

03 - 3. Dopamine

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© SPMM Course 3. Dopamine

• tyrosine → l-dopa → dopamine Source Source • tyrosine hydroxylase Rate limiting step Rate limiting step • Monoamine oxidase (MAO) & Catechol-o-methyl transferase (COMT). • MAO-A more selectively metabolizes norepinephrine and serotonin • MAO-B more selectively metabolizes dopamine. Breakdown enzymes Breakdown enzymes • Homovanillic acid Breakdown product Breakdown product • Dopamine transporter (cocaine inhibits this transported) Reuptake Reuptake • Motivation, novelty seeking, reward circuitry (addictions), arousal and motor movement gating in basal ganglia Function Function • 5 types; D1 to D5 . All are G protein coupled • D1-like → D1 & D5; increase adenylate cyclase (stimulatory). D1 exclusively postsynaptic; resistant to antagonism. D5 more limbic in distribution; 10 times higher dopamine affinity • D2-like → D2,3 & 4 ; decrease adenylate cyclase (inhibitory). D4 is found primarily in the frontal cortex and clozapine has a high affinity. D4-selective antagonists do not have antipsychotic efficacy. Receptors Receptors • Levels low in Parkinson's; high in psychosis especially at mesolimbic area; may be low in anhedonia and negative symptoms in mesocortical area. Disorders Disorders

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