

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 = isochromotropic effect but there is serotonin auto-receptors on the vagus nerve → bradycardia & hypotension
- GI: serotonin the primary neurotransmitter of the enteric NS → increases motility in the gut, but also has inhibitory effect
- Inflammation: platelet activation/aggregation, increase vascular permeability, chemottractant

Synthesis:

- Tryptophan → hydroxylated by Tryptophan hydroxylase → **L-tryptophan 5-hydroxytryptamine** (requires **5-HTT**) → **5-hydroxy-tryptamine, 5-HT**, **Serotonin** → secretory granules
- Serotonin can be converted into Melatonin (circadian rhythms) in **Pineal gland** → **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_{1/2} (no long-term effects of drowsiness)

Serotonin Receptors: **7 GPCR's except for 5HT2, varied distribution**

- 5HT1
 - o **G_i (inhibitory) → Ca²⁺ channels & K⁺ channels**
 - o 5HT1A & 5HT1D – auto-receptors, negative feedback to stop neurotransmitters
 - o (5HT1B subtype in vascular smooth muscle → vasoconstriction)
- 5HT2
 - o **G_q (IP₃/Ca²⁺ pathway) → smooth muscle contraction, platelet release of granules, CNS neurotransmission**
 - o 5HT2A & 5HT2C → **PLA₂ → cleaves arachidonic acid → inflammatory response**
 - **Methysergide**
 - **Antipsychotic drugs**
 - **LSD**
 - **Psilocybin**
 - **Ayahuasca**
- 5HT3
 - o **Ligand-gated Na⁺ channel → depolarisation**
 - **CNS, GI tract, vagal afference**
 - **Ondansetron**
- 5HT4
 - o **G_s (stimulatory) → increase secretions & peristalsis**
 - **CNS, smooth muscle, GI system**
- 5HT5,6
- 5HT7

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