# Prevalence of Cannabis Use Among Medical Students: A Systematic Review and Meta-analysis

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# ABSTRACT

OBJECTIVE: Cannabis is reported to be the most common illicit substance used among medical students; however, the number of related studies is limited and their results are not systematically reviewed. The aim of our study was to analyze the prevalence of lifetime and current use of cannabis among medical students worldwide.

METHODS: A systematic review and meta-analysis was performed with adherence to the PRISMA guidelines. The electronic databases PubMed, Scopus, and Cochrane library were searched for studies on the prevalence of cannabis use among medical students. Prevalence of lifetime, past-year, and past-month cannabis use was extracted. Pooled prevalence and relative risk for sex were calculated using the random effects model and subgroup analyses were conducted.

RESULTS: A total of 38 observational (cross-sectional and cohort) studies were included (total number of participants 19932), and most of them were conducted in Europe, Central and Southern America, and the United States. Overall pooled prevalence of lifetime cannabis use was 31.4% (95% confidence interval [CI]: 23.7%-39.6%), past-year use was 17.2% (95% CI: 10.8%-24.6%), and past-month use was 8.8% (95% CI: 5.6%-12.5%). Men displayed higher rates of cannabis use with a pooled relative risk of 1.55 (95% CI: 1.32-1.81). Heterogeneity was high ( $l^2 > 75\%$ ) and there were differences among continents in all outcomes (P < .001).

CONCLUSIONS: In conclusion, 1 in 3 medical students has used cannabis, whereas 8.8% were current users. Significant differences among continents were observed, but common finding was that male students tend to consume cannabis more often than female students.

KEYWORDS: Cannabis, medical students, prevalence

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# Introduction

Cannabis, both resin (hashish) and herbal (marijuana), is the illicit drug most likely to be used across all age groups.<sup>1,2</sup> Medical students are not the exception; cannabis is reported to be the most common illicit substance used, with a reported past-month use of 11.84%.<sup>3</sup> Exposure to diverse stressors, burnout, and relatively easy access to drugs set physicians and medical students vulnerable to substance use.<sup>4</sup> Substance use may affect students' current academic performance and also it may contribute to misjudgments and misperceptions of future physicians toward patients with substance use disorders.<sup>5</sup> It is reported that students' attitudes toward substance use behaviors influence their future preventive counseling practices.<sup>6</sup> Consequently, the study of lifetime and, especially, current cannabis use among medical students is of great importance.

Interestingly, although cannabis use and its consequences among adults is extensively studied worldwide and detailed analyses are published annually by the responsible offices of the United Nations (UN)<sup>1</sup> and the European Union,<sup>2</sup> there is a lack of recent studies on cannabis use among medical students. To our knowledge, there is only one published systematic review

that examined both legal and illegal substance use by medical students worldwide, where information regarding cannabis use could partly be retrieved.<sup>3</sup> Thus, the aim of our article was to systematically review and meta-analyze the literature on the epidemiology of cannabis use among medical students and to present the prevalence of lifetime, past-year, and past-month use by geographical area.

# **Methods**

# Search strategy

A systematic review was performed with adherence to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines.<sup>7</sup> The electronic databases PubMed, Scopus, and Cochrane library were searched from inception through October 31, 2017. The search strategy for PubMed included different combinations of Medical Subject Headings (MeSH) search terms and non-MeSH terms, integrating into the query: ("students, medical" [MeSH Terms] OR ("students" [All Fields] AND "medical" [All Fields]) OR "medical students" [All Fields] OR ("medical" [All Fields] AND "students" [All Fields]) AND ("cannabis" [MeSH

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Creative Commons Non Commercial CC BY-NC: This article is distributed under the terms of the Creative Commons Attribution-NonCommercial 4.0 License (http://www.creativecommons.org/licenses/by-nc/4.0/) which permits non-commercial use, reproduction and distribution of the work without further permission provided the original work is attributed as specified on the SAGE and Open Access pages (https://us.sagepub.com/en-us/nam/open-access-at-sage). Terms] OR "cannabis" [All Fields] OR "hashish" [All Fields] OR "marijuana" [All Fields] OR "illegal drugs" [All Fields] OR "illegal substances" [All Fields] OR "Psychoactive Substances" [All Fields] OR "Psychoactive Drugs" [All Fields]). All of the identified articles were then limited to English language articles to clearly evaluate the results and the methodology of the study, and only full-text articles were included ("Text Availability" filter: Full Text, "Language" filter: English). Similar search strategies were used for the Scopus and Cochrane Library database and duplicates were excluded.

#### Study selection

The inclusion criteria were as follows: (1) studies that presented the precise number of cannabis prevalence, distinguished from other substances use, to analyze data regarding only the use of cannabis; (2) studies that presented specific data of cannabis use only by medical students and not mixed together with other students' groups or graduates from medical school (junior doctors, master students, etc); (3) the number of participants who turned in fully completed and acceptable questionnaires had to be written alongside with the initial number of invited participants; and (4) full-text English language articles.

Two investigators (I.K. and I.T.) independently screened all the titles and abstracts, and discrepancies were solved by discussion with a third author (S.S.). The reference lists of relevant articles were also hand-searched. Records that considered as potentially relevant were retrieved in full text and preceded to evaluation. Articles published as editorials, letters, conferences, or meeting abstracts were excluded. The remaining articles were eligible for abstract review.

# Data extraction

From each study, data regarding the total number of participants, their year of studies, mean age, prevalence of cannabis use (lifetime, past-year, past-month) alongside with differences among the 2 sexes were extracted. For studies with a cohort of students, followed throughout their medical studies, only percentages from the last, in chronological order, survey were included, to evaluate their most recent use.

#### Assessment of quality

The quality of the selected articles was assessed using the Grading of Recommendations Assessment, Development and Evaluation system.<sup>8</sup> The grading scheme classifies quality of evidence as high (A), moderate (B), or low (C)

# Data synthesis

The narrative description was accompanied by the pooled prevalence (lifetime, past year, and past month) of cannabis use

and the pooled relative risk for sex, worldwide and stratified by continent. The 95% confidence intervals (CIs) of prevalence estimates were calculated using the Wilson methods.9 Prevalence data were transformed using the Freeman-Tukey double arcsine transformation, to stabilize the variance and avoid overestimation of prevalence extremities.<sup>10</sup> The relative risk and the accompanied 95% CIs were calculated to estimate the risk of sex for cannabis use. The random effects model was used to incorporate heterogeneity. The DerSimonian-Laird estimator for tau-squared was used, as well as the inverse variance method was used to calculate the pooled prevalence and the Mantel-Haenszel method to calculate the pooled relative risk. Heterogeneity was assessed with  $I^2$ , considering as high heterogeneity when  $I^2$  was equal to or greater than 75%. Subgroup analyses were conducted for continents, publication year, and sample size for overall studies and stratified by continent. Subgroup differences were assessed by the  $\chi^2$ -statistic. Significance was predefined at  $\alpha$  level .05 and for subgroup analyses at .01 to reduce falsepositive results due to multiple hypotheses testing. The statistical analysis was performed using the package meta version 4.9-011 on R version 3.3.2.12

# Results

### Study characteristics

The initial total number of studies after implementing the aforementioned filters was 331 from PubMed, 481 from Scopus, and 8 from the Cochrane Database. The screening of the titles and abstracts identified 51 potentially relevant articles that were retrieved in full-text format. A total of 13 full-text articles were excluded because they did not meet the inclusion criteria. Thus, after complete review, 38 studies were included in the systematic review (Figure 1). From them, 11 referring to Europe: 6 the United Kingdom,<sup>13–18</sup> 2 Spain,<sup>19,20</sup> 1 Croatia,<sup>21</sup> 1 France,<sup>22</sup> and 1 Serbia<sup>23</sup>; 13 to the United States<sup>24–36</sup>; 8 to the Central and Southern America: 7 Brazil<sup>37–43</sup> and 1 Honduras<sup>44</sup>; 4 to Asia: 2 India,<sup>45,46</sup> 1 Iran,<sup>47</sup> and 1 Nepal<sup>48</sup>; as well as 2 to Africa, Nigeria<sup>49,50</sup> (Table 1). No study was identified for Oceania.

The demographics of the selected studies, individual study quality, as well as the prevalence of cannabis use among medical students are listed in Table 1. The median publication year was 2004 (ranging from 1971 to 2017) and the median sample size was 400 (ranging from 110 to 2308). The total number of participants in the studies included in our review was 19932. Specifically, the studies included 6605 participants from Europe, 5951 from the United States, 3415 from Central and Southern America (92% came from Brazil), 3033 from Asia, and 928 participants from Africa (all of them from Nigeria). Because of the observational nature of the investigations, there were no studies that received a grade higher than C (Table 1).



# Prevalence of cannabis use

Lifetime cannabis use. The pooled prevalence of lifetime cannabis use among medical students worldwide was 31.4% (number of studies k=28, number of participants N=16061, prevalence=31.4% [95% CI: 23.7%-39.6%],  $I^2$ =99.2%; Figure 2A). There were differences among continents ( $\chi_4^2$  = 60.78, P < .001), with the lowest prevalence in Asia (k=3, N=2860, 6% [1.77%-12.44%]) and the highest in the United States (k=10, N=4656, 48.05% [36.72%-59.48%]). Publication year and sample size did not moderate lifetime prevalence of cannabis use, except for studies from Asia. Studies from Asia published after 2000 and having sample size larger than 400 reported higher prevalence (k=2, N=2645, 9.52% [4.09%-16.87%] versus an earlier and smaller study; k=1, N=215, 0.93% [0.01%-2.78%];  $\chi_1^2 = 10.05$ , P = .0015; see subgroup analyses in eAppendix 1).

*Past-year cannabis use.* Pooled prevalence of past-year cannabis use around the world was estimated to be 17.2% (k=15, N=5141, 17.2% [10.8%-24.6%],  $I^2$ =97.7%; Figure 2B). Again, there were subgroup differences regarding the continent ( $\chi_3^2 = 273.47$ , P < .001). There was no study reporting past-year use in Africa and only 1 study in Europe; this single European study reported surprisingly a very high past-year prevalence of 76.6%. Excluding this study from Europe,<sup>22</sup> the overall pooled prevalence was 13.9% (95% CI: 9.4%-19.2%,  $I^2$ =96%). Regarding the other continents, prevalence ranged between 5.8% in Asia (k=1, N=173, 5.8% [2.72%-9.82%])<sup>47</sup>

STUDY	COUNTRY	PARTICIPANTS GENDER: M/F	STUDY YEAR	AGE	GENDER OF USERS (M/F)	LIFETIME USE, %	PAST- YEAR USE, %	PAST- MONTH USE, %	QUALITY (GRADE)
Africa									
Ihezue <sup>49</sup>	Nigeria	728 M/F: 607/121	2nd-6th	$22.6\pm2.36$	M/F: 22/1	ns	ns	3.1%, n=23	С
James et al. 2013 <sup>50</sup>	Nigeria	200 M/F: 109/91	5th	$24.1\pm2.45$	ns	10.5%, n=21	ns	5%, n=10	С
Asia									
Ali and Vankar <sup>45</sup>	India	215 ns	1st, 2nd, 6th	19.7	M/F: 2/0	0.93%, n=2	ns	0	С
Jodati et al <sup>47</sup>	Iran	173 M/F: 173/0	ns	21.3	ns	ns	5.7%, n=10	ns	С
Rai et al46	India	2135 M/F: 1512/623	1st-6th	20.5	M/F: 129/12	6.0%, n=141	ns	1.4%, n=32	С
Budhathoki et al <sup>48</sup>	Nepal	510 M/F: 324/186	3rd	22.5	M/F: 62/5	13.1%, n=67	ns	ns	С
Central and	Southern Ame	erica							
Lambert Passos et al <sup>37</sup>	Brazil	1054 M/F: 499/553	1st-6th	$21.1\pm3.9$	M/F: 124/94	20.9%, n=218	ns	5.5%, n=59	С
Di Pietro et al <sup>38</sup>	Brazil	456 M/F: 247/209	1st-6th	$21.12\pm2.4$	ns	ns	ns	15%, n=68	С
Boniatti et al <sup>40</sup>	Brazil	183 M/F: 84/99	1st-6th	$22.5\pm2.4$	M/F: 31/26	31.1%, n=57	13.6%, n=25	7.6%, n=14	С
Buchanan and Pillon <sup>44</sup>	Honduras	260 M/F: 115/145	4th-5th	22.5	ns	ns	3.8%, n=10	1.9%, n=5	С
Carvalho et al <sup>41</sup>	Brazil	465 M/F: 265/200	1st-6th	21.5	ns	14.4%, n=67	4.5%, n=21	ns	С
Da Silveira et al <sup>39</sup>	Brazil	456 M/F: 247/209	1st-6th	21	M/F: 63/12	ns	16.4%, n=75	ns	С
De Oliveira et al <sup>42</sup>	Brazil	209 M/F: 111/98	6th	22	M/F: 34/17	24.3%, n=51	14.3%, n=30	10.5%, n=22	С
Petroianu et al <sup>43</sup>	Brazil	332 M/F: 160/172	1st-6th	23	M/F: 37/16	ns	16%, n=53	ns	С
Europe									
McKay et al <sup>13</sup>	United Kingdom	749 ns	1st-4th	20	M/F: 80/20	13.4%, n=100	ns	3.6%, n=27	С
Laporte et al <sup>19</sup>	Spain	808 M/F: 515/293	1st-6th	ns	M/F: 45/19	7.9%, n=64	ns	3.8%, n=31	С
Rodriguez et al. 1986 <sup>20</sup>	Spain	2308 M/F: 1112/1196	ns	ns	M/F: 285/193	20.7%, n=478	ns	ns	С
Ashton and Kamali <sup>14</sup>	United Kingdom	186 M/F: 76/109	ns	20.4±1.8	M/F: 26/30	30.3%, n=56	ns	ns	С
Webb et al <sup>15</sup>	United Kingdom	750 M/F: 333/417	2nd	ns	M/F: 180/165	46%, n=345	ns	10%, n=75	С
Newbury- Birch et al <sup>16</sup>	United Kingdom	194 M/F: 64/130	1st	$18.8\pm2.1$	M/F:32/54	44.3%, n=86	ns	19%, n=37	С

# Table 1. Studies of cannabis use among medical students worldwide.

#### Table 1. (Continued)

STUDY	COUNTRY	PARTICIPANTS GENDER: M/F	STUDY YEAR	AGE	GENDER OF USERS (M/F)	LIFETIME USE, %	PAST- YEAR USE, %	PAST- MONTH USE, %	QUALITY (GRADE)
Pickard et al <sup>17</sup>	United Kingdom	136 M/F: 46/90	2nd	ns	M/F: 11/30	30.1%, n=41	ns	ns	С
Newbury- Birch et al <sup>18</sup>	United Kingdom	110 M/F: 33/77	6th	ns	M/F: 22/50	65.5%, n=72	ns	23.6%, n=26	С
Trkulja et al <sup>21</sup>	Croatia	775 M/F: 290/485	1st-6th	18-24	M/F: 129/144	61.5%, n=273	ns	ns	С
Gignon et al <sup>22</sup>	France	171 ns	2nd-6th	$22.1\pm1.7$	ns	ns	77%, n=131	33%, n=56	С
Vujcic et al <sup>23</sup>	Serbia	418 M/F: 156/262	4th	$22.46\pm1.12$	M/F: 63/83	34.9%, n=146	ns	ns	С
USA									
Solursh et al <sup>24</sup>	USA	234 ns	3rd-Final	ns	ns	20.5%, n=48	16.6%, n=39	11.9%, n=28	С
Lipp et al <sup>25</sup>	USA	1063 ns	ns	ns	ns	52%, n=550	ns	30.9%, n=328	С
Rochford et al <sup>26</sup>	USA	134 ns	1st	22	ns	72%, n=96	ns	ns	С
Kory and Crandall <sup>27</sup>	USA	463 M/F: 421/42	1st-Final	23.6	ns	70%, n=302	ns	27.4%, n=127	С
McAuliffe et al <sup>28</sup>	USA	381 M/F: 225/156	1st-Final	25	ns	18.9%, n=72	5.77%, n=22	3.1%, n=12	С
Conard et al <sup>29</sup>	USA	589 M/F: 383/206	4th	27.6	ns	74%, n=436	31.9%, n=188	16.9%, n=100	С
Schwartz et al. 1990 <sup>30</sup>	USA	263 M/F: 170/93	2nd-3rd	ns	ns	48.3%, n=127	19%, n=50	5.3%, n=14	С
Croen et al <sup>31</sup>	USA	139 M/F: 79/60	3rd	<22	ns	ns	21.6%, n=30	ns	С
Choi et al <sup>32</sup>	USA	301 M/F: 109/192	1st-Final	ns	ns	ns	ns	1.3%, n=4	С
Zhou et al <sup>33</sup>	USA	431 M/F: 217/214	1st-Final	25	ns	31.5%, n=136	12.2%, n=53	9%, n=39	С
Chan et al <sup>34</sup>	USA	236 M/F: 119/115	1st-Final	25-29	ns	33.3%, n=127	ns	ns	С
Merlo et al <sup>35</sup>	USA	862 M/F: 369/491	1st-Final	ns	M/F: 190/224	46.8%, n=414	ns	4.1%, n=36	С
Ayala et al <sup>36</sup>	USA	855 M/F: 304/534	1st-Final	$25.6\pm3.30$	ns	ns	26.2%, n=224	11.7% n=100	С

When available, age is displayed with average and its standard deviation. M, male; F, female; ns, data not stated).

and 18.3% in the United States (k=7, N=2892, 18.3% [11.48%-26.32%], P=96.2%). Subgroup analyses suggested that publication year and sample sizes did not moderate prevalence of past-year cannabis use (see subgroup analyses in eAppendix 2).

Past-month cannabis use. A total of 25 studies reported pastmonth use of cannabis, with an estimated prevalence of 8.8%  $(k = 25, N = 13664, 8.8\% [5.6\%-12.5\%], I^2 = 98.1\%)$  worldwide (Figure 2C). Differences among continents were noticed  $(\chi_4^2 = 24.13, P < .001)$ , ranging from 0.6% in Asia  $(k=2, N=2350, 0.6\% [0\%-2.8\%], I^2 = 84.1\%)$  to 10.6% in the United States  $(k=10, N=5442, 10.6\% [5.13\%-17.74\%], I^2 = 98.3\%)$  and 13.4% in Europe  $(k=6, N=2782, 13.4\% [6.54\%-22.17\%], I^2 = 97.1\%)$ . Again, leaving out the recent study in France, which reported high past-month and past-year prevalence,<sup>22</sup>



the remaining overall pooled prevalence was similar, 8.1% (95% CI: 5.1%-11.7%) and 10.3% (95% CI: 5.1%-17.1%) in Europe. Subgroup analyses did not suggest any moderating effect except for Europe. Studies from Europe published after 2000 reported higher prevalence (k=2, N=281, 28.46% [20.04%-37.69%] versus earlier studies; k=4, N=2501, 7.98% [3.56%-13.91%];  $\chi_1^2 = 15.77$ , P < .001), as well as studies with sample size smaller than 400 participants (k=3, N=475, 24.94% [17.02%-33.8%] versus larger studies; k=3, N=2307, 5.5% [2.34%-9.87%];  $\chi_1^2 = 19.58$ , P < .001; see subgroup analyses in eAppendix 3).

#### Pooled relative risk for sex

Male medical students in comparison with female seem to be in higher risk for cannabis use, with an overall pooled relative risk of 1.55 (k=15, N=12149, 1.55 [1.32-1.81], I<sup>2</sup>=78.2%; Figure 3). There were differences among continents ( $\chi_4^2 = 40.29$ , P < .001), with higher risk for men demonstrated in Asia (k=2, N=2818, risk ratio [RR]=5.11 [3.13-8.32],  $I^2=0\%$ ) and Central and Southern America (k=5, N=2232, RR=1.99 [1.37-2.91], I<sup>2</sup>=72.9%). Only 1 study in the United States reported cannabis use stratified by sex but no association was identified (k=1, N=860, RR=1.13 [0.98-1.30]).<sup>35</sup> Regarding Europe, men were in higher risk with a relative risk of 1.32  $(k=9, N=5684, RR=1.32 [1.18-1.48], I^2=41.5\%)$ . Subgroup analyses suggested that publication year and sample sizes did not moderate the relative risk, except for Europe. Studies from Europe with sample size larger than 400 participants seem to have higher relative risk (k=5, N=5059, RR=1.45 [1.32-1.58], $I^2 = 0.0$  versus smaller studies; k = 4, N = 625, RR = 1.08 [0.90-1.30],  $I^2 = 0.0$ ;  $\chi_1^2 = 7.79$ , P = .0053; see subgroup analyses in eAppendix 4).

### Discussion

This is a first attempt to systematically review the published literature to examine lifetime and recent cannabis use among medical students' population. It is impressive that although cannabis use and its consequences for the adult population is extensively studied worldwide and detailed analyses are published annually by the responsible offices of the UN<sup>1</sup> and the European Union,<sup>2</sup> recent studies on the prevalence of cannabis use among medical students are lacking: only 8 studies were published worldwide within the past 5 years, whereas most of the studies (k=14) are more than 20 years old. This might explain some discrepancies found in our results compared with published data of the general population for some areas.

Concerning Europe, our results suggest that the lifetime prevalence among medical students was 31.4% and the past month was 13.4%, whereas the past-year use could not be estimated as only 1 study with a small sample size reported related information.<sup>22</sup> Unfortunately, official data for the young adult

population in Europe (aged 15-24) presented by the European Drug Report 2017 concern only past year use, which was 17.7% for 2017, with men outnumbering women by a factor of 2.<sup>2</sup> To the same report, cannabis is the illicit drug most likely to be used among European adults (aged 15-64) with a lifetime prevalence of 26.3%, whereas around 1% of European adults estimated to be daily or near-daily cannabis users. However, levels of use differ considerably between European countries ranging from the lowest 3.3% to the highest 22% concerning national estimates of past-year use.<sup>2</sup>

Of interest was our finding that the lifetime prevalence among US medical students was 48.1%, meaning that 1 in 2 medical students in the United States has experienced, even once, in lifetime the use of cannabis. This percentage on lifetime prevalence is hard to compare because the nationwide studies provide data only for past-year and past-month use. However, according to the UN Office on Drugs and Crime, cannabis use has been rising over the past decade in the Americas, most pronounced in the United States, with an overall increase of 43% in the number of past-year cannabis users and of 54% in the number of past-month users. Regarding Asia and Africa, the UN analysis noticed that cannabis use among the general adult population in Africa and in Asia is perceived to have continued to increase relatively rapidly in the past 5 years.1 However, precise data are missing not only for the medical students but also for the general population due to the lack of systematic research.

To our results, male students tend to consume cannabis almost 2-fold more often than female students because pooled relative risk worldwide was 1.55 and higher risk for men was demonstrated in Asia, Central and Southern America, as well as Europe. This is in accordance with the findings of a previously published narrative review where a male:female usage ratio of 2:1 among medical students was reported.<sup>3</sup> It is known that use of all drugs is generally higher among men and the recent European report found that among 88 million adult users in Europe, use of cannabis was more frequently reported by men (53.8 million) than women (34.1 million). Furthermore, an estimated 18.7 million young adults (aged 15-34) used drugs in the past year, with twice as many men as women.<sup>2</sup>

The major strength of this review was the number of studies included (k=38) and the total number of participants (N=19932). However, there were some limitations. Many of the included studies had sample sizes smaller than 400 and they were published before 2000, which may have had an impact on the exact prevalence of cannabis use. As in previous meta-analyses of prevalence of cannabis use among other population groups,<sup>51,52</sup> considerable heterogeneity between studies was detected. A meta-analysis of prevalence of cannabis use in Iranian students had considerable heterogeneity ( $I^2 > 94\%$ ), despite the inclusion of studies with a more homogeneous population.<sup>52</sup> Another meta-analysis on prevalence of cannabis use in patients with first episode psychosis found also

Study	Experim	ental Total	Events	Total	Risk Ratio	RR	95%-CI	Weight
continent = Africa Ihezue 1988 Random effects model Heterogeneity: not applical	<b>22</b>	<b>607</b> 607	1	<b>121</b> 121		— 4.39 — 4.39	[0.60; 32.23] [0.60; 32.23]	<b>0.6%</b> 0.6%
continent = Asia Dheeraj Rai 2008 Budhathoki 2010 Jodati 2007 Random effects model Heterogeneity: $I^2 = 0\%$ , $\tau^2$	129 62 10 = 0, p = 0.	1512 324 173 2009 38	12 5 0	623 186 0 809		4.43 7.12 5.11	[2.47; 7.94] [2.91; 17.39] [3.13; 8.32]	4.1% 2.3% 0.0% 6.4%
continent = CS America Lambert 2006 Boniatti 2007 Da Silveira 2008 De Oliveira 2009 Petroianu 2010 Random effects model Heterogeneity: $l^2 = 73\%$ , $\tau^2$	a 124 31 63 34 37 <sup>2</sup> = 0.1294	499 84 247 111 160 1101 , p < 0	94 26 12 17 16	553 99 209 98 172 1131		1.46 1.41 4.44 1.77 2.49 1.99	[1.15; 1.86] [0.91; 2.17] [2.46; 8.01] [1.06; 2.96] [1.44; 4.29] [1.37; 2.91]	7.4% 5.4% 4.0% 4.6% 4.4% 25.8%
continent = Europe Laporte 1977 Rodriguez 1986 Ashton 1995 Webb 1998 Birch 2000 Pickard 2000 Newbury-Birch 2001 Trkulja 2003 Vujcic 2017 Random effects model Heterogeneity: $l^2 = 41\%$ , $\tau^2$	45 285 26 180 32 11 22 129 63 2 = 0.0116	515 1112 76 333 64 46 33 290 156 2625 , $\rho = 0$	19 193 30 165 54 30 50 144 83	293 1196 109 417 130 90 77 485 262 3059		1.35 1.59 1.24 1.37 1.20 0.72 1.03 1.50 1.27 1.32	[0.80; 2.26] [1.35; 1.87] [0.80; 1.92] [1.17; 1.59] [0.88; 1.66] [0.40; 1.30] [0.77; 1.37] [1.24; 1.81] [0.98; 1.65] [1.18; 1.48]	4.6% 8.2% 5.3% 8.2% 6.6% 4.0% 6.9% 7.9% 7.2% 58.9%
continent = USA Merlo 2017 Random effects model Heterogeneity: not applicat	<b>190</b>	<b>369</b> 369	224	<b>491</b> 491	•	1.13 1.13	[0.98; 1.30] [0.98; 1.30]	<b>8.4%</b> 8.4%
<b>Random effects model</b> Heterogeneity: $I^2 = 78\%$ , $\tau^2$	<sup>2</sup> = 0.0740	<b>6711</b> , p < 0	.01	5611	0.1 0.5 1 2 10	1.55	[1.32; 1.81]	100.0%

**Figure 3.** Forest plot of pooled relative risk for sex of medical students for cannabis use by continent. Male medical students are referred as the experimental group and female students as the control group. The "Events" refer to the number of cannabis users, and "Total" refers the total number of males or females of the study. Black squares represent the relative risk of each study. The area of the square is proportional to weight of the study, which contributes to the pooled relative risk (calculated using the Mantel-Haenszel method). Horizontal lines show 95% confidence intervals. The diamond shows the pooled results and its width the 95% confidence interval. Between-study heterogeneity is represented by *I*<sup>2</sup>. The vertical dashed line indicates the overall pooled relative risk.

considerable heterogeneity between studies ( $I^2 = 92.1\%$ ).<sup>51</sup> It is suggested that prevalence of cannabis use could vary considerably at local level, even within geographical regions, reflecting diverge and changing patterns of cannabis cost and availability, as well as of attitudes toward cannabis use and awareness of the potential harms of cannabis use.<sup>51</sup> Methodologic differences among studies (such as different sampling of participants or questionnaires used) are also suggested as possible sources of heterogeneity.<sup>52</sup> As a result, the pooled worldwide and withincontinents results of our study should be interpreted with some caution. In addition, subgroup analyses had small statistical power and for some outcomes, they could not be conducted due to paucity of available data.

In conclusion, our results suggest a worldwide pooled lifetime prevalence of 31.4%, meaning that about 1 in 3 medical students has used cannabis one or more times during their life, whereas 8.8% of the students reported current use. However, significant differences were observed among continents, whereas Africa and Asia were underrepresented. Considering the future role of medical students toward patients' substance abuse behaviors, these numbers are not negligible. Further international studies among medical students are needed to strengthen the research on the epidemiology of cannabis use and to study their motivations and attitudes toward this risky behavior.

#### **Author Contributions**

GP: Study concept, design and supervision, critical revision of the manuscript. SS: Drafting of the manuscript, statistical analysis, interpretation of data. IT: Drafting of the manuscript, data and statistical analysis. IK: Study design, acquisition and analysis of data. TD:

Drafting of the manuscript, data interpretation. DK: Critical revision of the manuscript, study supervision.

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