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NEW POTENTIAL MOLECULAR TARGETS FOR CANNABIS ADDICTION

M. Vieira-Coelho^{1,2,3}, J. Azevedo³, M. Esteves³

¹Institute of Pharmacology and Therapeutics, Faculty of Medicine, Porto University, ²Institute for Molecular and Cell Biology, University of Porto, ³Department of Psychiatry, Hospital de S. João, Porto, Portugal

Cannabis is considered the most widely abused illicit drug in the world. The recent rising prevalence of cannabis use by young adults and increasing evidence of adverse health effects makes the search for new pharmacotherapy to reduce cannabis abuse extremely important. To date no medication has been approved for the treatment of cannabis addiction.

This study reviews recent results with potential interest for future pharmacological treatment of Cannabis dependence. Most of the relevant data obtained for treatment of cannabis dependence target the endocannabinoid or the central cholinergic systems, both involved and interact in brain systems implicated drug reinforcement. In laboratory animals blockade of cannabinoids CB₁ receptors reverses central effects of cannabinoids. Rimonabant is a selective, orally active, cannabinoid CB₁ receptor antagonist (inverse agonist) that has been shown in animals to modulate cannabinoid signaling in brain reward circuit. In humans, it has been shown that rimonabant, single or repeated oral doses blocked psychological and physiological effects of smoked marijuana (1). Although psychiatric adverse side effects like depression were reported with rimonabant, this compound was already approved for treatment of obesity and metabolic syndrome. Very recently, blockade of α_7 nicotinic receptors was shown to reverse abuse-related behavioral and neurochemical effects of cannabinoids in rats (2).

In conclusion, besides cannabinoid CB₁ receptor, the homomeric α_7 nicotinic receptors are novel molecular targets in the development of new drugs for treatment of cannabis addiction.

1. Solinas M., et al. (2007). *J Neurosci.* 27(21):5615-20.
2. Huestis MA, et al. (2007). *Psychopharmacol.* 194(4):505-15.