

## Thyroid Gland

- The largest endocrine gland, located in the anterior neck, consists of two lateral lobes connected by a median tissue mass called the *isthmus*
- Composed of follicles that produce the glycoprotein *thyroglobulin*
- Colloid (thyroglobulin + iodine) fills the lumen of the follicles and is the precursor of thyroid hormone

## Thyroid Hormone (TH)

- Thyroid hormone – the body’s major metabolic hormone
- Consists of two closely related iodine-containing compounds
  - $T_4$  – thyroxine; has two tyrosine molecules plus four bound iodine atoms
  - $T_3$  – triiodothyronine; has two tyrosines with three bound iodine atoms

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## Effects of Thyroid Hormone

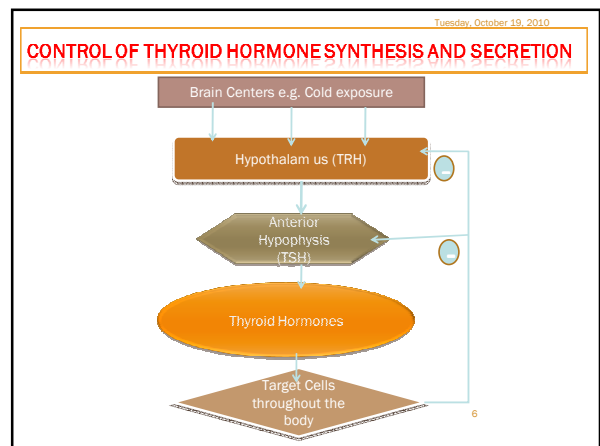
- TH is concerned with:
  - Glucose oxidation
  - Increasing metabolic rate
  - Heat production
- TH plays a role in:
  - Maintaining blood pressure
  - Regulating tissue growth
  - Developing skeletal and nervous systems
  - Maturation and reproductive capabilities

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## REGULATION OF THYROID HORMONE

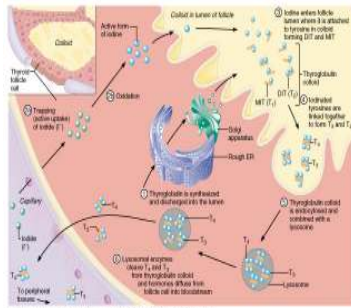
- $T_4$  and  $T_3$  bind to thyroxine-binding globulins (TBGs) produced by the liver
- Both bind to target receptors, but  $T_3$  is ten times more active than  $T_4$
- Peripheral tissues convert  $T_4$  to  $T_3$
- Mechanisms of activity are similar to steroids
- Regulation is by negative feedback
- Hypothalamic thyrotropin-releasing

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## Synthesis of Thyroid Hormone

- Thyroglobulin is synthesized and discharged into the lumen
- Iodides ( $I^-$ ) are actively taken into the cell, oxidized to iodine ( $I_2$ ), and released into the lumen
- Iodine attaches to tyrosine, mediated by peroxidase enzymes, forming  $T_1$  (monoiodotyrosine, or MIT), and  $T_2$  (diiodotyrosine, or DIT)
- Iodinated tyrosines link together to form  $T_3$  and  $T_4$
- Colloid is then endocytosed and

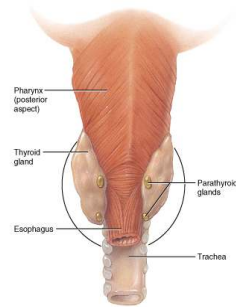


## Calcitonin

- A peptide hormone produced by the parafollicular, or C, cells
- Lowers blood calcium levels in children
- Antagonist to parathyroid hormone (PTH)
- Calcitonin targets the skeleton, where it:
  - Inhibits osteoclast activity and thus bone resorption and release of calcium from the bone matrix
  - Stimulates calcium uptake and incorporation into the bone matrix
- Regulated by a humoral (calcium ion)

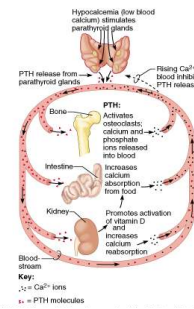
## Parathyroid Glands

- Tiny glands embedded in the posterior aspect of the thyroid
- Cells are arranged in cords containing oxyphil and chief cells
- Chief (principal) cells secrete PTH
- PTH (parathormone) regulates calcium



## Effects of Parathyroid Hormone

- PTH release increases  $Ca^{2+}$  in the blood as it:
  - Stimulates osteoclasts to digest bone matrix
  - Enhances the reabsorption of  $Ca^{2+}$  and the secretion of phosphate by the kidneys
  - Increases absorption of  $Ca^{2+}$  by intestinal mucosal cells
- Rising  $Ca^{2+}$  in the



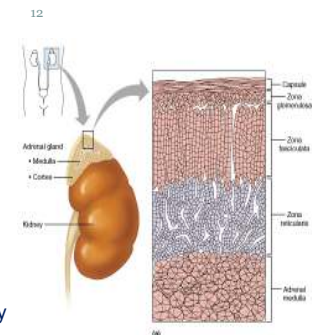
## ADRENAL (SUPRARENAL) GLANDS

- Adrenal glands – paired, pyramid-shaped organs atop the kidneys
- Structurally and functionally, they are two glands in one
  - Adrenal medulla – nervous tissue that acts as part of the SNS
  - Adrenal cortex – glandular tissue derived from embryonic mesoderm

Thomson, October 10, 2010

## Adrenal Cortex

- Synthesizes and releases steroid hormones called *corticosteroids*
- Different corticosteroids are produced in each of the three layers
  - Zona glomerulosa – mineralocorticoids (chiefly aldosterone)
  - Zona fasciculata – glucocorticoids (chiefly cortisol)
  - Zona reticularis –



## Mineralocorticoids

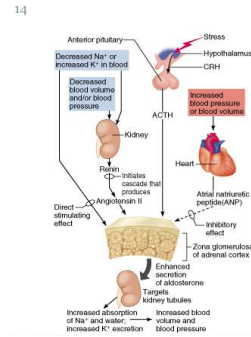
- Regulate the electrolyte concentrations of extracellular fluids
- Aldosterone – most important mineralocorticoid
  - Maintains  $\text{Na}^+$  balance by reducing excretion of sodium from the body
  - Stimulates reabsorption of  $\text{Na}^+$  by the kidneys
- Aldosterone secretion is stimulated by:
  - Rising blood levels of  $\text{K}^+$
  - Low blood  $\text{Na}^+$

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## The Four Mechanisms of Aldosterone Secretion

- Renin-angiotensin mechanism – kidneys release renin, which is converted into angiotensin II that in turn stimulates aldosterone release
- Plasma concentration of sodium and potassium – directly influences the zona glomerulosa cells
- ACTH – causes small increases of aldosterone during stress



## Glucocorticoids (Cortisol)

- Help the body resist stress by:
  - Keeping blood sugar levels relatively constant
  - Maintaining blood volume and preventing water shift into tissue
- Cortisol provokes:
  - Gluconeogenesis (formation of glucose from noncarbohydrates)
  - Rises in blood glucose, fatty acids, and amino acids

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## EXCESSIVE LEVELS OF GLUCOCORTICIDS

- Excessive levels of glucocorticoids:
  - Depress cartilage and bone formation
  - Inhibit inflammation
  - Depress the immune system
  - Promote changes in cardiovascular, neural, and gastrointestinal function

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## Gonadocorticoids (Sex Hormones)

- Most gonadocorticoids secreted are androgens (male sex hormones), and the most important one is testosterone
- Androgens contribute to:
  - The onset of puberty
  - The appearance of secondary sex characteristics
  - Sex drive in females
- Androgens can be converted into estrogens after menopause

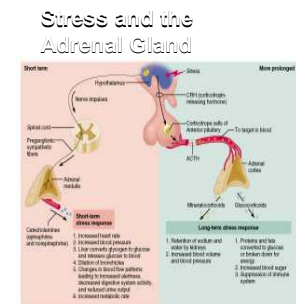
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## Adrenal Medulla

### Activity

- Made up of chromaffin cells that secrete epinephrine and norepinephrine
- Secretion of these hormones causes:
  - Blood glucose levels to rise
  - Blood vessels to constrict
  - The heart to beat faster
  - Blood to be diverted to the brain, heart, and skeletal muscle
- Epinephrine is the more potent stimulator of the heart and metabolic activities
- Norepinephrine is more influential on peripheral vasoconstriction



## Pancreas

- A triangular gland, which has both exocrine and endocrine cells, located behind the stomach
- Acinar cells produce an enzyme-rich juice used for digestion (exocrine product)
- Pancreatic islets (islets of Langerhans) produce hormones (endocrine products)
- The islets contain two major cell types:
  - Alpha ( $\alpha$ ) cells that produce glucagon
  - Beta ( $\beta$ ) cells that produce insulin

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## Hormones of the pancreas

### Glucagon

- A 29-amino-acid polypeptide hormone that is a potent hyperglycemic agent
- Its major target is the liver, where it promotes:
  - Glycogenolysis – the breakdown of glycogen to glucose
  - Gluconeogenesis – synthesis of glucose from lactic acid and noncarbohydrates
  - Release of glucose to the blood from liver cells

### Insulin

- A 51-amino-acid protein consisting of two amino acid chains linked by disulfide bonds
- Synthesized as part of proinsulin and then excised by enzymes, releasing functional insulin
- Insulin:
  - Lowers blood glucose levels
  - Enhances transport of glucose into body cells
  - Counters metabolic activity that would enhance blood glucose levels

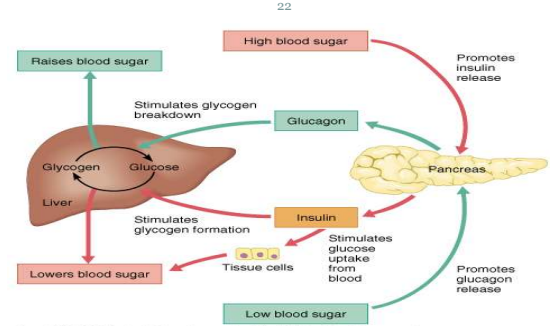
## Effects of Insulin Binding

- The insulin receptor is a tyrosine kinase enzyme
- After glucose enters a cell, insulin binding triggers enzymatic activity that:
  - Catalyzes the oxidation of glucose for ATP production
  - Polymerizes glucose to form glycogen
  - Converts glucose to fat (particularly in adipose tissue)

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## The hyperglycemic effects of glucagon and the hypoglycemic effects of insulin



## Diabetes Mellitus (DM)

- Results from hyposecretion or hypoactivity of insulin
- The three cardinal signs of DM are:
  - Polyuria – huge urine output
  - Polydipsia – excessive thirst
  - Polyphagia – excessive hunger and food consumption

Organ/tissue involved	Organ/tissue responses to insulin deficiency	Resulting condition of	Signs and symptoms
	Decreased glucose uptake and utilization	Hyperglycemia	Glycosuria
	Glycogenolysis	Diabetes Mellitus	Polyuria Polydipsia Polyphagia
	Protein catabolism and gluconeogenesis		
	Lipolysis and ketogenesis	Ketoneuria and ketonuria	Acidotic breath Hyperosmolar Spontaneous hypokalemia pain Cardiac irregularities Central nervous system depression, coma

Legend: = Muscle, = Adipose tissue, = Liver