

# OCULAR PHARMACOLOGY II

Miotics, Glaucoma Meds, Antibiotics,  
Corticosteroids, NSAIDs Antivirals,  
Antihistamines  
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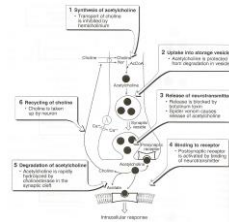
## CHOLINERGIC AND ANTICHOLINERGIC MEDS

### CHOLINERGIC DRUGS

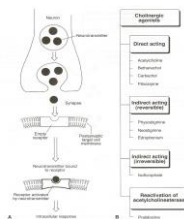
- Direct agonists
- Indirect agonist
- Antagonist

Generic Name	Trade Name	Strength
<b>Cholinergic drugs</b>		
Carbachol	Isopto Carbachol	1.5%, 2%
Pilocarpine HCl	Isopto Carpine	0.25%, 1%, 2%, 4%
	Available generically	0.5%, 1%, 2%, 3%, 4%, 6%
Pilocarpine HCl ointment	Pilopec HS gel	4%
<b>Cholinesterase inhibitors</b>		
Physostigmine	Available generically	1 mg/mL ampule
Edrophonium iodide	Phospholine Iodide	0.125%

### CHOLINERGIC NEURON



### CHOLINERGIC AGONISTS



### CHOLINERGIC DRUGS

- Direct agonist:
- Miosis, accommodation
- Increase aqueous outflow decrease IOP
- **Acetylcholine 1% carbachol 0.01% intracameral use** constrict pupil anterior segment surgery
- **Acetylcholine** rapid effect but short lived
- **Carbachol** 100 times more effective longer lasting 24 hours and decreases IOP

## CHOLINERGIC DRUGS DIRECT AGONIST

- **Pilocarpine:**
  - lowers IOP by increasing outflow
  - Open angle glaucoma treatment
- **Side effects:**
  - Miosis older patients with cataract difficulty in scotopic conditions
  - Cataractogenesis
  - Induced myopia and accommodation problems for younger patients; slow dissolving **pilocarpine gel** at bedtime

## CHOLINERGIC DRUGS: INDIRECT AGONIST CHOLINESTERASE INHIBITORS

- **Phospholine iodide**
  - No longer available for ophthalmic use in US
  - More potent than direct acting use twice a day
- Physostigmine, neostigmine, edrophonium

## CHOLINERGIC ANTAGONISTS

- **Atropine:** post-synaptic blockage of acetylcholine
- Pupil dilation, cycloplegia for iritis, accurate cycloplegic refraction
- Treat malignant glaucoma
- Systemic atropine given during ophthalmic surgery involving EOM manipulation to block oculocardiac reflex and prevent bradycardia and hypotension
- **Side effects of systemic absorption of atropine eye drops:**
  - Side effects treated with physostigmine
  - Other antagonists: tropicamide, cyclopentolate, homatropine, scopolamine, Flushing, tachycardia, constipation, urinary retention, delirium.

## INDIRECT ACTING ANTAGONIST

- **Edrophonium test:** increases acetylcholine at neuromuscular junction
- Myasthenia gravis: antibodies to acetylcholine receptor resulting in generalized weakness, ptosis and diplopia
- Edrophonium used to diagnose myasthenia gravis
- **Neostigmine** can be given IM for the same purpose and has a longer duration of activity
- Allows clinician more time to make orthoptic measurement for desired endpoint

## NEUROMUSCULAR BLOCKADE FOR GENERAL ANESTHESIA

- Globe laceration repair under general anesthetic
- Use of neuromuscular blocking drugs by anesthetist; i.e. succinylcholine is a "depolarizing" neuromuscular blocker and can cause contraction of EOMs on induction of general anesthetic and should not be used
- Exert force on open globe
- increase IOP in other cases where pressure measurement is desired such as in an examination under anesthesia

## MYDIATICS AND CYCLOPLEGICS

Table 10-2 Mydriatics and Cycloplegics

Generic Name	Trade Name	Strengths	Dose	Duration of Action
Phenylephrine HCl	Adrenalin	Solution, 2.5%, 10%	20-40 mm	10-20 min
	Albutin	Solution, 2.5%, 10%		
	Epinephrin	Solution, 2.5%		
	Neosyn	Solution, 2.5%		
	Neo-Synephrine	Solution, 2.5%, 10%		
Tropicamide	Atropine	Solution, 1%	40-100 mm	3-8 h
	Atropine	Solution, 1%		
	Atropine	Solution, 1%		
	Atropine	Solution, 1%		
	Atropine	Solution, 1%		
Cyclopentolate HCl	AK-Pentolate	Solution, 1%	30-40 mm	2 days
	Cyclopent	Solution, 0.2%, 2%		
	Cyclopent	Solution, 1%, 2%		
	Cyclopent	Solution, 1%, 2%		
	Cyclopent	Solution, 1%, 2%		
Homatropine	Homatropine	Solution, 0.5%	30-40 mm	4-7 days
	Homatropine	Solution, 0.5%		
	Homatropine	Solution, 0.5%		
	Homatropine	Solution, 0.5%		
	Homatropine	Solution, 0.5%		
Scopolamine	Scopolamine	Solution, 0.25%, 1%	30-40 mm	4-6 h
	Scopolamine	Solution, 0.25%, 1%		
	Scopolamine	Solution, 0.25%, 1%		
	Scopolamine	Solution, 0.25%, 1%		
	Scopolamine	Solution, 0.25%, 1%		
Cyclopentolate HCl	Cyclopentolate	Solution, 0.25%, 1%	30-40 mm	1-2 days
	Cyclopentolate	Solution, 0.25%, 1%		
	Cyclopentolate	Solution, 0.25%, 1%		
	Cyclopentolate	Solution, 0.25%, 1%		
	Cyclopentolate	Solution, 0.25%, 1%		



# GLAUCOMA MEDICATIONS

## ADRENERGIC DRUGS

- Norepinephrine (NE) is transmitter
- Alpha and beta receptors for NE
- Direct agonists
- Indirect agonist
- antagonists

Generic Name	Trade Name	Strength
<b>α<sub>1</sub>-Adrenergic agonists</b>		
Epinephrine HCl	Propine	0.1%
	Auralgan preservative	0.1%
	Not available in the United States	0.1%, 1%, 2%
<b>α<sub>2</sub>-Selective agonists</b>		
Apraclonidine HCl	Iopidine	0.1%, 1% (long-term treatment)
Brimonidine tartrate	Alphagan P	0.1%, 0.15%
	Auralgan preservative	0.2%
Brimonidine tartrate/brimonidine maleate	Combigan	0.2%/0.05%

## ALPHA-ADRENERGIC DRUGS

- Alpha-1 phenylephrine Mydriasis stimulate iris dilator muscle
- Naphazoline: decongestant with rebound vasodilation and hyperemia
- Elevate systemic blood pressure
- Caution: using 10% phenylephrine **topically 5mg per drop**; MI, stroke, cardiac arrest risks
- Systemic "pressor" dose bolus IV 50-100µgram

Primary Mechanism of Action	Drug Class	Examples
Decrease aqueous humor production	1. β <sub>1</sub> -Adrenergic antagonists	Timolol (bimatoprol), carboxylate
	2. α <sub>2</sub> -Adrenergic agonists	Apraclonidine, brimonidine
Increase trabecular outflow	1. α <sub>1</sub> -Adrenergic agonists	Phenylephrine
	2. α <sub>2</sub> -Adrenergic agonists	Brimonidine, apraclonidine
Increase uveoscleral outflow	1. Prostaglandins	Latanoprost, bimatoprost, travoprost, tafoprost
	2. α <sub>2</sub> -Adrenergic agonists	Apraclonidine, brimonidine

## ALPHA-2 ADRENERGIC AGONISTS

- Apraclonidine (iopidine)
  - Pre and post YAG cap and SLT, cataract extraction to manage increased IOP
  - Prevents NE release decrease pupil size
  - Decreases aqueous production and increases outflow
  - Topical sensitivity 40% and tachyphylaxis limit long term use
- Brimonidine (Alphagan)
  - Topical sensitivity 15%
  - 0.2% with BAK
  - 0.15% with polyquad
  - 0.1% with sodium chloride preservative Purite Alphagan-P
  - Avoid in infants less than 2 years: hypotension, hypothermia, bradycardia
  - CNS effects resulting from medication crossing the blood-brain barrier

## ALPHA-2 ADRENERGIC AGONISTS

- Caution in patients using apraclonidine or brimonidine :
  - Patients taking systemic MAO inhibitors or tricyclic antidepressants
  - Hypertensive risks

## INDIRECT ACTING ADRENERGIC AGONISTS

- Cocaine 4% or 10%
- Hydroxyamphetamine 1%
- Office diagnostic testing to confirm Horner syndrome
- Available through compounding pharmacies





## STEROIDS ADVERSE EFFECTS: SYSTEMIC ADMINISTRATION

- Suppression of pituitary-adrenal axis
- Hyperglycemia, muscle-wasting, osteoporosis
- Redistribution of fat from periphery to trunk
- Euphoria
- Insomnias
- Aseptic necrosis of the hip
- Peptic ulcer
- Diabetes mellitus
- psychosis

## STEROID-INDUCED ELEVATION OF IOP

- 4% develop an IOP > than 31 mm Hg after 6 weeks
- Reduce aqueous outflow is cause of IOP rise
- Dexamethasone > prednisolone > FML
- Reversible upon discontinuation of the drug if use is less than one year
- Permanent elevations of IOP if used more than 18 months
- Loteprednol 0.2%, 0.5% have lower incidences of increased IOP
- Fluocinolone implant for chronic uveitis
- Triamcinolone 40mg/ml preservative free for intravitreal use: increased rates of cataract and need for glaucoma treatment

## NONSTEROIDAL ANTI-INFLAMMATORY DRUGS (NSAIDS)

- **Flurbiprofen** 0.3% (Ocufen) first available topical NSAID used pre-op to reduce intraoperative miosis for cataract surgery
- **Diclofenac** 0.1% (Voltaren) prophylaxis and treatment of post-op inflammation and CME
- **Ketorolac** (Acular) post-op inflammation and allergic conjunctivitis treatment
- **Nepafenac** (Nevanac), **Bromfenac** (Xibrom) twice daily dosing treating postop cataract and retinal surgery inflammation
- NSAIDS have topical anesthetic properties: useful in short term management of: corneal abrasion, anterior segment procedures and refractive surgery



## ANTI-ALLERGY MEDICATIONS

## ANTI-ALLERGIC DRUGS: MAST CELL STABILIZERS AND ANTIHISTAMINES

Generic Name	Trade Name	Class
Acetylsalicylic acid	Aspirin	H <sub>1</sub> antagonist
Antihistamine H <sub>1</sub>	Benadryl	H <sub>1</sub> antagonist
Antihistamine H <sub>2</sub>	Cimetidine	H <sub>2</sub> antagonist
Antihistamine H <sub>3</sub>	Cimetidine	H <sub>3</sub> antagonist
Antihistamine H <sub>4</sub>	Cimetidine	H <sub>4</sub> antagonist
Antihistamine H <sub>5</sub>	Cimetidine	H <sub>5</sub> antagonist
Antihistamine H <sub>6</sub>	Cimetidine	H <sub>6</sub> antagonist
Antihistamine H <sub>7</sub>	Cimetidine	H <sub>7</sub> antagonist
Antihistamine H <sub>8</sub>	Cimetidine	H <sub>8</sub> antagonist
Antihistamine H <sub>9</sub>	Cimetidine	H <sub>9</sub> antagonist
Antihistamine H <sub>10</sub>	Cimetidine	H <sub>10</sub> antagonist
Antihistamine H <sub>11</sub>	Cimetidine	H <sub>11</sub> antagonist
Antihistamine H <sub>12</sub>	Cimetidine	H <sub>12</sub> antagonist
Antihistamine H <sub>13</sub>	Cimetidine	H <sub>13</sub> antagonist
Antihistamine H <sub>14</sub>	Cimetidine	H <sub>14</sub> antagonist
Antihistamine H <sub>15</sub>	Cimetidine	H <sub>15</sub> antagonist
Antihistamine H <sub>16</sub>	Cimetidine	H <sub>16</sub> antagonist
Antihistamine H <sub>17</sub>	Cimetidine	H <sub>17</sub> antagonist
Antihistamine H <sub>18</sub>	Cimetidine	H <sub>18</sub> antagonist
Antihistamine H <sub>19</sub>	Cimetidine	H <sub>19</sub> antagonist
Antihistamine H <sub>20</sub>	Cimetidine	H <sub>20</sub> antagonist

## MAST CELL STABILIZERS, ANTIHISTAMINES

- Human eye has approximately 50 million mast cells containing granules containing chemical mediators
- Immediate hypersensitivity reaction triggered when antigens combine with IgE on the surface of mast cells
- Mast cells release histamine and other factors
- Histamine increases capillary dilation, conjunctival swelling and injection
- Safer than steroids for chronic use



