

Cellular adaptations

HYPERTROPHY

HYPERPLASIA

ATROPHY

METAPLASIA

"Zindagi Kya Hai, Anasir Mein Zahoore Tarteeb;

Maut Kya Hai !! Inhi Ajza Ka Pareshan Hona





- Process of increase in size resulting from the synthesis of specific tissue components.
- Populations, individuals, organs, cells, or even subcellular organelles such as mitochondria. :

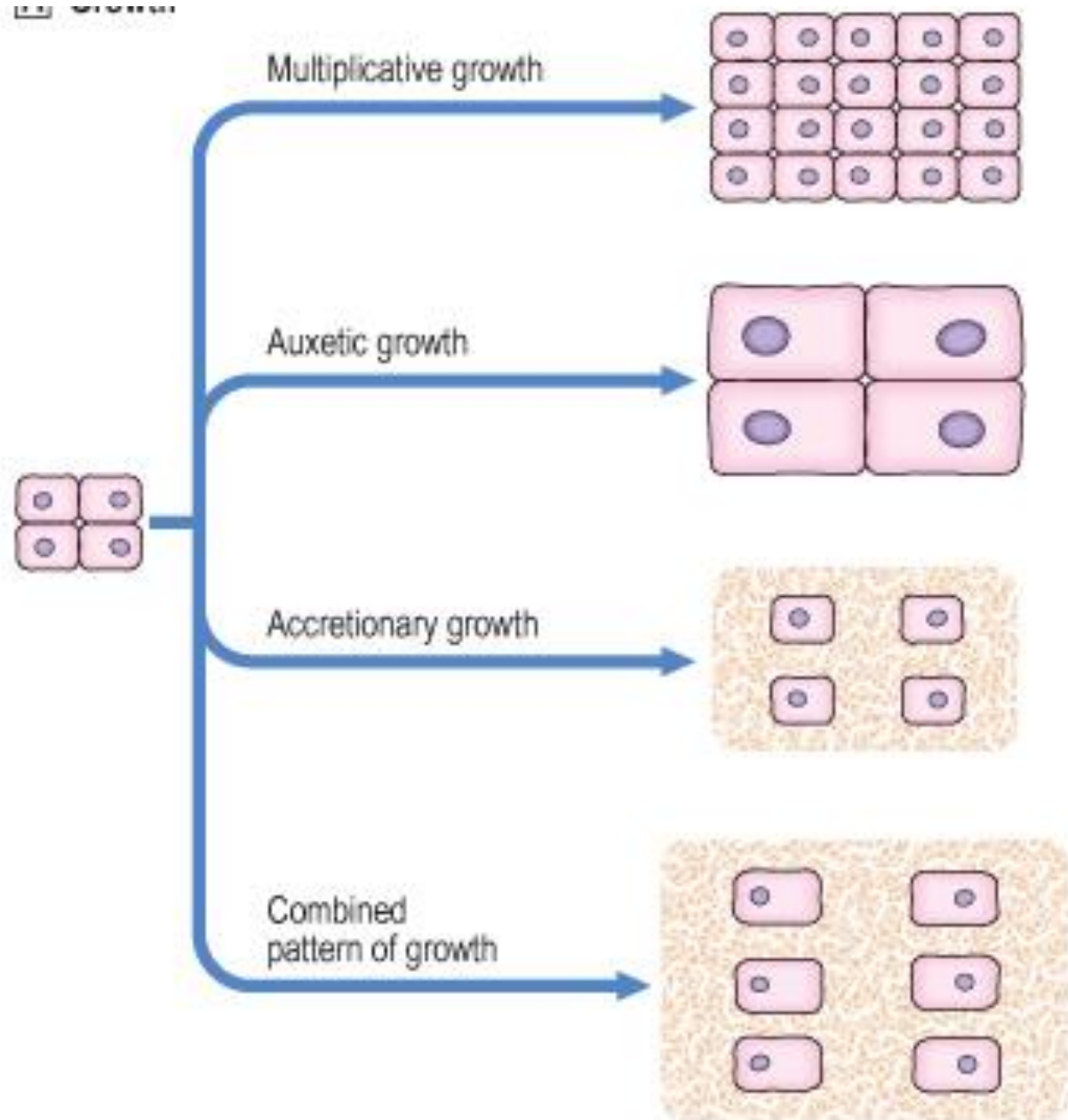
Types of growth in a tissue

Multiplicative

- Involving an increase in numbers of cells (or nuclei and associated cytoplasm in syncytia) by mitotic cell divisions)
- All tissues during embryogenesis

- *Auxetic*, increased size of individual cells (growing skeletal muscle)
- *Accretionary*, an increase in **intercellular tissue** components, as in bone and cartilage.

- *Combined patterns* of multiplicative, auxetic and accretionary growth as seen in **embryological development**

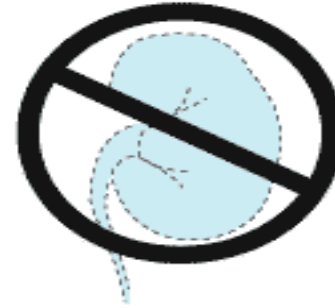


Normal Growth & Maturation

- Cells continue to **grow, divide, and differentiate** throughout life
- Labile stem cells
- Skin, as superficial keratinized cells
- Psoriasis,

PRIMITIVE EMBRYONIC CELLS
(ectoderm, mesoderm, endoderm)

AGENESIS
Failure of
development



ORGAN ANLAGE

**Cell
division**

Normal number of cell
divisions during development

Abnormal number of cell
divisions during development

**Cell
differentiation**

Normal
differentiation

Abnormal
differentiation

Normal
differentiation

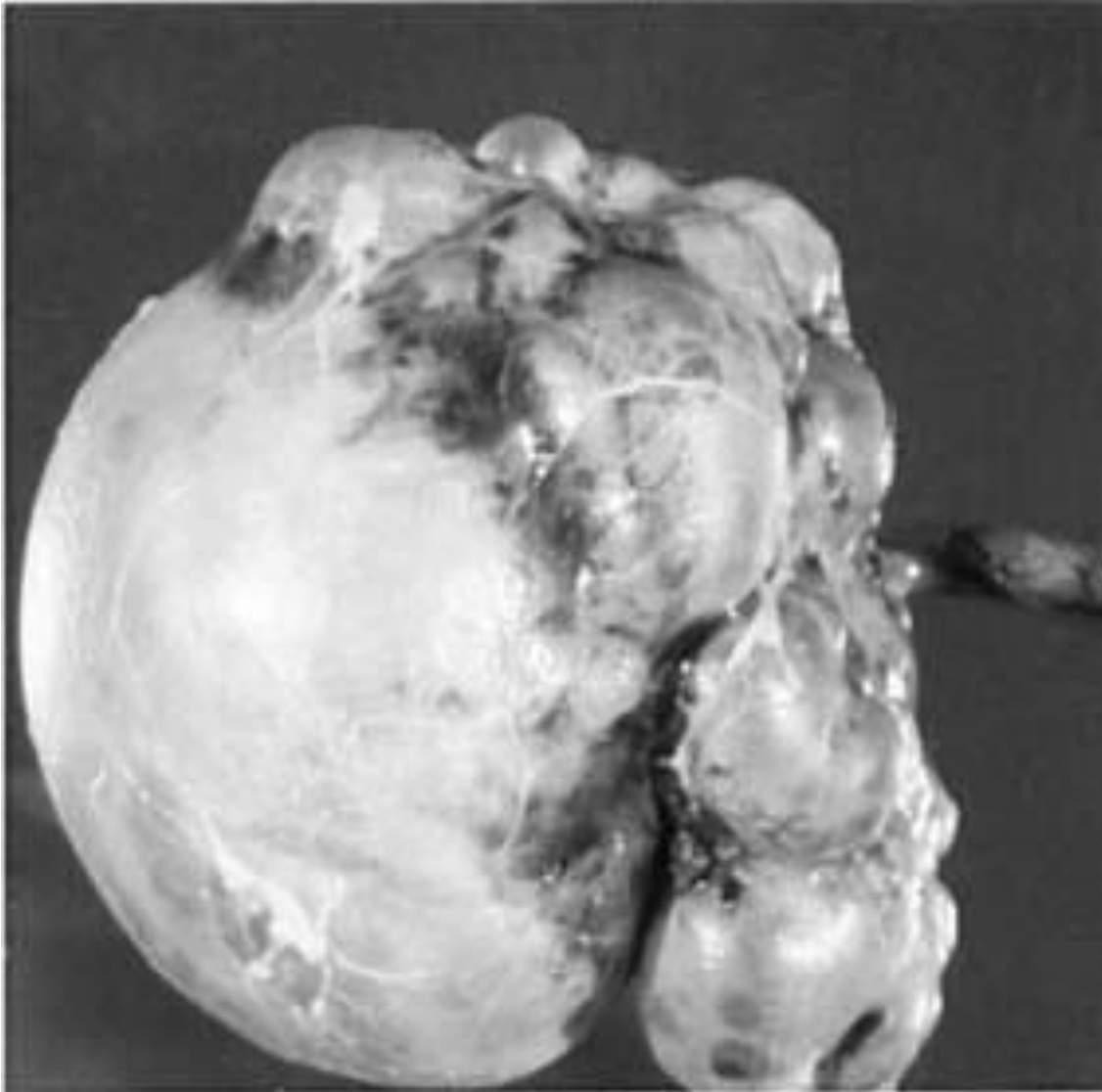
**Normal organ
development**

Dysgenesis

Hypoplasia



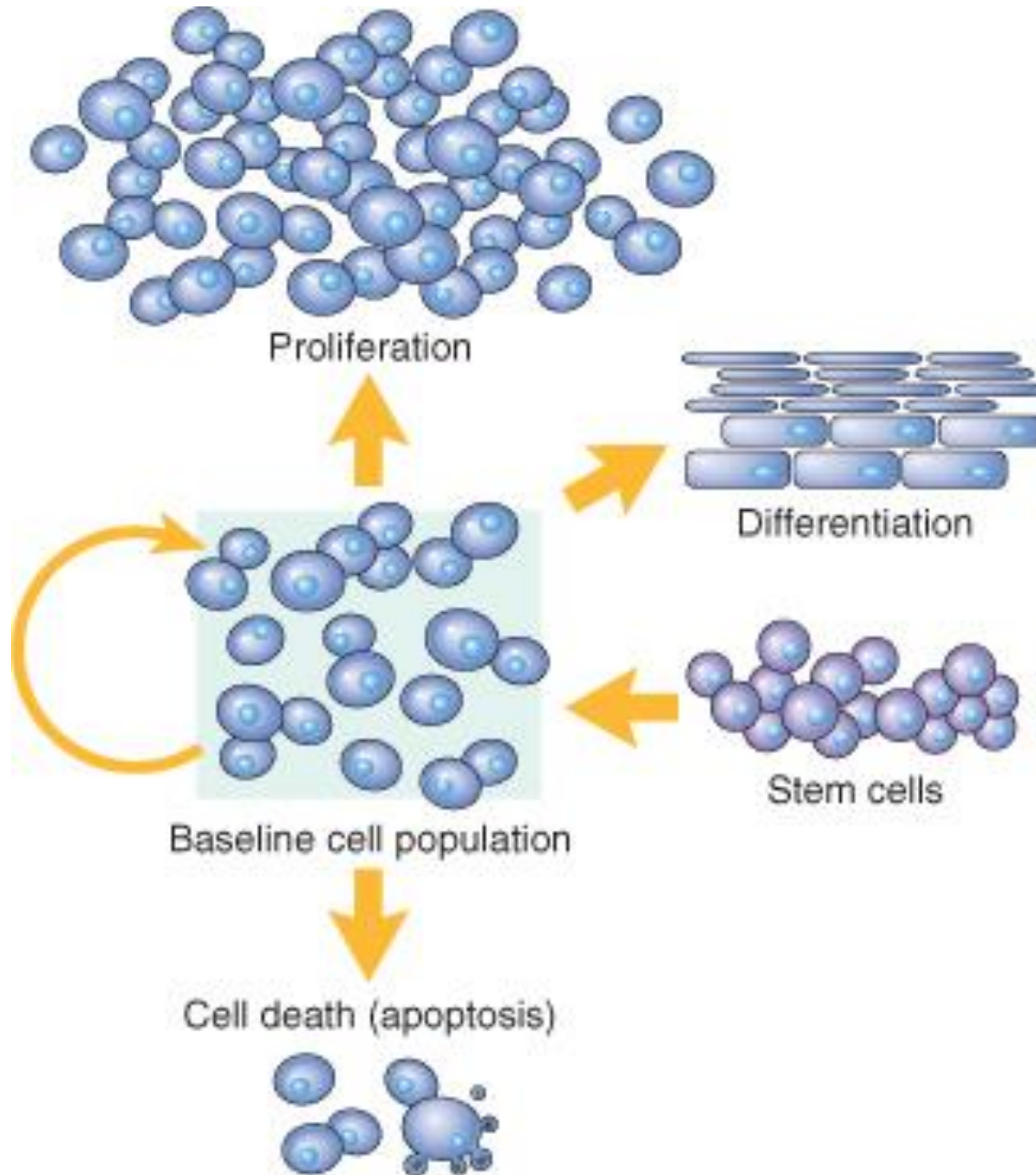
Renal dysgenesis



Control of Growth

- Net balance cell proliferation \Leftrightarrow cell differentiation, leading to cell death
- The rate of proliferation
- Many of the **cellular proto-oncogenes**

Mechanisms regulating cell populations



Growth Factors: Major Categories and Families

Major categories

Endocrine

The systemic hormones, eg, pituitary hormones, steroid hormones, thyroid hormones, insulin

Paracrine

Locally acting hormones, eg, the neuroendocrine system of the gastrointestinal tract

Autocrine

Same cell produces growth factor and corresponding receptor

Families

Epidermal growth factor (EGF)

Platelet-derived growth factor (PDGF)

Transforming growth factor (TGF)

Interleukin

Insulin-like growth factors

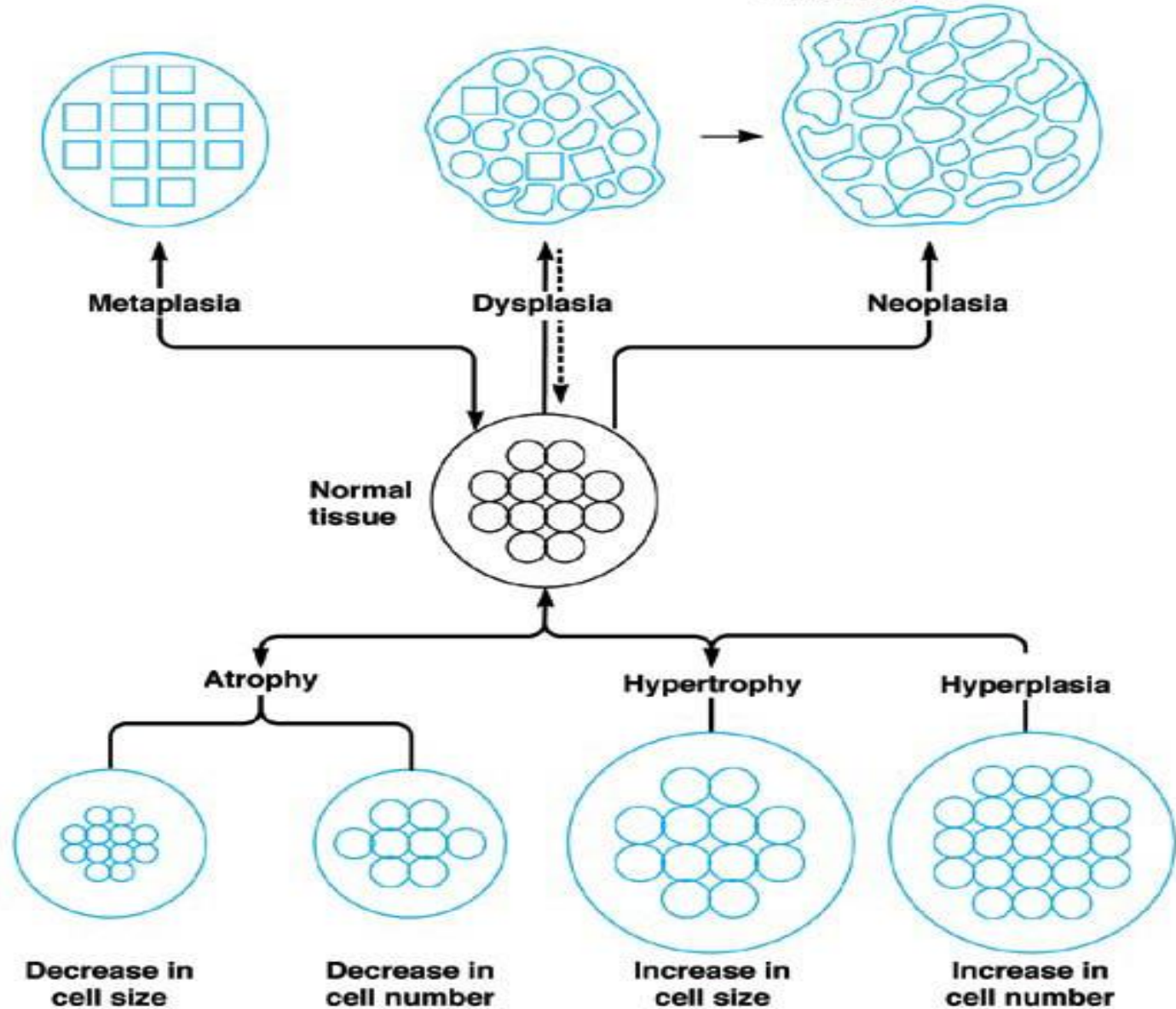
Fibroblast growth factors

The interferons

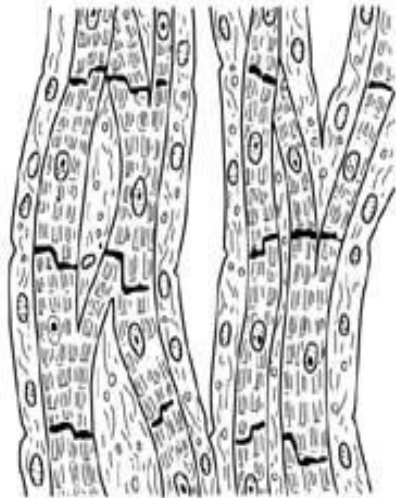
- Abnormal differentiation and maturation
- Replacement of mature cells of one type with cells of another type
- Regular organization of tissue maintained
- Reversible

- Abnormal differentiation and maturation
- Partial loss of control and organization
- Slight increase in cell number
- Cytologic abnormalities
- **Partially reversible**

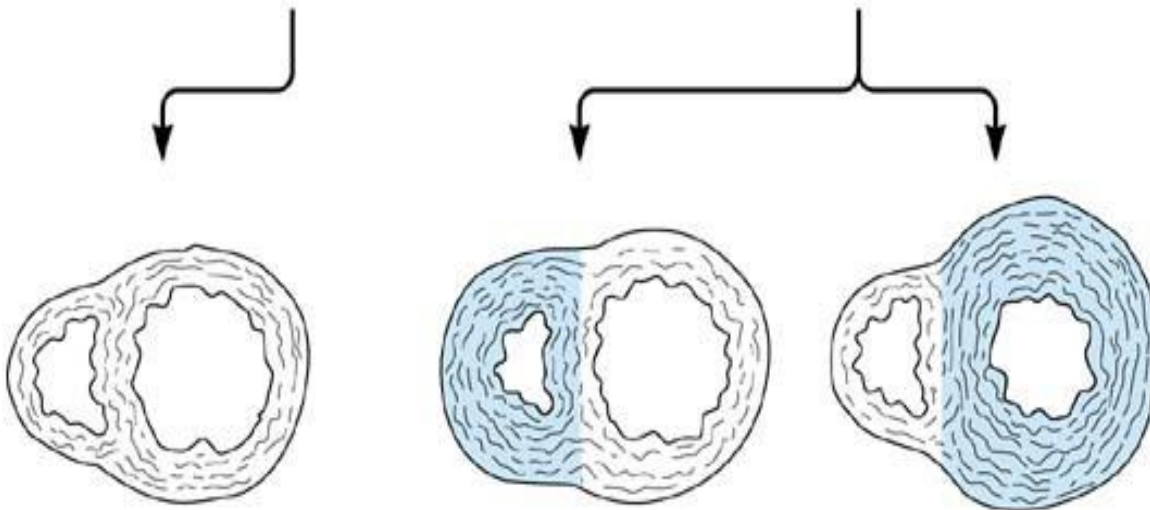
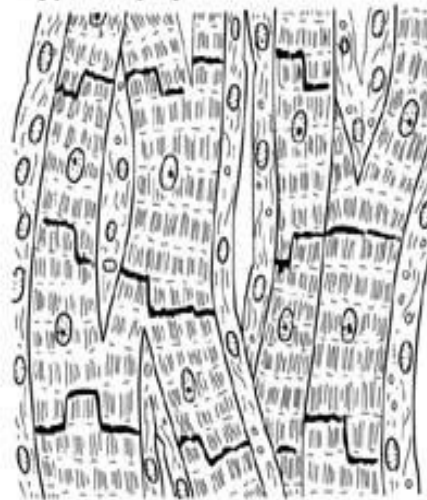
- Abnormal differentiation and maturation
- Marked increase in cell number
- Complete loss of control
- Variable loss of organization
- Cytologic abnormalities
- **Irreversible**



Normal



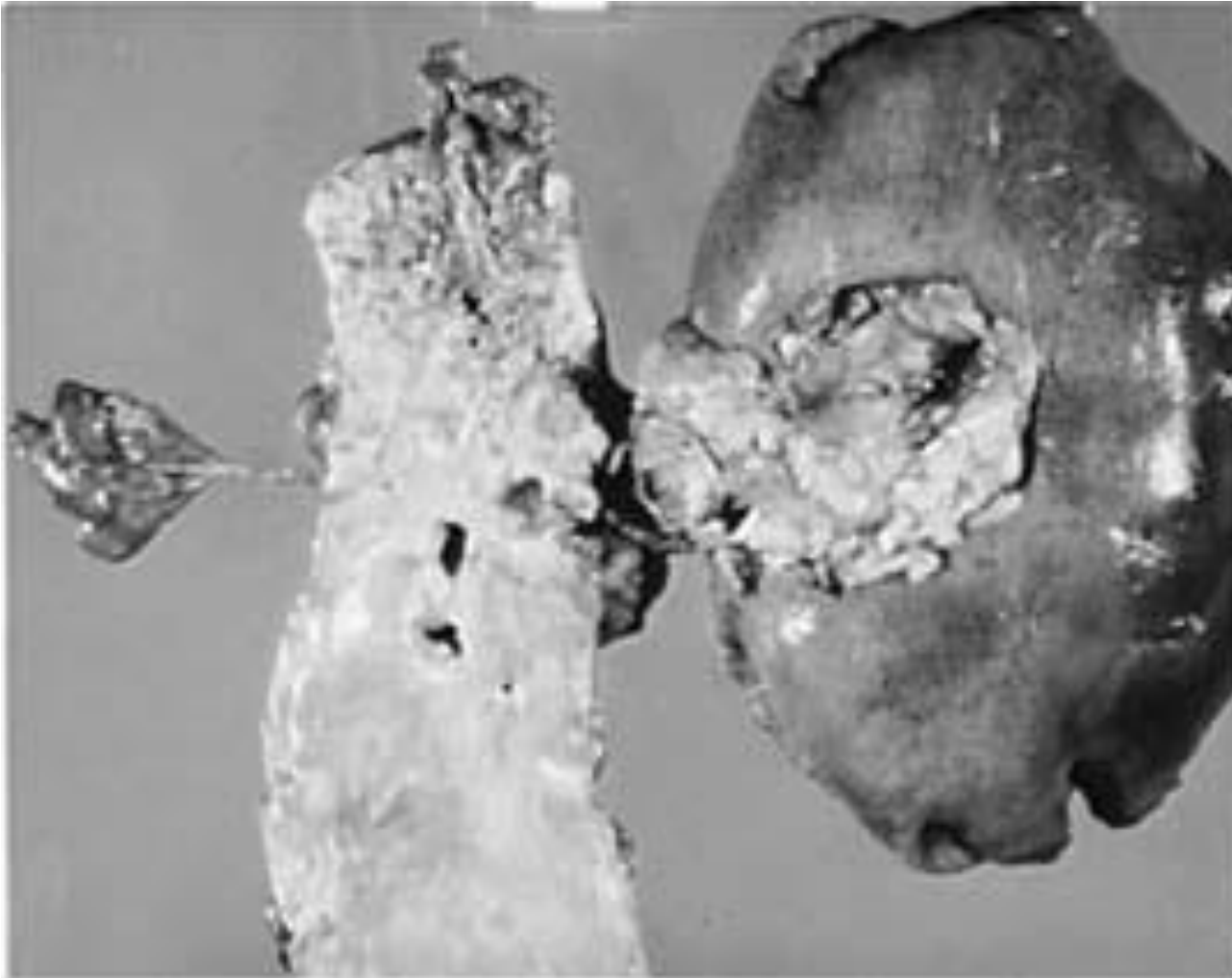
Hypertrophy

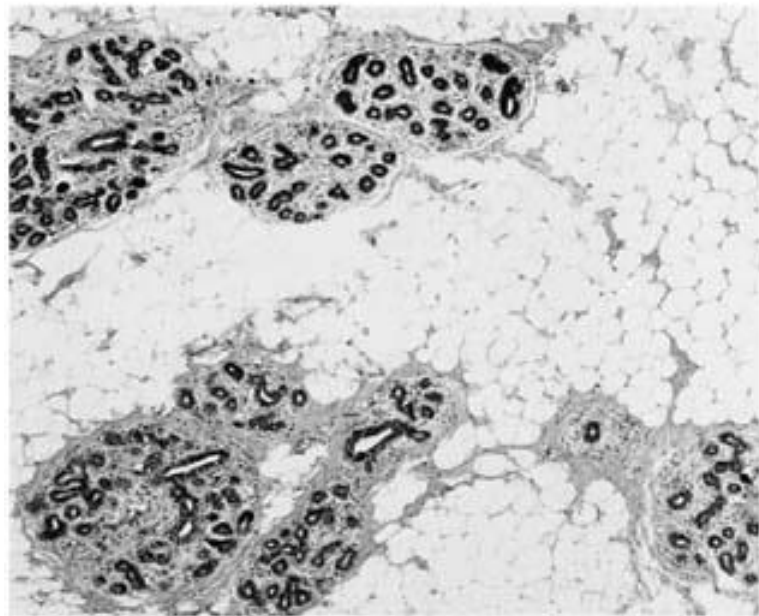
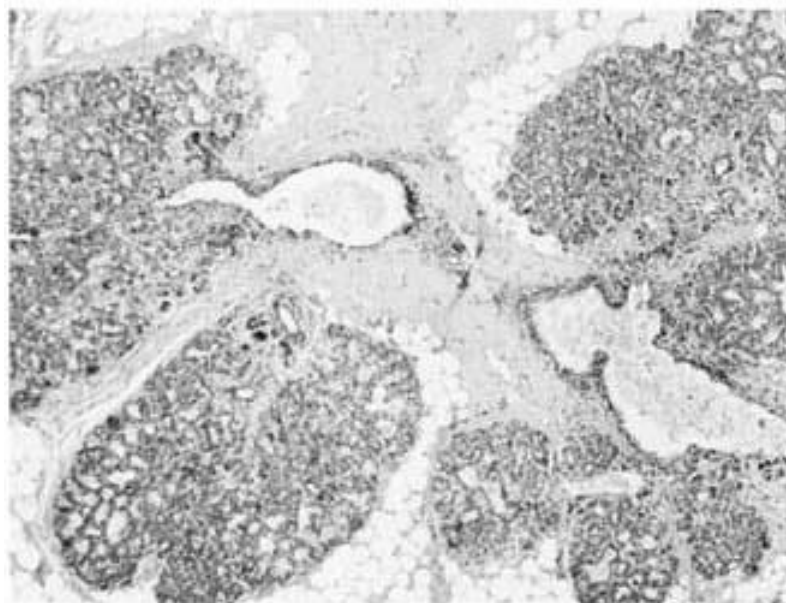


Hypertrophy and Hyperplasia of Organs.

Tissue	Cause of Increased Demand
Skeletal muscle hypertrophy	Physical activity, weight lifting
Cardiac muscle hypertrophy	Increased pressure load (high blood pressure, valve stenosis) or increased volume load (valve incompetence causing regurgitation of blood)
Smooth muscle (wall of intestine, urinary bladder) hypertrophy	Obstructive lesions
Uterine myometrial hypertrophy	Pregnancy (hormone-induced)
Bone marrow hyperplasia	
Erythroid hyperplasia	Increased destruction of erythrocytes (hemolytic process); prolonged hypoxia (living at high altitudes).
Breast hyperplasia	Pregnancy and lactation (hormone-induced)

Developmental hypoplasia of one kidney associated with marked compensatory hyperplasia of the other



A**B****C**

- Cells respond to increased demand and external stimulation by *hyperplasia* or *hypertrophy*,
- Respond to reduced supply of nutrients and growth factors by *atrophy*.
- Cells change from one type to another, a process called *metaplasia*.
- Some adaptations are induced by **direct stimulation of cells** by factors produced by the responding cells themselves or by other cells in the environment.
- Others are due to activation of various **cell surface receptors and downstream signaling pathways**.

Atrophy

- Decrease in the size of a tissue or organ, resulting from a decrease either in the size of individual cells or in the number of cells composing the tissue.
- Note that atrophy, which is a decrease in size of a normally formed organ, is distinct from agenesis, aplasia, and hypoplasia, which are abnormalities of organ development.

Causes of Atrophy

- Atrophy of Disuse
- Denervation Atrophy
- Atrophy Due to Loss of Trophic Hormones
- Atrophy Due to Lack of Nutrients
- Senile Atrophy
- Pressure Atrophy

Atrophy Due to Loss of Trophic Hormones

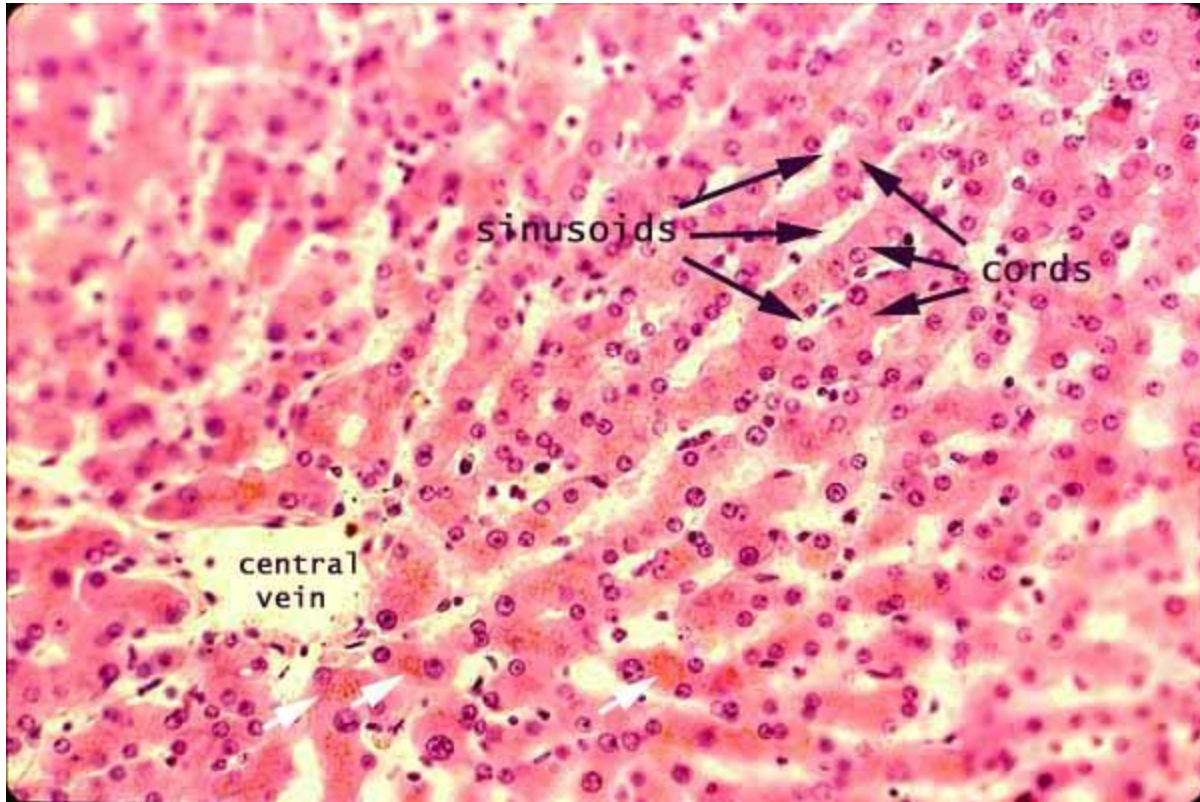
- **Estrogen** secretion by the ovary decreases at menopause, there is **physiologic atrophy of the endometrium**, vaginal epithelium, and breast.
- **Pituitary disease** associated with decreased secretion of **pituitary trophic hormones** results in **atrophy of the thyroid**, adrenals, and gonads.

- High-dose adrenal corticosteroid therapy, which is sometimes used for immunosuppression, causes **atrophy of the adrenal glands** because it suppresses pituitary corticotropin secretion.
- Such patients soon lose the ability to secrete cortisol and become dependent on exogenous steroids. **Withdrawal** of steroid therapy in such patients must **be gradual** enough to permit regeneration of the atrophied adrenal.

Reduction in the amount of cytoplasm and the number of cytoplasmic organelles

- Diminished metabolism.
- Degenerating organelles are taken up in lysosomal vacuoles for enzymatic degradation (**autophagy**).
- Residual organelle membranes often accumulate in the cytoplasm as **brown lipofuscin pigment**.

Liver, Kidney, Heart, Muscle, Nerve Cells



Hypertrophy and hyperplasia

- => Adaptation to increased demand.
- Controlled responses
- Demand is removed, the tissues revert toward normal.

Hypertrophy

- Increase in the size of a tissue due to increased size of individual cells
- Tissues made up of **permanent cells**, in which a demand for increased metabolic activity cannot be met through cell multiplication.

Hyperplasia

- Increase in the size of a tissue as a result of increased numbers of component cells
- Labile and stable cells.

- Not uncommonly, increased size of a tissue is due to a combination of hypertrophy and hyperplasia.

Causes of Hypertrophy & Hyperplasia

- **Hypertrophy** results from increased amounts of cytoplasm and cytoplasmic organelles
 - In secretory cells, the synthetic apparatus
 - In contractile cells such as muscle fibers, cytoplasmic myofibrils
- **Hyperplasia** results when cells of a tissue are stimulated to undergo mitotic division, thereby increasing the number of cells.

Pathologic Hypertrophy and Hyperplasia

- Absence of an appropriate stimulus of increased functional demand. **Myocardial hypertrophy**, if it occurs without recognizable cause
- Such hypertrophy is frequently associated with **abnormal cardiac function**, producing **cardiomyopathy**

Endometrial hyperplasia

- Increased estrogen stimulation
- Near menopause.
- Excessive trophic hormones => hyperplasia of the target organs
- Excessive secretion of ACTH => **bilateral adrenal hyperplasia.**
- Increased function. In the case of the adrenal gland,
- (Cushing's syndrome)

- **Thyroid hyperplasia (goiter; Graves' disease)** increased TSH stimulation of the thyroid or from the action of **autoantibodies** that are able to bind to TSH receptors in thyroid cell membranes

- **Hyperplasia of the prostate gland**

Abnormal Growth Principally Involving Differentiation: **Metaplasia**

- Varied potential for differentiation
- Germinative stem cells
- Differentiate in a manner that is abnormal for that location
- Following chronic physical or chemical irritation.

Squamous metaplasia

- —The most common type of epithelial metaplasia—
- Nonsquamous pseudostratified columnar/cuboidal epithelium => normal-appearing stratified squamous epithelium

Glandular metaplasia

- Esophagus, where the normal squamous epithelium => glandular, mucus-secreting epithelium
- Stomach and intestine,
- Intestinal metaplasia
- Gastric metaplasia

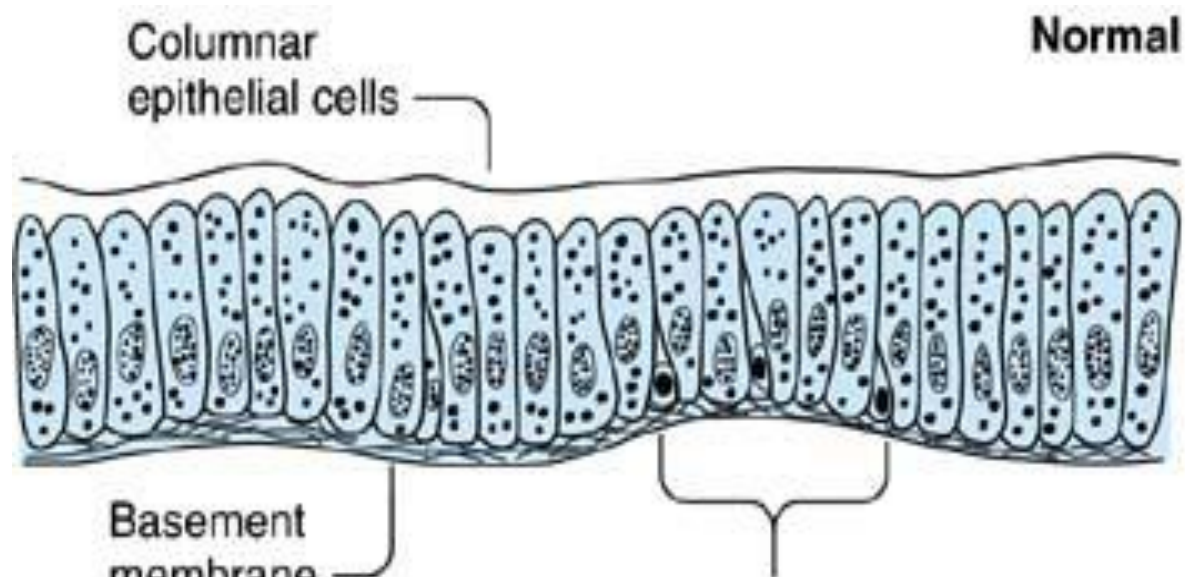
- Rarely occurs in **mesenchymal tissue** and is best exemplified by osseous metaplasia in scars and other fibroblastic proliferations.
- Potential for diverse differentiation of **mesenchymal stem cells**.

Functional deficits

- Loss of cilia and of mucus production in the bronchi may predispose to development of infection.
- Metaplastic tissue is structurally normal and itself carries no increased risk of development of cancer. However, **dysplastic changes** are often present as well , and cancer does occur in metaplastic epithelia under such circumstances.
- Squamous carcinoma develops in metaplastic squamous epithelium in the bronchus, and adenocarcinoma may arise in the esophagus from metaplastic glandular epithelium.

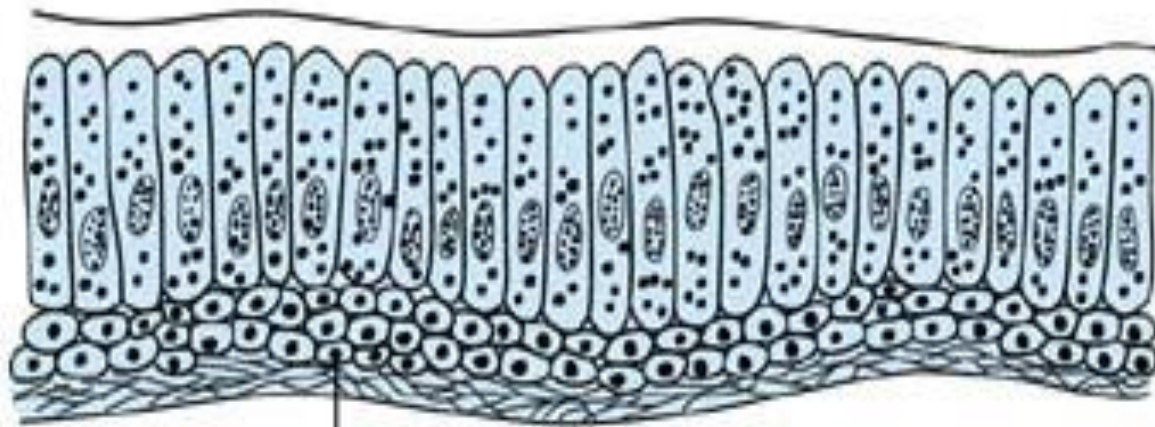
Abnormal Growth Principally Involving Differentiation: Metaplasia

Type of Metaplasia	Site	Causative Factors
Epithelial metaplasia		
Squamous metaplasia	Multiple sites Bronchus Endocervix Urinary bladder	Vitamin A deficiency Cigarette smoking, chronic inflammation Chronic inflammation Chronic inflammation, schistosomiasis
Intestinal metaplasia	Esophagus Stomach	Acid reflux Alkaline reflux, chronic inflammation
Gastric metaplasia	Esophagus Intestine	Acid reflux Unknown
Serous or mucinous metaplasia	Germinal epithelium of ovary	Trauma of multiple ovulation
Mesenchymal metaplasia		
Osseous metaplasia	Fibrous scars Areas of calcification	Unknown Unknown
Myeloid metaplasia	Spleen, liver	Unknown

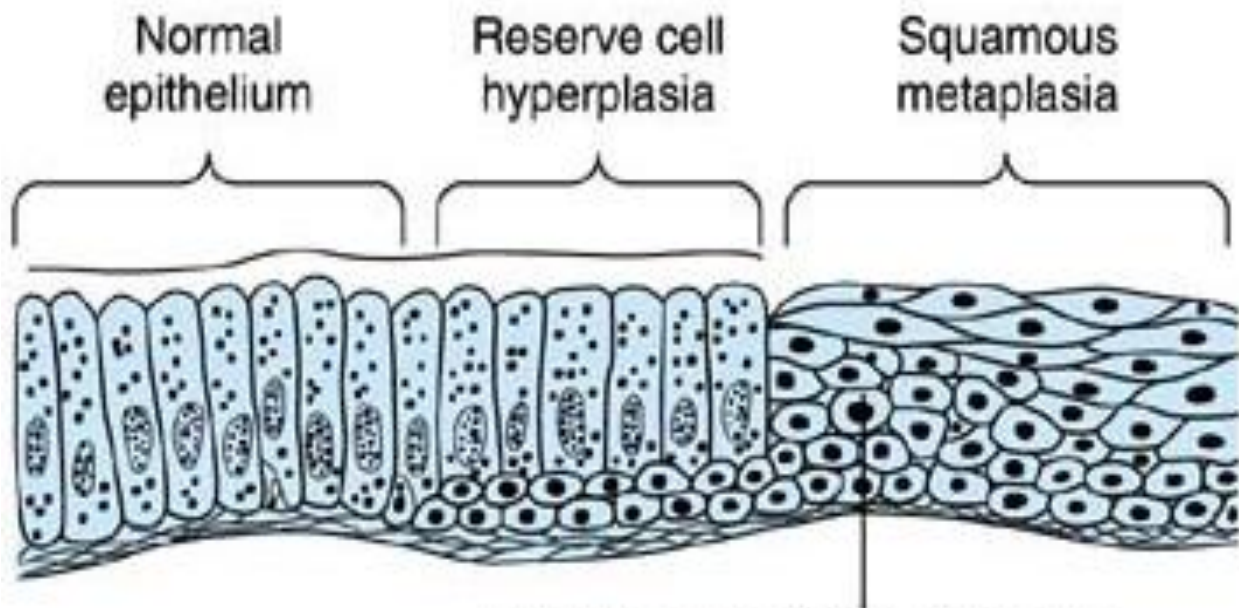


Reserve cells (stem cells of endocervical epithelium) normally divide and differentiate into glandular epithelial cells that replace exfoliated surface cells.

Hyperplasia

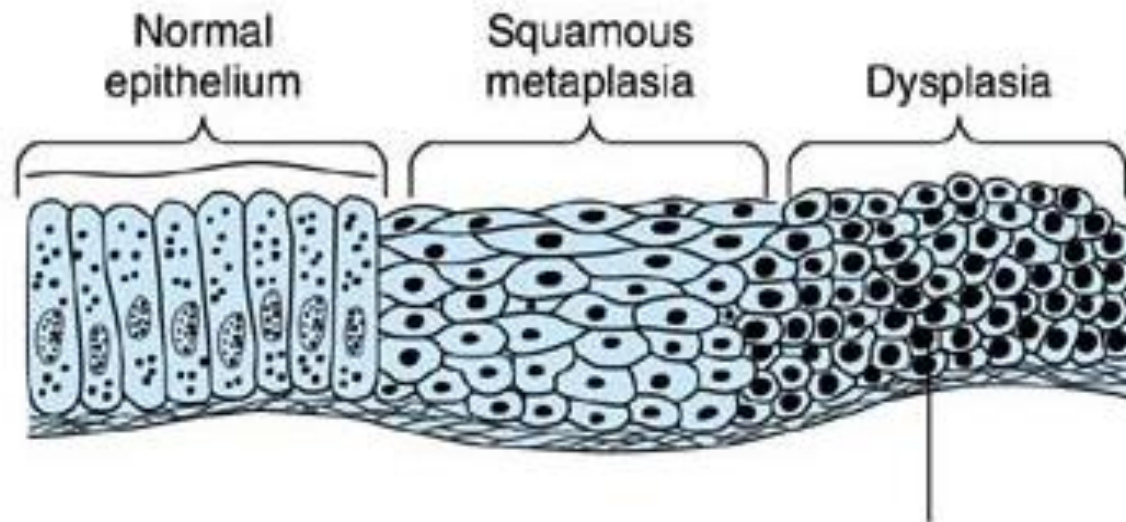


Hyperplastic reserve cells
differentiate into normal
glandular epithelial cells.



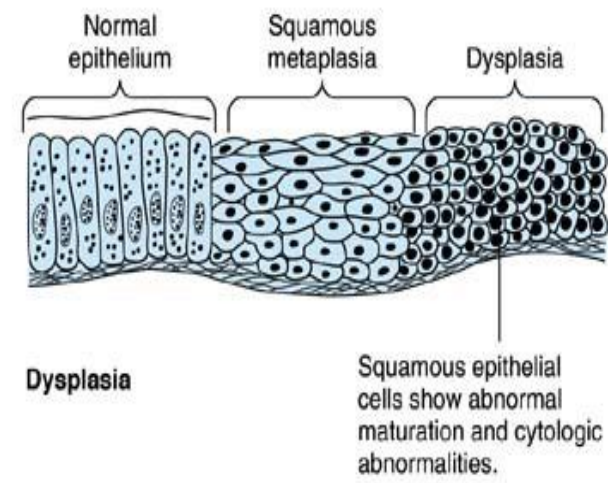
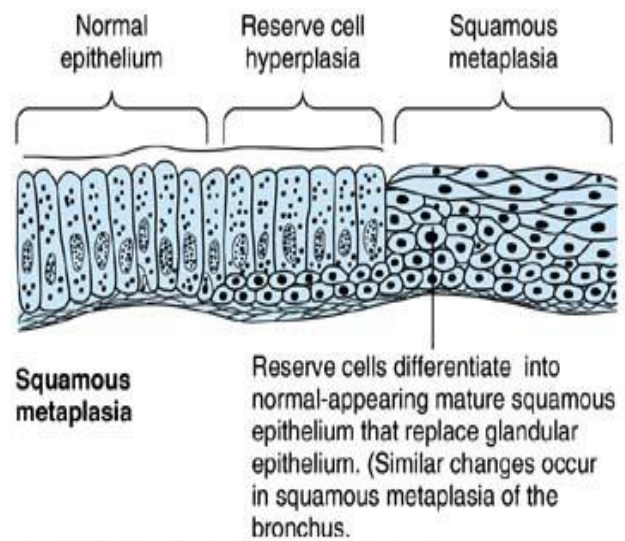
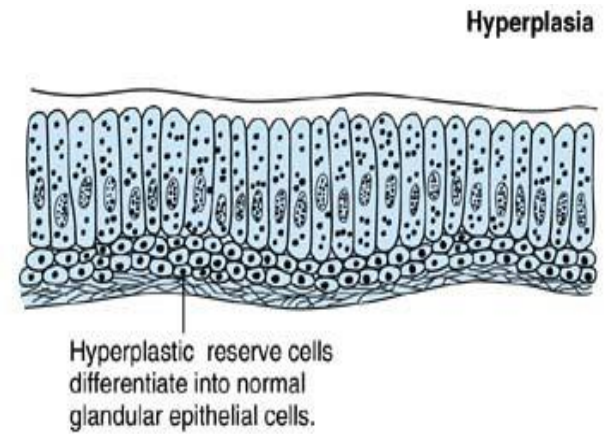
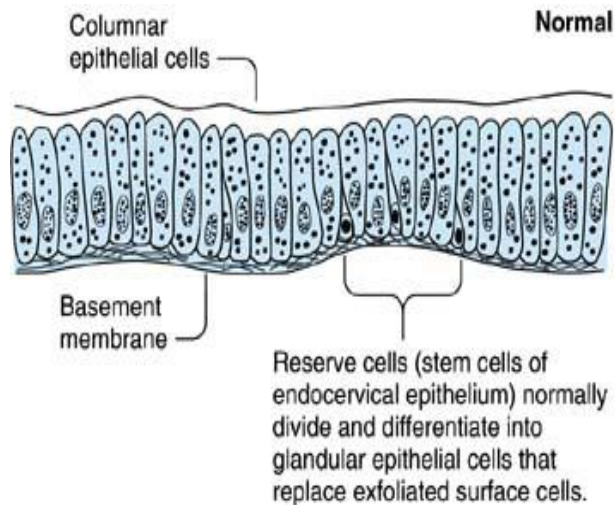
Squamous metaplasia

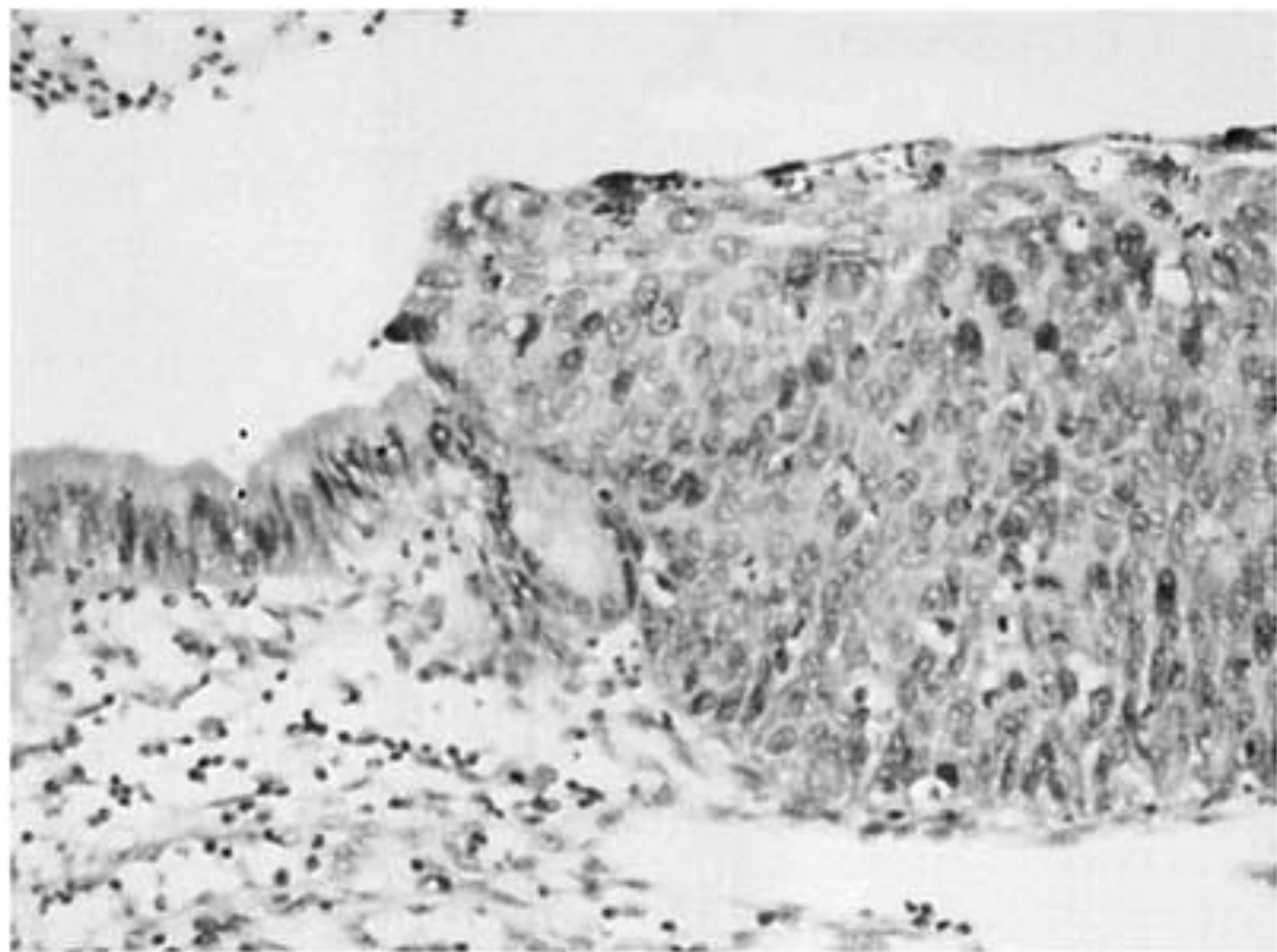
Reserve cells differentiate into normal-appearing mature squamous epithelium that replace glandular epithelium. (Similar changes occur in squamous metaplasia of the bronchus.)



Dysplasia

Squamous epithelial cells show abnormal maturation and cytologic abnormalities.





The world is a tragedy to those who feel, but a comedy to those who think

