## Pharmacological Interventions for dizziness

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#### First a caution

- Torok N. Old and new in Meniere's disease. Laryngoscope 87:1870-1877, 1977
- 600 treatments reviewed ranging from spinal fluid drainage to numerous medications.
- Nearly all had 60% efficacy (natural history)
- A lot of these medications may be placebo's

## Processes we might try to treat

- Vertigo (nystagmus)
- · Motion sickness, emesis
- Compensation

## Processes we might NOT try to treat with medications

- Sensory ataxia (such as ototoxicity)
- BPPV (best managed with physical treatments)
- Malingering (drug treatment facilitates them)



## Neurotransmitters



Neurotransmitter	Peripheral	Central
Acetylcholine	Excitatory	Excitatory
GABA	?	Inhibitory
Histamine	?	?
Dopamine	?	Excitatory
Norepinephrine	?	Modulator ?
Glutamate	?	Excitatory
Serotonin	(stomach)	Excitatory ?

## Main drug categories

- · Anticholinergic
- · GABA agonists
- · Everything else

### **Anticholinergics**

- · Block central and peripheral ACH
- · Reduce vertigo and nausea from peripheral vertigo
- Reduce central nystagmus (in very high doses)
- Numerous interesting side-effects →

#### Scopolamine Muscarinic antagonist



- Scopolamine (Transderm-Scop)
- Does not require ingestion (many other oral GI drugs do same thing Levsin for example)
- · Apply every 3 days to skin surface
- CNS effects side and Withdrawal syndrome limit use

# Anticholinergic side effects (Locoweed poisoning)

- Confusion (similar to drug induced Alzheimer's)
- Dry mouth, loss of sweating
- Urinary hesitancy/stoppage. Constipation
- · Blurry vision
- · Cardiac conduction block
- Addiction



Oxytropis lambertii

# H1-antihistamines with strong anticholinergic properties

- meclizine (Antivert)
- dimenhydrinate (Dramamine)
- diphenhydramine (Benadryl)

Antihistamines must cross BB barrier -- i.e. Claratin, Allegra would not work

#### Antihistamine side effects

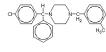
- Sleepiness
- · Weight gain

## Anticholinergic side effects

- Dry mouth and eyes
- · Constipation
- · Confusion

#### Meclizine (antivert)

- •12.5 TID or 25 TID. Lasts about 8 hours. Available OTC.
- Limitation is sedation and anticholinergic side effects
- Pregnancy: category B. May be best drug



C<sub>25</sub>H<sub>27</sub>CIN



# GABA agonists (benzodiazepines)

- · Modulate inhibitory transmitter GABA
- · Reduce vertigo and nausea from peripheral vertigo
- · Reduce nystagmus
- · Sedation, addiction limit usefulness
- ? May impede compensation (strangely, no evidence in humans for this – may actually do opposite)

### Benzodiazepines

- Diazepam (Valium, "Mothers little helper")
- Lorazepam (lorazepam)
- Klonapin (clonazepam)



### Dosing: beer scale 1 glass of beer =

- 2 mg of Valium
- 0.5 mg of Ativan
- 0.5 mg of Klonapin



### Benzodiazepines Bottom line

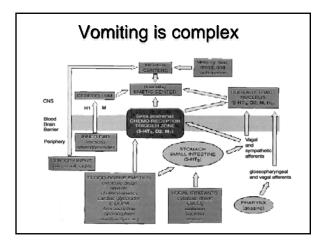
Extremely useful drugs Treat dizziness and anxiety Addiction is the big problem

#### **Diuretics**

- Dyazide and Maxide (Hctz+triamterine)
  - Menieres
- · Diamox (acetazolamide)
  - Menieres
  - Migraine
  - Periodic ataxia

## **Emesis**





## Drugs used for treatment of emesis

#### MOST IMPORTANT

- Dopamine blockers
- 5-HT3 antagonists
- Anticholinergics (OTC)
- · H1 antihistamines
- · Benzodiazepines

## Commonly used phenothiazine antiemetics dopamine blockers

prochlorperazine (Compazine) 5, 10 and 25 mg forms, including rectal suppositories. Pregnancy -- unknown

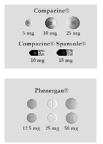
promethazine (Phenergan). 12.5, 25, 50 mg forms, including rectal suppositories 12.5 BID prn oral dose typical. Pregnancy Cat. C





## Commonly used phenothiazine antiemetics dopamine blockers

- •Powerful drugs
- •Major side effects
- •Use if you have a big vomiting problem



## odansetron (Zofran) 5HT3 receptor antagonist

- Dose: 32 mg IV, 4-8 mg PO. MLT form
- · Category B in pregnancy



Dr. Hain's drug of choice to use prior to nauseating PT session. Costs a LOT but generic is available

## Compensation

#### Compensation -- subtypes

- Static compensation recovery from tone imbalance (vertigo).
  - Largely automatic and not likely to be modified by drugs.
- Dynamic compensation (oscillopsia) readjust gain.
  - Takes some time, modifiable by medications.

#### Compensation -- goals

- Facilitate compensation for static vestibular lesions or central problems. (i.e. vestibular neuritis, bilateral loss)
- Halt compensation for transient vestibular lesions (i.e. Menieres attack).

## Drugs that accelerate dynamic compensation (in animals)

- · Amphetamines
- Bromocriptine (Dopamine agonist)
- ACTH (adreno-corticotrophic hormone)
- · Caffeine
- · Thyroid Releasing Hormone

Modified from Brandt, 1991

# Drugs that retard dynamic compensation in animals

- Phenobarbital (sedative, Barbituate)
- Dopamine antagonists (e.g. Lisuride, Thorazine)
- ACTH antagonists (e.g. steroids). Steroids seem to help in people!
- Diazepam, (GABA agonist, Valium). No evidence for this in people.

Modified from Brandt, 1991

### No pain – no gain ? or: Do rat studies apply to people ?

- Drugs that make people more comfortable often impede compensation in animals.
- Animal studies suggesting that medications impede compensation are generally not replicable in people.

### Summary

- · Large and complex pharmacology
  - Vertigo anticholinergics and benzodiazepines
  - Emesis phenothiazines and serotonin related drugs
  - $\\ Compensation sedatives \ and \ stimulants$

## More details

Hain TC, Yacovino D. Pharmacological Treatment of Dizziness. Continuum Neurotology Issue (Tusa R editor), 2006.

www.dizziness-and-balance.com