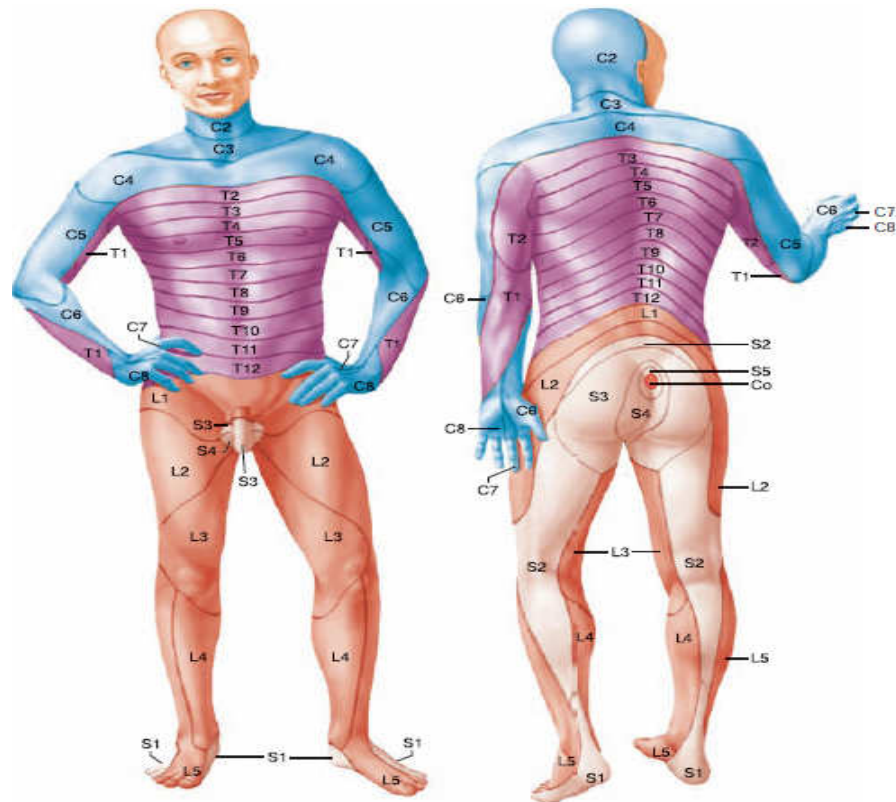
Distribution of nerves

There are 31 pairs of spinal nerves, each identified by a letter and a number. The letter indicates the region of the vertebral column from which the nerve emerges: C, cervical; T, thoracic; L, lumbar; and S, sacral. The single coccygeal nerve may be identified as Co, but typically receives no designation. The number indicates the location in each region where the nerve emerges from the vertebral column, with the smallest number always representing the most superior origin. For example, the most superior nerve exiting from the thoracic region of the vertebral column is designated T1. The cervical nerves are designated C1–C8, the thoracic nerves T1–T12, the lumbar nerves L1–L5, and the sacral nerves S1–S5. Note that the number of each type of nerve matches the number of each type of vertebrae, except for the cervical nerves. Although there are only 7 cervical vertebrae, there are 8 pairs of cervical spinal nerves because 2 pairs of nerves are associated with the first cervical vertebra. The first pair of cervical nerves exit the spinal cord between the skull and the first cervical vertebra and the second pair exit through the intervertebral foramina between the first and second cervical vertebrae. The nerves arising from each region of the spinal

cord and vertebral column supply specific regions of the body. Not surprisingly, the nerves supply skeletal muscles in a top to bottom pattern: the cervical nerves supply muscles of the head, neck, shoulders, and upper limbs; the thoracic nerves supply muscles of the upper limbs, thorax, vertebral column, and hips; the lumbar nerves supply muscles of the vertebral column, hips, and lower limbs; and the sacral nerves supply muscles of the lower limbs.

Each of the spinal nerves except C1 has a specific cutaneous sensory distribution. A **dermatome** is the area of skin supplied with sensory innervation by a pair of spinal nerves. A **dermatomal map** shows the distribution of all the dermatomes.





dermatomes of the skin

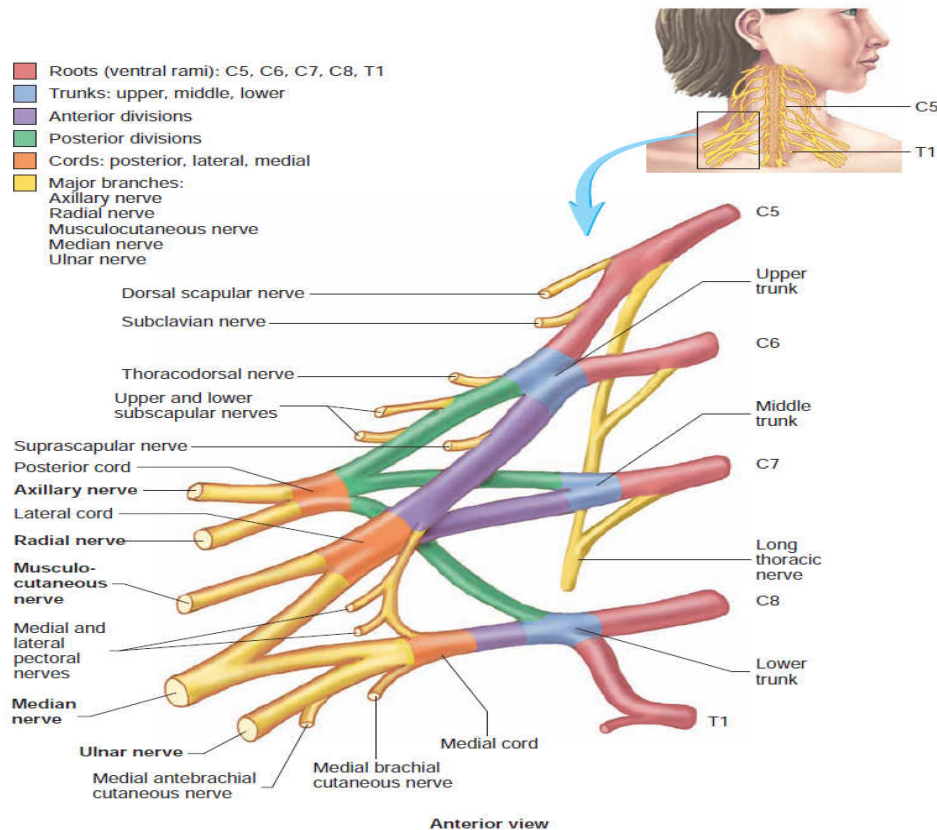
Major nerve plexuses

Spinal nerves arise from the spinal cord as rootlets. The rootlets combine to form the ventral and dorsal roots, which combine to form the spinal nerve. The spinal nerve divides to form branches called rami. Each spinal nerve has a dorsal and a ventral ramus. Additional communicating rami are associated with the sympathetic division of the ANS. The dorsal rami innervate most of the deep muscles of the dorsal trunk responsible for movement of the vertebral column. They also innervate the connective tissue and skin near the midline of the back.

The ventral rami are distributed in two ways. In the thoracic region, the ventral rami form intercostal (between ribs) nerves, which extend along the inferior margin of each rib and innervate the intercostal muscles and the skin over the thorax. The ventral rami of the cervical, lumbar, sacral, and coccygeal spinal nerves form plexuses. The term *plexus* means braid and describes the organization produced by the intermingling of the nerves. The brachial plexus will be used to illustrate how the spinal nerves intermingle within a plexus and then give rise to nerves that are distributed throughout the body. The brachial plexus is the most complicated plexus, and other plexuses have similar, but simpler, interconnections.

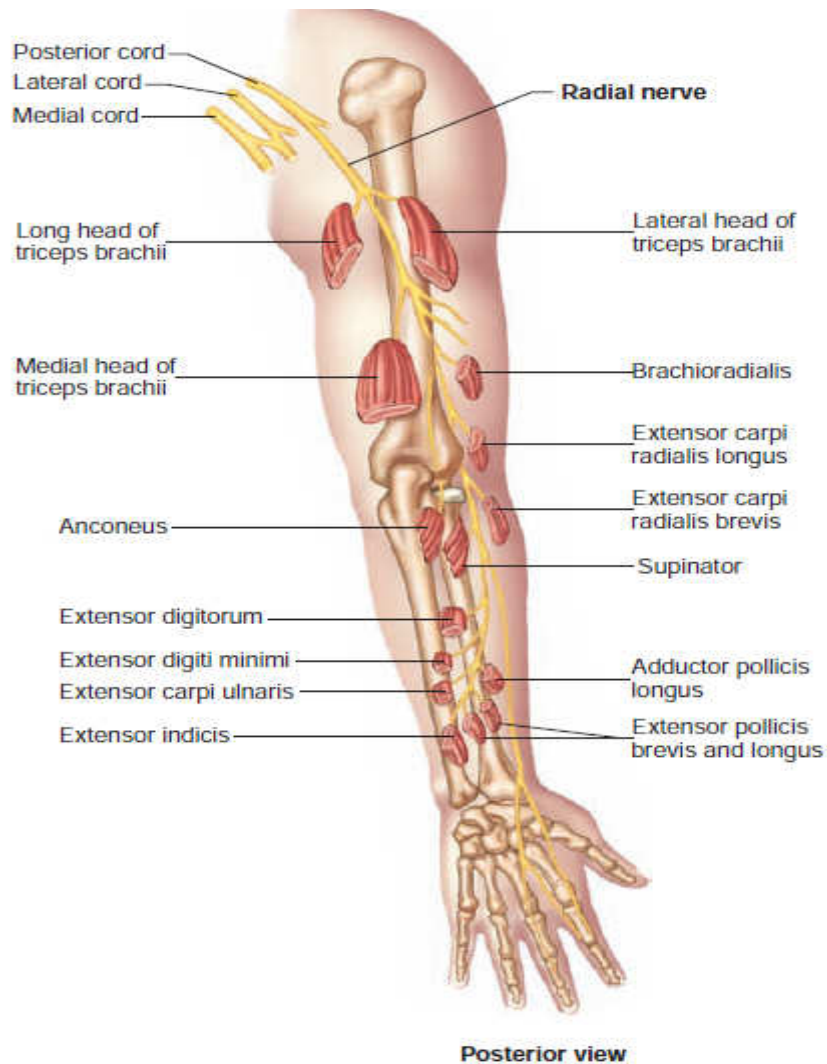
Brachial plexus

The brachial plexus originates from spinal nerves C5–T1. The ventral rami of the spinal nerves are called the roots of the plexus. These roots should not be confused with the dorsal and ventral roots from the spinal cord, which are more medial. The five roots join to form three trunks, which separate into six divisions and then join again to create three cords (posterior, lateral, and medial) from which five major **branches**, or nerves, emerge. The five major nerves are the **axillary, radial, musculocutaneous, ulnar, and median nerves**. They supply the upper limb.



The brachial plexus

The distribution of the radial nerve to skeletal muscles of the upper limb is shown below.



the radial nerve

In a similar fashion, nerves arising from plexuses are distributed to skeletal muscles throughout the body. Nerves arising from plexuses also supply the skin. Note that axons of spinal nerves C5–T1 can reach the radial nerve through the interconnections of the brachial plexus. Thus, axons in the radial nerve arise from levels C5–T1 of the spinal cord. The cutaneous distribution of a nerve arising from a plexus is different from the cutaneous distribution (dermatome) arising from a spinal nerve because the plexus nerve consists of axons from more than one level of the spinal cord.

Cervical Plexus

The **cervical plexus** is a relatively small plexus originating from spinal nerves C1–C4. Branches derived from this plexus innervate superficial neck structures, including several of the muscles attached to the hyoid bone. The cervical plexus innervates the skin of the neck and posterior portion of the head. The **phrenic nerve**, which originates from spinal nerves C3–C5, is derived from both the cervical and brachial plexus.

The phrenic nerves descend along each side of the neck to enter the thorax. They descend along the sides of the mediastinum to reach the diaphragm, which they innervate. Contraction of the diaphragm is largely responsible for the ability to breathe.

Lumbar and Sacral Plexuses

The **lumbar plexus** originates from the ventral rami of spinal nerves L1–L4 and the **sacral plexus** from L4 to S4. Because of their close, overlapping relationship and their similar distribution, however, the two plexuses often are considered together as a single **lumbosacral plexus** (L1–S4). Four major nerves exit the lumbosacral plexus and enter the lower limb: the **obturator, femoral, tibial, and common fibular (peroneal) nerves**. Other lumbosacral nerves supply muscles of the lower back, hip, and lower abdomen and the skin of the hip and thigh. The tibial and common fibular nerves originate from spinal segments L4–S3 and are bound together within a connective tissue sheath for the length of the thigh. The two nerves bound together are referred to jointly as the **sciatic nerve**. The sciatic nerve is by far the largest peripheral nerve in the body. It passes through the greater sciatic notch in the coxal bone and descends in the posterior thigh to the back of the knee, where the tibial and common fibular nerves separate from each other. The sciatic nerve supplies the posterior thigh muscles, the tibial nerve supplies the posterior compartment of the leg, and the common fibular supplies the anterior and lateral compartments. Branches of the two nerves combine to form the **sural nerve**, which supplies the skin on the posterior and lateral leg.

Coccygeal Plexus

The **coccygeal plexus** is a very small plexus formed from the ventral rami of spinal nerve S5 and the coccygeal nerve (Co). This small plexus supplies the muscles of the pelvic floor and the skin over the coccyx. The dorsal rami of the coccygeal nerves also innervate some skin over the coccyx.

Clinical correlates

Radial Nerve Damage

The radial nerve lies near the humerus in the axilla. When crutches are used improperly, the crutch is pushed tightly into the axilla. This can damage the radial nerve by compressing it against the humerus, resulting in **crutch paralysis**. In this disorder, muscles innervated by the radial nerve lose their function. The major symptom of radial nerve damage is **wrist drop**, an inability to extend the wrist when the pronated forearm is held parallel to the ground. Consequently, the wrist drops into a flexed position. The radial nerve can be permanently damaged by a

fracture of the humerus in the proximal part of the arm. A sharp edge of the broken bone may cut the nerve, resulting in permanent paralysis unless the nerve is surgically repaired. Because of potential damage to the radial nerve, a broken humerus should be treated very carefully.

Ulnar Nerve Damage

The ulnar nerve is the most easily damaged of all the peripheral nerves, but such damage is almost always temporary. Slight damage to the ulnar nerve may occur where it passes posterior to the medial epicondyle of the humerus. The nerve can be felt just below the skin at this point, and, if this region of the elbow is banged against a hard object, temporary ulnar nerve damage may occur. This damage results in painful tingling sensations radiating down the ulnar side of the forearm and hand. Because of this sensation, this area of the elbow is often called the **funny bone** or **crazy bone**.

Median Nerve Damage

Damage to the median nerve occurs most commonly where it enters the wrist through the **carpal tunnel**. This tunnel is created by the concave organization of the carpal bones and the retinaculum on the anterior surface of the wrist. None of the connective tissue components of the carpal tunnel expand readily. The tendons passing through the carpal tunnel may become inflamed and enlarged as a result of repetitive movements. This inflammation can produce pressure within the carpal tunnel, thereby compressing the median nerve and resulting in numbness, tingling, and pain in the fingers. The thenar muscles, innervated by the median nerve, have reduced function, resulting in weakness in thumb flexion and opposition. This condition is referred to as **carpal tunnel syndrome**. Carpal tunnel syndrome is common among people who perform repetitive movements of the wrists and fingers, such as keyboard operators. Surgery is often required to relieve the pressure.

4.0 SUMMARY

- Individual axons are surrounded by the endoneurium. Groups of axons, called fascicles, are bound together by the perineurium. The fascicles form the nerve and are held together by the epineurium.
- Eight cervical, 12 thoracic, 5 lumbar, 5 sacral pairs, and 1 coccygeal pair make up the spinal nerves. The distribution of spinal nerves to skeletal muscles has a top-to-bottom pattern in which superior nerves supply superior muscles and inferior nerves supply inferior muscles.

- Spinal nerves have specific cutaneous distributions called dermatomes. Spinal nerves branch to form rami. The dorsal rami supply the muscles and skin near the midline of the back.
- The five major plexuses are the cervical (C1–C4), brachial (C5–T1), lumbar (L1–L4), sacral (L4–S4), and coccygeal (S5 and coccygeal nerve). The lumbar and sacral plexuses are often considered together as the lumbosacral plexus. A major nerve of the cervical plexus is the phrenic nerve. The major nerves of the brachial plexus are the axillary, radial, musculocutaneous, ulnar, and median nerves.

Activity

- I. In the histology laboratory, observe and recognize the microscopic slides of a nerve and its connective tissue coverings.
- II. In the gross anatomy laboratory/ dissection room, identify the major spinal nerve plexuses – cervical, brachial, lumbar and sacral plexuses.

SELF – ASSESSMENT EXERCISE

1. Which of these is a correct count of the spinal nerves?
 - a. 9 cervical, 12 thoracic, 5 lumbar, 5 sacral, 1 coccygeal
 - b. 8 cervical, 12 thoracic, 5 lumbar, 5 sacral, 1 coccygeal
 - c. 7 cervical, 12 thoracic, 5 lumbar, 5 sacral, 1 coccygeal
 - d. 8 cervical, 11 thoracic, 4 lumbar, 6 sacral, 1 coccygeal
 - e. 7 cervical, 11 thoracic, 5 lumbar, 6 sacral, 1 coccygeal
2. Given these structures:
 1. dorsal ramus
 2. dorsal root
 3. plexus
 4. ventral ramus
 5. ventral root

Choose the arrangement that lists the structures in the order that an action potential passes through them, given that the action potential originates in the spinal cord and propagates to a peripheral nerve.

- a. 2,1,3
 - b. 2,3,1
 - c. 3,4,5
 - d. 5,3,4
 - e. 5,4,3
3. Damage to the dorsal ramus of a spinal nerve results in
 - a. loss of sensation.
 - b. loss of motor function.
 - c. both a and b.

4. A dermatome
 - a. is the area of skin supplied by a pair of spinal nerves.
 - b. exists for each spinal nerve except C1.
 - c. can be used to locate the site of spinal cord or nerve root damage.
 - d. all of the above.
5. A collection of spinal nerves that join together after leaving the spinal cord is called a
 - a. ganglion.
 - b. nucleus.
 - c. projection nerve.
 - d. plexus.
6. Which of these nerves arises from the cervical plexus?
 - a. median
 - b. musculocutaneous
 - c. phrenic
 - d. obturator
 - e. ulnar sheath.
7. The sciatic nerve is actually two nerves combined within the same. The two nerves are the
 - a. femoral and obturator.
 - b. femoral and gluteal.
 - c. common fibular (peroneal) and tibial.
 - d. common fibular (peroneal) and obturator.
 - e. tibial and gluteal.

UNIT 3 AUTONOMIC NERVOUS SYSTEM

CONTENTS

- 1.0 Introduction
- 2.0 Objectives
- 3.0 Main Content
 - 3.1 Difference between PNS and ANS
- 4.0 Summary

1.0 INTRODUCTION

During a picnic on a sunny spring day, it is easy to concentrate on the delicious food and the pleasant surroundings. The maintenance of homeostasis requires no conscious thought. The autonomic nervous system (ANS) helps keep body temperature at a constant level by controlling the activity of sweat glands and the amount of blood flowing through the skin. The ANS helps regulate the complex activities necessary for the digestion of food. The movement of absorbed nutrients to tissues is possible because the ANS controls heart rate, which helps maintain the blood pressure necessary to deliver blood to tissues. Without the ANS, all of the activities necessary to maintain homeostasis would be overwhelming.

A functional knowledge of the ANS enables you to predict general responses to a variety of stimuli, explain responses to changes in environmental conditions, comprehend symptoms that result from abnormal autonomic functions, and understand how drugs affect the ANS.

2.0 OBJECTIVES

At the end of this unit, you will be able to:

- distinguish further between the ANS and the somatic system.
- predict the effects of sympathetic versus parasympathetic stimulation on specific organs.

3.0 MAIN CONTENTS

3.1 Difference between PNS and ANS

The peripheral nervous system (PNS) is composed of sensory and motor neurons. Sensory neurons carry action potentials from the periphery to the central nervous system (CNS), and motor neurons carry action

potentials from the CNS to the periphery. Motor neurons are either somatic motor neurons, which innervate skeletal muscle, or autonomic motor neurons, which innervate smooth muscle, cardiac muscle, and glands.

Although axons of autonomic, somatic, and sensory neurons are in the same nerves, the proportion varies from nerve to nerve. For example, nerves innervating smooth muscle, cardiac muscle, and glands, such as the vagus nerves, consist primarily of axons of autonomic motor neurons and sensory neurons. Nerves innervating skeletal muscles, such as the sciatic nerves, consist primarily of axons of somatic motor neurons and sensory neurons. Some cranial nerves, such as the olfactory, optic, and vestibulocochlear nerves, are composed entirely of axons of sensory neurons. The cell bodies of somatic motor neurons are in the CNS, and their axons extend from the CNS to skeletal muscle. The ANS, on the other hand, has two neurons in a series extending between the CNS and the organs innervated.

The first neurons of the series are called preganglionic neurons. Their cell bodies are located in the CNS within either the brainstem or the lateral part of the spinal cord gray matter, and their axons extend to autonomic ganglia located outside the CNS. The autonomic ganglia contain the cell bodies of the second neurons of the series, which are called postganglionic neurons. The preganglionic neurons synapse with the postganglionic neurons in the autonomic ganglia. The axons of the postganglionic neurons extend from autonomic ganglia to effector organs, where they synapse with their target tissues.

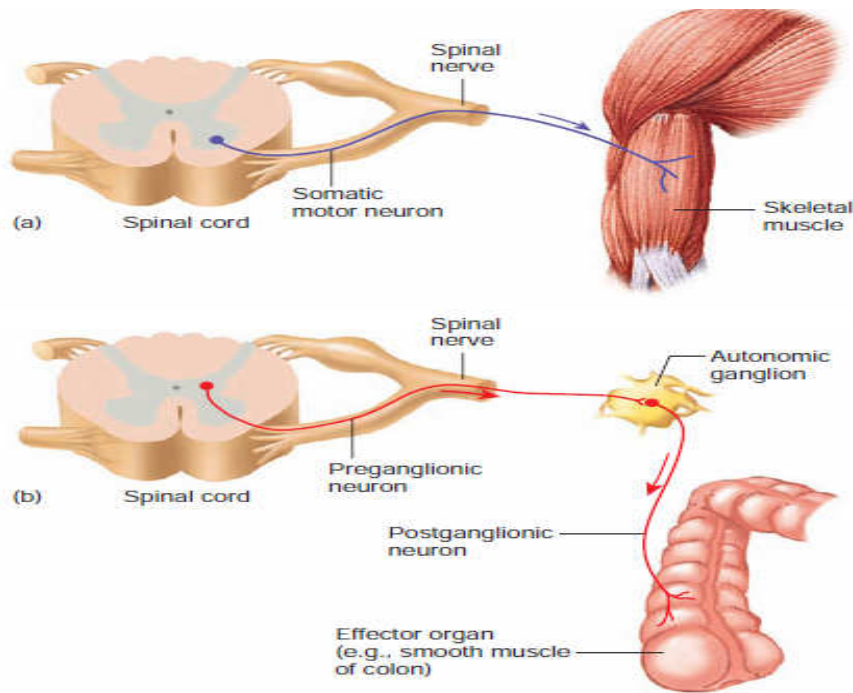


Figure 14.1 Organization of Somatic Motor and Autonomic Nervous System Neurons

(a) The cell body of the somatic motor neuron is in the CNS, and its axon extends to the skeletal muscle. (b) The cell body of the preganglionic neuron is in the CNS, and its axon extends to the autonomic ganglion and synapses with the postganglionic neuron. The postganglionic neuron extends to and synapses with its effector organ.

Anatomy of the ANS

The ANS is subdivided into the **sympathetic** and the **parasympathetic divisions** and the **enteric nervous system (ENS)**. The sympathetic and parasympathetic divisions differ structurally in (1) the location of their preganglionic neuron cell bodies within the CNS and (2) the location of their autonomic ganglia. The enteric nervous system is a complex network of neuron cell bodies and axons within the wall of the digestive tract. An important part of this network is sympathetic and parasympathetic neurons. For this reason, the enteric nervous system is considered to be part of the ANS.

Sympathetic Division

Cell bodies of sympathetic preganglionic neurons are in the lateral horns of the spinal cord gray matter between the first thoracic (T1) and the second lumbar (L2) segments. The sympathetic division is sometimes called the **thoracolumbar division** because of the location of the preganglionic cell bodies.

The axons of preganglionic neurons are small in diameter and myelinated. The short connection between a spinal nerve and a sympathetic chain ganglion through which the preganglionic axons pass is called a white ramus communicans because of the whitish colour of

the myelinated axons. Sympathetic axons exit the sympathetic chain ganglia by the following four routes:

1. *Spinal nerves* - Preganglionic axons synapse with postganglionic neurons in sympathetic chain ganglia. They can synapse at the same level that the preganglionic axons enter the sympathetic chain, or they can pass superiorly or inferiorly through one or more ganglia and synapse with postganglionic neurons in a sympathetic chain ganglion at a different level.

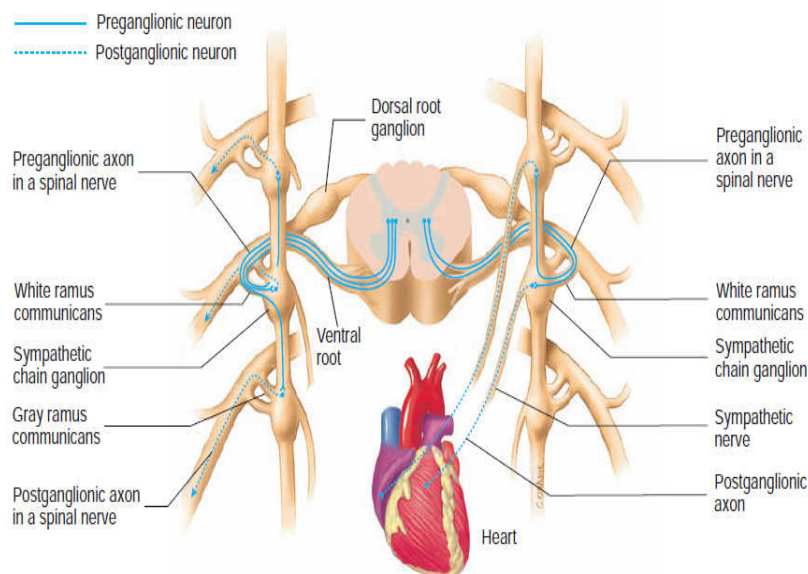
Axons of the postganglionic neurons pass through a gray ramus communicans and re-enter a spinal nerve. Postganglionic axons are not myelinated, thereby giving the gray ramus communicans its grayish colour. All spinal nerves receive postganglionic axons from a gray ramus communicans. The postganglionic axons then project through the spinal nerve to smooth muscle and glands located in the skin and skeletal muscles.

2. *Sympathetic nerves* - Preganglionic axons enter the sympathetic chain and synapse in a sympathetic chain ganglion at the same or a different level with postganglionic neurons. The postganglionic axons leaving the sympathetic chain ganglion form sympathetic nerves, which supply organs in the thoracic cavity.
3. *Splanchnic nerves* - Some preganglionic axons enter sympathetic chain ganglia and, without synapsing, exit at the same or a different level to form splanchnic nerves. Those preganglionic axons extend to collateral ganglia, where they synapse with postganglionic neurons. Axons of the postganglionic neurons leave the collateral ganglia through small nerves that extend to effector organs in the abdominopelvic cavity.
4. *Innervation to the adrenal gland* – The splanchnic nerve innervation to the adrenal glands is different from other ANS nerves because it consists of only preganglionic neurons. Axons of the preganglionic neurons do not synapse in sympathetic chain ganglia or in collateral ganglia. Instead, the axons pass through those ganglia and synapse with cells in the adrenal medulla. The adrenal medulla is the inner portion of the adrenal gland; it consists of specialized cells derived during embryonic development from neural crest cells, which are the same population of cells that gives rise to the postganglionic cells of the ANS. Adrenal medullary cells are round, have no axons or dendrites, and are divided into two groups. About 80% of the cells secrete epinephrine also called adrenaline and about 20% secrete norepinephrine, also called noradrenaline. Stimulation of

these cells by preganglionic axons causes the release of epinephrine and norepinephrine. These substances circulate in the blood and affect all tissues having receptors to which they can bind. The general response to epinephrine and norepinephrine released from the adrenal medulla is to prepare the individual for physical activity. Secretions of the adrenal medulla are considered hormones because they are released into the general circulation and travel some distance to the tissues in which they have their effect.

Parasympathetic Division

The cell bodies of parasympathetic preganglionic neurons are either within cranial nerve nuclei in the brainstem or within the lateral parts of the gray matter in the sacral region of the spinal cord from S2 to S4. For that reason, the parasympathetic division is sometimes called the **craniosacral division**. Axons of the parasympathetic preganglionic neurons from the brain are in **cranial nerves III, VII, IX, and X** and from the spinal cord in **pelvic splanchnic nerves**. The preganglionic axons course through these nerves to **terminal ganglia**, where they synapse with postganglionic neurons. The axons of the postganglionic neurons extend relatively short distances from the terminal ganglia to the effector organs. The terminal ganglia are either near or embedded within the walls of the organs innervated by the parasympathetic neurons. Many of the parasympathetic ganglia are small but some, such as those in the wall of the digestive tract, are large.



(a) Preganglionic axons from a spinal nerve pass through a white ramus communicans into a sympathetic chain ganglion. Some axons synapse with a postganglionic neuron at the level of entry; others ascend or descend to other levels before synapsing. Each postganglionic axon exits the sympathetic chain through a gray ramus communicans and enters a spinal nerve.

(b) Part (b) is like part (a), except that each postganglionic neuron exits a sympathetic chain ganglion through a sympathetic nerve.

Physiology of ANS

Neurotransmitters

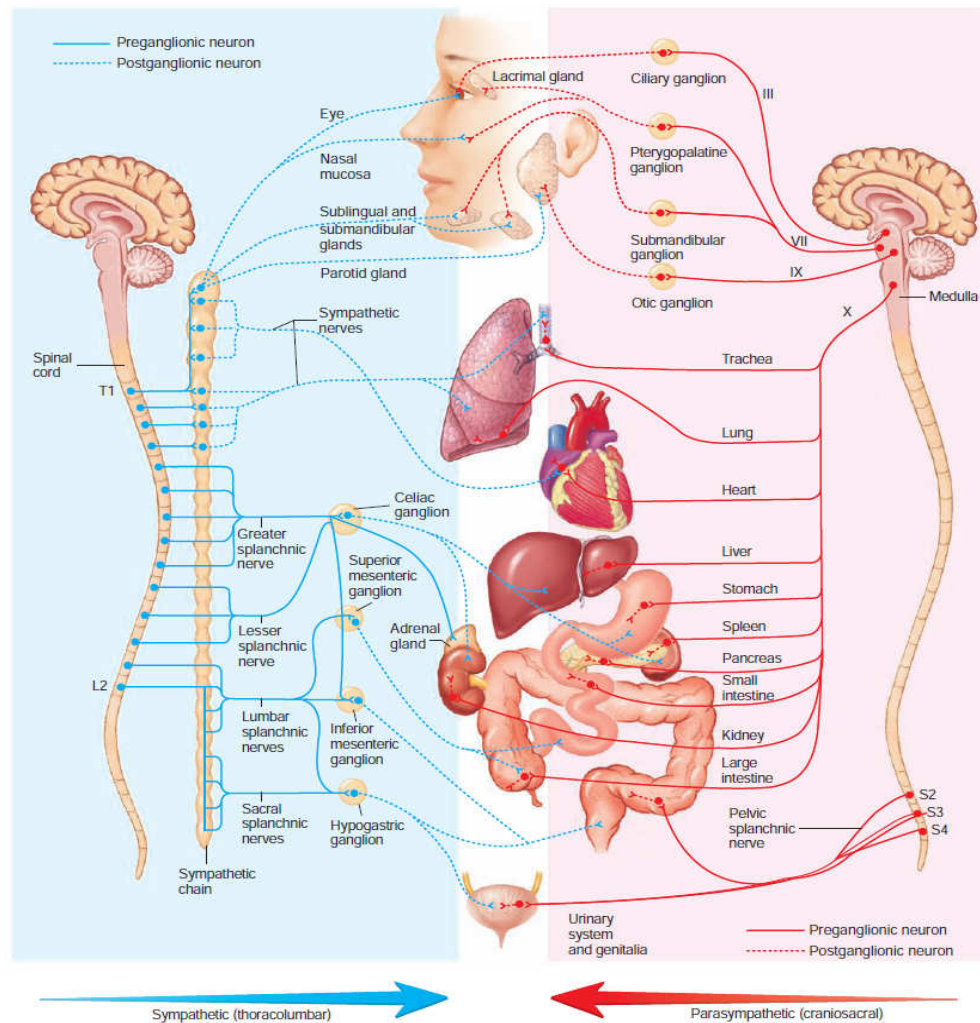
Sympathetic and parasympathetic nerve endings secrete one of two neurotransmitters. If the neuron secretes acetylcholine, it is a **cholinergic neuron**; if it secretes norepinephrine (or epinephrine), it is an **adrenergic neuron**. Adrenergic neurons are so named because at one time it was believed that they secreted adrenaline, which is another name for epinephrine. All preganglionic neurons of the sympathetic and parasympathetic divisions and all postganglionic neurons of the parasympathetic division are cholinergic. Almost all postganglionic neurons of the sympathetic division are adrenergic, but a few postganglionic neurons that innervate thermoregulatory sweat glands are cholinergic. In recent years, substances in addition to the regular neurotransmitters have been extracted from ANS neurons. These substances include nitric oxide; fatty acids, such as eicosanoids; peptides, such as gastrin, somatostatin, cholecystokinin, vasoactive intestinal peptide, enkephalins, and substance P; and monoamines, such as dopamine, serotonin, and histamine. The specific role that many of these compounds play in the regulation of the ANS is unclear, but they appear to function as either neurotransmitters or neuromodulator substances.

Receptors

Receptors for acetylcholine and norepinephrine are located in the plasma membrane of certain cells. The combination of neurotransmitter and receptor functions as a signal to cells, causing them to respond. Depending on the type of cell, the response is excitatory or inhibitory.

Cholinergic Receptors

Cholinergic receptors are receptors to which acetylcholine binds. They have two major, structurally different forms. **Nicotinic receptors** bind to nicotine, an alkaloid substance found in tobacco, and **muscarinic receptors** bind to muscarine, an alkaloid extracted from some poisonous mushrooms. Although nicotine and muscarine are not naturally in the human body, they demonstrate the differences between the two classes of cholinergic receptors. Nicotine binds to nicotinic receptors but not to muscarinic receptors, whereas muscarine binds to muscarinic receptors but not to nicotinic receptors. On the other hand, acetylcholine binds to and activates both types of receptors. The membranes of all postganglionic neurons in autonomic ganglia and the membranes of skeletal muscle cells have nicotinic receptors. The membranes of effector cells that respond to acetylcholine released from postganglionic neurons have muscarinic receptors.



ANS supply to organs of the body

Regulation of ANS

Much of the regulation of structures by the ANS occurs through autonomic reflexes, but input from the cerebrum, the hypothalamus, and other areas of the brain allows conscious thoughts and actions, emotions, and other CNS activities to influence autonomic functions. Without the regulatory activity of the ANS, an individual has limited ability to maintain homeostasis. **Autonomic reflexes**, like other reflexes, involve sensory receptors, sensory neurons, interneurons, motor neurons, and effector cells. For example, **baroreceptors** (stretch receptors) in the walls of large arteries near the heart detect changes in blood pressure, and sensory neurons transmit information from the baroreceptors through the glossopharyngeal and vagus nerves to the medulla oblongata. Interneurons in the medulla oblongata integrate the information, and action potentials are produced in autonomic neurons that extend to the heart. If baroreceptors detect a change in blood pressure, autonomic reflexes change heart rate, which returns blood pressure to normal.

A sudden increase in blood pressure initiates a parasympathetic reflex, which inhibits cardiac muscle cells and reduces heart rate, thus bringing blood pressure down toward its normal value. Conversely, a sudden decrease in blood pressure initiates a sympathetic reflex, which stimulates the heart to increase its rate and force of contraction, thus increasing blood pressure. Other autonomic reflexes participate in the regulation of blood pressure. For example, numerous sympathetic neurons transmit a low but relatively constant frequency of action potentials that stimulate blood vessels throughout the body, keeping them partially constricted. If the vessels constrict further, blood pressure increases; if they dilate, blood pressure decreases. Thus, altering the frequency of action potentials delivered to blood vessels along sympathetic neurons can either raise or lower blood pressure.

The brainstem and the spinal cord contain important autonomic reflex centers responsible for maintaining homeostasis. The hypothalamus, however, is in overall control of the ANS. Almost any type of autonomic response can be evoked by stimulating a part of the hypothalamus, which in turn stimulates ANS centers in the brainstem or spinal cord. Although there is overlap, stimulation of the posterior hypothalamus produces sympathetic responses, whereas stimulation of the anterior hypothalamus produces parasympathetic responses. In addition, the hypothalamus monitors and controls body temperature.

The hypothalamus has connections with the cerebrum and is an important part of the limbic system, which plays an important role in emotions. The hypothalamus integrates thoughts and emotions to produce ANS responses. Pleasant thoughts of a delicious banquet initiate increased secretion by salivary glands and by glands within the stomach and increased smooth muscle contractions within the digestive system. These responses are controlled by parasympathetic neurons. Emotions such as anger increase blood pressure by increasing heart rate and constricting blood vessels through sympathetic stimulation. The enteric nervous system is involved with autonomic and local reflexes that regulate the activity of the digestive tract. For example, in an ANS reflex, sensory neurons detecting stretch of the digestive tract wall send action potentials to the CNS. In response, the CNS sends action potentials out the ANS, causing smooth muscle in the digestive tract wall to contract.

The neurons of the enteric nervous system also operate independently of the CNS to produce local reflexes. A **local reflex** does not involve the CNS, but it does produce an involuntary, unconscious, stereotypic response to a stimulus. For example, sensory neurons not connected to the CNS detect stretch of the digestive tract wall. These sensory neurons

send action potentials through the enteric plexuses to motor neurons, causing smooth muscle contraction or relaxation.

Clinical correlates

I. Effects of Spinal Cord Injury on ANS Functions

Spinal cord injury can damage nerve tracts, resulting in the loss of sensation and motor control below the level of the injury. Spinal cord injury also interrupts the control of autonomic neurons by ANS centers in the brain. For the parasympathetic division, effector organs innervated through the sacral region of the spinal cord are affected, but most effector organs still have normal parasympathetic function because they are innervated by the vagus nerve. For the sympathetic division, brain control of sympathetic neurons is lost below the site of the injury. The higher the level of injury, the greater the number of body parts affected.

Immediately after spinal cord injury, spinal cord reflexes below the level of the injury are lost, including ANS reflexes. With time, the reflex centers in the spinal cord become functional again. This recovery is particularly important for reflexes involving urination and defecation. Autonomic reflexes mediated through the vagus nerves or the enteric nervous system are not affected by spinal cord injury.

II. Dopamine and the Treatment of Shock

Norepinephrine is produced from a precursor molecule called dopamine. Certain sympathetic neurons release dopamine, which binds to dopamine receptors. Dopamine is structurally similar to norepinephrine and it binds to beta receptors. Dopamine hydrochloride has been used successfully to treat circulatory shock because it can bind to dopamine receptors in kidney blood vessels. The resulting vasodilation increases blood flow to the kidneys and prevents kidney damage. At the same time, dopamine can bind to beta receptors in the heart, causing stronger contractions.

4.0 SUMMARY

- The cell bodies of somatic motor neurons are located in the CNS, and their axons extend to skeletal muscles, where they have an excitatory effect that usually is controlled consciously.
- The cell bodies of the preganglionic neurons of the ANS are located in the CNS and extend to ganglia, where they synapse with postganglionic neurons. The postganglionic axons extend to smooth muscle, cardiac muscle, or glands and have an excitatory or inhibitory effect, which usually is controlled unconsciously.

- Preganglionic cell bodies are in the lateral horns of the spinal cord gray matter from T1 to L2. Preganglionic axons pass through the ventral roots to the white rami communicantes to the sympathetic chain ganglia.
- Preganglionic cell bodies are in nuclei in the brainstem or the lateral parts of the spinal cord gray matter from S2 to S4. Preganglionic axons from the brain pass to ganglia through cranial nerves. Preganglionic axons from the sacral region pass through the pelvic splanchnic nerves to the ganglia. Preganglionic axons pass to terminal ganglia within the wall of or near the organ that is innervated.
- Sympathetic, parasympathetic, and sensory neurons intermingle in autonomic nerve plexuses. Sympathetic axons reach organs through spinal, sympathetic, and splanchnic nerves.
- Parasympathetic axons reach organs through cranial and pelvic splanchnic nerves. Sensory neurons run alongside sympathetic and parasympathetic neurons within nerves and nerve plexuses.
- Acetylcholine is released by cholinergic neurons (all preganglionic neurons, all parasympathetic postganglionic neurons, and some sympathetic postganglionic neurons).
- Norepinephrine is released by adrenergic neurons (most sympathetic postganglionic neurons). Acetylcholine binds to nicotinic receptors (found in all postganglionic neurons) and muscarinic receptors (found in all parasympathetic and some sympathetic effector organs). Norepinephrine and epinephrine bind to alpha and beta receptors (found in most sympathetic effector organs).
- Activation of nicotinic receptors is excitatory, whereas activation of muscarinic, alpha, and beta receptors is either excitatory or inhibitory.

Activity

In the anatomy museum, locate and examine any parts of the autonomic nervous system on models available.

SELF – ASSESSMENT EXERCISE

1. Given these phrases:
 1. neuron cell bodies in the nuclei of cranial nerves
 2. neuron cell bodies in the lateral gray matter of the spinal cord (S2–S4)
 3. two synapses between the CNS and effector organs
 4. regulates smooth muscle

Which of the phrases are true for the autonomic nervous system?

a. 1,3 b. 2,4 c. 1,2,3 d. 2,3,4 e. 1,2,3,4

2. Given these structures:

1. gray ramus communicans
2. white ramus communicans
3. sympathetic chain ganglion

Choose the arrangement that lists the structures in the order an action potential passes through them from a spinal nerve to an effector organ.

a. 1,2,3 b. 1,3,2 c. 2,1,3 d. 2,3,1 e. 3,2,1

3. Given these structures:

1. collateral ganglion
2. sympathetic chain ganglion
3. white ramus communicans
4. splanchnic nerve

Choose the arrangement that lists the structures in the order an action potential travels through them on the way from a spinal nerve to an effector organ.

a. 1,3,2,4 b. 1,4,2,3 c. 3,1,4,2 d. 3,2,4,1 e. 4,3,1,2

4. The white ramus communicans contains

- a. preganglionic sympathetic fibers.
- b. postganglionic sympathetic fibers.
- c. preganglionic parasympathetic fibers.
- d. postganglionic parasympathetic fibers.

5. The cell bodies of the postganglionic neurons of the sympathetic division are located in the

- a. sympathetic chain ganglia
- b. collateral ganglia.
- c. terminal ganglia.
- d. dorsal root ganglia.
- e. both a and b.

6. Splanchnic nerves

- a. are part of the parasympathetic division.
- b. have preganglionic neurons that synapse in the collateral ganglia.
- c. exit from the cervical region of the spinal cord.
- d. travel from the spinal cord to the sympathetic chain ganglia.
- e. all of the above.

7. Which of the following statements regarding the adrenal gland is true?

- a. The parasympathetic division stimulates the adrenal gland to release acetylcholine.
 - b. The parasympathetic division stimulates the adrenal gland to release epinephrine.
 - c. The sympathetic division stimulates the adrenal gland to release acetylcholine.
 - d. The sympathetic division stimulates the adrenal gland to release epinephrine.
8. The parasympathetic division
- a. is also called the craniosacral division.
 - b. has preganglionic axons in cranial nerves.
 - c. has preganglionic axons in pelvic splanchnic nerves.
 - d. has ganglia near or in the wall of effector organs.
 - e. all of the above.
9. Which of these is *not* a part of the enteric nervous system?
- a. ANS motor neurons
 - b. neurons located only in the digestive tract
 - c. sensory neurons
 - d. somatic motor neurons
10. Concerning the distribution of autonomic axons,
- a. autonomic nerve plexuses contain sympathetic and parasympathetic axons.
 - b. sympathetic axons reach the skin and skeletal muscles of most of the body through spinal nerves.
 - c. sympathetic axons reach thoracic organs and parts of the head and neck through sympathetic nerves.
 - d. sympathetic axons reach abdominopelvic organs through splanchnic nerves.
 - e. all of the above.