PSYC1022: The Psychology of Addiction

Topic 1: Origins & Manufacture of addictive drugs (III)

Dr. Helena Pacitti

Outline

- Others:
 - MDMA
 - Cannabis









1

MDMA

- 3,4-methylenedioxy-N-methylamphetamine
 - often classed within the 'amphetamine' group
 greater empathogenic effect than other stimulants
 - Also a Hallucinogen
 - Different experiences across individuals (hence "Other" classification)
- Primary precursor is Safrole, a light oily liquid extracted from the root-bark or the fruit of the sassafras tree.
 - numerous methods for synthesising safrole into MDMA.
- 1912: first synthesized by the German chemist, Anton Köllisch.
 - Employed by Merck to develop a haemostatic drug to stop abnormal bleeding (Bernschneider-Reifet. al., 2006)



Anton Köllisch

MDMA

- 1927: Merck scientist, Max Oberlin, reports MDMA's similarity to ephedrine.
- 1953: US military commissions studies with a range of psychoactive compounds including MDMA.
- 1958: independent synthesis of the same compound in Japan by Yutaka Kasuya.
- 1970: First report of recreational MDMA use in the US.
- 1970s: Pharmacist, Alexander Shulgin, synthesizes MDMA & publishes reports of its mood enhancing effects & improvement of therapeutic engagement.
- 1980s: MDMA becomes increasingly popular in dance/rave culture.
- 1985s: State prohibition of the compound is introduced.
- 2010s: MDMA usage decreasing due to scarcity of precursor chemical
 - academic documents to side-step prohibitive laws against of existing drugs.



Alexander Shulgin

3

Cannabis

- Believed to have originated in the mountainous regions northwest of the Himalayas.
- · Two varieties:
 - · Sativa: greater fibre content useful for making woven materials
 - high psychotomimetic effect
 - Indica: lower fibre content
 - low psychotomimetic effect
 - It is thought these two varieties were separated from the wild type through selective breeding
- Native populations made use of the plants for intoxication, clothes, nets, rope and string

Small & Cronquist (1976)



Cannabis

- 1000 BC: evidence for the earliest cannabis consumption
 - burial site in Turpan, North-West China.
 - The 'shaman' had 789g of cannabis on his person.
 - Site is especially important because it is the earliest recorded site in which researchers were able to conclusively demonstrate the presence of THC RUSSO et al. (2008)





5

Cannabis

- Earlier texts refer to the medicinal/intoxicant/spiritual effects of cannabis.
 - 2700 BC: Chinese herbal medicine the Pên-ts'ao Ching
 - 2000 BC: religious/philosophical text the Hindu Vedas.
 - Later references to cannabis in Japanese, West Asian, Middle Eastern, North African, Roman & Germanic texts
 - chart the spread of cannabis use across the ancient world following the trade routes of the steppe highway/silk road



Chinese Pên-ts'ao Ching

Cannabis

 Contemporary cultivation of cannabis is global (UNODC, 2014)

Country	Cultivated (ha)	Eradicated (ha)	Harvest- able (ha)	Production (tons)		Plants eradicated		Sites eradicated	
				Indoor	Outdoors	Indoors	Outdoors	Indoors	Outdoors
Afghanistan	10,000				1,400				
Albania					50		33,000		154
Australia						17,668	35,146	322	240
Azerbaijan	6	6	0		308		7,538		121
Bosnia and Herzegovina							2,807		3
Brazil		22			185		616,133		5
Bulgaria						6,913		42	
Chile						18,526	216,902	1,377	291
Costa Rica	8	8	0				965,320		129
Italy						7,706	4,114,911	458	1,318
Latvia						3,796	101	4	3
Lebanon	3,500	800	2,700						
Mexico		9,058			12,166				
Morocco	52,000	5,000	47,000	760	38,000				
New Zealand						21,202	119,059	783	
Philippines		21					1,224,738		188
Poland		4					58,156	687	627
Tajikistan							2,180,121		
Ukraine	529						2,200,000		
United States of America						302,377	3,631,582	2,596	6,470

7

Cannabis

- Cannabinoids are the main psychoactive component within the cannabis leaf.
- Phytocannabinoids: found in plants
- Synthetic cannabinoids: man made from other compounds.
- Around 85 phytocannabinoids have been isolated from the cannabis plant
 most research has focused on Δ9-tetrahydrocannabinol (THC) & cannabidiol (CBD).
- THC: primary psychoactive intoxicant, as indicated by the level of THC dose being associated with the subjectively reported drug effect (Miller et al. Pharmacol Biochem Behav 9, 573)
- The supposed potency of different forms of the drug have different THC concentrations:
 - Cannabis herb (leaves) 5%
 - Flowers (buds) 12%
 - Resin (Hashish) 20%
 - Oils (Hash oil) 60% (UNODC, 2009)
- There is initial evidence that although CBD has little psychoactive effect it may reduce the psychotomimetic effect of THC (Joy et al. 1999)



Cannabis as a therapeutic

(Grotenhermen et al. 2012)

Indication	Number of randomized controlled trials (some three-armed)	Positive studies	Negative studies	
Spasticity	n = 12 (dronabinol: [e1, e2, e4–e6]; cannabis: [e1–e3, e6–e12]) in multiple sclerosis	n = 9 (e4-e12)	n = 3 (e1-e3)	
	n = 3 (dronabinol: [e13–e14]; nabilone: [e15] in paraplegia)	n = 3 (e13-e15)	-	
Nausea and vomiting due to cytostatics	n = 41 (dronabinol: [e16-e34]; cannabis cigarettes: [e25]; cannabis extract: [e35]; nabilone: [e36-e52]; levonantradol: [e53-e56])	n = 40	n = 1 (e18)	
Loss of appetite/weight loss	n = 7 (dronabinol: [e59–e65]; cannabis cigarettes: [e63–e65]) in HIV/Aids	n = 7		
	n = 4 (dronabinol: [e66–e68]; cannabis extract: [e69]) in various tumor diseases	n = 3	n = 1 (e69)	
	n = 1 (dronabinol: [e70]) in Alzheimer's disease	n = 1	-	
Chronic pain	n = 14 (dronabinol: [e71–e74]; nabilone: [e75, e76]; cannabis extract: [e73, e74, e77–e79]; cannabis cigarettes: [e80–e83]; CT3 (ajulemic acid): [e84]) in neuropathic pain or pain in MS	n = 12 (e71, e73-e75, e77-e84)	n = 2 (e72, e76)	
	n = 12 (dronabinol: [e85–e87, e93]; NIB: [e88]; benzopyrano- peridine: [e89]; cannabis extract: [e87, e90, e94]; nabilone: [e91, e92, e96]; cannabis oigarettes: [e95]) in chronic pain (cancer, rheumatism, fibromyalgia)	n = 11 ([e85, e86, e87] cannabis ex- tract, [e88, e90–e96])	n = 2 ([e87] drona binol, [e89])	

^{*1} A complete list of clinical trials of cannabis medications can be found on the website of the IACM (24)

9

Medicinal Cannabis

- Still a prohibited substance in Australia although differences in state laws exist
 - Drug laws in Australia
 - NSW Drug misuse and trafficking Act
- But, can be legally used as a therapeutic in some cases and grown for medical/research use

 - TGA Special Access Scheme Medical cannabis cultivation
- Efficacy for treating:
 - Pain
 - Muscle spasticity
 - Pediatric epilepsy
 - Nausea/vomiting
 - Appetite stimulant
- Research is ongoing...



Synthetic Cannabis

- John William Huffman, organic chemist
 - synthesised a whole series (JWH-018)
 - funded by the National Institute on Drug Abuse (NIDA)
- Prof. Raphael Mechoulam, Hebrew University
 - developed HU-210
- Pfizer developed CP 47,497
- These synthetic cannabinoids are considered designer drugs
 - mixed herbs infused with synthetic cannabinoids.



11

Topic Summary

- Knowledge of historical origins of various depressant, stimulant and other drugs.
- Understanding of drug discovery process, natural vs. synthetic compounds and manufacture.
- Recreational use and abuse aside, drugs have a wide range of therapeutic applications.
- Substance use is embedded in human culture and has been since antiquity.

