1	Invasive Pneumococcal Disease in Calgary During the Sars-CoV-2 Pandemic 2020
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14	Abstract – 97 words
15	During the Sars-CoV-2 pandemic in 2020, many countries shut down schools and
16	businesses in an effort to slow transmission of the virus. As some businesses reopened,
17	increased public health protocols, mask wearing, hand sanitizer, and personal protective
18	equipment use remained. The shut down and public health restrictions for person-to-person
19	interaction, resulted in a decline of other transmissible diseases as well as Sars-CoV-2. In
20	Calgary, the incidence of invasive disease due to Streptococcus pneumoniae declined to much
21	lower levels than would be expected in early spring and remained low until December 2020.
22	This decline occurred despite no changes in vaccine use and uptake.

- 23 Introduction
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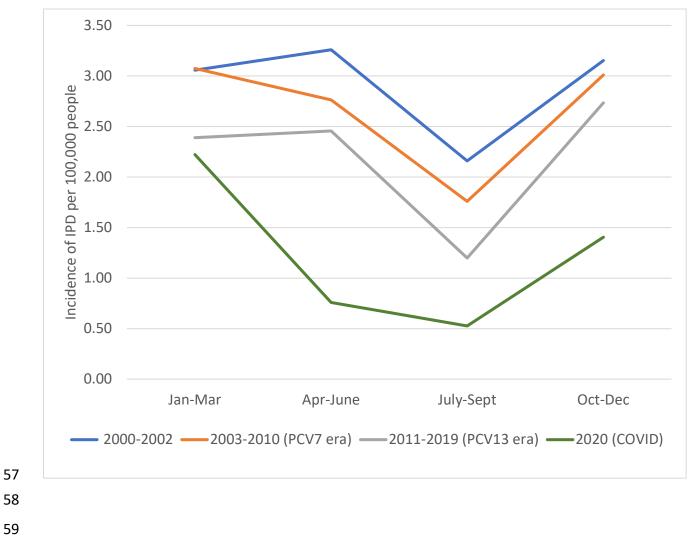
25 On March 11, 2020 the World Health Organization declared the Sars-CoV-2 outbreak to 26 be a pandemic (1). As cases spread globally, Alberta began to lock down by shutting down 27 schools on March 16, 2020 in an attempt to slow the spread of the novel Sars-CoV-2 virus. This 28 shut down of schools, followed by further lock down of businesses, and increased use of 29 personal protective equipment and hand sanitizer in public places, led to a decline in other 30 infectious diseases. We were interested in what affect the public health measures would have 31 on invasive pneumococcal disease (IPD) in Calgary during a time when there were no changes in 32 vaccination rates or vaccines offered. The 7-valent pneumococcal conjugate vaccine (PCV7) 33 was introduced in Alberta in 2002 and the 13-valent pneumococcal conjugate vaccine (PCV13) 34 was introduced in 2010 (2). Therefore, both vaccines were widely used by 2020. Normally, 35 with pneumococcal disease, some seasonal variation is present with the lowest incidence 36 occurring in summer and highest in winter months, similar to many infectious diseases.

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38 Methods

39 The Calgary Area Streptococcus pneumoniae Epidemiology Research team (CASPER) has 40 been collecting information on invasive pneumococcal disease in the Calgary Zone of Alberta 41 Health Services for over two decades (3-6). Our surveillance is reported through a central 42 laboratory system and is population-based. When the Calgary Laboratory services notifies us of 43 a case positive by culture for IPD, a research nurse conducts a chart review to collect clinical 44 information. We also receive microbial susceptibility and serotyping data. Serotyping is done through the Quellung reaction (7). All cases of IPD occurring in the Calgary Zone between 2000 45 46 and 2020 were included the analysis. Here we report our quarterly incidence of invasive 47 disease in the era of the Sars-CoV-2 pandemic in 2020 compared to quarterly incidence in the years pre and post conjugate vaccine introduction. Incidence was calculated using the number 48 49 of cases per 100,000 people. Population estimates for each year came from the Alberta 50 Interactive Health Data application (8). We took the average incidence per quarter for 2000-51 2002 to (the pre-vaccine era), 2003-2010 (PCV7 era), and 2011-2019 (PCV13 era) and 2020

- (Sars-CoV-2 pandemic) and graphed them to see how the average incidence per quarter in 2020
 compared to past eras. We also stratified by age groups (<18 and >=18 years old).
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- 55 **Figure 1.** Average Incidence per quarter of IPD for pre-vaccine (2000-2002), PCV7 era (2003-
- 56 2010), PCV13 era (2011-2019) and Sars-CoV-2 pandemic (2020), all ages.

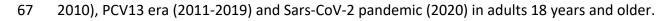


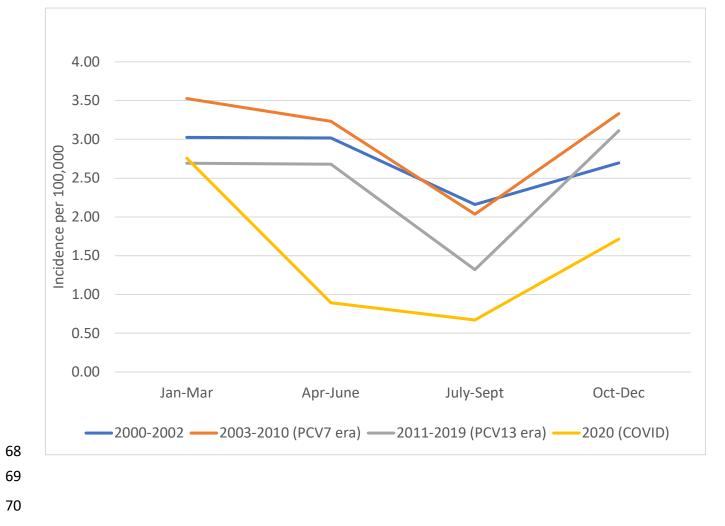
60 Figure 2a. Average Incidence per quarter of IPD for pre-vaccine (2000-2002), PCV7 era (2003-

61 2010), PCV13 era (2011-2019) and Sars-CoV-2 pandemic (2020) in children less than 18 years of

- 62 age. 5.00 4.50 4.00 3.50 3.00 3.00 2.50 2.00 1.50 1.00 0.50 0.00 Jan-Mar Apr-June July-Sept Oct-Dec 63 64
- 65

66 Figure 2b. Average Incidence per quarter of IPD for pre-vaccine (2000-2002), PCV7 era (2003-





71 Results and Discussion

72 In Calgary, January to March of 2020 began with normal post-PCV incidence rates of 73 invasive pneumococcal disease (IPD), followed by a sharp decline in incidence for guarters 2 74 and 3 in April to September. The incidence rate ratio for 2020 compared to 2019 incidence in 75 the first quarter was 1.2 (95% CI: 0.8 to 2.1; p=0.3675) for the second quarter the incidence rate 76 ratio was 0.22 (95% CI 0.1-0.4); p<0.001). Indicating that 2020 started off with normal incidence 77 in the first quarter, but declined to significantly lower incidence in 2020 compared to 2019 for 78 April to June. Incidence increased in the October to December quarter, but not to normal rates. 79 The incidence rate ratio for October to December 2020 compared to 2019 was 0.6 (95%CI: 0.3-80 1.0; p=0.0402). Incidence in children, which was already much lower in the vaccine era 81 compared to pre-vaccine, declined to almost nothing for 2020 and was low for the entire year, 82 with a total of 3 cases occurring in under 18-year-olds. The decline was seen in adults as well 83 but in adults it occurred in the second quarter of the year. This change in incidence occurred 84 within the context of normal, continued surveillance for pneumococcal disease and normal vaccine uptake within the community. Likely the decline was a result of the combination of 85 decreased transmission due to the closure of schools and businesses as well as other public 86 87 health measures. But there may have also been a reduction in presentation to hospital for 88 those with less severe IPD, as many people avoided hospitals during the beginning of the Sars-89 CoV-2 pandemic (9).

90 The public health efforts to mitigate the spread of Sars-CoV-2 also resulted in historically 91 low rates of influenza in multiple countries (10), including in Canada (11). A prospective cohort 92 study examining co-infection of pneumococcus and Sars-CoV-2 also reported declines in 93 pneumococcal disease following the initial lock down during March to June 2020 (12). Similar 94 reports came from Taiwan and Singapore (13, 14). Surveillance in the United Kingdom showed 95 large reductions in IPD in all age groups during March to June 2020 when lockdowns were 96 initiated (15). In Calgary, rates were normal in the first quarter of the year then dropped 97 substantially in the second quarter. The normal seasonal effect of IPD would have cases staying 98 high through April to June and then declining in the summer quarter, but in 2020 the decline 99 happened immediately after public health measures took effect in March. There was an

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increase again in the last quarter, but not to pre-COVID rates. This may be due to a combination
 of some businesses re-opening through the summer months when Sars-CoV-2 was more
 controlled, and due to the normal seasonal rise of invasive pneumococcal disease seen in the
 winter months.

Association of pneumococcal infection with Sars-CoV-2 is uncommon but not impossible from what has been described in the literature so far (15-20). There is an increasing number of reports to show this lack of association (15, 19, 21). In a systematic review of coinfection by Sars-CoV-2 and other respiratory pathogens, the results showed influenza A to be the most common pathogen to appear in co-infection with Sars-CoV-2 (16). However, Amin-Chowdhury et al. showed the odds of death to be over 7-fold higher in cases where co-infection with pneumococcus did occur, particularly in older adults (15).

We were limited in this analysis in that we only have population numbers for each year, not per quarter, so we had to assume the same population for every quarter of that year. We also have low numbers of pneumococcal disease within the Calgary region each year; therefore, we did not have the numbers to look more closely at children.

115 This ecological variation in IPD incidence is important in the vaccine era. The United 116 Kingdom modified their vaccine schedule to 1+1 doses in 2020; therefore, data will need to be 117 cautiously interpreted from these years of vaccine schedule change in the climate of the Sars-118 CoV-2 pandemic (22). This dramatic decline in incidence in a Calgary, where vaccination has not 119 changed, suggests that 2020 data will not be reliable in a place where schedules may have been 120 modified for 2020.

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