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Journal article

Predictors of herpes zoster vaccination among Australian adults aged 65 and over

Ricks, Thomas, Trent, Mallory J. and MacIntyre, C. Raina

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Abstract

Objective(s): To estimate HZ vaccine coverage in Australia among older Australians and to identify potential barriers to vaccination

Design: Analysis of data from three cross-sectional surveys administered online between 2019 and 2020

Setting and participants: Adults aged 65 and over residing in Australia

Main outcome measures: Self-reported herpes zoster vaccination

Results: Among the 744 adults aged 65 and over in this sample, 32% reported being vaccinated for HZ, including 23% of participants aged 65-74, 55% of participants aged 75-84, and 0% for participants aged 85 and above. Those who are vaccinated with other immunisations are more likely to have received HZ vaccine, including seasonal influenza (OR = 4.41, 95% CI: 2.44 - 7.98) and pneumococcal vaccines (OR = 4.43, 95% CI: 2.92 – 6.75). Participants with a history of certain conditions, such as stroke (OR = 2.26, 95% CI: 1.13 – 4.49), were more likely to be vaccinated against HZ. Participants that reported smoking tobacco daily were less likely to be vaccinated against HZ (OR = 0.48, 95% CI: 0.26 – 0.89). Participants were less likely to be vaccinated against HZ if they preferred to develop immunity 'naturally' (OR = 0.29, 95% CI: 0.15 – 0.57) or expressed distrust of vaccines (OR = 0.34, 95% CI: 0.13 – 0.91).

Conclusion(s): Further research is required to understand the barriers to HZ vaccine uptake. Increasing the funding eligibility for those who are at risk of complications from shingles, or lowering the age of eligibility, may increase vaccine coverage.

Predictors of herpes zoster vaccination among Australian adults aged 65 and over

Authors

Thomas Ricks

Biosecurity Program, The Kirby Institute, University of New South Wales, Sydney, Australia

Mallory J Trent

Biosecurity Program, The Kirby Institute, University of New South Wales, Sydney, Australia

C Raina MacIntyre

Biosecurity Program, The Kirby Institute, University of New South Wales, Sydney, Australia

Author for Correspondence:

Mallory J Trent, Biosecurity Program, The Kirby Institute, University of New South Wales, Sydney, Australia

Address: The Kirby Institute, Wallace Wurth Building Room 644, University of New South Wales, Kensington NSW 2052, Australia

Email: mallory.trent@unsw.edu.au; Tel: +61 4 3402 4017

1 Introduction

2 Herpes zoster (HZ), also known as shingles, is an infectious disease caused by the reactivation of
3 varicella zoster virus (VZV), commonly known as chickenpox. After the initial stage of VZV resolves,
4 the virus lies latent in the dorsal root ganglia and can reactivate decades later causing painful, itchy
5 vesicular rash and blisters usually around the thoracic region of the body (1). Statistically, about 1 in 3
6 people who previously had chickenpox will develop shingles at some point in their lives (2).

7 Although most cases of herpes zoster resolve after 3 to 5 weeks, approximately 10-18% of cases result
8 in post-herpetic neuralgia (PHN), a condition where a person develops neuropathic pain and vesicular
9 rash that often results in hospitalisation (3, 4). Despite antiviral treatment, PHN can last for many years
10 after onset and heavily affects quality of life (5).

11 Although the majority of the population is at risk of VZV reactivation, over 70% of shingles cases occur
12 in adults aged 50 and over (6). The cause of VZV reactivation is not well understood, but is likely due
13 to immunosenescence, the progressive decline of the immune system after the age of 50 (7). Certain
14 health conditions increase the risk of developing HZ and post-zoster complications, such as human
15 immunodeficiency virus (HIV), immunosuppressive conditions, stroke, myocardial infarction, chronic
16 obstructive pulmonary disease (COPD), asthma, and emphysema (8-10). There has also been a
17 reported association with COVID-19 vaccines and reactivation of HZ (11, 12).

18 The Australian Immunisation Handbook (AIH), currently recommends HZ vaccination for people aged
19 60 and over, regardless of their history of VZV infection (13) HZ vaccine was added to the National
20 Immunisation Program (NIP) in 2016, which provides funding for all adults 70 years of age, as well as
21 a catch up program for those aged 71-79 until 2023 (14). In Australia, the only available and registered
22 HZ vaccines are the live-attenuated vaccine, Zostavax (Oka strain) and the inactivated, adjuvanted
23 recombinant vaccine, Shingrix (15, 16).

24 A recent study by Lin et al (2020) concluded that only 47% coverage was achieved for people aged 70-
25 79 years old in Australia between 2016 and 2018. They compared their data extracted from the
26 Australian Immunisation Registry (AIR) with the distribution of vaccine and concluded that the HZ
27 vaccine coverage rate might be underestimated due to the uneven distribution of vaccines (17).
28 However, this is considerably lower than coverage for other adult vaccines on the National
29 Immunisation Program such as influenza and pneumococcal vaccines. The estimates of influenza and
30 pneumococcal vaccine coverage in adults aged 65 and over are reported to be 82% and 56%,
31 respectively, in 2015 in Australia (18).

32 Given the aging population and increasing chronic disease prevalence, the burden of HZ is likely to
33 increase over time. Therefore, in this study we aim to estimate the current HZ vaccine coverage rate
34 among older Australians and identify potential barriers to vaccination.

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39 Methods

40 This paper reports data on HZ vaccination collected from three different cross-sectional surveys: (1)
41 survey on adult immunisation (AIS) in Australia in October of 2019; (2) phase 1 of a study on 'Bushfire,
42 Respiratory protection, Emphysema, bronchitis, and Asthma Triggering Health Effects' (BREATHE);
43 and (3) a survey on mask use survey (MUS) during the COVID-19 pandemic in Australia, the United
44 States, and the United Kingdom. This analysis includes participants that are over the age of 65. All
45 three surveys were reviewed and approved by the University of New South Wales Human Research
46 Ethics Committee (HC #190617, #200460, and # 200477).

47

48 Data collection and participant selection

49 Out data was collected via three cross-sectional surveys administered by two consumer research
50 panels, Lucid (<https://luc.id/>) and Dynata (19). All participants gave informed consent prior to
51 beginning a survey. Failure to complete the survey was considered revocation of consent and data
52 was not used.

53 Survey 1 was the Australian Immunisation Survey (AIS). Data was collected via Lucid (<https://luc.id/>)
54 in October 2019. Participants were included if they were aged 18 or over and residing in Australia. The
55 sample was stratified to be representative of Australia in terms of age, gender, and state or territory
56 of residence. The survey aimed to evaluate participants' vaccination status as well as the general
57 attitudes about immunisation. To understand participants attitude towards immunisation attitude,
58 we provided statements about vaccine attitude and asked the participants whether they agree or
59 disagree with the statements using 2- and 4-point Likert scales (20). The statements include "I don't
60 trust vaccines", "Myself or someone I know has had a bad experience with a vaccine", "I prefer to
61 develop immunity naturally, rather than take vaccines." "I am afraid of needles", and "I have
62 difficulties getting an appointment with my doctor to get vaccinated". Additional details on survey
63 methods have been published previously (21, 22).

64 Survey 2 collected data on health outcomes during the 2019-20 bushfire season in Australia.
65 Participants aged 18 and over from bushfire-prone postcodes were recruited from Dynata's consumer
66 research panel in July 2020. The survey specifically targeted 50% participants to be with chronic lung
67 diseases and 50% without. More detailed survey methods have been published elsewhere (23).

68 Survey 3 collected data on attitudes towards mask use and vaccination during the COVID-19 pandemic.
69 Participants were recruited from Dynata's consumer research panel. The survey was distributed the
70 survey to a random sample of adults in Melbourne and Sydney via email. Detailed survey methods can
71 be found elsewhere (23).

72 All three surveys used the same question regarding HZ vaccination. We asked the question: "have you
73 ever received a vaccine for shingles (zoster)", with response options "yes", "no", and "not sure". We
74 combined the "no" and "not sure" responses into one category during data analysis.

75 All three surveys included questions about participants demographic characteristics such as their age,
76 gender, country of birth, education level and health insurance status. Questions on their health history
77 were also asked in all three surveys. These questions include whether the participants have self-
78 reported heart disease, lung disease, cancer or history of cancer, stroke, weak immunity, and whether
79 they are regular smokers of tobacco. The details of these questions can be found in the supplementary
80 tables.

81

82 **Data Analysis**

83 Descriptive statistics were calculated for all characteristics of interest, such as gender, age, education
84 level, country of birth, private health insurance status, HZ vaccination status, as well as vaccination
85 attitudes for the AIS Survey only. To account for differences in the sampling frames for the three
86 surveys, sampling weights were calculated using data for age group, gender, and the prevalence of
87 chronic lung disease using data from the Australian Bureau of Statistics (24). These were used to
88 calculate weighted estimates of demographic characteristics of the combined sample, which have
89 been published previously (25).

90 In the unweighted sample, we used Pearson's chi-square test to determine whether any
91 sociodemographic characteristics were associated with self-reported HZ vaccine status. In the
92 weighted sample, we used Pearson's chi-square test with the Rao and Scott correction for survey data
93 (26). We also used Pearson's chi-square test to identify perceived barriers associated with self-
94 reported HZ vaccination in the AIS (survey 1) data. Data were analysed using STATA version 16 IE by
95 StataCorp LLC (TX, USA).

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98 Results

99 A total of 3,162 participants responded to the three surveys and 744 (23%) participants were aged 65
100 and over and thus included in this analysis, including 339 (28%) from survey 1, 285 (17%) from the
101 survey 2, and 120 (23%) from survey 3. Weighted and unweighted demographic characteristics for this
102 study sample have been published previously (25).

103 For participants aged 65 and above, the weighted estimate for HZ vaccine coverage is 32%, ranging
104 between 28% in the survey 3 and 32% in survey 1. Estimates of HZ vaccine coverage by age are: 23%
105 of participants aged 65-74; 55% of participants aged 75-84; and 0% for participants aged 85 and above.

106 Rates of HZ vaccination were higher among those with a history of stroke ($p=0.05$) or cancer ($p=0.003$),
107 but there was no association with cardiovascular disease ($p=0.21$), chronic lung disease ($p=0.2$),
108 immunocompromising conditions ($p=0.32$), or smoking status ($p=0.14$). Participants that reported
109 smoking tobacco daily were less likely to be vaccinated against HZ ($p=0.02$). We did not observe any
110 association between HZ vaccination and gender ($p=0.21$), being born overseas ($p=0.1$), education level
111 (0.65), or having private health insurance ($p=0.43$). The HZ vaccination rate was higher among those
112 who received influenza and pneumococcal vaccines with high significance value ($p<0.001$).

113 Perceived barriers that were significantly associated with self-reported HZ vaccination included
114 distrust of vaccines in general ($p<0.001$) and preferring to develop immunity naturally ($p<0.001$). We
115 did not observe any association between HZ vaccination and any of the following perceived barriers:
116 being afraid of needles ($p=0.63$), having difficulties getting an appointment with their doctor ($p=0.92$),
117 having a bad experience with a vaccine (0.93), having mobility issues that make it difficult to visit GP
118 ($p=0.25$), or not having time to visit the GP to get vaccinated ($p=0.06$).

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121 Discussion

122 After HZ vaccine was added to the National Immunisation Program in 2016, the number of
123 prescriptions for HZ antivirals declined significantly among adults aged 70-79, suggesting that the
124 program is effective at decreasing the incidence of HZ (27). However, HZ vaccine uptake has been slow.
125 In Australia, the coverage of HZ vaccine between 2006 to 2013 was less than 2% for adults aged 60
126 and over (28). According to the NCIRS, in the first 17 months after the funded program was introduced,
127 HZ vaccine coverage was below 30% for adults aged 70-79 (29). In 2019-2020, we found that
128 approximately 30% of adults aged 65 and over have received HZ. We estimated 55% coverage among
129 Australians aged 75-84, which was comparable to findings by Lin et al, who found there was 47%
130 coverage among adults aged 70-79 (17). HZ vaccine coverage in Australia is low compared to other
131 western countries with similar health systems such as the UK, which has coverage as high as 77%
132 among adults 76 years of age (30). Several factors identified in literature review may have been
133 contributing factors for low HZ vaccination coverage, including uneven distribution of the vaccine and
134 restricted funding of the program for people aged 70 and above only (29).

135 Surprisingly, we did not observe any association between self-reported HZ vaccination and perceived
136 logistical barriers to vaccination, such as having mobility issues, having difficulties getting an
137 appointment with a GP, or not having enough time to get vaccinated. However, there were several
138 participants (10%) that agreed or strongly agreed with the statement of "I don't trust vaccines", and
139 nearly 1 in 4 agreed with the statement "I prefer to develop immunity naturally". Unsurprisingly, both
140 attitudes were associated with lower rates of HZ vaccination. This could possibly be a result of
141 increased social media and internet influence in the past two decades contributing to health decision-
142 making (31).

143 The need to increase HZ vaccination uptake has never been greater. There has been a significant
144 increase in HZ incidence for the last several decades. Between 2002 and 2012, the incidence of HZ in
145 Australia increased from 11.9 to 15.4 per 1,000 persons (32). Although it is often regarded as a mild
146 illness, HZ results in thousands of hospitalizations in Australia each year (33). In addition, treatment
147 and hospitalization for post-herpetic neuralgia results in significant economic burden (34). Given the
148 aging population of Australia and increasing rates of immunosuppression, the incidence of HZ is likely
149 to continue to increase without intervention. There is also emerging evidence that mRNA COVID-19
150 vaccines may trigger HZ, as well as the reactivation of human herpesvirus-6, -7 and Epstein-Barr virus
151 (11, 12). As of January 2020, more than 30 million doses of mRNA COVID-19 vaccines have been
152 administered in Australia (35).

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154 Limitations

155 This study was not without limitations. Firstly, this analysis used data from three separate studies that
156 used different sampling frames, meaning that the three study populations had key demographic
157 differences and are not necessarily representative of the general Australian population. Furthermore,
158 consumer research panels are not always representative of the general population, particularly for
159 older adults [48]. Another limitation is that survey 3 sampled only from the Sydney and Melbourne
160 metropolitan areas, so regional Australia is potentially underrepresented in this analysis. We utilized
161 survey weights in this analysis to mitigate potential biases due to the sampling frames used. Lastly,
162 the question regarding age was asked categorically for two out of three surveys, with response options:
163 "55-64", "65-74", "75-84", and "85 and over". Thus, we were unable to determine the coverage rate
164 among adults aged 60-69 or 70-79. This is notable, as these age ranges align with HZ vaccine
165 recommendations in Australia.

166

167 Conclusions

168 The overall HZ vaccination rate in Australia between 2019-2020 is suboptimal, despite the government
169 funded HZ vaccine program and the recommendation from Australian Technical Advisory Group on
170 Immunisation. We recognised that the government funded HZ vaccine program is only aim at people
171 aged 70-79 in Australia, however, compared to countries with similar health system such as the UK,
172 the coverage rate is still low. Further research is required to evaluate drivers and barriers to HZ
173 vaccination, including health access equity, socioeconomics, and awareness of HZ vaccine.

174

175 Declaration of Competing Interest

176 The authors have no competing interests to declare.

177

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Table 1. Demographic characteristics associated with self-reported herpes zoster vaccination

	Self-reported vaccination for herpes zoster							
	No/Not sure	Yes	Odds ratio (unweighted)			Odds ratio (weighted)		
	n (%)	n (%)	OR	95% CI	p-value	OR	95% CI	p-value
Age Group								
65-74	433 (78%)	125 (22%)	ref	-	-	-	-	-
75-84	80 (45%)	98 (55%)	4.24	(2.97, 6.06)	<0.01	4.12	(2.71, 6.23)	<0.01
85 or over	8 (100%)	0 (0%)	-	-	-	-	-	-
Gender								
Male	306 (68%)	142 (32%)						
Female	215 (73%)	81 (27%)	0.81	(0.59, 1.12)	0.21	0.98	(0.67, 1.42)	0.90
Country of Birth								
Overseas	139 (75%)	47 (25%)						
Australia	380 (68%)	176 (32%)	1.36	(0.94, 1.98)	0.11	1.09	(0.69, 1.70)	0.72
Education Level								
High school or less	195 (71%)	79 (29%)	ref	-	-	ref	-	-
Technical diploma	188 (70%)	79 (30%)	1.04	(0.72, 1.50)	0.85	1.01	(0.65, 1.57)	0.96
Tertiary degree	134 (67%)	65 (33%)	1.19	(0.80, 1.76)	0.40	1.19	(0.74, 1.90)	0.47
Private Health Insurance Status								
No	242 (71%)	97 (29%)						
Yes	277 (69%)	126 (31%)	1.14	(0.83, 1.56)	0.43	1.14	(0.78, 1.65)	0.51
Heart Disease								
No	432 (71%)	177 (29%)						
Yes	87 (65%)	46 (35%)	1.35	(0.90, 2.03)	0.14	1.14	(0.70, 1.88)	0.59
Lung Disease								
No	369 (71%)	148 (29%)						
Yes	150 (67%)	76 (33%)	1.32	(0.94, 1.84)	0.11	1.25	(0.84, 1.88)	0.27
Immunocompromising condition								
No	500 (70%)	218 (30%)						
Yes	19 (79%)	5 (21%)	0.46	(0.13, 1.61)	0.22	0.56	(0.13, 2.37)	0.43
Stroke								
No	494 (71%)	204 (29%)						
Yes	25 (57%)	19 (43%)	1.86	(1.00, 3.46)	0.05	2.26	(1.13, 4.49)	0.02
Cancer (past or current)								
No	429 (72%)	163 (28%)						
Yes	90 (60%)	60 (40%)	1.74	(1.20, 2.52)	<0.01	1.29	(0.81, 2.05)	0.27
Smokes tobacco daily								
No	449 (69%)	205 (31%)						
Yes	70 (80%)	17 (20%)	0.53	(0.31, 0.93)	0.03	0.48	(0.26, 0.89)	0.02
Influenza Vaccine								
No/Not Sure	130 (88%)	17 (12%)						
Yes	389 (65%)	206 (35%)	4.41	(2.47, 6.81)	<0.01	4.41	(2.44, 7.98)	<0.01
Pneumococcal Vaccine								
No/Not Sure	307 (85%)	56 (15%)						
Yes	212 (56%)	167 (44%)	4.31	(3.04, 6.11)	<0.01	4.43	(2.92, 6.75)	<0.01

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Table 2. Perceived barriers associated with self-reported herpes zoster vaccine status

Has received herpes zoster vaccine (self-reported)					
Odds Ratio					
Perceived barriers	No/Not sure (n=235), %	Yes (n=102), %	OR	95% CI	p-value†
I do not trust vaccines					
Disagree/strongly disagree	205 (68%)	97 (32%)			
Agree/strongly agree	30 (86%)	5 (14%)	0.34	(0.13, 0.91)	0.03
I prefer to develop immunity naturally					
Disagree/strongly disagree	166 (65%)	91 (35%)			
Agree/strongly agree	69 (86%)	11 (14%)	0.29	(0.15, 0.57)	0.001
I am afraid of needles					
Disagree/strongly disagree	193 (70%)	86 (31%)			
Agree/strongly agree	42 (62%)	16 (28%)	0.86	(0.46, 1.62)	0.65
I have difficulties getting an appointment with my doctor to get vaccinated					
Disagree/strongly disagree	230 (70%)	100 (30%)			
Agree/strongly agree	5 (61%)	2 (29%)	0.93	(0.18, 4.86)	0.92
Myself or someone I know has had a bad experience with a vaccine					
Disagree	188 (70%)	82 (30%)			
Agree	47 (70%)	20 (30%)	0.99	(0.55, 1.77)	0.93
I have mobility issues that make it difficult to visit my GP					
Disagree	218 (69%)	98 (31%)			
Agree	17 (81%)	4 (19%)	0.53	(0.17, 1.61)	0.26
I do not have time to visit my GP to get vaccinated					
Disagree	227 (69%)	102 (31%)			
Agree	8 (100%)	0 (0%)	-	-	-

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CRedit author statement

Mallory J Trent: Conceptualization, Methodology, Formal analysis, Investigation, Writing - Editing, Visualization, Supervision

Thomas Ricks: Formal analysis, Writing – Original Draft

C Raina MacIntyre: Conceptualization, Methodology, Writing - Review & Editing, Investigation, Supervision, Funding acquisition