# **Supplementary Online Content**

Gedin F, Blomé S, Pontén M, et al. Placebo response and media attention in randomized clinical trials assessing cannabis-based therapies for pain: a systematic review and meta-analysis. *JAMA Netw Open.* 2022;5(11):e2243848. doi:10.1001/jamanetworkopen.2022.43848

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This supplementary material has been provided by the authors to give readers additional information about their work.

# eTable 1. Search Strategies

# Medline (Ovid)

1	Cannabis/	8926
2	exp "Marijuana Use"/	5301
3	Medical Marijuana/	1116
4	exp Cannabinoids/	13758
5	(bhang* or cannabi* or ganja* or hash* or hemp* or marijuana* or marihuana* or marijuana).ti,ab,kf.	55503
6	or/1-5	60051
7	exp Pain/	389370
8	(pain* or neuralgia*).ti,ab,kf.	687785
9	or/7-8	831552
10	Randomized Controlled <u>Trial.pt</u> .	501769
11	Controlled Clinical <u>Trial.pt</u> .	93575
12	randomized.ab.	472899
13	placebo.ab.	205951
14	Clinical Trials as Topic.sh.	190378
15	randomly.ab.	329139
16	trial.ti.	214952
17	or/10-16	1275136
18	exp Animals/ not Humans.sh.	4678248
19	17 not 18	1173020
20	6 and 9 and 19	424

### Embase (embase.com)

('cannabis'/de OR 'cannabis use'/exp OR 'medical cannabis'/de OR 'cannabinoid'/exp OR bhang\*:ti,ab,kw OR cannabi\*:ti,ab,kw OR ganja\*:ti,ab,kw OR hash\*:ti,ab,kw OR hemp\*:ti,ab,kw OR marijuana\*:ti,ab,kw OR marijuana\*:ti,ab,kw)

AND

('pain'/exp OR pain\*:ti,ab,kw OR neuralgia\*:ti,ab,kw)

AND

('controlled clinical trial'/exp OR random\*:ab OR placebo:ab OR 'clinical trial (topic)'/exp OR trial:ti) NOT ([animals]/lim NOT [humans]/lim)

NOT [medline]/lim = 487

#### **Updated literature search**

The updated literature search was performed in September of 2021. There were no modifications made to the literature search strategy and the purpose of the updated search was only to include newly published articles.

### Embase (embase.com)

('cannabis'/de OR 'cannabis use'/exp OR 'medical cannabis'/de OR 'cannabinoid'/exp OR bhang\*:ti,ab,kw OR cannabi\*:ti,ab,kw OR ganja\*:ti,ab,kw OR hash\*:ti,ab,kw OR hemp\*:ti,ab,kw OR marijuana\*:ti,ab,kw OR marijuana\*:ti,ab,kw OR marijuana\*:ti,ab,kw)

AND

('pain'/exp OR pain\*:ti,ab,kw OR neuralgia\*:ti,ab,kw)

AND

('controlled clinical trial'/exp OR random\*:ab OR placebo:ab OR 'clinical trial (topic)'/exp OR trial:ti) NOT ([animals]/lim NOT [humans]/lim)

NOT [medline]/lim

AND [randomized controlled trial]/lim AND [2020-2021]/py

= 42

### Medline ovid

1	Cannabis/	7822
2	exp "Marijuana Use"/	6458
3	Medical Marijuana/	1449
4	exp Cannabinoids/	15548
5	(bhang* or cannabi* or ganja* or hash* or hemp* or marijuana* or marihuana* or marijuana).ti,ab,kf.	63608
6	or/1-5	67143
7	exp Pain/	418520
8	(pain* or neuralgia*).ti,ab,kf.	764157
9	or/7-8	913068
10	Randomized Controlled <u>Trial.pt</u> .	542796
11	Controlled Clinical <u>Trial.pt</u> .	94373
12	randomized.ab.	532993
13	placebo.ab.	221223
14	Clinical Trials as Topic.sh.	197233
15	randomly.ab.	365375
16	trial.ti.	247058
17	or/10-16	1392241
18	6 and 9 and 17	538
21	limit 18 to (humans and yr="2020 - 2021")	56

eTable 2. Excluded Articles and Reason for Exclusion

Publication	Author	Reason for exclusion							
Year		Population	Intervention	Control	Outcome	Other			
2007	Abrams	HIV							
2019	Almog					Abstract			
2009	Ambler					Abstract			
2018	Amerongen	Healthy							
1974	Angelico					Letter			
2019	Babalonis	Healthy							
2017	Babalonis	Healthy							
2014	Barbara					Abstract			
2009	Chung					Abstract			
2012	Cooper					Abstract			
2017	Cooper					Abstract			
2013	Cooper	Healthy							
2018	Cooper	Healthy							
2009	Ellis	HIV							
2007	Esfandyari	Healthy							
2016	Fionda					Abstract			
1977	Gottschalk				Did not				
					measure pain				
2004	Grotenhermen					Commentary			
2018	Hunter					Abstract			
2012	Kalliomaki	Healthy							
2013	Kalliomaki	Healthy							
2013	Kalliomaki	Healthy							
1975	Karnol				Did not				
					measure pain				
2011	Klooker				Discomfort,				
					not pain				
2008	Kraft	Healthy							
2014	Leocani					Abstract			
2017	Levin	Not pain							
2013	Lötsch					Abstract			
2017	Marinelli					Protocol paper			
1975	Milstein	Healthy							
2003	Naef	Healthy							
2004	Naef	Healthy							
2009	Notcutt					Abstract			

2011	Ostenf	eld,	Dentist						
			patients						
2012 Pinsge		r					Abstract		
2009/2010 Pittler							Commentary		
2008	Redmo	ond	Healthy						
2006	Robert	S	Healthy						
2007	Rog				Uncontrolled				
					follow-up				
2005	Salim						Secondary analysis of		
							another article in study		
2019	Schind	ller,					Abstract		
2013	Sewell			-			Conference abstract		
2009	Turcot	te					Poster abstract		
2011	Turcot	te				Poster abstract			
2017	Turner	•					Conference abstract		
2006	Wade						Follow-up study of Wade		
							et al. (2004)		
2013	Wallac	e					Abstract		
2017	Wallac	e					Conference abstract		
2019	Walter	•				Discrimination			
						not intensity			
2010B	Ware				Active				
					control				
2009	Wasan						Conference abstract		
2018	Werth						Conference abstract		
2019	Werth						Conference abstract		
2016	Wilsey	7					Same participants as		
							Wilsey et al. (2016)		
2009	Zajick						Abstract		
2015	Zalai						Abstract		
2020	Covell	i	No pain at						
			baseline						
	1		Artic	les that did not	provide sufficie	nt data	1		
Year		Author							
2020		Abrams							
2020		Almog							
2017		Amerongen							
2006		Beaulieu							
2021		Bebee							
2006		Blake							
2008		Conte							
2016		Cote							

2007	Hagenbach	
1973	Hill	
1978	Jochimsen	
2010	Johnson	
2003	Karst	
2018	Lichtman	
2014	Lynch	
2020	Meuth	
2008	Narang	
1975	Noyes	
2012	Portenoy	
1977	Raft	
2010	Selvarajah	
2014	Serpell	
1978	Staquet	
2004	svendsen	
2015	Turcott	
2019	van de Donk	
2004	Wade	
2008	Wilsey	
2013	Wilsey	
2016	Wilsey,	
2006	Wissel	
2003	Zajicek	
2008	Nietecka-Buchta	

### eTable 3. Risk of Bias Assessments

The Risk-of-bias tool 2 (ROB 2) criteria were used with the addition of a sixth category for blinding.

Source	1. Bias arising from the randomization process	2. Bias due to deviations from intended interventions	3. Bias due to missing outcome data	4. Bias in measurement of the outcome	5. Bias in selection of the reported result	Bias due to blinding	Overall risk of bias
Berman, 2004 <sup>12</sup>	Low	Low	Low	Low	High	Low	Moderate
Buggy, 2003 <sup>13</sup>	Low	Low	Low	Low	High	Low	Moderate
Chaves, 2020 <sup>14</sup>	Low	Low	Low	Low	Low	Low	Low
Corey-Bloom, 2012	Low	Low	Moderate	Low	Low	Low	Moderate
De Vries, 2017 <sup>16</sup>	Low	Low	Moderate	Low	Moderate	Low	Moderate
Issa, 2014 <sup>17</sup>	Low	Low	Low	Low	High	Low	Moderate
Langford, 2013 <sup>23</sup>	Low	Low	Low	Low	Low	Low	Low
Malik,2017 <sup>24</sup>	Low	Low	Moderate	Low	Moderate	Low	Moderate
Nurmikko, 2007 <sup>25</sup>	Low	Low	Low	Low	Moderate	Low	Moderate
Rog, 2015 <sup>22</sup>	Low	Low	Low	Low	Moderate	Moderate	Moderate
Schimrigk, 2017 <sup>18</sup>	Low	Low	Low	Low	Moderate	Moderate	Moderate
Skrabek, 2008 <sup>19</sup>	Moderate	Moderate	Moderate	Moderate	High	Moderate	High
Toth 2012 <sup>20</sup>	Moderate	Moderate	Low	Low	High	Moderate	High
Turcott 2018 <sup>21</sup>	Moderate	High	High	Moderate	Moderate	High	High
Wade 2003 <sup>27</sup>	Moderate	Moderate	Low	Moderate	High	Moderate	High
Wallace, 2015 <sup>26</sup>	Low	Low	Low	Low	Low	Moderate	Moderate
Ware, 2010 <sup>28</sup>	Low	Low	Low	Low	Low	Moderate	Moderate
Weizman, 2018 <sup>29</sup>	Low	Moderate	Low	Low	Moderate	Moderate	Moderate
Zadikoff, 2011 <sup>30</sup>	Low	Moderate	Low	Low	Low	Moderate	Moderate
Zaijcek, 2012 <sup>31</sup>	Low	Low	Low	Low	Moderate	Moderate	Moderate

eTable 4. Each Trial's Scientific Citations, Journal Impact Factor, and Altmetric Scores

	Crossref	Journal	Altmetric score	Altmetric
Source	Academic	Impact	Non-academic	Age adjusted
	citations	Factor	citations	percentile
Berman, 2004 12	219	6	20	91
Buggy, 2003 13	120	6	4	65
Chaves, 2020 14	5	3	169	99
Corey-Bloom, 2012 15	107	7	329	99
De Vries, 2017 16	37	5	26	91
Issa, 2014 <sup>17</sup>	22	3	5	76
Langford, 2013 <sup>23</sup>	129	5	33	96
Malik, 2017 <sup>24</sup>	23	3	3	61
Nurmikko, 2007 <sup>25</sup>	234	7	24	94
Rog, 2005 <sup>22</sup>	458	10	93	99
Schimrigk, 2017 18	41	6	15	87
Skrabek, 2008 19	208	6	42	97
Toth 2012 <sup>20</sup>	96	6	15	92
Turcott 2018 <sup>21</sup>	35	3	6	72
Wade 2003 <sup>27</sup>	263	3	59	97
Wallace, 2015 <sup>26</sup>	98	5	285	99
Ware, 2010 <sup>28</sup>	232	7	378	99
Weizman, 2018 <sup>29</sup>	25	8	212	98
Zadikoff, 2011 <sup>30</sup>	23	1	3	58
Zajicek, 2012 <sup>31</sup>	123	8	154	99
Mean (SD)	125 (112)	5 (2)	89 (115)	88 (14)

eFigure 1. Funnel Plot for the Risk of Publication Bias

