

## Final Pharmacology Test Study Guide

### Antipsychotics & Serotonin

Serotonin – synthesised from Tryptophan

Found in the:

- CNS: sleep & wake cycles... many functions within the CNS
- CV: vasoconstrictor in splanchnic, renal, pulmonary beds, but also stimulates NO which has a vasodilatory effect, also has  $\alpha$ -adrenoceptor effect but there is serotonin auto-receptors on the vagus nerve  $\rightarrow$  bradycardia & hypotension
- GI: serotonin the primary neurotransmitter of the enteric NS  $\rightarrow$  increases motility in the gut, but also has inhibitory effect
- Inflammation: platelet activation/aggregation, increase vascular permeability, chemottractant

Synthesis:

- Tryptophan  $\rightarrow$  hydroxylated by Tryptophan hydroxylase  $\rightarrow$  **L-tryptophan 5-hydroxy-L-tryptophan** ( $\rightarrow$  requires **K<sub>2</sub>B<sub>6</sub>**)  $\rightarrow$  **5-hydroxy-tryptamine, 5-HT**, **Serotonin**  $\rightarrow$  secretory granules
- Serotonin can be converted into Melatonin (circadian rhythms) in **Pineal gland**  $\rightarrow$  **Suprachiasmatic nucleus**, (rich in melatonin receptors) regulates sleep/wake cycles

**Ramelteon** – prescribed for insomnia, agonist at the melatonin receptors in the suprachiasmatic nucleus, has short t<sub>1/2</sub> (no long-term effects of drowsiness)

Serotonin Receptors:  **$\rightarrow$  7 GPCR's except for 5HT<sub>2</sub>, varied distribution**

- 5HT<sub>1</sub>
  - o **G<sub>i</sub> (inhibitory)**  $\rightarrow$  **Ca<sup>2+</sup> channels & K<sup>+</sup> channels**
  - o 5HT<sub>1A</sub> & 5HT<sub>1D</sub> – auto-receptors, negative feedback to stop neurotransmitters
  - o (5HT<sub>1B</sub> subtype on vascular smooth muscle  $\rightarrow$  vasoconstriction)
- 5HT<sub>2</sub>
  - o **G<sub>q</sub> (IP<sub>3</sub>/Ca<sup>2+</sup> pathway)**  $\rightarrow$  smooth muscle contraction, platelet release of granules, CNS neurotransmission
  - o 5HT<sub>2A</sub> & 5HT<sub>2C</sub>  $\rightarrow$  **PLA<sub>2</sub>**  $\rightarrow$  cleaves arachidonic acid  $\rightarrow$  inflammatory response
    - Methysergide
    - Antipsychotic drugs
    - LSD
    - Psilocybin
    - Ayahuasca
- 5HT<sub>3</sub>
  - o **Ligand-gated Na<sup>+</sup> channel**  $\rightarrow$  depolarisation
    - CNS, GI tract, vagal afference
    - Ondansetron
- 5HT<sub>4</sub>
  - o **G<sub>s</sub> (stimulatory)**  $\rightarrow$  increase secretions & peristalsis
    - CNS, smooth muscle, GI system
- 5HT<sub>5,6</sub>
- 5HT<sub>7</sub>

# Mcneese State University Pharmacology Final Study Guide

**A Gutmann**



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