Research

SARS-CoV-2, Tdap, and influenza vaccination during pregnancy from 2019 to 2022 in Ontario, Canada: a population-based retrospective cohort study

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Abstract

Background: Hesitancy about vaccination during pregnancy posed challenges to SARS-CoV-2 vaccination efforts. We aimed to examine rates of SARS-CoV-2 vaccination among Ontario residents who gave birth in early 2022, and to compare rates of SARS-CoV-2 vaccine uptake with rates of tetanus, diphtheria, and pertussis (Tdap) and influenza vaccination during pregnancy in 2019, 2021, and 2022.

Methods: We conducted a populationbased retrospective cohort study to describe vaccination rates among pregnant and comparable nonpregnant populations in Ontario using linked administrative data. Provincially insured females who had a live, in-hospital birth from Jan. 1 to Mar. 31 in 2019, 2021, or 2022 were our primary cohort. Using log-binomial regression, we tested associations between SARS-CoV-2 (2022) and Tdap and influenza (2019, 2021, 2022) vaccination status, with birth group and covariates. We compared SARS-CoV-2 vaccination status with the status of a matched cohort of nonpregnant females and conducted subgroup analyses by age and prenatal clinician type.

Results: Among birthing people, 78.7% received their first SARS-CoV-2 vaccine dose and 74.2% received a second dose. The rate was significantly higher among nonpregnant comparators (dose 1: relative risk [RR] 0.94, 95% confidence interval [CI] 0.93–0.94; dose 2: RR 0.91, 95% CI 0.90–0.91). However, the rate of SARS-CoV-2 vaccination uptake among birthing people was higher than uptake of Tdap or influenza vaccination. Tetanus, diphtheria, and pertussis vaccination increased over

time from 22.2% in 2019 to 32.6% in 2022, and influenza vaccination rose to 35.3% in 2021 but returned to prepandemic levels in 2022 (27.7%). Vaccination rates were lower among pregnant people who were young, multiparous, or residents of rural or economically deprived areas for all 3 vaccines.

Interpretation: Rates of SARS-CoV-2 vaccination were lower among pregnant people than among nonpregnant comparators but were higher than rates of routinely recommended Tdap and influenza vaccinations. Pandemic urgency may have overcome a great deal of hesitancy about vaccinating against SARS-CoV-2 during pregnancy in 2022, but uptake of routinely recommended vaccines in pregnancy remains a challenge. **Trial registration:** Clinicaltrials.gov, no. NCT05663762.

Although pregnancy is associated with a higher risk of severe COVID-19,¹⁻⁸ pregnant people were not included in premarket clinical trials of SARS-CoV-2 vaccines.⁹⁻¹² On the basis of international vaccination registry data indicating safety of the messenger RNA (mRNA) vaccines against SARS-CoV-2, vaccination in pregnancy was recommended by several national organizations,^{13,14} and Ontario prioritized pregnant people for vaccine access beginning Apr. 23, 2021.¹⁵ Despite growing evidence of the vaccines' safety and effectiveness during pregnancy,¹⁶⁻²⁰ SARS-CoV-2 vaccine uptake was challenged by concerns about the safety of new vaccines and mRNA technology during pregnancy^{21,22} and fertility-focused misinformation, some of which

specifically targeted marginalized communities.^{23–27} Early analysis of SARS-CoV-2 vaccination during pregnancy in Ontario indicated relatively low SARS-CoV-2 vaccination coverage among those giving birth in 2021, with a trend of steadily increasing coverage over the course of that year;²⁸ many who initially did not receive the vaccination during pregnancy received it after their pregnancy.²⁹ Additional research is required to investigate reduced uptake and disparities in uptake among pregnant people who are eligible for vaccination.

Uptake of SARS-CoV-2 vaccination in pregnancy may be informed by assessing the rates of other routinely recommended vaccinations during pregnancy (seasonal influenza and

tetanus, diphtheria and pertussis [Tdap]), which have been low among pregnant people in Canada compared with rates among children or seniors,³⁰ or pregnant populations in other countries.³¹⁻³³ Barriers to vaccination receipt during pregnancy may both reflect and widen social and health disparities.^{29,30,34,35} Strong pan-Canadian recommendations³⁶ may be improving uptake,³⁷ although this progress may have been disrupted by the COVID-19 pandemic.^{38,39} In 2007, the Canadian National Advisory Committee on Immunization (NACI) strengthened its recommendation for influenza vaccination during pregnancy to specify that vaccination for all pregnant people was "particularly recommended" because the group is "at high-risk of influenza-related complications."40 Ontario publicly funds influenza vaccination during pregnancy through the province's Universal Influenza Immunization Program, which began in 2000.41 In 2018, NACI and the Society of Obstetricians and Gynaecologists of Canada recommended pertussis vaccination (in the form of the Tdap vaccine) in every pregnancy;³⁶ however, funding for this recommendation was not immediate in all provinces.⁴² Ontario started to publicly fund a routine dose of the Tdap vaccine in every pregnancy in April 2022.43

In this study, we aimed to examine whether Ontario residents who gave birth in early 2022, and who would have been eligible for vaccination during pregnancy, had lower receipt of SARS-CoV-2 vaccines than nonpregnant comparators, and to investigate whether they may have delayed vaccination until the postpartum period. Our secondary aim was to compare rates of Tdap and influenza vaccination among 2019, 2021, and 2022 birth groups, and to explore similarities and differences between Tdap and SARS-CoV-2 vaccination uptake. We hypothesized that rates of SARS-CoV-2 vaccination would be lower among pregnant people than among their comparators, that the 2019 group would have higher uptake of Tdap and influenza vaccination than the pandemic groups, and that pregnant people who received Tdap vaccination would be more likely to receive SARS-CoV-2 vaccination than those who did not receive Tdap vaccination.

Methods

We conducted a population-based retrospective cohort study to describe and compare rates of SARS-CoV-2, Tdap, and influenza vaccination among pregnant and comparable nonpregnant populations. This study is part of a convergent parallel mixed-methods study examining changes in health service use in pregnancy during the COVID-19 pandemic (Appendix 1, available at www.cmaj.ca/lookup/doi/10.1503/cmaj.231522/tab-related-content); the protocol has been published⁴⁴ and prospectively registered (NCT05663762). We report the results in accordance with the Reporting of Studies Conducted Using Observational Routinely-collected Health Data statement.⁴⁵

Setting

We conducted the study in Ontario, Canada, which maintains multiple health administrative data holdings for publicly funded health services, including those for about 140000 live births each year.⁴⁶

Data sources and study population

We linked data sets using encoded identifiers and analyzed them at ICES. Data sets included a derived data set linking infants born in Ontario hospitals to birthing people (MOMBABY), Ontario's central SARS-CoV-2 vaccination database (COVaxON), and the Ontario Health Insurance Plan (OHIP) claims database for Tdap and influenza vaccinations. Full data sources are listed in Appendix 2 (available at www.cmaj.ca/lookup/doi/10.1503/cmaj.231522/tab -related-content); details of data sets and cohort creation have been published.⁴⁴ Our primary cohort included provincially insured females who had a live, in-hospital birth between Jan. 1 and Mar. 31 in 2019, 2021, or 2022. The 3 birth years represent distinct periods of the COVID-19 pandemic with no overlapping gestational periods.⁴⁴ Eligible patents had a valid ICES key number, delivery date, and birth date. We assigned date of childbirth as the index event; the lookback window spanned gestation (Appendix 3, available at www.cmaj.ca/lookup/doi/10.1503/cmaj.231522/tab -related-content). We established a comparator cohort of nonpregnant females aged 18-44 who received health care in Ontario between Jan. 1 and Mar. 31, 2022, for matched analysis.

Exposure

Primary exposures were antenatal periods for the 2019, 2021, and 2022 birth groups.⁴⁴ Births between Jan. 1 and Mar. 31, 2019, were pregnancies and births before the COVID-19 pandemic; those between Jan. 1 and Mar. 31, 2021, occurred during the pandemic but before widespread vaccine availability; and those between Jan. 1 and Mar. 31, 2022, were patients who were pregnant and gave birth after widespread vaccine availability.

Outcomes

SARS-CoV-2, Tdap, and influenza vaccinations were the outcomes of interest. We assessed the SARS-CoV-2 outcome only for the 2022 birth group, who would have had access to vaccines during the entirety of their gestational period.¹⁵ We observed first and second dose coverage from the release of SARS-CoV-2 vaccines (December 2020) up to 3 months after the study's latest birth date (June 2022), and observed timing of these doses in relation to pregnancy. We classified vaccination timing by comparing gestational age (reported or estimated for the n = 23 records without gestational age by subtracting 40 wk from childbirth date), birth date, and vaccination date. We assessed Tdap and influenza vaccination receipt during gestation for all patients in our primary cohort by examining the relevant physician billing claims in OHIP.

Covariates, potential confounders, and effect modifiers

Sociodemographic characteristics and health service use may affect vaccination status. We selected a priori 10 independent measures collected in administrative health data by consulting experts and reviewing literature (Appendix 2): parity, use of assisted reproductive technology, gestational length, singleton or multiple birth, perinatal care provider type, early perinatal care visits, delivery type, age, rurality, and socioeconomic "deprivation" (reported by dissemination area–level Canadian Index of Multiple Deprivation [CIMD] ethnocultural composition, residential instability, and economic dependency).⁴⁷



Figure 1: Flow diagram showing primary cohort creation and exclusions. Note: IKN = ICES key number, MOMBABY = Ontario Mother–Baby linked data set, OHIP = Ontario Health Insurance Plan.

Statistical analysis

Descriptive analysis

We report sociodemographic characteristics, use of perinatal health services, and birth outcomes using measures of general frequency (counts and percentages), central tendency (means and medians), and dispersion (standard deviations and interquartile ranges [IQRs]). To account for minimal missingness in rurality and CIMD values (< 0.5%), we performed a single imputation using predictive mean matching.

Inferential analysis

We tested associations between outcomes, primary exposure (birth group), and other covariates using log-binomial regression models with robust variance estimation to estimate relative risk (RR) and associated confidence intervals (CIs). For our primary outcome (SARS-CoV-2 vaccination in pregnant and nonpregnant groups), we used propensity score matching to minimize effects of confounding.48 We restricted the 2022 birth group to ages 18-44 years and performed a propensity score-matched analysis with a cohort of similarly aged nonpregnant females. We estimated the propensity score by logistic regression using age, rurality, parity, and the 3 CIMD factors to estimate the probability of being in the pregnant female group. We greedy matched the 2 cohorts 1:1 using a caliper of 0.2 times the standard deviation of the logit of the propensity score. For the remaining multivariable analyses, we forced all factors into our models to reflect the conscientious process by which they were selected and our exploratory objectives. We performed all analyses using SAS software, version 9.4.

Sensitivity analysis

We conducted prespecified subgroup analyses to examine differences by 2 factors identified in our literature review as potentially influential: prenatal clinician type and age (which we categorized into birthing people aged < 25 yr v. \ge 25 yr) following the same methods previously described. When comparing SARS-CoV-2 vaccinations by age, we re-ran the matching procedure with an additional hard match on age. We also stratified our 2022 group by birth month to identify any differences in SARS-CoV-2 vaccine uptake.

Ethics approval

Section 45 of Ontario's *Personal Health Information Protection Act* allows ICES to collect and analyze health care and demographic data, without consent, for health system evaluation and improvement. Study approval was waived by the Hamilton Integrated Research Ethics Board and granted by the University of British Columbia Behavioural Research Ethics Board (H22-01905).

Results

We identified 86815 people who gave birth between Jan. 1 and Mar. 31 in 2019 (n = 29181), 2021 (n = 29258), and 2022 (n = 28376) (Figure 1). The median age was 32 years (Table 1). Obstetricians most frequently provided prenatal care, and the median number of prenatal visits before 32 weeks of gestation was 5 (IQR 3–7). A total of 43.6% gave birth for the first time and 3.5% used assisted reproductive technology. The comparator cohort contained 2576817 eligible nonpregnant females aged 18–44 years; propensity score matching resulted in 28177 people in each group (Table 2). The standardized differences of all descriptive factors were 0.1 or less, suggesting comparability between groups.

Among those in the 2022 birth group, 22 508 (79.6%) received an initial SARS-CoV-2 vaccine dose and 21 435 (75.5%) received a second dose by June 30, 2022. Of those who were vaccinated, nearly half (49.9%) received the first dose before pregnancy, and 3.9% delayed until after delivery. Among the nonpregnant

Table 1 (part 1 of 2): Sociodemographic characteristics, perinatal health services contact, and vaccination information among patients in the primary birth cohorts

	No. (%)*			
Variable	Overall n = 86 815	2019 birth group n = 29 181	2021 birth group n = 29 258	2022 birth group n = 28 376
Sociodemographic characteristics				
Age, yr, median (IQR)	32 (28–35)	31 (28–35)	31 (28–35)	32 (29–35)
Rural residence	6202 (7.1)	2094 (7.2)	2066 (7.1)	2042 (7.2)
Neighbourhood-level ethnocultural composition†				
1 (least diversity)	13 533 (15.6)	4556 (15.6)	4604 (15.7)	4373 (15.4)
2	13 997 (16.1)	4600 (15.8)	4795 (16.4)	4602 (16.2)
3	15 527 (17.9)	5128 (17.6)	5211 (17.8)	5188 (18.3)
4	19 044 (21.9)	6387 (21.9)	6361 (21.7)	6296 (22.2)
5 (greatest diversity)	24 714 (28.5)	8510 (29.2)	8287 (28.3)	7917 (27.9)
Neighbourhood-level economic dependency†				
1 (least dependency)	24 963 (28.8)	8340 (28.6)	8435 (28.8)	8188 (28.9)
2	18 099 (20.8)	5980 (20.5)	6162 (21.1)	5957 (21.0)
3	16 161 (18.6)	5519 (18.9)	5278 (18.0)	5364 (18.9)
4	14 839 (17.1)	5001 (17.1)	5055 (17.3)	4783 (16.9)
5 (greatest dependency)	12 753 (14.7)	4341 (14.9)	4328 (14.8)	4084 (14.4)
Neighbourhood-level residential instability†				
1 (least instability)	12 601 (14.5)	4190 (14.4)	4353 (14.9)	4058 (14.3)
2	16 387 (18.9)	5374 (18.4)	5604 (19.2)	5409 (19.1)
3	19 047 (21.9)	6425 (22.0)	6311 (21.6)	6311 (22.2)
4	18 645 (21.5)	6159 (21.1)	6274 (21.4)	6212 (21.9)
5 (greatest instability)	20 135 (23.2)	7033 (24.1)	6716 (23.0)	6386 (22.5)
Perinatal health services contact				
Use of assisted reproductive technology for current pregnancy	3061 (3.5)	1128 (3.9)	595 (2.0)	1338 (4.7)
Early prenatal care visits (count), median (IQR)‡	5 (3–7)	6 (4–8)	5 (3–7)	5 (3–7)
Prenatal care provider type				
Obstetrician	56 717 (65.3)	18 931 (64.9)	19 330 (66.1)	18 456 (65.0)
Family physician	12 308 (14.2)	4459 (15.3)	3966 (13.6)	3883 (13.7)
Shared care§	2280 (2.6)	741 (2.5)	758 (2.6)	781 (2.8)
Midwife	14 372 (16.6)	4761 (16.3)	4794 (16.4)	4817 (17.0)
No care or other	1138 (1.3)	289 (1.0)	410 (1.4)	439 (1.5)
Parity				
0	37 846 (43.6)	12 390 (42.5)	12 973 (44.3)	12 483 (44.0)
1	31 365 (36.1)	10 737 (36.8)	10 416 (35.6)	10 212 (36.0)
2	11 276 (13.0)	3874 (13.3)	3747 (12.8)	3655 (12.9)
≥3	6328 (7.3)	2180 (7.5)	2122 (7.3)	2026 (7.1)
≥ 1 infant(s) aged ≤ 24 mo in household	11 433 (13.2)	3882 (13.3)	3824 (13.1)	3727 (13.1)

matched cohort, 82.6% received an initial dose and 80.5% a second dose in the same period (Table 2); pregnant people were less likely than nonpregnant females to be vaccinated with either SARS-CoV-2 dose (dose 1: RR 0.94, 95% Cl 0.93–0.94; dose 2: RR 0.91, 95% Cl 0.90–0.91) (Table 3 and Figure 2). Rural residence and increasing neighbourhood-level economic dependency were

negatively associated with receiving each SARS-CoV-2 vaccine dose during pregnancy (Table 3). We did not observe differences by birthing month (Appendix 4, available at www.cmaj.ca/lookup/doi/10.1503/cmaj.231522/tab-related-content).

Overall, 24923 (28.7%) birthing people received Tdap vaccination during pregnancy; the proportion vaccinated increased

Table 1 (part 2 of 2): Sociodemographic characteristics, perinatal health services contact, and vaccination information among patients in the primary birth cohorts

	No. (%)*			
Variable	Overall n = 86 815	2019 birth group n = 29 181	2021 birth group n = 29 258	2022 birth group n = 28 376
SARS-CoV-2 vaccination (2022 birth group only)				
First dose				
Overall¶	-	-	-	22 581 (79.6)
Timing				
Before conception	-	-	-	11 264 (49.9)
During pregnancy	-	-	-	10 435 (46.2)
After giving birth	-	-	-	882 (3.9)
Second dose				
Overall§	-	-	-	21 425 (75.5)
Timing				
Before conception	-	-	-	2929 (13.7)
During pregnancy	-	-	-	17 595 (82.1)
After giving birth	-	-	-	901 (4.2)
SARS-CoV-2 and Tdap vaccination (2022 birth group only)				
Received dose 1 and Tdap vaccine	-	-	-	8347 (29.4)
Received dose 1 but not Tdap vaccine	-	-	-	14 234 (50.2)
Received Tdap vaccine but not dose 1	-	-	-	916 (3.2)
Received neither Tdap vaccine nor dose 1	-	-	-	4879 (17.2)
Received dose 2 and Tdap vaccine	-	-	-	8086 (28.5)
Received dose 2 but not Tdap vaccine	-	-	-	13 339 (47.0)
Received Tdap vaccine but not dose 2	-	-	-	1177 (4.1)
Received neither Tdap vaccine nor dose 2	-	-	-	5774 (20.3)
Tdap and influenza vaccination				
Received both Tdap and influenza vaccine	12 877 (14.8)	3734 (12.8)	5186 (17.7)	3957 (13.9)
Received influenza but no Tdap vaccine	13 676 (15.8)	4614 (15.8)	5152 (17.6)	3910 (13.8)
Received Tdap but no influenza vaccine	12 046 (13.9)	2742 (9.4)	3998 (13.7)	5306 (18.7)
Received neither influenza nor Tdap vaccine	48 216 (55.5)	18 091 (62.0)	14 922 (51.0)	15 203 (53.6)

Note: IQR = interquartile range; Tdap = tetanus, diphtheria and pertussis.

*Unless stated otherwise.

†Measured as quintiles.

‡Early prenatal care visits occurred before 32 weeks of gestation.

§Includes patients with an equal number of minor prenatal assessment billings by family physicians and obstetricians between conception date and index date.

Vaccine records were examined for each birthing person between Dec. 1, 2020, and up to June 30, 2022.

between 2019 and 2022 (Table 1). After multivariable adjustment, the temporal effect remained significant (Table 4). All variables except assisted reproductive technology were significantly associated with Tdap vaccination.

Pregnant people who received Tdap vaccination (n = 9263) were more likely to also receive either dose of SARS-CoV-2 vaccination (dose 1, n = 8347, 90.1%; dose 2, n = 8086, 87.3%) than those who did not receive Tdap vaccination (n = 19113) (dose 1, n = 14234, 74.5%; dose 2, n = 13339, 69.8%).

Overall, influenza vaccination was received by 26553 (30.6%) birthing persons, with higher uptake in 2021 than either 2019 or 2022 (Table 1).

The factors associated with SARS-CoV-2 vaccination for birthing people across type of prenatal clinician are shown in Table 5. Age was significantly associated with uptake of both doses across all clinician types. Among those cared for by family physicians, primary care model was not associated with significant differences.

Across all clinician types, Tdap vaccination increased from 2019 to 2022 (Table 6). In multivariable analysis, later birth group, older age, and more early prenatal care visits were associated with greater uptake, and rurality and parity with lower uptake. Among patients of family physicians, Tdap vaccination differed by primary care model Table 2: Sociodemographic characteristics, perinatal health services contact, and receipt of SARS-COV-2 vaccination of matched birth cohort and nonpregnant comparator cohort

	No.					
Variable	2022 birth cohort n = 28 177	Matched nonpregnant cohort n = 28 177	Standardized difference			
Sociodemograph	ic characteristic	:s				
Age, yr, median (IQR)	32 (29–35)	32 (29–35)	0.00			
Rural residence	2023 (7.2)	2012 (7.1)	0.00			
Neighbourhood-le	evel ethnocultura	al composition†				
1 (least diversity)	4338 (15.4)	4346 (15.4)	0.00			
2	4576 (16.2)	4577 (16.2)	0.00			
3	5150 (18.3)	5131 (18.2)	0.00			
4	6249 (22.2)	6248 (22.2)	0.00			
5 (greatest diversity)	7864 (27.9)	7875 (27.9)	0.00			
Neighbourhood-le	evel economic de	ependency†				
1 (least dependency)	8142 (28.9)	8144 (28.9)	0.00			
2	5925 (21.0)	5925 (21.0)	0.00			
3	5330 (18.9)	5330 (18.9)	0.00			
4	4747 (16.8)	4748 (16.9)	0.00			
5 (greatest dependency)	4033 (14.3)	4028 (14.3)	0.00			
Neighbourhood-le	evel residential ir	nstability†				
1 (least instability)	4035 (14.3)	4029 (14.3)	0.00			
2	5373 (19.1)	5369 (19.1)	0.00			
3	6275 (22.3)	6284 (22.3)	0.00			
4	6168 (21.9)	6170 (21.9)	0.00			
5 (greatest instability)	6326 (22.5)	6325 (22.4)	0.00			
Perinatal health	services contact	:				
Parity						
0	12 363 (43.9)	12 361 (43.9)	0.00			
1	10 170 (36.1)	10 172 (36.1)	0.00			
2	3640 (12.9)	3640 (12.9)	0.00			
≥3	2004 (7.1)	2004 (7.1)	0.00			
SARS-CoV-2 vacc	ination					
First dose‡	22 431 (79.6)	23 282 (82.6)	NA			
Second dose‡	21 289 (75.6)	22 682 (80.5)	NA			
Note: IQR = interquartile range, NA = not applicable. *Unless stated otherwise.						

[‡]Vaccine records were examined between Dec. 1, 2020, and June 30, 2022.

†Measured as quintiles.

Table 3: Factors associated with SARS-CoV-2 vaccination among pregnant people compared with matched nonpregnant females

	Adjusted RR‡ (95% CI)			
Variable	First SARS-CoV-2 vaccine dose	Second SARS-Cov-2 vaccine dose		
SARS-CoV-2 vaccir	nation			
Pregnancy status*				
Pregnant	0.94 (0.93-0.94)	0.91 (0.90-0.91)		
Nonpregnant†	-	-		
Sociodemographi	c characteristics			
Age, yr	1.00 (1.00-1.00)	1.00 (1.00-1.00)		
Rural residence	0.94 (0.94–0.94)	0.94 (0.93-0.94)		
Neighbourhood-lev	vel ethnocultural composi	ition		
1 (least diversity)†	-	-		
2	1.00 (1.00-1.00)	1.00 (1.00-1.00)		
3	1.01 (1.00-1.01)	1.00 (1.00-1.01)		
4	1.00 (1.00-1.00)	1.00 (1.00-1.00)		
5 (greatest diversity)	1.00 (1.00-1.00)	1.00 (1.00-1.00)		
Neighbourhood-lev	vel economic dependency	/		
1 (least dependency)†	-	-		
2	0.98 (0.98–0.98)	0.98 (0.98-0.98)		
3	0.98 (0.98–0.98)	0.98 (0.97–0.98)		
4	0.97 (0.97–0.97)	0.97 (0.97–0.97)		
5 (greatest diversity)	0.98 (0.97–0.97)	0.97 (0.96–0.98)		
Neighbourhood-lev	vel residential instability			
1 (least instability)†	-	-		
2	1.00 (1.00-1.00)	1.02 (0.99–1.05)		
3	0.99 (0.99–0.99)	1.01 (0.98-1.04)		
4	0.99 (0.99–0.99)	1.01 (0.98-1.04)		
5 (greatest instability)	0.97 (0.97–0.97)	0.99 (0.96–1.02)		
Perinatal health s	ervices contact			
Parity				
0†	-	-		
1	0.99 (0.98–0.99)	0.98 (0.98–0.98)		
2	1.02 (1.02–1.02)	1.02 (1.02–1.02)		
≥3	0.94 (0.94–0.95)	0.93 (0.93–0.93)		
Noto: CI = confidence	intonual BB = relativo sisk			

Note: CI = confidence interval, RR = relative risk.

*Vaccine records were examined between Dec. 1, 2020, and June 30, 2022. †Reference group.

Adjusted for maternal age, rurality, Canadian Index of Multiple Deprivation measures (i.e., ethnocultural composition, economic dependency, and residential instability), and parity.

SARS-CoV-2 vaccination dose 1 receipt: pregnant v. nonpregnant



SARS-CoV-2 vaccination dose 2 receipt: pregnant v. nonpregnant



Figure 2: Timing of SARS-CoV-2 vaccinations, by pregnant versus nonpregnant status. The vertical dashed lines represent the birthing windows of the cohort (i.e., Jan. 1 to Mar. 31).

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Table 4: Factors associated with tetanus, diphtheria, andpertussis vaccination among pregnant people

Variable	Adjusted RR* (95% CI)
Birthing group	
2019 birth†	-
2021 birth	1.46 (1.42-1.50)
2022 birth	1.50 (1.46-1.54)
Sociodemographic characteristics	
Age, yr	1.02 (1.02-1.03)
Rural residence	0.83 (0.79-0.88)
Neighbourhood-level ethnocultural compos	ition
1 (least diversity)†	-
2	1.08 (1.04-1.13)
3	1.17 (1.13–1.22)
4	1.19 (1.15–1.24)
5 (greatest diversity)	1.18 (1.14-1.23)
Neighbourhood-level economic dependency	/
1 (least dependency)†	-
2	0.96 (0.93–0.99)
3	0.91 (0.89-0.94)
4	0.88 (0.85-0.91)
5 (greatest dependency)	0.86 (0.83-0.89)
Neighbourhood-level residential instability	
1 (least instability)†	-
2	0.98 (0.94-1.01)
3	0.95 (0.91-0.98)
4	0.91 (0.88-0.94)
5 (greatest instability)	0.89 (0.86-0.93)
Perinatal health services contact	
Use of assisted reproductive technology for current pregnancy	0.95 (0.90-1.00)
Early prenatal care visits‡	1.04 (1.03-1.04)
Prenatal care provider type	
Obstetrician†	-
Family physician	1.38 (1.34–1.42)
Shared care	1.20 (1.13–1.27)
Midwife	1.05 (1.01–1.09)
No care or other	0.51 (0.43-0.60)
Parity	
0†	-
1	0.81 (0.79–0.83)
2	0.62 (0.60-0.65)
≥3	0.45 (0.42-0.47)

Note: CI = confidence interval, RR = relative risk.

*Adjusted for birth group, maternal age, rurality, Canadian Index of Multiple Deprivation measures (i.e., ethnocultural composition, economic dependency, and residential instability), use of assisted reproductive technology, early prenatal care visits, type of prenatal care provider, and parity. TReference group.

‡Early prenatal care visits occurred before 32 weeks of gestation.

The overall birthing cohort contained 7916 (9.1%) people younger than age 25 years. Of the 28376 people who gave birth in the 2022 group, 2275 (8.0%) were younger than 25 years. Crude analysis showed that they were significantly less likely to receive either the first or second SARS-CoV-2 vaccine than birthing people aged 25 years and older. After model adjustment, this difference decreased but remained significant (Appendix 5, available at www.cmaj.ca/lookup/doi/10.1503/cmaj.231522/ tab-related-content).

The comparator cohort contained 585 009 nonpregnant females whom we hard-matched by age to the cohort of pregnant people younger than 25 years, resulting in 2191 matches. This analysis showed that young people who gave birth were significantly less likely than matched comparators to receive either dose of SARS-CoV-2 vaccination in both the unadjusted and matched models (Appendix 6, available at www.cmaj.ca/lookup/ doi/10.1503/cmaj.231522/tab-related-content).

Interpretation

Among the first group of Ontario residents eligible for SARS-CoV-2 vaccination during their entire pregnancies, most were vaccinated (78.7%) and completed the 2-dose series (74.2%). This is a high uptake for a new vaccine or any vaccine during pregnancy, likely driven by a pandemic-related sense of urgency, including emerging evidence on the risks of SARS-CoV-2 infection to pregnant people and the possibility of benefit to the newborn from transplacental passage of antibodies after vaccination.^{22,49,50} As hypothesized, uptake of SARS-CoV-2 vaccination was slightly lower among pregnant people than among matched nonpregnant females in Ontario, despite priority access for pregnant people. That said, only 3.9% delayed vaccination until after delivery, suggesting that the decision not to take the vaccine was either not strongly tied to specific concerns about pregnancy or continued owing to hesitancy about vaccination while breastfeeding. Given that SARS-CoV-2 vaccination was largely administered in special vaccination clinics rather than during routine office visits in 2022, it is unsurprising that prenatal clinician type was not associated with vaccine receipt.

In contrast to the rate of SARS-CoV-2 vaccination, a minority of pregnant people received Tdap (28.7%) or influenza (30.6%) vaccination during pregnancy, suggesting that they weighed the benefits and risks of the vaccines differently. Contrary to our second hypothesis that the pandemic would have a negative impact on Tdap and influenza vaccination, Tdap uptake increased considerably, from 22.2% in 2019 to 31.4% in 2021 and 32.6% in 2022. Whereas an increase of Tdap vaccination has been observed by others using a similar data set from 2011/12 to 2019/20,³⁷ the large increase from 2019 to 2021 may have been associated with increased attention to vaccination related to the release of SARS-CoV-2 vaccines, as influenza vaccination also rose in 2021. However, uptake of influenza vaccination did not follow the same trend as Tdap vaccination uptake, falling back to prepandemic levels in 2022. Our third hypothesis, that those who received Tdap vaccination would be more likely to receive SARS-CoV-2 vaccination was supported by this analysis, suggesting

	Obstetrician;		Family physician;		Midwife;	
	adjusted RR* (95% CI)		adjusted RR*† (95% CI)		adjusted RR* (95% CI)	
	n = 18 456		n = 3883		n = 4817	
Variable	First dose	Second dose	First dose	Second dose	First dose	Second dose
Sociodemographic characteristics						
Age, yr	1.01	1.01	1.00	1.00	1.01	1.01
	(1.01-1.01)	(1.01-1.01)	(1.00-1.01)	(1.00-1.01)	(1.01-1.01)	(1.01–1.02)
Rural residence	0.95	0.93	0.98	0.98	0.96	0.95
	(0.92–0.99)	(0.89–0.97)	(0.95–1.01)	(0.95–1.01)	(0.88–1.04)	(0.85–1.07)
Neighbourhood-level ethnocultural composition						
1 (least diversity)‡	-	-	-	-	-	-
2	0.99	0.98	0.99	1.00	1.00	1.02
	(0.97–1.01)	(0.96–1.01)	(0.96–1.03)	(0.97–1.03)	(0.94–1.06)	(0.94–1.10)
3	0.99	0.98	0.99	0.99	1.00	1.01
	(0.97–1.01)	(0.96–1.01)	(0.96–1.02)	(0.96–1.02)	(0.94–1.06)	(0.9–1.09)
4	1.00	0.99	0.99	0.99	0.98	0.99
	(0.98–1.02)	(0.96–1.01)	(0.96–1.02)	(0.95–1.02)	(0.92–1.04)	(0.91–1.07)
5 (greatest diversity)	1.01	0.99	0.98	0.98	0.97	0.95
	(0.99–1.03)	(0.97–1.01)	(0.95–1.02)	(0.94–1.01)	(0.90-1.04)	(0.87–1.05)
Neighbourhood-level economic dependency						
1 (least dependency)‡	-	-	-	-	-	-
2	0.98	0.98	0.99	0.99	1.01	1.02
	(0.97–1.00)	(0.96–1.00)	(0.96–1.03)	(0.96–1.02)	(0.96–1.06)	(0.96–1.08)
3	0.98	0.97	0.99	0.98	0.97	0.96
	(0.96–0.99)	(0.95–0.99)	(0.95–1.02)	(0.95–1.01)	(0.92–1.02)	(0.90–1.029)
4	0.97	0.96	0.98	0.98	0.95	0.92
	(0.95–0.99)	(0.94–0.98)	(0.95–1.01)	(0.95–1.01)	(0.90-1.01)	(0.85–1.00)
5 (greatest dependency)	0.97	0.96	0.98	0.98	0.96	0.95
	(0.95–0.99)	(0.94–0.98)	(0.95–1.02)	(0.94–1.01)	(0.90–1.02)	(0.88–1.03)
Neighbourhood-level residential instability						
1 (least instability)‡	-	-	-	-	-	-
2	1.01	1.01	0.99	0.99	1.02	1.00
	(0.99–1.03)	(0.99–1.04)	(0.96–1.03)	(0.96–1.02)	(0.96–1.09)	(0.93–1.08)
3	1.01	1.01	0.99	0.99	1.03	1.04
	(0.99–1.03)	(0.99–1.03)	(0.96–1.03)	(0.96–1.02)	(0.97–1.09)	(0.97–1.12)
4	0.99	0.99	1.00	0.99	1.05	1.05
	(0.97–1.01)	(0.97–1.02)	(0.96–1.03)	(0.96–1.03)	(0.99–1.11)	(0.98–1.13)
5 (greatest instability)	0.99	0.99	0.99	0.99	0.99	0.99
	(0.97–1.01)	(0.96–1.01)	(0.95–1.03)	(0.95–1.02)	(0.93–1.06)	(0.90–1.08)
Use of assisted reproductive technology for current pregnancy	1.00	0.99	1.00	1.00	1.05	1.01
	(0.98–1.02)	(0.97–1.02)	(0.92–1.09)	(0.93–1.08)	(0.95–1.15)	(0.90-1.13)
Early prenatal care visits§	1.00	1.01	1.00	1.01	1.01	1.01
	(1.00-1.01)	(1.00-1.01)	(1.00-1.01)	(1.00-1.01)	(0.99–1.02)	(0.90-1.13)
Parity						
0‡	-	-	-	-	-	-
1	0.98	0.97	0.99	0.98	0.99	0.99
	(0.97–0.99)	(0.96–0.98)	(0.96–1.01)	(0.96–1.00)	(0.95–1.03)	(0.94–1.04)
2	0.92	0.88	0.96	0.95	0.94	0.86
	(0.90–0.93)	(0.86–0.90)	(0.93–0.99)	(0.92–0.98)	(0.89-1.01)	(0.79–0.94)
≥3	0.84	0.77	0.94	0.92	0.77	0.64

Table 5 (part 2 of 2): Factors associated with SARS-CoV-2 vaccination among pregnant people, by type of prenatal care provider

	Obstetrician; adjusted RR* (95% CI) n = 18 456		Family physician; adjusted RR*† (95% CI) n = 3883		Midwife; adjusted RR* (95% CI) n = 4817	
Variable	First dose	Second dose	First dose	Second dose	First dose	Second dose
Primary care contact						
Rostered to family physician	-	-	1.00 (0.97–1.02)	1.00 (0.98–1.02)	-	-
Primary care model¶						
Capitation‡	-	-	-	-	-	-
Comprehensive care model	-	-	1.02 (0.95–1.09)	1.00 (0.93–1.06)	-	-
Missing	-	-	1.00 (0.97–1.04)	1.00 (0.97–1.04)	-	-
Other	-	-	1.01 (0.98–1.04)	1.01 (0.98–1.04)	-	-

Note: CI = confidence interval, RR = relative risk.

*Adjusted for maternal age, rurality, Canadian Index of Multiple Deprivation measures (i.e., ethnocultural composition, economic dependency, and residential instability), use of assisted reproductive technology, early prenatal care visits, and parity.

†Also adjusted for whether the pregnant person was attached or rostered to their family physician and the physicians' practice model.

‡Reference group.

§Early prenatal care visits occurred before 32 weeks of gestation.

¶Capitation-based models include teams of 6 or more physicians who are compensated primarily through capitation payments but also receive fee-for-service payments. Comprehensive care model physicians are compensated primarily through fee-for-service. Both capitation and comprehensive care model physicians are eligible for specific bonuses and premiums based on patient enrolment. Other patient enrolment models were grouped together owing to small cell sizes and include family health groups and specialized models (such as general practice- or focused practice-physicians with alternative funding plans).

that those who did not receive routine vaccinations during pregnancy were at greater risk of also not receiving pandemic vaccinations. However, given different data sources for SARS-CoV-2 vaccination and Tdap and influenza vaccinations, we believe this finding requires further research.

As has been previously observed with Tdap and influenza vaccination during pregnancy,³⁰ those seen by family physicians were more likely to receive Tdap vaccination than those attended by obstetricians or midwives,^{30,34,35} possibly owing to family practice offices being equipped to offer vaccination onsite. These findings suggest that stocking and administering recommended vaccinations in obstetric and midwifery practices could increase uptake among pregnant people. Pregnant people younger than 25 years, along with rural residents, those who had previously given birth, and those residing in neighbourhoods with greater economic dependency were significantly less likely to receive vaccination during pregnancy, suggesting that inequities that posed barriers to Tdap and influenza vaccination in the past continued during the COVID-19 pandemic despite greater focus on vaccination and increased overall receipt of vaccination during pregnancy. Addressing disparities in access to vaccination may help close this gap.

Future research should examine uptake of SARS-CoV-2 vaccination among pregnant people over time and compare jurisdictions with different vaccine implementation approaches and social or epidemiologic conditions. Qualitative research examining pregnant people's experiences with vaccine decisions during the pandemic could help explain the patterns observed. People who are pregnant or planning to become pregnant should be included in clinical trials of vaccines intended for their use, to provide highguality evidence to support clinical and public health communication and decision-making.⁵¹ Despite the challenges this may pose for safety surveillance of new vaccines, offering coadministration of vaccines (e.g., Tdap and influenza vaccination at the same time as SARS-CoV-2 vaccination, rather than potentially requiring patients to prioritize and make multiple appointments) could improve uptake. Because additional vaccines may be recommended in Canada during pregnancy (e.g., the newly approved vaccine against respiratory syncytial virus,⁵² potential future vaccines against group B streptococcal disease,⁵³ or against future pandemic pathogens), issues around collecting safety surveillance data for new vaccines and coadministration with other vaccinations should be considered. Although the predictors of receiving 1 or 2 doses were similar in this analysis, future research should investigate whether there are observable differences (e.g., in timing during or in relation to pregnancy between the minority of vaccine recipients who received only 1 dose and the majority who received both doses, and seek to identify predictors of receiving subsequent doses among those who received a first dose. Given low uptake of additional doses that have been recommended beyond the initial 2-dose series, this may be increasingly of interest if SARS-CoV-2 vaccination is eventually recommended in every pregnancy, similar to Tdap and influenza vaccinations.

Limitations

Ascertaining Tdap and influenza vaccination status from OHIP billing claims likely underestimated the total number of vaccinated

Table 6 (part 1 of 2): Crude and adjusted analyses of tetanus, diphtheria, and pertussis vaccination, by prenatal care provider type

Type of prenatal care provider	Obstetrician <i>n</i> = 56 717	Family physician n = 12 308	Midwife n = 14 372
Crude analysis		RR (95% CI)	
Birthing group			
2019 birth*	-	-	-
2021 birth	1.46 (1.41–1.51)	1.28 (1.21-1.36)	1.42 (1.32–1.54)
2022 birth	1.54 (1.49-1.60)	1.22 (1.15-1.30)	1.55 (1.44-1.67)
Adjusted analysis		Adjusted RR§ (95% CI)	
Birthing group			
2019 birth*	-	-	-
2021 birth	1.49 (1.45–1.55)	1.32 (1.24–1.40)	1.46 (1.35–1.57)
2022 birth	1.56 (1.51–1.61)	1.26 (1.09–1.33)	1.56 (1.45–1.68)
Sociodemographic characteristics			
Age, yr	1.02 (1.02–1.03)	1.02 (1.01–1.02)	1.04 (1.03–1.04)
Rural residence	0.82 (0.75–0.90)	0.81 (0.75-0.88)	0.77 (0.67–0.89)
Neighbourhood-level ethnocultural composition			
1 (least diversity)*	-	-	-
2	1.06 (1.00-1.12)	1.11 (1.03–1.20)	1.13 (1.02–1.25)
3	1.18 (1.11–1.24)	1.09 (1.00–1.17)	1.29 (1.17–1.42)
4	1.21 (1.15–1.28)	1.10 (1.01–1.19)	1.28 (1.16–1.41)
5 (greatest diversity)	1.24 (1.17–1.30)	1.03 (0.95–1.11)	1.01 (0.90-1.13)
Neighbourhood-level economic dependency			
1 (least dependency)*	-	-	-
2	0.98 (0.94–1.01)	0.97 (0.91–1.04)	0.87 (0.81–0.94)
3	0.91 (0.88–0.95)	1.00 (0.94–1.08)	0.82 (0.76-0.90)
4	0.88 (0.85–0.92)	0.95 (0.88–1.02)	0.82 (0.75–0.89)
5 (greatest dependency)	0.87 (0.83-0.91)	0.92 (0.85–1.00)	0.75 (0.68–0.83)
Neighbourhood-level residential instability			
1 (least instability)*	-	-	-
2	0.97 (0.92–1.01)	1.00 (0.92–1.08)	0.98 (0.89–1.08)
3	0.94 (0.90–0.98)	0.96 (0.89–1.03)	0.92 (0.84–1.01)
4	0.92 (0.88–0.96)	0.88 (0.82–0.95)	0.95 (0.87–1.04)
5 (greatest instability)	0.89 (0.85–0.93)	0.92 (0.85–1.00)	0.89 (0.81-0.98)
Perinatal health services contact			
Use of assisted reproductive technology for current pregnancy	0.92 (0.87–0.98)	0.96 (0.81–1.14)	1.06 (0.92–1.22)
Early prenatal care visits†	1.03 (1.03-1.04)	1.05 (1.04–1.06)	1.04 (1.03–1.05)
Parity			
0*	-	-	-
1	0.80 (0.78–0.82)	0.88 (0.84–0.93)	0.78 (0.73–0.83)
2	0.60 (0.57–0.62)	0.75 (0.69–0.81)	0.57 (0.51-0.64)
≥3	0.43 (0.40-0.47)	0.59 (0.23–0.67)	0.39 (0.32–0.46)

Table 6 (part 2 of 2): Crude and adjusted analyses of tetanus, diphtheria, and pertussis vaccination, by prenatal care provider type

Type of prenatal care provider	Obstetrician <i>n</i> = 56 717	Family physician n = 12 308	Midwife n = 14 372
Primary care contact			
Rostered to family physician	-	0.99 (0.94–1.04)	-
Primary care model‡			
Capitation*	-	-	-
Comprehensive care model	-	0.67 (0.53–0.85)	-
Missing	-	0.63 (0.56-0.70)	-
Other	-	1.09 (1.03-1.16)	-

Note: CI = confidence interval, RR = relative risk.

*Reference group.

†Early prenatal care visits occurred before 32 weeks of gestation.

‡Capitation-based models include teams of 6 or more physicians who are compensated primarily through capitation payments but also receive fee-for-service payments. Comprehensive care model physicians are compensated primarily through fee-for-service. Both capitation and comprehensive care model physicians are eligible for specific bonuses and premiums based on patient enrolment. Other patient enrolment models were grouped together owing to small cell sizes and include family health groups and specialized models (such as general practice– or focused practice–physicians with alternative funding plans).

\$Adjusted for birth group, maternal age, rurality, Canadian Index of Multiple Deprivation measures (i.e., ethnocultural composition, economic dependency, residential instability), use of assisted reproductive technology, early prenatal care visits, and parity.

people.^{35,37} Prior estimates of OHIP vaccination data compared with self-report of influenza vaccination using the Canadian Community Health Survey found a sensitivity of around 50% and a specificity of 95.7%–98%.^{54,55} On the other hand, evidence suggests that the COVaxON database from which we obtained SARS-CoV-2 vaccination data is more complete.⁵⁶ We were unable to assess some factors influencing vaccine receipt among pregnant people, such as individual- or community-level access to vaccination and severity of COVID-19 outbreaks locally. People who gave birth in January 2022 may have had poorer access to SARS-CoV-2 vaccines than those who gave birth in March, although data (Appendix 3) suggest minimal overall impact of time within our 3-month window. Additionally, some people may have received vaccination before becoming pregnant or before realizing they were pregnant. In addition, the CIMD data were created from the 2016 Census, which may not reflect the birthing person's actual residence or social position in 2019–2022, and there are known limitations of using area-level CIMD guintiles to make individual-level inferences.^{47,57} However, with our large data set and propensity score-matching methods, we were able to control for many known confounders.

Conclusion

Among pregnant people in Ontario in early 2022, 78.7% received 1 dose of SARS-CoV-2 vaccine and 74.2% received a second dose. Although this was a lower rate than among nonpregnant comparators, it was higher than the rate of either Tdap (32.6%) or influenza (27.7%) vaccination in the same year. Rates of Tdap vaccination rose considerably from 2019 to 2022. Influenza vaccination was higher in 2021 but returned to 2019 levels the following year. Vaccination rates were lower among pregnant people who were young, multiparous, or lived in rural or economically deprived areas. Pandemic urgency may have overcome a great deal of hesitancy about vaccinating against SARS-CoV-2 during pregnancy in 2022, but uptake of routinely recommended vaccines in pregnancy remains a challenge.

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Data sharing: The administrative data set for this study is held securely in coded form at ICES. While legal data-sharing agreements between these data stewards and data providers prohibit ICES from making the data set publicly available, access may be granted to those who meet prespecified criteria for confidential access. Readers are welcome to contact the research team for further information. Data queries to ICES can be directed to Data and Analytic Services at das@ices.on.ca.

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