

Differences in prescribed medicinal cannabis use by cannabinoid product composition: Findings from the cannabis as medicine survey 2020 Australia-wide study

Presented by Benjamin Trevitt

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Acknowledgement of country

I would like to acknowledge the Traditional Custodians of the lands on which we are meeting here today:

I pay my respects to Elders past, present and emerging and celebrate the diversity of Aboriginal peoples and their ongoing cultures and connections to the lands and waters of Australia.

I also would like to acknowledge and pay my respects to our Aboriginal and Torres Strait Islander colleagues joining us here today.



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Background (1)

- Medical practitioners have been able to legally prescribe cannabinoids since November 2016 using the compassionate access regulatory pathways (Special Access (SAS) and Authorised Prescriber Schemes)
- Guidelines are available which highlight the evidence available for using prescribed medicinal cannabis to treat a range of conditions (e.g. chronic pain, palliative care, chemotherapy and epilepsy)

BUT.....

- **In practice, clinicians can prescribe medicinal cannabis for ANY clinical reason as long as they can provide some justification to the regulatory body based on available evidence**



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Background (2)

- Prescribed medicinal cannabis products consist of two major cannabinoid products:
 - Tetrahydrocannabinol (THC)
 - Cannabidiol (CBD)
- THC and CBD differ considerably in terms of their:
 - Pharmacological actions
 - Clinical rationale
 - Safety
 - Effectiveness



Background (3)

- **Primary aims of the study**

- i) To describe the demographics of prescribed medicinal cannabis consumers by cannabinoid product composition
- ii) To examine how patient-reported cost, patterns and reasons for use differ based on product composition
- iii) To compare patient reported effectiveness and side-effect profiles of the prescribed products
- iv) To review the safety profiles of these prescribed products based on current medication use and driving patterns



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Methods (1)

- **Study Type:**
 - Online cross-sectional survey. Data was collected via REDCap
- **Eligibility:**
 - To be eligible for inclusion participants had to:
 - i) Be Australian adults
 - ii) Used cannabis for therapeutic or medical reasons within last 12 months
- **Excluded:**
 - Participants who:
 - i) Identified as only using illicit medicinal cannabis
 - ii) Reported prescribed medicinal cannabis varied between batches/unaware of its constituents



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Methods (2)

- **Variables of interest were:**
 - Demographic characteristics
 - Primary medical condition treated by the prescribed cannabinoid
 - Side-effects
 - Improvement in symptoms and concurrent use of other medications over last 12 months
 - Cost of prescribed medicinal cannabis products
 - Time between prescription cannabinoid consumption and driving



Methods (3)

- **Participants were grouped based on prescribed cannabinoid product's composition:**
 - i) **THC-dominant group**
 - These participants received prescribed cannabinoid products containing >98% THC
 - ii) **Mixed THC-CBD group**
 - These participants received prescribed cannabinoid products containing >2% THC AND >2% CBD
 - iii) **CBD-dominant group**
 - These participants received prescribed cannabinoid products containing >98% CBD

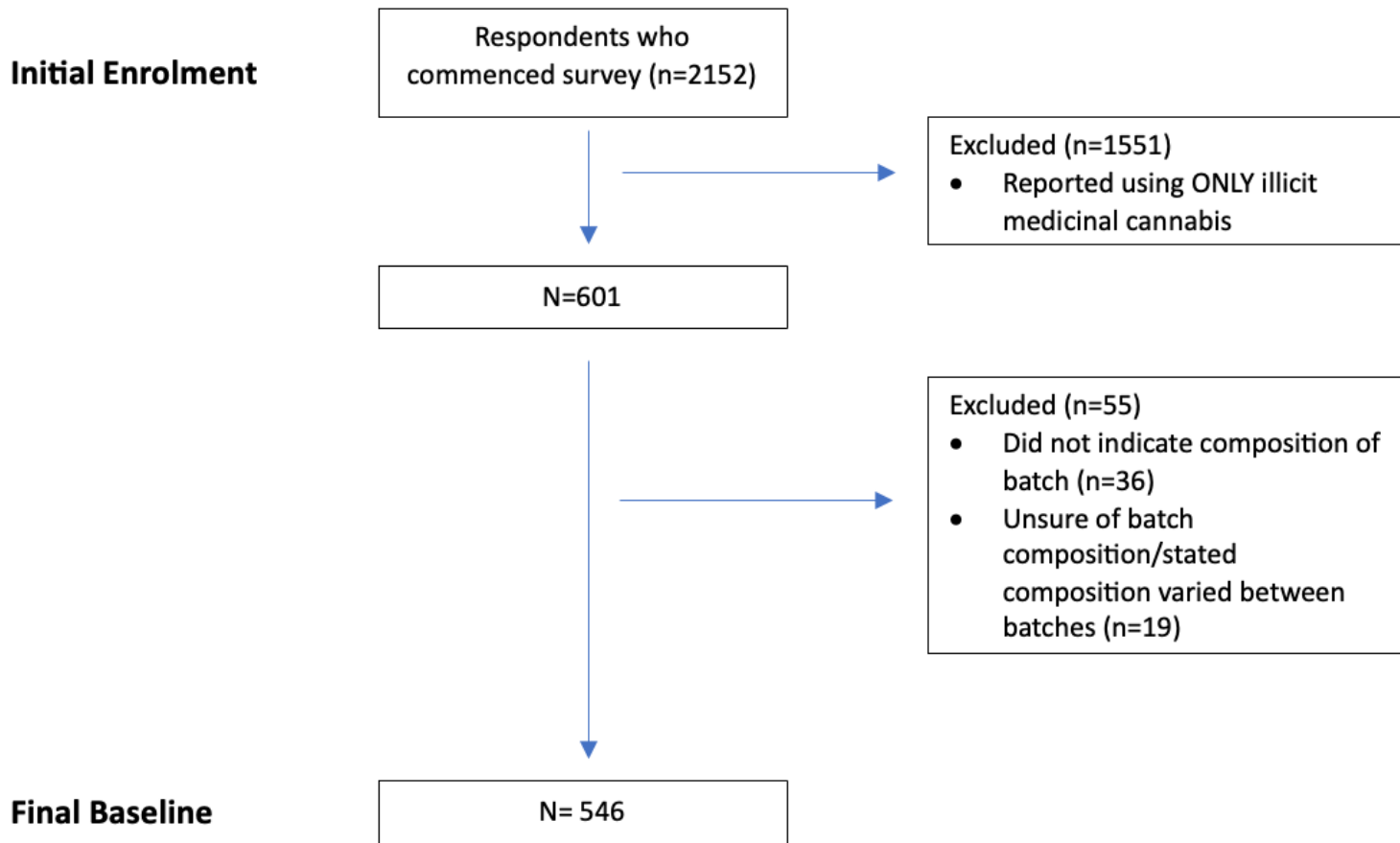
Methods (4)

- We used various regression models to analyse the data
 - i) ***Gaussian models*** for continuous outcomes (e.g. age, weekly cost of prescribed medicinal cannabis, frequency of use)
 - ii) ***Binary logistic regression models*** for binary categorical outcomes (e.g. relationship status and past use of illicit cannabis)
 - iii) ***Multinomial logistic regression models*** for multilevel outcomes (e.g. route of administration)
 - iv) ***Ordinal regression models*** for ordinal outcomes (e.g. change in symptoms following treatment with prescribed medicinal cannabis)
- Primary predictor of interest was prescribed medicinal cannabis composition



Results: Participation

Figure 1: Flow chart of eligibility requirements for inclusion in the study



Results: Characteristics of prescribed MC users by cannabinoid product composition

Table 1: Characteristics of prescribed medicinal cannabis users by cannabinoid product composition

Characteristic	THC only (TO) (n=144)	Mixed (T+C) (n=227)	CBD only (CO) (n=175)	Total (n=546)	Coefficient	Comparisons estimate (95% CI)	p values
Age, numeric, in y, M (SD)	42.5 (13.2)	46.3 (13.6)	47.5 (14.9)	45.6 (14.1)	Beta	TO-T+C: -3.8 (-7.4, -0.2) TO-CO: -5.0 (-8.8, -1.2) T+C - CO: -1.2 (-4.6, 2.2)	0.005 0.03 1.00
Gender, binary ^b (not-male [ref] vs male), n (%) male	97 (67%)	133 (59%)	56 (32%)	286 (52%)	Odds ratio	TO-T+C: 1.5 (0.9, 2.5) TO-CO: 4.4 (2.5, 7.8) T+C-CO: 3.0 (1.8, 5.0)	0.27 <0.001 <0.001
Highest level of education, binary, (school [ref] vs tertiary) n (%) Tertiary	110 (76%)	189 (84%)	140 (80%)	439 (80%)	Odds ratio	- TO-T+C: 0.7 (0.3, 1.2) TO-CO: 0.8 (0.4, 1.6) T+C-CO: 1.2 (0.7, 2.3)	0.31 1.00 1.00
Employment, binary ^c , not-employed [ref] vs employed, n (%) employed	61 (42%)	114 (50%)	90 (51%)	265 (49%)	Odds ratio	TO-T+C: 0.7 (0.4, 1.2) TO-CO: 0.7 (0.4, 1.2) T+C-CO: 1.0 (0.6, 1.5)	0.42 0.32 1.00
Used non-medical cannabis regularly prior to commencing MC, binary, No [ref] vs Yes, n (%) Yes	118 (94%)	194 (94%)	135 (85%)	447 (91%)	Odds ratio	TO-T+C: 1.1 (0.4, 3.6) TO-CO: 3.0 (1.0, 8.8) T+C-CO: 2.7 (1.1, 6.3)	1.00 0.04 0.02
Weekly Cost of prescribed medicinal cannabis ^e , numeric, in \$AU, M (SD)	113.7 (78.4)	68.8 (53.1)	74.2 (49.2)	79.6 (59.1)	Beta	TO-T+C: 45.0 (20.8, 69.1) TO-CO: 39.6 (16.2, 63.0) T+C-CO: -5.4 (-24.4, 13.6)	<0.001 <0.001 0.58
Route of Administration, categorical ^f , oral [ref], n (%) Vaporiser	69 (50%)	45 (20%)	2 (1%)	116 (22%)	Odds ratio	TO-T+C: 4.8 (3.0, 7.8) TO-CO: 109.2 (25.91, 460.42) T+C-CO: 22.8 (5.4, 95.3)	<0.001 <0.001 <0.001
Inhaled	16 (12%)	12 (5%)	0 (0%)	28 (5%)		TO-T+C: 4.2 (1.9, 9.4) TO-CO: <i>Not enough data</i> T+C-CO: <i>Not enough data</i>	0.001 NA NA
Frequency of use, numeric, days used in past 28 days, M (SD)	22.8 (9.2)	23.3 (8.4)	22.5 (9.4)	22.9 (8.9)	Beta	TO-T+C: -0.5 (-2.9, 1.9) TO-CO: 0.3 (-2.2, 2.9) T+C-CO: 0.8 (-1.5, 3.1)	1.00 1.00 1.00

Results: Primary reasons for prescribed MC

Table 2: Main reasons for use of prescribed medicinal cannabis users by cannabinoid product production composition

Indication	THC only (TO) (n=144)	Mixed (T+C) (n=227)	CBD only (CO) (n=175)	Comparisons estimate OR (95% CI) ^r	Adjusted P value
Pain (total), binary, No [ref] vs Yes, n(%) Yes (n=263)	75 (29%)	111 (42%)	77 (29%)	TO-T+C: 1.1 (0.6,1.8) TO-CO: 1.3 (0.7,2.3) T+C-CO: 1.2 (0.7,2.6)	1.00 0.73 0.92
MHSU^c (total), binary, No [ref] vs Yes, n(%) Yes (n=133)	32 (24%)	52 (39%)	49 (37%)	TO-T+C: 0.9 (0.5,1.7) TO-CO: 0.7 (0.4,1.3) T+C-CO: 0.8 (0.4, 1.3)	1.00 0.49 0.70
Neurological disorder, binary, No [ref] vs Yes, n(%) Yes (n=34)	7 (21%)	16 (47%)	11 (32%)	TO -T+C: 0.6 (0.2,2.0) TO-CO: 0.7 (0.2,2.4) T+C-CO: 1.1 (0.4, 3.0)	1.00 1.00 1.00
Sleep, binary, No [ref] vs Yes, n (%) Yes (n=29)	11 (38%)	9 (31%)	9 (31%)	TO -T+C: 1.9 (0.6,5.9) TO-CO: 1.5 (0.5,4.5) T+C-CO: 0.8 (0.2, 2.4)	0.47 1.00 1.00
Gastrointestinal disorder, binary, No [ref] vs Yes, n (%) Yes (n=8)	1 (13%)	5 (63%)	2 (25%)	TO -T+C: 0.3 (0.02,4.2) TO-CO: 0.6 (0.03,11.1) T+C-CO: 2.0 (0.3, 14.7)	0.82 1.00 1.00
Cancer, binary, No [ref] vs Yes, n (%) Yes (n=9)	2 (22%)	6 (67%)	1 (11%)	TO -T+C: 0.5 (0.07,3.6) TO-CO: 2.4 (0.1,44.9) T+C-CO: 4.7 (0.4, 63.6)	1.00 1.00 0.46
Other, binary, No [ref] vs Yes, n (%) Yes (n=22)	7 (32%)	7 (27%)	9 (41%)	TO -T+C: 1.6 (0.4,5.9) TO-CO: 0.9 (0.3,3.2) T+C-CO: 0.6 (0.2, 2.0)	1.00 1.00 0.90

- Pain and MHSU were the two most common primary reasons indicated by participants for prescribed medicinal cannabis
- There were no significant differences in likelihood of patients being prescribed any of the three MC compositions to manage above umbrella presentations

Results: Side effects

Table 3: Most commonly reported side effects in participants receiving prescribed MC

Side Effect	THC containing products (n=313)	CBD dominant products (n=152)	Total (n=465)	Coefficient	Comparisons estimate (95% CI)	p-values
Dry Mouth , No [ref] vs Yes, n(%)	197 (63%)	64 (42%)	261 (56%)	Odds Ratio	THC-CO: 2.3 (1.6, 3.5)	<0.001
Drowsiness , No [ref] vs Yes, n(%)	179 (57%)	60 (39%)	239 (51%)	Odds Ratio	THC-CO: 2.0 (1.4,3.0)	<0.001
Fatigue , No [ref] vs Yes, n(%)	88 (28%)	41 (27%)	129 (28%)	Odds Ratio	THC-CO: 1.1 (0.7, 1.6)	0.80
Eye Irritation , No [ref] vs Yes, n(%)	92 (29%)	21 (14%)	113 (24%)	Odds Ratio	THC-CO: 2.6 (1.5, 4.4)	<0.001
Anxiety , No [ref] vs Yes, n(%)	55 (18%)	32 (21%)	87 (19%)	Odds Ratio	THC-CO: 0.8 (0.5, 1.3)	0.37
Dizziness , No [ref] vs Yes, n(%)	61 (19%)	21 (14%)	82 (18%)	Odds Ratio	THC-CO: 1.5 (0.9, 2.6)	0.14
Bad Taste , No [ref] vs Yes, n(%)	47 (15%)	24 (15%)	71 (15%)	Odds Ratio	THC-CO: 0.9 (0.6, 1.6)	0.83
Confusion , No [ref] vs Yes, n(%)	42 (13%)	17 (11%)	59 (13%)	Odds Ratio	THC-CO: 1.2 (0.7, 2.2)	0.50

- Dry mouth and drowsiness were the two most commonly reported side-effects.
- Participants on prescribed THC-containing products were significantly more likely to experience dry mouth, drowsiness and eye irritation than those on CBD-dominant products

Results: Perceived effect of prescribed medicinal cannabis on main condition

Table 4: Reported improvement in main condition: comparing prescribed cannabinoids using PGIC scores

Degree of Improvement in main condition	THC only (TO) (n=144)	Mixed (T+C) (n=227)	CBD only (CO) (n=175) ^a	Odds of a greater degree of improvement	Adjusted p values
Pain , numeric, M (SD)	6.1 (0.8)	6.1 (0.9)	5.8 (0.9)	TO-T+C: 0.6 (0.3, 1.3) TO-CO: 1.3 (0.6,2.9) T+C-CO: 2.2 (1.1,4.7)	0.28 1.00 0.03
MHSU^b numeric, M (SD)	6.4 (0.7)	6.2 (1.0)	5.9 (1.0)	TO-T+C: 1.5 (0.5,4.1) TO-CO: 3.1 (1.1,9.0) T+C-CO: 2.1 (0.9,5.3)	1.00 0.03 0.14
Anxiety , numeric, M (SD)	6.3 (0.7)	6.5 (0.6)	5.9 (0.9)	TO-T+C: 0.6 (0.2,2.3) TO-CO: 2.8 (0.8,10.3) T+C-CO: 4.5 (1.4,14.9)	1.00 0.15 0.007
Depression , numeric, M (SD)	5.8 (0.5)	6.2 (0.8)	6 (0)	TO-T+C: 0.2 (0.0,7.5) TO-CO: 0.5 (0.0,71.8) T+C-CO: 2.2 (0.0, 339.4)	0.89 1.00 1.00
Neuro , numeric, M (SD)	6.1 (0.7)	6.1 (0.9)	5.8 (0.8)	TO-T+C: 0.9 (0.1,14.0) TO-CO: 2.1 (0.3,17.5) T+C-CO: 2.3 (0.4,14.0)	1.00 1.00 0.79
Sleep , numeric, M (SD)	6.3 (0.7)	6.7 (0.5)	5.3 (1.4)	TO-T+C: 0.9 (0.1,8.2) TO-CO: 5.6 (0.6,48.1) T+C-CO: 6.3 (0.6,61.9)	1.00 0.17 0.17
Back pain , numeric, M (SD)	6.1 (0.6)	6.0 (0.8)	5.6 (1.1)	TO-T+C: 1.2 (0.3,4.6) TO-CO: 2.9 (0.5, 17.9) T+C-CO: 2.3 (0.4,14.0)	1.00 0.49 0.79
Arthritis , numeric, M (SD)	5.8 (1.1)	6.2 (0.7)	6.2 (0.7)	TO-T+C: 0.5 (0.1,3.6) TO-CO: 0.5 (0.1, 3.3) T+C-CO: 1.0 (0.3,4.0)	1.00 1.00 1.00

- 95% of participants reported improvements in pain (n=209); 100% in depression (n=10); 97% in anxiety (n=85) and 83% in sleep (n=30)
- Patients prescribed THC containing products were more likely to report improvements in pain, MHSU and anxiety than those prescribed CBD-dominant products



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Results: Concurrent use of other medications versus prescribed cannabinoid

Table 5: Concurrent medications used versus prescribed cannabinoids

Other medication use	THC only (TO) (n=144)	Mixed (T+C) (n=227)	CBD only (CO) (n=175) ^a	Total (n=546)	Comparisons estimate (95% CI)	Adjusted p value
Opioids , binary, No [ref] vs Yes, n(%)	78 (54%)	142 (63%)	90 (51%)	310 (56%)	TO-T+C: 0.7 (0.4,1.2) TO-CO: 1.1 (0.7,1.9) T+C-CO: 1.6 (1.0,2.6)	0.33 1.00 0.05
Benzodiazepines , binary, No [ref] vs Yes, n(%)	74 (51%)	111 (49%)	81 (46%)	266 (50%)	TO-T+C: 1.1 (0.7,1.8) TO-CO: 1.2 (0.7,2.1) T+C-CO: 1.1 (0.7,1.8)	1.00 1.00 1.00
Antidepressants , binary, No [ref] vs Yes, n(%)	80 (56%)	114 (50%)	93 (53%)	287 (53%)	TO-T+C: 1.2 (0.7,2.1) TO-CO: 1.1 (0.6, 1.9) T+C-CO: 0.9 (0.5,1.4)	0.95 1.00 1.00
Antipsychotics , binary, No [ref] vs Yes, n(%)	30 (21%)	25 (11%)	16 (9%)	71 (13%)	TO-T+C: 2.1 (1.0,4.3) TO-CO: 2.6 (1.2,5.8) T+C-CO: 1.2 (0.5, 2.8)	0.03 0.01 1.00
Anticonvulsants , binary, No [ref] vs Yes, n(%)	16 (11%)	17 (7%)	17 (10%)	50 (9%)	TO-T+C: 1.5 (0.6, 3.7) TO-CO: 1.2 (0.5,2.8) T+C-CO: 0.8 (0.3,1.8)	0.71 1.00 1.00
Gabapentinoids , binary, No [ref] vs Yes, n(%)	33 (23%)	50 (22%)	34 (19%)	117 (21%)	TO-T+C: 1.1 (0.6,1.9) TO-CO: 1.2 (0.6,2.4) T+C-CO: 1.2 (0.6,2.1)	1.00 1.00 1.00
Non-opioid analgesics , binary, No [ref] vs Yes, n(%)	72 (50%)	126 (56%)	88 (50%)	285 (52%)	TO-T+C: 0.8 (0.5, 1.3) TO-CO: 1.0 (0.6,1.7) T+C-CO: 1.2 (0.8,2.0)	0.90 1.00 0.90
Other , binary, No [ref] vs Yes, n(%)	11 (8%)	10 (4%)	12 (7%)	33 (6%)	TO-T+C: 1.8 (0.6,5.3) TO-CO: 1.1 (0.4,3.2) T+C-CO: 0.6 (0.2,1.8)	0.58 1.00 0.86

Across each of the three groups:

- $\geq 50\%$ used concurrent opioids
- $\geq 45\%$ used concurrent benzodiazepines
- $\geq 50\%$ used concurrent antidepressants
- $\geq 50\%$ used concurrent non-opioid analgesics

Participants prescribed THC dominant products were more likely to also be prescribed antipsychotics than participants on CBD-dominant products

- Participants prescribed mixed THC-CBD products were more likely to be prescribed opioids than participants on CBD-dominant products



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Results: Driving patterns versus prescribed cannabinoid

Table 6: Driving patterns of participants prescribed products containing THC vs prescribed products containing CBD only

Driving activities	THC containing products (n=241)	CBD only products (n=123)	Total (n=364)	Coefficient	Comparisons estimate (95% CI)	p value
Driving within 24 hours of consumption, No [ref] vs Yes, n(%)	210 (87%)	113 (92%)	323 (89%)	Odds Ratio	0.6 (0.3,1.3)	0.18
Driving within 12 hours of consumption, No [ref] vs Yes, n(%)	180 (75%)	98 (80%)	278 (76%)	Odds Ratio	0.8 (0.4,1.3)	0.29
Driving within 6 hours of consumption, No [ref] vs Yes, n(%)	117 (49%)	69 (56%)	186 (51%)	Odds Ratio	0.7 (0.5, 1.2)	0.17
Driving within 3 hours of consumption, No [ref] vs Yes, n(%)	85 (35%)	57 (46%)	142 (39%)	Odds Ratio	0.6 (0.4,1.0)	0.04
Driving within 1 hour of consumption, No [ref] vs Yes, n(%)	36 (15%)	40 (33%)	76 (21%)	Odds Ratio	0.4 (0.2,0.6)	<0.001

- No difference in odds of participants prescribed THC-containing products vs CBD-dominant products driving within 24, 12, or 6 hours of consumption
- Those prescribed THC containing products were significantly less likely to drive within 3 or 1 hour of MC consumption than those prescribed CBD dominant products



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Discussion (1)

- Compared to those prescribed CBD-dominant products, individuals prescribed THC dominant products were:
 - Younger
 - More likely to be male
 - More likely to consume via inhalation
 - Tended to spend more money on prescribed MC per week
 - No difference in frequency of use



Discussion (2)

- Systematic reviews conducted by the TGA have found:
 - THC and CBD are effective in treatment chronic non-cancer related pain and neuropathic pain
 - *But tend to favour THC containing products*
 - CBD used as an adjunct to antiepileptic medications may reduce seizure frequency in people under 25



Discussion (3)

- Recent RCT conducted in Australia found evidence supporting cannabinoids in treating chemotherapy induced N&V
- However, evidence for role of cannabinoid to treat MH and palliative care conditions is limited.
- **Almost half the participants received prescribed MC products for conditions other than pain, epilepsy and chemotherapy induced N&V**



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Discussion (4)

- Unlike CBD, THC is well known for its increased risk of impairment, particularly when combined with other sedatives

BUT....

- Patients on prescribed opioids and/or antipsychotics were significantly more likely to be receiving THC containing products than CBD-dominant products



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Discussion (5)

- Unlike THC, the primary concern with CBD is its risk of interaction with CYP450 metabolised drugs (e.g. many benzodiazepines)

BUT....

- Almost 50% of patients on prescribed CBD were receiving concurrent benzodiazepines



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Limitations

- Data is self reported and thus there may be inaccuracies around diagnostic conditions, reported effectiveness and reported adverse events
- Was collected by an anonymous survey - sample may not be representative of prescribed MC users generally



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Conclusion

- Compared to CBD-dominant consumers, consumers of prescribed THC-dominant medical cannabis products tend to be:
 - Younger
 - More likely to have used illicit cannabis in past
- Products containing THC were favoured over CBD-dominant products for management of
 - Pain
 - Mental Health
 - Sleep
- Crucial to educate medical practitioners on evidence based reasons for cannabinoid product prescription + potential drug interactions



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Questions?



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