II. NEURO-EMBRYOLOGY

DEVELOPMENT OF NEURAL TUBE



- Neural Plate: thickening of embryonic ectoderm, Day 18.
- Notochord: Just ventral to the neural plate. It induces formation of the Neural Tube.
- Neural Folds are formed on the Neural Plate. Next, they begin to move toward each other, forming a Neural Groove.
- NEURAL TUBE: Is formed from the primitive Neural Groove. *This occurs first in the midsection of the embryo and then proceeds rostrally and caudally.*
- o Rostral Neuropore and Caudal Neuropore are the open ends of the Neural Tube.
- NEURAL TUBE DEFECTS:
- SPINA BIFIDA: Failed closure of the caudal part of the Neural Tube. But, the nervous system continues to develop normally.
- Meningoceles (outpocketings of meninges) and Meningomyeloceles (outpocketings of meninges + nervous tissue) will result.
- o ANENCEPHALY: Failure of rostral closure of neural tube and subsequent differentiation.

NEURAL CREST CELLS: Lie on either side of the Neural Groove and are pinched off by closure of the Neural Tube. They form a number of important structures.

- Dorsal Root Ganglia and portions of sensory ganglia that are like the Dorsal Root (V, VII, VIII, IX, X)
- Sympathetic Ganglia
- Parasympathetic (Enteric) Ganglia
- Pia and Arachnoid Mater
- Schwann Cells
- Melanocytes Proliferation in Neural Tube:
- Cells start connected both to the internal and external limiting membranes of the neural tube, but ultimately remain connected only to the internal limiting membrane.
- **NEURAL BIRTHDAY:** Occurs when a cell-line has had its last division and remains in the same structure terminally thereafter. Neurons from the same structure tend to have the same Neural Birthdays.
- THREE LAYERS of Proliferating Tube:



- Ventricular Layer: Contains dividing cells.
- Mantle Layer: Postmitotic neuronal cells bodies (after their birthday)
- Marginal Layer: Axoplasmic extensions of the mantle layer.

• NEURAL VESICLES: At 3 weeks, three distinct outpocketings can be made out. These are the classical three vesicles out of which entire nervous system grows:



- **Rhombencephalon** (Hindbrain)
- o Mesencephalon (Midbrain) -----> Midbrain
- **Prosencephalon** (Forebrain) -----> Diencephalon + Telencephalon
 - Thalamus
 - Epithalamus
 - Hypothalamus
 - Subthalamus
- VENTRICLES will arise from the Central Canal of the Neural Tube.
- FLEXURES: Characteristic flexures create the shape of the CNS
- o 26 Days: Mesencephalic and Cervical Flexures.
- \circ 35-50 Days: Pontine Flexure brings the Cerebellum to lie dorsal to the Pons.

GROSS BRAIN DEVELOPMENT





- o (2) Diencephalon
- Mesencephalon (Midbrain) -----> (3) Midbrain
- Rhombencephalon (Hindbrain)
 - Metencephalon -----> (4) Pons, Cerebellum
 - Myelencephalon -----> (5) Medulla Oblongata

SPINAL CORD DEVELOPMENT

- Sulcus Limitans: It appears along the Neural Tube, and separates dorsal and ventral regions of the spinal cord.
- ALAR (DORSAL) PLATE: Neurons become specialized for sensory.
- BASAL (VENTRAL) PLATE: Neurons become specialized for motor.

BRAINSTEM / CEREBELLUM DEVELOPMENT

- MEDULLA
 - o CN NUCLEI are arranged in Columns in the medulla.
 - CLOSED MEDULLA:
 - OPEN MEDULLA: *The Alar Plate is displaced laterally*. So, sensory stuff is now lateral to motor stuff, which tends to be more medial.
- PONS: It maintains the alar / basal plate distinction between sensory / motor.
- CEREBELLUM: Formed from the **Rhombic Lips** of the Alar Plate of the Pons.
- o These lips fold medially to cover the Pons, so that Pons is ventral to Cerebellum.
- o There are two proliferative zones present during development:

TELENCEPHALON: The neurons develop in an "inside-out" fashion. The earliest neuronal birthdays occur closest to the medullary center, then neurons migrate beyond that.

CELLULAR EVENTS IN DEVELOPMENT

- Making Neuronal Connections:
 - Sometimes a neuron will reel out its axon as it grows.
- At other times, a neuron will use physical or chemical (chemotaxis) cues to grow toward a target.
- Synaptic Plasticity: Modifications to neuronal connections made after development is complete.
- They can be made as an alternative pathway following damage to a connection.
- They can be made in the process of "learning."
- Programmed Cell Death: More neurons than are needed are made during development.
- Neurons that are unsuccessful at making their connections are then lost (killed, DEAD) by a preprogrammed neuronal cell death.