Autonomic Nervous System 3

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Catecholamines:

The autonomic fibers that secrete catecholamines are known as adrenergic fibers. They secrete noradrenaline, adrenaline and dopamine (may be chemical transmitters in some neurons of the CNS).

✓ Site of catecholamines release:-

- Noradrenaline is the chemical transmitter of all sympathetic postganglionic fibers (not adrenaline) except sympathetic postganglionic to sweat gland and skeletal muscle blood vessels (cholinergic).
- Catecholamines (adrenaline 80% and noradrenaline 20%) are also secreted by adrenal medulla in emergency conditions.

Functions of the adrenal medulla;

- It augments the activity of ANS in conditions of emergency by increasing the amounts of catecholamines in plasma carried by the blood to different tissues.
- The released catecholamines from adrenal medulla have longer effect (10 times) than that of the sympathetic stimulation because it is slowly removed from blood.
- Adrenaline is secreted mainly by the adrenal medulla than by sympathetic nerve endings which secrete mainly noradrenaline.
- Chemical transmitter of adrenal medulla is acetylcholine that acts on the nicotinic receptors of the chromaffin cells to release adrenaline mainly (chemical secretion).

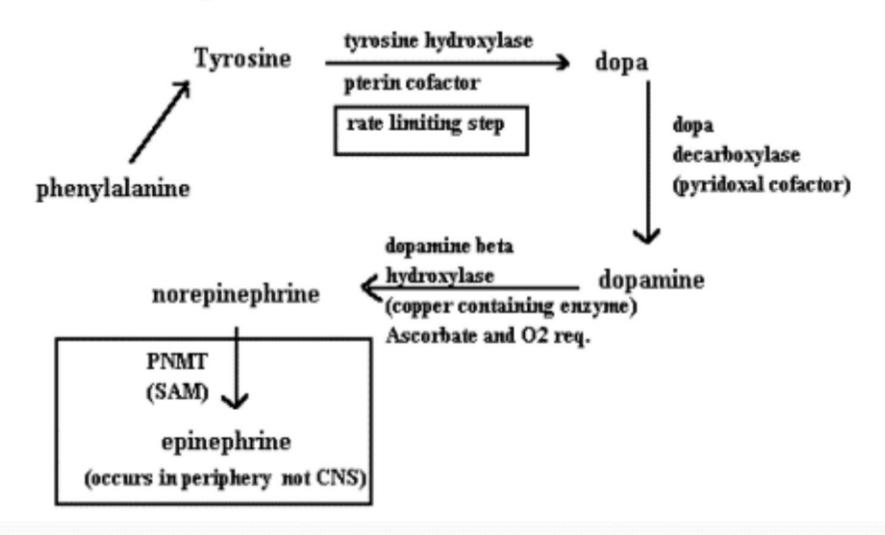
Adrenaline	Noradrenaline.
 Contain methyl group. 	 Devoid from methyl
	group.
• Released mainly from	 Released mainly from
adrenal medulla.	postganglionic
	adrenergic
	sympathetic fibers.
 Have more affinity to 	\bullet Have more affinity to α
β receptors.	receptors.
 Strong metabolic and 	 Vascular action mainly
cardiac actions.	(vasoconstriction)

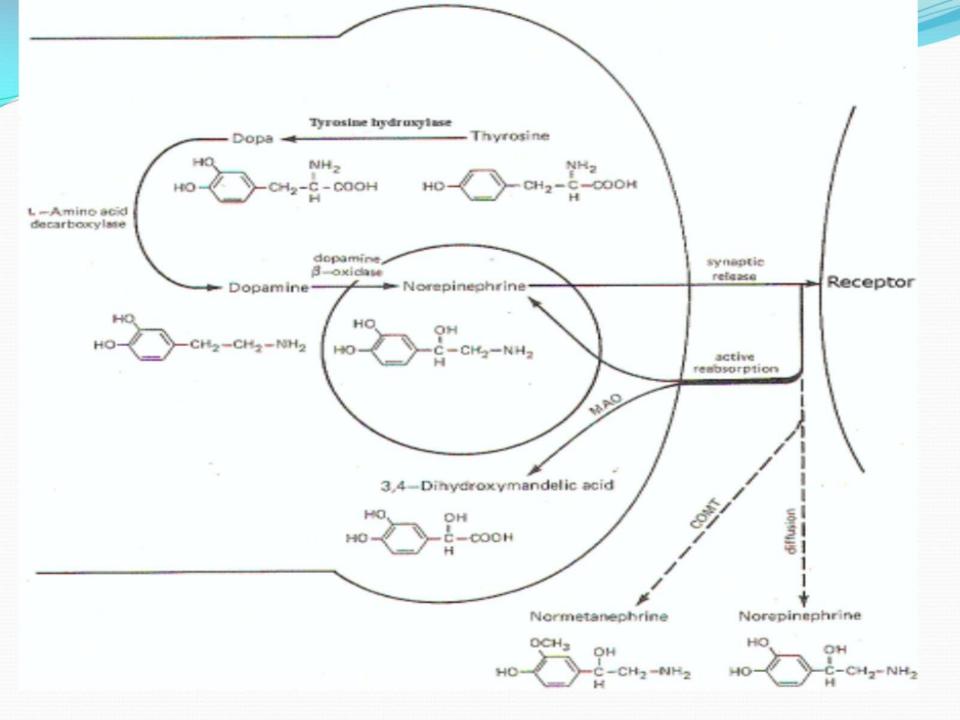
✓ Biosynthesis of catecholamines;

- They are formed in adrenergic neurons and the adrenal medulla from amino acid tyrosine (from diet or synthesized from amino acid phenylalanine).
- CA are stored inside special vesicles in the adrenergic nerve terminals or adrenomedullary cells called chromaffin granules.

Catecholamines (dopamine and norepinephrine)

synthesis and metabolism





✓ Methods of inactivation:-

- After release of CA and production of their effects, they are rapidly inactivated by:
- Active reuptake into the vesicles of the nerve endings, inhibited by Reserpine.
- Oxidation by an enzyme (Monamine oxidase, MAO) that present in high condentration in the mitochondria of adrenergic nerve fibers → Vanilyl mandelic acid derivatives (VMA, 24h. urinary VMA is an indication of plasma CA secretion) → excreted in urine.
- <u>Orthomethylation</u> in the liver by an enzyme (catecholamine orthomethyl transferase, COMT).

√Adrenergic receptors (Metabotropic receptors):

They are of two types: alpha and beta receptors.

- Stimulation of α-receptors produces mainly excitatory effects except in GIT while β-receptors produces inhibitory effects except on the heart.
- Adrenaline stimulates both α, β receptors,
 predominantly β-receptors while
 Noradrenaline predominantly α-receptors.

Items	Alpha (α)	Beta (β)
- Types	- α_1 , α_2	- β ₁ , β ₂
- Action	- Mainly excitatory	- Mainly inhibitory (β ₂)
	(α_1) as it is	as it is produced muscle
	produced muscle	relaxation (GIT and
	contraction	urinary bladder),
	(sphincters),	bronchodilatation and
	mydrasis and blood	blood vessels
	vessels	vasodilatation but in
	vasoconstriction	heart, it is excitatory
	except in intestine	(β_1) .
	(α_2) inhibitory.	- Lipolysis (β ₃)
	- Presynaptic α_2 are	 Glycogenolysis (β₂)
	called autoreceptors.	
- Response to	Sensitive to	Sensitive to
catecholamines	noradrenaline more than	adrenaline more than
	adrenaline	noradrenaline

▼ The sympathomimetics drugs:

- These are the drugs that have an effect similar to sympathetic stimulation, they include;
- Ganglion stimulants: small doses of Nicotine.
- Drugs that stimulate α-receptors directly: Phenylephrine (is better than atropine in fundus examination as it have short duration of action), adrenaline and noradrenaline (life saving in bronchial asthma).
- Drugs that stimulate β-receptors directly: Isoprenaline (used in cases of severe bradycardia).

✓ The sympatholytics drugs:

These are drugs which block the actions of sympathetic nervous system, they include:

- Ganglion blockers: *large dose of nicotine*.
- Drugs that prevent storage: Reserpine (given to irritable patient but must be sure that liver function is normal, why?
- Drugs that prevent synthesis: α -methyl-p-tyrosine.
- Alpha-blockers (anti-hypertensive):
 Phentolamine.
- Beta-blockers (act on heart to treat tachycardia, arrhythmia): Propranolol.