ANTIDEPRESSANTS: MAOIs (p.1)

1. Mono-Amine Oxidase Inhibitors (MAOIs)

developed in the 1950s are very effective (as effective as the tri-cyclic antidepressants/TCAs) their use is limited by serious (even fatal) side-effects if taken with other catecholamine agonists major concern is **HBP** --- **cardiovascular accidents** ("strokes")

2. <u>Current Uses for MAOIs</u>

are very effective can be **safer than TCAs** (**esp. re. cardiac arrhythmias**)

can work in pts. in whom TCAs/SSRIs have failed to work

are esp. useful in pts. with "atypical depression"

are esp. useful in pts. with **mixed depression & anxiety**

are good for tx of **anorexia**, **bulimia**

are good for tx of **bipolar depression & dysthymia**

are **not** good for tx. for depression in **elderly patients**...why?

are good for tx. of **panic attacks** and for **phobias**

ANTIDEPRESSANTS: MAOIs (p.2)

3. MAOI Effects on NS/Body

there are actually **two MAO enzymes**:

MAOA – found in NE and 5HT axon terminals MAOB – found in DA axox terminals, GI tract, liver

MAOIs block both enzymes --- beneficial main effects & neg. SEs

4. Non-Selective & Irreversible MAOIs

non-selective = block both MAOA and B enzymes irreversible = the MAOI enzyme is permanently blocked only when new molecules of MAO (A/B) are synthesized can they again metabolize monoamines (NE, 5HT, DA)

issues re. switching from one drug to another

e.g. MAOI to an SSRI or TCA

5. <u>Particular MAOIs</u> (non-selective, irreversible) phenelzine (Nardil), tranylcypromine (Parnate), isocarboxazid (Marplan)

6. Pharmacokinetics:

- **Parnate** $-\frac{1}{2}$ life = 2 hours, but is irreversible (is actually active metabolite that actually binds to MAO molecule)
 - as the S keeps taking Parnate, over one week about 70% of the MAO is blocked, slowly reaches a "**steady state**" between blocked MAO molecules and newly formed unblocked molecules of MAO
 - if S D/Cs the Parnate, it may take several weeks for the MAO levels to recover to normal/baseline levels
 - thus, may need **2 to 3 weeks of washout period** before starting a new (monoamine agonistic) antidepressant

neg. SEs:

ANTIDEPRESSANTS: MAOIs (p.3)

7. <u>Reversible & Selective MAOIs</u>

are not available in USA yet...but will be are short acting & highly selective, esp. for MAOA have little effect on MAOBs, and thus eating tyramine-rich foods is not a problem because are short acting, multiple doses are necessary no long-term deactivation of MAOA, no effect on MAOB minimal neg. SEs

e.g. brofaromine toloxatone pirlindole meclobemide (Aurorix) deprenyl, selegiline (Eldepryl)

8. Miscellaneous

effects on sleep architecture

"cheese reaction" & how to by-pass same

Potentiation of other drugs