Original quantitative research

The trends and determinants of seasonal influenza vaccination after cardiovascular events in Canada: a repeated, pan-Canadian, cross-sectional study

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Abstract

Introduction: Annual influenza vaccination is recommended for individuals with a history of cardiovascular disease (CVD) events. We aimed to examine (1) the time trends for influenza vaccination among Canadians with a CVD event history between 2009 and 2018, and (2) the determinants of receiving the vaccination in this population over the

Methods: We used data from the Canadian Community Health Survey (CCHS). The study sample included respondents from 2009 to 2018 who were 30 years of age or more with a CVD event (heart attack or stroke) and who indicated their flu vaccination status. Weighted analysis was used to determine the trend of vaccination rate. We used linear regression analysis to examine the trend and multivariate logistic regression analysis to examine determinants of influenza vaccination, including sociodemographic factors, clinical characteristics, health behaviour and health system variables.

Results: Over the study period, in our sample of 42 400, the influenza vaccination rate was overall stable around 58.9%. Several determinants for vaccination were identified, including older age (adjusted odds ratio [aOR] = 4.28; 95% confidence interval [95% CI]: 4.24–4.32], having a regular health care provider (aOR = 2.39; 95% CI: 2.37–2.41), and being a nonsmoker (aOR = 1.48; 95% CI: 1.47-1.49). Factors associated with decreased likelihood of vaccination included working full time (aOR = 0.72; 95% CI: 0.72-0.72).

Conclusion: Influenza vaccination is still at less than the recommended level in patients with CVD. Future research should consider the impact of interventions to improve vaccination uptake in this population.

Keywords: cardiovascular diseases, influenza vaccines, utilization, secondary prevention, trend, determinants

Introduction

Annual vaccination against influenza is recommended for all individuals with a history of an ischemic cardiovascular disease (CVD) event.1-5 Seasonal influenza infection further elevates the already increased risk of recurrent CVD events and deaths in this particular population. 6-9 Although the exact mechanism is unknown, this increased risk from influenza infection is believed to be the result of viral particles activating inflammatory pathways, which can contribute to dysfunctions in arterial endothelium and lipid metabolism and lead to coronary atherosclerotic events such as myocardial infarction and stroke. 6,10,11 Empirical evidence supports the efficacy of influenza vaccine in the secondary prevention context. 12,13 In

Highlights

- Vaccine uptake in the post-CVD Canadian population from 2009 to 2018 was found to be suboptimal and is a potential area for optimization of health outcomes in these patients.
- Factors associated with increased likelihood of vaccination include older age, having a regular health care provider and being a nonsmoker.

a systematic review of randomized clinical trials, influenza vaccination was associated with a 36.0% reduction in future CVD events, with a relative risk of 0.6 (95% confidence interval [CI]: 0.5–0.9). 12

In Canada, annual influenza vaccines are widely available in pharmacies, physician offices and local public health units.14 Public funding of the vaccine is also provided for those with chronic conditions, including CVD, in all 13 Canadian jurisdictions.¹⁵ Despite this availability, the uptake of influenza vaccine among patients with CVD remains low.15-18 Data from the 2019/20 influenza season revealed the proportion of vaccinated Canadian adults with one or more chronic conditions (including CVD) was 44.0%, well below the 80.0% target set by the National Advisory Committee on Immunizations (NACI).18 However, time trends of vaccination rates in Canadians specifically with a previous CVD event history are unknown.¹⁸

There is also inadequate evidence pertaining to the determinants of vaccine uptake

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in patients with CVD. Studies usually focus on determinants either in the general population or in patients suffering from chronic conditions as a whole, but not CVD specifically.19-23 Older age was found to be significantly associated with a higher rate of vaccine uptake in the general population of the United States, Canada, Italy and Portugal. 19-21,23 In countries such as the United States, where the cost of influenza vaccinations may not be covered by the government, individuals with higher occupational and educational status were more likely to be vaccinated than those with lower incomes.22,24 Aside from cost, other factors such as systemic racism and a reduced degree of prioritization by clinicians and the health care system may hinder patients' access to the influenza vaccine. Although these factors may also exist in Canada, findings from the United States are not directly applicable to the Canadian population, due to differences in demographics and health care coverage.19

It is important to identify the time trend and determinants of influenza vaccination in Canadian patients after CVD events to inform effective strategies, to help determine whether current policies are sufficient and to what extent there is a need to further improve influenza vaccine uptake among these high-risk patients. ^{25,26} The aim of this study was to identify the trend for the period of 2009 to 2018, as well as the determinants of receiving influenza vaccine among Canadian patients with a previous CVD event. We hypothesized that influenza vaccine uptake is increasing in Canada.

Methods

Data source

We used data from the Public Use Microdata Files (PUMF) of the Canadian Community Health Survey (CCHS)27 to conduct this study. We accessed the data through the Ontario Data Documentation, Extraction Service and Infrastructure ("odesi") tool.28 The CCHS is a voluntary, cross-sectional survey of noninstitutional Canadian residents aged 12 years and older, used to obtain health-related information representative of different health regions across Canada.27 The data are collected year-round. Subject areas of interest include various health conditions, utilization of health care services, lifestyle factors and mental health.27

The CCHS utilizes a complex, two-stage, stratified cluster design to sample those 18 years of age and over in the Labour Force Survey while using a simple, random sample to query children aged 12 to 17 years.²⁷ A letter from Statistics Canada inviting participation in the survey is mailed to respondents; those who agree are then directed to an online questionnaire.27 Together, those excluded from the sample make up less than 3% of the Canadian population.27 The PUMF from CCHS compiles responses from approximately 130 000 individuals over a twoyear period, published as a microdata file biennially.^{27,29}

For this study, we included CCHS data from the 2009–2010, 2011–2012, 2013–2014, 2015-2016 and 2017-2018 cycles. The variables concerning influenza vaccination and all exposure variables used in this study were core content in the CCHS documentation, signifying that the variables were asked in all Canadian provinces and territories.27 We applied weights provided in the Statistics Canada datasets to all data analyzed and presented in this study.27 As per Statistics Canada, the survey weights are determined by a combination of modelling probabilities of response at the household and person levels, and correlates to the number of persons in the Canadian population represented by each respondent.27

Study population

We included respondents from the 2009 to 2018 CCHS who indicated that they were 30 years of age or older, had experienced a CVD event and who answered questions pertaining to influenza vaccinations. CVD event history was assessed using the survey questions "Do you have heart disease?" and "Do you suffer from the effects of a stroke?" Respondents who answered "yes" to either one or both questions were included in the study. Although it was not established through hospitalization records that all respondents indicating presence of heart disease or stroke had in fact experienced an ischemic CVD event, it was considered a reasonable indicator to be used for this study. Individuals below 30 years of age were excluded from this study due to the extremely low prevalence of heart disease in this age group and differences in etiology compared to older adults (i.e. higher proportion of non-atherosclerotic causes of CVD).30

Vaccination status

Respondents were considered vaccinated for the influenza season if they have indicated "yes" to the question "Have you ever had a flu shot?" and have also indicated " < 1 year ago" to the question "When did you last receive the vaccine?" As the influenza vaccine is recommended annually, respondents who had had the flu shot but indicated "1-2 years ago" or "2 or more years ago" as the last time they received the vaccine, as well as those who had indicated "no" to the question "Have you ever had a flu shot?", were considered unvaccinated in this study. The remaining respondents who indicated "don't know" or "unsure" to any of the above questions, or refused to answer, were all considered unvaccinated.

Measurement and confounding variables

Various independent variables were chosen to identify potential determinants of the outcome of interest (i.e. being vaccinated). We included sociodemographic factors related to age, sex, marital status, income, education level, immigrant status and employment status, based on previous findings for their correlation to vaccination among the general population. 19-21,23 In addition, we included the cycle year and chronic diseases variables. We also included variables pertaining to smoking status and body mass index (BMI)-calculated by Statistics Canada—to study the impact of various health-related factors, as well as a variable assessing selfperceived health, with responses ranging from "poor" to "excellent."31 Further, the variables of having a regular health care provider and requiring help with personal care were included to evaluate health care and external aid utilization. Whether or not respondents resided in provinces or territories allowing pharmacists to provide immunizations was also assessed, due to recent evidence showing that Canadian jurisdictions that had implemented this policy saw increased influenza vaccination rates.32 Details about included variables are available in Appendix 1.

Data analysis

The weighted rate of respondents in the overall CVD population who received influenza vaccination (i.e. the proportion of respondents vaccinated) across the study period from 2009 to 2018 was first plotted, along with the confidence interval.

The same procedure was then repeated to plot the vaccination rate of respondents in the overall CVD population over time stratified by province. These data plots were then analyzed utilizing the linear regression analysis test in Microsoft Excel version 16.43 (Microsoft Corp., Redmond, WA, US) to determine the significance of any trends in receiving vaccination over the study years.

Next, descriptive statistics were calculated for patients who were vaccinated versus those who were unvaccinated. The association between each independent variable and receipt of influenza vaccine was examined using the chi-square (χ^2) test of independence. Similar to previous research, a weighted multivariable logistic regression model was then fitted using a stepwise forward-selection model.33 The independent variables in the final model were included based on significance (p < 0.05) from the Wald statistic and goodness of fit using the Akaike information criterion. Selected variables-cycle year, age and sex-were included in the model regardless of statistical significance. Patients with missing data were dealt with first by listwise deletion approach (where only respondents with complete data in all variables were kept in the analysis).34 In a sensitivity analysis, we used the educatedguessing approach, in which variables with missing values are replaced with "no" in binary variables or by the lowest level (in ordinal variables).

Using weighted results, a second sensitivity analysis was performed in order to evaluate the robustness of the study definition for vaccination status. In the main model, respondents were considered vaccinated only if they had indicated having received the flu shot less than one year ago. This definition, however, excludes those who received the flu shot exactly one year ago or just over one year ago but who would still go on to be vaccinated for the upcoming influenza season. Therefore, in this sensitivity analysis, we considered respondents vaccinated as long as they had received the influenza vaccine less than two years before the survey date. This is because influenza vaccination is recommended annually during the influenza season to protect against new strains of the influenza virus. However, because the CCHS collects data annually, the questions pertaining to vaccination status may be referring to either year of a two-year cycle.

Several subgroup analyses were also performed to identify any differences in determinants for vaccination based on age group and type of CVD event (e.g. stroke only). The same independent variables and statistical procedure applied to the main model were used to perform the subgroup analyses.

SAS University Edition (SAS Studio version 3.8, SAS version 9.4; SAS Institute Inc., Cary, NC, US) was utilized to analyze the survey data. Due to the fact that the data are publicly available through Statistics Canada,²⁷ there was no need for research ethics board approval to conduct this study. All numbers presented are rounded to the closest 100, as per Statistics Canada rounding guidelines.³⁵

Results

Descriptive statistics

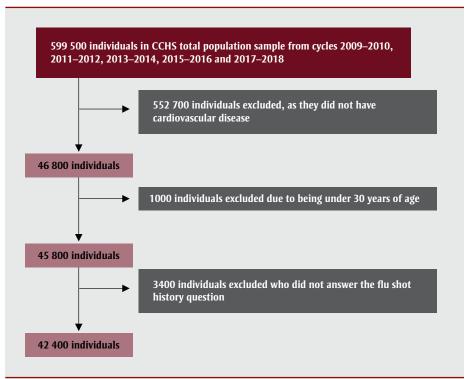
The study sample included a total of 42 400 respondents, representing a weighted population of 7 148 500 Canadians, from the CCHS cycles 2009–2010 to 2017–2018, residing in all 10 Canadian provinces and 3 territories. Figure 1 illustrates the process taken to determine the final study sample. Most respondents (81.0%) had

heart disease only, 13.0% had a history of a stroke and the remaining 6.0% had both. In the total weighted sample of 42 400 respondents, 58.9% had received influenza vaccination. More than half of the sample population (58.0%) were aged 65 and older, and 56.0% were males. Table 1 describes the baseline characteristics of the study weighted sample. Vaccinated individuals were generally older in age, were married and had other comorbidities. A low level of missingness was observed (Appendix 2) in all independent variables (< 3%) and most variables had less than 1% missingness.

Trends in vaccination

Figure 2 illustrates the weighted proportion of respondents with CVD events vaccinated against influenza from 2009 to 2018. Over the ten-year study period, there was a general downward trend in the proportion of vaccinated individuals with a history of a CVD event, from 59.3% (95% CI: 59.2–59.4) in the 2009–2010 cycle to 55.5% (95% CI: 55.4–55.6) in the 2017–2018 cycle. Vaccination rates peaked in the 2013–2014 cycle, with 61.5% (95% CI: 61.4–61.6) of respondents indicating vaccination against influenza. However, the trend line was not significant (*p*

FIGURE 1 Selection process of study respondents



Data source: Canadian Community Health Study (CCHS), 2009–2010 to 2017–2018.

TABLE 1 Characteristics of the weighted study sample

Characteristics of t				
	— P values from			
Unvaccinated n = 2 939 900 (column %)	Vaccinated n = 4 208 600 (column %)	Total N = 7 148 400 ^b (column %)	chi-square test of independence	
			< 0.0001	
267 300 (9%)	107 200 (3%)	374 400 (5%)		
1 445 900 (49%)	1 163 200 (28%)	2 609 100 (36%)		
1 226 700 (42%)	2 938 200 (70%)	4 164 900 (58%)		
			< 0.0001	
2 327 200 (79%)	3 475 800 (83%)	5 803 000 (81%)		
			< 0.0001	
426 600 (15%)	465 900 (11%)	892 500 (13%)		
			0.9829	
186 200 (6%)	266 800 (6%)	453 000 (6%)		
			0.0002	
1 239 700 (42%)	1 922 000 (46%)	3 161 700 (44%)		
			0.0007	
576 000 (20%)	837 700 (20%)	1 413 600 (20%)		
587 000 (20%)	877 900 (21%)	1 464 900 (20%)		
567 900 (19%)	906 800 (22%)	1 474 700 (21%)		
578 100 (20%)	799 600 (19%)	1 377 700 (19%)		
631 000 (21%)	786 600 (19%)	1 417 600 (20%)		
			< 0.0001	
82 700 (3%)	172 900 (4%)	255 600 (4%)		
			< 0.0001	
562 000 (19%)	441 500 (10%)	1 003 500 (14%)		
109 800 (4%)	95 000 (2%)	204 800 (3%)		
1 484 400 (50%)	2 416 000 (57%)	3 900 400 (55%)		
454 400 (15%)	756 000 (18%)	1 210 300 (17%)		
			0.3403	
2 552 300 (87%)	3 633 100 (86%)	6 185 300 (87%)		
377 000 (13%)	565 200 (13%)	942 200 (13%)		
			< 0.0001	
574 700 (20%)	1 129 000 (27%)	1 703 700 (24%)		
			< 0.0001	
287 100 (10%)	523 700 (12%)	810 700 (11%)		
			< 0.0001	
283 100 (10%)	558 000 (13%)	841 100 (12%)		
			< 0.0001	
2 663 300 (91%)	4 068 100 (97%)	6 731 500 (94%)		
			0.4614	
	Unvaccinated n = 2 939 900 (column %) 267 300 (9%) 1 445 900 (49%) 1 226 700 (42%) 2 327 200 (79%) 426 600 (15%) 1 239 700 (42%) 576 000 (20%) 587 000 (20%) 567 900 (19%) 578 100 (20%) 631 000 (21%) 82 700 (3%) 562 000 (19%) 109 800 (4%) 1 484 400 (50%) 454 400 (15%) 2 552 300 (87%) 377 000 (13%) 574 700 (20%) 287 100 (10%)	Unvaccinated n = 2 939 900 (column %) Vaccinated n = 4 208 600 (column %) 267 300 (9%) 107 200 (3%) 1 445 900 (49%) 1 163 200 (28%) 1 226 700 (42%) 2 938 200 (70%) 2 327 200 (79%) 3 475 800 (83%) 426 600 (15%) 465 900 (11%) 186 200 (6%) 266 800 (6%) 1 239 700 (42%) 1 922 000 (46%) 576 000 (20%) 837 700 (20%) 587 000 (20%) 877 900 (21%) 567 900 (19%) 906 800 (22%) 578 100 (20%) 799 600 (19%) 631 000 (21%) 786 600 (19%) 82 700 (3%) 172 900 (4%) 562 000 (19%) 441 500 (10%) 109 800 (4%) 95 000 (2%) 1 484 400 (50%) 2 416 000 (57%) 454 400 (15%) 756 000 (18%) 2 552 300 (87%) 3 633 100 (86%) 377 000 (13%) 565 200 (13%) 574 700 (20%) 1 129 000 (27%) 283 100 (10%) 558 000 (13%)	Unvaccinated n = 2 939 900 (column %) (colum	

TABLE 1 (continued)
Characteristics of the weighted study sample

		Davidson from			
Respondent characteristics	Unvaccinated n = 2 939 900 (column %)	Vaccinated n = 4 208 600 (column %)	Total N = 7 148 400 ^b (column %)	 P values from chi-square test of independence 	
Marital status				0.0048	
Single/widowed/divorced	1 073 000 (36%)	1 437 700 (34%)	2 510 700 (35%)		
Married	1 858 600 (63%)	2 764 400 (66%)	4 623 100 (65%)		
Highest educational attainment				0.0367	
Secondary and lower	1 278 400 (43%)	1 916 600 (46%)	3 195 000 (45%)		
Postsecondary and higher	1 568 900 (53%)	2 169 800 (52%)	3 738 700 (52%)		
Pharmacist immunization in province of residence				< 0.0001	
Yes	1 552 500 (53%)	2 497 400 (59%)	4 049 900 (57%)		
Province of residence				< 0.0001	
British Columbia	334 400 (11%)	531 700 (13%)	866 000 (12%)		
Alberta	247 400 (8%)	346 300 (8%)	593 800 (8%)		
Saskatchewan	88 500 (3%)	124 700 (3%)	213 200 (3%)		
Manitoba	96 600 (3%)	125 500 (3%)	222 300 (3%)		
Ontario	1 024 800 (35%)	1 761 000 (42%)	2 785 800 (39%)		
Quebec	916 800 (31%)	901 800 (21%)	1 818 700 (25%)		
Atlantic provinces	223 700 (8%)	409 100 (10%)	632 800 (9%)		
Territories	7 600 (0%)	8 300 (0%)	15 900 (0%)		
ВМІ				0.9008	
< 25	996 400 (34%)	1 429 800 (34%)	2 426 200 (34%)		
≥ 25	1 877 600 (64%)	2 680 300 (64%)	4 557 900 (64%)		
Immigrant status				0.8963	
Yes	639 700 (22%)	925 200 (22%)	1 564 900 (22%)		
Full-time worker				< 0.0001	
Yes	916 000 (31%)	651 000 (15%)	1 567 100 (22%)		

Data source: Canadian Community Health Study, 2009–2010 to 2017–2018.

Abbreviations: BMI, body mass index; COPD, chronic obstructive pulmonary disease.

value = 0.12). Figure 3 illustrates the breakdown of vaccination trends within the Canadian provinces. Ontario, Quebec and British Columbia saw a general decrease in vaccination rates, while Alberta experienced an overall increase over the study period. Quebec consistently remained the province with the lowest percentage of respondents vaccinated.

Determinants of receiving influenza vaccination

The variables that were retained in the multivariate logistic regression analysis, in

addition to age, cycle year and sex, were smoking status, presence of comorbidities (diabetes, asthma and chronic obstructive pulmonary disease [COPD]), marital status, working status, highest educational attainment, requiring help for personal care and having a regular health care provider (Appendix 3). The adjusted odds ratios (aORs) of the variables controlled for in the final main model are listed in Table 2. Age of 65 years or older was associated with the greatest odds of receiving influenza vaccination, with an aOR of 4.28 (95% CI: 4.24–4.32) and having a

regular health care provider was also associated with increased odds (aOR = 2.39; 95% CI: 2.37-2.41).

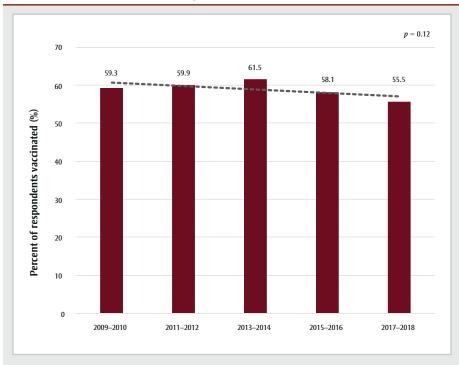
Subgroup analysis

Results of the subgroup analyses stratified by age revealed some differences in the aORs for the following variables: COPD, requiring help with personal care and working status. Respondents in the youngest age group (aged 30–44) were approximately four times more likely to receive the influenza vaccination if they had COPD than those without COPD (aOR = 4.6;

^a Respondents were considered vaccinated if they indicated having received the flu shot less than one year ago.

^b Number has been rounded.

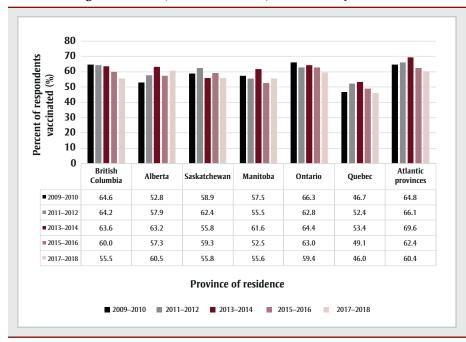
FIGURE 2 Weighted percentage of Canadians with a CVD event history vaccinated against influenza, from 2009 to 2018



Data source: Canadian Community Health Survey.

Abbreviation: CVD, cardiovascular disease.

FIGURE 3
Weighted percentage of Canadians with a CVD event history vaccinated against influenza, from 2009 to 2018, across various provinces^a



Data source: Canadian Community Health Survey.

95% CI: 4.4-4.8), while respondents aged 45 and above with COPD had similar odds of vaccination as in the main model (aOR = 1.2; 95% CI: 1.1-1.2). Respondents in the youngest age group requiring help with personal care had an aOR of 2.7 (95% CI: 2.6-2.9), while requiring help with personal care was not associated with vaccination in respondents aged 45 to 64. The odds of vaccination were also increased in respondents working full time between 30 and 44 years of age (aOR = 2.5; 95% CI: 2.5-2.6) while respondents working full time older than 45 years of age had a decreased likelihood of vaccination (aOR = 0.8; 95% CI: 0.8-0.8). The remaining variables remained consistent with the results of the main model across all age groups (Table 3). In a second subgroup analysis on respondents suffering from the effects of a stroke only, findings were similar to the results of the main model (Table 4).

Sensitivity analysis

In the sensitivity analysis, in which respondents were considered vaccinated as long as they indicated vaccination less than 2 years ago, there were no extensive changes to the odds ratios compared to the main model (Appendix 4). Similarly, the educated-guessing approach for missing data yielded similar estimates (Appendix 5).

Discussion

We examined the trends and determinants of receiving influenza vaccination in individuals with a history of a CVD event in a representative sample from the Canadian population from 2009 to 2018. Over the study period, the percentage of respondents vaccinated each year remained generally stable (ranging from 55.5%-61.5%) and experienced no significant change (p = 0.12). Despite various attempts to improve vaccine uptake through national influenza vaccination campaigns and increased accessibility of vaccines through local pharmacies, vaccination rates remained below NACI's 80% target for Canadians with chronic conditions. 18,32 This is a concerning finding, as annual influenza vaccinations are an easily accessible and cost-effective measure to reduce morbidity and mortality from CVD events.25,32 The data demonstrate that, as in other jurisdictions, not enough post-CVD patients are utilizing this cardioprotective strategy in Canada.23 Therefore, while national influenza

^a Territories are not shown due to the very small number of respondents.

vaccination campaigns distribute information for the general population, additional strategies to distribute information tailored to high-risk populations may be required.³⁶

Within the study period, vaccination rates peaked in 2013/14. A possible reason may be attributed to the implementation of funding and policy allowing pharmacists to administer influenza vaccines in Manitoba and Atlantic provinces that year. 19,32 This is also supported by the finding that Quebec had the lowest rates of vaccination throughout the study period, potentially explained by the absence of universal funding and pharmacist immunization policy for influenza vaccinations in this province.32 However, there was no significant improvement in vaccine uptake from 2009, and in none of the study years was the target of 80% reached.18

Consistent with previous studies, increasing age was associated with higher influenza vaccination rates. ^{19-21,23} Likewise, the presence of other comorbidities was another strong predictor for vaccination, and comorbidities also become more prevalent with age. ¹⁸ Older individuals and

TABLE 2
Adjusted odds ratios of being vaccinated^a from multivariate logistic regression analysis (main model)^b

Effect	Adjusted odds ratio (aOR) estimation	95% Wald confidence limits	
2011–2012 vs. 2009–2010	1.05	1.04	1.05
2013–2014 vs. 2009–2010	1.10	1.09	1.10
2015–2016 vs. 2009–2010	0.96	0.95	0.96
2017–2018 vs. 2009–2010	0.80	0.79	0.80
Sex: female vs. male	1.06	1.06	1.07
Age: 45–64 vs. 30–44	1.73	1.72	1.75
Age: 65 and older vs. 30–44	4.28	4.24	4.32
Require help with personal care: yes vs. no	1.11	1.10	1.11
Smoking: no vs. yes	1.48	1.47	1.49
Diabetes: yes vs. no	1.37	1.37	1.38
Asthma: yes vs. no	1.36	1.35	1.37
COPD: yes vs. no	1.32	1.31	1.32
Marital status: married vs. single/widowed/divorced	1.25	1.25	1.26
Work full time: yes vs. no	0.72	0.72	0.72
Has a regular HCP: yes vs. no	2.39	2.37	2.41
Educational attainment: postsecondary and higher vs. secondary and lower	1.10	1.09	1.10

Abbreviations: COPD, chronic obstructive pulmonary disease; HCP, health care provider.

TABLE 3
Adjusted odds ratios of subgroup model stratified by age

	Ages 30–44 y		Ages 45–64 y			Ages 65 y and above			
Effect	Adjusted odds ratio estimation		Wald nce limits	Adjusted odds ratio estimation	95% confiden	Wald ce limits	Adjusted odds ratio estimation		Wald nce limits
2011–2012 vs. 2009–2010	0.38	0.37	0.40	1.10	1.09	1.11	1.36	1.35	1.38
2013–2014 vs. 2009–2010	0.68	0.66	0.70	0.98	0.97	0.99	1.24	1.23	1.25
2015–2016 vs. 2009–2010	0.94	0.92	0.97	0.92	0.91	0.93	1.00	0.99	1.00
2017–2018 vs. 2009–2010	0.44	0.43	0.46	0.70	0.69	0.70	0.91	0.91	0.92
Sex: female vs. male	1.48	1.46	1.51	1.11	1.10	1.12	1.02	1.01	1.02
Require help with personal care: yes vs. no	2.34	2.22	2.45	1.02	1.00	1.04	1.13	1.11	1.14
Smoking: no vs. yes	1.35	1.32	1.39	1.25	1.24	1.25	1.78	1.76	1.79
Diabetes: yes vs. no	2.02	1.97	2.08	1.44	1.43	1.45	1.27	1.26	1.28
Asthma: yes vs. no	1.19	1.16	1.22	1.58	1.57	1.60	1.16	1.15	1.17
COPD: yes vs. no	3.16	3.04	3.28	1.18	1.17	1.20	1.36	1.35	1.37
Has a regular HCP: yes vs. no	3.00	2.91	3.11	2.55	2.52	2.58	2.36	2.33	2.39
Marital status: married vs. single/widowed/divorced	0.99	0.98	1.02	1.16	1.16	1.17	1.30	1.30	1.31
Work full time: yes vs. no	1.53	1.50	1.56	0.79	0.79	0.80	0.47	0.46	0.47
Educational attainment: postsecondary and higher vs. secondary and lower	1.12	1.10	1.15	1.15	1.15	1.16	1.09	1.09	1.10

Abbreviations: COPD, chronic obstructive pulmonary disease; HCP, health care provider; y, years.

^a Respondents were considered vaccinated if they indicated having received the flu shot less than one year ago.

^b All variables in the table were included in the multivariate model.

TABLE 4
Adjusted odds ratio of subgroup sample of respondents who had stroke only

Effect	Adjusted odds ratio estimation	95% Wald con	fidence limits
Ages (yr): 45–64 vs. 30–44	1.60	1.56	1.63
Ages (yr): 65 and older vs. 30-44	3.68	3.60	3.77
2011–2012 vs. 2009–2010	0.67	0.66	0.69
2013–2014 vs. 2009–2010	0.79	0.78	0.80
2015–2016 vs. 2009–2010	1.06	1.05	1.08
2017–2018 vs. 2009–2010	0.68	0.67	0.69
Sex: female vs. male	1.20	1.19	1.21
Require help with personal care: yes vs. no	1.26	1.24	1.29
Smoking: no vs. yes	1.44	1.42	1.46
Diabetes: yes vs. no	1.55	1.53	1.57
Asthma: yes vs. no	1.64	1.61	1.66
COPD: yes vs. no	0.89	0.88	0.91
Has a regular HCP: yes vs. no	2.25	2.12	2.30
Marital status: married vs. single/widowed	1.28	1.27	1.29
Work full time: yes vs. no	0.52	0.51	0.53
Educational attainment: postsecondary and higher vs. secondary and lower	1.32	1.30	1.33

Abbreviations: COPD, chronic obstructive pulmonary disease; HCP, health care provider.

those with greater comorbidities are more readily perceived by health care providers to be at higher risk for complications from influenza, leading to greater frequency of recommendations and higher vaccination rates.³⁷ Increasing age may also be associated with increased self-perceived risk to complications of influenza infection, thereby influencing self-motivated vaccine uptake.³⁸

Individuals with a regular health care provider were more than twice as likely as those without one to receive vaccination. This supports findings that health care utilization is an important determinant for vaccination.21,39 Yet, while 94% of individuals in our study reported having a regular health care provider, almost 40% were not vaccinated against influenza. This suggests a potential gap in communication between health care providers and patients regarding the cardioprotective benefits of the influenza vaccine.25 Considering the significant impact of health care provider recommendations on vaccine uptake as demonstrated by numerous studies, a greater focus on patient education on vaccine benefits during all points of contact with the health care system (e.g. hospitalizations, followup visits) is warranted.40-42

We also found that nonsmokers across all age groups were more likely to receive influenza vaccination than smokers (OR = 1.5; 95% CI: 1.4-1.5). While there are some discrepancies in the literature, 20,21,37 57.0% of our vaccinated study population were noted to be former smokers. It is possible that former smokers who made the decision to guit smoking after a CVD event may be more inclined to take part in other preventative measures such as influenza vaccinations.43 However, it is the current smokers who are at a higher risk for CVD events and have a higher incidence of CVD mortality than former smokers, and would therefore derive greater benefit from vaccination.43

We found that Canadians with a CVD event aged 65 and older with higher educational attainment were more likely to be vaccinated. This supports the findings of several Canadian studies showing that higher educational status is a determinant for vaccination in the elderly.^{37,44} On the other hand, higher educational attainment was linked to a decrease in odds of vaccination for those under 65 years of age, which is in line with the findings from previous studies in other countries.^{20,45} This can be potentially explained by the

association between higher education status and greater likelihood of working, rendering these individuals busier and potentially less able to conveniently access vaccination than those who are not working.39 Lastly, our results suggest that future vaccination campaigns could benefit from directing efforts to the working population. Working full-time was associated with a decreased likelihood of vaccination among middle-aged respondents aged 45 to 64. Full-time workers may potentially be busier than their unemployed counterparts, contributing to greater difficulties with booking health care appointments or taking part in vaccination programs.39

Strengths and limitations

Our study utilized representative data from the Canadian population collected over ten years. This enabled us to examine the trend determinants for vaccination and vaccine receipt in the past decade. However, some limitations should be noted.

First, CCHS relies on self-reporting, in which the responses may be subject to recall bias. Nevertheless, the CCHS questions pertaining to heart disease and stroke were validated and found to be robust. Lix et al. reported that these guestions have very high specificity (> 96%) and negative predictive value (> 98%),46 which would support the existence of heart disease in CCHS respondents who reported that they have heart disease. Regarding vaccination status, some respondents may have stated their last vaccination to be one to two years ago, when in actuality it was less than one year ago. This would have led them to be categorized as unvaccinated in the study, leading to an underestimation of the actual vaccination rate. It should be noted, however, that in the sensitivity analysis, expanding the window of vaccination to two years did not have an impact on the results.

Second, there were no specific questions asked in CCHS concerning history of CVD events. The question "Do you have heart disease?" encompasses many heart diseases, such as atrial fibrillation or heart failure, while the aim of this study was to look at only those with a history of an atherosclerotic cardiacvascular or cerebrovascular event.

Nevertheless, our results can be generalized to the Canadian public, as our sample was large, and the data collected over an extended period of time. In addition,

sample weights provided by Statistics Canada provide a robust estimation of vaccination level among patients with heart disease.

Conclusion

In spite of the morbidity and mortality benefits of the annual influenza vaccination in patients with a history of a CVD event, influenza vaccination rates among Canadians are still suboptimal, and were found to be overall stable over the tenyear study period from 2009 to 2018.18 Major determinants associated with vaccine uptake include increasing age, having a regular health care provider, having concurrent comorbidities, requiring help with personal care and being a nonsmoker. Future influenza vaccination campaigns should include messages directed at post-CVD patients, as well as groups associated with lower odds of vaccination, such as those employed full-time in the workforce and individuals under 65 years of age. The results of this study also re-emphasize the important role clinicians play in patient education and the recommendation of influenza vaccinations for improved vaccine uptake and health outcomes in the Canadian CVD population.1

Conflicts of interest

None.

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Authors' contributions and statement

HC: data acquisition, data analysis, writing—original draft, writing—review and editing. SH: conceptualization, methodology, writing—review and editing. WA: conceptualization, methodology, data acquisition, data analysis, writing—review and editing.

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References

 Davis MM, Taubert K, Benin AL, et al. Influenza vaccination as secondary prevention for cardiovascular disease. J Am Coll Cardiol. 2006;48(7):1498-502. https://doi.org/10.1016/j.jacc.2006 .09.004

- 2. Fitchett DH, Theroux P, Brophy JM, et al. Assessment and management of acute coronary syndromes (ACS): a Canadian perspective on current guideline-recommended treatment part 1: non-ST–segment elevation ACS. Can J Cardiol. 2011;27(Suppl A):S387-S401. https://doi.org/10.1016/j.cjca.2011.08.110
- 8. Grohskopf LA, Alyanak E, Broder KR, et al. Prevention and control of seasonal influenza with vaccines: recommendations of the Advisory Committee on Immunization Practices United States, 2020-21 influenza season. MMWR Recomm Rep. 2020;69(8):1-24. https://doi.org/10.15585/mmwr.rr6908a1
- 4. Young K, Gemmill I, Harrison R. Summary of the NACI seasonal influenza vaccine statement for 2020-2021. Can Commun Dis Rep. 2020;46(5):132-7. https://doi.org/10.14745/ccdr.v46i05a06
- . Piepoli MF, Hoes AW, Agewall S, et al. 2016 European guidelines on cardio-vascular disease prevention in clinical practice: the Sixth Joint Task Force of the European Society of Cardiology and Other Societies on Cardiovascular Disease Prevention in Clinical Practice (constituted by representatives of 10 societies and by invited experts). Developed with the special contribution of the European Association for Cardiovascular Prevention & Rehabilitation (EACPR). Eur Heart J. 2016; 37(29):2315-81. https://doi.org/10.1093/eurheartj/ehw106
- Nichol KL, Nordin J, Mullooly J, Lask R, Fillbrandt K, Iwane M. Influenza vaccination and reduction in hospitalizations for cardiac disease and stroke among the elderly. N Engl J Med. 2003;348(14):1322-32. https://doi.org/10.1056/NEJMoa025028
- 7. Hebsur S, Vakil E, Oetgen WJ, Kumar PN, Lazarous DF. Influenza and coronary artery disease: exploring a clinical association with myocardial infarction and analyzing the utility of vaccination in prevention of myocardial infarction. Rev Cardiovasc Med. 2014; 15(2):168-75. https://doi.org/10.3909/ricm0692
- Kwong JC, Schwartz KL, Campitelli MA, et al. Acute myocardial infarction

- after laboratory-confirmed influenza infection. N Engl J Med. 2018: 378(4):345-53. https://doi.org/10.1056/NEJMoa1702090
- Govender RD, Al-Shamsi S, Soteriades ES, Regmi D. Incidence and risk factors for recurrent cardiovascular disease in middle-eastern adults: a retrospective study. BMC Cardiovasc Disord. 2019;19(1):253. https://doi.org/10.1186/s12872-019-1231-z
- 10. van der Wal AC, Becker AE. Atherosclerotic plaque rupture—pathologic basis of plaque stability and instability. Cardiovasc Res. 1999;41:334-44. https://doi.org/10.1016/S0008-6363 (98)00276-4
- 11. Cowan LT, Lutsey PL, Pankow JS, Matsushita K, Ishigami J, Lakshminarayan K. Inpatient and outpatient infection as a trigger of cardiovascular disease: the ARIC study. J Am Heart Assoc. 2018;7(22):e009683. https://doi.org/10.1161/JAHA.118.009683
- 12. Udell JA, Zawi R, Bhatt DL, et al. Association between influenza vaccination and cardiovascular outcomes in high-risk patients: a meta-analysis. JAMA. 2013;310(16):1711-20. https://doi.org/10.1001/jama.2013.279206
- 13. Phrommintikul A, Kuanprasert S, Wongcharoen W, Kanjanavanit R, Chaiwarith R, Sukonthasarn A. Influenza vaccination reduces cardiovascular events in patients with acute coronary syndrome. Eur Heart J. 2011; 32(14):1730-5. https://doi.org/10.1093/eurheartj/ehr004
- 14. Government of Canada. Flu clinics across Canada [Internet]. Ottawa (ON): Government of Canada; 2020 [cited 2021 Oct]. Available from: https://www.canada.ca/en/public-health/services/diseases/flu-influenza/flu-clinics-across-canada.html
- 15. Government of Canada. Public funding for influenza vaccination by province/territory (as of August 2020) [Internet]. Ottawa (ON): Government of Canada; 2020 [cited 2021 Oct]. Available from: https://www.canada.ca/en/public-health/services/provincial-territorial-immunization-information/public-funding-influenza-vaccination-province-territory.html

- 16. Centers for Disease Control and Prevention (CDC). Flu vaccination coverage, United States, 2019–20 influenza season [Internet]. Atlanta (GA): CDC; 2020 [cited 2021 Oct]. Available from: https://www.cdc.gov/flu/fluvaxview/coverage-1920estimates.htm
- 17. Statistics Canada. Flu vaccination rates in Canada [Internet]. Ottawa (ON): Statistics Canada; 2015 [cited 2021 Oct]. Available from: https://www150.statcan.gc.ca/n1/pub/82-624-x/2015001/article/14218-eng.htm
- 18. Government of Canada. Seasonal influenza vaccination coverage in Canada, 2019-2020 [Internet]. Ottawa (ON): Government of Canada; 2020 [modified 2022 Jul 11; cited 2022 Oct]. Available from: https://www.canada.ca/en/public-health/services/immunization-vaccines/vaccination-coverage/2019-2020-seasonal-influenza-flu-vaccine-coverage.html
- 19. Buchan SA, Kwong JC. Trends in influenza vaccine coverage and vaccine hesitancy in Canada, 2006/07 to 2013/14: results from cross-sectional survey data. CMAJ Open. 2016;4(3): E455–E462. https://doi.org/10.9778/cmajo.20160050
- Chiatti C, Barbadoro P, Lamura G, et al. Influenza vaccine uptake among community-dwelling Italian elderly: results from a large cross-sectional study. BMC Public Health. 2011;11: 207. https://doi.org/10.1186/1471-2458-11-207
- Machado A, Kislaya I, Santos AJ, et al. Factors associated to repeated influenza vaccination in the Portuguese adults with chronic conditions. Vaccine. 2018;36(35):5265-72. https://doi.org/10.1016/j.vaccine.2018.07.041
- 22. Williams WW, Lu P-J, O'Halloran A, et al. Surveillance of vaccination coverage among adult populations United States, 2014. MMWR Surveill Summ. 2016;65(1):1–36. http://doi.org/10.15585/mmwr.ss6501a1
- 23. Madjid M, Alfred A, Sahai A, Conyers JL, Casscells SW. Factors contributing to suboptimal vaccination against influenza: results of a nationwide telephone survey of persons with cardiovascular disease. Texas Heart Inst J. 2009;36(6):546-52.

- 24. Grandhi GR, Mszar R, Vahidy F, et al. Sociodemographic disparities in influenza vaccination among adults with atherosclerotic cardiovascular disease in the United States. JAMA Cardiol. 2021;6(1):87-91. https://doi.org/10.1001/jamacardio.2020.3978
- 25. Gurfinkel EP, Leon de la Fuente R, Mendiz O, Mautner B. Flu vaccination in acute coronary syndromes and planned percutaneous coronary interventions (FLUVACS) Study. Eur Heart J. 2004;25(1):25-31. https://doi.org/10.1016/j.ehj.2003.10.018
- 26. Ciszewski A, Bilinska ZT, Brydak LB, et al. Influenza vaccination in secondary prevention from coronary ischaemic events in coronary artery disease: FLUCAD study. Eur Heart J. 2008; 29(11):1350-8. https://doi.org/10.1093/eurheartj/ehm581
- 27. Statistics Canada. Canadian Community
 Health Survey Annual Component
 (CCHS) [Internet]. Ottawa (ON):
 Statistics Canada; [updated 2022; cited
 2021 Oct]. Available from: https://www23.statcan.gc.ca/imdb/p2SV.pl
 ?Function = getSurvey&SDDS = 3226
- 28. Scholars Portal. About odesi [Internet]. Toronto (ON): Ontario Council of University Libraries; date unknown [cited 2021 Oct]. Available from: https://learn.scholarsportal.info/all-guides/odesi/#: ~:text = is%20a%20 web%2D,basic%20tabulation%20 and%20analysis%20online
- 29. Statistics Canada. Canadian Community Health Survey: Public Use Microdata File [Internet]. Ottawa (ON): Statistics Canada [Catalogue No.: 82M0013X]; 2022 [cited 2022 Oct]. Available from: https://www150.statcan.gc.ca/n1/en/catalogue/82M0013X
- 30. Rubin JB, Borden WB. Coronary heart disease in young adults. Curr Atheroscler Rep. 2012;14(2):140-9. https://doi.org/10.1007/s11883-012-0226-3
- 31. Schmid P, Rauber D, Betsch C, Lidolt G, Denker M-L. Barriers of influenza vaccination intention and behavior a systematic review of influenza vaccine hesitancy, 2005-2016. PLoS ONE. 2017;12(1): e0170550. https://doi.org/10.1371/journal.pone.0170550

- 32. Buchan SA, Rosella LC, Finkelstein M, et al. Impact of pharmacist administration of influenza vaccines on uptake in Canada. CMAJ. 2017;189(4): E146–E152. https://doi.org/10.1503/cmai.151027
- 33. Amoud R, Grindrod K, Cooke M, Alsabbagh Mhd W. The impact of prescription medication cost coverage on oral medication use for hypertension and type 2 diabetes mellitus. Healthc Policy. 2020;16(2):82-100. https://doi.org/10.12927/hcpol.2020.26351
- 34. Allison P. Missing Data. In: Millsap RE, Maydeu-Olivares A, editors. The SAGE handbook of quantitative methods in psychology. Thousand Oaks (CA): SAGE Publications Ltd; 2009:72-89. https://doi.org/10.4135/9780857020994.n4
- 35. Statistics Canada. 7.0 Guidelines for analysis and presentation [Internet]. Ottawa (ON): Statistics Canada; 2009 [cited 2022 Oct]. Available from: https://www150.statcan.gc.ca/n1/pub/13f0026m/2007001/ch7-eng.htm
- 36. Immunize Canada. Influenza and pneumococcal immunization awareness campaign [Internet]. Ottawa (ON): Immunize Canada; unknown date [cited 2021 Oct]. Available from: https://immunize.ca/influenza-campaign
- 37. Andrew MK, McNeil S, Merry H, Rockwood K. Rates of influenza vaccination in older adults and factors associated with vaccine use: a secondary analysis of the Canadian Study of Health and Aging. BMC Public Health. 2004;4:36. https://doi.org/10.1186/1471-2458-4-36
- 38. Kan T, Zhang J. Factors influencing seasonal influenza vaccination behaviour among elderly people: a systematic review. Public Health. 2018; 156:67-78. https://doi.org/10.1016/j.puhe.2017.12.007
- 39. Singleton JA, Wortley P, Lu P-J. Influenza vaccination of persons with cardiovascular disease in the United States. Tex Heart Inst J. 2004;31(1): 22-7.

- 40. Winston CA, Wortley PM, Lees KA. Factors associated with vaccination of Medicare beneficiaries in five U.S. communities: results from the Racial and Ethnic Adult Disparities in Immunization Initiative survey, 2003. J Am Geriatr Soc. 2006;54(2):303-10. https://doi.org/10.1111/j.1532-5415.2005.00585.x
- 41. Lau JTF, Kim JH, Choi KC, Tsui HY, Yang X. Changes in prevalence of influenza vaccination and strength of association of factors predicting influenza vaccination over time—results of two population-based surveys. Vaccine. 2007;25(49):8279-89. https://doi.org/10.1016/j.vaccine.2007.09.047
- 42. Avelino-Silva VI, Avelino-Silva TJ, Miraglia JL, Miyaji KT, Jacob-Filho W, Lopes MH. Campaign, counseling and compliance with influenza vaccine among older persons. Clinics. 2011; 66(12):2031-5. https://doi.org/10.1590/S1807-59322011001200006
- 43. Ockene IS, Miller NH. Cigarette smoking, cardiovascular disease, and stroke: a statement for healthcare professionals from the American Heart Association. Circulation. 1997;96:3243-7. https://doi.org/10.1161/01.cir.96.9.3243
- 44. Farmanara N, Sherrard L, Dubé È, Gilbert NL. Determinants of non-vaccination against seasonal influenza in Canadian adults: findings from the 2015–2016 Influenza Immunization Coverage Survey. Can J Public Health. 2018;109(3):369-78. https://doi.org/10.17269/s41997-018-0018-9
- 45. Dyda A, Karki S, Hayen A, et al. Influenza and pneumococcal vaccination in Australian adults: a systematic review of coverage and factors associated with uptake. BMC Infect Dis. 2016;16(1):515. https://doi.org/10.1186/s12879-016-1820-8
- 46. Lix L, Yogendran M, Burchill C, et al. Defining and validating chronic diseases: an administrative data approach. Winnipeg (MB): Manitoba Centre for Health Policy; 2006. 217 p.