

Sympatholytic drugs;

- Adrenergic Blockers.
- Adrenergic antagonists.
- Adrenergic receptor antagonists.
 - Adrenoceptor antagonists.

by

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Clinical Uses of \(\beta \) blockers;

- Cardiovascular Uses;
 - Hypertension,
 - -IHD,
 - -MI,
 - Cardiac arrhythmias,
 - Chronic (not acute) heart failure,
 - Dissecting aortic aneurysm,
 - Hypertrophic cardiomyopathy.

- Other Uses;
- Pheochromocytoma,
- Thyrotoxicosis,
- Migraine,
- Anxiety (to reduce somatic manifestations),
- Essential tremors,
- Glaucoma,
- Cirrhosis (to reduce portal vein pressure).

Cardiovascular Uses;

> Hypertension;

- β blockers are relatively mild antihypertensives.
- All β blockers are nearly equally effective.
- First choice antihypertensive drugs in patients of IHD.
- Can be given in combination with other antihypertensives.
- Combination with verapamil and diltiazem is contraindicated.

> IHD;

- All β blockers **benefit angina of effort.**
- Interpretation of the following of the follo
- ↑ exercise tolerance.
 - Cardiac work and oxygen consumption are reduced.

- Undesirable effects of β blocking drugs in angina;
- Total coronary flow is \downarrow (subepicardial region) due to blockade of β_2 and unopposed α_1 receptors.
- These deleterious effects of β blocking agents can be balanced by the concomitant use of nitrates.

Myocardial Infarction;

- Given I/V within 4-6 hours of an acute MI;
 - May **limit infarct size** by reducing O2 consumption.
 - Prevent arrhythmias including VF.
- Secondary prophylaxis of MI (after recovery from acute MI);
 - ↓ mortality by 20 %
 - » By preventing reinfarction.
 - » By preventing sudden VF at second attack of MI.
- Not to be given in;
 - » shock, bradycardia or heart block.

Cardiac arrhythmias;

- Class II anti-arrhythmic drugs.
 (Vaughan Williams classification)
- Suppress adrenergic mediated ectopic activity;
 - Very useful in treating inappropriate sinus tachycardia, atrial and nodal Extrasystoles provoked by emotions or exercise.
 - Thyrotoxicosis
 - $-\beta$ blockers suppress extrasystoles and tachycardia.
- They control ventricular rate in Atrial Fibrillation and A. Flutter but only occasionally restores sinus rhythm.
- **PSVT** (adenosine, verapamil) --- Esmolol I/V.

Chronic (not acute) heart failure;

- β blockers may worsen heart failure.
- Beneficial hemodynamic effects at low doses.
 - Over activity of cardiac β_1 receptors exert toxic effects on heart.
 - Low dose β blockade antagonize the sympathetic over activity on myocardium.

Not to be given in patients;

- Acute heart failure,
- With marked fluid retention,
- Those requiring I/v vasodilatation,
- Those requiring I/v ionotropic drugs.

Dissecting aneurysm of aorta;

- Reduce cardiac contractile force and aortic pulsation.
- Decrease rate of development of systolic pressure.

Hypertrophic cardiomyopathy;

- β blockers improve diastolic dysfunction and LV compliance.
- β blockers improve cardiac output during exercise, but have little effect at rest.

> FALLOT'S TETRALOGY;

 To reduce right ventricular infundibular spasm in Tetralogy of Fallot with Pulmonary stenosis.

Other Uses;

> Pheochromocytoma;

- To control tachycardia and arrhythmia.
- Should never be given before α blocker.
 - Dangerous rise in BP
- Suppress cardiomyopathy caused by excess catecholamines.

> Thyrotoxicosis;

- Propranolol rapidly control symptoms without significantly affecting the thyroid status.
 - Palpitation, nervousness, tremors, severe myopathy and sweating.
- Highly valuable during thyroid storm;
 - Inhibit peripheral conversion of T4 to T3.
- Used preoperatively and while awaiting response to antithyroid drugs / Radioactive iodine.

→ Migraine;

Propranolol is used in prophylaxis of migraine.

>Anxiety;

- Propranolol exert antianxiety effect under conditions which provoke nervousness and panic attacks.
 - Block peripheral manifestation of anxiety (palpitation, tremors)
 - Reduce somatic manifestations.
 - Ineffective in anxiety neurosis.

≻ Cirrhosis of liver

To reduce portal vein pressure.

Essential tremors;

- Non selective β blockers are effective.
- Do not benefit Parkinsonian tremors.

≻Glaucoma;

- Timolol and other ocular β blockers are first choice drugs for chronic simple (wide angle) glaucoma.
- Used as adjuvant in angle closure glaucoma.

Adverse Effects (propranolol);

CNS:

 Sedation, sleep disturbances, depression, increased dreaming, nightmares and rarely hallucinations.

CVS:

- Bradycardia, cardiac conduction abnormalities,
- Cardiac failure;
 - Precipitate CCF and edema by blocking sympathetic support to heart.
- Peripheral arterial insufficiency,
 - Cold hands and feet,
 - Tiredness and reduced exercise capacity,
 - due to blunting of β2 mediated increase in blood flow to exercising muscles and attenuation of glycogenolysis and lipolysis.
- Exacerbate variant (prinzmetal's) angina due to unopposed α mediated coronary constriction.

Adverse Effects;

Respiratory Tract:

Bronchoconstriction, Can precipitate an attack of bronchial asthma.

• GIT:

Nausea, vomiting, constipation, diarrhea.

Metabolism:

- Hypoglycemia.
- Impair carbohydrates tolerance in prediabetics.
- Alter plasma lipid profile;
 - Total triglycerides and LDL-cholesterol tend to increase while HDLcholesterol falls.
 - Cardioselective β blockers and those with intrinsic sympathomimetic activity do not affect blood lipids.

Allergy:

- Rash, fever, purpura.
- Impotence in males.

Effects of sudden withdrawal;

- Worsening of angina,
- Rebound hypertension,
- Tachycardia,
- cardiac arrhythmias and even sudden death.
 - This is due to super sensitivity of β receptors occurring as a result of long term reduction of agonist stimulation.

Precautions/ contraindications;

Precautions;

- Patients with asthma.
- patients with diabetes mellitus esp. IDDM.
 - Non selective Beta blockers mask the effects of hypoglycemia.

Contraindications;

- Cardiogenic shock,
- Right ventricular failure secondary to pulmonary hypertension,
- Congestive cardiac failure (acute),
- Greater than 1st degree heart block,
- Hypotension,
- Peripheral arterial insufficiency e.g., Raynaud's phenomenon.
- Severe Bronchial Asthma and COPD.
- Patients on MAO inhibitors.

