

Arthritis Talks Medical Cannabis: What's new and changing

Dr. Jason Busse McMaster University Hamilton, Ontario

November 9, 2022

Presenters



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Arthritis Society
(Moderator)



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Hamilton, Ontario



Webinar tips

- Use the Q&A section to ask the presenters your questions. Some of the questions will be chosen for the live question period at the end of the webinar.
- Click on the Chat box to connect with other participants and the Arthritis Society's chat moderator.
- If you have further issues, email arthritistalks@arthritis.ca



question



Overview

[1]
New Evidence for Medical
Cannabis and Chronic Pain



[2] Health Canada Guidelines



[3] Questions & Answers





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What's the latest information and research related to medical cannabis for people affected by arthritis?





Background

- Cannabis has been legally available for select medical conditions in Canada since 2001.
- Licensed healthcare practitioners can provide authorization for patients to acquire medical cannabis.
- ▼ The number of Canadians authorized to use medical cannabis increased from <u>23,930</u> in June 2015 to <u>377,024</u> by September 2020.





Original Research

Risks and Benefits of Marijuana Use A National Survey of U.S. Adults

Salomeh Keyhani, MD, MPH; Stacey Steigerwald, MSSA; Julie Ishida, MD, MAS; Marzieh Va Deborah Hasin, PhD; Camille Dollinger, BS; Sodahm R. Yoo, BS; and Beth E. Cohen, MD, M.

Background: Despite insufficient evidence regarding its risks and benefits, marijuana is increasingly available and is aggressively marketed to the public.

Objective: To understand the public's views on the risks and benefits of marijuana use.

Design: Probability-based online survey.

Setting: United States, 2017.

Participants: 16 280 U.S. adults.

Measurements: Proportion of U.S. adults who agreed with a statement.

Results: The response rate was 55.3% (n = 9003). Approximately 14.6% of U.S. adults reported using marijuana in the past year. About 81% of U.S. adults believe marijuana has at least 1 benefit, whereas 17% believe it has no benefit. The most common benefit cited was pain management (66%), followed by treatment of diseases, such as epilepsy and multiple sclerosis (48%), and relief from anxiety, stress, and depression (47%). About 91% of U.S. adults believe marijuana has at least 1 risk,

Annals of Internal Medicine

- 81% believe cannabis has health benefits
- 9% believe it has no risks
- 22% believe it is not addictive

that it is somewhat or complete the spondents, 7.3% agree the spanna use is somewhat or completely safe during megnancy. About 22.4% of U.S. adults believe that manuana is not at all addictive.

Limitation: Wording of the questions may have affected interpretation.

Conclusion: Americans' view of marijuana use is more favorable than existing evidence supports.

Primary Funding Source: National Heart, Lung, and Blood Institute.

Ann Intern Med. 2018;169:282-290. doi:10.7326/M18-0810 Annals.org
For author affiliations, see end of text.

This article was published at Annals.org on 24 July 2018.



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THE WALL STREET JOURNAL.

OPINION | COMMENTARY

If Weed Is Medicine, So Is Budweiser

Legalize marijuana, but don't pretend it's therapeutic.

By Peter B. Bach

Jan. 17, 2019 7:23 p.m. ET

Actual medicines have research behind them, enumerating their benefits, characterizing their harms, and ensuring the former supersedes the latter. Marijuana doesn't. It's a toxin, not a medicine. It impairs judgment and driving ability. It increases the risk of psychosis and schizophrenia. Smoking it damages the respiratory tract. A 2017



Cannabis legalization may be associated with reduced prescription drug use

Where medicinal cannabis is legal, U.S. states saw a significant reduction in the use of prescription drugs.

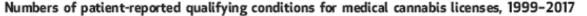
- E.g. anxiety, depression, nausea, pain, sleep disorders and spasticity
- Users may substitute cannabis for prescription medication, suggesting the possibility of therapeutic benefits

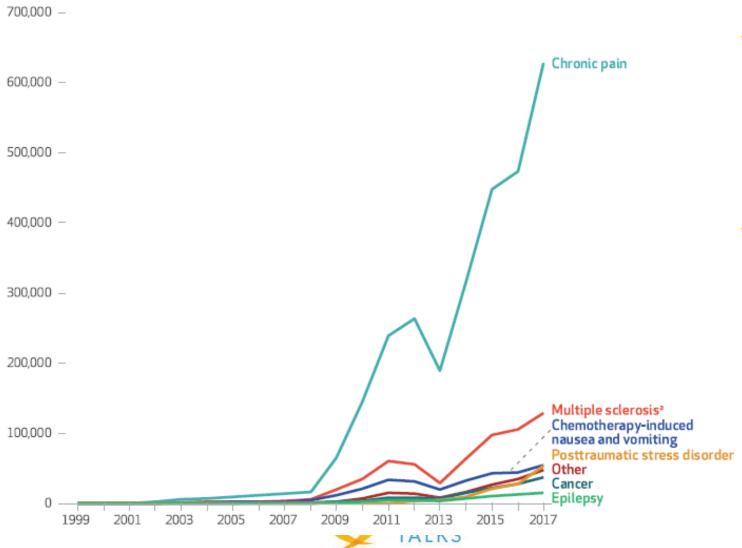




Qualifying Conditions Of Medical Cannabis License Holders In The United States

HEALTH AFFAIRS 38, NO. 2 (2019): 295–302 ©2019 Project HOPE— The People-to-People Health Foundation, Inc.

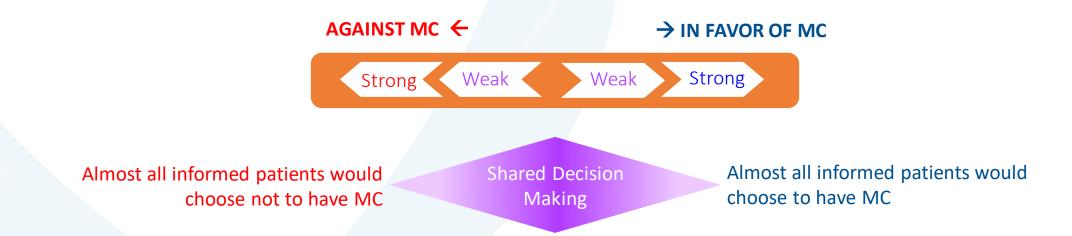




- 2017: 813,917
 medical cannabis
 licence holders in the
 U.S.
- Chronic pain was the qualifying condition reported by 62% of patients

Our clinical question

In adult and adolescent patients living with cancer or non-cancer chronic pain, should we recommend/suggest a trial of medical cannabis (MC) or cannabinoids vs. continued care without medical cannabis or cannabinoids?



Strong recommendations can be quality of care standards.





Strong recommendations

Weak recommendations

1. Clear imbalance



- benefits clearly outweigh risks/hassle/cost
- risk/hassle/cost clearly outweighs benefits
- 2. Sufficient certainty in estimates
- 3. Patients' values & preferences:
- almost all same choice



1. Close balance



- Close call between benefits and risks/hassle/cost
- Therefore, more preference-sensitive
- 2. Lower certainty in estimates
- 3. Patients' values & preferences:
- choice varies appreciably (or is very uncertain)





Guideline perspective



Patient-centered perspective

Not: funder-centred or societal-perspective

Guideline Panel

- 21 members
- Clinical experts
- Methodologists
- 3 patient partners living with chronic pain, including 1 veteran
- No financial or important intellectual conflicts of interest



Values and preferences

We conducted a systematic review for <u>studies that</u> reported the values and preferences of patients with chronic pain, who walues and preferences are values and preferences.

- 1) The relativ
- 2) The attitumethods,
- 3) Factors that influen

15 studies were eligible for review:

- All reported adult patient values and preferences (no carers, children or adolescents)
- 9 enrolled U.S. patients (2 UK, 2 Israel, 1 Canada, 1 Australia)
- All conducted from 2000 onwards
- Both cancer and non-cancer patients were represented

n routes, ingestion



Values and preferences: Results

Moderate- to High-Certainty Evidence

- Many valued the effectiveness of medical cannabis for symptom management ever experiencing adverse events related to concentration, memory or fatigue.
- Greater preference for high CBD (cannabidiol) or balanced ratios of TH of medicinal cannabis
- Cannabis use was influenced by both positive social consequences (family) and negative social consequences (e.g. stigma surrounding)
- Concerns about medical cannabis included side effects, addiction, tolerance strangely, and were related to unwillingness to use cannabis

Neither NICE nor the IASP systematically explored patients' values & preferences

Low-Certainty Evidence

- Highly variable values towards medical cannabis among people living with chronic pain
- Patients were motivated to use medical cannabis to reduce use of prescription medication, and felt it was 'safer' than opioids

Systematic review of randomized clinical trials (RCTs) of cannabis for chronic pain

- 32 eligible trials: 28 non-cancer and 4 cancer chronic pain
- Adult populations, median age among trials: 54
- Length of follow-up ranged from 28 to 140 days
- Placebo was the most common control
- Types of cannabis:
 - Endocannabinoids (PEA) 5
 - Phytocannabinoids 20
 - Synthetics 12
- Mode of administration: oral (16), spray (13), sublingual (1), transdermal (2)





Review of RCTs: Cannabis vs. placebo (benefits)

Event	Follow-up	Treatmen	t Effects	Certainty	Lay Summary	
(#RCTs, #pts)	(months)	placebo	cannabis	(GRADE)		
Pain, 10cm VAS (27 RCTs, 3,939 pts) MID: 1cm	1 - 4	952 (52%)	1,309 (62%)	Moderate	Cannabis probably results in a small increase in the proportion of patients	
		Risk difference +1	.0% (5% to 15%)	Due to inconsistency (I ² = 75%)		
		WMD -0.50cm (-0.75 to -0.25)		experiencing an important reduction in pain	
SF-36 Physical	1 - 4	294 (28%)	447 (32%)		Cannabis results in a very	
functioning subscale, 0-100 (16		RD +4% (0% to 7%)		High	small increase in the	
RCTs, 2,473 pts) MID: 10-points		WMD 1.57 (0.001 to 3.14)			proportion of patients experiencing improvement in physical functioning	
SF-36 Emotional	1 - 4	276 (31%) 403 (33%)				
functioning subscale, 0-100 (10		RD +2% (-2	% to 4%)	High	Cannabis does not improve	
RCTs, 2,115 pts) MID: 10-points		WMD 0.53 (-0.67 to 1.73)			emotional functioning	



Review of RCTs: Cannabis vs. placebo (penefits)

Event	Follow-up	Treatme	ent Effects	Certainty	Lay Summary	
(#RCTs, #pts)	(months)	placebo	cannabis	(GRADE)		
Role physical, 0-100 SF-36 subscale (7 RCTs, 1128 pts) MID: 10 points		1957 4 trials reporting		Cannabis does not improve role functioning		
Social functioning, 0- 100 SF-36 subscale (8 RCTs, 1405 pts) MID: 10 points	Rev	ients to maintain iew of 6 observa RY LOW certainty	nnabis does not improve social functioning			
Sleep quality, 0-10cm (9 RCTs, 2652 pts) MID: 1cm		oid dose among D, 95%CI -33 to	cannabis results in a small increase in the proportion of patients experiencing improved sleep quality			
Daily opioid dose (MED) (4 RCTs, 1359 pts)	1.25 – 1.75		to 1.63)	Very Low * (>20% LTFU) (<u>indirect</u>)	It is uncertain whether cannabis may reduce opioid use	





What are the risks associated with medical cannabis use?





Side effects

- **▼ Most often, THC is dose dependent and dissipate over time through tolerance.**
- **▼** Many can be prevented, or at least mitigated, with low-dose initiation and slow titration.
- Common adverse events include:
 - drowsiness/fatigue
 - dizziness
 - dry mouth
 - nausea
 - effects on cognitive function
 - deficits in motor
 - diarrhea



Event	Follow-up (months)	Treatmer	nt Effect	Certainty	Lay Summary	
(#RCTs, #pts)		placebo	cannabis	(GRADE)		
Cognitive		7 (1%)	21 (3%)		Cannabis probably	
impairment (5 RCTs, 1033 pts)	1.3 – 3.5	RD +2% (0.1% to 6%)		Moderate (imprecise)	results in a very small increase in the proportion of patients experiencing	
(5 NC13, 1033 pt3)		RR 2.39 (1.06 to 5.38)			cognitive impairment	
Vomiting		61 (6%)	117 (9%)	Moderate	Cannabis probably	
(9 RCTs, 2284 pts)	1-3.5	RD +3% (0.4% to 6%)		(imprecise)	results in a very small increase in the proportion of patients experiencing	
		RR 1.46 (1.07 to 1.99)			vomiting	
Drowsiness	1-3.5	53 (5%)	151 (9%)	Moderate	Cannabis probably	
(16 RCTs, 2553 pts)		RD +4% (2% to 8%)		(imprecise)	results in a very small increase in the proportion of patients experiencing	
pts)		RR 2.01 (1.44 to 2.81)			drowsiness	
Dizziness	1-1.9	79 (8%)	472 (16%)	Moderate	Cannabis probably increases the proportion	
<3m F/U (11 RCTs, 2270 pts)		RD +8% (4% to 12%)		(imprecise)	of patients experiencing dizziness at 1-2 months	
		RR 2.10 (1.56 to 2.60)			IIIOIILIIS	
3+m F/U (7 RCTs, 1595 pts)	3 - 4	63 (9%)	173 (42%)		Cannabis probably results in a substantial	
		RD +33% (21% to 50%)		Moderate (indirectness*)	increase in the proportion of patients experiencing dizziness	
		RR 4.64 (3.31 to 6.51)		(maneciness)	at 3-4 months	

Oral cannabis vs. placebo (adverse events)

Event	Follow-up	Treatmen	nt Effects	Certainty	Lay Summary	
(#RCTs, #pts)	(months)	placebo	cannabis	(GRADE)		
Impaired attention (7 RCTs, 895 pts)	1 - 4	4 (1%)	25 (4%)	Moderate	Cannabis probably results in a very	
		RD +3% (1% to 8%)		(imprecise)	small increase in the proportion of	
		RR 4.04 (1.67 to 9.74)			patients experiencing impaired attention	
Diarrhea (7 RCTs, 2005 pts)	1 - 4	30 (4%)	119 (8%)	Moderate	Cannabis probably results in a very	
		RD +4% (2% to 8%)		(imprecise)	small increase in the proportion of	
		RR 2.19 (1.49 to 3.22)			patients experiencing diarrhea	
Nausea (14 RCTs, 2877 pts)	1 - 4	208 (8%)	213 (13%)	Moderate	Cannabis probably results in a very	
		RD +5% (2% to 8%)		(imprecise)	small increase in the proportion of	
		RR 1.59 (1.2	28 to 1.99)		patients experiencing nausea	

^{* 1} RCT of topical cannabis reported "no adverse events"

Systematic review of observational studies

Event (# studies, #pts)	Follow-up (months)	Risk Difference or Prevalence	Certainty (GRADE)	Lay Summary
Cannabis dependence (n=3, 1824 pts)	3	prevalence: 5% (95%CI 0% to 20%)	Very Low	We are very uncertain
Road traffic accident causing injury (n=1, 431 pts)	12	RD 14 more/100,000 (95%Cl 6 fewer to 523 more)		regarding the prevalence of dependence or effect of cannabis
Falls (n=1, 431 pts)	12	RD 0 more/1,000 (16 fewer to 56 more)	Very Low	on motor vehicle accidents and falls
Suicide (n=1, 431 pts)	12	RD: 3 fewer/1,000 (5 fewer to 36 more)	Very Low	We are very
Suicidal thoughts (n=3, 3066 pts)	12	prevalence: 0.2% (0% to 0.4%)	Very Low	uncertain regarding the prevalence of suicidal thoughts, or the effect of cannabis on suicide



Cannabis use and driving-related performance in young recreational users: a within-subject randomized clinical trial

Tatiana Ogourtsova PhD OT(c), Maja Kalaba MPH, Isabelle Gelinas PhD OT(c), Nicol Korner-Bitensky PhD OT(c), Mark A. Ware MBBS MSc

- Participants completed tests in the no-cannabis state and at 1, 3 and 5 hours after inhalation of a standard 100mg dose of cannabis.
- Cannabis use showed no effect on simple driving-related tasks, but there was significant impairment on complex tasks, especially when these were novel.
- These effects, along with lower self-perceived driving ability and safety, lasted up to 5 hours after use.



To whom the recommendation applies

- ▼ No trial eligible for our systematic review explored the effect of inhaled forms of medical cannabis, or enrolled patients involved in palliative care.
 - Our recommendation does not apply to smoked or vaporized forms of cannabis, cannabis provided for recreational purposes, or patients receiving end-of-life care.
- Trials eligible for our reviews largely excluded chronic pain patients with concurrent mental illness, or those receiving disability benefits; the generalizability of our recommendations to these populations is therefore uncertain.





Strong recommendations

1. Clear imbalance



- benefits clearly outweigh risks/hassle/cost
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Weak recommendations

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- choice varies appreciably (or is very uncertain)





Final recommendation

PRACTICE



For numbered affiliations see end of article.

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RAPID RECOMMENDATIONS

Medical cannabis or cannabinoids for chronic pain: a clinical practice guideline

Jason W Busse, ^{1,2,3,4} Patrick Vankrunkelsven, ^{5,6} Linan Zeng, ^{2,7} Anja Fog Heen, ⁸ Arnaud Merglen, ⁹ Fiona Campbell, ¹⁰ Lars Petter, ¹¹ Bert Aertgeerts, ^{12,13} Rachelle Buchbinder, ^{14,15} Matteo Coen, ^{16,17} David Juurlink, ^{18,19} Caroline Samer, ^{20,21} Reed A C Siemieniuk, ² Nimisha Kumar, ²² Lynn Cooper, ²³ John Brown, ⁴ Lyubov Lytvyn, ² Dena Zeraatkar, ^{2,24} Li Wang, ^{2,3} Gordon H Guyatt, ² Per O Vandvik, ⁸ Thomas Agoritsas^{2,25}





More comprehensive guidelines are underway

- Recent funding by Health Canada has facilitated more comprehensive guideline development, including:
 - Updating of all prior evidence syntheses
 - Benefits and harms of inhaled forms of cannabis
 - Relative effectiveness of medical cannabis vs. opioids
 - Impact of cannabis on driving
 - Guidance of cannabis tapering
 - Evidence on risk for developing cannabis use disorder
 - Primary study of patients values and preferences regarding cannabis for chronic pain





How can medical cannabis be accessed?





What are the steps for accessing medical cannabis?

1. A **medical document** (prescription) is written by an MD or NP and is sent to a **Health Canada-regulated License Holder** (your pharmacy for medical cannabis) by secure fax or physical mail.

2. You must fill out a **registration document for each License Holder** you order from online or by hand.

3. After your registration document is **approved** by the License Holder, **you can then place an order online or over the phone**.



Accessing medical cannabis

- Patients authorized by their healthcare provider may access cannabis for medical purposes by:
 - buying directly from a federally licensed seller
 - registering with Health Canada to produce a limited amount of cannabis for their own medical purposes
 - designating someone to produce it for them

https://www.canada.ca/en/health-canada/services/drugs-medication/cannabis/medical-use-cannabis.html





What is the Cannabis Act, why is it up for review, and how will its review affect medical cannabis users?





The Cannabis Act

- The Cannabis Act has two objectives:
 - 1. Aims to protect the health and safety of Canadians while adapting and responding to the ongoing and emerging needs of Canadians.
 - 2. Provides for the establishment of a competitive legal industry to displace the illicit market.
- A review of the Act has recently been announced, whose goal is to ensure that the Act adapts to the current situation and continues to meet Canadians needs and expectations.





What resources are available to learn more about medical cannabis?





Most physicians were reluctant to authorize cannabis due to:

- lack of knowledge,
 - There is no formal training in medical school about medical cannabis
 - Most expressed interest in continuing medical education on cannabis
- concerns about limited evidence,
 - Conflicting guidelines
- potential harms
 - Increased risk for older patients (e.g., falls)
 - Drug interactions with cannabis: There are 392 drugs known to interact with cannabis, 27 are major and 365 are moderate:

https://www.drugs.com/drug-interactions/cannabis.html



Research

Attitudes toward medical cannabis among family physicians practising in Ontario, Canada: a qualitative research study

Jeremy Y. Ng MSc, Kevin Gilotra, Sana Usman BSc, Yaping Chang PhD, Jason W. Busse DC PhD

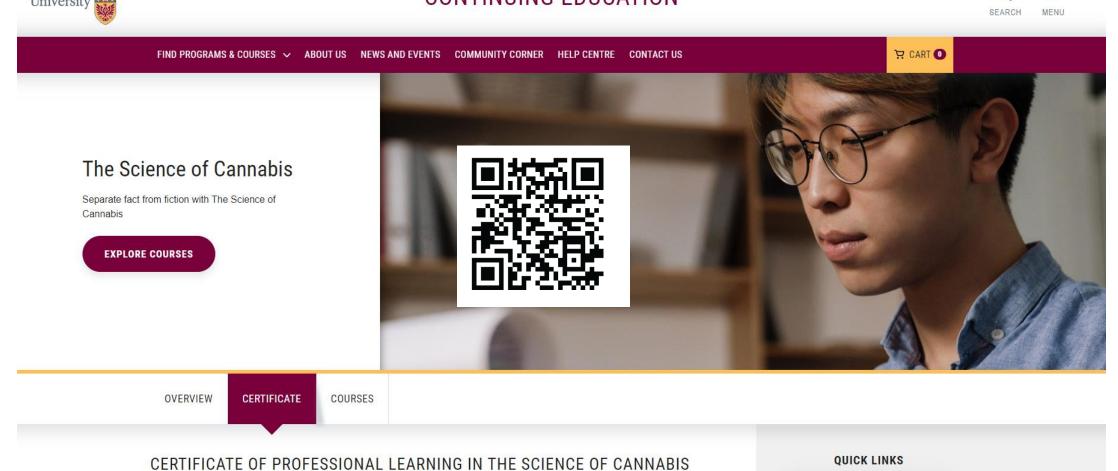






Curating the Evidence Base

McMaster University CONTINUING EDUCATION



https://continuing.mcmaster.ca/programs/health-social-services/the-science-of-cannabis/#tab-content-certificate

Medical Cannabis Learning Hub available at arthritis.ca

▼ Learn more about:

- CBD & THC
- Risks of cannabis use
- Accessing and using medical cannabis
- Current research







Any final thoughts or recommendations?





Questions





Tell us what you think...







