



CJC Open 4 (2022) 206-213

Original Article

Increasing Prevalence and Incidence of Atherosclerotic Cardiovascular Disease in Adult Patients in Ontario, Canada From 2002 to 2018

Erin S. Mackinnon, PhD,^a Ron Goeree, BA, MA,^{b,c} Shaun G. Goodman, MD, MSc,^d Raina M. Rogoza, MSc,^a Millicent Packalen, BSc(Hons),^a Louisa Pericleous, PhD,^a Ponda Motsepe-Ditshego, MD,^a and Paul Oh, MD, MSc, FRCPC^e

^a Amgen Canada Inc, Mississauga, Ontario, Canada

^b Goeree Consulting Ltd, Hamilton, Ontario, Canada

^c Professor Emeritus, McMaster University, Hamilton, Ontario, Canada

^d St. Michael's Hospital, University of Toronto, Toronto, Ontario, Canada

^e Toronto Rehabilitation Institute, Toronto, Ontario, Canada

ABSTRACT

Background: Cardiovascular disease is the second-leading cause of death in Canada. However, limited data are available on the prevalence of atherosclerotic cardiovascular disease (ASCVD) in Canada. The study objective was to describe the incidence and prevalence of ASCVD in adult patients in Ontario, Canada, and to evaluate temporal trends for subsequent ASCVD events among those with new-onset ASCVD.

Methods: This retrospective, observational study identified ASCVD incidence and prevalence data from the Institute for Clinical Evaluative Sciences Data Repository for adults from Ontario. Overall prevalence was established for the period from 2002 to 2018. Incident cases from April 1, 2005 to March 2016 were then identified, and followed

RÉSUMÉ

Contexte : Les maladies cardiovasculaires constituent la deuxième cause de décès au Canada. Toutefois, on dispose de peu de données sur la prévalence de la maladie cardiovasculaire athéroscléreuse (MCVAS) au Canada. L'étude avait pour objectifs de décrire l'incidence et la prévalence de la MCVAS chez les patients adultes en Ontario (Canada) et d'évaluer les tendances temporelles des manifestations subséquentes de MCVAS chez les personnes ayant une MCVAS d'apparition récente.

Méthodologie : Cette étude observationnelle rétrospective a permis de colliger les données sur l'incidence et la prévalence de la MCVAS chez les adultes ontariens consignées dans le référentiel de l'Institut des sciences cliniques évaluatives. La prévalence globale a été établie

Cardiovascular disease (CVD) is the leading cause of death worldwide; an estimated 17.9 million people died from CVD in 2016, representing 31% of all global mortality.¹ Atherosclerotic cardiovascular disease (ASCVD) typically encompasses a range of cerebral, cardiac, and peripheral vascular diagnoses, events, and revascularization procedures.^{2,3} Of these, myocardial infarction (MI) is highly prevalent and is one of the predominant causes of all inpatient hospitalizations

Received for publication August 3, 2021. Accepted October 5, 2021.

E-mail: emackinn@amgen.com

See page 213 for disclosure information.

in Canada.^{4,5} Patients with ASCVD have a high risk of cardiovascular events and recurrent events.⁶⁻⁸ Indeed, atherosclerosis is responsible for ischemic heart disease and stroke, which were identified as the leading causes of mortality after cancer in Canada and accounted for approximately 1 of 4 deaths in 2018.⁹

Globally over the past few decades, CVD event and mortality rates have decreased.^{10,11} In Canada, although the mortality rate due to ischemic heart disease has been steadily decreasing since 2000, the age-standardized prevalence of diagnosed ischemic heart disease has remained stable, being 7.1% in 2000 and 7.8% in 2015.¹² A study conducted by the Cardiovascular Health in Ambulatory Care Research Team (CANHEART) initiative in Ontario, Canada, showed that despite improvements in ASCVD diagnosis and treatment guidelines, the average decline in annual rates during 1994-2012 in hospitalizations and deaths due to major

https://doi.org/10.1016/j.cjco.2021.10.003

Ethics Statement: The study protocol was reviewed and approved by the Advarra Institutional Review Board.

Corresponding author: Dr Erin S. Mackinnon, Amgen Canada Inc, 6775 Financial Dr, Suite 100, Mississauga, Ontario L5N 0A4, Canada. Tel.: +1-905-285-3478.

²⁵⁸⁹⁻⁷⁹⁰X/© 2021 The Authors. Published by Elsevier Inc. on behalf of the Canadian Cardiovascular Society. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/).

up to 2018. Primary outcomes were date and type of index event/ procedure, patient characteristics/baseline demographics, and comorbidities. Secondary outcomes assessed were time from first to second ASCVD event, subsequent event(s) and/or mortality, and type of subsequent event(s) relative to the type of index/primary event. Results: A total of 1,042,621 eligible prevalent ASCVD cases were identified; of these, 743,309 patients (69%) were newly diagnosed with incident ASCVD. The 10-year prevalence rates for all ASCVD subtypes increased over the study period. Overall event incidence rates per 1000 person-years were mostly stable or increased. Among incident cases, 50% experienced subsequent events over the study period. Conclusions: This observational study demonstrated increasing prevalence and high incidence of new ASCVD diagnoses in adults from Ontario, over the study period. These data, together with the substantial number of subsequent events in ASCVD patients, demonstrate significant clinical burden of this disease in Ontario.

ASCVD events was < 2% in adults aged < 50 years, accompanied by a slight decrease in self-reported prevalence rates of heart disease and stroke.¹³ Another Canadian study from Quebec demonstrated a decline in the age-standardized incidence and associated mortality of vascular diseases from 2000 to 2015, but an increase in their prevalence.¹⁴

A recent prospective study from Ontario investigated the implications of failure to achieve guideline-recommended lowdensity lipoprotein cholesterol (LDL-C) levels in patients who have undergone percutaneous coronary intervention (PCI). This study demonstrated a significant association between higher levels of LDL-C and increased incidence of CV events.¹⁵ Such data describing the prevalence and incidence rates (IRs) associated with individual ASCVD subtypes and temporal trends in disease progression in Canada are limited. Therefore, the objective of the present study was to describe the incidence and prevalence of ASCVD in adult patients in Ontario, and to characterize and evaluate temporal trends for subsequent ASCVD events.

Materials and Methods

Study design, population, and data source

This retrospective, observational study used administrative health data from the province of Ontario, Canada maintained at the Institute for Clinical Evaluative Sciences (ICES) Data Repository (Supplemental Table S1). This study focused on identification and characterization of prevalence and incidence data of ASCVD events and procedures.

Incidence data for all adult patients with an initial (index) ASCVD event between April 1, 2005 and March 31, 2016 (Supplemental Fig. S1) were identified from the ICES pour la période allant de 2002 à 2018. Les cas incidents survenus du 1^{er} avril 2005 à mars 2016 ont ensuite été recensés et suivis jusqu'en 2018. Les paramètres d'évaluation principaux étaient : date et nature de la manifestation index et de l'intervention; caractéristiques des patients et données démographiques initiales; comorbidités. Les éléments suivants constituaient les paramètres d'évaluation secondaires : temps écoulé entre la première et la deuxième manifestation de MCVAS, la ou les manifestations subséquentes par rapport à celle de la manifestation index ou primaire.

Résultats : En tout, 1042 621 cas de MCVAS prévalents admissibles ont été dénombrés; parmi ceux-ci, il y avait 743 309 (69 %) cas incidents de MCVAS nouvellement diagnostiqués. Les taux de prévalence à 10 ans de tous les sous-types de MCVAS ont augmenté au cours de la période étudiée. Les taux globaux d'incidence des manifestations par 1000 années-personnes étaient généralement stables ou accrus. Cinquante pour cent des cas incidents ont été associés à des manifestations subséquentes au cours de la période étudiée.

Conclusions : Cette étude observationnelle a démontré une prévalence croissante et une incidence élevée de nouveaux diagnostics de MCVAS chez les adultes en Ontario au cours de la période étudiée. Les données à cet égard, ainsi que le grand nombre de manifestations subséquentes de la maladie, démontrent que la MCVAS constitue un fardeau clinique considérable en Ontario.

database and included in this analysis. To determine if these were new (incident) events, patients with a diagnostic code for an ASCVD event within a 2-year lookback period (from April 1, 2003) were excluded from the incident cases. This exclusion was used to ensure that only index events (ie, first ASCVD event) were included; it also provided patient baseline characteristics for this cohort. ASCVD incidence was then calculated for the time period 2005 to 2016. The incidence of ASCVD was described by year as the number of new cases of ASCVD identified from April 1 to March 31 of the following year as a proportion of the population present within the ICES database (adults in Ontario aged \geq 18 years) who did not have ASCVD. Follow-up on these incident cases was done until 2018, to gain an understanding of subsequent ASCVD events.

Overall prevalence for the study cohort was established by identification of patients given a diagnosis of ASCVD between April 1, 2002 and March 31, 2018 (Supplemental Figure S1). Prevalent ASCVD cases were then assessed and reported as 5- and 10-year prevalence. The overall prevalence over time of ASCVD was described as the number of cases (existing and new) of ASCVD identified from April 1 to March 31 (numerator) of the given period as a proportion of the population present within the entire ICES database (adults in Ontario aged \geq 18 years; denominator).

Based on the definition in the 2016 Canadian Cardiovascular Society (CCS) dyslipidemia management guidelines,² in this study, ASCVD included coronary, cerebrovascular, and peripheral events. Coronary events included MI, PCI, and coronary artery bypass graft surgery (CABG), as well as a hospitalization or emergency room visit for unstable angina. Cerebrovascular events included ischemic stroke, transient ischemic attack (TIA), and carotid endarterectomy/stenting. Peripheral events included symptomatic peripheral artery disease (PAD), peripheral artery angioplasty/stenting, peripheral artery bypass surgery, peripheral artery endarterectomy, and aortic aneurysm.

Study variables and outcomes

Variables of interest were extracted from the ICES databases up to 2 years prior to the date of the index event.

Date and type of event were identified using diagnostic codes from the International Classification of Diseases, versions 9 and 10 (ICD-9 and -10; Supplemental Table S2). Patient characteristics at the time of hospitalization for events were collected, including average age at the time of the index event/procedure, gender, and LDL-C test results. Diabetes, hypertension, and hypercholesterolemia comorbidities were identified by ICD coding and ICES predefined cohorts/databases; respective Charlson Comorbidity Index (CCI)¹⁶ scores were extracted.

Secondary outcomes assessed were the time from the first ASCVD event to the second event, subsequent event(s) and/ or mortality, and type of second and/or subsequent event(s) relative to the type of index/primary event. Event types within each of the groups (coronary, cerebrovascular, and peripheral events/procedures) were considered related. Related events occurring within 30 days were counted as one event, with precedence given to the event that came first in the hierarchical order listed above. For instances in which unrelated events occurred within 30 days of each other, both events were counted (the first was recorded as the index event, and the second as a subsequent event). Data were collected for deaths occurring within the given study period. For patients with a date of death, the time from the index event to the date of death was recorded. Deaths (all-cause mortality) were extracted as subsequent events only. Deaths that occurred within 30 days of an MI, aortic aneurysm, or ischemic stroke were considered related to those events and were not counted as a subsequent event. Deaths that occurred more than 30 days after these related events, and deaths that occurred within 30 days of unrelated events, were counted as subsequent events.

Statistical methods and ethics approval

Descriptive statistics were used to describe the study outcomes, including means, medians, and ranges. Event rates were calculated per 1000 person-years. No statistical analyses were performed to assess significance. The study protocol was reviewed and approved by the Advarra Institutional Review Board.

Results

Description of study cohort

A schematic overview of the patients described in this study is presented in Figure 1. Based on ICD diagnostic codes, 1,042,621 eligible prevalent cases were identified from the ICES database with a diagnosis of ASCVD between April 1, 2002 and March 31, 2018, providing a strong sense of overall ASCVD prevalence in this population. Of these, 743,309 patients (69%) were newly diagnosed with incident ASCVD between April 1, 2005 and March 31, 2016. These latter incidence data were analyzed separately to describe annual outcomes relative to the index event.

Patient characteristics

Characteristics of patients included in this study are described in Table 1. The majority of patients who experienced an ASCVD event were ≥ 65 years of age, among both prevalent cases (634,928 of 1,042,621 [61%]) and incident cases (458,569 of 743,309 [62%]). Among the incident ASCVD cases, the majority were male (59%), the median age was 69 years (interquartile range [IQR]: 59-79 years), and the most frequent events/procedures were MI (30%), unstable angina (17%), TIA (16%), stroke (11%), PCI (9%), PAD (5%), and CABG (5%). Similar data were reported for the prevalent ASCVD cases, shown in Table 1. Concomitant hypertension (71%-73%) and/or diabetes (31%-32%) were common among both the prevalent and incident ASCVD cases.

Incidence of ASCVD subtypes in Ontario

The overall event IRs per 1000 person-years (IRs) were calculated for each ASCVD subtype over the study period; IRs for all events ranged from 7.2 to 8.8 over the study period (Fig. 2; Supplemental Fig. S2). The IRs for MI, PAD, and aortic aneurysm remained stable at approximately 2.5, 0.7, and 0.4, respectively, over the study period; however, the IRs increased steadily for PCI/CABG (from 2.5 in 2005 to 3.0 in 2016), TIA (from 0.9 in 2005 to 1.5 in 2016), and stroke (from 0.7 in 2005 to 1.5 in 2016) every year across the study period (Fig. 2). IRs for unstable angina decreased over the course of the study period (from 3.3 in 2005 to 1.7 in 2016).

The annual IR for the first event was calculated per 1000 individuals and analyzed for the total incident cases and stratified by age (< 65 years and ≥ 65 years; Fig. 3; Supplemental Fig. S3). The IRs for the first event were higher among patients aged ≥ 65 years than those aged < 65 years; however, the trends were largely similar in the 2 age groups. The IRs of index MI were consistently higher than those for other ASCVD subtypes across the 11-year study period (Fig. 3). Of note, the annual incidence of index MI events was nearly halved in older patients aged ≥ 65 years (from 8.4 per 1000 individuals in 2005 to 4.8 per 1000 individuals in 2015; Fig. 3B), whereas in patients aged < 65 years, it remained stable (1.0 per 1000 individuals in 2005 to 0.9 per 1000 individuals in 2015; Fig. 3C). The annual IR for index unstable angina also decreased substantially in patients aged \geq 65 years (from 7.5 per 1000 individuals in 2005 to 2.1 per 1000 individuals in 2015; Fig. 3B), as well as in patients aged < 65 years (from 0.9 per 1000 individuals in 2005 to 0.4 per 1000 individuals in 2015; Fig. 3C). However, IRs doubled for index TIA (from 0.26 per 1000 individuals in 2005 to 0.5 per 1000 individuals in 2015) and stroke (from 0.2 per 1000 individuals in 2005 to 0.4 per 1000 individuals in 2015) in patients aged < 65 years (Fig. 3C).

Prevalence of ASCVD subtypes in Ontario

The 10-year prevalence rates for all ASCVD subtypes increased over the study period (Fig. 4; Supplemental Fig. S4). The highest rates were seen for PCI/CABG, MI, and unstable angina, which respectively increased from 24.5 to 31.8, 23.5

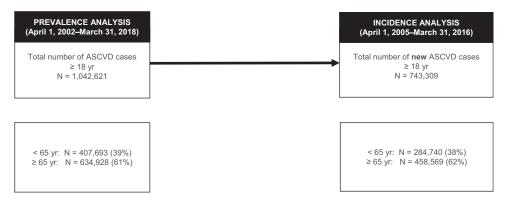


Figure 1. Number of eligible adult patients with atherosclerotic cardiovascular disease (ASCVD) in Ontario for whom data were extracted. The data were extracted from the ICES databases up to 2 years prior to the date of the index event (ie, lookback date from April 1, 2002) and included all data available in the database at the time of extraction (maximum follow-up date March 31, 2018).

to 26.9, and 22.1 to 23.7 per 1000 individuals, between the periods 2004-2013 and 2008-2017 (Fig. 4). The 10-year prevalence rates for all events changed from 64.2 per 1000 individuals to 75.5 per 1000 individuals, respectively (data not shown). The prevalence of ASCVD per 1000 individuals in Ontario for the period 2004-2017 was 78.2 (95% confidence interval [CI]; 78.1, 78.4; Supplemental Table S3). The prevalence per 1000 individuals was greater for coronary events (eg, MI, unstable angina, PCI) and cerebrovascular events (eg, TIA, stroke), compared with that of peripheral artery events. Similarly, 5-year prevalence rates for all ASCVD subtypes increased over the study period, changing from 43.7 per 1000 individuals for the period 2012-2016 and 69.1 per 1000 individuals for the period 2013-2017 (data not shown).

Subsequent events and mortality after index incident ASCVD event

Half (374,715 of 743,309) of all incident cases experienced only one event over the study period, whereas 33% (242,698 of 743,309) had 2 ASCVD events, and 17% (125,896 of 743,309) had 3 or more ASCVD events (Table 2). Analysis of index cerebrovascular or coronary events revealed that approximately half of all patients had a subsequent event within the study period for the following subtypes by index event: MI (48%; 106,057 of 221,429); unstable angina (53%; 68,459 of 128,490); TIA (44%; 51,903 of 117,421); and stroke (48%; 40,368 of 84,739; Fig. 5, A and B). Overall, 19% (143,649 of 743,309) of patients died following their index ASCVD event within the study period (data not

Characteristic	ASCVD prevalence (N = $1,042,621$)	ASCVD incidence (N = $743,309$)	
Female sex	432,300 (41.5)	306,954 (41.3)	
Age at index/primary ASCVD event, $y^{*,\dagger}$			
Median (IQR)	69 (58-79)	69 (59-79)	
18-64	407,693 (39.1)	284,740 (38.3)	
65-74	260,454 (25.0)	181,780 (24.5)	
75-84	249,886 (24.0)	180,756 (24.3)	
≥ 85	124,588 (11.9)	96,033 (12.9)	
Selected comorbidities			
Hypertension	738,861 (70.9)	539,787 (72.6)	
Diabetes	322,926 (31.0)	239,529 (32.2)	
Index ASCVD event*			
MI	304,326 (29.2)	221,429 (29.8)	
Unstable angina	197,481 (18.9)	128,490 (17.3)	
PCI/CABG surgery	138,494 (13.3)	100,590 (13.5)	
Aortic aneurysm	42,632 (4.1)	30,878 (4.2)	
PAD	57,247 (5.5)	39,543 (5.3)	
Peripheral artery angioplasty/ stenting	12,001 (1.2)	9241 (1.2)	
Peripheral artery bypass surgery	4483 (0.4)	3357 (0.5)	
Peripheral artery endarterectomy	1283 (0.1)	952 (0.1)	
TIA	157,066 (15.1)	117,421 (15.8)	
Stroke	118,015 (11.3)	84,739 (11.4)	
Carotid endarterectomy/stenting	9593 (0.9)	6669 (0.9)	

Table 1. Characteristics of patients in the study

Values are n (%), unless otherwise indicated. Study methods used precluded statistical comparisons being performed.

ASCVD, atherosclerotic cardiovascular disease; CABG, coronary artery bypass graft; IQR, interquartile range; MI, myocardial infraction; PAD, peripheral arterial disease; PCI, percutaneous coronary intervention; TIA, transient ischemic attack.

* An event was considered an index event if it occurred in a patient aged \geq 18 years, from April 1, 2005 to March 31, 2016.

^{\dagger} Proportion of patients aged \geq 65 years: 61% (634,928 of 1,042,621) with prevalent ASCVD; 62% (458,569 of 743,309) with incident ASCVD.

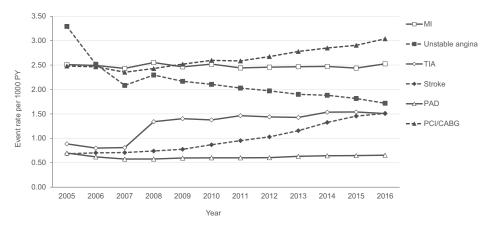


Figure 2. Incident atherosclerotic cardiovascular disease event rate per 1000 person-years (PY) in Ontario (2005/2006 to 2016/2017; for subtypes > 5% of all cases). CABG, coronary bypass graft; MI, myocardial infarction; PAD, peripheral artery disease; PCI, percutaneous coronary intervention; TIA, transient ischemic attack.

shown). Rates of mortality as a second event were highest for those with index events of PAD (26.8%; n = 10,589), stroke (27.1%; n = 22,990), unstable angina (18.3%; n = 23,478), and MI (17.8%; n = 39,320).

Type of subsequent event relative to index event

Second ASCVD events were most common among patients whose index event was MI (n = 106,057), followed by unstable angina (n = 68,459), TIA (n = 51,903), stroke (n =40,368), and PAD (n = 27,151; Supplemental Fig. S5). Subtypes of subsequent events are reported in Supplemental Figure S6, A-C. In general, index events commonly led to subsequent coronary events; of note, experiencing an incident cerebrovascular or peripheral event conferred a risk of having a second coronary event. For example, among those with a peripheral index procedure, 12.2% had a subsequent coronary event. Analysis by individual subtype revealed that patients most frequently had an identical diagnosis for both first and second events (data not shown).

Time to event/death

Of the incident cases, the median time between the index and second event was 1.66 years (IQR: 0.35-4.16 years; 368,594 of 743,309 patients); the median time between the second and third event was 1.10 years (IQR: 0.29-3.00 years; 125,896 of 743,309 patients) (Supplemental Fig. S7). The median time between the index event and death (including deaths occurring after the third event) was 2.16 years (IQR: 0.25-5.11 years), and between the last event and death, it was 2.08 years (IQR: 0.55-4.57 years).

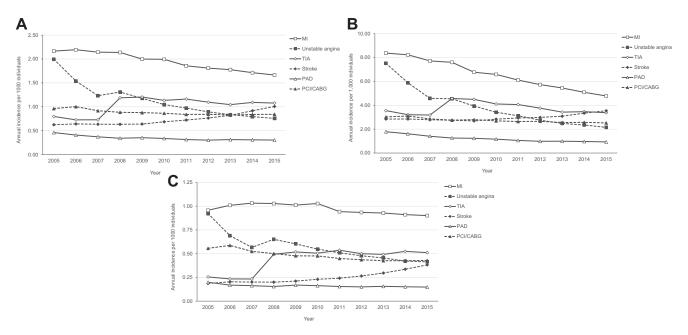


Figure 3. Annual incidence of index atherosclerotic cardiovascular disease event per 1000 individuals in Ontario (2005/2006 to 2015/2016; for subtypes > 5% of all cases), for (A) all incident cases; (B) patients aged ≥ 65 years; (C) patients aged < 65 years. CABG, coronary bypass graft; MI, myocardial infarction; PAD, peripheral artery disease; PCI, percutaneous coronary intervention; TIA, transient ischemic attack.

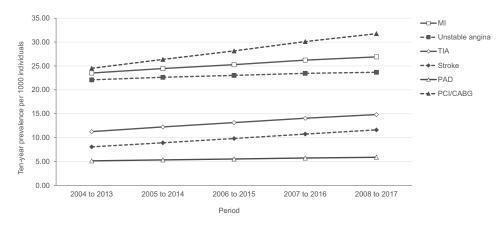


Figure 4. Ten-year prevalence of atherosclerotic cardiovascular disease in Ontario (for subtypes > 5% of all cases). Prevalence rates are for the entire population. CABG, coronary artery bypass graft; MI, myocardial infarction; PAD, peripheral artery disease; PCI, percutaneous coronary intervention; TIA, transient ischemic attack.

Discussion

This is the first comprehensive analysis of ASCVD prevalence and incidence in a cohort of adult patients in Ontario, Canada. This study spans a substantial period of time, demonstrating the progression of 12 subtypes of ASCVD events and procedures that have not been reported to date in Ontario. Further, the findings of this study demonstrate increased 10-year prevalence rates over time for all ASCVD subtypes, with stable or increased annual IRs observed for most events. A recent cohort study of adult patients within a large, diverse US healthcare system demonstrated an ASCVD IR of 8.67 (95% confidence interval 8.61-8.74) per 1000 person-years, similar to the observed range of ASCVD incidence in this study (7.2-8.8 per 1000 person-years). In the present study, 10-year prevalence rates increased from 64.2 per 1000 individuals in 2004-2013 to 75.5 per 1000 individuals in 2008-2017, which represents a 17.6% increase. The trend of increased ASCVD prevalence over time mirrors that of another recent Canadian study that showed a 21.4% increase in the age-standardized prevalence of vascular diseases between 2000 and 2015.

The rate of subsequent events among the incident cases over the follow-up period was approximately 50%—those with ASCVD and additional cardiovascular risk enhancers, including polyvascular disease, symptomatic PAD, history of MI, MI in the past 2 years, and previous CABG surgery, per the 2021 CCS Guidelines for the Management of Dyslipidemia for the Prevention of Cardiovascular Disease in the Adult.¹⁸ In our study, the second event was more likely to be of the same class as the first (ie, coronary, cerebrovascular, peripheral). However, regardless of the index event, these patients had a substantial risk of having a subsequent coronary or cerebrovascular event, further supporting their classification as very high risk, as outlined by the CCS.¹⁸ Overall, 1 in 5 patients diagnosed with ASCVD in the incidence cohort died over the study period. Of the 143,649 deaths, 1 in 4 occurred within 3 months of the patient's first ASCVD event. In line with CCS and other global guidelines, these findings indicate that patients who experience an event are at increased risk of death (especially within the first 3 months after the event), and survivors are at risk of further subsequent events. Of note, patients who experienced peripheral events/procedures had the greatest risk of subsequent ASCVD diagnoses or death, highlighting a care gap that requires urgent attention. The median time between the index event and death was approximately 2.2 years, whereas the median time between the last event and death was approximately 2.1 years. This increased risk of mortality within 2 years post-ASCVD event is consistent with that found in other real-world analyses,¹⁹ and it reinforces the urgency of treating these patients using guideline-recommended targets.

Over the 11-year study period, MI was the most common index event, followed by unstable angina, TIA, stroke, PCI, and PAD. Patients with an index MI event had the greatest

Table 2. Number of ASCVD events reported over the study period, by age group

Characteristic	All events* (N = $743,309$)	1 event [†] (n = 374,715)	2 events ^{\dagger} (n = 242,698)	\geq 3 events [†] (n = 125,896)
Female sex	306,954 (41.3)	153,622 (41.0)	104,115 (42.9)	49,217 (39.1)
Age at the time of index ASCVD				
event, y				
Median (IQR)	69 (59-79)	65 (55-75)	75 (64-83)	72 (62-80)
18-64	284,740 (38.3)	184,628 (49.3)	62,938 (25.9)	37,174 (29.5)
65-74	181,780 (24.5)	92,374 (24.7)	55,775 (23.0)	33,631 (26.7)
75-84	180,756 (24.3)	67,893 (18.1)	74,364 (30.6)	38,499 (30.6)
≥ 85	96,033 (12.9)	29,820 (8.0)	49,621 (20.4)	16,592 (13.2)

Values are n (%), unless otherwise indicated.

ASCVD, atherosclerotic cardiovascular disease; IQR, interquartile range.

* For the period from 2005/2006 to 2015/2016.

[†] For the period from 2005/2006 to 2017/2018.

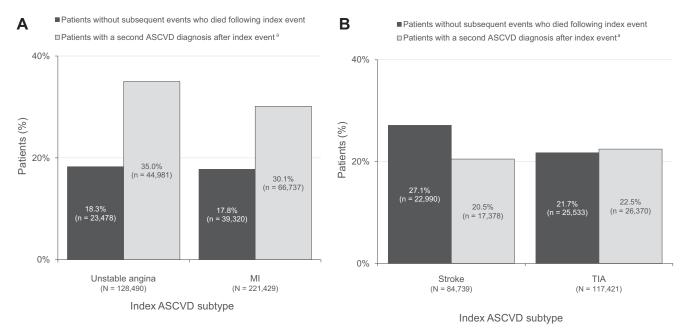


Figure 5. Second atherosclerotic cardiovascular disease (ASCVD) events based on index ASCVD subtype: (A) index coronary event; (B) index cerebrovascular event. MI, myocardial infarction; TIA, transient ischemic attack. ^aData are plotted for patients with additional ASCVD diagnoses after index event who survived the second event.

number of second and third events of any type, and the mortality rate was 17.8% (39,320 of 221,429). These findings are consistent with a recent study demonstrating that almost one-third of patients with a first event had a subsequent event; in those with an initial MI, subsequent MI was the most common secondary event.²⁰ Furthermore, these findings support previous real-world evidence of increased cardiovascular risk in patients post-MI,8 and they emphasize the burden of disease and the importance of addressing MI with aggressive treatment. CCS guideline-recommended treatment strategies can be implemented using LDL-C levels to guide therapeutic goals,² as well as other secondary prevention strategies. Of note, higher annual MI incidence was observed in earlier years for the patients aged \leq years. However, this measure only captured index MI events; as a result, those patients who had greatest residual risk and went on to have further events were excluded from this analysis. The lower incidence in MI events in younger patients may be explained by the fact that events likely are not as common in the younger group, and many of these patients may experience single events only.

Limitations

Limitations of this retrospective study include the quality and accuracy issues inherently associated with collecting data in a real-world setting. In addition, patient data may be incomplete due to their relocation to other provinces and/or countries; consequently, the study may not have included data for all eligible Ontario residents. Similarly, the incidence and prevalence of ASCVD in Ontario may be under-represented due to missed events if a patient with ASCVD did not present to a hospital or visit a physician. The data collection for the index ASCVD event began in 2005, with a lookback of 2 years for prior events; however, the index events herein are not guaranteed to be the first events for each patient. When describing the incidence of index ASCVD event by type, patients are limited to a single "first event," with more common events such as unstable angina potentially limiting the reporting of other subsequent ASCVD event types, due to their hierarchical order (as outlined in the Materials and Methods section), and death was excluded as an index event. Patients who were not as healthy could experience these initial first events early in the study time period, and hence, they were removed from the pool of patients able to have first ASCVD events, and potentially events of greater severity, in subsequent years. ASCVD event rates were calculated per 1000 person-years, but they were not standardized for age, which may lead to inaccuracies in reporting trends over time. Finally, the comparisons performed in this study are descriptive in nature.

Conclusions

This retrospective, observational study explored the prevalence and incidence of ASCVD in Ontario, Canada in patients aged \geq 18 years throughout 2002-2018, demonstrating a prevalence of more than 1 million cases of ASCVD. Prevalence rates are increasing, and there is a high IR for new ASCVD diagnoses, highlighting opportunities to continue to close the care gap and improve outcomes in ASCVD patients in Canada.

Acknowledgements

This study made use of de-identified data from the ICES Data Repository, with support from its funders and partners: Canada's Strategy for Patient-Oriented Research (SPOR), the Ontario SPOR Support Unit, the Canadian Institutes of Health Research, Canada, and the Government of Ontario. The opinions, results, and conclusions reported are those of the authors. No endorsement by ICES or any of its funders or partners is intended or should be inferred. Parts of this material are based on data and/or information compiled and provided by the Canadian Institute for Health Information (CIHI). However, the analyses, conclusions, opinions, and statements expressed in the material are those of the author(s), and not necessarily those of CIHI. Medical writing support was provided by Natalie Nkwor, and sponsored by Amgen Canada Inc.

Funding Sources

This study was funded by Amgen Canada Inc. (Mississauga, Ontario).

Disclosures

Shaun G. Goodman reports receiving research grant support (eg, steering committee or data and safety monitoring committee) and/or speaker/consulting honoraria (eg, advisory boards) from Amgen, AstraZeneca, Bayer, Boehringer Ingelheim, Bristol Myers Squibb, CSL Behring, Daiichi-Sankyo/ American Regent, Eli Lilly, Esperion, Ferring Pharmaceuticals, GlaxoSmithKline, HLS Therapeutics, JAMP Pharma, Janssen/Johnson & Johnson, Merck, Novartis, Novo Nordisk A/C, Pendopharm, Pfizer, Regeneron, Sanofi, Servier, and Valeo Pharma; and salary support/honoraria from the Heart and Stroke Foundation of Ontario/University of Toronto (Polo) Chair, Canadian Heart Research Centre and MD Primer, Canadian VIGOUR Centre, Duke Clinical Research Institute, New York University Clinical Coordinating Centre, and PERFUSE Research Institute. Paul Oh reports receiving speaker/consulting honoraria (eg, advisory boards) from Amgen, AstraZeneca, Bayer, Eli Lilly, GlaxoSmithKline, and HLS Therapeutics. Erin S. Mackinnon, Raina M. Rogoza, Millicent Packalen, Louisa Pericleous, and Ponda Motsepe-Ditshego are employees of Amgen Canada Inc and may own Amgen stock and/or stock options. Ron Goeree has no conflicts of interest to disclose.

References

- World Health Organization. Cardiovascular diseases (CVDs) fact sheet. Available at: http://www.who.int/mediacentre/factsheets/fs317/en/. Accessed February 22, 2021.
- 2. Anderson TJ, Gregoire J, Pearson GJ, et al. 2016 Canadian Cardiovascular Society guidelines for the management of dyslipidemia for the prevention of cardiovascular disease in the adult. Can J Cardiol 2016;32: 1263-82.
- Lloyd-Jones DM, Morris PB, Ballantyne CM, et al. 2016 ACC expert consensus decision pathway on the role of non-statin therapies for LDLcholesterol lowering in the management of atherosclerotic cardiovascular disease risk: a report of the American College of Cardiology Task Force on Clinical Expert Consensus Documents. J Am Coll Cardiol 2016;68: 92-125.
- Canadian Institute for Health Information. Inpatient hospitalization, surgery and newborn statistics. Available at: https://www.cihi.ca/sites/ default/files/document/dad-hmdb-childbirth-quick-stats-2017-2018-enweb.xlsx. Accessed February 22, 2021.

- Charles River Associates. The burden of acute coronary syndrome: the second leading cause of death in Canada. Available at: http://www.crai. com/sites/default/files/publications/Burden-of-acute-coronary-syndromein-Canada.pdf. Accessed February 22, 2021.
- Mach F, Baigent C, Catapano AL, et al. 2019 ESC/EAS guidelines for the management of dyslipidaemias: lipid modification to reduce cardiovascular risk. Eur Heart J 2020;41:111-88.
- Sanchis-Gomar F, Perez-Quilis C, Leischik R, Lucia A. Epidemiology of coronary heart disease and acute coronary syndrome. Ann Transl Med 2016;4:256.
- 8. Jernberg T, Hasvold P, Henriksson M, et al. Cardiovascular risk in post-myocardial infarction patients: Nationwide real world data demonstrate the importance of a long-term perspective. Eur Heart J 2015;36:1163-70.
- Statistics Canada. Table 13-10-0394-01 Leading causes of death, total population, by age group. Available at: https://www150.statcan.gc.ca/t1/ tbl1/en/tv.action?pid=1310039401. Accessed February 22, 2021.
- Benjamin EJ, Virani SS, Callaway CW, et al. Heart disease and stroke statistics-2018 update: a report from the American Heart Association. Circulation 2018;137:e67-492.
- Roth GA, Johnson C, Abajobir A, et al. Global, regional, and national burden of cardiovascular diseases for 10 causes, 1990 to 2015. J Am Coll Cardiol 2017;70:1-25.
- Government of Canada. Canadian Chronic Disease Surveillance System. Available at: https://health-infobase.canada.ca/ccdss/data-tool/. Accessed February 22, 2021.
- Tu JV, Khan AM, Ng K, Chu A. Recent temporal changes in atherosclerotic cardiovascular diseases in Ontario: clinical and health systems impact. Can J Cardiol 2017;33:378-84.
- Blais C, Rochette L, Ouellet S, Huynh T. Complex evolution of epidemiology of vascular diseases, including increased disease burden: from 2000 to 2015. Can J Cardiol 2020;36:740-6.
- Sud M, Han L, Koh M, et al. Low-density lipoprotein cholesterol and adverse cardiovascular events after percutaneous coronary intervention. J Am Coll Cardiol 2020;76:1440-50.
- Charlson ME, Pompei P, Ales KL, MacKenzie CR. A new method of classifying prognostic comorbidity in longitudinal studies: development and validation. J Chronic Dis 1987;40:373-83.
- Vassy JL, Lu B, Ho Y-L, et al. Estimation of atherosclerotic cardiovascular disease risk among patients in the Veterans Affairs health care system. JAMA Netw Open 2020;3:e208236.
- Pearson GJ, Thanassoulis G, Anderson TJ, et al. 2021 Canadian Cardiovascular Society guidelines for the management of dyslipidemia for the prevention of cardiovascular disease in adults. Can J Cardiol 2021;37: 1129-50.
- Li S, Peng Y, Wang X, et al. Cardiovascular events and death after myocardial infarction or ischemic stroke in an older Medicare population. Clin Cardiol 2019;42:391-9.
- Hess CN, Clare RM, Neely ML, et al. Differential occurrence, profile, and impact of first recurrent cardiovascular events after an acute coronary syndrome. Am Heart J 2017;187:194-203.

Supplementary Material

To access the supplementary material accompanying this article, visit *CJC Open* at https://www.cjcopen.ca/ and at https://doi.org/10.1016/j.cjco.2021.10.003.