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Cannabis-Based Products

• Medicinal cannabis products contain tetrahydrocannabinol (THC) and/or cannabidiol (CBD) in differing concentrations.

• THC and CBD are pharmacologically distinct: they do not have the same mechanisms of action or clinical effects (beneficial or

adverse). THC is the main psychoactive constituent responsible for the 'high' derived from cannabis, whereas CBD is less psychoactive.
All medicinal cannabis products are subject to the requirements of Aotearoa's <u>Medicinal Cannabis Scheme</u> and must meet <u>minimum</u> <u>quality standards</u>. The Ministry of Health maintains a <u>list of products</u> that meet these standards, which can be prescribed by any doctor. Meeting the standard does not mean they have been approved by Medsafe; safety and efficacy have not been assessed for any product unless it has been approved. Currently the only approved products are Sativex[®] spray (CBD+THC) and Epidyolex[®] liquid (CBD).

Overall, the evidence for efficacy of cannabis-based products is very limited, and none are funded.

	THC	CBD
Indications	Spasms in multiple sclerosis	Paediatric epilepsy syndromes
Evidence	Moderate evidence of slight benefit over placebo in RCTs.	Moderate evidence of benefit in RCTs.
Dose	Sativex [®] spray: typical starting dose 1-2 sprays/day with	Epidyolex [®] liquid: 2.5-10 mg/kg BD in trials;
	maintenance dose in trials 8 sprays/day (maximum 12).	however, specialists often use smaller doses
		in clinical practice.
	Chemotherapy-induced nausea and vomiting	Anxiety
	Weak evidence for benefit over placebo but no better than	Weak evidence for benefit in RCTs and case
	conventional antiemetics.	series.
	Starting doses in trials 2.5-5 mg BD, titrated up to 20 mg/day	
	(given 3-4 times daily).	
	Chronic pain	Pain
	Weak evidence for slight benefit in RCTs.	RCTs have not shown benefit over placebo.
	Starting dose in trials 2.5 mg BD, titrated up to 15 mg BD.	
Common adverse	Incidence >10%	Incidence >10%
reactions	Drowsiness, dizziness, dry mouth, anxiety, nausea and	Drowsiness, diarrhoea or vomiting, loss of
	vomiting, impaired concentration, memory and coordination	appetite, weight loss, increased
	Incidence 1-10%	transaminases
	Euphoria, blurred vision, increased appetite	
Pharmacokinetics	Metabolised by CYP2C9, 2C19 and 3A.	Metabolised by CYP3A and 2C19, and
	Half-life 25-36 hours.	UGT1A7/9 and 2B7 (to active metabolite).
		Half-life 60 hours.
Interactions	Additive CNS adverse effects with other CNS depressants.	May inhibit CYP2C19, 2C8/9, 3A4/5 and 1A2
	THC concentrations may be increased or decreased by CYP2C9,	P-glycoprotein, and UGT1A9 and 2B7, and
	2C19 and 3A inhibitors or inducers, respectively.	therefore may increase concentrations of
		substrates for these enzymes/transporters.
		CBD concentrations may be increased or
		decreased by CYP3A and 2C19 inhibitors or
		inducers, respectively.
Legislation	Cannabis oral liquids/spray	Oral liquids/spray
	THC >2% of total product (with or without CBD): Class B1 CD.	Prescription medicine: not a CD.
	Prescribe on CD prescription.	Prescribe on conventional prescription form
	90-day period of supply dispensed monthly.	90-day period of supply and can dispense
	Store in CD cabinet (except for Sativex [®] : store in fridge).	stat.
	Record in CD register.	Store on dispensary shelf.
	Dried cannabis plant	
	THC and CBD: Class C1 CD.	
	Prescribe on conventional prescription form.	
	90-day period of supply dispensed monthly.	
	Store in CD cabinet.	
	Best practice to record in CD register but no legal requirement.	

RCTs = Randomised Controlled Trials; CNS = Central Nervous System; CD = Controlled Drug



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