

B. Pharm - IIIrd yr. 6th Sem.
(ROP-363)

PHARMACOLOGY - II

Topic:- Pharmacology of CVS. (Cardiovascular System)

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UNIT-1 Pharmacology-2

CARDIAC GLYCOSIDES

These are glycosidic drugs having cardiac inotropic property. They ↑ myocardial contractility and output in a hypodynamic heart - increases a ↑ in O₂ consumption. Also ↑ myocardial efficiency ↑.

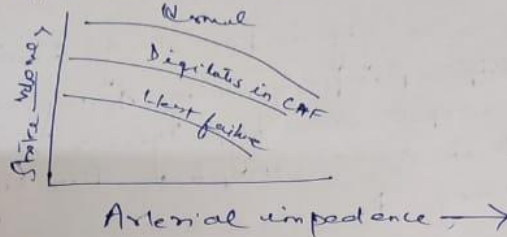
Cardiac glycosides & their source :-

Source	Glycosides
1) <i>Digitalis purpurea</i> (leaf)	Digitalin, Gitoxin.
2) <i>Digitalis lanata</i> (leaf)	Digitalin, Digoxin.
3) <i>Strophanthus kumbe</i> (bud)	Strophanthin K.
4) <i>Strophanthus greggii</i> (bud)	Strophanthin-G.
5) <i>Convallaria majalis</i>	Convallotoxin
6) <i>Bufo vulgaris</i>	Bufo toxin
7) Semi synthetic	Acetyl-digoxin Acetyl-strophanthin

Pharmacological Actions

1) Heart:- *Digitalis* has direct effect on myocardial contractility & electrophysiological properties. In addition it has vagomimetic action, reflex effects due to activation in baroreceptors & effect of direct sympathetic activity.

① causes adrenergic dependent ↑ in force of contraction of heart's positive inotropic action.



Relationship b/w peripheral resistance & cardiac output in normal & failing heart. Action of digitals in failing heart.

Mechanism of Action:

Digitals ↑ the force of cardiac contraction by a direct action independent of innervation. It selectively binds to extracellular face of cell membrane associated Na^+K^+ ATPase of myocardial fibres and inhibits this enzyme. Inhibition of this cation pump results in decrease accumulation of Na^+ intracellularly. This indirectly results in intracellular Ca^{2+} accumulation.

② Blood vessels: - ① has mild direct vasoconstrictor action

② has prominent effect on BP. Systemic may ↑ & diastolic may fall in CHF patients.

③ Kidney:- Diuretics in CHF patients. ③

④ CNS:- Higher doses cause CTZ stimulation.
Lower higher doses causes Nausea Vomiting,
diarrhoea, ataxia, visual disturbances.

Pharmacokinetics :- Digoxin metabolized in liver
via hepatic enterohepatic circulation.

Uses:-

- 1) Congestive heart failure.
- 2) Cardiac Arrhythmias.

Precautions of Cardiac glycosides

- 1) Hypokalaemia.
- 2) elderly, renal or severe hepatic disease.
- 3) Ventricular tachycardia.
- 4) Acute Myocardial infarction.
- 5) Thyrotoxicosis.

Antiarrhythmic drugs (4)

These are the drugs used to prevent or treat irregularities of cardiac rhythm.

Classification Antiarrhythmic drugs act by blocking myocardial Na^+ , K^+ or Ca^{2+} channels:

Membrane stabilizing agents (Ca^{2+} channels blockers) — Quinidine.
Procainamide.
Disopyramide.
Moricizine.
Lidocaine, Mexiletine.
Tocainide, Phenytoin.
Flecainide, Encainide.
Propafenone.

Antidrenergic agents (β -blockers) — Propranolol, Esmolol.
Sotalol.

Agents widening AP (Prolong repolarization) — Amiodarone.
Bretylium.

Calcium channel blockers — Verapamil.
Diltiazem.

Quinidine : alkaloid from cinchona bark.
dextro isomer of quinine.

Pharmacological actions :- 1) Heart :
Antivagal action.
 \downarrow automaticity in

Purkinje fibres and other ectopic foci. ⓐ

ⓐ ↑ threshold for excitation.

~~MOA~~ Skeletal muscle: - ↓ contractility.

GIT: - can reduce vomiting, diarrhoea.

CNS: - Neurological effects ↑ dose.

Uterus: - ↑ uterine contractile force.

MOA: - ⓐ Block myocardial Na^+ channels on the open state → ↓ automaticity &

Maximal rate of 0 phase depolarization as a frequency. Prolongation of APD (Action potential duration) due to K^+ channel block while slighting of ERP caused by moderate effect of recovery of Na^+ & K^+ channels.

ADRs: - GIT intolerance.

hypersensitivity.

Idiosyncrasy.

fear, asthma, vascular collapse.

Uses: -

ⓐ Uses in atrial & ventricular arrhythmias used to maintain sinus rhythm after AF has been terminated by DC shock. & to prevent recurrences of VT. (ventricular tachycardia).

Antianginal Drugs

Antianginal Drugs are those that prevent, abort or diminish attacks of anginal pectoris.

AP - Anginal Pectoris is a pain syndrome due to ischaemia of an adverse oxygen supply/demand situation in a portion of Myocardium.

Classification :-

1) Nitrate :

a) Short Acting :- glyceryl trinitrate. (Nitroglycerin)

b) Long Acting :- Isosorbide dinitrate
Isosorbide Mononitrate
Erythrityl tetranitrate
Penterythritol tetranitrate

2) β -Blockers :- Propranolol.
Metoprolol.
Atenolol.

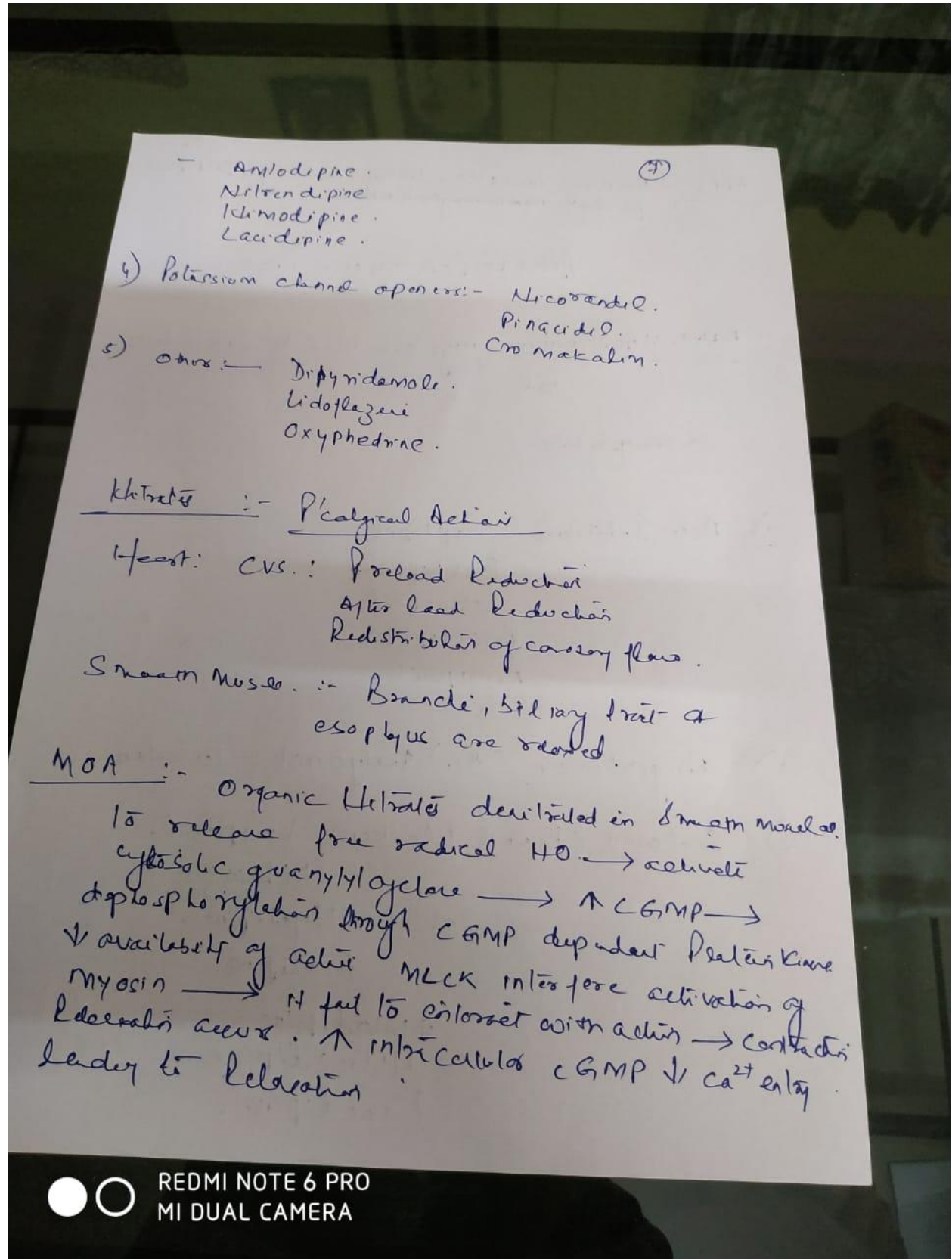
3) Calcium Channel Blockers.

a) Phenyl alkyl amine :- Verapamil.

b) Benzothiazepine Diltiazem.

c) Dihydropyridines :- Nifedipine
Felodipine.
Amlodipine





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ADP's throbbing headache, flushing, ⁽⁸⁾swelling, ⁽⁸⁾dizziness, ⁽⁸⁾fatigue.
~~metformin, glimepiride~~

ANTI HYPERTENSIVE DRUGS

ANTI-HYPERTENSIVES Drugs are used to lower down B.P.

Classification :-

- 1) ACE Inhibitors :- Captopril.
Enalapril.
Lisinopril.
Perindopril.
Ramipril.
- 2) Angiotensin (AT₁) antagonist :- Losartan
- 3) Calcium Channel Blockers :- Verapamil.
Diltiazem.
Nifedipine.
Fludipine
Amlodipine
Hydrochloride
Lacidipine.



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4) Diuretics:

Thiazides:- Hydrochlorothiazide.
Chlorthalidone.

High ceiling:- Furosemide.

K⁺ sparing:- Spironolactone.
Triamterene.
Amiloride.

5) β -Adrenergic Blockers:- Propranolol.
Metoprolol.
Atenolol.

Labetalol ($\alpha + \beta$ Blockers)

6) α -Adrenergic Blockers:- Prazosin.

Terazosin

Phentolamine

Pheoxybenzamine.

7) Central Sympatholytics:- Clonidine

Methyldopa

8) Vasodilators:- Arterial:- Hydralazine.

Minoxidil

Diazoxide.

Arterial + Venous:- Sodium Nitroprusside.

Prinidipin

Cromakalim.



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