

Introduction

Adrenergic drugs release acetylcholine at the post synaptic level. Only post ganglionic sympathetic nerves are adrenergic as they use epinephrine as neuron transmitters.

Function Norepinephrine is a neurotransmitter of adrenergic synapse, coming from tyrosine → DOPA → Dopamine, stored in vesicles until it needs to be released, when impulses from CNS send the signals. They use exocytosis so the norepinephrine gets into the synaptic cleft. In synaptic cleft, norepinephrine needs to act on post synaptic receptors.

Norepinephrine is released from the presynaptic membrane

- 1) Most of it is taken back w/out any changes to be reused (uptake 1),
- 2) Taken by factors cells AKA non neuronal uptake (uptake 2)
- 3) Diffuse
- 4) Degrade (by enzyme present in the synaptic left, minor).

Introduction (cont)

Norepinephrine can act on presynaptic receptors, regulate release of Norepinephrine from sympathetic nerve endings (AKA alpha 2 receptors), regulates -ive feedback mechanism (uptake 1)

Drugs - based on potency

α-adrenoc-eptors Norepinephrine → epinephrine → isoprenaline

β-adrenoc-eptors Isoprenaline → epinephrine → norepinephrine

α,β adrenoc-eptors Epinephrine + norepinephrine

Groups of drugs

Subtypes Mechanism of action

α1 adrenoc-eptors **Postsynaptic** - mediate effect on sympathetic nerve system (smooth muscle tissues + organs + blood vessels))

α2 adrenoc-eptors **Presynaptic** - out of synapse (mediate effects on catecholamines released from adrenal medulla)

β1 adrenoc-eptors **Postsynaptic** - mediate effect on sympathetic nerve system (heart)

β2 adrenoc-eptors **Presynaptic** - out of synapse (mediate effects of catecholamines circulating in blood leads to vasodilation)

Adreno-positive drugs

α, β-adrenomimetics (natural - used intravenously) **Epinephrine, norepinephrine**

α-adrenomimetics (selective) **α1-adrenomimetics = phenylephrine** (increases BP locally + effective orally)
α2-adrenomimetics = Naphazoline, clonidine (decreases BP, antihypertensive drug)

β-adrenomimetics (selective) **β1,β2-adrenomimetics = isoprenaline**
β1-adrenomimetics = dobutamine (if patient doesn't have hypoxia or ischemic disease)
β2-adrenomimetics = salbutamol (short acting drug), **phenoterol, salmeterol** (slow/-prolonged acting drug)

Sympathomimetics (indirect binding on receptors) **Ephedrine** (acts on sympathetic endings, stimulates release of norepinephrine, indirectly produces effects on post synaptic receptors)

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Adreno-negative drugs

Adrenoblockers/-adrenoceptors antagonists (direct binding on receptors)

α, β-adrenoblockers (natural) - **Labetalol**

α-adrenoblockers - **α1,α2-adrenoblockers** (non-selective) - **Phentolamine**, dihydroergotamine

α1-adrenoblockers - **α1a-adrenoblockers** - **tamsulosin**

α1b-adrenoblockers - **prazosin**

β- Adrenoblockers / adrenoceptor antagonists

β1,β2-Adrenoblockers (non-selective) - **Propranolol**, timolol

β1-adrenoblockers - **atenolol, metrolol**

Sympatholytics (indirect binding on receptors)

Reserpine, guanethidine

Indications + side effects

α adreno-mimetics

Pharmacological effects - Vasoconstriction of blood vessels

Mydriasis

Decrease NA peripheral

Indications + side effects (cont)

β adreno-mimetics

Indications - Bronchial asthma, uterine relaxation (preserving pregnancy)

pharmacological effects - cause dilation of bronchial passages, Vasodilation in muscle and liver, Relaxation of uterine muscle, Release of insulin

α adreno-blockers

Side effects - hypotension

Tachycardia

β adreno-blockers

Mechanism of action of antihypertensive action of beta adrenoblockers - decrease cardiac output

Decrease renin secretion

reduce central sympathetic activity (selective BB))

Side effects

Side effects β2 AB - *Heart insufficiency, bronchoconstriction, *hypoglycemia**, fatigue, dizziness, nausea, diarrhea

Side effects β1 AB -

Specific drugs

Phentolamine (antagonises α1+2 adrenoceptors)

Decreases BP caused by adrenaline, affects -ive feedback mechanism (α2-adrenoceptors) in the synapse

Indications - Pheochromocytoma (adrenal gland tumour) + endarteritis (inflammation of arteries, legs)

Side effects - Orthostatic (standing) collapse (severe drop in BP) + tachycardia

reverse effects of adrenaline on BP

Specific drugs (cont)

Tamsulosin (Selectively anatogonises/-blocks α1A-adrenoceptors)

Indication - benign prostate hyperplasia (increased cell production in a normal tissue or organ)

Relax smooth muscle of prostate gland

Help to decrease these symptoms

Prazosin (Selectively antagonises/blocks α1B-adrenoceptors in blood vessels)

Does not affect the -ive feedback mechanism in synapse

Indication - arterial hypertension

Side effect - orthostatic collapse

Specific drugs

Propranolol - Non-selective β1,β2-adrenoblocker

Decrease heart output, decrease activity of SA + AV nodes, decrease BP due to action on:

Heart - decreases heart output

Kidney (propranolol decreases production of renin in the kidney)

CNS (decrease sympathetic activity on PNS))

Atenolol, Metoprolol (Selective β1-Adrenoblocker)

Cardioselective in therapeutic doses

drugs of choice in cardiac patients

