

Carbonic Anhydrase Inhibitor

pCT

Acetazolamide
Dichlorophenamide
Methazolamide

↓ HCO_3^- reab

Alkaline diuresis

↓ aqueous humor [ciliary body]

↓ CSF

Metabolic Acidosis

Glucoma

Metabolic Alkalosis

rarely Diuretics

↑ Cl^- Metabolic Acidosis

calcium phosphate stones

↓ K^+

hypersensitive rxn

↑ NH_4^+

mountain sickness

Loop diuretics

Thick ascending
[loop of Henle]

Sulfonamide
most

Furosemide
Bumetanide
Torsemide

high ceiling [↑ dose ↑ effect]
most efficacious
No Tolerance

phenoxyacetic acid Ethacrynic Acid

least

↓ $\text{Na}^{+2}/\text{K}^+/2\text{Cl}^-$ cotransporter

Lumen (+) potential

↑ loss of Na^{+2} , Cl^- , K^+ , Mg^{+2} , Ca^{+2}

PTH → ↑ Ca^{+2} rabs.... [hypocalcemia rare]

↑ PTHs2 → vasodilation → RBF → diuresis

↓ K^+ Metabolic Alkalosis

↓ Mg^{+2} ↓ Na^{+2} ↓ Ca^{+2}

ototoxicity [reversible] [dose related]

Allergic rxns

severe dehydration

hyperuricemia → hypovolemia { uric acid metab...
hyperglycemia → ↓ K^+ → insulins

Rapid abso.. [orally]

Drug - Drug intns [plasma protein]

↳ ↑ free ↑ toxicity + elimination

eliminated

↳ Tubular Secretion
glomerular filtration

furosemid + ethacrynic [partially metab...]

↳ ↓ pulmonary congestion

↳ ↑ systemic venous

furosemide + bumetanide

↳ ↓ left ventricular filling
↳ ↓ CA

Torsemide active meta

$T_{1/2}$ longer

* Acute Pulmonary edema

Congestive HF → venous caputitatione

* hyperkalemia * R bilure

They diff in meta...

other renally

orally

Thiazide Hydrochlorothiazide

Chlorothiazide Injected / parenterally [water soluble] high solubility

Chlorthalidone slow abs... long duration

Thiazide Distal Tube

Thiazide-like Indapamide excreted Biliary system

Metolazone

distal

two

two

$\downarrow K^+$ metabolic Alkalosis $\uparrow Na^{+2}$ reabs $\rightarrow K^+$ sec..
 $\downarrow Na^{+2}$

$\downarrow Na^{+2}/Cl^-$ co-transporter

$\uparrow Na^{+2}/Ca^{+2}$ exchange

$\uparrow Ca^{+2}$ reab...

↓ PGI₂ synthesis

$\downarrow CA$

Nephrolithiasis $\rightarrow \uparrow Ca^{+2}$

Impotence, weakness, fatigue, paresthesia

photosensitivity

hyperuricemia

hyperglycemia

hyperlipidemia $\uparrow LDL$ give lipid drug with it

hypersensitivity anuria, thrombo..., -itis, Bown's

first line \rightarrow Hypertension [HTN]

Edema

Nephrolithiasis $\rightarrow \uparrow Ca^{+2}$

Nephrogenic diabetes insipidus unresponsive to ADH

($\rightarrow Na^{+2}$ restriction $\rightarrow \uparrow$ therapeutic effect)

Potassium-sparing Diuretics



minely to hypokalemia

Aldosterone Antagonists

spironolactone

↓ libido → Anti-androgenic oral enterohepatic cycling ↓ Dose in hepatic failure active metabolite canrenone drug-drug intx

Eplerenone

drug intx ketoconazole, itraconazole ↓ Bioavailability ↓ CYP3A4 ↑ Eplerenone in Blood

ENaC Inhibitors

Amiloride unchanged in urine, no specific dose in hepatic disease.

Triaterene meta in liver renally exr... shorter $t_{1/2}$ crystalluria

$\uparrow K^+$
Both ↑ Metabolic acidosis

Block Aldosterone receptors

↓ Na^{+} water reab...

↓ K^+ exr...

H^+ intercalated cells → depend on PGE₂ production

Spironolactone Gynecomastia

BPH

impotence, GI upset

Triamterene leg cramps, azotemia nephrolithiasis, interstitial nephritis

Acute Renal Failure

glucose intolerance

photosensitivity

Block Na^+ entry through selective channels

spare $K^+ + H^+$ ~ no Na^+

depends on renal PGE₂ production

ENaC Inhibitors nausea, vomiting headache

Mineralocorticoid excess

main → combined with other diuretics to prevent potassium loss, treat hypokalemia

10-20° hyperaldosteronism

Ectopic Aldosterone production

Osmotic Diuretics

PCT + Descending

* Mannitol naturally, not metab., osmotic diarrhea
excreted by glomerular filtration

urea

Glycerin

Isosorbide

act in water not Na^+

water \rightsquigarrow lumen

ADH \rightsquigarrow oppose in collecting tube \rightarrow urine volume

\uparrow water exc ... \rightsquigarrow $\uparrow \text{Na}^{+}$, dehydration

\uparrow ECV

dilutional hyponatremia

pulmonary edema

headache, nausea

congestive HF

$\uparrow \text{K}^+$

Acute renal failure \rightarrow PIs \rightsquigarrow vasodilation \rightsquigarrow $\uparrow \text{RBF}$ \rightsquigarrow prevent ARF, pigment load

hemolysis by hemoglobin rhabdomyolysis by Myoglobin

\downarrow intracranial \downarrow intraocular pressure

ADH Antagonists

collecting duct

selective Conivaptan ^{ok} nonpeptide IV, oral possible $\uparrow \frac{t_{1/2}}{2} = 5-10$ hrs

non-selective Demeclocycline [tetracycline]

Lithium not use as ADHR antagonist

Block V₂ receptors

\downarrow

\downarrow cAMP

\downarrow

\downarrow water reab....



so ↓ ADH by vasopressin Block

Severe hypernatremia $\uparrow\uparrow \text{Na}^+$

nephrogenic diabetes insipidus

dry mouth

hypotension

* syndrome of Inappropriate ADH secretion

[SIADH]

سُبْحَانَ اللَّهِ

الْحَمْدُ لِلَّهِ

لَا إِلَهَ إِلَّا اللَّهُ

اللَّهُ أَكْبَرُ

يَا أَيُّهَا الَّذِينَ آمَنُوا اذْكُرُو اللَّهَ ذِكْرًا كَثِيرًا

«اللَّهُمَّ إِنِّي أَسأَلُكَ الثَّباتَ فِي الْأَمْرِ، وَالْعَزِيمَةَ
عَلَى الرُّشْدِ، وَأَسأَلُكَ مُوْجَاتَ رَحْمَتِكَ، وَعَزَائِمَ
مَغْفِرَتِكَ، وَأَسأَلُكَ شُكْرَ نِعْمَتِكَ، وَحُسْنَ عِبَادَتِكَ،
وَأَسأَلُكَ قُلْبًا سَلِيمًا، وَلِسَانًا صَادِقًا، وَأَسأَلُكَ مِنْ
خَيْرِ مَا تَعْلَمُ، وَأَعُوذُ بِكَ مِنْ شَرِّ مَا تَعْلَمُ، وَأَسْتَغْفِرُكَ
لِمَا تَعْلَمُ، إِنَّكَ أَنْتَ عَلَامُ الْغُيُوبِ»^(١).