

PHARMACOLOGY OF PERIPHERAL NERVOUS SYSTEM

AUTONOMIC NERVOUS SYSTEM

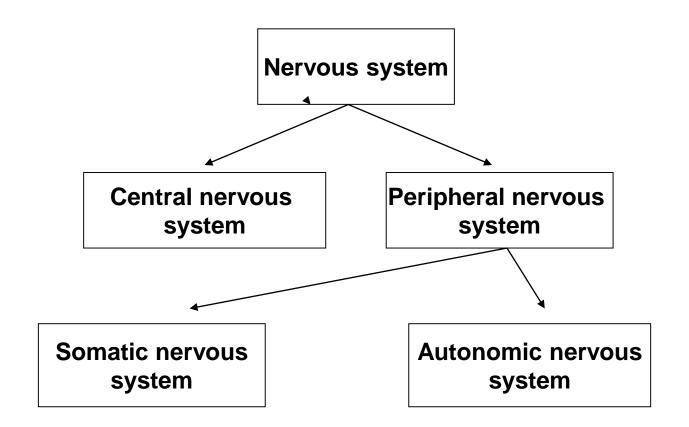
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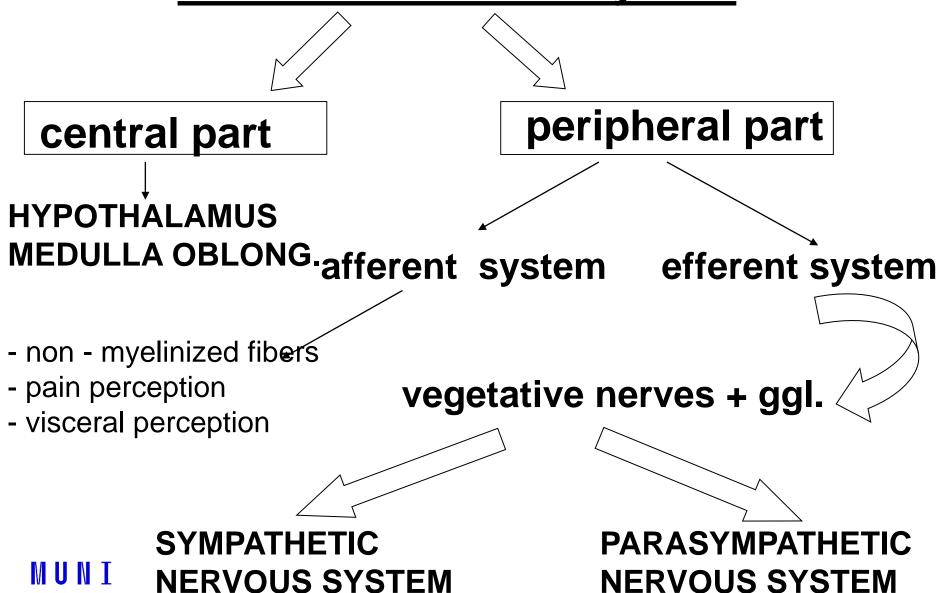
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Department of Pharmacology





Autonomic nervous system



Main functions of ANS

- contractions and relaxations of smooth muscles
- function of all exocrine and some of endocrine glands
- heart functions
- metabolic functions



ANS

Sympathetic

- = adrenergnic system
- thoracolumbal s.
- fight or flight
- noradrenaline(NA)
- α a ß receptors

Parasympathetic

- = cholinergnic system
- craniosacral s.
- rest and digest
- acetylcholine
- N a M receptors

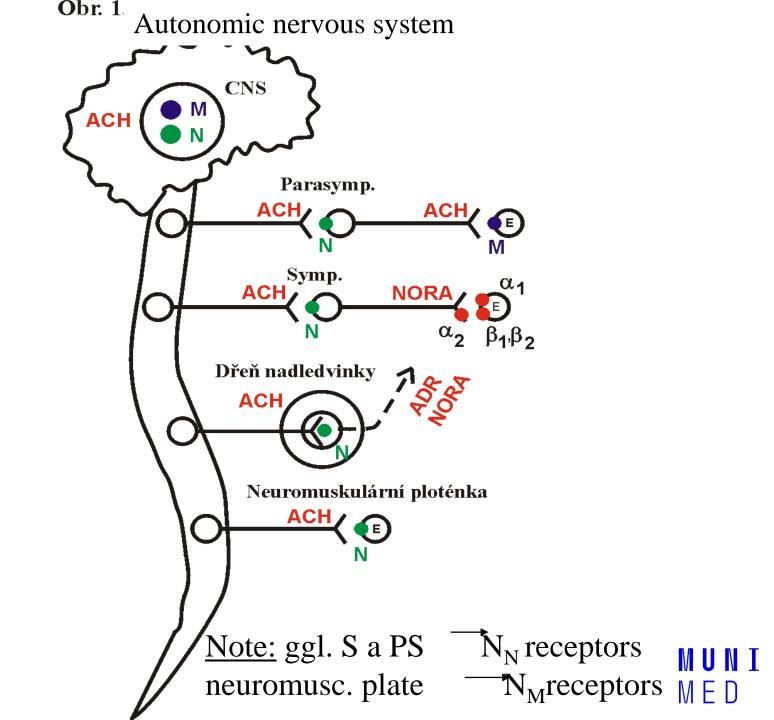


Autonomic nervous system

The activity is mutually regulated

- heterotropic interactions
- homotropic interactions
- most of visceral organs is inervated by both S and PS
- opposite activity bronchi, heart, bladder,,...
- similar action salivary glands
- only S blood vessels





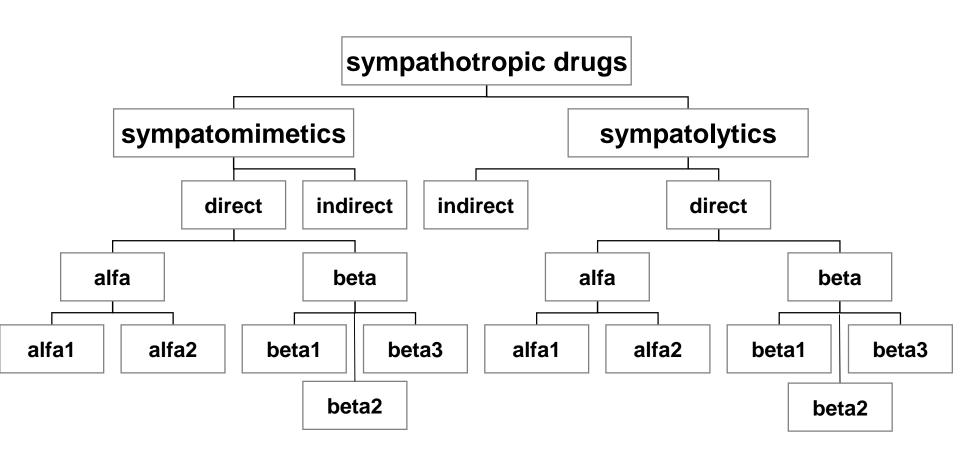
Autonomic acting pharmaceuticals

On the basis of mechanism of action - drugs:

- 1. **binding to the receptors** for Ach or NA:
 - a) starting reaction = agonist DIRECT MIMETICS
 - b) receptor blockade = antagonist DIRECT LYTICS
- 2. changing the synaptic concentration of NT intervene in the fate of the Ach or NA (affect the synthesis, storage, release from nerve endings, inactivation); do not bind directly to receptors on the effector organs
 - a) increase of NT effect = **INDIRECT MIMETICS**
 - b) decrease of NT effect = **INDIRECT LYTICS**

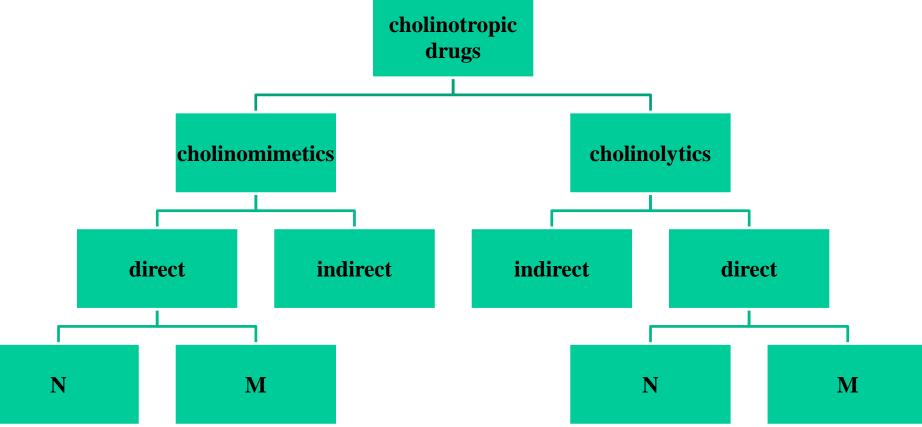


Vegetative acting drugs 2. sympatotropic



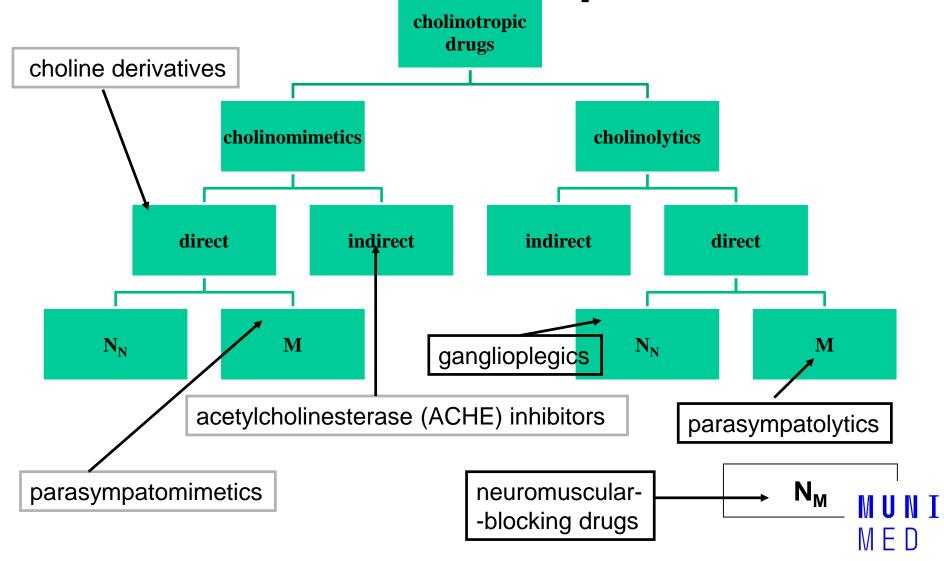


Vegetative acting drugs 2. cholinotropic



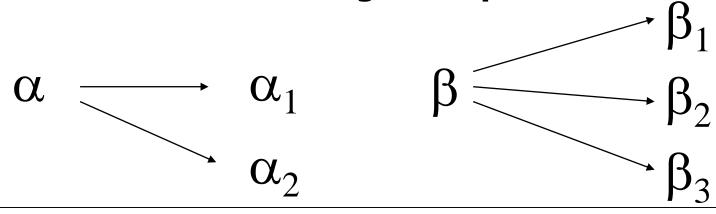


Vegetative acting drugs 2. cholinotropic



ANS RECEPTORS

adrenergic receptors



cholinergic receptors



NICOTINE: N

-skeletal muscle N_M

-vegetative ganglia N_N

-(<u>CNS</u>)



MUSCARINIC:

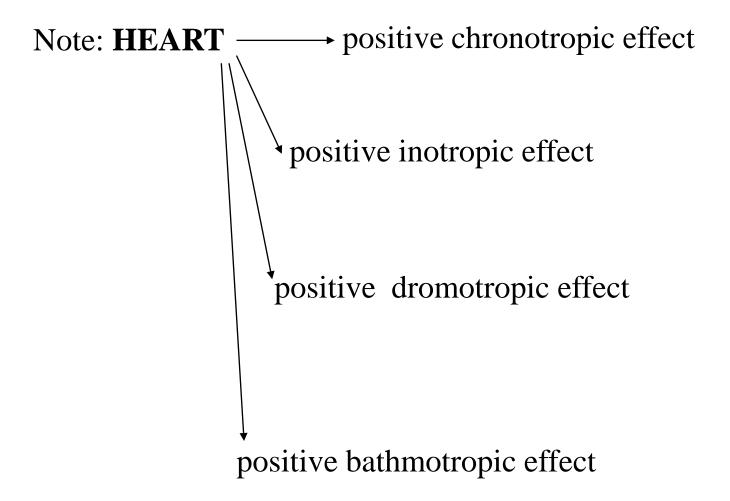
 M_1, M_2, M_3, M_4, M_5



organ	receptor		sympathetic system	parasympathetic system
heart	ß ₁	M	+ chrono, dromo, bathmo, inotropic	- chrono, dromo bathmo, inotrop.
eye	α_1 β_2	M	mydriasis acomodation into the distance	miosis acom.to close
respiratory tract	(α_1) $\underline{\beta_2}$	M	bronchoconstriction bronchodilatation	bronchoconstriction ↑secretion
blood vessels	α_1 (α_2) β_2	M	vasoconstriction vasoconstr. dilatation (coronary, blood vessels in skeletal muscles)	dilatation MUNI MED

organ	rece	ptor	sympathetic system	parasympathetic system
GIT	$\frac{\alpha_1}{\alpha_2}$ $\underline{\beta_2} > \beta_1$	M	↓ motility and tone sphincter contraction secretion inhibition	↑ motility sphincter relaxation secretion stimulation ↑ gastr. secretion
urinary bladder	α_1 $\beta_{2,}$ β_3	M_3	sphinct. contraction relax. of the bladder wall	sphinct. relaxation contract. of the bladder wall
kidney	<u>β</u> ₁ >β ₂		↑ renin secretion	
uterus	$egin{array}{c} lpha_1 \ eta_2 \end{array}$		contraction relaxation-tocolysis	MUN MED

			1
organ	receptor	sympathetic system	parasympathetic system
liver	α_1, β_2	glycogenolysis gluconeogenesis	
pancreas	α_2 β_2	↓insulin secretion ↑insulin secretion	
sexual organs	α_1 M	ejaculation	erection
glands	α_1 M β_2	sparse secretion viscous secretion	sparse significantly increased secretion
			M U N M E D





Adrenergic receptors

- metabotropic
- α_1 , α_2 a β_1 , β_2 a β_3
- stimulated by noradrenaline (norepinephrine)



Receptor α_1 stimulation:

- <u>vasoconstriction</u> (skin, mucous membranes, splanchnic area,..)
- mydriasis

(+ ↓ intraocular pressure)

- contraction of pregnant uterus
- ejaculation
- urinary bladder sphincter contraction, GIT sphincter contraction
- glycogenolysis and gluconeogenesis stimulation
- (reduce secretion of bronchial glands)



Receptor α_2 stimulation:

- (presynaptic) <u>increased NA release</u> (espec. in CNS)
- stimulation of platelet aggregation
- <u>vasoconstriction in local application</u>, otherwise the influence of stimulation of central receptors to reduce sympathetic tone and BP
- <u>hypotensive effect of central mechanism</u>
- inhibition GIT secretion
- inhibition of lipolysis, increased fat storage



Receptor β_1 stimulation :

heart:

- 1 HR (+ chronotropic effect) SA node
- 1 automaticity (+ bathmotropic) AV node, ventricles
- 1 force of heart contraction (inotropic effect)
- 1 conduction (dromotropic effect)
- 1 oxygen consumption

kidney:

• 1 renin secretion

Receptor β_2 stimulation:

- <u>vasodilatation</u>, <u>espec.</u> in <u>skeletal muscles</u> ("preparation for fight or flight"), ↓ diastol. BP, <u>vasodilatation in coronar</u> <u>blood vessels</u>
- bronchodilatation
- relaxation of uterus (indic. in impending preterm birth)
- intestine wall relaxation
- intestinal passage decrease
- urinary bladder wall relaxation
- **glycogenolysis** ↑ glycemia, increased insulin secretion
- <u>blockade of mast cells degranulation</u>



Receptor β_3 stimulation:

- lipolysis
- urinary bladder wall relaxation (m. detrusor)



Cholinergic receptors

MUSCARINIC:

- M₁("neural") CNS, peripheral neurons, parietal cells of stomach, (glands with external secretion)
- M₂ ("heart") heart (SA, atria, AV, ventricles), (smooth muscle (GIT), neuronal tissue), presynapt. neur.endings
- M₃ glands, blood vessels (smooth muscle, hl. sval, endothelium), smooth muscles: bronchial muscles, GIT, urinary bladder, eye
- M₄ salivary glands, GIT (muscles), eye, CNS
- M₅ lungs, CNS

Cholinergic receptors

- M metabotropic
- stimulated by acetylcholine
- N coupled with ion channels
- stimulated by nicotine

