INTRODUCTION

Definitions

- Organic Disorders of Communication
  - Any kind of anomaly in the physical structures responsible for speech production and/or language processing
    - From problems in
      - Respiration
      - Phonation
      - Articulation
      - Hearing
      - Neurology
- Neurogenic Disorders of Communication
  - Impairments stemming from damage to the central or peripheral nervous system
- Functional Disorders of Communication
  - No physical bases for the communication problem
- May Coexist!

Prevalence of Organic and Neurogenic Disorders

- 42 millions in the US affected by a communication disorder
  - 28 million hearing loss
  - 14 million speech, voice, or language disorder
- 26 million/year – Neurologic
  - Stroke: 1.5 million
  - Dementia:
    - 10% of over 65yrs
    - 80% of over 85yrs
  - Dysphagia:
    - 15 million Americans have dysphagia
  - Trauma:
    - 2 million/year have head trauma
      - 100,000 endure permanent communication disability
- Numbers likely to increase, why?
  - Aging population that is living longer
  - Medically fragile infants are now surviving

Impact of Organic & Neurogenic Disorders of Communication

- Quality of life
  - Individual level
  - Society

- Can be alleviated through communicative, educational, and medical services
  - Costly? Yes!!
    - Lack of these services is costlier in the long run!!

Technological and Theoretical Advances

- Advancements of the knowledge in the physiological bases of communication
- Instrumentation to analyze and quantify
  - **Physiological aspects of communication**
    - Videofluroscopy
      - Dysphagia
      - VPI
    - Stroboscopic endoscopy
      - Voice
    - Brain imaging techniques
      - Positron Emission Tomography (PET)
      - Magnetic Resonance Imaging (MRI)
  - **Acoustic aspects of communication**
    - Computerized Speech Lab (CSL)
    - Time-Frequency analysis (TF32)
    - Etc.
  - **Analytic Discourse Techniques**

Scope of Clinical Practice

- SLPs working in
  - Hospitals
  - Rehabilitation centers
    - Long-term rehab
      - To maximize individual's ability to communicate independently
      - Individual's readjustment to society (community reentering programs)
  - Acute care facilities
    - Assessment
    - Short-term treatment
    - Counseling patient and family regarding long-term rehabilitation
  - Neonatal nurseries
  - Early intervention programs
  - Schools
    - Mainstreaming of children
      - Medically fragile
      - At risk for developmental delay
      - Autism
      - Multiple handicaps
      - Hearing loss
      - Etc.
    - Preschools
      - Early intervention
        - » Feeding therapy
        - » Intensive language intervention
  - Home care
  - Private practice

Developmental Requirements

- Anything that interferes with the child's ability to interact with the environment in a normal manner → Developmental delay factor
  - Ability
    - Anything that interferes with child's ability

  Might be biological, environmental, or both
    - These factors can be used for early identification of children with potential risk for developmental delay
      - The earlier the identification and intervention, the better the outcome of the service → more efficacious

Hypoxic-Ischemic Encephalopathy

- **the most prominent cause of neonatal mortality and long term neurologic morbidity often referred to as cerebral palsy.**
- **It is noted in approximately 1 to 2 per 1000 deliveries.**

Pathogenesis

Impaired cerebral blood flow (CBF) is the principal pathogenetic mechanism underlying most of the neuropathology attributed to perinatal brain injury.

It is most likely to occur as a consequence of interruption of placental blood flow and gas exchange; a state that is referred to as asphyxia.

Definitions

- Hypoxia - refers to an abnormal reduction in oxygen delivery to the tissue
- Ischemia - refers to a reduction in blood flow to the tissue
- Asphyxia - refers to progressive hypoxia, hypercarbia and acidosis.

- Acidemia a cord pH $\leq$ 7.00. Might be fatal

Characteristics of Hypoxic-Ischemic Brain Damage

- Hypoxic-ischemic brain injury is an evolving process that begins during the insult and extends into a recovery period - reperfusion period
- Tissue injury takes the form of :

  Necrosis - characterized by tissue swelling and membrane disruption

  Apoptosis occurs with less severe cases. Called “programmed cell death” characterized by cellular and nuclear shrinkage, chromatin condensationand DNA fragmentation.

Early Identification of High Risk Infants

1) Evidence of an Acute Perinatal Insult

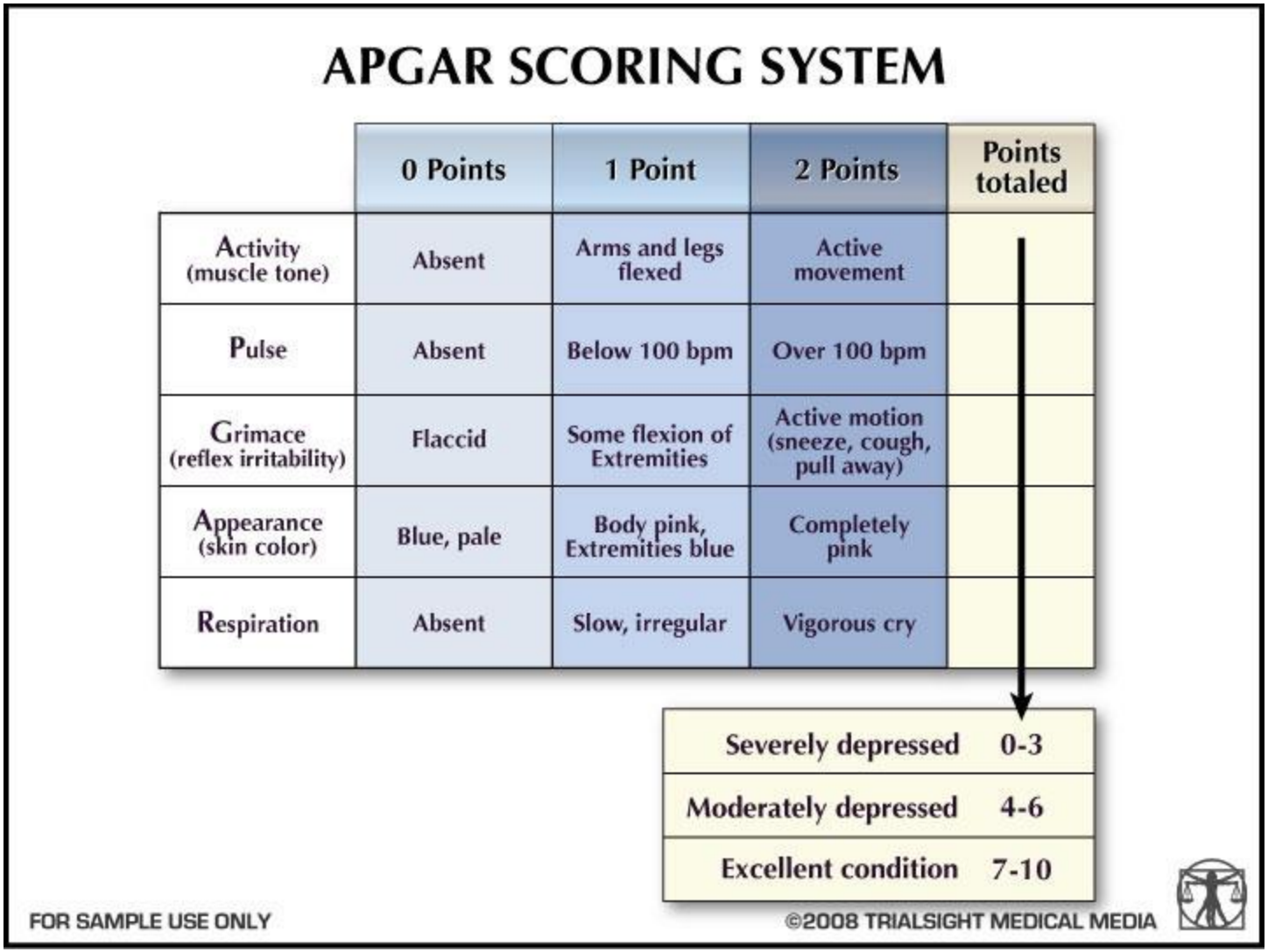
 * Indicated by a combination of markers

- Delivery room resuscitation
- 5 Minute Apgar score ≤ 5
- Cord arterial pH ≤ 7.00

2) Postnatal evidence of encephalopathy

- Clinical
- EEG

APGAR score


APGAR SCORING SYSTEM
0 Points
1 Point
2 Points
Points totaled
Activity (muscle tone)
Absent
Arms and legs flexed
Active movement
Pulse
Absent
Below 100 bpm
Over 100 bpm
Grimace (reflex irritability)
Flaccid
Some flexion of Extremities
Active motion (sneeze, cough, pull away)
Appearance (skin color)
Blue, pale
Body pink, Extremities blue
Completely pink
Respiration
Absent
Slow, irregular
Vigorous cry
Severely depressed 0-3
Moderately depressed 4-6
Excellent condition 7-10
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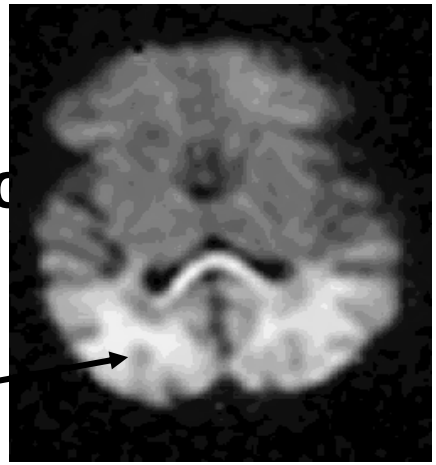
General Measures beyond delivery room

- Ventilation
- Fluid Status
- Oliguria
- Hypotension
- Glucose status
- Seizures
- Cerebral edema

Role of Glucose

- **Both hyper and hypoglycemia may be seen in the post resuscitative phase.**
- **Both may exacerbate neuronal injury**
- **Hyperglycemia may contribute to ↑ levels o lactate and thus to continuing acidosis**
- **Hypoglycemia may contribute to injury particularly in parieto-occipito cortex**
- **The goal should be to maintain glucose levels in the normal range**



Infant Mortality

- Major risk factors:
  - Gender
    - Males higher infant mortality than females
  - Gestational weight
    - Infant mortality increases with decreased BW
  - Live Birth Order
    - Second born have lower infant mortality than others
  - Maternal Age
    - Decreases with increasing maternal age up to 30-34yrs
    - Increases after maternal age of 35 yrs
  - Maternal Education
    - Declines with increased maternal education
  - Prenatal Care
    - With prenatal care starting in first trimester, infant mortality decreases
      - Especially for BW of 1500gms and more

Low Birth Weight (LBW) and Prematurity

- Premature infant:
  - A child born at or before the 36th week of gestation
    - One month early
  - <5% of all live births
    - <1% of whom have BW <1,500gms
  - Mam's status
    - 20% are born to adolescent mams
    - Highest proportion among women of low socioeconomic status
  - SO?
    - Higher incidence of developmental disability
    - Majority – NO handicap
    - 1/3 developing according to age expectations
    - Factors affecting this:
      - Gestational age
      - Status of neonatal care

Respiratory Distress Syndrome (RDS)

- Main cause of death among premature/LBW infants

- RDS common in premature infants
  - Most common illness in NICU, one of most common causes of death
  - Affects 20% of premature infants in the first few days of life
  - Cause: deficiency of a material called surfactant, needed for infants lungs to prevent air sacs collapse
    - Adequate surfactant amount produced by 36wks GA → presence of RDS depends on child's degree of prematurity
  - Manifestations:
    - Grunting on expiration
    - Frequent breathing (panting) with increased effort
    - Flaring of nostrils
    - Retraction of muscles between ribs and below the rib cage
  - Medical Management:
    - Increasing concentration of inspired oxygen
    - Continuous Positive Air Pressure (CPAP)
    - Intubation and ventilation
    - Surfactant Replacement – Synthetic
      - » 65% survival rate compared to 26% if not treated
    - Cant nurse or drink – IV or NGT
    - 90% survive – mostly with no long-term deficit
      - » Small group demonstrate developmental delay

Bronchopulmonary Dysplasia (BPD)

- AKA: chronic lung disease of the premature infant
- Characteristics
  - Chronic lung changes
    - Decreased lung capacity → increased risk for respiratory disease
- Occurs in
  - Infants with hx of RDS and/or prolonged mechanical ventilation and high concentration of oxygen
  - Full-terms with meconium aspiration, pneumonia, or other causes of respiratory distress
  - The greater the prematurity the greater the likelihood of BPD
    - 80% of BW<1000 gm have BPD
    - 10% of BW>1500gms
- Mortality in 20% of cases in the first year of life
- Causes:
  - Initial injury to lungs secondary to prematurity
  - Standard treatment to this injury
- Lasts months – years with mild-severe long-term complications
  - Infections
  - Kidney stones
  - Fragile bones
  - Tracheal abnormalities
  - Feeding problems
  - Poor growth
  - High blood pressure
  - Psychosocial problems

Intraventricular Hemorrhage (IVH)

- Bleeding into the fluid-filled spaces of the brain
  - A risk associated with respiratory disease
- Incidence: 35-45% for infants < 35 weeks of gestation
- Outcome: depends on the extent of the IVH
  - Mild IVH → outcome generally good
  - Severe IVH →
    - MR
    - Spastic quadriplegia
    - Possible hydrocephalus
- Intervention:
  - Most effective: prevention
- Detection: Ultrasound
  - Max severity at 48 hours

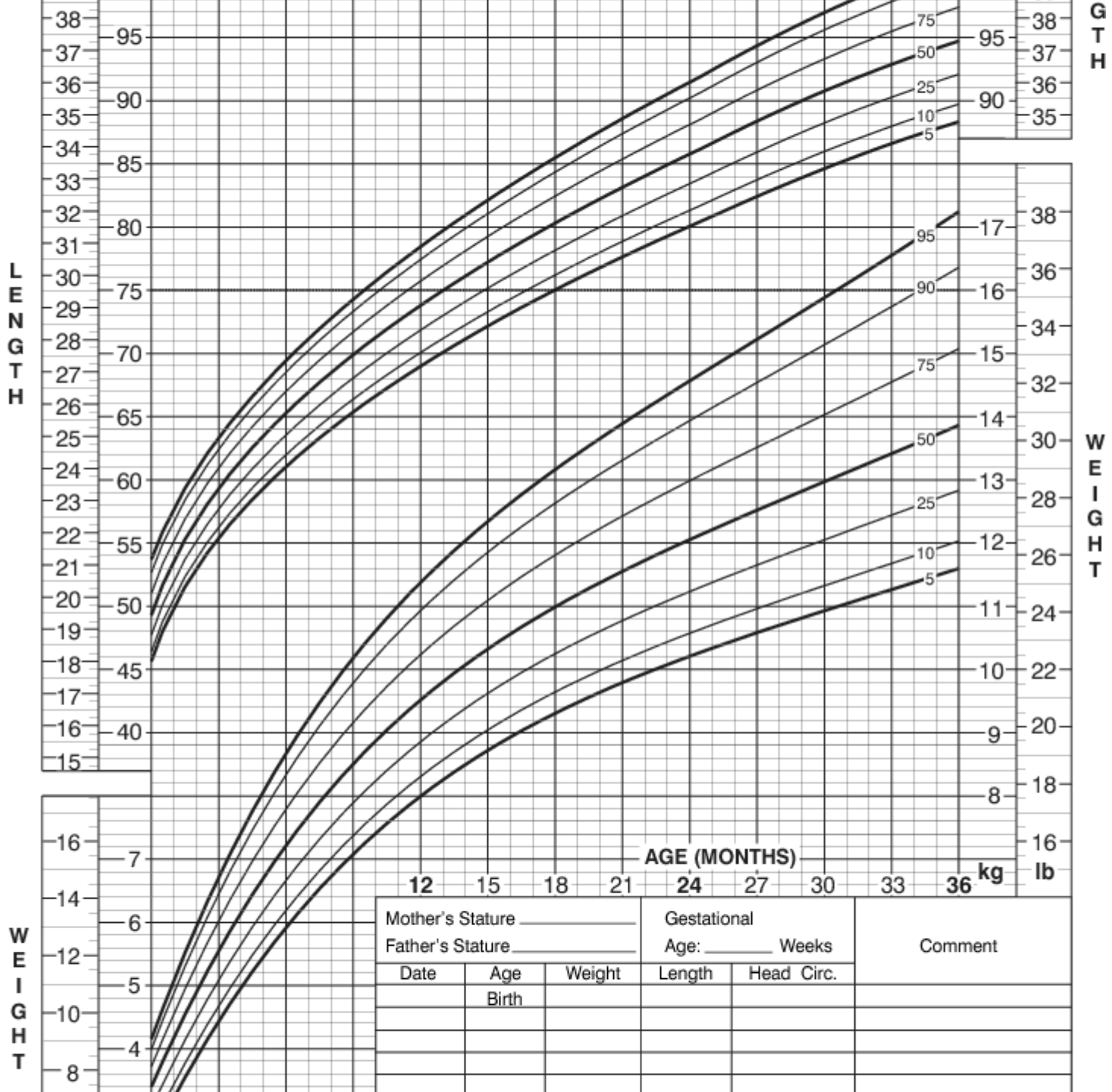
Necrotizing Enterocolitis (NEC)

- A serious intestinal disorder
- Causes
  - Injury to intestinal wall
  - Bacteria
  - Early formula feedings when infant's gut is immature
- Incidence: 10% of infants with BW < 1,500 gms
- Course:
  - 50% of infants with NEC require surgery to remove diseased bowel sections
    - 10% would have short gut syndrome (remaining bowel insufficient, diarrhea, poor nutrition)
  - Overall failure to thrive
  - Medically fragile
  - Affected developmental status

Growth and development

Introduction

- Children during their first year grow more quickly than any other age group
- Some kids follow the standard growth rate, however other children fail to do so
- This condition is know as “failure to thrive”


LENGTH
WEIGHT
AGE (MONTHS)
12 15 18 21 24 27 30 33 36
kg
lb
95
90
75
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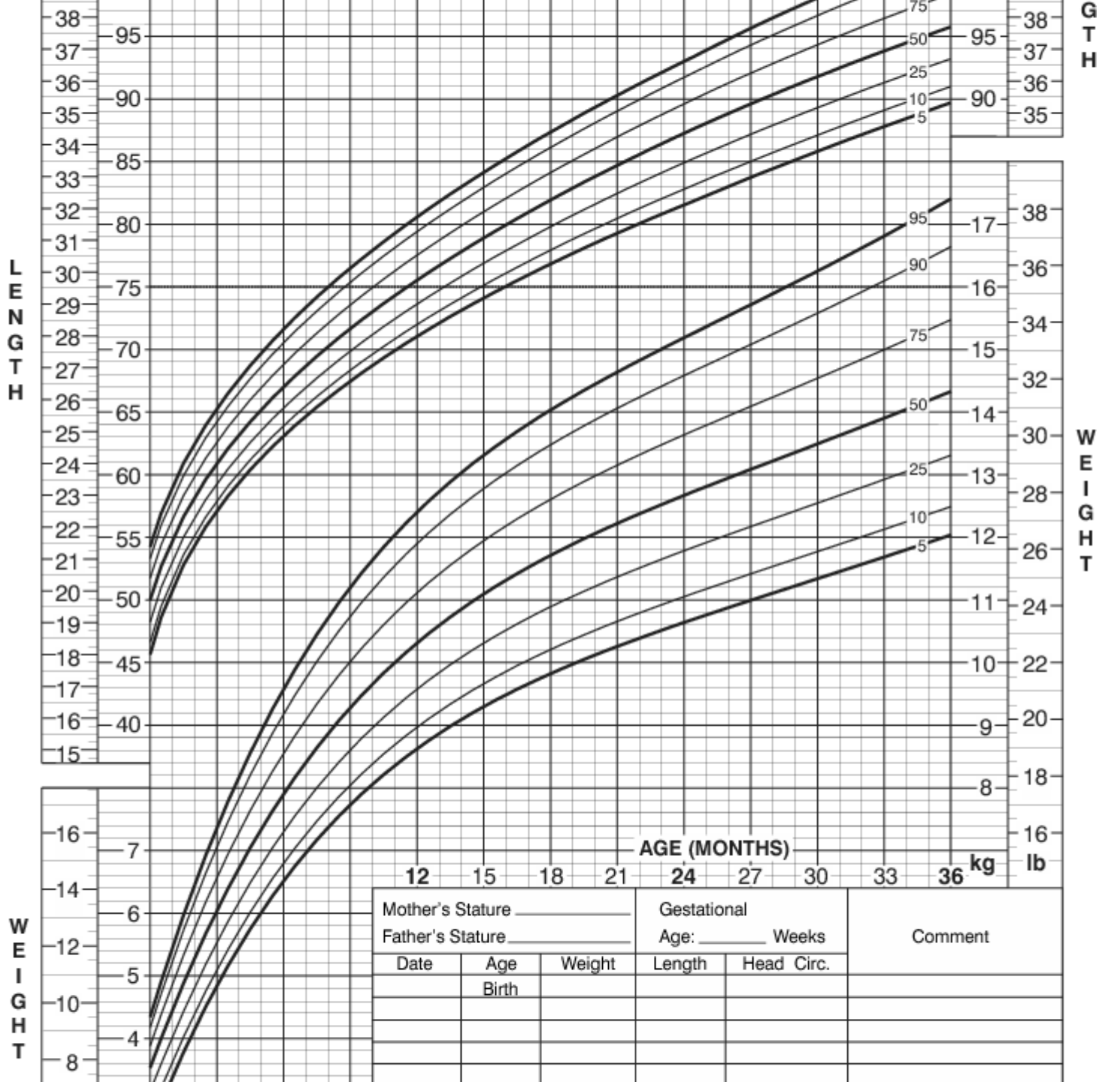
25
10
5
Mother's Stature
Father's Stature
Gestational
Age: Weeks
Comment
Date
Age
Weight
Length
Head Circ.
Birth


LENGTH
WEIGHT
AGE (MONTHS)
12 15 18 21 24 27 30 33 36
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5
Mother's Stature
Father's Stature
Gestational
Age: Weeks
Comment
Date
Age
Weight
Length
Head Circ.
Birth

Consequences

- Children’s brains’ maximum growth occurs during the first year of life
- Failure in nutrition may lead to irreversible brain damage
- Results may include negative effects on mental development

Consequences

- Undernourished kids may also:
- Become disinterested in their environment
- Avoid eye contact
- Become irritable
- Fail to reach developmental milestones (sitting, walking, and so on)

Causes

- Social factors:
- Mothers with post partum depression

- Child neglect

- SES (poverty)

- Cultural and regional factors

Causes

- Medical factors:
- GERD (gastroesophageal reflux disease)
- Chronic diarrhea
- Cystic fibrosis
- Chronic liver disease
- Celiac disease

Causes

- In GERD, kids are irritated by feeding so they no more interested in feeding
- Severe diarrhea prevent the body to get a benefit of nutritious materials
- Liver disease, cystic fibrosis, and celiac disease limit body’s ability to absorb nutrients

Causes

- Other medical reasons:
- Cleft lip or palate
- Cardiac disease
- Asthma and other respiratory problems
- Endocrinologic diseases

* These conditions may affect development by increasing the calorie need of the body

Causes

- Infections:
- Parasites
- Urinary tract infection
- Tuberculosis
- These infections increase the energy demands of the body, fasten nutrients usage, and decrease appetite
- These infections may cause short term or long term failure of thrive

Diagnosis

- Inability of the child to gain weight for 3 consecutive months during the first year of life
- Measures used:

- Development charts
- Head circumference
- Lab tests
- Calorie count

Genetics: Basis for Development and Disorders

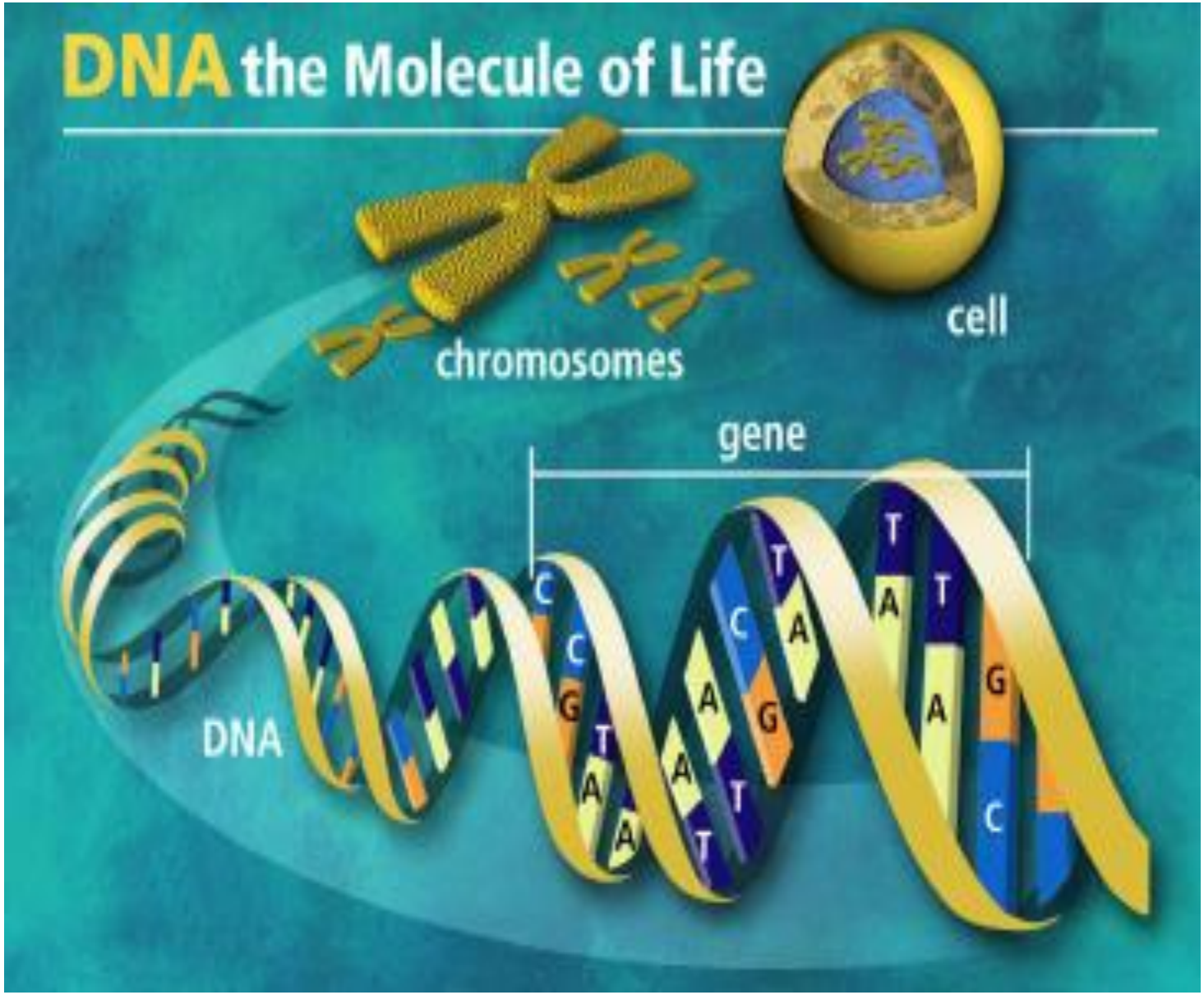
Definitions

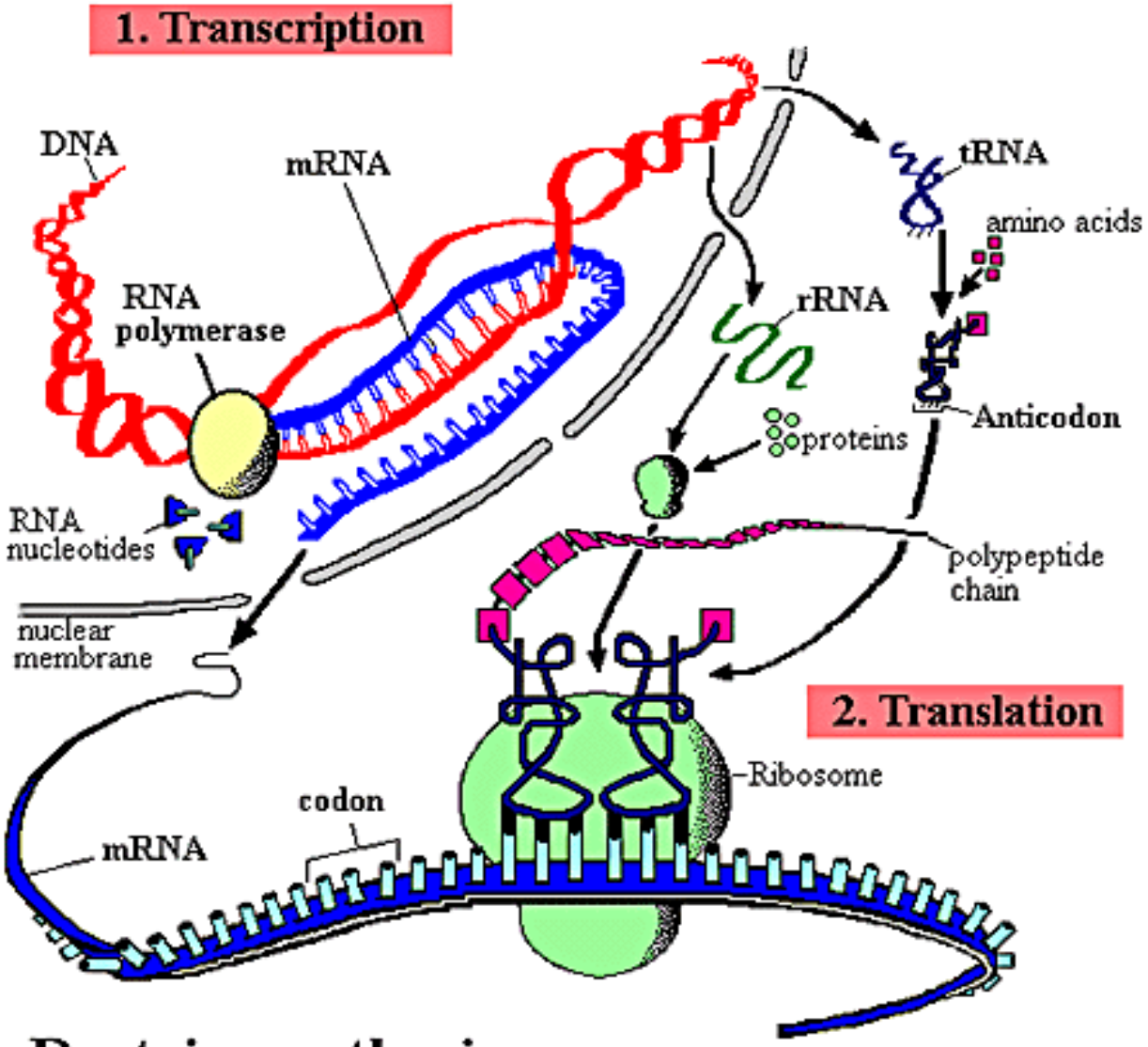
- Genetics: is a branch of biology that studies inheritance of genes.
- Genes are the units of heredity in living organisms. It is composed of DNA and RNA. Coding genes forms 3% of the DNA.
- DNA (Deoxyribonucleic acid) is a nucleic acid that contains the genetic instructions monitoring the biological development of all cellular forms of life (usually double helix)

- RNA Ribonucleic acid is a single-stranded. The sequences of most RNA molecules are translated to make proteins.
- Human genome is the sum of all genes.
- Genomics is the study of the entire set of DNA sequences, both coding and non-coding DNA.
- Chromosome is a large macromolecule into which DNA is normally packaged in a cell.

Importance of genes in life

- DNA is the coding system humans have to reproduce and develop through the life circle.
- Most cells of the human body contain 23 pairs of chromosomes, half of which are inherited from each parent.
- the sperm cells in males and the ovum in females, have 23 *individual* chromosomes,


DNA the Molecule of Life
chromosomes
cell
gene
DNA



Protein synthesis

Inheritance patterns

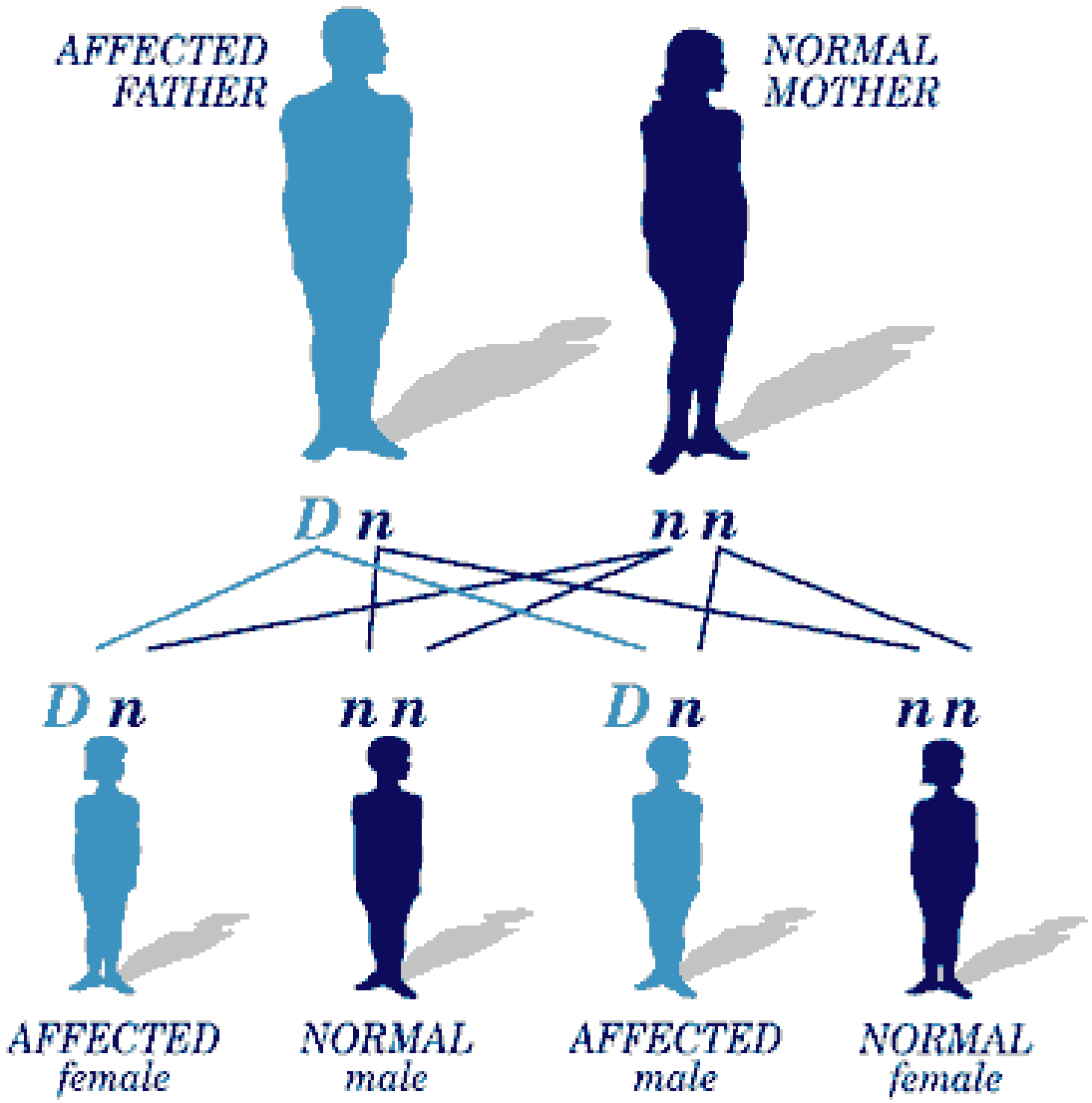
- Mendelian:

_ discovered by Gregor Monk

_ traits are passed from each parent to each child:

_ recessive: both parents carry affected gene

_ dominant (only one parent passes the gene)


AFFECTED
FATHER
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D n
n n
AFFECTED
female
NORMAL
male
AFFECTED
male
NORMAL
female

Non Mendelian inheritance

*uniparental

*Maternal and parental imprinting

*Anticipation

*Mitochondrial inheritance.

Multifactorial inheritance

- Complex inheritance pattern. Most communication disorders have this type
- Genotype: includes the genes and genomes
- Phenotypes: physical and behavioral characteristics resulted from the interaction between the genotype and the environment.

Down Syndrome

- Down syndrome is the most frequent genetic cause of mild to moderate mental retardation
- Chromosomal disorder caused by an error in cell division that results in the presence of an additional third chromosome 21 or "trisomy 21.“
- It occurs in one out of 800 live.

- It is not attributable to any behavioral activity of the parents or environmental factors.

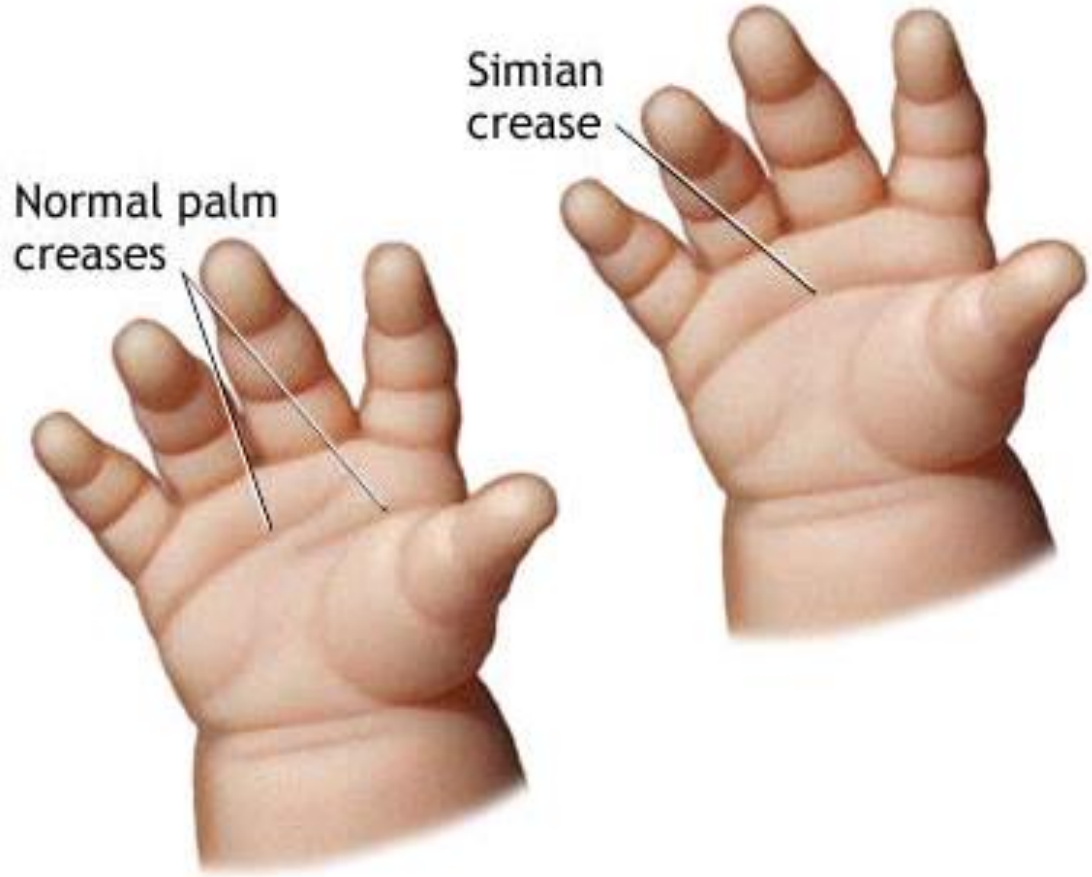
- The probability that another child with Down syndrome will be born in a subsequent pregnancy is about 1 percent, regardless of maternal age.

Down syndrome and maternal age

- The older the mother gets, the higher the probability that she will have a child with down syndrome.

Some physical aspects

Flat facial profile, an upward slant to the eye, a short neck, abnormally shaped ears, and a single, deep transverse crease on the palm of the hand.


Simian
crease
Normal palm
creases

Down Syndrome and Associated Medical Disorders

*Hearing loss

_ More than 75% of young children with Down syndrome are found to have at least a mild hearing problem at sometime in childhood.

_ The hearing problems are fluctuating.

*Congenital heart diseases.

_more than 50% of the children.

Newborns

- Hypotonia and tongue protruding makes it take longer to feed the child with Down syndrome.

Infants and Preschool Children

- Show developmental delay linguistically and in motor terms.
- Meet the development milestones but with delay.
- Development could not be complete at all.

Adolescence

- Hormonal changes occur.

- Males have reduces reproduction abilities while females retain there abilities.

verbal language characteristics

- Slower development of language relative to other cognitive skills.
- language comprehension is better than language production.
- The vocabulary is better than the grammar of the language.
- Continued progress in speech, language and communication should be expected beyond early childhood.

Factors affecting progress

- hearing status.
- speech-motor function status.
- nonverbal cognitive skills and every day experience.

Fragile X Syndrome

- Called so because of a fragile site on the X-chromosome.
- Multi factorial transmitted syndrome
- It is one of the most commonly inherited form of mental retardation.

FXS

- Often in X-linked disorders only females are carriers and only males are affected. However, in Fragile X both males and females can be carriers (and both males and females can be affected).
- This occurs because the changes in this gene go through different stages as it is passed down in a family

- These stages are commonly called a premutation and a full mutation. The differences in the stages are determined by the number of CGG repeats (repeats of a DNA pattern) and the degree of methylation (whether the gene is turned on or off).

- An FMR1 gene that is methylated is turned off and does not make an adequate amount of an important brain protein called FMRP. Fragile X syndrome occurs due to the fact that FMRP is absent or significantly reduced in an individual with a full mutation.

Gene carriers

- Traditionally, a carrier of a genetic mutation was defined as an individual who inherited an altered form of a gene but had no effects or symptoms of that gene change or mutation
- There is a different story in FXS!

Gene carriers

- as carriers of a premutation have a risk to develop:

_ fragile X- associated tremor ataxia syndrome

_ fragile X-related premature ovarian failure

_ D.D., social and emotional disturbances

- the most common cause of *inherited* mental impairment.

- This impairment can range from learning disabilities to more severe cognitive or intellectual disabilities
- It may affect balance, tremor and memory in some older male gene carriers.

Initial signs

- Developmental delay, including speech
- short attention span or hyperactivity
- frequent temper tantrums.
- autistic-like behaviors such as rocking, talking to oneself, spinning, unusual hand movements.
- poor motor coordination.

physical features

2-3 years:

Long and/or wide and/or protruding ears.

prominent jaw or long face

flattened nasal bridge

prominent forehead.

Speech and Language Characteristics in Males

- Attention, memory, and auditory comprehension problems.
- Speech production may be characterized by imprecise articulation, a fast rate of speech, and cluttering, all of which affect intelligibility. Voice characteristics may include a loud volume, with unusual, high pitch, and harshness.

- Syntax, or grammar, is usually reported as in keeping with overall cognitive level.
- Pragmatics, or conversational skills, are a major area of concern for boys with fragile X syndrome. Poor eye contact, perseverative speech, poor topic maintenance, and self-talk often characterize the conversational attempts of boys with fragile X.

Speech and Language Characteristics in Females

- Generally, much better than boys. Almost in normal ranges.
- Most difficulties are reported in the area of pragmatics. Because of anxiety and shyness.

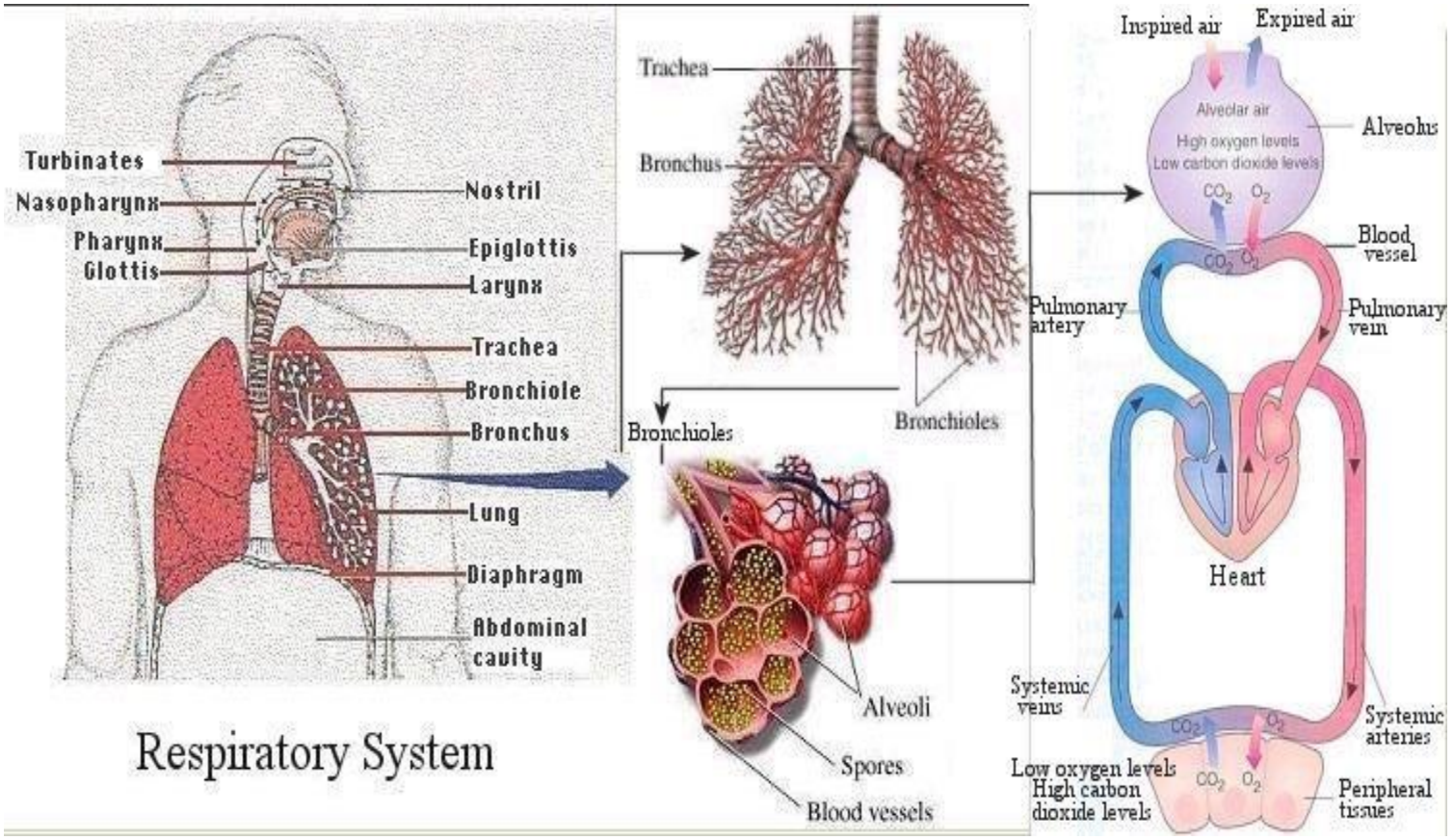
Respiratory system diseases

Normal respiration

- Physiology of respiration
- Neural support
- Muscles involved in respiration
- Abnormal respiration drive

Normal respiration

- Ventilation: Moving air inside and outside alveoli
- Pulmonary gas exchange: Change gases between alveoli and pulmonary capillaries
- Gas transport: Moving gases through pulmonary capillaries to circulation to peripheral capillaries in organs
- Peripheral gas exchange: gas change between tissue capillaries and the tissue or organ



Respiratory System

Pneumonia

Symptoms could include:

- Fever
- Cough
- Chest pain
- Shortness of breath
- Productive cough
- Pain

Symptoms could include:

- Fatigue
- Muscle pain
- Loss of appetite
- Wheezing

Pneumonia

Causes

- Streptococcus pneumoniae
- Respiratory syncytial virus
- Mycoplasma pneumoniae
- Herpes Simplex virus
- And many others

Pneumonia

Risk factors

- Elderly
- Alcohol abuse
- Young children
- Smoking
- Hospitalization!
- Improper immune system

Pneumonia

- Complications

Bronchiectasis

Acute Respiratory Distress Syndrome (ARDS)

Lung abscess

Empyema of lung

Respiratory failure

Bronchiectasis

- destruction and widening of the large airways
- Could be congenital or acquired
- Symptoms often develop gradually, and may occur months or years after the event that causes the bronchiectasis.

Bronchiectasis

- Signs

Bluish color

Breath odor

Cough that gets worse when lying on one side

Fatigue

Shortness of breath

Weight loss

Acute respiratory distress syndrome

- is a life-threatening lung condition that prevents enough oxygen from getting into the blood

  ARDS can be caused by any major swelling (inflammation) or injury to the lung.

- Some common causes include:
- Breathing vomit into the lungs (aspiration)
- Inhaling chemicals
- Pneumonia
- Septic shock
- Trauma

- ARDS leads to a buildup of fluid in the air sacs. This fluid prevents enough oxygen from passing into the bloodstream.
- The fluid buildup also makes the lungs heavy and stiff, and decreases the lungs' ability to expand. The level of oxygen in the blood can stay dangerously low, even if the person receives oxygen from a breathing machine through a breathing tube (endotracheal tube).

- ARDS often occurs along with the failure of other organ systems, such as the liver or the kidneys. Cigarette smoking and heavy alcohol use may be risk factors.

ARDS

- Symptoms

- Labored, rapid breathing
- Low blood pressure and organ failure
- Shortness of breath
- Symptoms usually develop within 24 to 48 hours of the original injury or illness. Often, people with ARDS are so sick they are unable to complain of symptoms.

Lung Abscess

- acute or chronic infection of the lung, marked by a localized collection of pus, inflammation, and destruction of tissue.

  Lung abscess is the end result of a number of different disease processes ranging from fungal and bacterial infection to cancer.

Lung Abscess

- It can affect anyone at any age.
- Patients who are most vulnerable include:

- Those weakened by cancer and other chronic diseases
- Patients with a history of substance abuse, diabetes, epilepsy, or poor dental hygiene.
- patients who have recently had operations under anesthesia and stroke patients.
- Children with weakened immune systems, malnutrition, or blunt injuries to the chest.

Lung Abscess

- The phagocytes release chemicals that contribute to inflammation and eventual necrosis or death of a part of the lung tissue
- Related problems:

- Aspiration
- Bronchial obstruction
- Spread of infection

Empyema

- a collection of pus in the space between the lung and the inside of the chest wall (pleural space).

  Empyema is caused by an infection that spreads from the lung. It leads to a buildup of pus in the pleural space.

Empyema

- Risk factors include:
- Bacterial pneumonia
- Lung abscess
- Thoracic surgery
- Chest injury or trauma
- Complications:
- Pleural thickening
- Reduced lung function

Empyema

- Expectations (prognosis):
- When empyema complicates pneumonia, the risk of permanent lung damage and death goes up.
- Patients will need long-term treatment with antibiotics and drainage. However, most people fully recover from empyema.

Respiratory Failure

- is nearly any condition that affects breathing function or the lungs themselves and can result in failure of the lungs to function properly
- Level of oxygen in the blood becomes dangerously low, and/or the level of $C0_2$ becomes dangerously high.
- This happens in two ways: Either gas exchange brake down or improper ventilation

hypoxemic respiratory failure

- The problem is in gas change
- Result in decreased amount of oxygen in blood
- All body organs and tissues are affected

hypoxemic respiratory failure

- Hypoxemia also may result from
- Spending time at high altitudes
- Various forms of lung disease that separate oxygen from blood in the lungs; severe anemia; and blood vessel disorders that shunt blood away from the lungs, thus precluding the lungs from picking up oxygen.

Ventilatory failure

- In this type, breathing is not strong enough to rid the body of $C0_2$.
- CO_2 builds up in the blood (hypercapnia).
- Ventilatory failure can result from

- Failure of the respiratory center in the brainstem fails to drive breathing
- Muscle disease
- COPD
- Cystic fibrosis

Treatment of muscle failure

- Nearly all patients are given oxygen as the first treatment.
- Then the underlying cause of respiratory failure must be treated. For example, antibiotics for lung infection
- A patient whose breathing remains very poor will require a ventilator to aid breathing.

Chronic obstructive pulmonary disease

- COPD) is one of the most common lung diseases. It makes it difficult to breathe.
- There are two main forms of COPD:

- Chronic bronchitis, defined by a long-term cough with mucus

- Emphysema, defined by destruction of the lungs over time

- Most people with COPD have a combination of both conditions.

Causes and risk factors

- Smoking is the leading cause of COPD. The more a person smokes, the more likely that person will develop COPD
- In rare cases, nonsmokers who lack a protein called alpha-1 antitrypsin can develop emphysema.

- Other risk factors for COPD are:

Exposure to certain gases or fumes in the workplace

Exposure to heavy amounts of secondhand smoke and pollution

Frequent use of cooking gas without proper ventilation

Symptoms

- Cough with mucus
- Shortness of breath that gets worse with mild activity
- Fatigue
- Frequent respiratory infections
- Wheezing

Complications

- Irregular heart beats (arrhythmias)
- Ventilator dependency
- Right-sided heart failure
- Cor pulmonale (heart swelling)
- Pneumonia
- Severe weight loss and malnutrition

Treatment

- There is no cure for COPD.
- Treatment aims to relieve symptoms and keep the disease from getting worse.
- Stop smoking is a must

- Medications used to treat COPD include:
- Inhalers (bronchodilators) to open the airways
- Inhaled steroids to reduce lung inflammation
- Intravenous steroids in severe cases
- Antibiotics to prevent infections
- Oxygen therapy

Pulmonary rehabilitation

Does not cure the lung disease, but it can teach patients to breathe in a different way so they can stay active and maintain strength

- Additional needed behaviors:

- Avoiding very cold air
- Reducing air pollution
- Eat a healthy diet with fish, poultry, or lean meat, as well as fruits

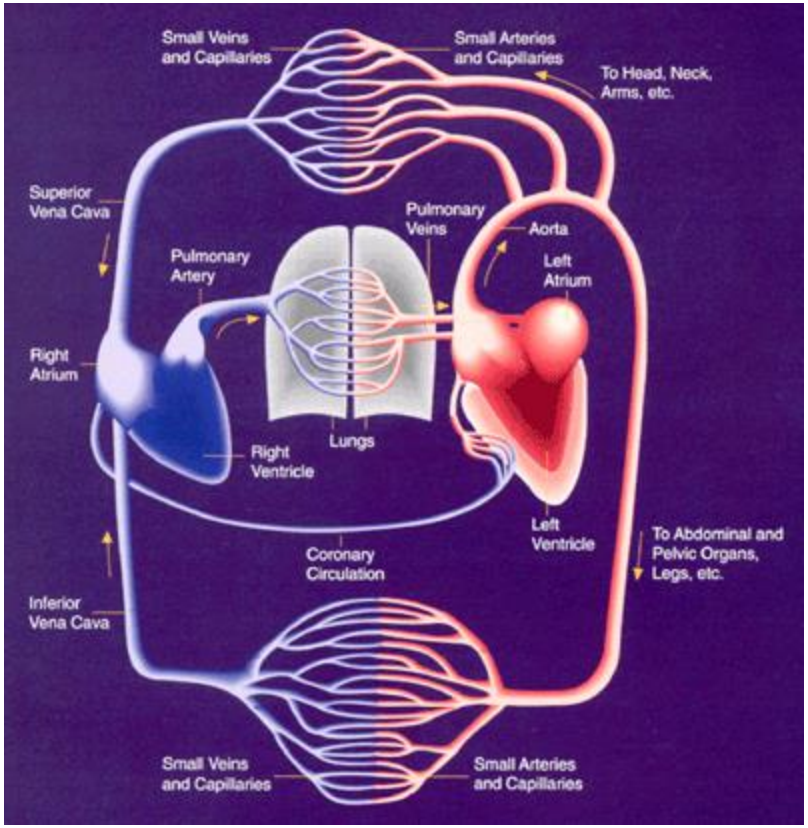
Surgical treatments may include:

- Surgery to remove parts of the diseased lung
- Lung transplant for severe cases

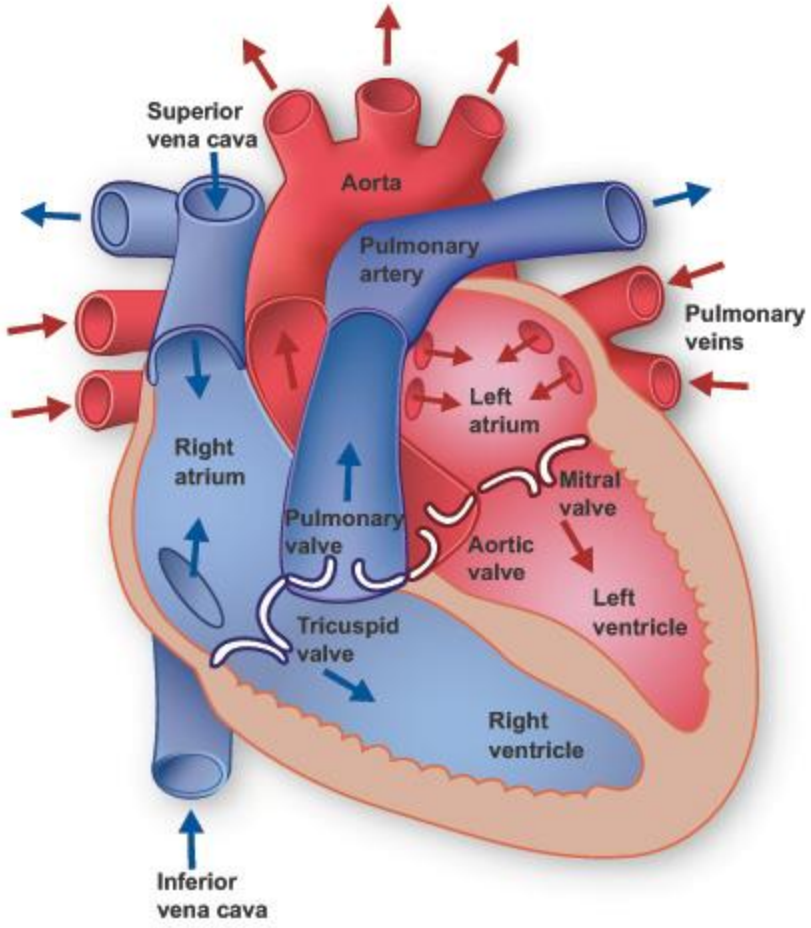
Cardiovascular system

Cardiovascular system

- Cardiovascular system’s function is to maintain blood flow in the body
- It Consists of heart and blood vesicles
- Arteries
- Veins
- Arterioles
- Venules
- Capillaries


Small Veins and Capillaries
Small Arteries and Capillaries
To Head, Neck, Arms, etc.
Superior Vena Cava
Pulmonary Veins
Aorta
Pulmonary Artery
Left Atrium
Right Atrium
Lungs
Right Ventricle
Left Ventricle
To Abdominal and Pelvic Organs, Legs, etc.
Coronary Circulation
Inferior Vena Cava
Small Veins and Capillaries
Small Arteries and Capillaries

The heart


Superior vena cava
Aorta
Pulmonary artery
Pulmonary veins
Left atrium
Right atrium
Mitral valve
Pulmonary valve
Aortic valve
Left ventricle
Tricuspid valve
Right ventricle
Inferior vena cava

The heart

- Weighs 200-450 gms
- Covered by double layered membrane called the pericardium
- The outer layer is attached to the vertebral column
- The inner layer is connected to the heart’s muscle
- The space between the two layers is filled with fluids

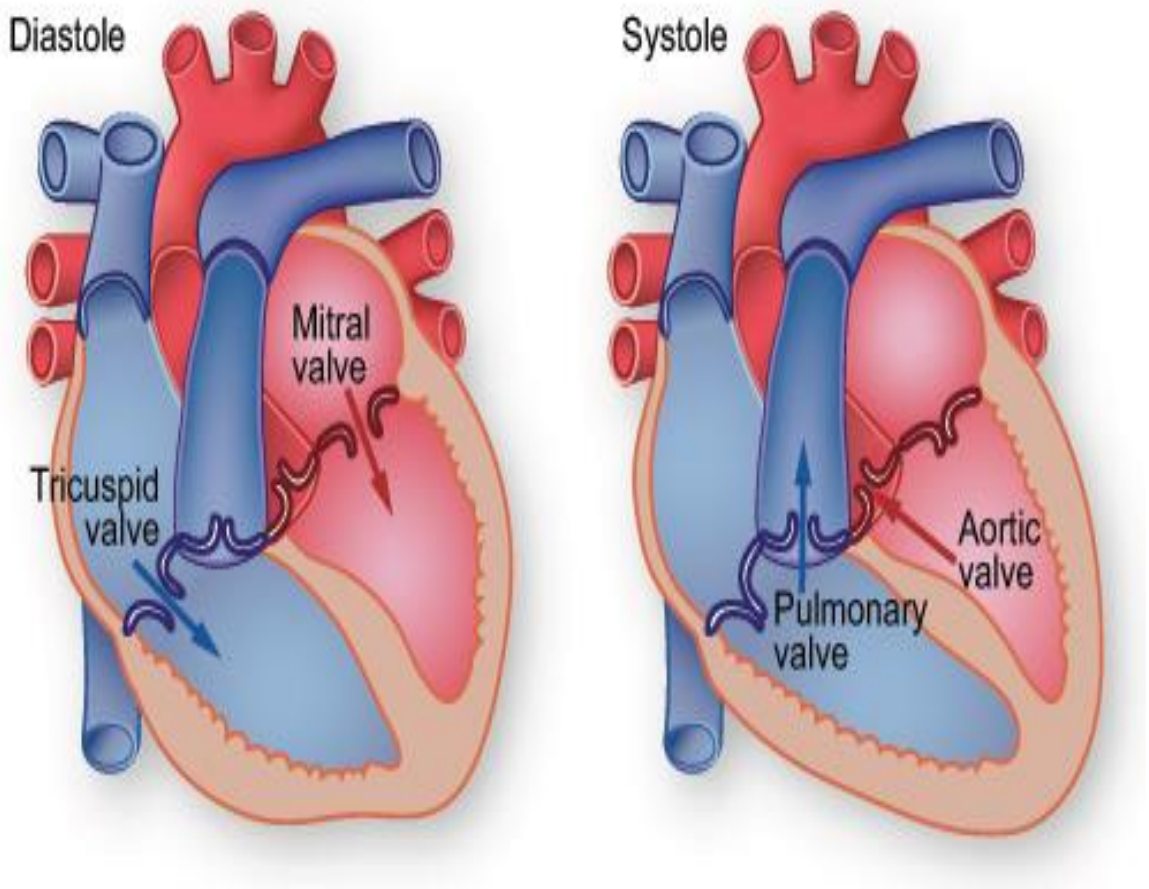
The heart

- Consist of four champers; two atria and two ventricles

- Left and right heart is separated by a wall of muscle called the septum

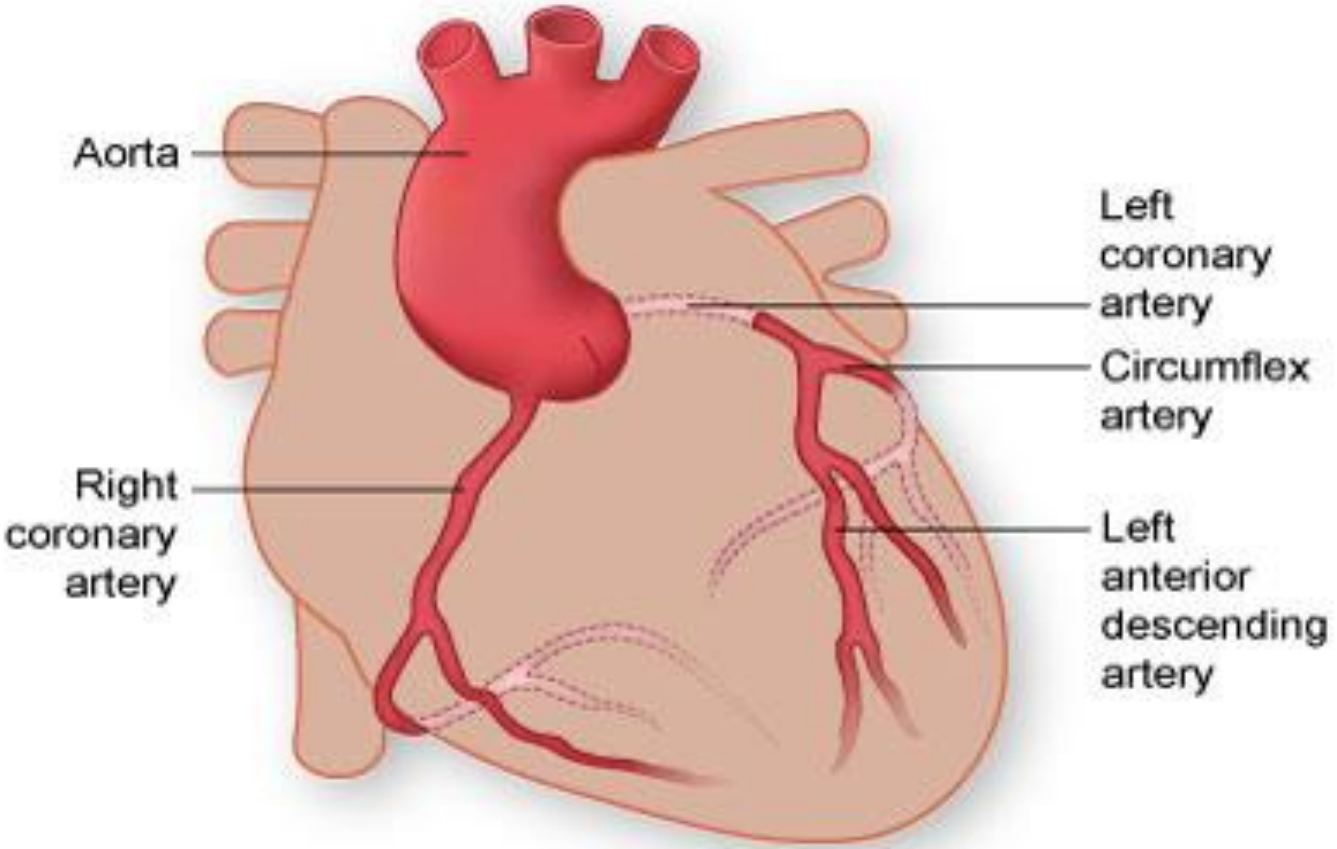
Valves of the heart

- Tricuspid: right atrium and right ventricle
- Pulmonary: right ventricle and pulmonary artery
- Mitral: left atrium and left ventricle
- Aortic: left ventricle into the Aorta


Diastole
Mitral valve
Tricuspid valve
Systole
Aortic valve
Pulmonary valve

Coronary circulation

- The arterial system that’s supplies the heart with blood
- Two coronary branches (right and left) stems from the Aorta
- Left coronary artery branches into descending artery and circumflex artery


Aorta
Left coronary artery
Circumflex artery
Right coronary artery
Left anterior descending artery

Heart beat

- Diastole:
- Upper part full of blood (atria)
- SA node makes an electrical signal cause contraction of atria
- Blood goes through tricuspid and mitral valves into resting ventricles

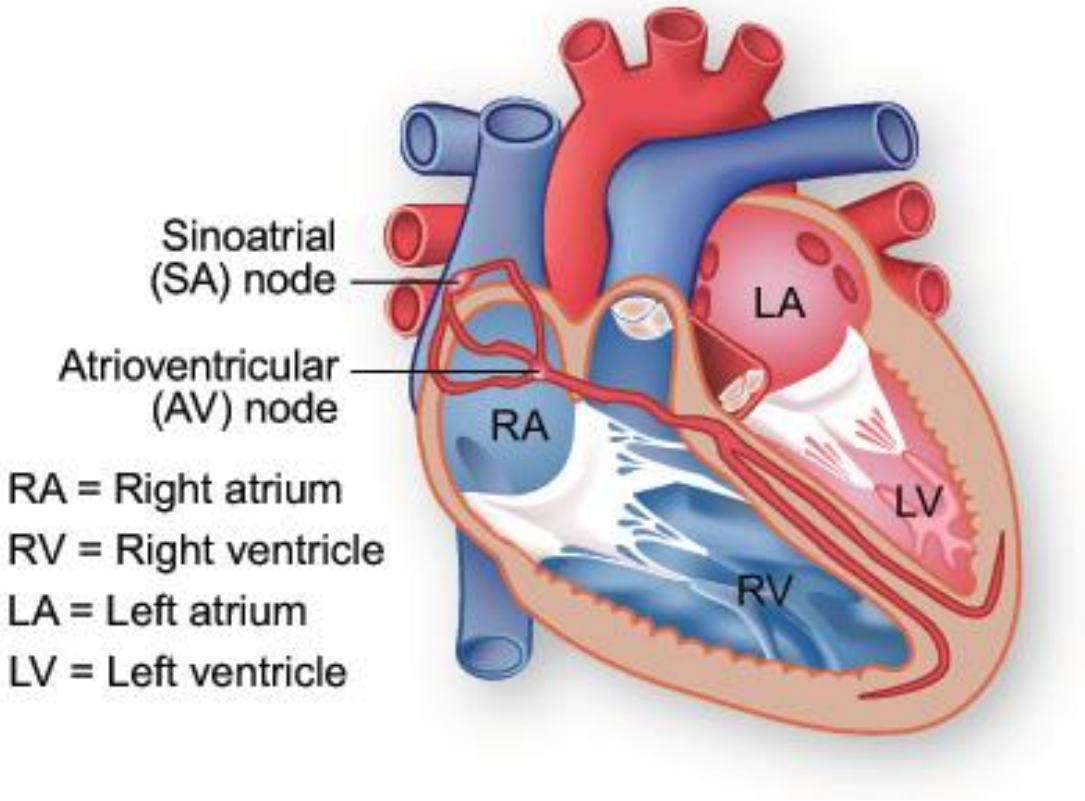
Heart beat

- Systole:
- Ventricles are full of blood
- Electrical signal (initiated by SA node) travels to the ventricles
- Ventricles contract pushing blood into pulmonary and Aortic valves
- Tricuspid and mitral valves are closed
- Then, ventricles relax and valves are close

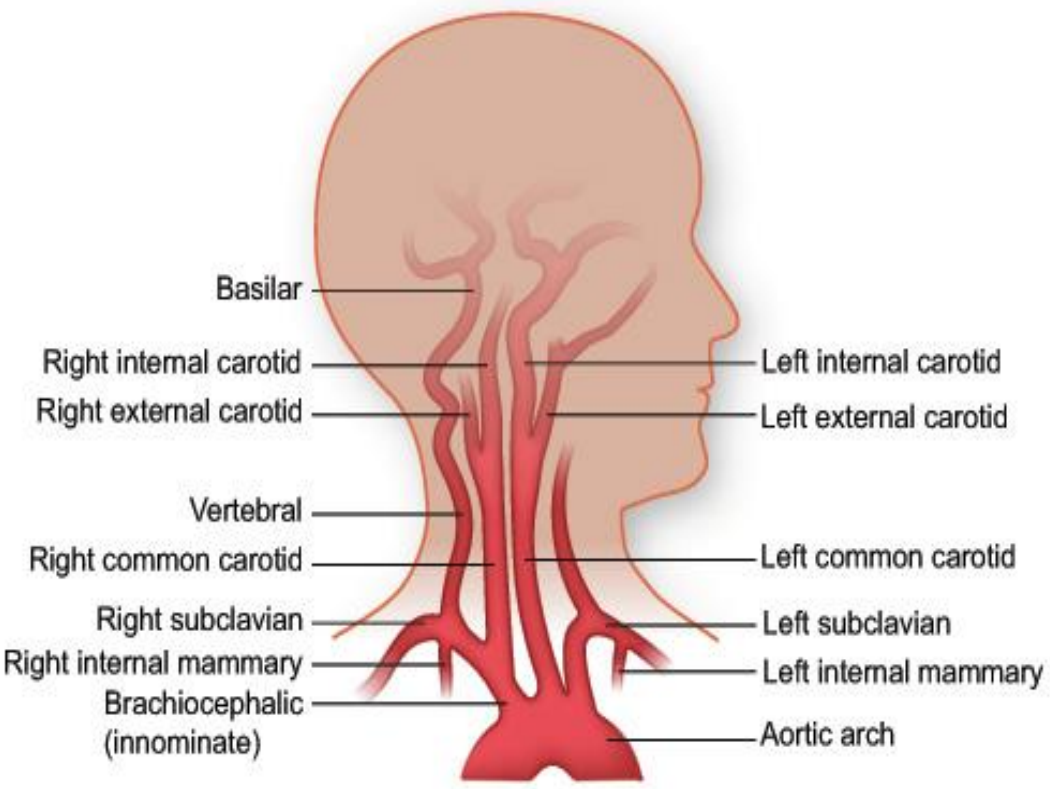
Conduction system

- The electrical signal begins in the SA node (Sinoatrial) on the top of the right atrium

- The signal travels into the AV node (Atrioventricular), which pass the signal into the tissues of ventricles


Sinoatrial
(SA) node
Atrioventricular
(AV) node
RA = Right atrium
RV = Right ventricle
LA = Left atrium
LV = Left ventricle
LA
RA
LV
RV

Arteries of head



Arteries

- Blood vesicles that transfer oxygenated blood to the body
- Outer walls contains smooth muscle fibers that contract and relax by the control of sympathetic system
- Have small lumen and include more muscle (than veins)
- Have no valves

Veins

- Carry blood back to heart
- Walls consist of three layers that are thinner and less elastic than arteries
- Transport blood with less pressure than arteries
- Major veins include valves that help carrying blood to the heart

Arterioles

- Transport blood from arteries to capillaries
- Dilate and constrict by the control of sympathetic nervous system
- Considered as the main regulators of blood flow and blood pressure

Venules

- Drain blood from capillaries into veins
- Small venules unit to form veins

Hypertension

Blood pressure

- Simplest definition: The pressure of blood against walls of arteries
- Determiners of blood pressure:

1. Cardiac output (heart beats and stroke size)
2. Resistance of the peripheral vesicles

Blood pressure

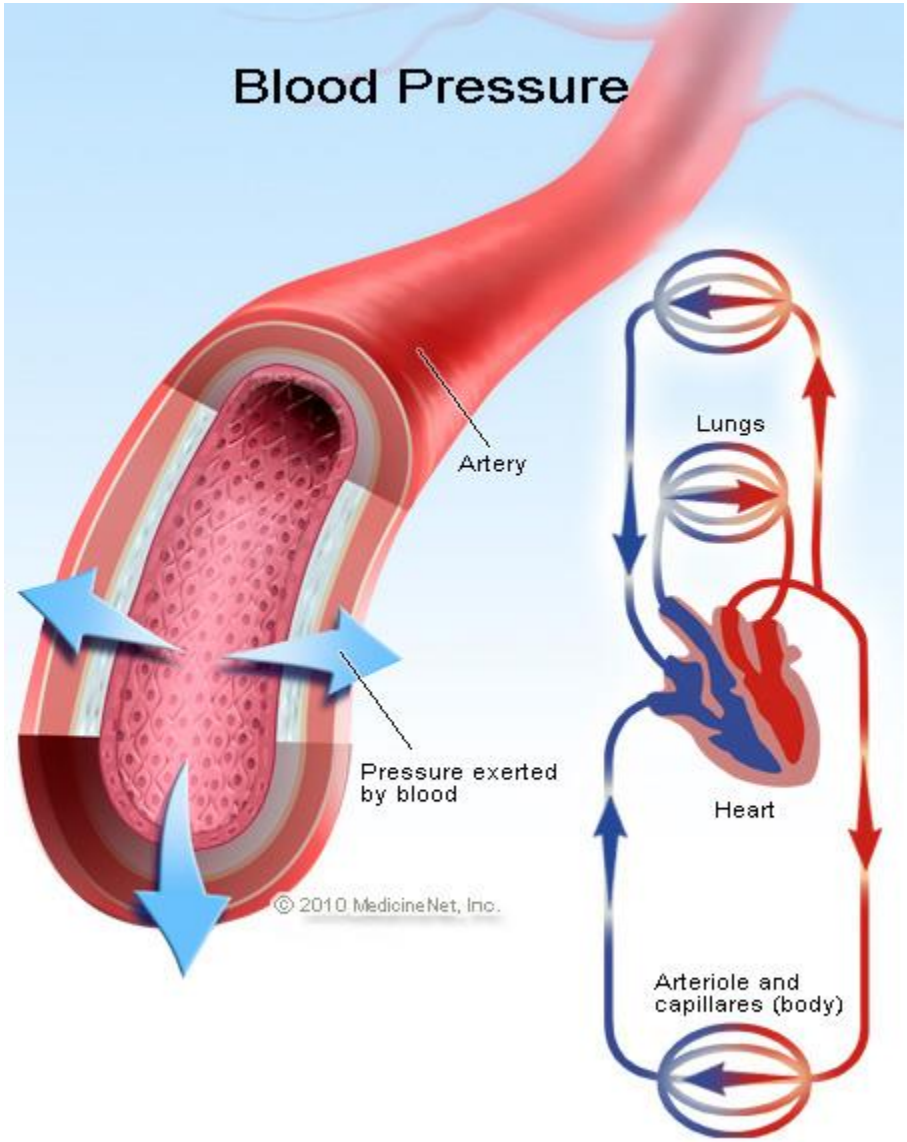
- Two numbers in the reading.
- The higher number (systolic) indicate pressure while heart pumps blood into blood vesicles
- The lower number (diastolic) indicate pressure while heart relaxes between two beats

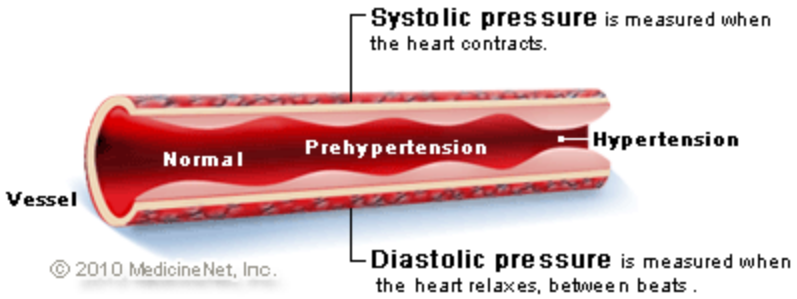
- Normal blood pressure is below 120/80; blood pressure between 120/80 and 139/89 is called "pre-hypertension", and a blood pressure of 140/90 or above is considered high.

- An elevation of the systolic and/or diastolic blood pressure increases the risk of developing heart (cardiac) disease, kidney (renal) disease, hardening of the arteries (atherosclerosis or arteriosclerosis), eye damage, and stroke (brain damage).

- These complications of hypertension are often referred to as end-organ damage because damage to these organs is the end result of chronic (long duration) high blood pressure.
- For that reason, the diagnosis of high blood pressure is important so efforts can be made to normalize blood pressure and prevent complications

- It was previously thought that rises in diastolic blood pressure were a more important risk factor than systolic elevations, but it is now known that in people 50 years or older systolic hypertension represents a greater risk.


Blood Pressure
Artery
Pressure exerted by blood
© 2010 MedicineNet, Inc.
Lungs
Heart
Arteriole and capillares (body)


Systolic pressure is measured when the heart contracts.
Normal
Prehypertension
Hypertension
Vessel
© 2010 MedicineNet, Inc.
Diastolic pressure is measured when the heart relaxes, between beats .

- For some people, blood pressure readings lower than 140/90 may be a more appropriate normal cut-off level. For example, in certain situations, such as in patients with long duration (chronic) kidney diseases that spill (lose) protein into the urine (proteinuria), the blood pressure is ideally kept at 130/80, or even lower. The purpose of reducing the blood pressure to this level in these patients is to slow the progression of kidney damage

- Patients with diabetes (diabetes mellitus) may also benefit from blood pressure that is maintained at a level lower than 130/80.

Isolated systolic high blood pressure

- A systolic blood pressure that is persistently higher than 140 mm Hg is usually considered elevated, especially when associated with an elevated diastolic pressure (over 90).

- Isolated systolic hypertension, however, is defined as a systolic pressure that is above 140 mm Hg with a diastolic pressure that still is below 90. This disorder primarily affects older people and is characterized by an increased (wide) pulse pressure. The pulse pressure is the difference between the systolic and diastolic blood pressures

- An elevation of the systolic pressure without an elevation of the diastolic pressure, as in isolated systolic hypertension, therefore, increases the pulse pressure. Stiffening of the arteries contributes to this widening of the pulse pressure.

- Isolated systolic hypertension is associated with a two to four times increased future risk of an enlarged heart, a heart attack, a stroke (brain damage), and death from heart disease or a stroke.

Borderline high blood pressure

- Borderline hypertension is defined as mildly elevated blood pressure higher than 140/90 mm Hg at some times, and lower than that at other times. Patients with borderline hypertension need to have their blood pressure taken on several occasions and their end-organ damage assessed in order to establish whether their hypertension is significant.

- People with borderline hypertension may have a tendency as they get older to develop more sustained or higher elevations of blood pressure. They have a modestly increased risk of developing heart-related (cardiovascular) disease.

Causes

- Two forms of high blood pressure have been described: essential (or primary) hypertension and secondary hypertension. Essential hypertension is a far more common condition and accounts for 95% of hypertension.

Essential hypertension

- affects approximately 72 million Americans
- The cause is multifactorial.
- Basic cause is unknown
- However, associations have been made

Essential hypertension

- Essential hypertension develops in groups or societies that have a fairly high intake of salt, exceeding 5.8 grams daily.
- Excess salt may be involved in the hypertension that is associated with advancing age, obesity, hereditary (genetic) susceptibility, and kidney failure (renal insufficiency).

- Genetic factors are thought to play a prominent role in the development of essential hypertension.
- Approximately 30% of cases of essential hypertension are attributable to genetic factors
- However, the genes for hypertension have not yet been identified.

- The current research in this area is focused on the genetic factors that affect the renin-angiotensin-aldosterone system. This system helps to regulate blood pressure by controlling salt balance and the tone (state of elasticity) of the arteries.

Essential hypertension

- The vast majority of patients with essential hypertension have in common a particular abnormality of the arteries: an increased resistance (stiffness or lack of elasticity) in the tiny arteries that are most distant from the heart (peripheral arteries or arterioles).

Secondary hypertension

- Secondary to a specific abnormality in one of the organs or systems of the body.

- Abnormality may be in kidney, adrenal gland, or aortic artery.

Renal (kidney) hypertension

- Hypertension secondary to a renal disorder.
- One important cause of renal hypertension is narrowing (stenosis) of the artery that supplies blood to the kidneys (renal artery).

- In younger individuals, usually women, the narrowing is caused by a thickening of the muscular wall of the arteries going to the kidney (fibromuscular hyperplasia). In older individuals, the narrowing generally is due to hard, fat-containing (atherosclerotic) plaques that are blocking the renal artery.

- Narrowed renal artery impairs the circulation of blood to the affected kidney.
- That stimulates the kidney to produce the hormones, renin and angiotensin.
- These hormones, along with aldosterone from the adrenal gland, cause a constriction and increased stiffness (resistance) in the peripheral arteries throughout the body.

- Renal hypertension is usually first suspected when high blood pressure is found in a young individual or a new onset of high blood pressure is discovered in an older person.
- A narrowing of the renal artery may be treated by balloon angioplasty.

- Any of the other types of chronic kidney disease that reduces the function of the kidneys can also cause hypertension due to hormonal disturbances and/or retention of salt.
- Hypertension can also cause kidney disease. Therefore, all patients with high blood pressure should be evaluated for the presence of kidney disease

Adrenal gland tumors

- Tumors of the adrenal glands are less common, secondary causes of hypertension. The adrenal glands sit right on top of the kidneys. Both of these tumors produce excessive amounts of adrenal hormones that cause high blood pressure.
- Also, certain rare genetic disorders affecting the hormones of the adrenal gland can cause secondary

Coarctation of the aorta

- Rare hereditary disorder that is one of the most common causes of hypertension in children.
- Characterized by a narrowing of a segment of the aorta.
- The narrowed segment (coarctation) of the aorta generally occurs above the renal arteries, which causes a reduced blood flow to the kidneys.
- This lack of blood to the kidneys prompts the renin-angiotensin-aldosterone hormonal system to elevate the blood pressure.
- Treatment of the coarctation is usually the surgical correction of the narrowed segment of the aorta.

Symptoms of high blood pressure

- Uncomplicated high blood pressure usually occurs without any symptoms (silently) and so hypertension has been labeled "the silent killer."
- Uncomplicated hypertension may be present and remain unnoticed for many years, or even decades.

- Some people with uncomplicated hypertension, however, may experience symptoms such as headache, dizziness, shortness of breath, and blurred vision.
- Often, however, a person's first contact with a physician may be after significant damage to the end-organs has occurred.
- That includes stroke, kidney failure, and impaired vision

- Patients with accelerated or malignant hypertension (severe high blood pressure) have more obvious symptoms:

_ Severe headache

_ Nausea

_ Visual symptoms

_ Dizziness

End-organ damage

- Damage of organs fed by the circulatory system due to uncontrolled hypertension.

- Include enlarged heart, kidney failure, brain or neurological damage, and changes in the retina at the back of the eyes.

- Increased stiffness, or resistance, in the peripheral arteries Increase the load on heart resulting in abnormalities including enlargement
- Heart enlargement may be a step leading to heart failure, coronary (heart) artery disease, and cardiac arrhythmias.

- Uncontrolled hypertension can cause strokes.
- Strokes are usually due to a hemorrhage or a blood clot (thrombosis) of the blood vessels that supply blood to the brain.
- A stroke can cause weakness, or paralysis of the arms or legs and difficulties with speech or vision
- Multiple small strokes can lead to dementia

Congenital heart disease

Congenital heart disease

- Most Common group of life threatening anomalies
- 8/1000 live births
- VSD form 35-50% of cases

Causes of CHD

- Mainly causes are unknown. However, these diseases occur with:

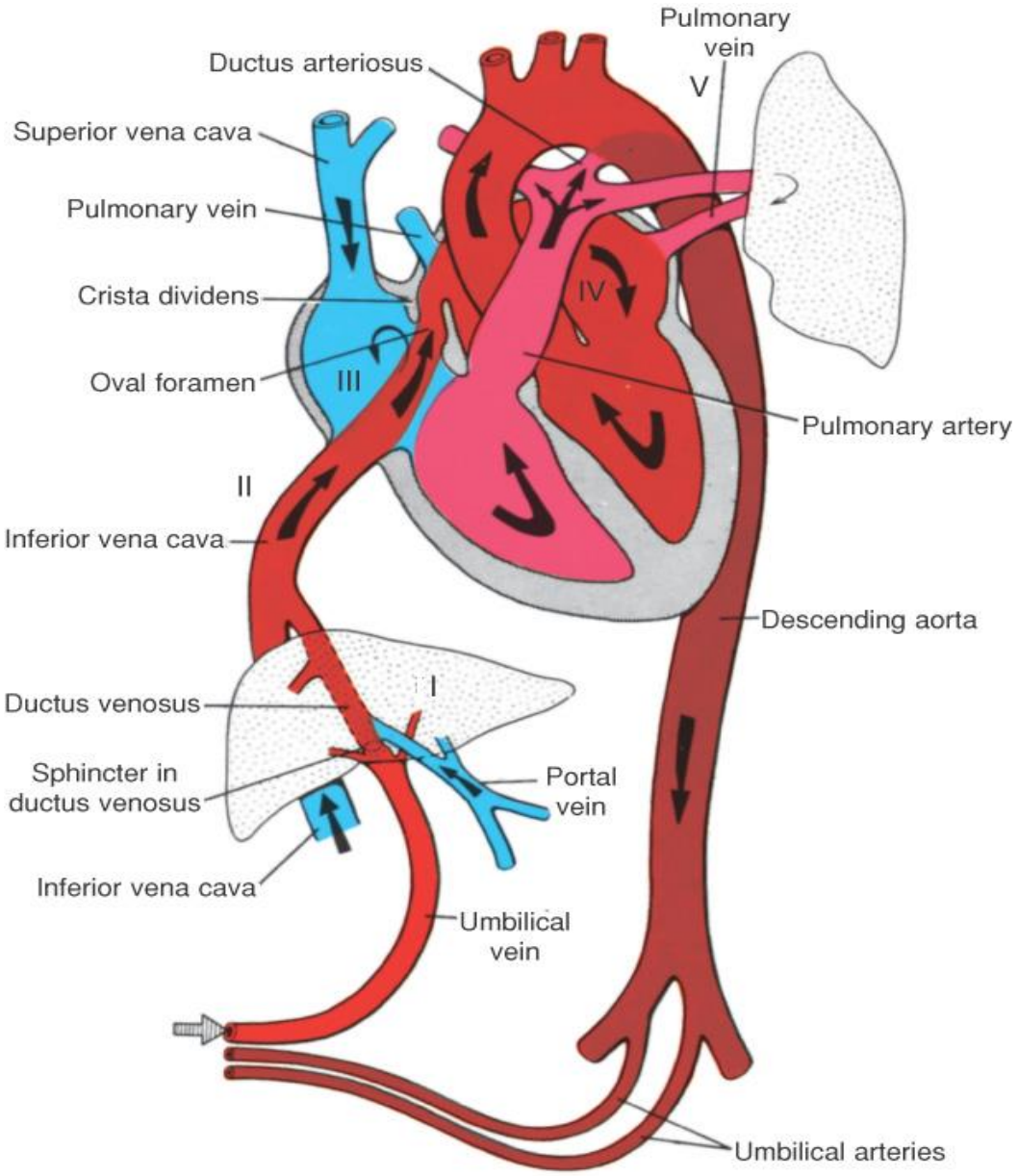
- Genetic or chromosomal defects such as Down syndrome
- Substance abuse during pregnancy
- Maternal infections during the first trimester

Classifications

1. Cyanotic/ a cyanotic
2. Disease of valve/ walls/ muscle
3. Diseases that increase/ decrease pulmonary artery flow

Fetal circulation

- Right-to-left shunting at atrial level (Foramen ovale) and at arterial level (ductus arteriosus)
- High pulmonary vascular resistance
- Little pulmonary blood flow
- Ventricles work in parallel


Pulmonary vein
Ductus arteriosus
V
Superior vena cava
Pulmonary vein
Crista dividens
IV
Oval foramen
III
Pulmonary artery
II
Inferior vena cava
Descending aorta
I
Ductus venosus
Sphincter in ductus venosus
Portal vein
Inferior vena cava
Umbilical vein
Umbilical arteries

Transition From the Fetal Circulation

- Pulmonary vascular resistance falls
- Ductus venosus and ductus arteriosus close
- Right-to-left shunting through foramen ovale ceases

Congenital heart disease

- Obstructive congenital heart lesions
- Congenital heart lesions that increase pulmonary arterial blood flow
- Congenital heart lesions that decrease pulmonary arterial blood flow

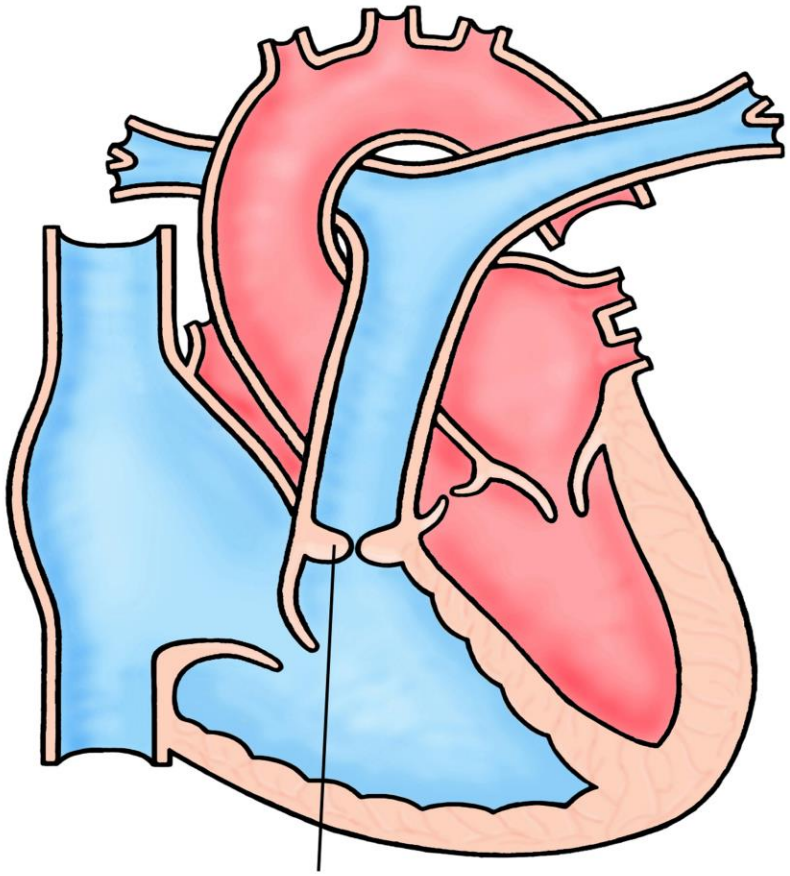
Obstructive congenital heart lesions

- Impede the forward flow of blood

- Pulmonary stenosis
- Aortic stenosis
- Coarctation of the Aorta

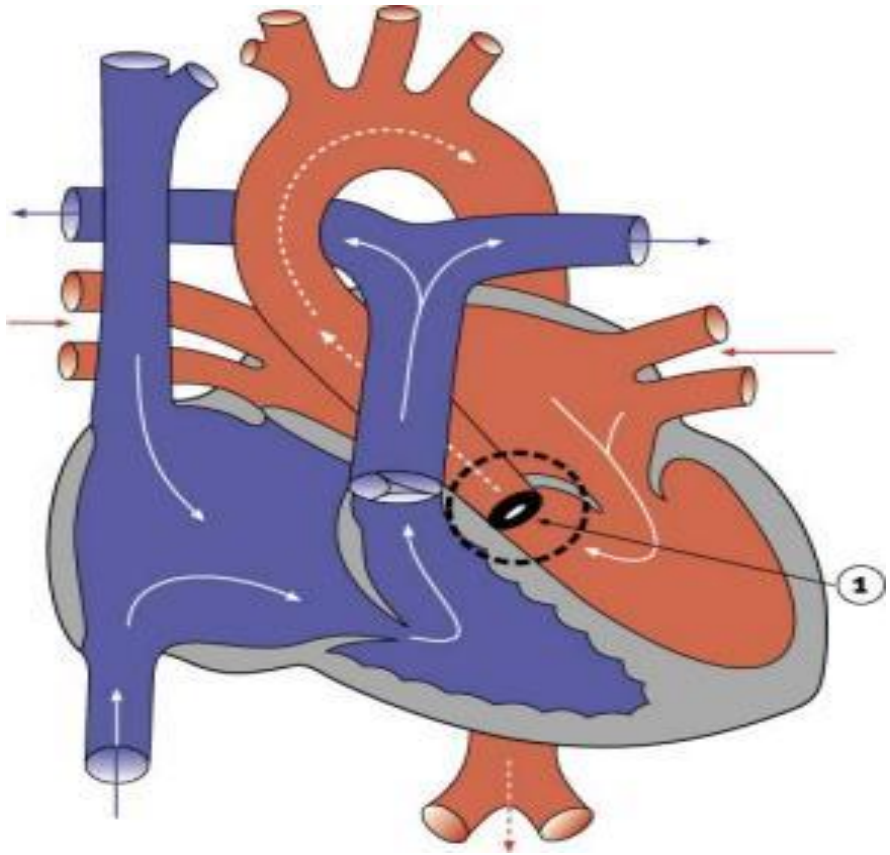
Pulmonary stenosis

- Obstruction that prevents blood to flow from the right ventricle to the pulmonary artery
- Result from valve obstruction usually
- However, it can be due to subvalvular or suprevalvular stenosis
- Silent until the case is severe
- Results in right side heart failure and cyanosis


Narrowed pulmonary valve

Aortic stenosis

- Valvular Aortic stenosis
- Subaortic stenosis
- Supravalvular Aortic stenosis
- Asymmetric septal hypertrophy


1

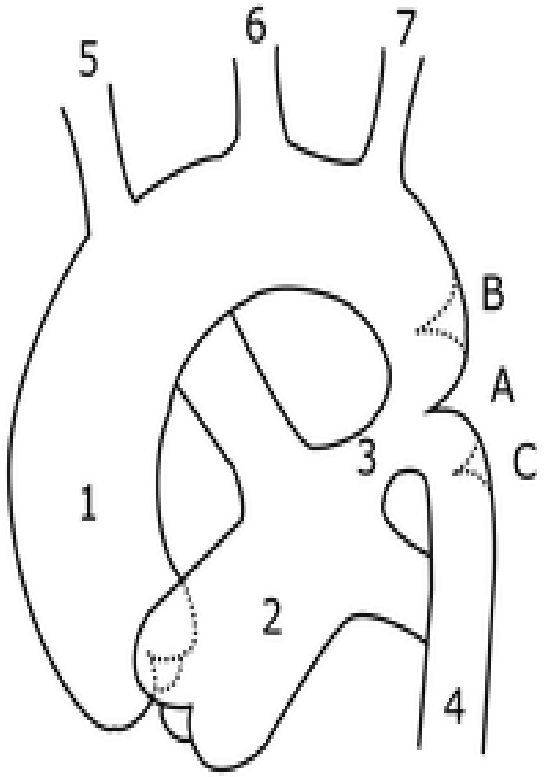
Valvular Aortic stenosis

- Most common
- Asymptomatic in children usually
- Narrow pulse pressure
- Result in heart failure
- Thickened, malformed aortic leaflets

Coarctation of the Aorta

- Narrowing of Aorta at the level of Ductus arteriosus
- Caused by both external narrowing and intraluminal membrane
- Systolic pressure higher in upper extremities than in lower extremities; diastolic pressures are similar

- Absent or weak femoral pulses
- Blood flow maintained through collateral vessels

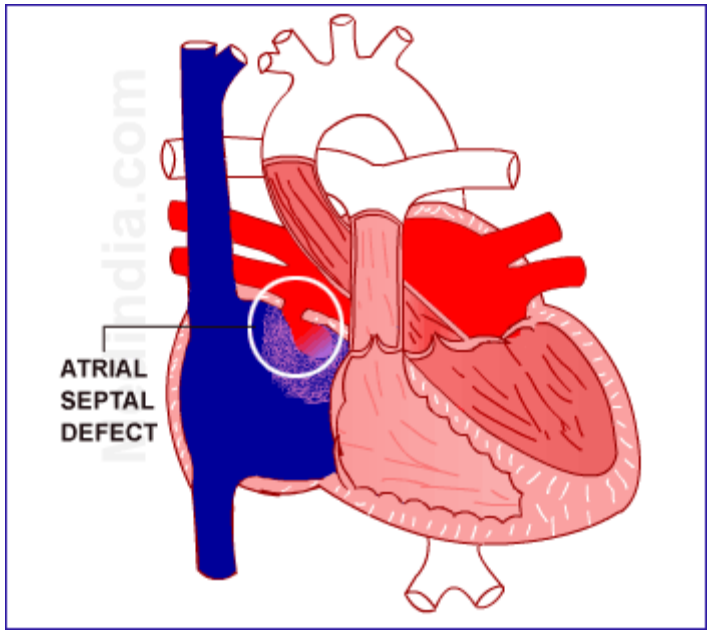

5
6
7
B
A
C
3
1
2
4

CHD increasing pulmonary flow

- Atrial septal defect
- Ventricular septal defect
- Patent ductus arteriosis
- Trancus arteriosus

Atrial septal defect

- Sometimes called patent foramen ovale
- Result in right ventricular failure, dysrhythmia, and pulmonary vascular disease

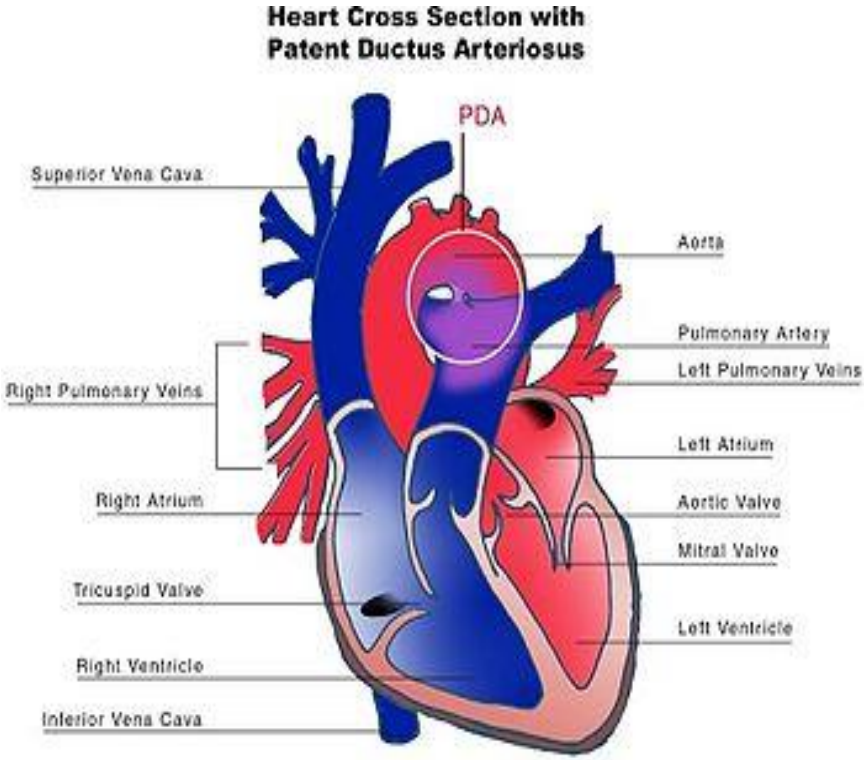

ATRIAL
SEPTAL
DEFECT

Ventricular septal defect

- Asymptomatic if small
- Result in heart failure with dyspnea with frequent respiratory infections and poor growth
- Usually a part of more complex congenital heart lesion
- Cardiomegally is expected
- Many defects close at age of 7-8

Patent ductus arteriosis

- Failure of closure after birth
- Results in respiratory distress and frequent respiratory infections and heart failure


Heart Cross Section with
Patent Ductus Arteriosus
PDA
Superior Vena Cava
Aorta
Pulmonary Artery
Left Pulmonary Veins
Right Pulmonary Veins
Left Atrium
Right Atrium
Aortic Valve
Mitral Valve
Tricuspid Valve
Left Ventricle
Right Ventricle
Inferior Vena Cava

Truncus arteriosus

- Rare form of CHD

- The structure known as truncus arteriosus fails to divide into pulmonary artery and aorta
- Accompanied with large VSD
- Cyanosis presents at birth

CHD decrease pulmonary flow

- Tetralogy of Fallot
- Transposition of the great arteries
- Tricuspid atresia

Tetralogy of Fallot

1. Pulmonary stenosis
2. VSD
3. Overriding aorta
4. Right ventricular hypertrophy

Hypoxic and cyanosis occurs usually

Transposition of the great arteries

- Aorta from right ventricle, pulmonary artery from left ventricle
- Cyanosis from birth
- Heart failure often present
- Cardiac enlargement usually occurs
- VSD, ASD, PDA must exist

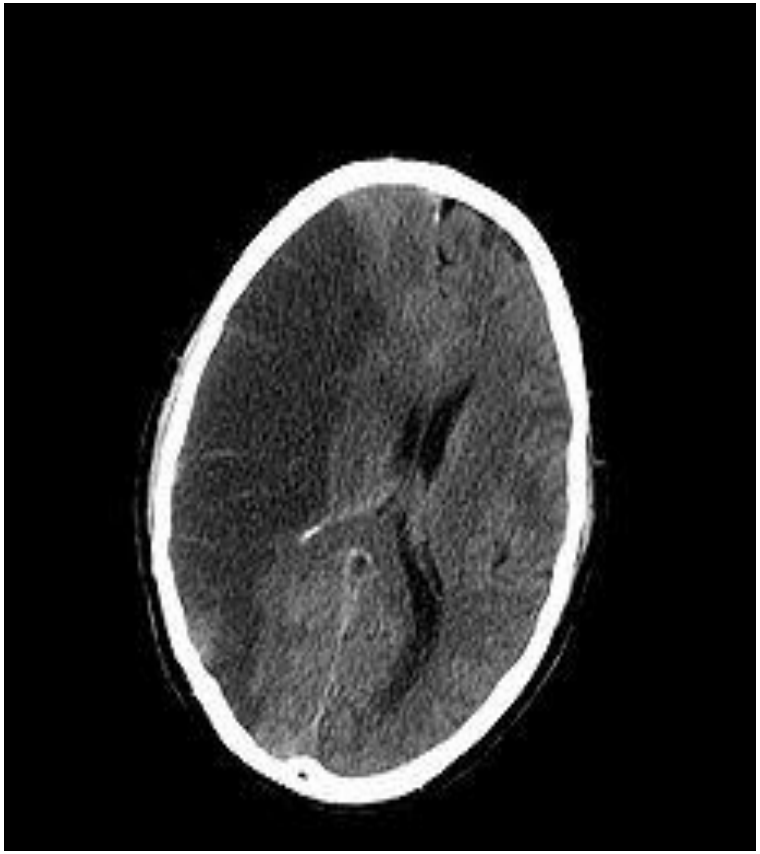
Tricuspid atresia

- Tricuspid valve is completely absent
- Result in early cyanosis

Signs of heart failure

- Fatigue
- Poor feeding- failure to thrive
- Tachypnoea
- Tachycardia
- Hepatomegally
- Sweating

CVA (Stroke)



- developing loss of brain function(s) due to disturbance in the blood supply
- CVA is considered as a medical emergency that may cause a permanent damage
- CVA is the leading cause of disability in adults and the 2nd reason of death worldwide

Definition

- WHO: "neurological deficit of cerebrovascular cause that persists beyond 24 hours or is interrupted by death within 24 hours.
- According to this definition, problem that last less than 24 hours is considered as TIA

Classification

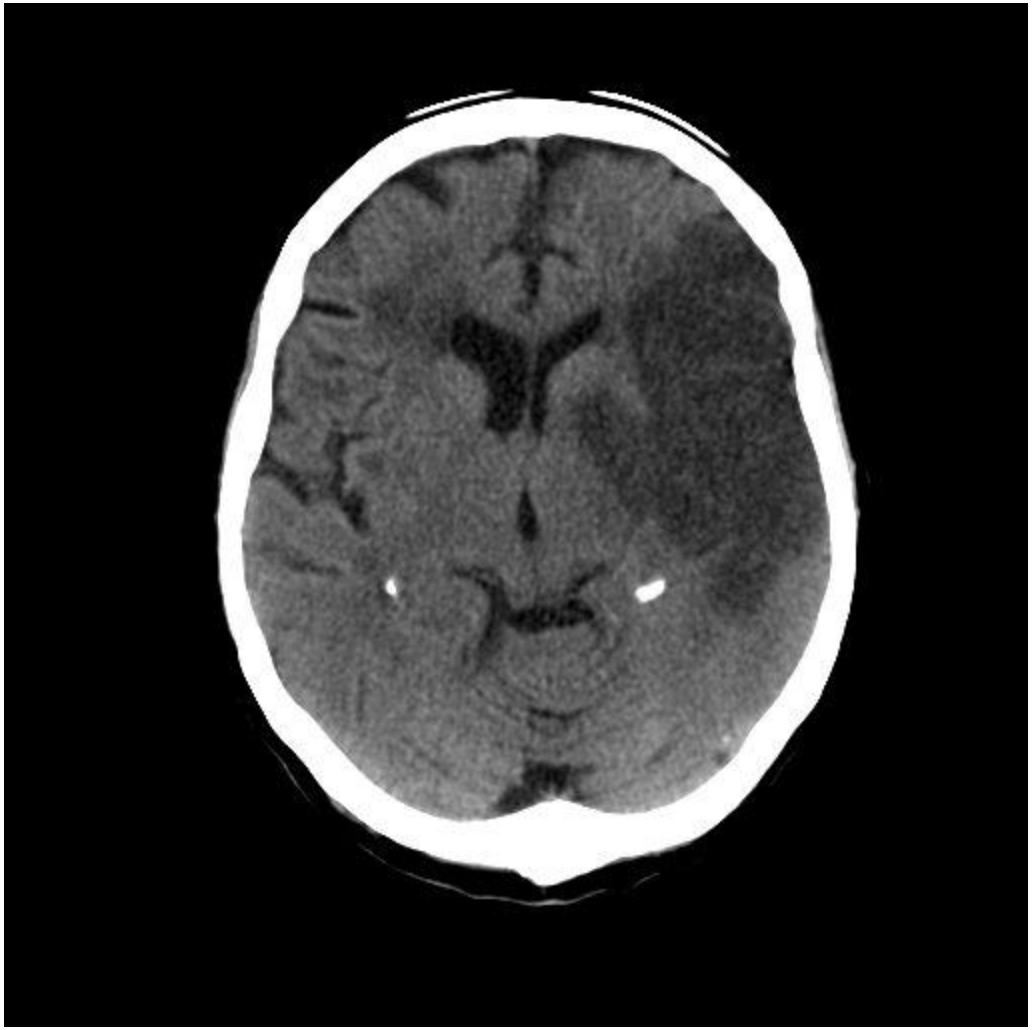
- Strokes are divided into ischemic (caused by interruption of blood supply to the brain) and hemorrhagic (resulting from a rupture in a blood vessel or abnormal vascular structure)

Ischemic CVAs

- Decreased blood supply to areas of the brain.
- Causes:

1. Thrombosis
2. Embolism
3. Systemic hypoperfusion
4. Venus thrombosis
5. Cryptogenic stroke (30-40% of ischemic strokes)

Ischemia

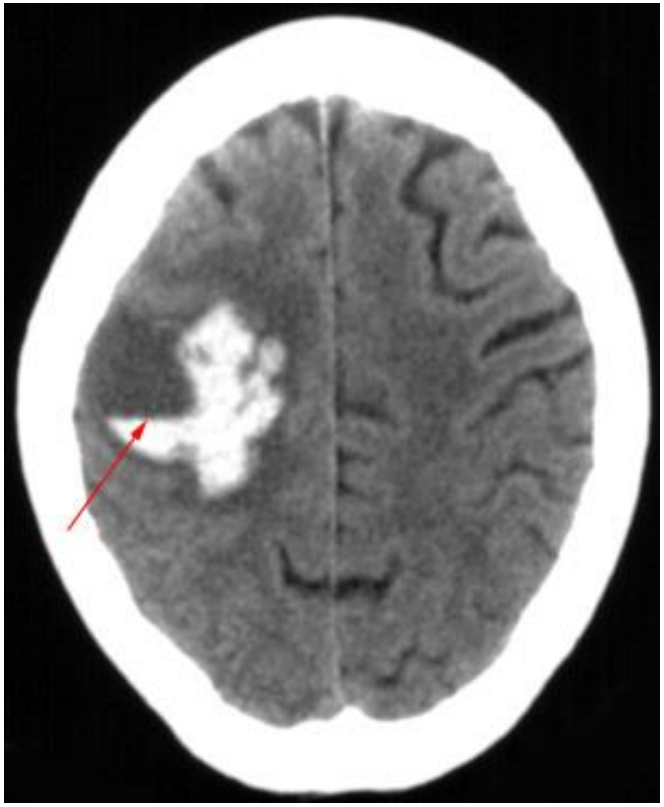


Hemorrhagic CVAs

- Intracranial hemorrhage: accumolation of blood anywhere within the skull vault
- Two types of brain hemorrhage

1. Intra axial hemorrhage (inside the brain)
2. Extra axial hemorrhage (epidural, dural, subdural, subarachnoid)

Hemorrhage



Signs and symptoms

- Symptoms of stroke are occur suddenly
- Symptoms vary depending on the affected part of the brain
- Symptoms may include headache, blurred vision, weakness of face and extremities, and seizures
- Loss of consciousness, vomiting, and headache occur more frequently in hemorrhage

Subtypes

- Stroke may affect CNS pathways, Spinothalamic tract, and Corticospinal tract and result in
- Hemiplegia
- Numbness
- Reduced sensation

Brainstem strokes

- Strokes resulting in alteration of cranial nerve functions and may include:

- Altered sensations
- Decreased reflexes (including gag and swallowing reflexes)
- Balance problems
- Altered breathing and heart rate

Cortical symptoms

- Aphasia
- Dysarthria
- Apraxia
- Memory deficits
- Hemineglect
- Confusion and disorganized thinking

Cerebellum symptoms

- Problems in walking
- Altered movement coordination
- Vertigo and disequilibrium

Thrombotic stroke

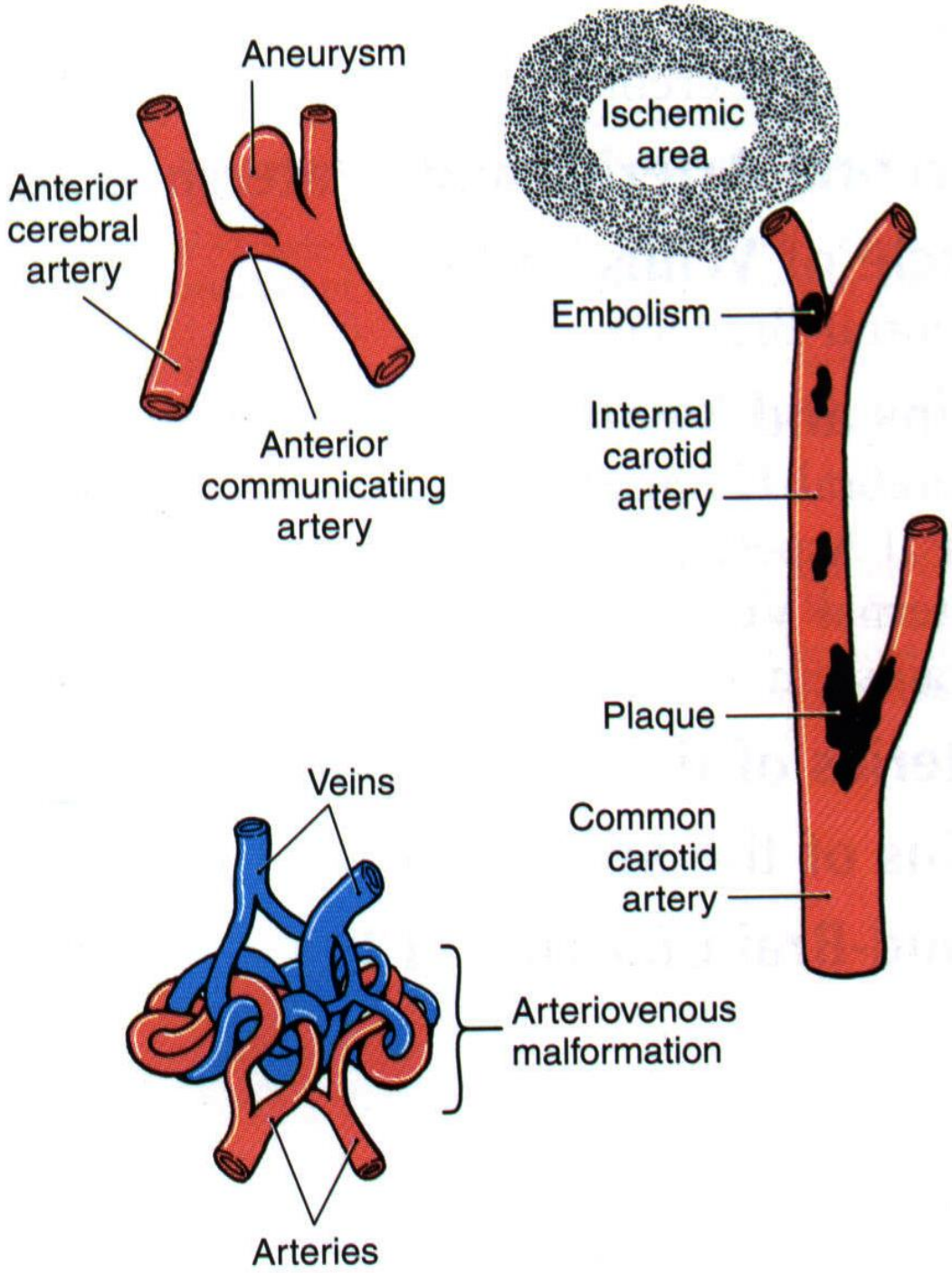
- a thrombus (blood clot) usually forms around atherosclerotic plaques. Since blockage of the artery is gradual, onset of symptomatic thrombotic strokes is slower.

- A thrombus itself can lead to an embolic stroke if the thrombus breaks off

Large vessel disease

- involves the common and internal carotids, vertebral, and the Circle of Willis.
- Diseases that may form thrombi in the large vessels include:

1. atherosclerosis, 2. vasoconstriction 3. aortic, carotid or vertebral artery dissection, 4. inflammatory diseases of the blood vessel wall (giant cell arteritis, vasculitis), 5. noninflammatory vasculopathy


Aneurysm
Anterior cerebral artery
Anterior communicating artery
Ischemic area
Embolism
Internal carotid artery
Plaque
Common carotid artery
Veins
Arteriovenous malformation
Arteries

Small vessel disease

- Involve smaller arteries in the brain: branches of the circle of Willis, middle cerebral artery, stem, and arteries arising from the distal vertebral and basilar artery
- lipohyalinosis (build-up of fatty hyaline matter in the blood vessel as a result of high blood pressure and aging) that cause lacunar infarcts and microatheroma (small atherosclerotic plaques
- Sickle cell anemia, which can cause blood cells to clump up and block blood vessels, can also lead to stroke. A stroke is the second leading killer of people under 20 who suffer from sickle-cell anemia

Atherosclerosis

- Thickening of artery walls as the result of a build-up of fatty materials such as cholesterol.
- It affects arterial blood vessels, a chronic inflammatory response in the walls of arteries, in large part due to the accumulation of macrophage white blood cells and promoted by low-density lipoproteins without adequate removal of fats and cholesterol from the macrophages by functional high density lipoproteins (HDL).
- It is commonly referred to as a hardening of the arteries.

Atherosclerosis

- The atheromatous plaque is divided into three distinct components:

-The atheroma which is the nodular accumulation of a soft, flaky, yellowish material at the center of large plaques

-Underlying areas of cholesterol crystals

-Calcification at the outer base of older/more advanced lesions.

Atherosclerosis

- Divided into stable and unstable plaques

Stable: usually asymptomatic. It is rich in extracellular matrix and smooth muscle cells

Unstable: Rich in macrophages and foam cells

the extracellular matrix separating the lesion from the arterial lumen is usually weak and prone to rupture

- Ruptures of the fibrous cap, expose thrombogenic material, such as collagen to the circulation and eventually induce thrombus formation in the lumen

Atherosclerosis

- Modifiable risk factors:

_ D.M.

_ Disturbed serum proteins

_ Hypertension

_ Smoking

_ Vitamin B6 deficiency

Atherosclerosis

- Nonmodifiable risk factors

_ Aging

_ Gender

_ Familial history

_ Genetic abnormality

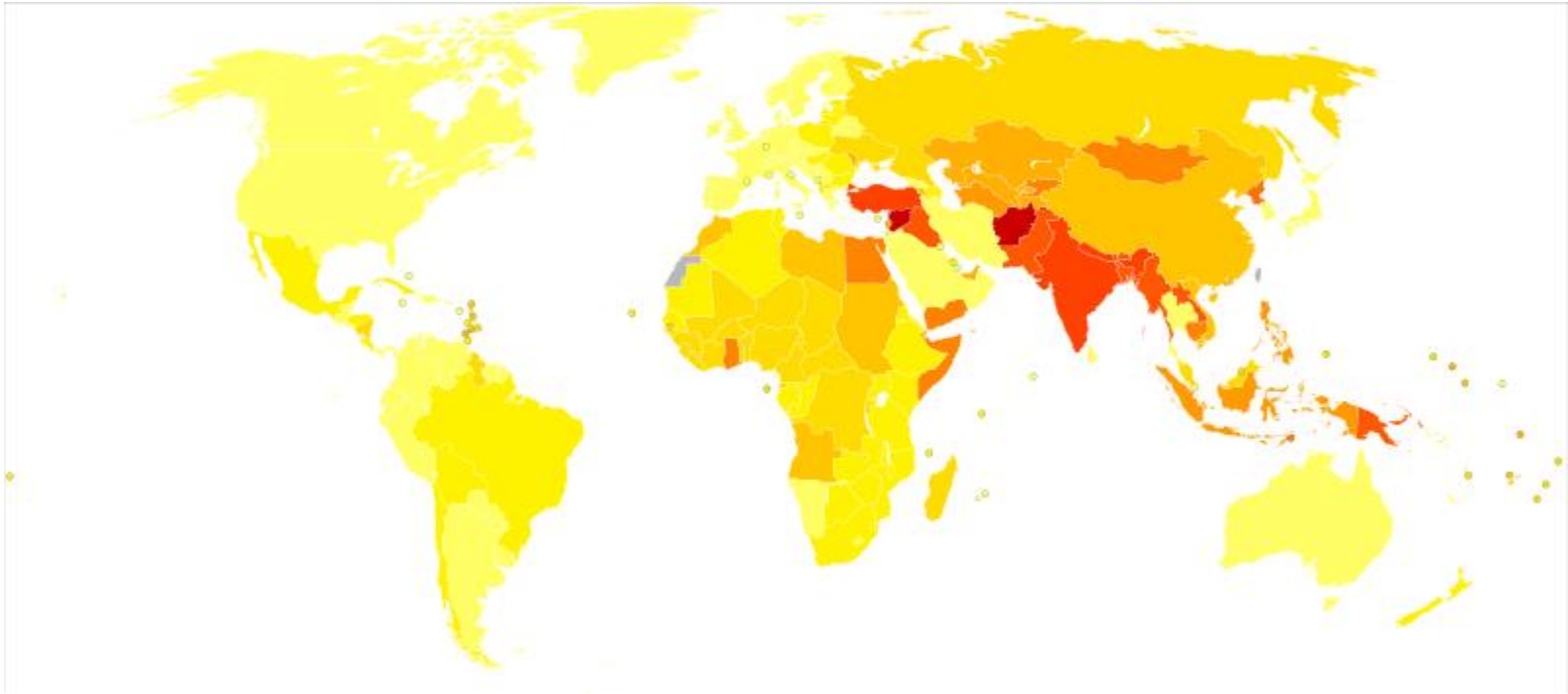
Embolic stroke

- Blockage of an artery by an embolus
- Embolus is mostly a thrombus and can be any other material such as fat (e.g. bone marrow), air, cancer cells, clumps of bacteria that is a result of endocarditis
- Heart is one major origin of embolus especially in case of atrial fibrillation
- Deep vein thrombus may cause embolic stroke in cases of ASD and VSD
- Symptoms are usually maximum at beginning

High risk cardiac causes

- Atrial fibrillation
- Rheumatic disease of the mitral or aortic valve disease
- Artificial heart valves
- Sustained atrial flutter
- Recent myocardial infarction, chronic myocardial infarction together with ejection fraction <28 percent, symptomatic congestive heart failure with ejection fraction <30 percent, dilated cardiomyopathy, endocarditis.
- Coronary artery bypass graft (CABG) surgery

Rheumatic heart disease map



Low risk cardiac causes

- Calcification of the mitral valve

- Patent foramen ovale (PFO), atrial septal aneurysm, atrial septal aneurysm *with* patent foramen ovale.

Systemic hypoperfusion

- Result in reduction of blood flow to all body organs
- Result mainly from cardiac pump failure due to cardiac arrest or disrhythmias

Reduced blood output due to myocardial infarction, pulmonary embolism, pericardial effusion, or bleeding

- Hypoxemia increase the effect of hypoperfusion

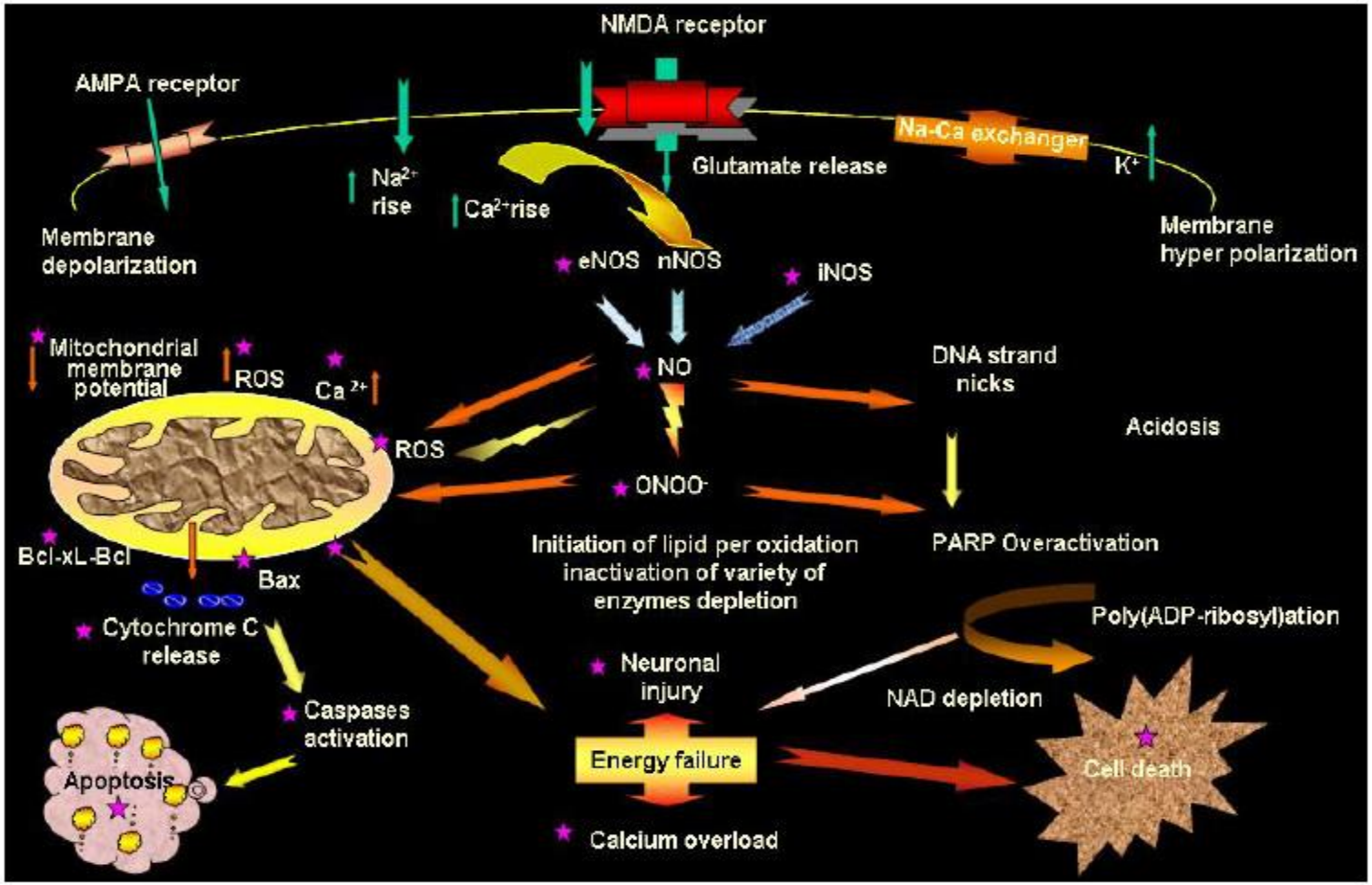
Venous thrombosis

- Cerebral venous sinus thrombosis leads to stroke due to locally increased venous pressure, which exceeds the pressure generated by the arteries. Infarcts are more likely to undergo hemorrhagic transformation (leaking of blood into the damaged area) than other types of ischemic

Intracerebral hemorrhage

- It generally occurs in small arteries or arterioles
- Commonly due to hypertension, intracranial vascular malformations (including arteriovenous malformations), or infarcts
- Other causes may be trauma, bleeding disorders, drugs (e.g. amphetamines or cocaine).
- The hematoma enlarges until pressure from surrounding tissue limits its growth, or until it decompresses by emptying into the ventricular system or other cavities

Ischemic sacade



Pathophysiology

- Ischemic stroke occurs due to a loss of blood supply to part of the brain, initiating the ischemic cascade
- Brain tissue ceases to function if deprived of oxygen for more than 60 to 90 seconds and after approximately three minutes, will suffer irreversible injury possibly leading to death of the tissue, i.e., infarction.
- Since blood vessels in the brain are now occluded, the brain becomes low in energy, and thus it resorts into using anaerobic respiration within the region of brain tissue affected by ischemia.
- This kind of respiration produces less adenosine triphosphate (ATP) but releases lactic acid which could potentially destroy cells since it is an acid and disrupts the normal acid-base balance in the brain.

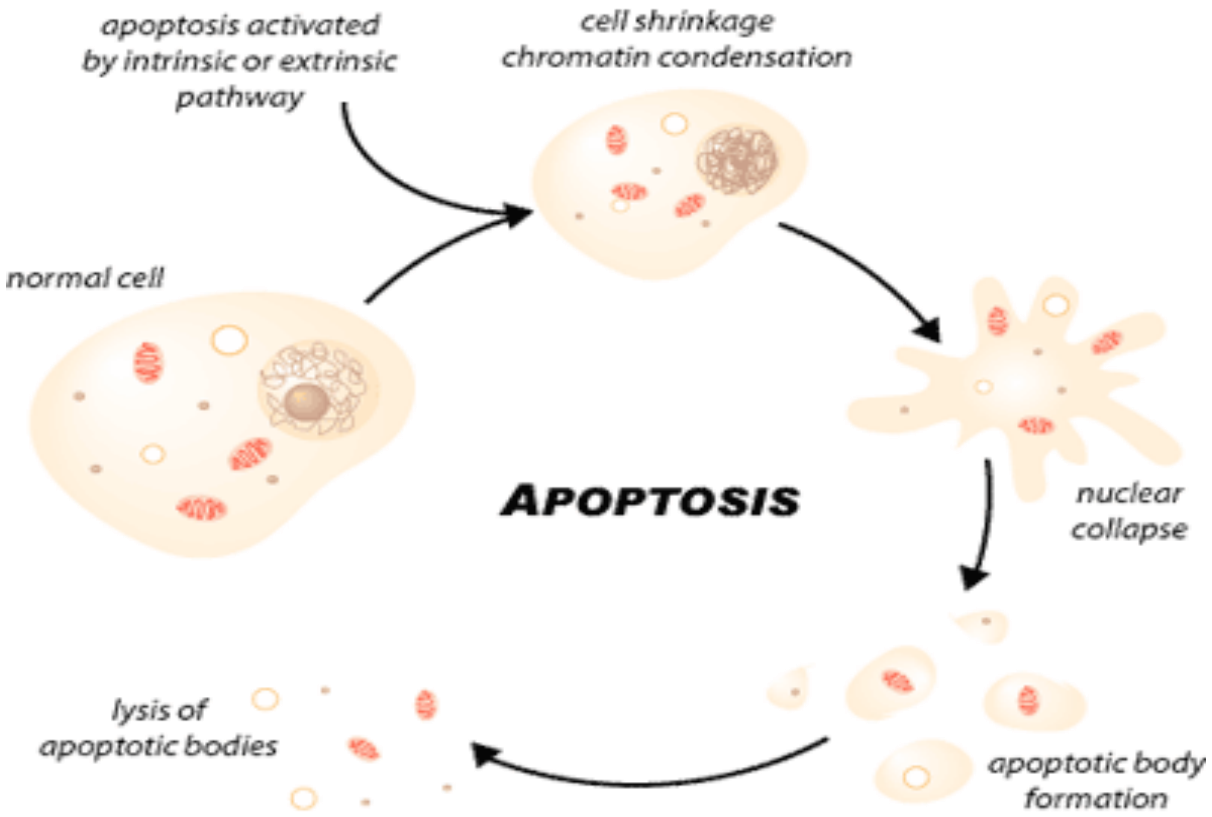
- Failure of the production of high energy phosphate compounds such as adenosine triphosphate (ATP) leads to failure of energy-dependent processes (such as ion pumping) necessary for tissue cell survival. This sets off a series of interrelated events that result in cellular injury and death.

- A major cause of neuronal injury is release of Glutamate. The concentration of glutamate outside the cells of the nervous system is normally kept low by so-called uptake carriers, which are powered by the concentration gradients of ions (mainly Na+) across the cell membrane. However, stroke cuts off the supply of oxygen and glucose which powers the ion pumps maintaining these gradients. As a result the transmembrane ion gradients run down, and glutamate transporters reverse their direction, releasing glutamate into the extracellular space.

- Glutamate acts on receptors in nerve cells (especially NMDA receptors), producing an influx of calcium which activates enzymes that digest the cells' proteins, lipids and nuclear material. Calcium influx can also lead to the failure of mitochondria, which can lead further toward energy depletion and may trigger cell death.

- Ischemia also induces production of oxygen free radicals. These react with and damage a number of cellular and extracellular elements. Damage to the blood vessel lining or endothelium is particularly important.
- Free radicals also directly initiate elements of the apoptosis cascad.

Apoptosis


apoptosis activated by intrinsic or extrinsic pathway
cell shrinkage chromatin condensation
normal cell
APOPTOSIS
nuclear collapse
lysis of apoptotic bodies
apoptotic body formation

- In addition to injurious effects on brain cells, ischemia and infarction can result in loss of structural integrity of brain tissue and blood vessels
- Release of matrix metalloproteases, which are zinc- and calcium-dependent enzymes that break down collagen, and other elements of connective tissue.
- The loss of vascular structural integrity results in a breakdown of the protective blood brain barrier that contributes to cerebral edema, which can cause secondary progression of the brain injury.

- As is the case with any type of brain injury, the immune system is activated by cerebral infarction and may under some circumstances exacerbate the injury caused by the infarction.
- Inhibition of the inflammatory response has been shown experimentally to reduce tissue injury due to cerebral infarction, but this has not proved out in clinical field

Hemorrhagic stroke

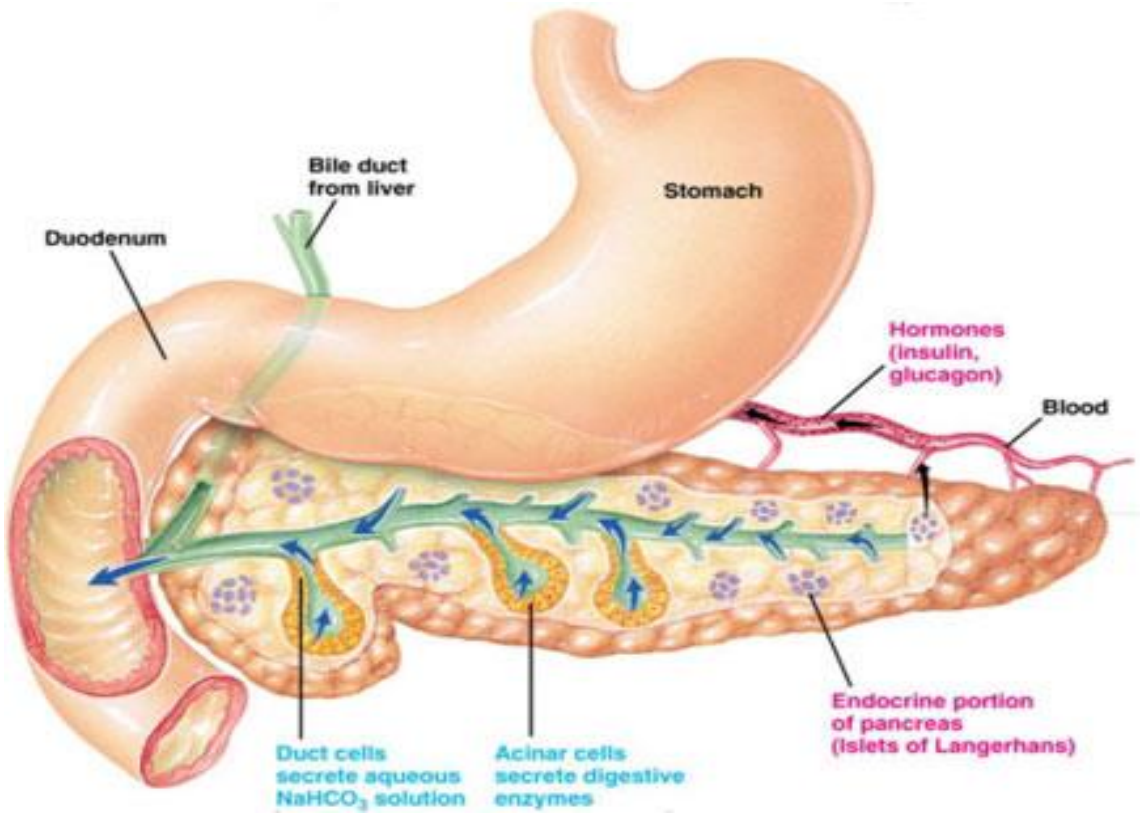
- Hemorrhagic strokes result in tissue injury by causing compression of tissue from an expanding hematoma or hematomas.
- This can distort and injure tissue. In addition, the pressure may lead to a loss of blood supply to affected tissue with resulting infarction, and the blood released by brain hemorrhage appears to have direct toxic effects on brain tissue and vasculature

Risk factors

- Hypertension
- Atrial fibrillation
- Increased cholesterol levels
- D.M.
- Smoking
- Heavy alcohol consumption
- Drugs

Diabetes

Pancreas


Bile duct
from liver
Stomach
Duodenum
Hormones
(insulin,
glucagon)
Blood
Endocrine portion
of pancreas
(Islets of Langerhans)
Duct cells
secrete aqueous
$NaHCO_3$ solution
Acinar cells
secrete digestive
enzymes

Definition

- Diabetes is a chronic metabolic disease associated with abnormally high levels of sugar (glucose) in the blood either because the body does not produce enough insulin, or because cells do not respond to the insulin that is produced.
- This high blood sugar produces the classical symptoms of polyuria (frequent urination), polydipsia (increased thirst) and polyphagia (increased hunger).

Types

- Type 1 diabetes: results from the body's failure to produce insulin, and presently requires the person to inject insulin. (Also referred to as *insulin-dependent* diabetes mellitus, and *juvenile* diabetes.)
- Type 2 diabetes: results from insulin resistance, a condition in which cells fail to use insulin properly, sometimes combined with an absolute insulin deficiency. (Formerly referred to as *non-insulin-dependent* diabetes mellitus, and *adult-onset* diabetes.)
- Gestational diabetes: is when pregnant women, who have never had diabetes before, have a high blood glucose level during pregnancy. It may precede development of type 2 DM.

Type 1 diabetes

- Characterized by loss of the insulin-producing beta cells of the islets of Langerhans in the pancreas leading to insulin deficiency.
- This type of diabetes can be further classified as immune-mediated or idiopathic.
- The majority of type 1 diabetes is of the immune-mediated nature, where beta cell loss is a T-cell mediated autoimmune attack
- There is no known preventive measure against type 1 diabetes, which causes approximately 10% of diabetes mellitus cases in North America and Europe.
- Most affected people are otherwise healthy and of a healthy weight when onset occurs.
- Type 1 diabetes can affect children or adults but was traditionally termed "juvenile diabetes" because it represents a majority of the diabetes cases in children.

Type 2 diabetes

- Type 2 diabetes mellitus is characterized by insulin resistance which may be combined with relatively reduced insulin secretion.
- The defective responsiveness of body tissues to insulin is believed to involve the insulin receptor. However, the specific defects are not known.
- Type 2 diabetes is the most common type.

Type 2 diabetes

- The release of insulin by the pancreas may also be defective and suboptimal.
- In addition to steady decline in beta cell production of insulin
- Finally, the liver in these patients continues to produce glucose through a process called gluconeogenesis despite elevated glucose levels.

Type 2 diabetes risk factors

- Strong genetic component
- Obesity in children as well as adults. Chance to develop diabetes doubles for every 20% increase over desirable body weight.
- Age
- Ethnicity
- Gestational diabetes

Secondary diabetes

- Elevated blood sugar levels from another medical condition.
- May develop when the pancreatic tissue responsible for the production of insulin is destroyed by disease, such as chronic pancreatitis (inflammation of the pancreas by toxins like excessive alcohol), trauma, or surgical removal of the pancreas.

- Diabetes can also result from other hormonal disturbances:

- Acromegaly: a pituitary gland tumor causes excessive production of growth hormone, leading to hyperglycemia.
- Cushing's syndrome: The adrenal glands produce an excess of cortisol, which promotes blood sugar elevation.

Gestational diabetes

- Gestational diabetes mellitus (GDM) resembles type 2 diabetes in several respects, involving a combination of relatively inadequate insulin secretion and responsiveness.
- It occurs in about 2%–5% of all pregnancies and may improve or disappear after delivery.
- About 20%–50% of affected women develop type 2 diabetes later in life.
- Risks to the baby include macrosomia (high birth weight), congenital cardiac and central nervous system anomalies, and skeletal muscle malformations. Increased fetal insulin may inhibit fetal surfactant production and cause respiratory distress syndrome.
- In severe cases, perinatal death may occur, most commonly as a result of poor placental perfusion due to vascular impairment.

Pathophysiology

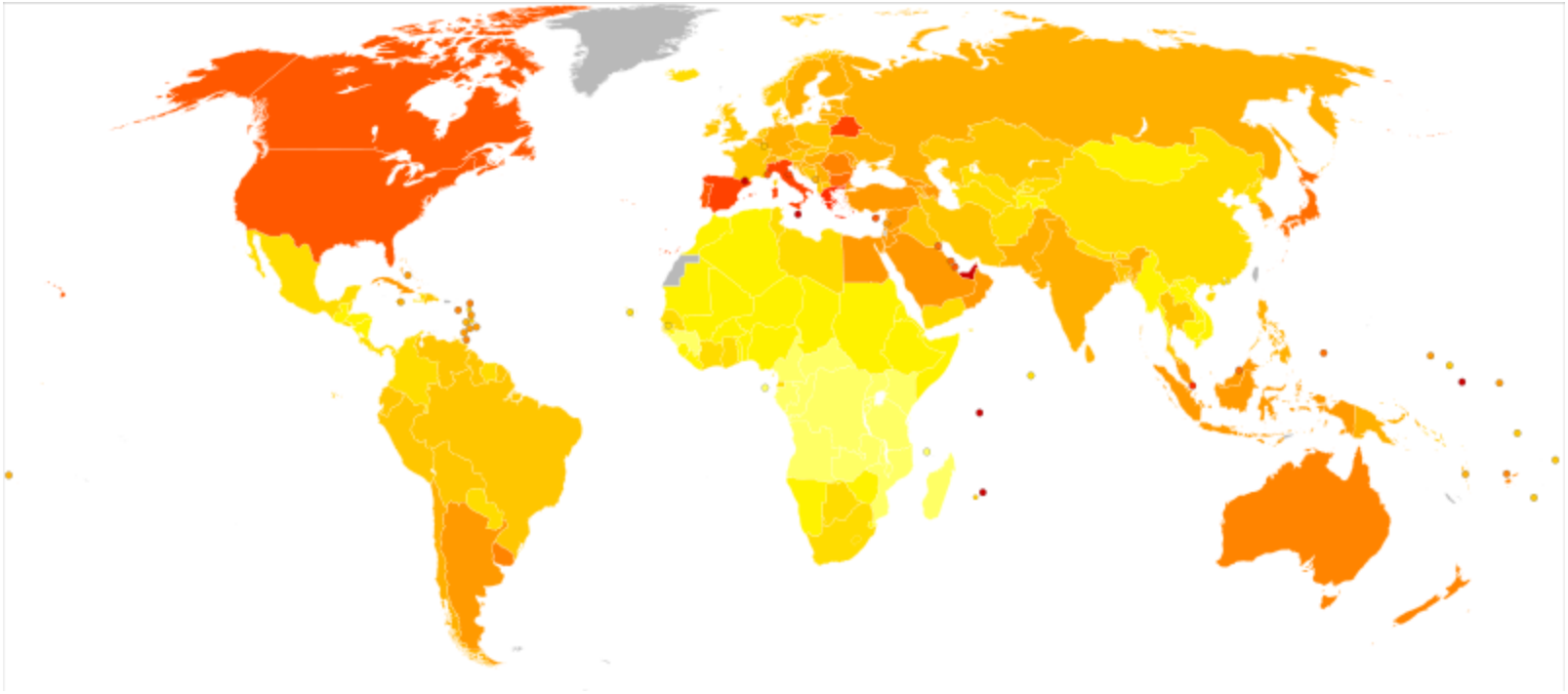
- Carbohydrates are converted to simpler forms of monosaccharide glucose, the principal energy source used by the body.

- Insulin is the principal hormone that regulates uptake of glucose from the blood into most cells (primarily muscle and fat cells, but not central nervous system cells).
- Insulin is released into the blood by beta cells (β-cells), found in the Islets of Langerhans in the pancreas, in response to rising levels of blood glucose, typically after eating.
- cells use glucose as fuel, for conversion to other needed molecules, or for storage.

- Lowered glucose levels result both in the reduced release of insulin from the beta cells and in the reverse conversion of glycogen to glucose when glucose levels fall.
- This is mainly controlled by the hormone glucagon which acts in the opposite manner to insulin.
- Glucose thus forcibly produced from internal liver cell stores (as glycogen) re-enters the bloodstream

- Lack of insulin result in improper glucose absorption by body cells and will not be stored in the liver and muscles.
- This result in persistent high levels of blood glucose, poor protein synthesis, and other metabolic derangements, such as acidosis.
- When the glucose concentration in the blood is raised beyond its renal threshold (about 10 mmol/L, although this may be altered in certain conditions, such as pregnancy), reabsorption of glucose in the proximal renal tubuli is incomplete, and part of the glucose remains in the urine (glycosuria).
- This increases the osmotic pressure of the urine and inhibits reabsorption of water by the kidney, resulting in increased urine production (polyuria) and increased fluid loss.
- Lost blood volume will be replaced osmotically from water held in body cells and other body compartments, causing dehydration and increased thirst.

Epidemiology



Signs and symptoms

- The classical symptoms of diabetes are polyuria (frequent urination), polydipsia (increased thirst) and polyphagia (increased hunger).
- Weight loss despite an increase in appetite.
- Patients with diabetes are prone to developing infections of the bladder, and skin.
- Fluctuations in blood glucose levels can lead to blurred vision. Extremely elevated glucose levels can lead to lethargy and coma.

Diagnosis

- The fasting blood glucose (sugar) test is the preferred way to diagnose diabetes.
- After the person has fasted overnight (at least 8 hours), blood level of glucose is measured

- Normal fasting plasma glucose levels are less than 100 milligrams per deciliter (mg/dl).

- Fasting plasma glucose levels of more than 126 mg/dl on two or more tests on different days indicate diabetes.

- A random blood glucose test can also be used to diagnose diabetes. A blood glucose level of 200 mg/dl or higher indicates diabetes.

Oral glucose tolerance test

- A gold standard for making the diagnosis of type 2 diabetes. It is still commonly used for diagnosing gestational diabetes.
- The person fasts overnight (at least eight but not more than 16 hours). Then first, the fasting plasma glucose is tested. After this test, the person receives 75 grams of glucose (100 grams for pregnant women).
- Blood samples are taken at specific intervals to measure the blood glucose.

Acute complications

- Severely elevated blood sugar levels due to an actual lack of insulin or a relative deficiency of insulin.

- Abnormally low blood sugar levels due to too much insulin or other glucose-lowering medications.

Diabetic ketoacidosis

- Result from severely elevated blood sugar levels.
- This leads to increased urine glucose, which in turn leads to excessive loss of fluid and electrolytes in the urine.
- Lack of insulin also causes the inability to store fat and protein along with breakdown of existing fat and protein stores.
- This dysregulation, results in the process of ketosis and the release of ketones into the blood.
- Ketones turn the blood acidic, a condition called diabetic ketoacidosis (DKA).
- Symptoms of diabetic ketoacidosis include nausea, vomiting, and abdominal pain.

Diabetic ketoacidosis

- Urgent treatment of diabetic ketoacidosis involves the intravenous administration of fluid, electrolytes, and insulin, usually in a hospital intensive care unit.
- Dehydration can be very severe, and it is not unusual to need to replace 6-7 liters of fluid
- Without prompt medical treatment, patients with diabetic ketoacidosis can rapidly go into shock, coma, and even death.

Hyperosmolar state

- Result from severe blood sugar elevation accompanied by dehydration in patients with type 2 diabetes.
- This condition can lead to coma (hyperosmolar coma).
- Immediate treatment with intravenous fluid and insulin is important in reversing the hyperosmolar state.
- Concomitant medical conditions are more likely to exist, and these patients may have poorer health.
- Death rates from hyperosmolar coma is thus higher than in DKA.

Hypoglycemia

- Abnormally low blood sugar (glucose).
- The most common cause of low blood sugar is excessive use of insulin or other glucose-lowering medications, with delayed or absent meal.
- The condition is called an insulin reaction.

Hypogycemia

- Low blood sugar can lead to central nervous system symptoms such as:
  - dizziness
  - Confusion
  - Weakness
  - Tremors
- Usually it occurs when blood sugars are less than 65 mg/dl.
- Untreated, severely low blood sugar levels can lead to coma, seizures, and irreversible brain death. At this point, the brain is suffering from a lack of sugar, and this usually occurs somewhere around levels of <40 mg/dl.

Chronic complications

- These diabetes complications are related to blood vessel diseases
- Generally classified into:
- Small vessel disease (microvascular disease), such as those involving the eyes, kidneys and nerves
- Large vessel disease (macrovascular disease) involving the heart and blood vessels via atherosclerosis

* That leads to coronary heart disease (angina or heart attack), and strokes

Eye complications

- Diabetic retinopathy: occurs in patients who have had diabetes for at least five years. Diseased small blood vessels in the back of the eye cause the leakage of protein and blood in the retina, causes the formation of small aneurysms (microaneurysms), and new but weak blood vessels (neovascularization).
- Spontaneous bleeding from the new and weak blood vessels can lead to retinal scarring and retinal detachment, thus impairing vision.
- Approximately 50% of patients with diabetes will develop some degree of diabetic retinopathy after 10 years of diabetes, and 80% of diabetics have retinopathy after 15 years of the disease.

- Cataracts and glaucoma are also more common among diabetics.
- It is also important to note that since the lens of the eye lets water through, if blood sugar concentrations vary a lot, the lens of the eye will shrink and swell with fluid accordingly. As a result, blurry vision is very common in poorly controlled diabetes.

Kidney damage

- Called diabetic nephropathy.
- The onset of kidney disease and its progression is extremely variable.
- Initially, diseased small blood vessels in the kidneys cause the leakage of protein in the urine. Later on, the kidneys lose their ability to cleanse and filter blood.
- The accumulation of toxic waste products in the blood leads to the need for dialysis.
- The progression of nephropathy in patients can be significantly slowed by controlling high blood pressure, and by aggressively treating high blood sugar levels.

Nerve damage

- Called diabetic neuropathy.
- Symptoms of diabetic nerve damage include numbness, burning, and aching of the feet and lower extremities. Sometimes there is a complete loss of sensation
- Because of poor blood circulation, diabetic foot injuries may not heal. Sometimes, minor foot injuries can lead to serious infection, ulcers, and even gangrene, necessitating surgical amputation of toes, feet, and other infected parts.
- Diabetic nerve damage can affect the sexual ability of affected men.
- Diabetic neuropathy can also affect nerves to the stomach and intestines, causing nausea, weight loss, diarrhea, and other symptoms of gastroparesis

Space occupying lesions

What Is Cancer?

- Cancer is the general name for a group of more than 100 diseases in which cells in a part of the body begin to grow out of control. Although there are many kinds of cancer, they all start because abnormal cells grow out of control. Untreated cancers can cause serious illness and even death.

Normal cells in the body

- The body is made up of trillions of living cells. Normal body cells grow, divide, and die in an orderly fashion. During the early years of a person's life, normal cells divide faster to allow the person to grow. After the person becomes an adult, most cells divide only to replace worn-out or dying cells or to repair injuries

How cancer starts

- Cancer starts when cells in a part of the body start to grow out of control. There are many kinds of cancer, but they all start because of out-of-control growth of abnormal cells.
- Cancer cell growth is different from normal cell growth. Instead of dying, cancer cells continue to grow and form new, abnormal cells. Cancer cells can also invade (grow into) other tissues, something that normal cells cannot do. Growing out of control and invading other tissues are what makes a cell a cancer cell.
- Cells become cancer cells because of damage to DNA. DNA is in every cell and directs all its actions. In a normal cell, when DNA gets damaged the cell either repairs the damage or the cell dies. In cancer cells, the damaged DNA is not repaired, but the cell doesn't die like it should. Instead, this cell goes on making new cells that the body does not need. These new cells will all have the same damaged DNA as the first cell does.
- People can inherit damaged DNA, but most DNA damage is caused by mistakes that happen while the normal cell is reproducing or by something in our environment. Sometimes the cause of the DNA damage is something obvious, like cigarette smoking. But often no clear cause is found.
- In most cases the cancer cells form a tumor. Some cancers, like leukemia, rarely form tumors. Instead, these cancer cells involve the blood and blood-forming organs and circulate through other tissues where they grow.

How cancer spreads

- Cancer cells often travel to other parts of the body, where they begin to grow and form new tumors that replace normal tissue. This process is called metastasis. It happens when the cancer cells get into the bloodstream or lymph vessels of our body.

How cancers differ

- No matter where a cancer may spread, it is always named for the place where it started. For example, breast cancer that has spread to the liver is still called breast cancer, not liver cancer. Likewise, prostate cancer that has spread to the bone is metastatic prostate cancer, not bone cancer.
- Different types of cancer can behave very differently. For example, lung cancer and breast cancer are very different diseases. They grow at different rates and respond to different treatments. That is why people with cancer need treatment that is aimed at their particular kind of cancer

Tumors that are not cancer

- Not all tumors are cancerous. Tumors that aren’t cancer are called benign. Benign tumors can cause problems – they can grow very large and press on healthy organs and tissues. But they cannot grow into (invade) other tissues. Because they can’t invade, they also can’t spread to other parts of the body (metastasize). These tumors are almost never life threatening.

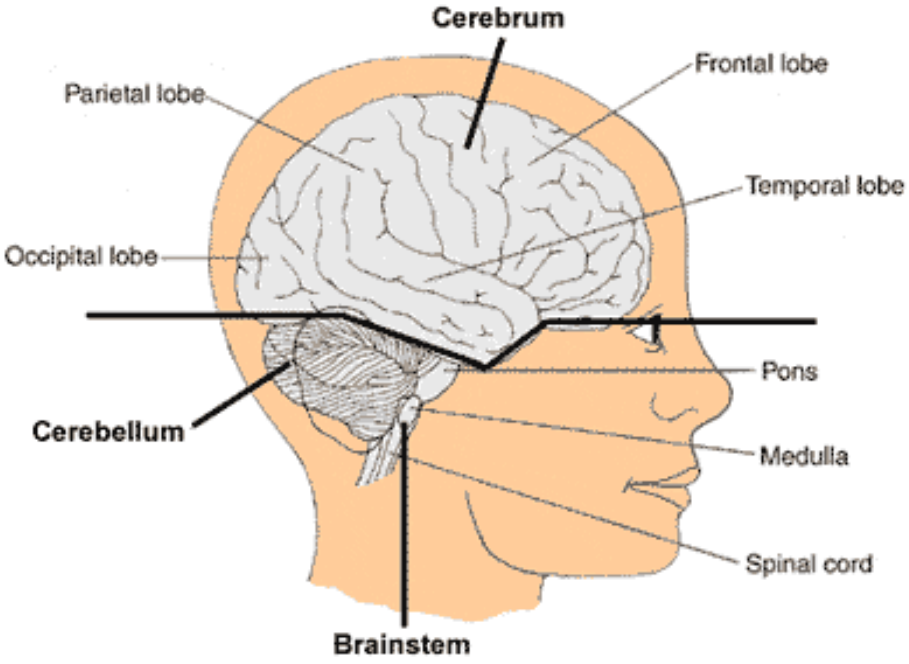
How common is cancer?

- Half of all men and one-third of all women in the US will develop cancer during their lifetimes.
- Today, millions of people are living with cancer or have had cancer. The risk of developing most types of cancer can be reduced by changes in a person's lifestyle, for example, by quitting smoking, limiting time in the sun, being physically active, and eating a better diet. The sooner a cancer is found and treated, the better the chances are for living for many years.

Brain Cancer

Significance

- The brain is the center of thoughts, emotions, memory and speech.
- Brain also control muscle movements and interpretation of sensory information (sight, sound, touch, taste, pain etc)



Significance

- Tumors can effect any part of the brain and depending on what part(s) of the brain it affects can have a number of symptoms.
  - Seizures
  - Difficulty with language
  - Mood changes
  - Change of personality
  - Changes in vision, hearing, and sensation.
  - Difficulty with muscle movement
  - Difficulty with coordination control

Background

- Estimated 18,400 primary malignant brain tumors will be diagnosed in 2004 — 10,540 in men & 7,860 in women.
- Approximately 12,690 people will die from these tumors in 2004.
- Accounts for 1.4% of all cancers
- Accounts for 2.4% of all cancer related deaths

Background

- In adults over 45 years of age 90% of all brain tumors are Gliomas
  - Gliomas: A general category of cancer that includes astrocytomas, oligodendrogliomas, and ependymomas

Astrocytoma

- Astrocytes brain cells abnormally dividing causing tumors called astrocytomas.
- Astrocytes are glial cells that help nourish neurons– they help repair damage
- How the astroytomas are classified
  - How close the cells are together within the tumor
  - How abnormal the cells are
  - How many of the cells are proliferating
  - Whether or not there are blood vessels growing near the tumor
  - Whether or not some of the cancer cells have degenerated or not

Astrocytomas--Treatments

- If tumors have not infiltrated normal brain tissue then surgery can be a cure
- Low-grade Astrocytomas are not curable by surgery. However through surgery as much of the tumor as possible is removed and then the patient usually goes through radiation treatment.

Astrocytomas--Treatment

- High-grade Astrocytomas are not curable by surgery. After surgery has removed as much of the tumor as possible the patient can go through radiation treatment and chemotherapy.

Oligodendrogliomas

- These tumors start in mutated oligodendrocyte brain cells
- Oligodendrocytes make myelin which help neurons transmit signals through the axons
- These tumors may spread through cerebrospinal fluid pathways but typically do not usually spread to locations outside of the brain or spinal cord.

Oligodendrogliomas--Treatments

- Because these tumors infiltrate normal brain tissue these tumors are not cured through surgery. However removal of part of the tumors can relieve some symptoms and prolong life.
- If the tumor is causing disabilities to the patient after surgery the patient may go through chemotherapy, perhaps followed by radiation treatments.

Ependymomas

- Mutated ependymal cells
- Ependymal cells line the ventricles in the central area of the brain and they line part of the pathway through which the cerebrospinal fluid travels
- Theses mutated cells may block the cerebrospinal fluid from exiting the ventricles causing the ventricles to enlarge (hydrocephalus)

Ependymomas--Treatments

- These tumors do not usually infiltrate normal brain tissue and are therefore curable through surgery.
- If surgery is unable to completely remove the tumors the patient may try radiation therapy.

Diagnosis

- These tumors can be detected through a MRI, CT scan or a PET scan.
- Once detected, depending on where the tumor is located, a biopsy officially is used to diagnosis cancer.

Risk Factors

- Most brain cancers happen for reasons unknown, however some small risk factors are
  - Radiation exposure
  - Exposure to vinyl chloride
  - Immune system disorders

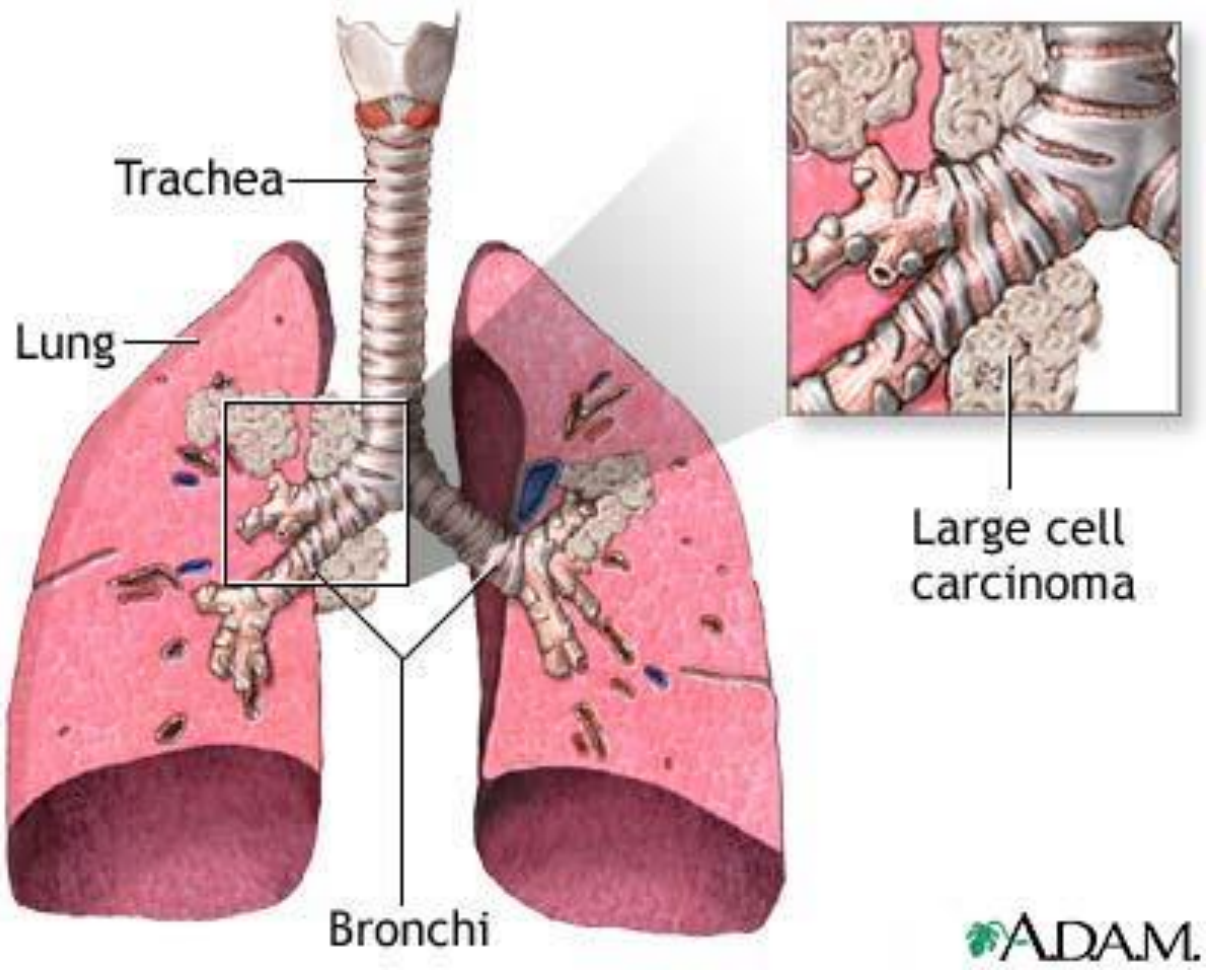
Prognosis

- For people ages 15-44 five year survival rate is 55%
- For people ages 45-64 five year survival rate is 16%
- For people over 65 five year survival rate is 5%

Lung Cancer

- Uncontrolled growth of malignant cells in one or both lungs and tracheo-bronchial tree
- A result of repeated carcinogenic irritation causing increased rates of cell replication
- Proliferation of abnormal cells leads to hyperplasia, dysplasia or carcinoma in situ

Picture of the Lungs

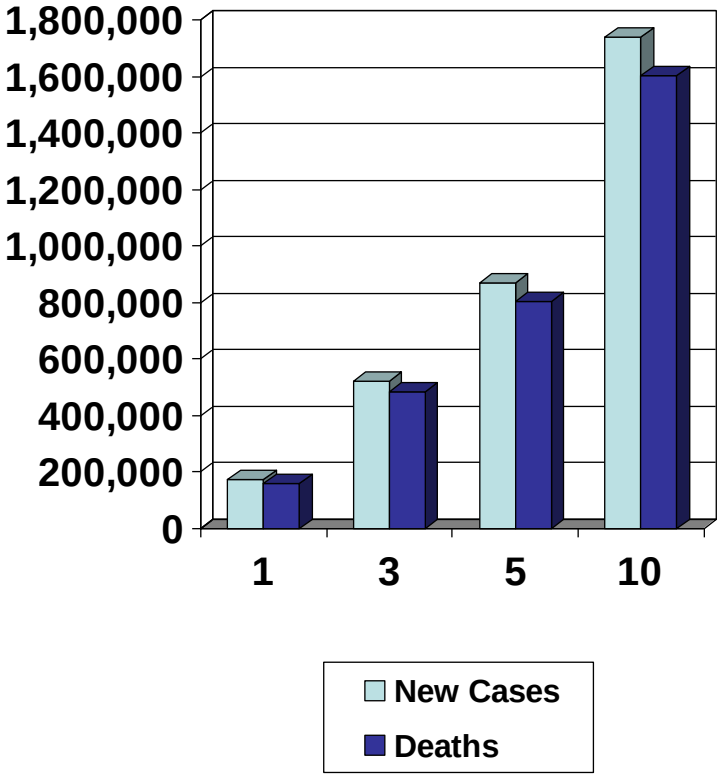


Lung Cancer in the US

- According to 2004 statistics, there were

  173,770 new cases and

  160,440 deaths yearly
- More deaths from lung cancer than prostate, breast and colorectal cancers ***combined***
- Decreasing incidence and deaths in men; continued increase in women



Women & Lung Cancer

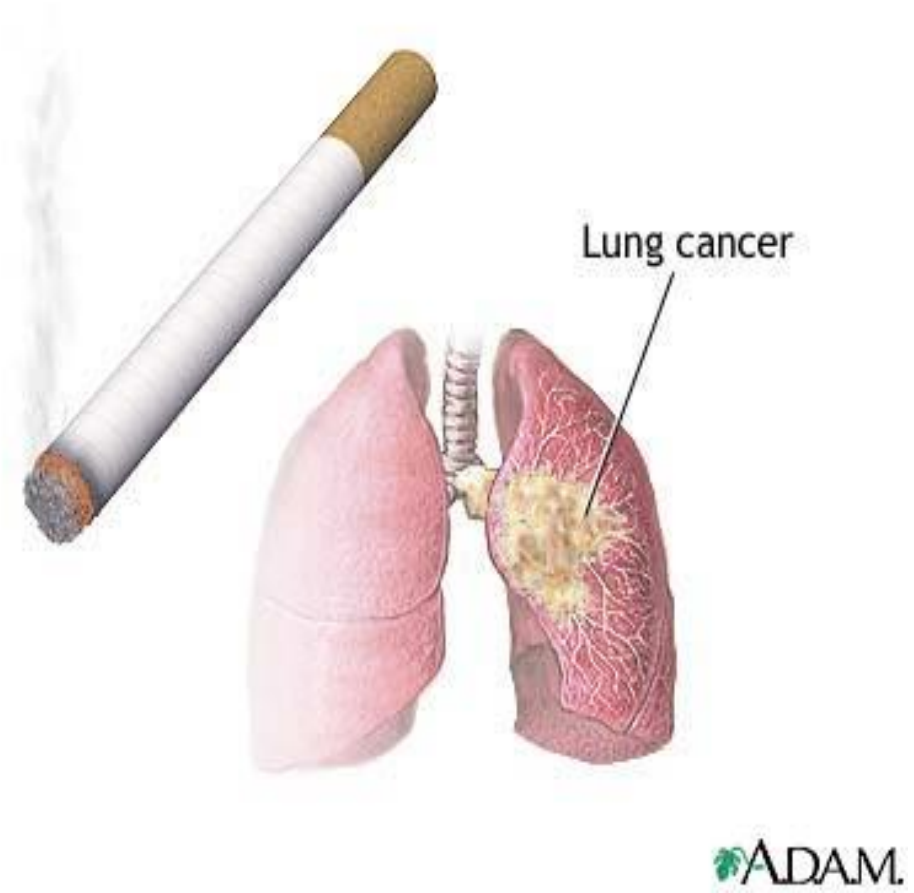
- 80,660 new cases were reported in 2004
  - Account for 12 % of all new cases
- 68,510 deaths were reported in 2004
  - An increase of 150% between 1974 and 1994
- Women are more prone to tobacco effects - 1.5 times more likely to develop lung cancer than men with same smoking habits

Where Does it Come From?

- Radiation Exposure
- Smoking
- Environmental/ Occupational Exposure
  - Asbestos
  - Radon
  - Passive smoke

Smoking Facts

- Tobacco use is the leading cause of lung cancer
- 87% of lung cancers are related to smoking
- Risk related to:
  - age of smoking onset
  - amount smoked
  - gender
  - product smoked
  - depth of inhalation



Where does it travel?

- Lymph Nodes, Brain, Liver, Adrenal, Gland, Bones

- 40% of metastasis occurs in the Adrenal Gland

Diagnosis

- History and Physical exam
- Diagnostic tests
  - Chest x-ray
  - Biopsy (bronchoscopy, needle biopsy, surgery)
- Staging tests
  - CT chest/abdomen
  - Bone scan
  - Bone marrow aspiration
  - PET scan

Symptoms

- cough
- dyspnea
- hemoptysis
- recurrent infections
- chest pain

Syndromes/Symptoms secondary to regional metastases:

- Esophageal compression ☑ dysphagia
- Laryngeal nerve paralysis ☑ hoarseness
- Symptomatic nerve paralysis ☑ Horner's syndrome
- Cervical/thoracic nerve invasion ☑ Pancoast syndrome
- Lymphatic obstruction ☑ pleural effusion
- Vascular obstruction ☑ SVC syndrome
- Pericardial/cardiac extension ☑ effusion, tamponade

Two Lung Cancer Cells, Classified

Non Small Cell Lung Cancer (NSCLC)

- Adenocarcinoma
- Squamous Cell Carcinoma
- Large Cell Carcinoma

Small Cell Lung Cancer (SCLC)

- Oat Cell
- Intermediate
- Combined

Treatment and Staging NSCLC

Stage	Description	Treatment Options
Stage I a/b	Tumor of any size is found only in the lung	Surgery
Stage II a/b	Tumor has spread to lymph nodes associated with the lung	Surgery
Stage III a	Tumor has spread to the lymph nodes in the tracheal area, including chest wall and diaphragm	Chemotherapy followed by radiation or surgery
Stage III b	Tumor has spread to the lymph nodes on the opposite lung or in the neck	Combination of chemotherapy and radiation
Stage IV	Tumor has spread beyond the chest	Chemotherapy and/or palliative (maintenance) care

SCLC

- **Limited Stage**

  Defined as tumor involvement of one lung, the mediastinum and ipsilateral and/or contralateral supraclavicular lymph nodes or disease that can be encompassed in a single radiotherapy port.

- **Extensive Stage**

  Defined as tumor that has spread beyond one lung, mediastinum, and supraclavicular lymph nodes. Common distant sites of metastases are the adrenals, bone, liver, bone marrow, and brain.

Conclusion

- Smoking cessation is essential for prevention of lung cancer.
- New screening tools under way.
- Clinical trials under way.
- New treatments under way.
- Treatment can palliate symptoms and improve quality of life.
- Read first bullet again!!

Oral Cavity and Oropharyngeal Cancer

Background Info.

- Oral Cavity = Mouth
  - Lips, inside lining of the lips and cheeks, the teeth, the gums, the front two-thirds of the tongue, the floor of the mouth below the tongue, the bony roof of the mouth (hard palate), and the area behind the wisdom teeth.

More Info…

- Oropharyngeal cancer develops in the part of the throat just behind the mouth, called the oropharynx. The oropharynx begins where the oral cavity stops. It includes the base of tongue (the back third of the tongue), the soft palate, the tonsils and tonsillar pillars, and the back wall of the throat.

More Info…

- The oral cavity and oropharynx assist with breathing, talking, eating, chewing, and swallowing. Minor salivary glands located throughout the oral cavity and oropharynx make saliva that keeps the mouth moist and helps digest food.
- contain several types of tissue and each of these tissues contains several types of cells.

Leukoplakia, Erythroplakia, and Dysplasia

Leukoplakia and **Erythroplakia**: an abnormal area in the mouth or throat.

Leukoplakia: is a white area.

Erythroplakia: is a slightly raised, red area that bleeds easily if scraped.

These white or red areas may be a cancer, or they may be a precancerous condition called dysplasia. They could also be some relatively harmless condition.

Malignant Oral Cavity and Oropharyngeal Tumors

- More than 90% of cancers of the oral cavity and oropharynx are **squamous cell carcinomas,** also called squamous cell cancer.
- Squamous cells are flat, scale-like cells that normally form the lining of the oral cavity and oropharynx.
- Invasive squamous cell cancer means that the cancer cells have spread beyond this layer into deeper layers of the oral cavity or oropharynx.

Risk Factors

- **Tobacco:** About 90% of people with oral cavity and oropharyngeal cancer use tobacco
- **Alcohol:** Drinking alcohol strongly increases a smoker's risk of developing oral cavity and oropharyngeal cancer.
- **Ultraviolet light:** More than 30% of patients with cancers of the lip have outdoor occupations associated with prolonged exposure to sunlight.
- **Irritation:** Long-term irritation to the lining of the mouth caused by poorly fitting dentures

Risk Factors Cont…

- **Poor nutrition:** A diet low in fruits and vegetables is associated with an increased risk
- **Mouthwash:** Some studies have suggested that mouthwash with a high alcohol content
- **Human papillomavirus (HPV) infection:**
- **Immune system suppression**:
- **Age:** The likelihood of developing oral and oropharyngeal cancer increases with age, especially after age 35.
- **Gender:** Oral and oropharyngeal cancer is twice as common in men as in women

Tobacco & Alcohol

- We know that tobacco and alcohol can damage cells in the lining of the oral cavity and oropharynx, and that cells in this layer must grow more rapidly to repair this damage. Many of the chemicals found in tobacco cause damage to DNA, which contains the cell's instructions for repair and growth.
- Scientists are not sure whether alcohol directly damages DNA, but they have shown that alcohol increases penetration of many DNA-damaging chemicals into cells

How to Detect and Diagnose Oral Cancer

- Many cancers of the oral cavity and oropharynx can be found early, during routine screening examinations by a doctor or dentist, or by self-examination.

Symptoms

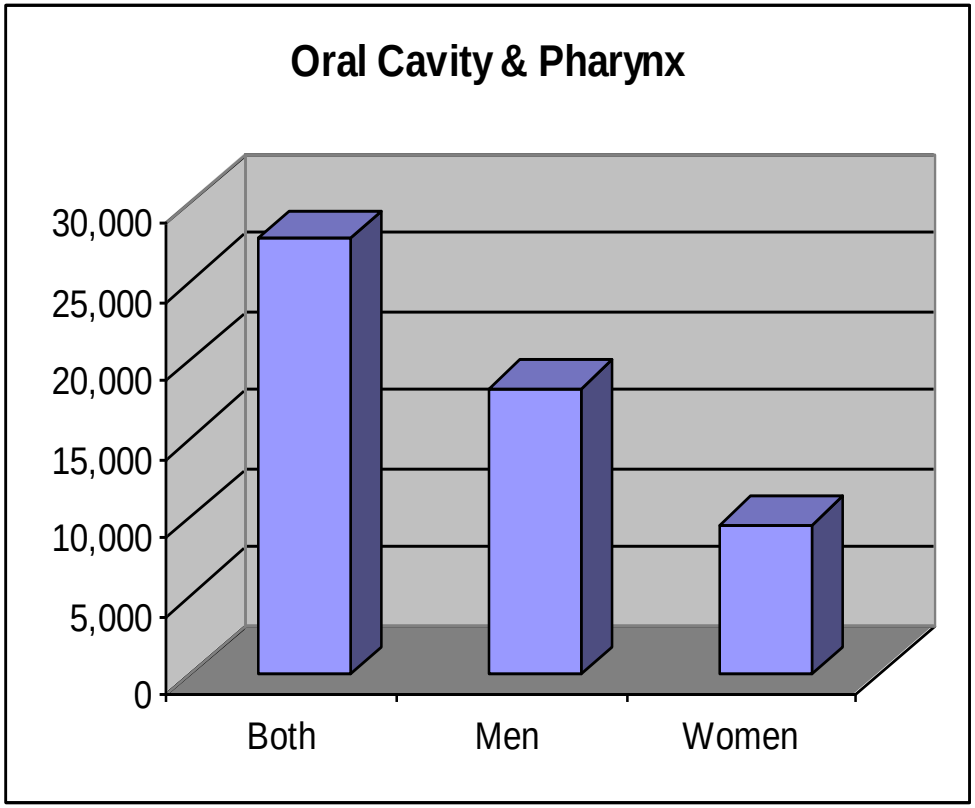
- a sore in the mouth that does not heal (most common symptom)
- pain in the mouth that doesn't go away (also very common)
- a persistent lump or thickening in the cheek
- a persistent white or red patch on the gums, tongue, tonsil, or lining of the mouth
- a sore throat or a feeling that something is caught in the throat that doesn't go away

More Symptoms

- difficulty chewing or swallowing
- difficulty moving the jaw or tongue
- swelling of the jaw that causes dentures to fit poorly or become uncomfortable
- loosening of the teeth or pain around the teeth or jaw
- voice changes
- a lump or mass in the neck
- weight loss
- persistent bad breath

Estimated New Cancer Cases

(2003)



Treatment & Survival

Radiation therapy and surgery are standard treatments. In advanced disease, chemotherapy may be useful as an adjunct to surgery and or radiation.

Survival for all stages combined, about 81% of oral cavity and pharynx cancer patients survive 1 year after diagnosis. The 5 year and 10 year survival rates are 56%and 41% respectively

Treatment

- Surgery
  - Most common with radiation
  - General health of mouth is analyzed and affects treatment
- Chemotherapy
  - Sometimes added to decrease possibility of distant micro-metastasis
  - Loss of taste and smell
- Removal of portion of mandible
  - Severe cases
  - Facial reconstruction is needed
  - Later assistance needed with speech and chewing

What happens after Treatment?

- Speech and Swallowing Therapy
- Follow-up tests
- Chemoprevention
- Watch for new symptoms
- General health considerations

What's new in oral cavity and oropharyngeal cancer research and treatment?

- **DNA changes:**

  One of the changes often found in DNA of oral cancer cells is a mutation of the p53 gene. Recent studies suggest that tests to detect these p53 gene alterations may allow very early detection of oral and oropharyngeal tumors. These tests may also be used to better define surgical margins

What’s New…

- **Tumor growth factors:**

  Oral and oropharyngeal cancers with too many EGF receptors tend to be especially aggressive. New drugs that specifically recognize cells with too many EGF receptors are now being tested in clinical trials. These drugs work by preventing EGF from promoting reproduction of cancer cells, and may also help the patient's immune system recognize and attack the cancer

What's New…

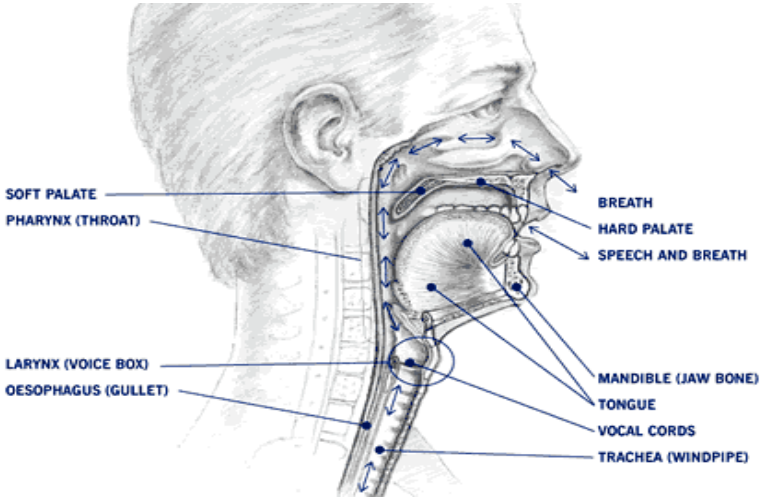
- **New chemotherapy**
- **New radiotherapy methods**
- **Vaccines:** Some oral and oropharyngeal cancers contain DNA from human papillomaviruses, vaccines against these viruses are being studied as a treatment for these cancers .
- **Gene therapy:** Another type of gene therapy adds new genes to the cancer cells to make them more susceptible to being killed by certain drugs

Throat Cancer

What is Throat Cancer and what does it encompass?

-AKA Cancer of the Pharynx.
*-Carcinoma (Mostly squamous cell). Cancer of the pharynx region usually stays local in the lining of the epithelial tissue. Can also be a Sarcoma (**Type of cancer that invades supportive tissue such as bone, cartilage, nerves, muscle, and fat).***
-Abnormal, cancerous cell growth within the lining tissue of the pharynx or throat or neck region.

- Cancer of the Pharynx
- Cancer of the Larynx
- Epiglottis
- Upper Oesophagus



History

- ***Head and neck cancer. (linked to oral, tongue, salivary gland, thyroid, pharyngeal, and epiglottal, and oesophageal cancers)***
- ***3% U.S. cancers, throat cancer less***
- ***More common amongst men, especially 50+***
- ***Between 30-38,000 new cases of head and neck cancer per year in U.S.***

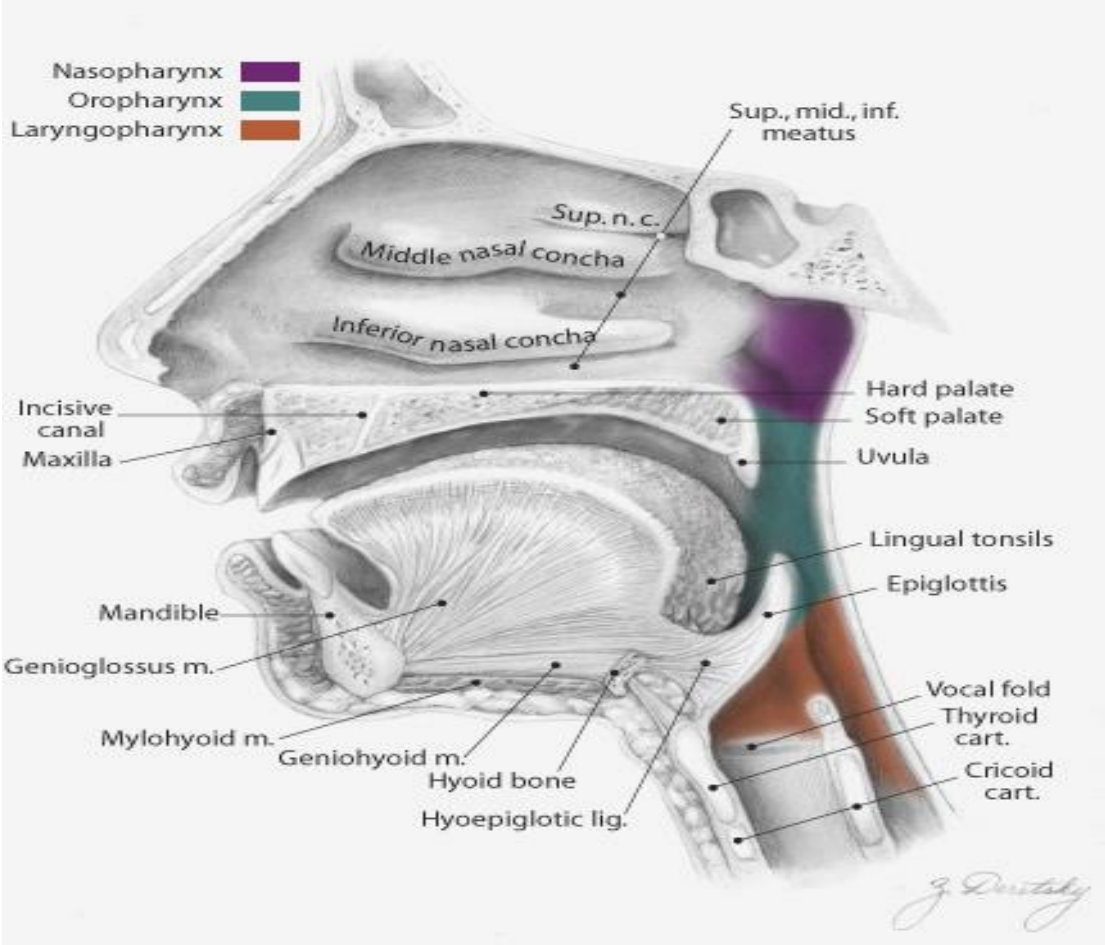
Risk Factors

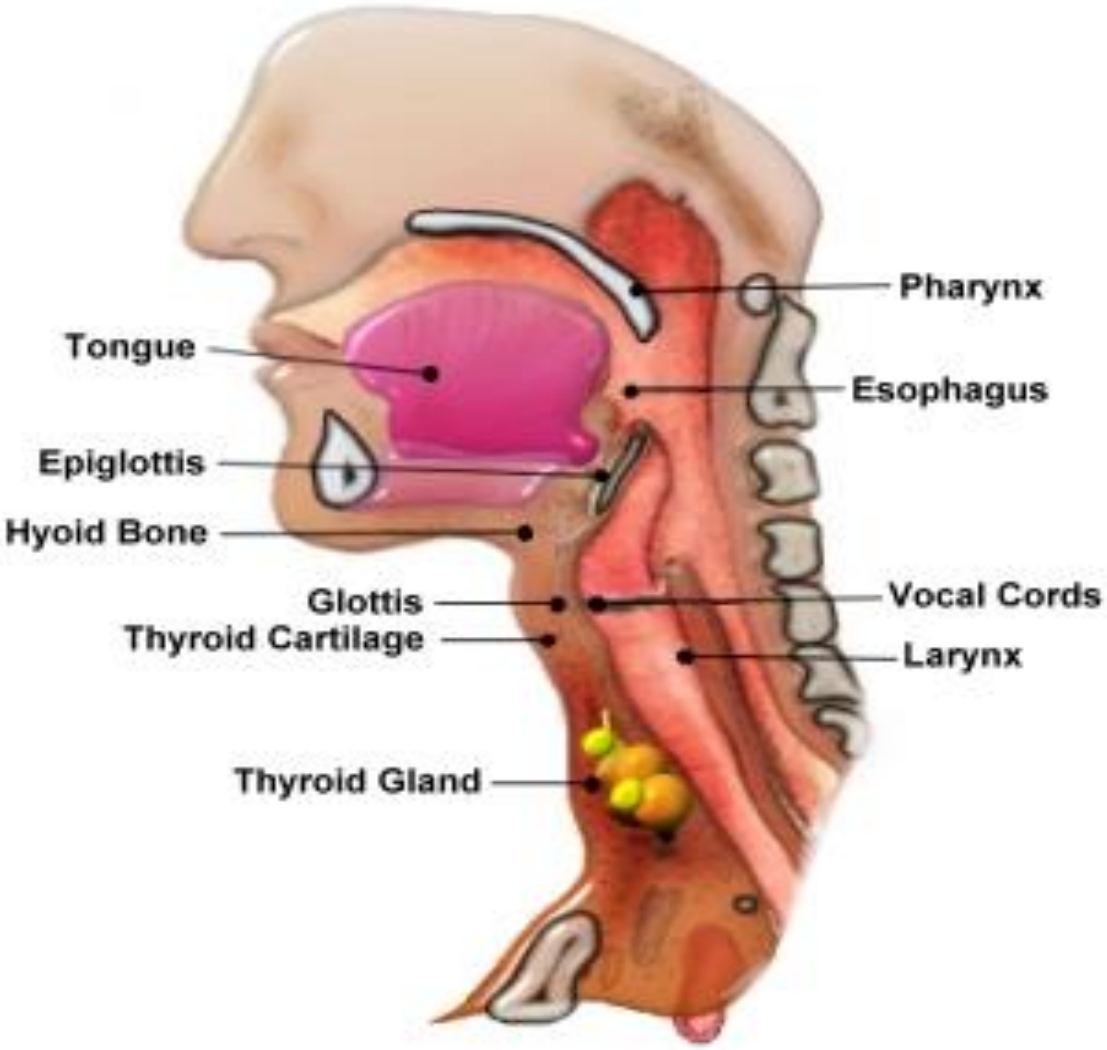
- Tobacco Products
- Alcohol Consumption
- Overexposure to harmful chemicals
  (Carbon Monoxide or second-hand smoke)
- Poor oral hygiene
- Epstein-Barr virus
- SES status

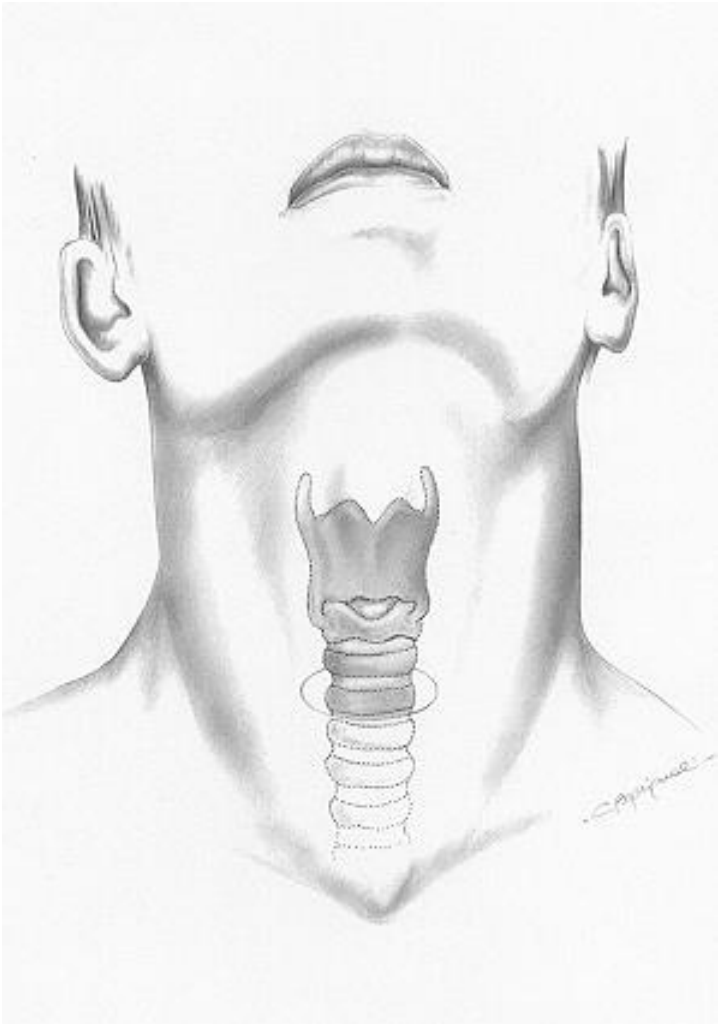
- Any type of harmful foreign substance to the body can act as a carcinogen & lead to throat cancer

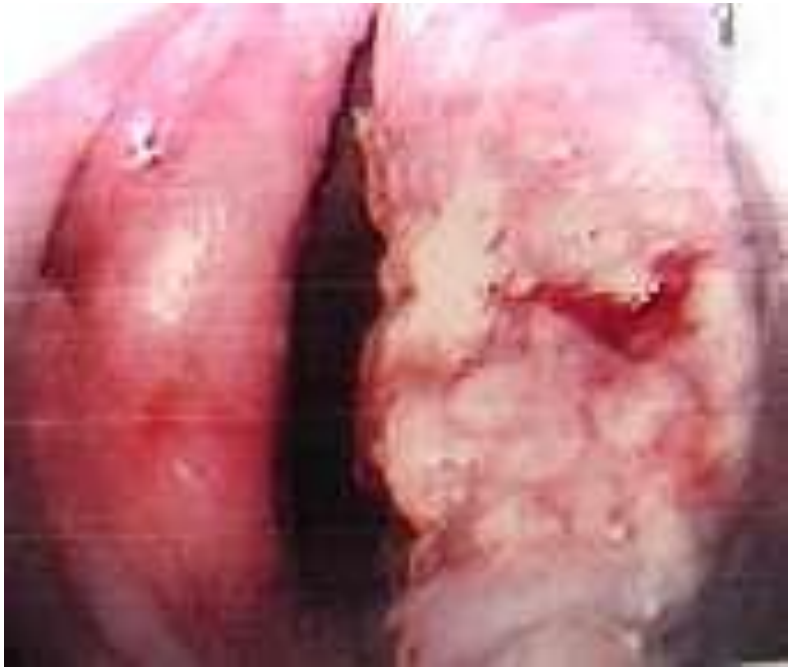
Identifying Throat Cancer (signs & symptoms)

- Adam's apple appears to be abnormally large or swelled up
- A small lump beginning to grow anywhere on or around the neck. This is beginning steps of a tumor taking form
- Having laryngitis for an extended period of time (consider seeing a doctor if laryngitis continues for longer that 2 weeks)
- Prolonged soreness in the neck
- Hoarseness that persists 3+ weeks
- Troubled breathing and painful swallowing of food
- Heat flashes


Nasopharynx
Oropharynx
Laryngopharynx
Sup., mid., inf. meatus
Sup. n. c.
Middle nasal concha
Inferior nasal concha
Hard palate
Soft palate
Uvula
Incisive canal
Maxilla
Lingual tonsils
Epiglottis
Mandible
Genioglossus m.
Vocal fold
Thyroid cart.
Cricoid cart.
Mylohyoid m.
Geniohyoid m.
Hyoid bone
Hyoepiglotic lig.

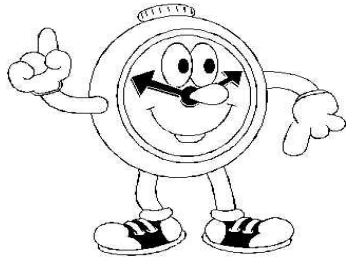

Pharynx
Tongue
Esophagus
Epiglottis
Hyoid Bone
Glottis
Vocal Cords
Thyroid Cartilage
Larynx
Thyroid Gland





Effects

- *Early detection vs. not so early detection*



The neck is a very critical part of the body!!

What's in your neck?

-Voice box
-Epiglottis
-Oesophagus
-Lymph nodes
-Carotid artery
-Jugular vein
-Spinal cord
-Windpipe
-Blood vessels galore
-Thyroid gland which produces essential hormones thyroxine, triiodothyronine, and calcitonin (assist w/ growth and metabolism)

Just like any other cancer, Throat Cancer starts out small but can invade any of these listed above and can spread quickly

Doctor Diagnosis

- Requires immediate treatment
- Physical examination looking for listed symptoms as well as enlarged lymph nodes

•*Staging process begins. Throat cancer is diagnosed in 4 stages*

Stage 1 and 2: Mild, usually curable. Radiation therapy followed by surgery if necessary

Stage 3: chances the cancer has spread are high. Radiotherapy alongside chemotherapy. Surgery usually precedes radiotherapy, but not always.

Stage 4: Can be treated but not likely curable. Same treatment as Stage 3. Microsurgery is usually performed followed by a clinical trial of hyperthermia and radiation therapy.

Treatments and related Side effects

Radiation Therapy: (2 Options)

High energy X-Rays to kill cells (External Beam Radiation Therapy)

Placing radioisotopes on the infected area killing all cells (Internal Radiation Therapy)

Side Effects: (Will vary patient to patient)

-fatigue, dry skin, discolored skin, itching, nausea and vomiting, loss of hair, and of course many good/ unaffected cells killed as well as cancerous ones.

Treatments and related side effects

Chemotherapy:

Drugs taken to kill the cancer (Via pills or injection)

Side Effects:

Very similar to those of radiation treatment

Hyperthermia Treatment:

Used to raise the temperature of the tissue and/or blood to hopefully receive therapeutic benefits

Side Effects:

blood coagulation, kidney problems, sometimes severe pain and occasional burn marks

Treatments and related side effects

Surgery:

Surgery is done on the patient to remove the malignant tumor.

Depending on the cancer, surgery is not always used. Not always 100% accurate.

Side Effects:

1. *Scarring on the patient. Physically and emotionally*
2. *Soreness*
3. *Bedridden*
4. *Unable to perform physical activities for various lengths of time*

Preventive measures

- *Avoid alcohol and tobacco products*
- *Diet*
- *Always be cautious of the air you are breathing and what might be in it. Don't overexpose yourself to it longer than necessary*
- *See a doctor upon slightest suspicion of abnormality or soreness*

MENINGITIS

Definition

- Meningitis is an inflammatory process of the leptomeninges and CSF

Classification

- 1. acute pyogenic (bacterial) meningitis
- 2.acute aseptic (viral) meningitis
- 3.acute focal suppurative infection (brain abscess,subdural and extradural empyema)
- 4.chronic bacterial infection (tuberculosis).

Acute pyogenic bacterial meningitis

- Most important
- Can be fatal if untreated
- Organisms:

  E.coli ---------- neonates

  Streptococci B ---------- neonantes

  H. influenzae----------adolescents

  Neisseria meningitidis---------- young adults

  Streptococcus pneumonia--------- elderly

Clinical signs

- Signs of infection (fever,malaise,rigor….)
- Signs of meningeal irritation:
  1.headache
  2.neck stiffness
  3.photophobia
  4.irritability

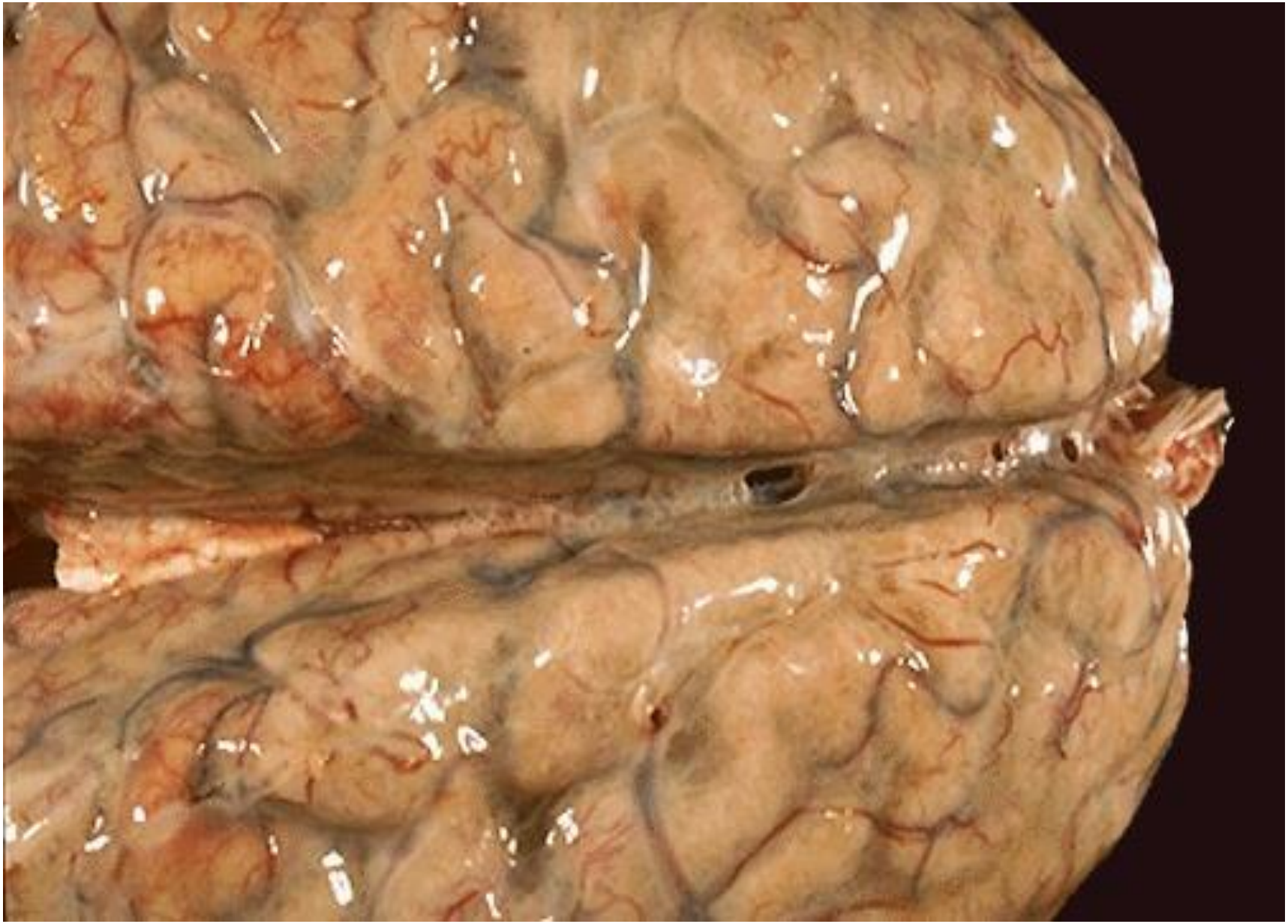
C.S.F by lumbar puncture shows :

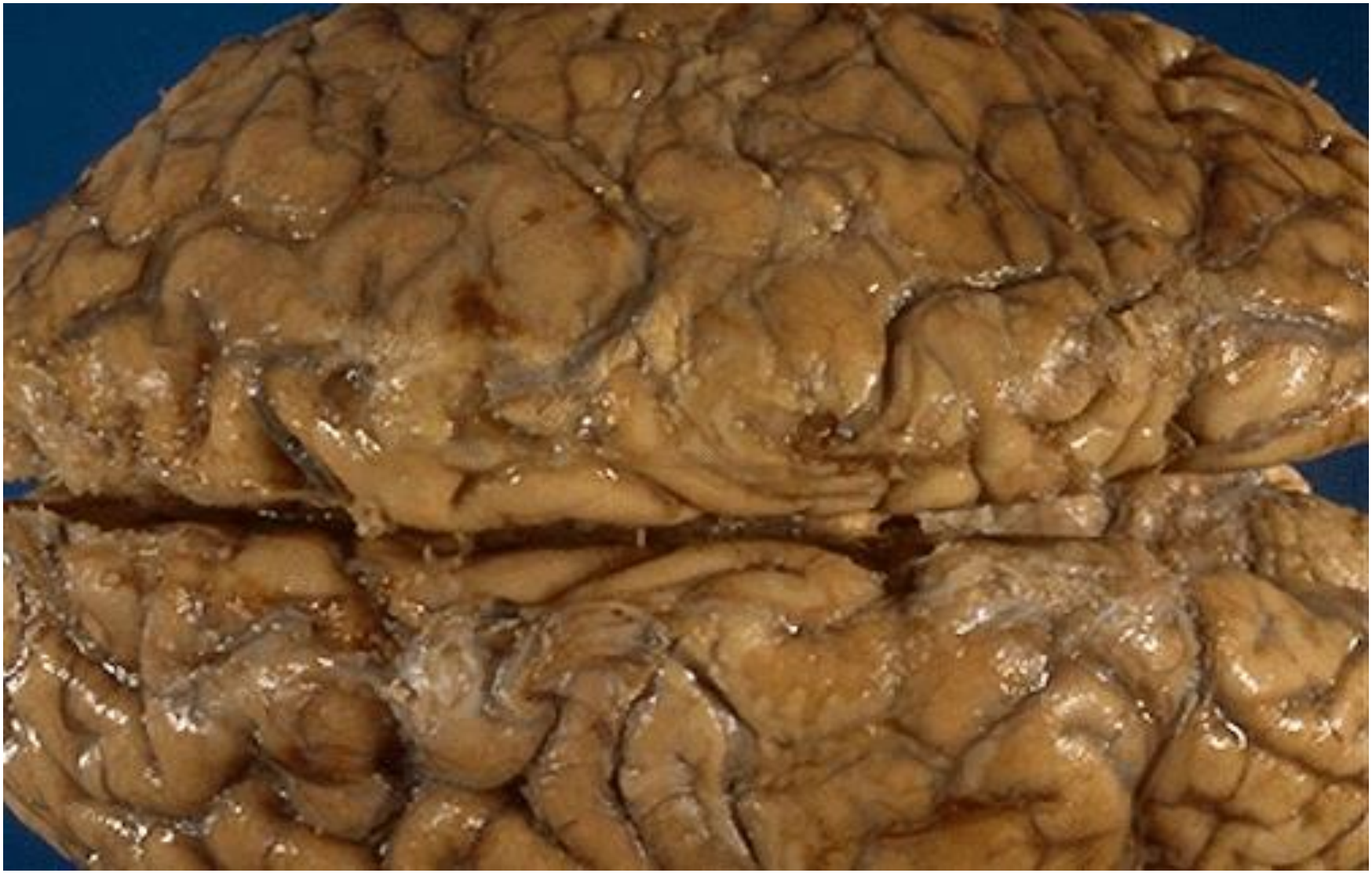
a.cloudy purulent csf
b.abundant neutrophils > 90,000/mm3
c.high protein level and
d.reduced glucose level.

Morphology

- Grossly , pyogenic meningitis shows a thick layer of suppurative exudate covers the leptomeninges over the surface of the brain.
- Exudate in basal surface--- H.INFLUENZAE
- Exudate in covexity surface--- P.MENINGT
- Microscopically :

  neutrophils in the subarachnoid space





Complications

- Antibiotic treatment------ full recovery
- Delayed or untreated cases--- can be fatal
- Healing by fibrosis cause obliteration of subarachenoid space--- HYDROCEPHALUS
- Brain abscess
- Septic shock and skin rashes, why ?

Skin rashes

- Is due to small skin bleed
- All parts of the body are affeced
- The rashes do not fade under pressure
- Pathogenesis:

a. Septicemia

b. wide spread endothelial damage

c. activation of coagulation

d. thrombosis and platelets aggregation

e. reduction of platelets (cosumption)

f. BLEEDING 1.skin rashes

2.adrenal hemorrhage

Arenal hemorrhage is called Waterhouse-Friderichsen Syndrome.It cause acute adrenal insufficiency and is uaually fatal

Acute Aseptic (Viral) Meningitis

- Can follow any viral infection
- Less danger
- CSF shows :

  1.lymphocytes

  2. mild increase in protein

  3. normal glucose level

Viral meningitis is usually self-limiting and treated symptomatically.

Brain abscess

- Causes :
  1. complication of bacterial meningitis
  2. bacterial endocarditis
  3. pulmonary sepsis : peumonia……etc
  4. other sepsis

Brain abscess cause a space occupying lesion in the brain