

# Crude estimates of cannabis-attributable mortality and morbidity in Canada—implications for public health focused intervention priorities

**Benedikt Fischer<sup>1,2,3</sup>, Sameer Imtiaz<sup>1,4</sup>, Katherine Rudzinski<sup>1,5</sup>, Jürgen Rehm<sup>1,3,4,5,6,7</sup>**

<sup>1</sup>Social and Epidemiological Research Department, Centre for Addiction and Mental Health, Toronto, Canada M5S 2S1

<sup>2</sup>Centre for Applied Research in Mental Health and Addiction, Faculty of Health Sciences, Simon Fraser University, Vancouver, Canada V6B 5K3

<sup>3</sup>Department of Psychiatry, University of Toronto, Toronto, Canada M5T 1R8

<sup>4</sup>Institute of Medical Science, University of Toronto, Toronto, Canada M5S 1A8

<sup>5</sup>Dalla Lana School of Public Health, University of Toronto, Toronto, Canada M5T 3M7

<sup>6</sup>WHO/PAHO Collaborating Centre for Mental Health and Addiction, Toronto, Canada M5S 2S1

<sup>7</sup>Klinische Psychologie und Psychotherapie, Technische Universität Dresden, Dresden 01069, Germany

Address correspondence to Benedikt Fischer, E-mail: bfischer@sfu.ca

## ABSTRACT

**Background** Cannabis is the most commonly used drug in Canada; while its use is currently controlled by criminal prohibition, debates about potential control reforms are intensifying. There is substantive evidence about cannabis-related risks to health in various key outcome domains; however, little is known about the actual extent of these harms specifically in Canada.

**Methods** Based on epidemiological data (e.g. prevalence of relevant cannabis use rates and relevant risk behaviors; risk ratios; and annual numbers of morbidity/mortality cases in relevant domains), and applying the methodology of comparative risk assessment, we estimated attributable fractions for cannabis-related morbidity and mortality, specifically for: (i) motor-vehicle accidents (MVs); (ii) use disorders; (iii) mental health (psychosis) and (iv) lung cancer.

**Results** MVs and lung cancer are the only domains where cannabis-attributable mortality is estimated to occur. While cannabis use results in morbidity in all domains, MVs and use disorders by far outweigh the other domains in the number of cases; the popularly debated mental health consequences (e.g., psychosis) translate into relatively small case numbers.

**Conclusions** The present crude estimates should guide and help prioritize public health-oriented interventions for the cannabis-related health burden in the population in Canada; formal burden of disease calculations should be conducted.

**Keywords** attributable fractions, Canada, cannabis use, health risks, morbidity, mortality, public health

## Introduction

Cannabis is the most commonly used illegal drug in Canada. Some 10–13% of the general adult population are current cannabis users (i.e. past year-use), translating into an estimated total user population of 2.5–3.0 million cannabis users; use rates are considerably elevated (20–30%) in adolescent and young adult age groups (i.e. 16–25 years).<sup>1,2</sup> While cannabis use is criminalized in Canada, there are ongoing debates about potential control or policy reforms, including proposals for ‘decriminalization’ or ‘legalization’; these debates are amplified by recent policy developments throughout the Americas, where, for example, select US states as well as the country of

Uruguay have moved towards implementation of legalization regimes for cannabis.<sup>3–5</sup>

Alcohol, tobacco and illicit drug use are among the top 10 risk factors for disease burden in North America.<sup>6</sup> While it is well documented in seminal reviews that cannabis is

**Benedikt Fischer**, Senior Scientist, Professor & CIHR/PHAC Applied Public Health Chair

**Sameer Imtiaz**, Doctoral Research Student

**Katherine Rudzinski**, Doctoral Research Student

**Jürgen Rehm**, Addiction Policy Chair, Senior Scientist & Professor

associated with elevated risks for detrimental health outcomes in various domains,<sup>5,7–9</sup> there have been few attempts to actually quantify or compare cannabis-related harm (e.g. morbidity, mortality) outcomes on a population level for Canada [for exceptions see refs (10,11)]; such efforts would not only be useful to assess the public health burden of cannabis vis-à-vis other drugs, yet also to inform and guide policy and intervention priorities on the basis of actual harms.<sup>12–14</sup>

Based on the above reviews, there are four main substantive areas of health risks identified for cannabis use: (i) accidents/injury (e.g. motor-vehicle accidents, MVAs); (ii) use disorders (including dependence); (iii) mental health (e.g. psychosis); (iv) cancer (e.g. lung cancer); while most of these problems occur predominantly among frequent (e.g. daily or near-daily users, e.g. 25–30% of total users) and/or long-term/chronic cannabis users, accidents/injuries can occur acutely, i.e. following single episode-use.<sup>7,8,12–16</sup> Given these elevated cannabis-related risk profiles, what do we actually know about the extent of tangible health—i.e. morbidity or mortality—outcomes on a population level in Canada? The simple answer is: Not much, especially since the last attempt for quantification as part of the Canadian ‘Cost of Substance Abuse’ study in 2002.<sup>10,11,17</sup>

Facing this major gap, we conducted a small and simple projection exercise to estimate cannabis-attributable morbidity and mortality in the above key domains in Canada, with the principal objective to suggest basic relative quantification in these different areas of potential cannabis-related harms which can then be used to inform and prioritize interventions.

## Methods

We applied the assumptions of a comparative risk assessment (CRA), with a counterfactual scenario of ‘no cannabis use at all’.<sup>18</sup> In other words, we estimated the proportion of mortality and morbidity which could be avoided in the absence of cannabis use in Canada.<sup>19</sup> The estimates were based on cannabis-attributable population impact fractions (PIFs), a generalized form of attributable fractions,<sup>20,21</sup> with the basic formula of:

$$\text{PIF} = \frac{\sum_{i=1}^n P_i \text{RR}_i - \sum_{i=1}^n P'_i \text{RR}_i}{\sum_{i=1}^n P_i \text{RR}_i}$$

where  $i$  is the exposure-level category;  $\text{RR}_i$  is the relative risk at exposure level  $i$ ;  $P_i$  is the prevalence of the  $i$ th category of exposure under the current situation in Canada; and  $P'_i$  is the prevalence of the  $i$ th category under the counterfactual scenario, i.e. 100% of Canadians with a relative risk of 1 for no cannabis use.

The exposure levels were based on theoretical considerations: for motor-vehicle accidents/injuries as acute effects, we

used cannabis exposure immediately prior to traffic participation; for the other risks, we used regular cannabis use over time. As cannabis use is more prevalent at younger ages,<sup>1,2</sup> but some of the health problem outcomes (e.g. cancer) are more prevalent later in life, we used sex- and age-stratified analysis, i.e. the PIFs were calculated for each sex–age group and applied to the respective disease outcomes. Further disease-specific considerations are given below.

## Results

The results of our estimations for the four main areas of cannabis-related harms are presented in the following.

### Motor-vehicle accidents/injuries

Recent data from general population and roadside surveys suggest that some 2–4% of drivers have had recent driving episodes following cannabis use.<sup>2,22,23</sup> Seminal reviews concluded that, based on relevant cognitive, memory/reaction and psychomotor ability impairments, drivers under the influence of cannabis have a two to three times elevated risk of being involved in (fatal or non-fatal) MVA.<sup>24–26</sup> Recent Canadian studies based on blood samples collected from emergency room populations in several provinces found that 11–12% of drivers admitted for MVAs had used cannabis prior to driving and the derived relative risk was  $>4$ .<sup>27,28</sup> While concrete data for Canada are still limited, it can thus be estimated that 4–12% of MVA-related fatalities and/or injuries may involve driving under the influence of cannabis.<sup>25,29,30</sup> There were 2227 MVA-related fatalities and 170 629 injuries (all) in Canada in 2010.<sup>31</sup> On this basis, there may be 89–267 cannabis-related MVA fatalities, and 6825–20 475 cannabis-related MVA injuries in 2010. Beyond MVAs, cannabis may contribute to other kinds of injuries (i.e. falls, other traffic injuries involving bicyclists, pedestrians or boating), so that the reported data are likely underestimates.<sup>32</sup>

### Use disorders

It has been estimated that some 5–10% of cannabis users develop dependence, but a much higher proportion develops cannabis use disorders, i.e. dependence or abuse as defined in DSM-IV; dependence, as the more severe outcome, interferes with social and economic functioning and, in many instances, requires treatment.<sup>33–36</sup> Importantly for problem outcomes, use disorders can be assumed to result in morbidity only, and do not lead to direct mortality outcomes. Based on most recent Canadian survey data (2012), 1.3% of the general adult population (15 years+) qualified for cannabis use disorders,<sup>37</sup> corresponding to 380 000 Canadians with cannabis use

disorders in a given year. About 20–25% of this population (33 000) were registered—on a prorated basis—in Ontario’s publicly funded treatment system in 2012/13<sup>38</sup> which would translate into a Canada-wide treatment population of 76 000–95 000; however, an additional substantive number can be expected to receive private treatment care.

### Mental health (Psychosis)

Psychoactive substance use and related disorders are associated with most mental disorders, and cannabis is no exception.<sup>39,40</sup> Some of these associations are causal, but there may also be reverse causality or third factors such as a genetic predisposition underlying both the substance use and/or the mental disorders.<sup>41</sup> This complex causal web makes it virtually impossible to quantify the causal impact of cannabis use on mental health disorders; similar circumstances have led to a situation where, even though causality had been established, the impact of alcohol on mental health disorders (e.g. depression) could not be estimated within a CRA framework.<sup>6,42</sup> For cannabis, the only mental health condition for which a causal impact has been established within the most recent CRA is psychosis. Recent systematic reviews suggested that regular cannabis users have a 1.5–4 times elevated risk of psychosis.<sup>43–45</sup> The resulting attributable fractions would be overestimates, as many underlying studies were not adjusted for confounders, and as the respective formulae assumed that all causality is uni-directional.<sup>46</sup> Allowing for a correction factor of 50% of the effect as usual in CRA,<sup>47</sup> we estimate a cannabis-attributable fraction for psychosis between 2 and 3%. Based on the estimates by Dealberto<sup>48</sup> and on the assumption of at least weekly cannabis use, this would correspond to 106–186 estimated cannabis-attributable incident cases of schizophrenia in Canada. This number is conservative in the sense that all other possible cannabis-attributable mental health problem outcomes are neglected; for psychosis specifically, this number may over- or underestimate the actual cannabis-attributable incidence.

### Cancer

While cannabis use may be associated with risk for other cancers, the main concern is lung cancer—the most common form of cancer in Canada. There are some 26 000 newly diagnosed cases of lung cancer in Canada per year, with a mortality rate of 80–85%.<sup>49</sup> While most studies find elevated risk for lung cancer, the quantification of the risk is difficult,<sup>50–53</sup> due to the following factors: (i) the high overlap with tobacco smoking—the primary risk factor for lung cancer, and the impossibility to fully adjust for confounding; (ii) most studies did not assess frequency and quantity of cannabis use; (iii) the

substantial lag-effect between risk and disease outcomes (~20 years). We based our estimates on Callaghan *et al.*<sup>53</sup> who included a follow-up of 40 years and adjustment for tobacco smoking (RR slightly >2 based on at least 50 cannabis use occasions). This would result in cannabis-attributable fractions of 1–2%, corresponding to 130–280 lung cancer deaths in Canada. Based on biological pathways, we would expect more cases of cannabis-attributable cancer in Canada,<sup>54,55</sup> but we are lacking the necessary data to quantify these relationships.

## Discussion

### Main findings of this study

In the above, we offered simple and crude estimates of the extent of cannabis-attributable morbidity and mortality in key risk domains for Canada. Main findings are that the largest number of cases with cannabis-related harm consequences appears to occur through motor-vehicle accidents and use disorders, whereas the number of cases of lung cancer and psychosis occur at a markedly smaller scale. In addition, only two of the domains—motor-vehicle accidents and lung cancer—result in mortality outcomes. Based on these considerations, we conclude on the basis of our estimates that motor-vehicle accidents and use disorders form the majority of instances of cannabis-attributable disease burden in Canada, whereas lung cancer—despite a limited amount of mortality in this category—and psychosis lead to substantially less extensive burdens.

### What is already known on this topic

Numerous previous studies have offered indicators of cannabis use-related relative risks for disease outcome in the main domains [e.g. refs (7,8)], although very few concrete and comprehensive estimates of actual disease burden exist which also allow for the comparison of actual outcome burden between the main domains. Previous studies from Canada and elsewhere have concluded that, when compared with other illicit drugs, the cannabis-related disease burden is comparably small despite relatively large population-level exposure.<sup>8,10,11,56</sup> For partial inter-domain comparisons, recent Global Burden of Disease (GBD) estimates focusing on cannabis use disorders have this burden estimated to be ~100 times larger than that for cannabis-related psychosis outcomes on a global level.<sup>57</sup>

### What this study adds

Our crude estimates at least point to a couple of key implications regarding the estimated extent of cannabis use-attributable disease outcomes in Canada. First, while cannabis use is not associated with direct acute mortality (e.g. from overdose/

poisoning), it may lead to fatalities from MVAs and cancer.<sup>58</sup> While these problems are rather different phenomena in terms of risk-behavioral characteristics—for example, cannabis use can be separated from the risk of impaired driving, while this is more difficult for cancer risks, at least in the predominant context of smoked cannabis—these should be key areas of attention for interventions given the (potentially extensive) mortality involved. Yet, while not involving attributable mortality outcomes, the extent of estimated cannabis use disorders is large, and therefore requires targeted and adequate interventions. In Canada's largest province of Ontario cannabis-related cases constitute a major proportion—30%—of overall admissions for substance use treatment, and treatment options and resources are limited.<sup>38</sup> Compared with the other domains, the extent of cannabis-related instances of psychosis—while undoubtedly a serious disease outcome in individual cases—can safely be assumed to constitute the smallest amount of health problem burden of the four domains assessed on a general population level in Canada. This is very much contrary to trends in popular media reporting and social debates, also concerning the potential consequences of liberalizing cannabis control policies.<sup>59–62</sup>

The overall assessment of the actual burden of cannabis-attributable health problems on a population level should be further developed and refined. While our calculations are relatively simple and crude, priority-setting and resource allocation for interventions should be informed by the above public health-oriented estimations. For example, the effectiveness and practical application of interventions to detect and deter cannabis-impaired driving need to be improved, and treatment options for cannabis use disorder need to be expanded in variety and availability.<sup>33,63–66</sup> At the same time, recent public health-oriented intervention tools have offered evidence-based recommendations on how to reduce user-behavioral risks for key cannabis-attributable harm outcomes, and other recommendations for public health-oriented cannabis interventions and policy development are available.<sup>12,14–16</sup>

### Limitations of this study

Our study was based on relatively crude and simple estimates, more detailed calculations for cannabis-attributable disease burden based on state-of-the-art methods in Canada need to be done.<sup>56</sup> These would include formal systematic reviews of the causality and level of the risk relations, focusing on evidence from Canada, and more detailed characterization of the relevant exposure dimensions. As many of the variables for estimating the actual extent of cannabis-attributable health harms are context specific, our estimate results are not generalizable to other settings or countries; however, similar exercises should be conducted elsewhere based on similar approaches.

### Acknowledgements

The authors acknowledge the skilled assistance of Chantal Burnett and Yoko Murphy in assembling this manuscript.

### Funding

This work was, in part, supported by a CIHR/PHAC Applied Public Health Chair award to B.F. (Canadian Institutes of Health Research, CIHR; grant number CPP-85657).

### References

- 1 Health Canada. Canadian Alcohol and Drug Use Monitoring Survey (CADUMS): Summary of Results for 2012. 2014; [http://www.hc-sc.gc.ca/hc-ps/drugs-drogues/stat/\\_2012/summary-sommaire-eng.php](http://www.hc-sc.gc.ca/hc-ps/drugs-drogues/stat/_2012/summary-sommaire-eng.php) (1 May 2014, date last accessed).
- 2 Ialomiteanu A, Adlaf EM, Hamilton H *et al.* 2011 *CAMH Monitor EReport: Addiction and Mental Health Indicators Among Ontario Adults 1977–2011*. Toronto: Centre for Addiction and Mental Health, 2012.
- 3 Room R. Legalising a market for cannabis for pleasure: Colorado, Washington, Uruguay and beyond. *Addiction* 2014;**109**(3):345–51.
- 4 Caulkins JP. *Marijuana Legalization: What Everyone Needs to Know*. New York: Oxford University Press, 2012.
- 5 Room R, Fischer B, Hall W *et al.* *Cannabis Policy: Moving Beyond Stalemate*. New York: Oxford University Press, 2010.
- 6 Lim SS, Aryee M, Laden F *et al.* A comparative risk assessment of burden of disease and injury attributable to 67 risk factors and risk factor clusters in 21 regions, 1990–2010: a systematic analysis for the Global Burden of Disease Study 2010. *Lancet* 2012;**380**(9859):2224–60.
- 7 Volkow ND, Baler RD, Compton WM *et al.* Adverse health effects of marijuana use. *N Engl J Med* 2014;**370**(23):2219–27.
- 8 Hall W, Degenhardt L. Adverse health effects of non-medical cannabis use. *Lancet* 2009;**374**(9698):1383–91.
- 9 Gordon AJ, Conley JW, Gordon JM. Medical consequences of marijuana use: a review of current literature. *Curr Psychiatry Rep* 2013;**15**(12):1–11.
- 10 Popova S, Rehm J, Patra J *et al.* Illegal drug-attributable morbidity in Canada 2002. *Drug Alcohol Rev* 2007;**26**:251–63.
- 11 Popova S, Rehm J, Patra J. Illegal drug-attributable mortality and potential years of life lost in Canada 2002: implications for prevention and policy. *Contemp Drug Probl* 2006;**33**(3):343–66.
- 12 Pacula RL, Kilmer B, Wagenaar AC *et al.* Developing public health regulations for marijuana: lessons from alcohol and tobacco. *Am J Public Health* 2014;**104**(6):1021–8.
- 13 Fischer B, Rehm J, Hall W. Cannabis use in Canada: the need for a 'public health' approach. *Can J Public Health* 2009;**100**(2):101–3.
- 14 Hall W, Degenhardt L. What are the policy implications of the evidence on cannabis and psychosis? *Can J Psychiatry* 2006;**51**(9):566–74.
- 15 Hall W, Pacula RL. *Cannabis Use and Dependence: Public Health and Public Policy*. Melbourne: Cambridge University Press, 2003.

- 16 Fischer B, Jeffries V, Hall W *et al.* Lower Risk Cannabis use Guidelines for Canada (LRCUG): a narrative review of evidence and recommendations. *Can J Public Health* 2011;**102**(5):324–7.
- 17 Rehm J, Baliunas D, Brochu S *et al.* *The Costs of Substance Abuse in Canada 2002*. Ottawa: Canadian Centre on Substance Abuse, 2006.
- 18 Ezzati M, Lopez AD, Rodgers A *et al.* *Comparative Quantification of Health Risks: Global and Regional Burden of Disease Attributable to Selected Major Risk Factors*. Geneva: World Health Organization, 2004.
- 19 Rehm J, Taylor B, Patra J *et al.* Avoidable burden of disease: conceptual and methodological issues in substance abuse epidemiology. *Int J Methods Psychiatr Res* 2006;**15**(4):181–91.
- 20 Walter SD. The estimation and interpretation of attributable risk in health research. *Biometrics* 1976;**32**:829–49.
- 21 Rockhill B, Newman B, Weinberg C. Use and misuse of population attributable fractions. *Am J Public Health* 1998;**88**(1):15–9.
- 22 Beirness DJ, Beasley EE. A roadside survey of alcohol and drug use among drivers in British Columbia. *Traffic Inj Prev* 2010;**11**(3):215–21.
- 23 Beasley EE, Beirness DJ. *Alcohol and Drug Use Among Drivers Following the Introduction of Immediate Roadside Prohibitions in British Columbia: Findings From the 2012 Roadside Survey*. Ottawa: Beirness & Associates, Inc., 2012.
- 24 Ramaekers JG, Berghaus G, van Laar M *et al.* Dose related risk of motor vehicle crashes after cannabis use. *Drug Alcohol Depend* 2004;**73**(2):109–19.
- 25 Asbridge M, Hayden JA, Cartwright JL. Acute cannabis consumption and motor vehicle collision risk: systematic review of observational studies and meta-analysis. *BMJ* 2012;**344**:e536.
- 26 Li M, Brady JE, DiMaggio CJ *et al.* Marijuana use and motor vehicle crashes. *Epidemiol Rev* 2012;**34**(1):65–72.
- 27 Asbridge M, Mann R, Cusimano MD *et al.* Cannabis and traffic collision risk: findings from a case-crossover study of injured drivers presenting to emergency departments. *Int J Public Health* 2014;**59**(2):395–404.
- 28 Brubacher JR, Chan H, Martz W *et al.* Recent drug use in injured British Columbia drivers. In: Presented at the Canadian Multi-disciplinary Road Safety Conference, Vancouver, BC, 2014.
- 29 Brubacher JR. Cannabis and motor vehicle crashes. *BCMj* 2011;**53**(6):292–3.
- 30 Pitel S, Solomon R. *Estimating the Number and Cost of Impairment-Related Traffic Crashes in Canada: 1999 to 2009*. Oakville: MADD Canada, 2012.
- 31 Transport Canada. *Canadian Motor Vehicle Traffic Collision Statistics, 2011*. Ottawa: Public Works and Government Services Canada, 2013.
- 32 Asbridge M, Mann R, Cusimano MD *et al.* Cycling-related crash risk and the role of cannabis and alcohol: a case-crossover study. *Prev Med* 2014;**66**:80–6.
- 33 Nordstrom BR, Levin FR. Treatment of cannabis use disorder: a review of the literature. *Am J Addict* 2007;**16**(5):331–42.
- 34 Copeland J, Swift W. Cannabis use disorder: epidemiology and management. *Int Rev Psychiatry* 2009;**21**(2):96–103.
- 35 Anthony J. The epidemiology of cannabis dependence. In: Roffman R, Stephens R (eds). *Cannabis Dependence: It's Nature, Consequences and Treatment*. Cambridge, UK: Cambridge University Press, 2006.
- 36 Lev-Ran S, Imtiaz S, Rehm J *et al.* Exploring the association between lifetime prevalence of mental illness and transition from substance use to substance use disorders: results from the National Epidemiologic Survey of Alcohol and Related Conditions (NESARC). *Am J Addict* 2013;**22**(2):93–8.
- 37 Pearson C, Janz T, Ali J. *Health at a Glance: Mental and Substance Use Disorders in Canada*. Ottawa: Minister of Industry, 2013.
- 38 Drug and Alcohol Treatment Information System (DATIS). *Substance Abuse Statistical Tables: Fiscal Year 2007/2008 to Fiscal Year 2012/2013*. Toronto: Centre for Addiction and Mental Health, 2013.
- 39 Grant BF, Stinson FS, Dawson DA *et al.* Prevalence and co-occurrence of substance use disorders and independent mood and anxiety disorders: results from the National Epidemiologic Survey on Alcohol and Related Conditions. *Arch Gen Psychiatry* 2004;**61**:807–16.
- 40 Jane-Llopis E, Matytsina I. Mental health and alcohol, drugs and tobacco: a review of the comorbidity between mental disorders and the use of alcohol, tobacco and illicit drugs. *Drug Alcohol Rev* 2006;**25**(6):515–36.
- 41 Rehm J, Room R, Monteiro M *et al.* Alcohol use. In: Ezzati M, Lopez AD, Rodgers A, Murray CJL (eds). *Comparative Quantification of Health Risks. Global and Regional Burden of Disease Attributable to Selected Major Risk Factors*, Vol. 1. Geneva: World Health Organization, 2004, 959–1108.
- 42 Rehm J, Baliunas D, Borges GLG *et al.* The relation between different dimensions of alcohol consumption and burden of disease—an overview. *Addiction* 2010;**105**(5):817–43.
- 43 Moore THM, Zammit S, Lingford-Hughes A *et al.* Cannabis use and risk of psychotic or affective mental health outcomes: a systematic review. *Lancet* 2007;**370**(9584):319–28.
- 44 Zammit S, Moore THM, Lingford-Hughes A *et al.* Effects of cannabis use on outcomes of psychotic disorders: systematic review. *Br J Psychiatry* 2008;**193**(5):357–63.
- 45 D'Souza DC, Sewell RA, Ranganathan M. Cannabis and psychosis/schizophrenia: human studies. *Eur Arch Psychiatry Clin Neurosci* 2009;**259**(7):413–31.
- 46 Degenhardt L, Hall WD, Lynskey M *et al.* Should burden of disease estimates include cannabis use as a risk factor for psychosis? *PLoS Med* 2009;**6**(9):e1000133.
- 47 Peto R, Boreham J, Lopez AD *et al.* Mortality from tobacco in developed countries: indirect estimation from national vital statistics. *Lancet* 1992;**339**(8804):1268–78.
- 48 Dealberto M. Are the rates of schizophrenia unusually high in Canada? A comparison of Canadian and international data. *Psychiatry Res* 2013;**209**(3):259–65.
- 49 Canadian Cancer Society's Advisory Committee on Cancer Statistics. *Canadian Cancer Statistics 2014*. Toronto: Canadian Cancer Society, 2014.
- 50 Mehra R, Moore BA, Crothers K *et al.* The association between marijuana smoking and lung cancer: a systematic review. *Arch Intern Med* 2006;**166**(13):1359–67.
- 51 Hashibe M, Straif K, Tashkin DP *et al.* Epidemiologic review of marijuana use and cancer risk: a systematic review. *Alcohol* 2005;**35**(3):265–75.
- 52 Berthiller J, Bartsch M, Ayed FB *et al.* Cannabis smoking and risk of lung cancer in men: a pooled analysis of three studies in Maghreb. *J Thorac Oncol* 2008;**3**(12):1398–403.

- 53 Callaghan RC, Allebeck P, Sidorchuk A. Marijuana use and risk of lung cancer: a 40-year cohort study. *Cancer Causes Control* 2013; **24(10)**:1811–20.
- 54 Sturgis EM, Cinciripini PM. Trends in head and neck cancer incidence in relation to smoking prevalence. *Cancer* 2007;**110(7)**:1429–35.
- 55 Hall W, MacPhee D. Cannabis use and cancer. *Addiction* 2002; **97(3)**:243–7.
- 56 Degenhardt L, Hall W. Extent of illicit drug use and dependence, and their contribution to the global burden of disease. *Lancet* 2012; **379(9810)**:55–70.
- 57 Degenhardt L, Whiteford HA, Vos T *et al.* The global epidemiology and contribution of cannabis use and dependence to the global burden of disease: results from the GBD 2010 study. *PLoS One* 2013;**8(10)**:e76635.
- 58 Calabria B, Degenhardt L, Hall W *et al.* Does cannabis use increase the risk of death? Systematic review of epidemiological evidence on adverse effects of cannabis use. *Drug Alcohol Rev* 2010; **29(3)**:318–30.
- 59 Schizophrenia Society of Canada. Cannabis & Psychosis: Exploring the link. 2011; <http://cannabisandpsychosis.ca/> (1 October 2014, date last accessed).
- 60 Cole K. Canadian study links cannabis and psychosis in youth. 2013; <http://www.madinamerica.com/2013/12/canadian-study-links-cannabis-psychosis/> (1 October 2014, date last accessed).
- 61 CBC News. Information/facts about marijuana use and mental illness. 2013; <http://www.cbc.ca/natureofthings/features/information-facts-about-marijuana-use-mental-illness> (1 October 2014, date last accessed).
- 62 Canadian Centre on Substance Abuse. Clearing the smoke on cannabis series: highlights. 2014; <http://www.ccsa.ca/Resource%20Library/CCSA-Clearing-Smoke-on-Cannabis-Highlights-2013-en.pdf> (1 October 2014, date last accessed).
- 63 Freeman J, Watling C, Davey J *et al.* Perceptual deterrence versus current behaviours: a study into factors influencing drug driving in Queensland. *Road Trans Res* 2010;**19(3)**:3–13.
- 64 DuPont RL, Voas RB, Walsh JM *et al.* The need for drugged driving per se laws: a commentary. *Traffic Inj Prev* 2012;**13(1)**:31–42.
- 65 McRae A, Budney A, Brady K. Treatment of marijuana dependence: a review of the literature. *J Subst Abuse Treat* 2003;**24(4)**:369–76.
- 66 Hart CL. Increasing treatment options for cannabis dependence: a review of potential pharmacotherapies. *Drug Alcohol Depend* 2005; **80(2)**:147–59.

## Advertising and Corporate Services

In association with our leading journals, we offer tailored services to industry, including:

- Print, online and app advertising, with options to suit every campaign and budget
- Customized bulk article reprints and digital ePrints: excellent as handouts at conferences and to distribute via email or your website
- Sponsored peer-reviewed supplements and subscriptions
- Sponsored educational websites (microsites)
- Recruitment advertising in the *Journals Career Network*
- Custom book publishing - from our prestigious portfolio of over 2,000 medical titles, we can provide branded, tailored, or even bespoke publications, specific to your target market

Please contact Corporate Services:

- [www.oupmediainfo.com](http://www.oupmediainfo.com)
- [corporate.services@oup.com](mailto:corporate.services@oup.com)