

Adolescent vaccination

Australian Vaccinology Course

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Presentation overview

- Beyond the childhood schedule – why adolescents?
 - Risk, immunology, Australia's schedule
 - What do adolescents think about vaccines?
- HPV vaccines primer
- Meningococcal vaccines primer
- Safety of adolescent vaccinations
- Where are adolescents vaccinated in Australia?
- Coverage – where to now?
- Resources to support adolescent vaccination conversations
- Emerging research in adolescent vaccination



Image from

https://www.health.wa.gov.au/Articles/A_E/campaign-adolescent-immunisation

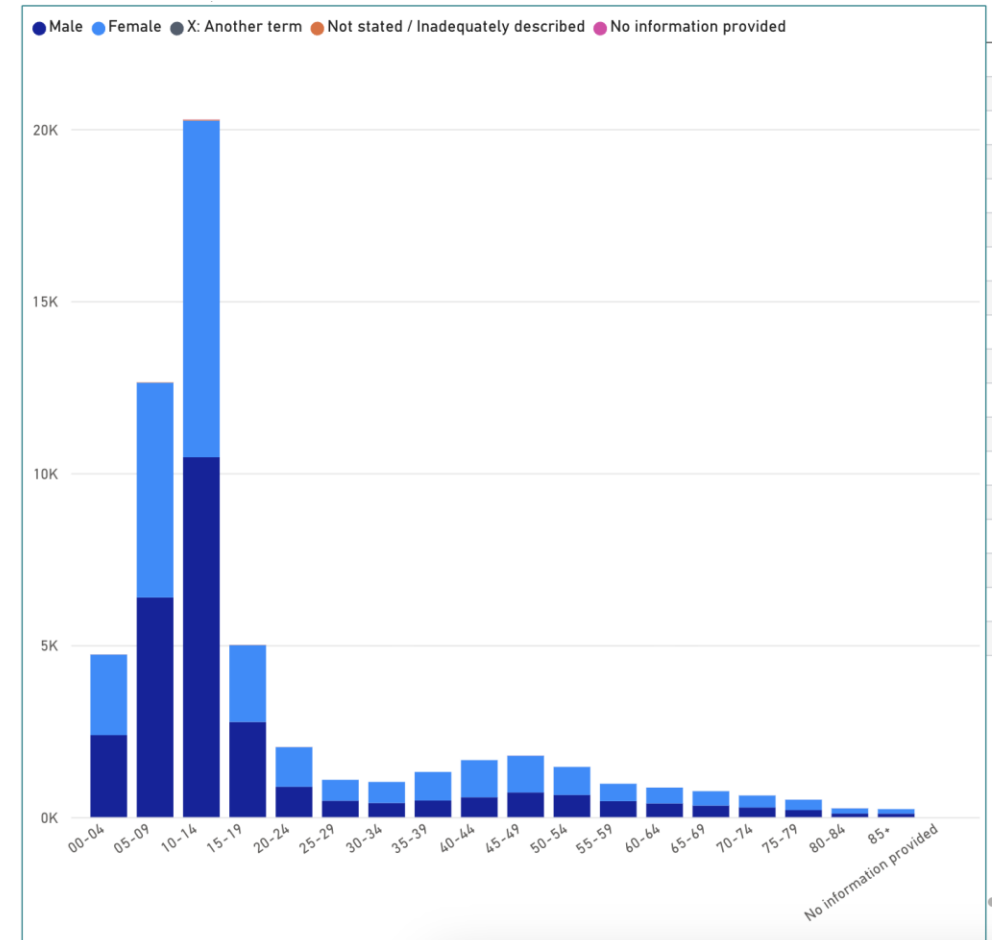


Beyond the childhood schedule – why adolescents?

- Waning of protection from some childhood immunisations for diseases that occur throughout life (eg pertussis, tetanus)
- Important for individual protection but also for herd protection eg of infants, elderly
- New exposures create new risk of other vaccine preventable diseases (eg HPV, meningococcal disease)



Pertussis notifications, 2024, Australia



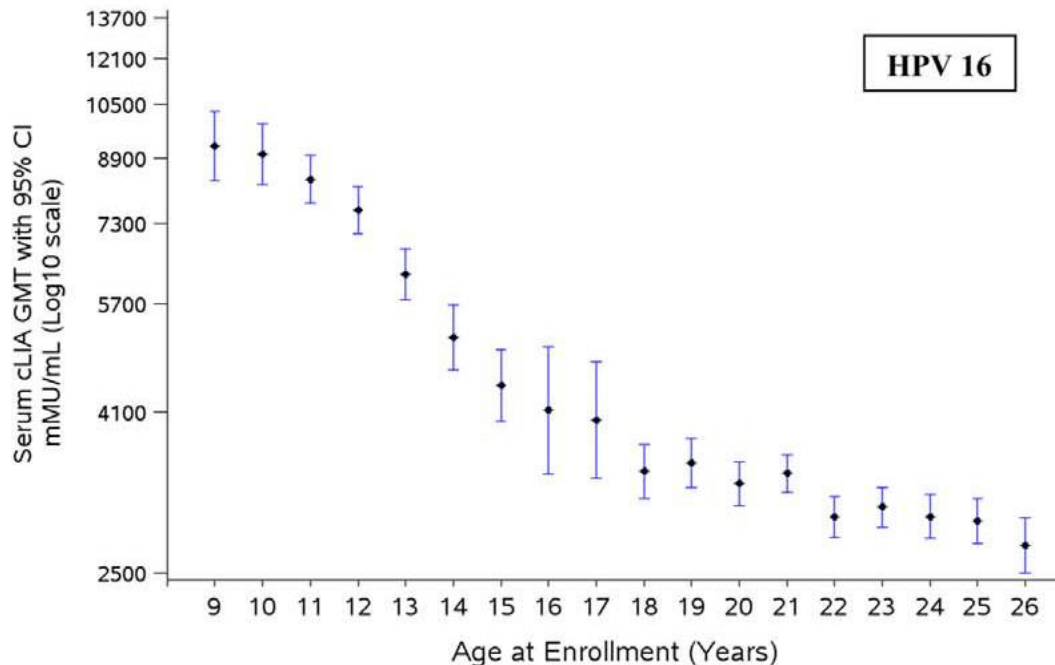
Source: NNDS, downloaded 24/8/2025

<https://nindss.health.gov.au/pbi-dashboard/>



Robust responses to immunisation in adolescence

9vHPV vaccine (primary) – GMT lower with age



Petersen LK, et al. Impact of baseline covariates on the immunogenicity of the 9-valent HPV vaccine - A combined analysis of five phase III clinical trials. *Papillomavirus Res.* 2017 Jun;3:105-115. doi: 10.1016/j.pvr.2017.03.002.

Adolescent vaccination

Pertussis responses (boosting)

- In a study (UK, Finland, Netherlands) of responses to an acellular pertussis booster in children (7-10 yrs) , adolescents (11-15 yrs), young adults (20-34 yrs) and older adults (60-70yrs):
 - Geometric mean (GMTs) were highest at day 28 in adolescents for:
 - Memory B cell frequencies for pertussis antigens
 - IgG antibody concentrations to pertussis toxin and filamentous haemagglutinin
 - Whereas, pertactin and IgA responses increased with age

Versteegen P et al. Memory B Cell Activation Induced by Pertussis Booster Vaccination in Four Age Groups of Three Countries. *Front Immunol.* 2022 May 23;13:864674; Versteegen P, et al. Responses to an acellular pertussis booster vaccination in children, adolescents, and young and older adults: A collaborative study in Finland, the Netherlands, and the United Kingdom. *EBioMedicine.* 2021 Mar;65:103247.

What do adolescents think about vaccines?



Systematic review of qualitative studies n= 59 (54 from HIC, 75% about HPV vaccine)

Adolescents' understanding of vaccines	Vaccine information sources described by adolescents	Vaccine decision-making
Purpose of vaccines: <ul style="list-style-type: none"> • Understand general purpose, but often unclear about specific vaccines • May be unsure whether vaccines preventative or therapeutic • Tend to think about vaccines in terms of personal rather than public protection 	Parents or caretakers: <ul style="list-style-type: none"> • Usually main source of information • Generally perceived by adolescents as being responsible for obtaining and evaluating vaccine information • Vaccines usually discussed at home at the time of vaccine consent 	Lead by parents or caretakers: <ul style="list-style-type: none"> • Parents generally required to provide vaccine consent • Adolescents may have pre-formulated opinions about vaccines, but final decision is generally parental, even when adolescent self-consent is an option
Vaccine preventable diseases: <ul style="list-style-type: none"> • Difficulties understanding and differentiating present and future risk • May be unfamiliar with diseases and disease transmission 	Schools: <ul style="list-style-type: none"> • Many adolescents wish for more vaccine information through schools • Vaccine information sheets not sufficient, adolescents wish for interactive sessions 	Lead by adolescents: <ul style="list-style-type: none"> • Self-consent often not offered in practice • Sub-groups of adolescents from marginalized and minority groups may exercise higher degree of autonomy when it comes to vaccine decisions
Concerns about vaccines: <ul style="list-style-type: none"> • Pain with vaccination • Lack of privacy during vaccination • Short and long-term side effects • Questionable effectiveness 	Healthcare providers: <ul style="list-style-type: none"> • Trusted information resource, but not all providers initiate vaccine conversations • Adolescents perceive providers address parents rather than adolescents when discussing vaccines 	
	Peers: <ul style="list-style-type: none"> • Not commonly described as key information source • Can be source of support during school vaccine clinics 	
	Media <ul style="list-style-type: none"> • Increasing use of internet and social media • Unclear how adolescents assess and triangulate media information • In-person communication remains important 	



National Immunisation Schedule

Adolescent vaccination

(also see vaccination for people with medical risk conditions)

Age	Diseases	Vaccine Brand
All ages	<ul style="list-style-type: none">● Influenza (adolescents with specified medical risk conditions)● Influenza (Aboriginal and Torres Strait Islander adolescents)● Pneumococcal (adolescents with specified medical risk conditions)	Age appropriate Age appropriate Prevenar 13® and Pneumovax 23®
12–13 years (Year 7 or age equivalent)	<ul style="list-style-type: none">● Human papillomavirus (HPV)● Diphtheria, tetanus, pertussis (whooping cough)	Gardasil®9 Boostrix® or Adacel®
14–16 years (Year 10 or age equivalent)	<ul style="list-style-type: none">● Meningococcal ACWY	MenQuadfi®

State and Territory additional scheduled adolescent vaccines



National Immunisation Program

February 2025 South Australia Schedule

Age	Antigen	Vaccine Brand	Route	Important Information
9-14 years Aboriginal**	23vPPV	Pneumovax 23®	IM	5-10 years after the previous dose of 23vPPV
Year 7 students	dTpa HPV	Boostrix® OR Adacel® Gardasil® 9	IM IM	1 dose of Gardasil 9 is recommended unless person is immunocompromised
Year 10 students	4vMenCV (ACWY) Meningococcal B	MenQuadfi™ Bexsero®	IM IM	2 doses Bexsero with a minimum 8 week interval



NT Immunisation Schedule

Children and Adolescents

Up to 19 years

July 2025

89228315

Immunisation program | NT Health

All people		Aboriginal and Torres Strait Islander people		Specified medical risk conditions	
12 years year 7 students	I N F L U E N Z A	Human papillomavirus (HPV)	Gardasil 9® (IM)	1	People who are immunocompromised Human papillomavirus (HPV) AIH
		Diphtheria, tetanus, pertussis	Boostrix® (IM) or Adacel® (IM)	1	
14 - 19 years	I N F L U E N Z A	Meningococcal ACWY	MenQuadfi® (IM)	2	Recommended to receive one dose of Men ACWY vaccine 14 years and over regardless of previous doses
		Meningococcal B	Bexsero® (IM)	1 & 2	Catch-up schedule for adolescents 15-19 years till 31 Dec 2026. 8 weeks between doses *
		Pneumococcal	Pneumovax 23® (IM)	2	See NT Pneumococcal schedule

*Adolescents 14 years and over are recommended to receive 2 doses of meningococcal B vaccine 8 weeks apart, regardless of previous doses of Men B vaccine received under 14 years of age

Immunisation Schedule Queensland - Adolescent and Adult Immunisation

Year 10 students (or age equivalent)	Meningococcal ACWY	MenQuadfi	R	IM / DL	Meningococcal ACWY is funded for Year 10 Students (or age equivalent) AND adolescents aged 15-19 years inclusive.
	Meningococcal B	Bexsero		IM / DL	2 doses given a minimum of 8 weeks apart. Meningococcal B is funded for Year 10 students (or age equivalent) AND adolescents aged 15-19 years inclusive.

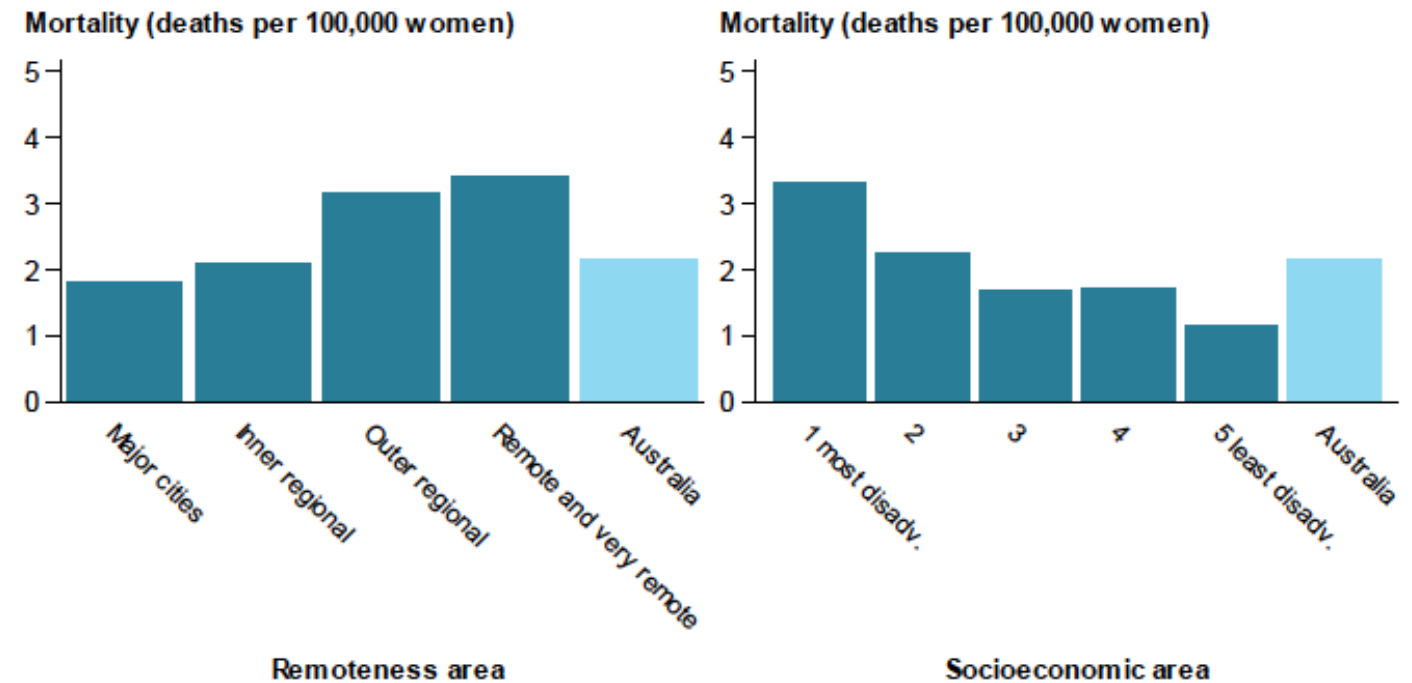


Cervical cancer is a disease of inequity

- 4th commonest cancer globally but 2nd most common in females 15-44 years
- Treatable if diagnosed early
- Late-stage disease associated with severe suffering, and often highly stigmatised and socially isolating
- Incidence and mortality, and their ratio, strongly correlated with human development index
- Inequities exists within Australia too, despite cervical cancer being *almost completely preventable*.

Bruni et al, *Lancet Glob Health* 2023 ;
Singh et al. *Int J MCH AIDS* 2012; AIHW 2024

Figure 3.20.2: Cervical cancer mortality, by remoteness area and socioeconomic area, women aged 25–74, 2018–2022

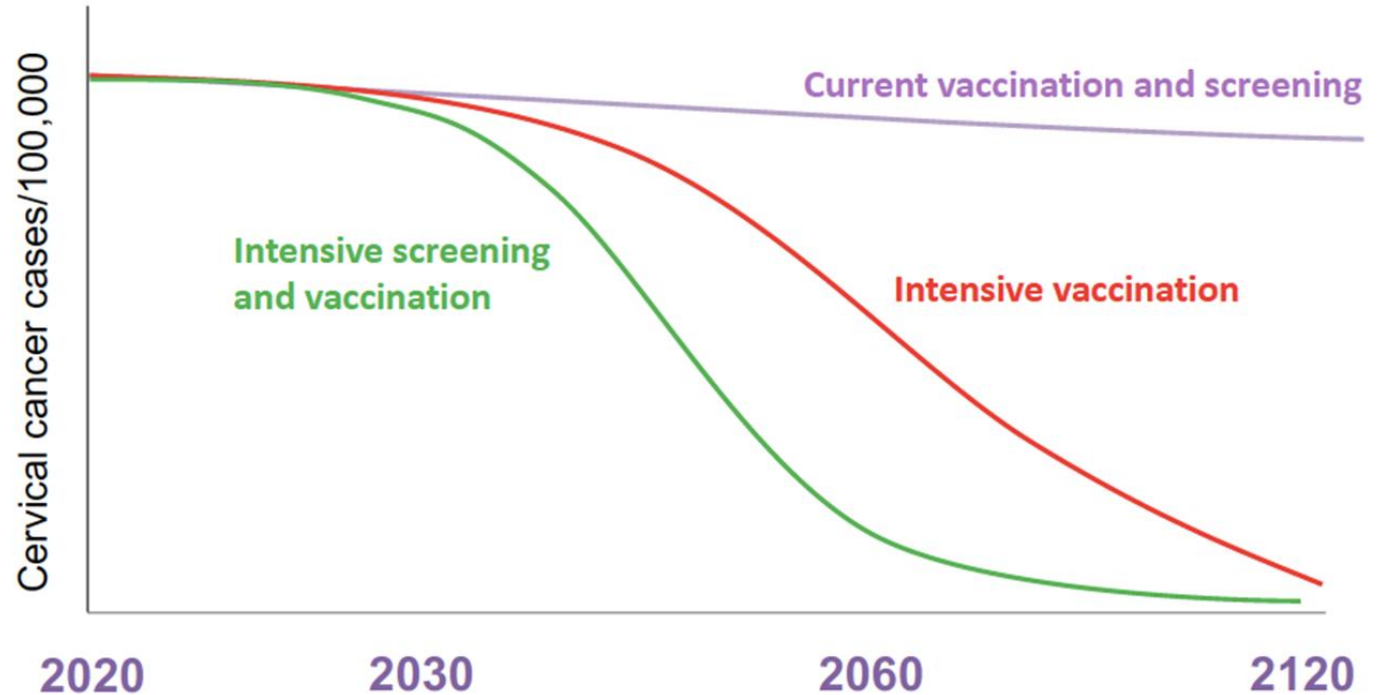




We are at an incredible moment in time...

- We understand the natural history of HPV and cervical cancer
- Primary prevention with HPV vaccines incredibly effective and safe (19+ years of rigorous global data)
- Secondary prevention with HPV screening far more effective than previous methods
- Even once or twice in a lifetime screening greatly reduces risk

CERVICAL CANCER ELIMINATION: CONCEPTUAL FRAMEWORK

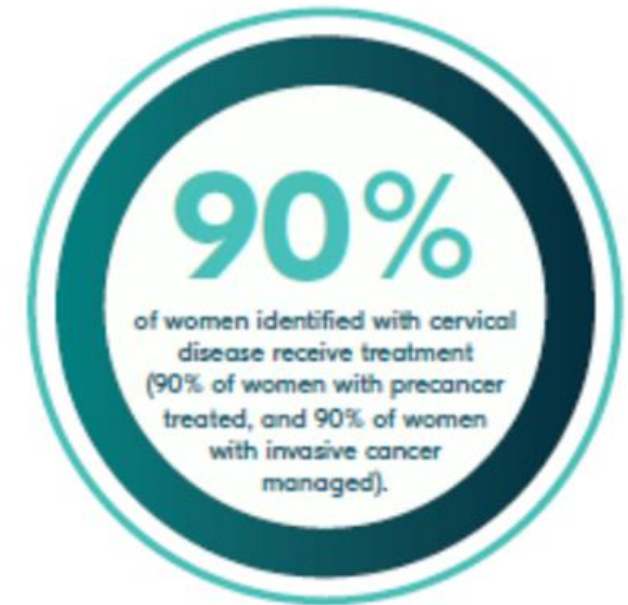


The Global Effort to Eliminate Cervical Cancer



This global strategy to eliminate cervical cancer proposes:

- a vision of a world where cervical cancer is eliminated as a public health problem;
- a threshold of 4 per 100 000 women-years for elimination as a public health problem;
- the following 90-70-90 targets that must be met by 2030 for countries to be on the path towards cervical cancer elimination:



Predicted to avert

- >74 million cases
- >62 million cervical cancer deaths

A strategy focused on equity

An Australia where preventable cervical cancer is a disease of the past, in which Australia's diverse communities have equitable access to information and to culturally safe and inclusive vaccination, screening and treatment services.

This Strategy recognises the right to equitable prevention and care for all people, including but not limited to:

- Aboriginal and Torres Strait Islander people
- People from culturally and linguistically diverse backgrounds (including immigrants, refugees, and asylum seekers)
- People who identify as lesbian, gay, bisexual, transgender, queer, and/or asexual or who are intersex
- People with disability, and
- People living in remote and rural areas.

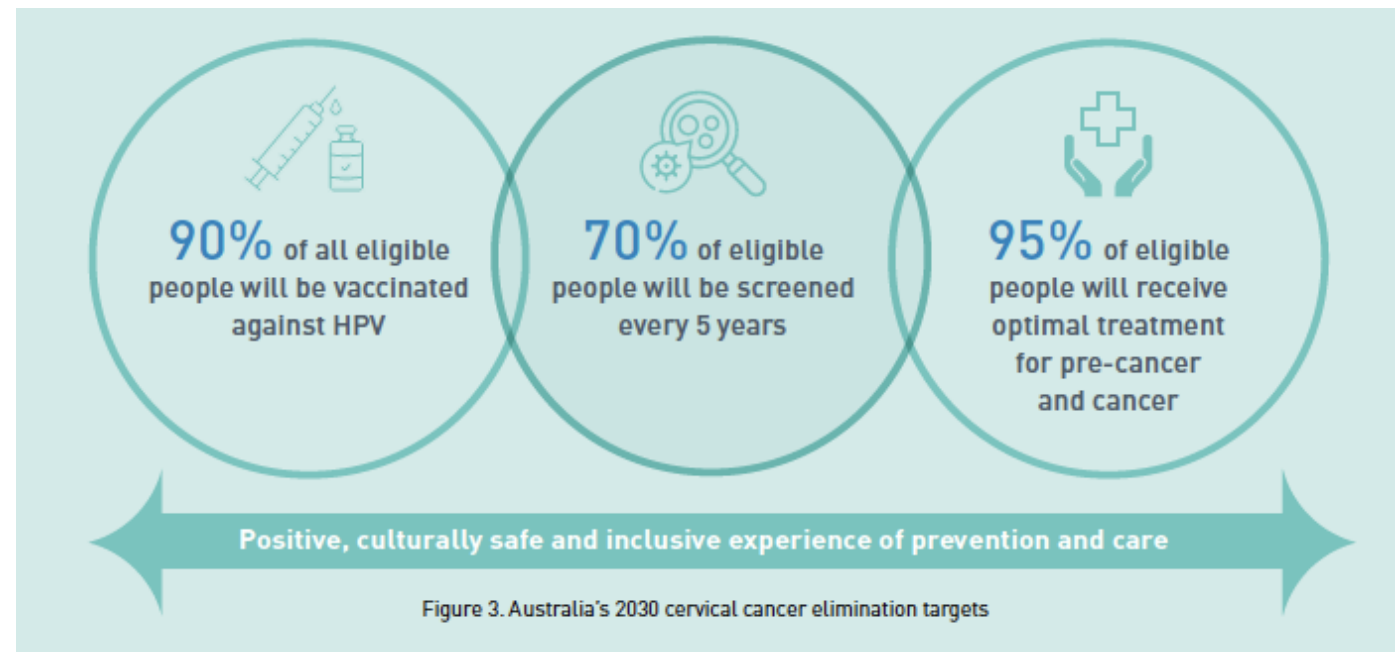
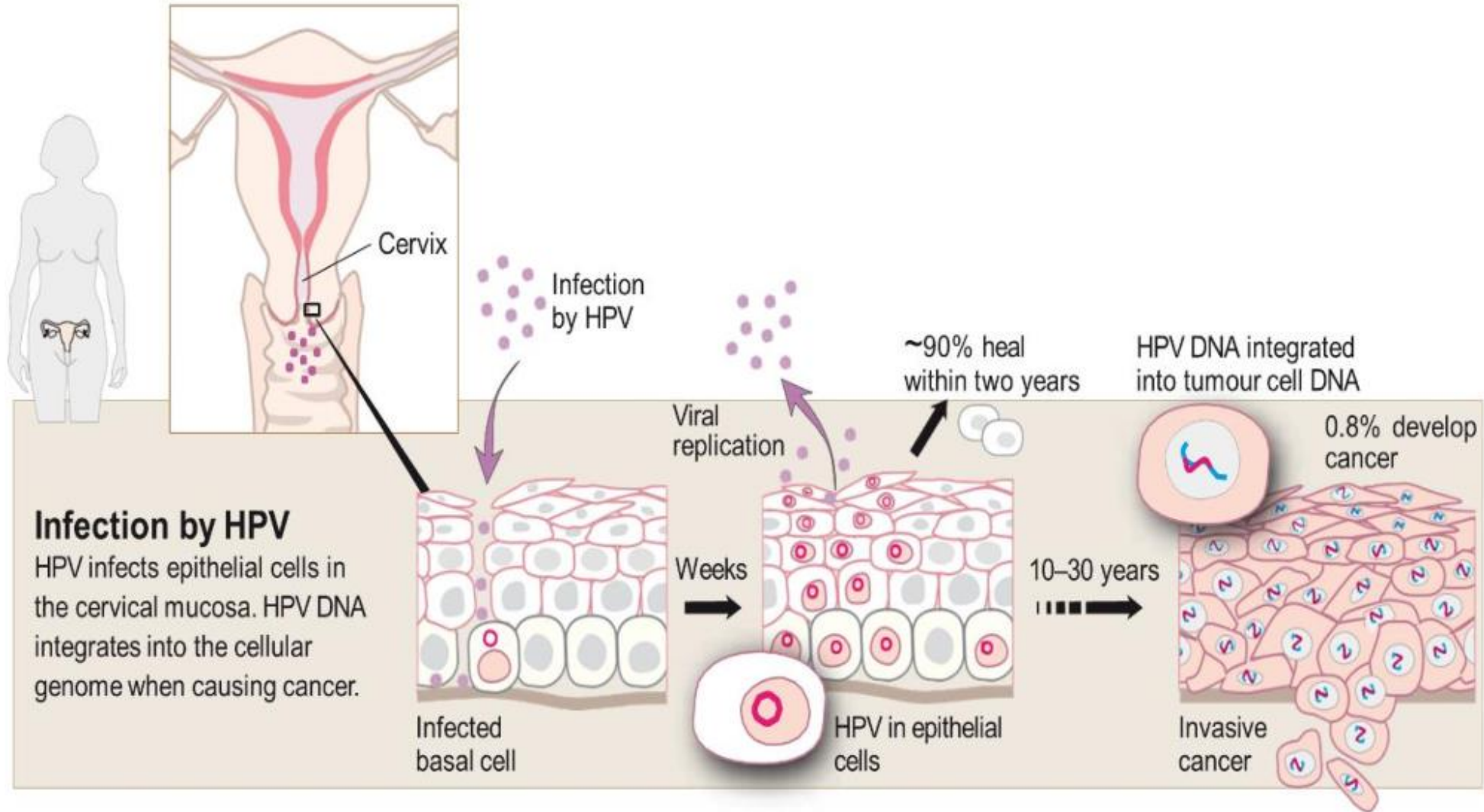


Figure 3. Australia's 2030 cervical cancer elimination targets



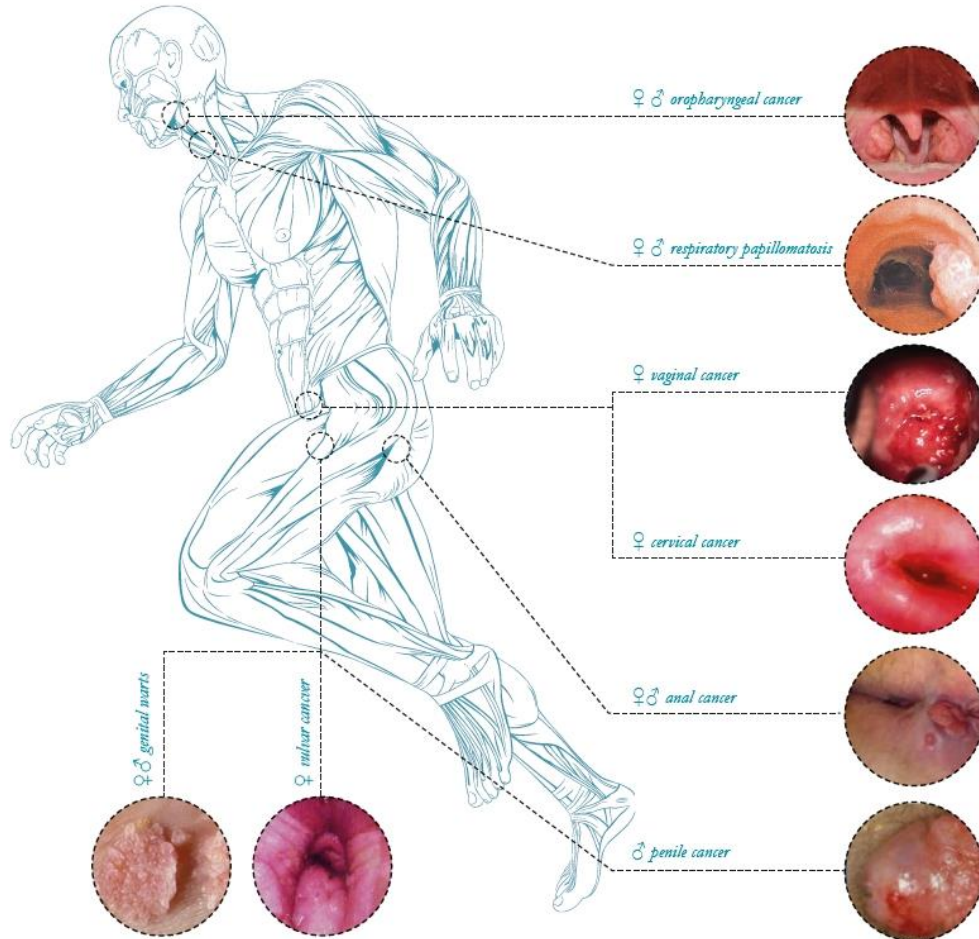


Pathogenesis of HPV in cervical cancer





Other HPV-related diseases



Annually, in women HPV infections cause 530,000 cancer cases in the cervix, 18,000 in the anus, 8,500 in the vulva, 12,000 in the vagina and 5,500 in the oropharynx. In men, HPV infections cause 17,000 cancer cases in the anus, 13,000 in the penis and 24,000 in the oropharynx. Ref. (1)

HPV attribution

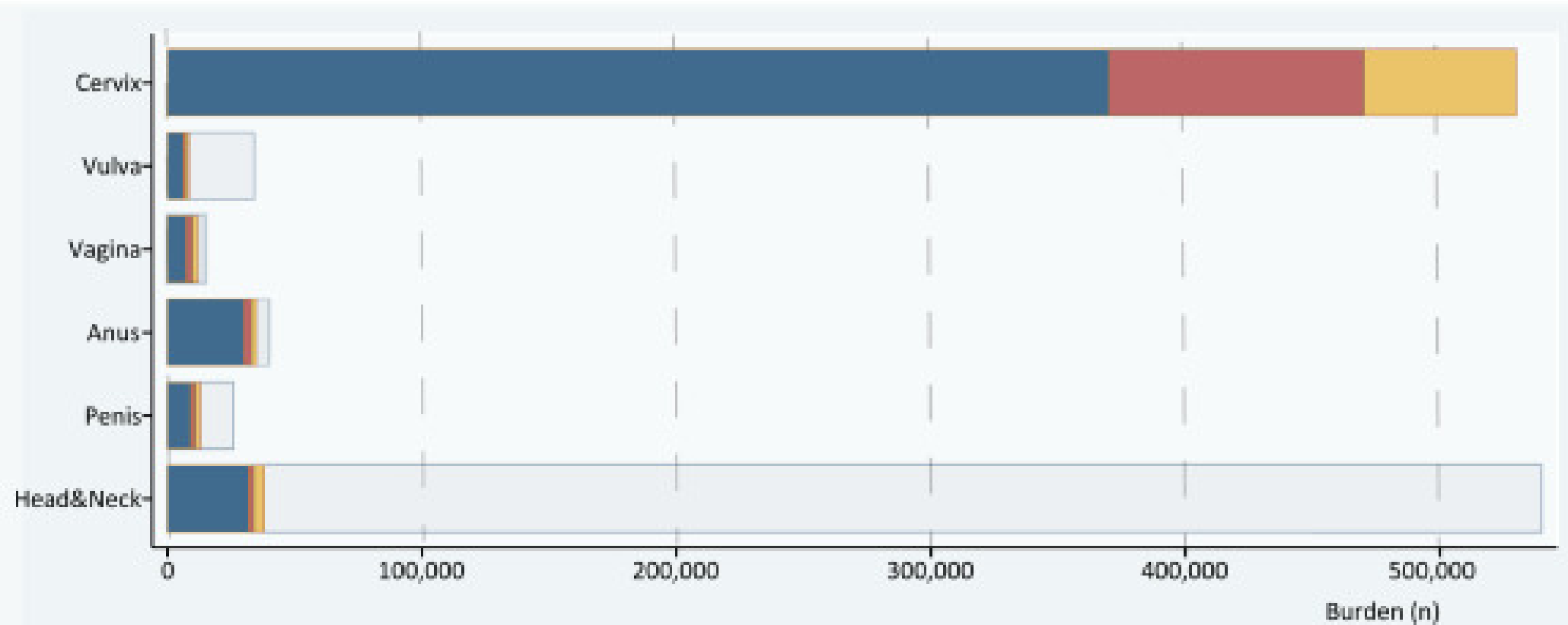
- Anogenital warts -100%
- Recurrent respiratory papillomatosis - 100%
- Anal cancer – 88%
- Vaginal cancer – 78%
- Penile cancer -50%
- Vulval cancer – 25%
- Oropharyngeal cancer – 30% globally. Up to 80% in some settings.

de Martel C, et al Worldwide burden of cancer attributable to HPV by site, country and HPV type. Int J Cancer 2017; 141(4): 664-70.

de Martel C, et al. Global burden of cancer attributable to infections in 2018: a worldwide incidence analysis. Lancet Glob Health 2020; 8(2): e180-e90.



Global HPV related cancer burden by HPV type



	Cervix	Vulva	Vagina	Anus	Penis	Head&Neck
HPV16/18	370,000	6,200	7,400	30,000	9,100	32,000
HPV31/33/45/52/58/6/11	100,000	1,200	2,500	3,000	1,900	2,000
Other HPV	60,000	1,100	2,100	2,000	2,000	4,000
No HPV	0	25,000	3,000	5,000	13,000	502,000



How do HPV vaccines work?

HPV vaccines *prevent* infection with targeted types by antibody binding to the virus, preventing it from entering the basal epithelial cell

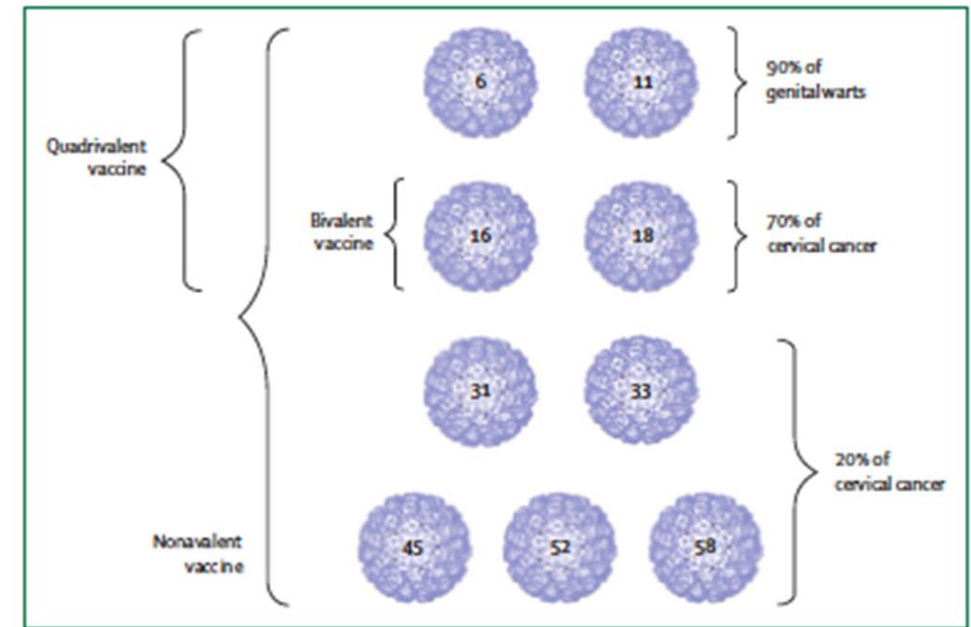
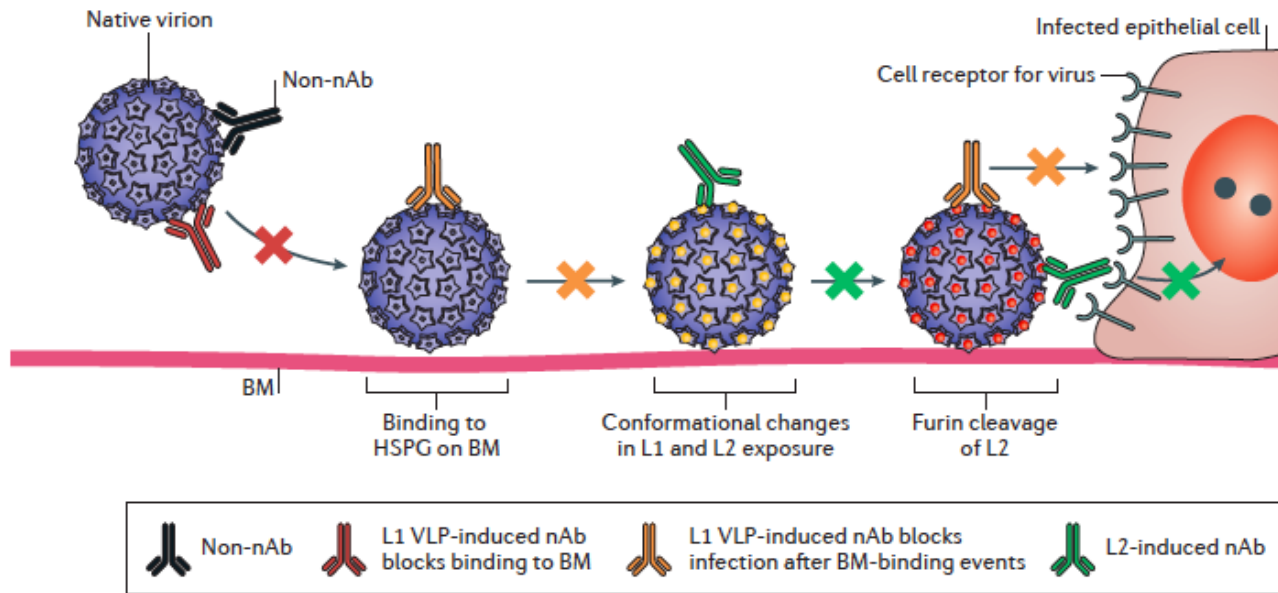


Figure 2: HPV VLP types in the nonavalent VLP vaccine
VLPs in the bivalent, quadrivalent, and the nonavalent vaccines are shown with the proportion of neoplastic disease attributed to each group. HPV=human papillomavirus. VLP=virus-like particle.

HPV vaccines are very effective

Initial RCTs - 94% +efficacy against persistent infection in those uninfected at baseline (2vHPV 94%, 4vHPV and 9vHPV 96%)

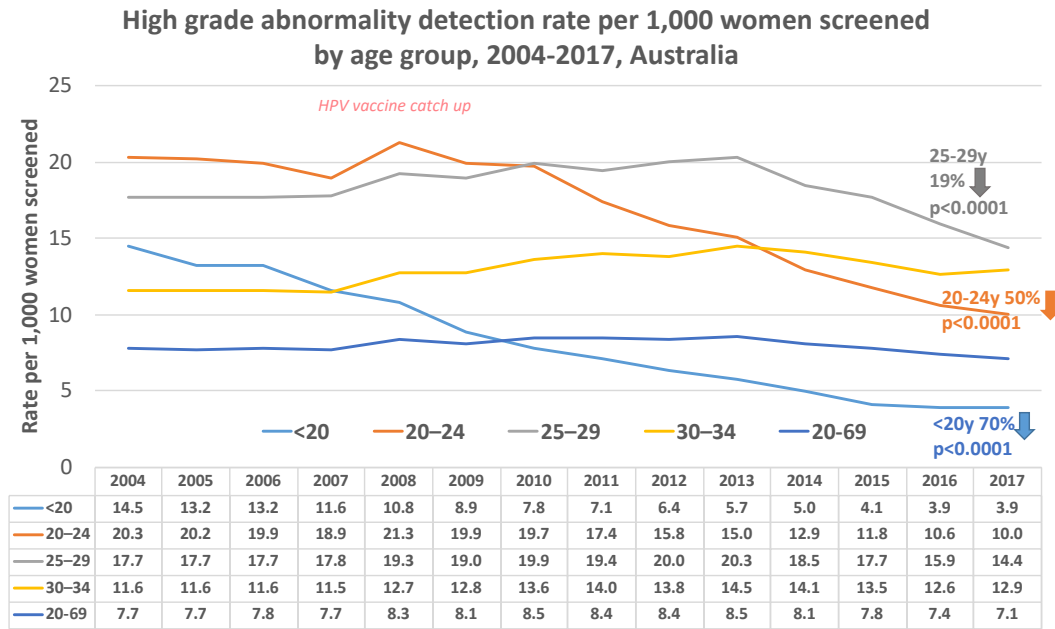
In Australia (and elsewhere) large declines seen in

- HPV infection
- Genital warts
- Cervical precancer
- Recurrent respiratory papillomatosis

Impact accelerated by multicohort vaccination at introduction

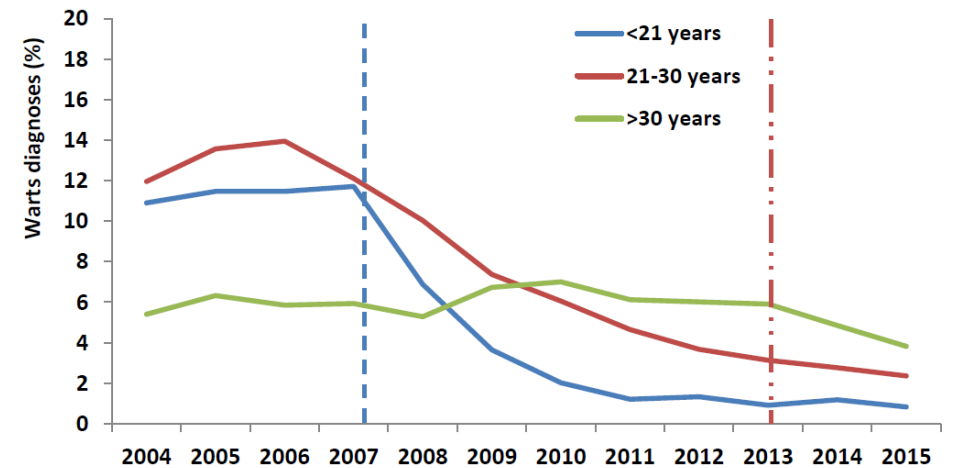
WHO recommends vaccination of 9-14 year old girls, with an initial multi-age cohort catch up at commencement

Harper Gyne Onc 2017; Patel et al EuroSurveillance 2018



Data source: AIHW 2020

Figure 1: Proportion of Australian born women diagnosed with genital warts at first visit, by age group, 2004-2015



* The first dotted line represents the start of the national HPV vaccination program for women in mid-2007 and the second dotted line represents the start of the national HPV vaccination program for boys in 2013

Source: Genital Warts Surveillance Network report 2016

Evidence of cervical cancer prevention



Lei et al. HPV Vaccination and the Risk of Invasive Cervical Cancer

N Engl J Med 2020;383:1340-8

->1.6 million females in Sweden

IRR vax <17 years

0.12 (95%CI 0.00-0.34)

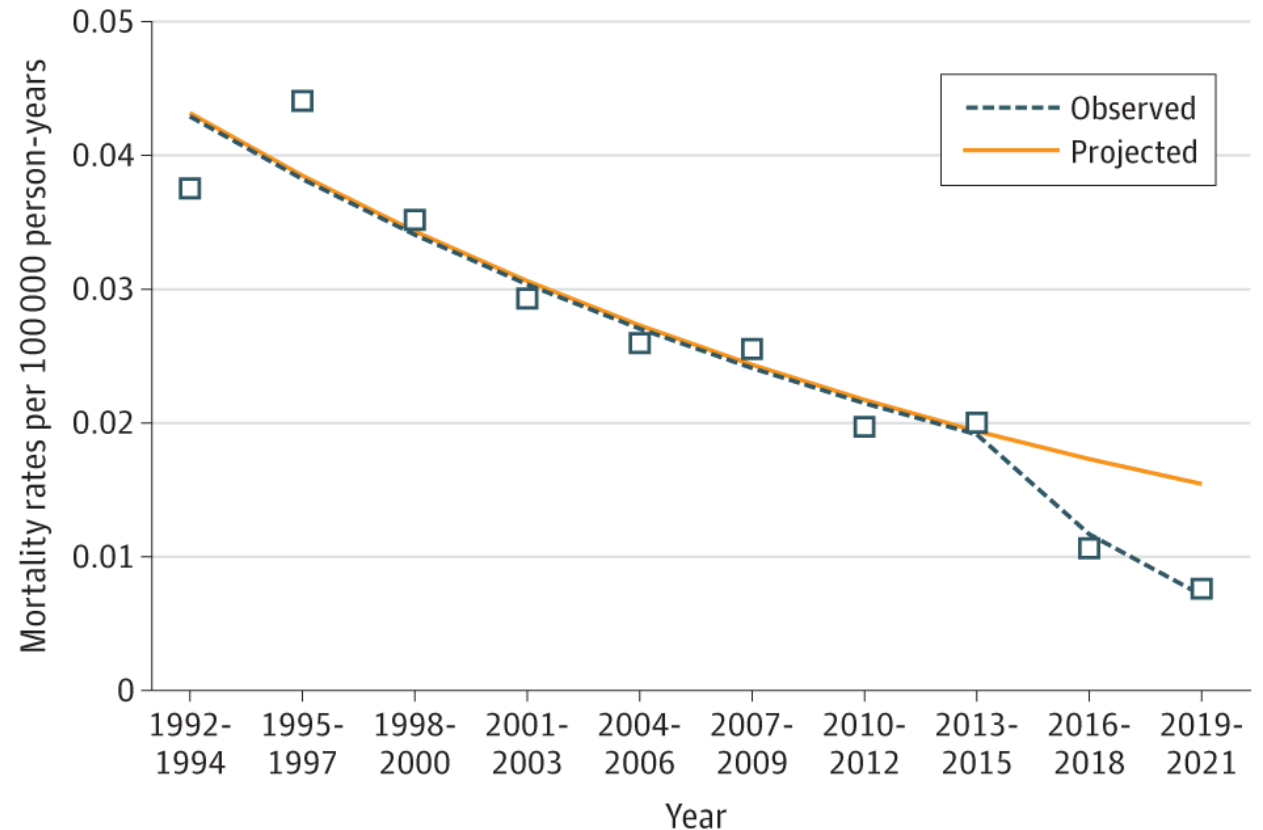
IRR vax 17-30 years

0.47 (95%CI 0.27-0.75)

ALSO SHOWN IN DENMARK, UK,
NETHERLANDS and USA

Recent US data shows significant decline in cervical cancer mortality in women under 25

A Cervical cancer mortality rates



Dorali P et al. Cervical Cancer Mortality Among US Women Younger Than 25 Years, 1992-2021. JAMA. 2025;333(2):165-166.

A single dose is enough...high and equal efficacy

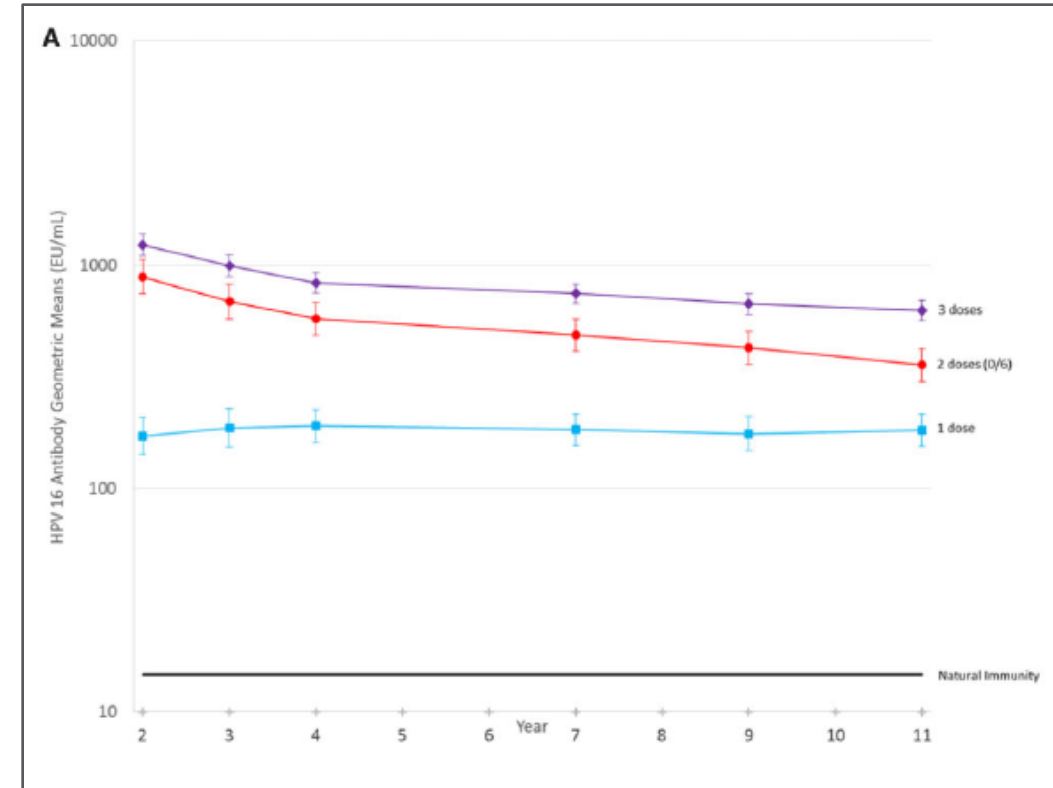


- Single dose HPV vaccination endorsed by WHO up to age 20 years (Dec 2022)
 - (2 doses for age 21+, 3 doses for immunocompromised)

See: Human papillomavirus vaccines: WHO position paper, December 2022. Weekly Epidemiological Record No 50, 2022, 97, 645–672

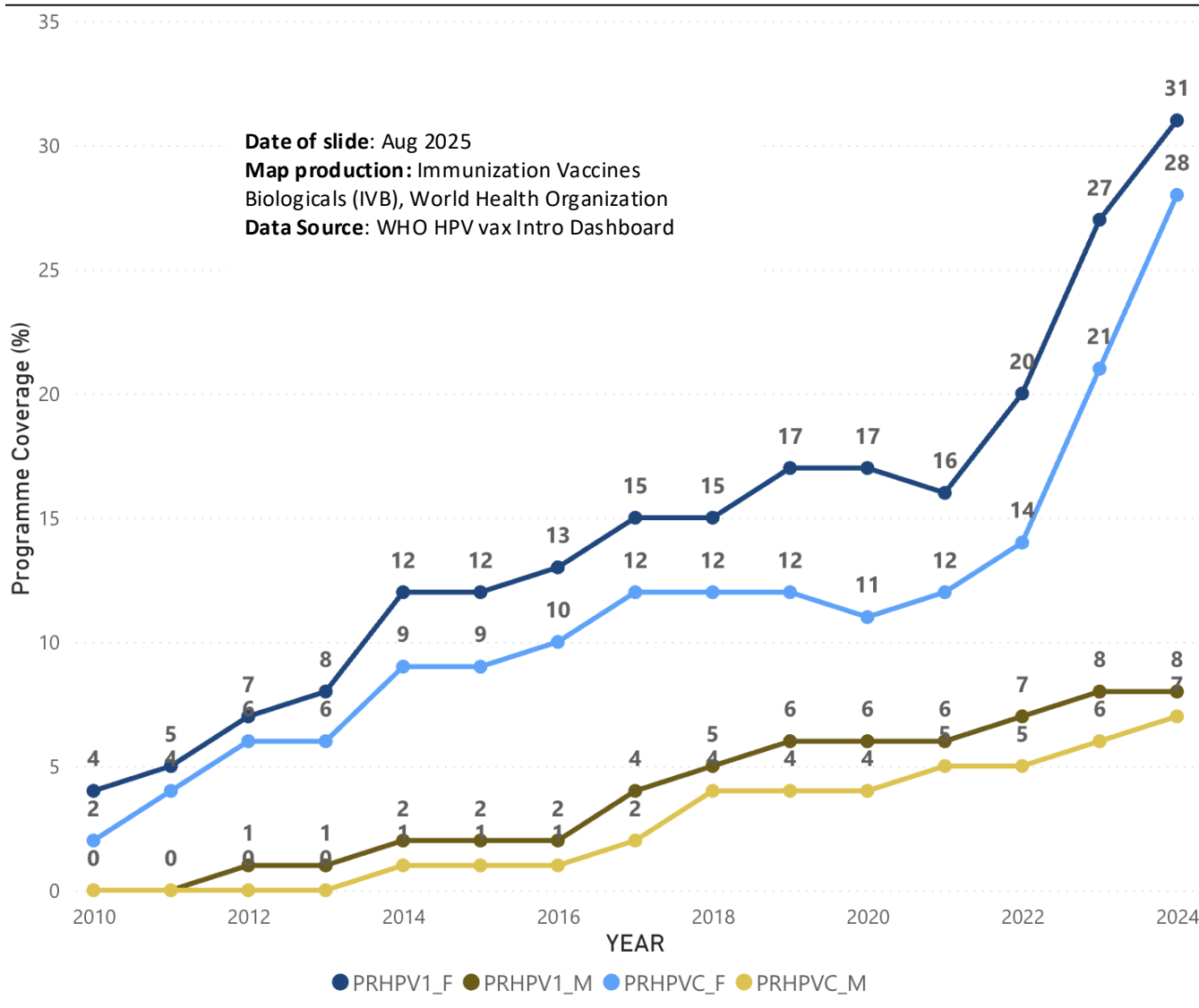
Key evidence

- Kenya RCT women 15-20 years, 2vHPV or 9vHPV vs meningococcal (*Barnabas et al, 2023*)
- 3 yr incident persistent HPV16/18 infection VE 98% 2vHPV, 99% 9vHPV; 16/18/31/33/45/52/58 infection 96% 9vHPV)
- India IARC study- girls aged 10-18 at vax 4vHPV (*Basu et al, 2021; Joshi et al 2023, Malvi et al 2025*) 10 yr incident persistent 16/18 infection (detected ≥ 10 months apart) VE **95.4%** (95% CI 85.0-99.9%)
- DoRIS trial 2vHPV, 9vHPV non-inferiority and immunobridging 10-14 yr olds, Tanzania (*Watson-Jones et al 2022, 2025; Baisley et al 2022, 2024*)



Sustained high level antibody after 1 dose 2vHPV vaccine and associated VE from Costa Rica trial (Kreimer et al JNCI 2020. See also Porras et al, JNCI Monograph 2024)

HPV vaccine global coverage



New introductions

2024 n=4

2023 n=13

2022 n=14

Support from GAVI and HAPPI consortium

- GAVI aims to reach 86 million girls by 2025

- HPV Vaccine Acceleration Program Partners Initiative (HAPPI) Consortium (BMGF funded) for LMICs

Need to optimise the use and distribution of the available supply

- Supply constraints now easing
- New vaccines entering market from India and China
- GLOBAL: 2vHPV (Cervarix), 4vHPV (Gardasil), 9vHPV (Gardasil9)
- CHINA: 2vHPV (Cecolin), 2vHPV (Walrinvax)
- INDIA: 4vHPV (Cervavac)
- 5 prequalified by WHO at Oct 24
 - 4 for use as single dose

150 Countries have HPV vaccine in national program



Date of slide: August 2025

Map production: Immunization Vaccines
Biologicals (IVB), World Health Organization

Data Source: WHO HPV vax Intro Dashboard

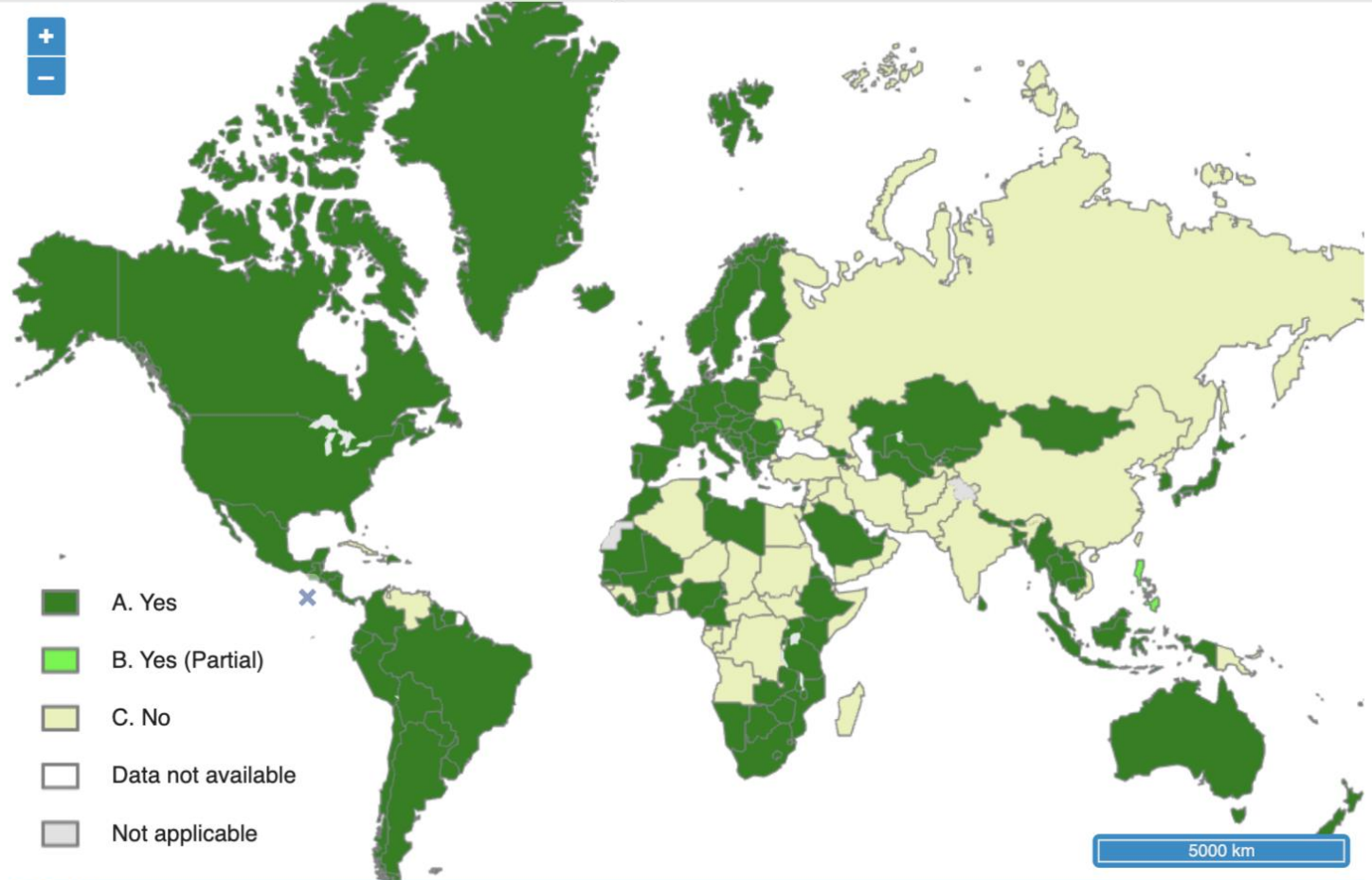
2030 Target: 194 countries

194
Total countries reported

HPV national schedule	No. of countries
A. Yes	148
B. Yes (Partial)	2
C. No	44

150 (77%)
44 (23%)

HPV vaccine included in national immunization programme



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Last update:

8/22/2025 10:12:27 AM

77 Countries have one dose in national program

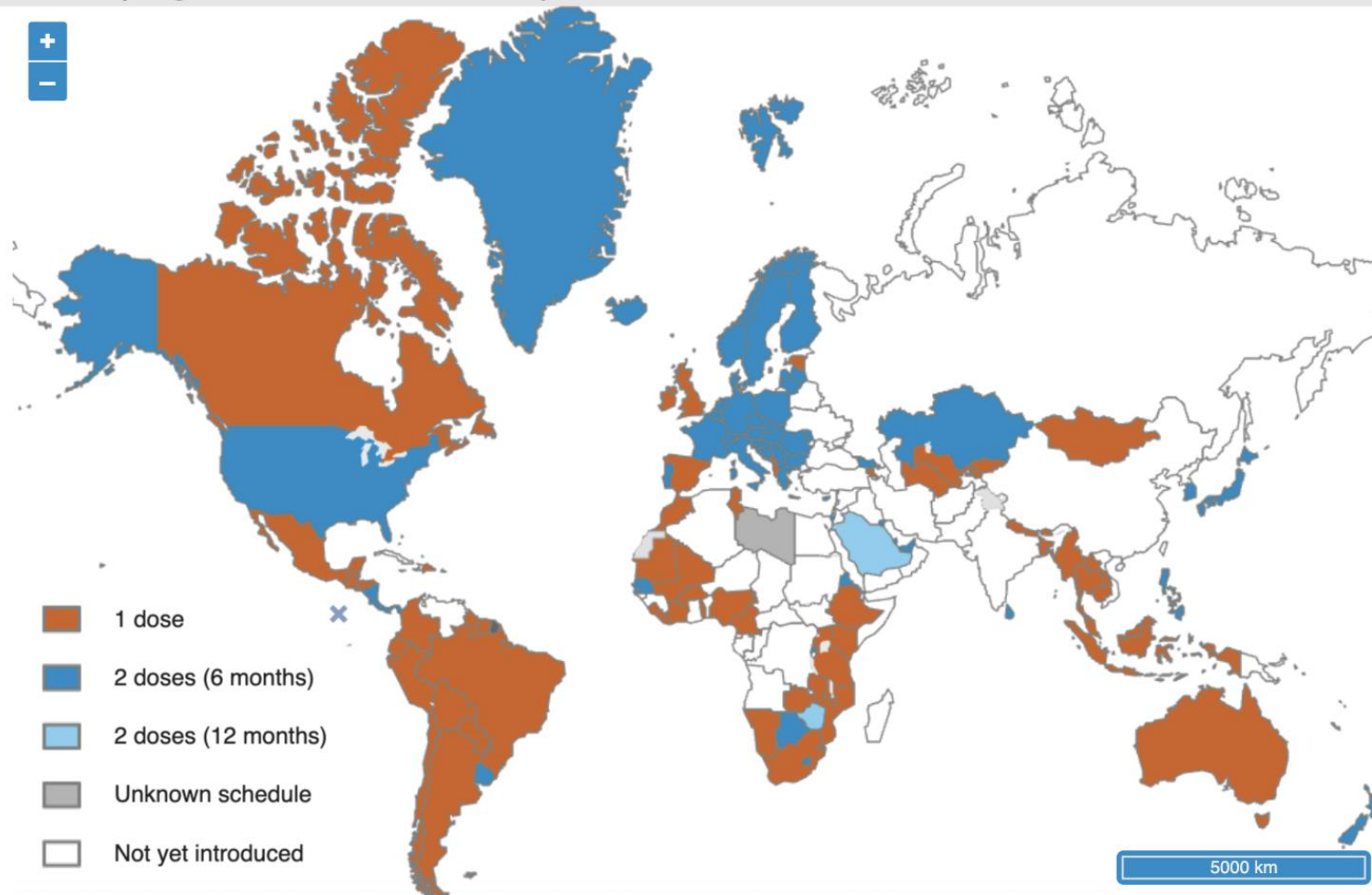


HPV vaccination programme schedule (9-14 years old)

Date of slide: Aug 2025

Map production: Immunization Vaccines
Biologicals (IVB), World Health Organization
Data Source: WHO HPV vax Intro Dashboard

Interval_doses	No. of countries
1 dose	77
2 doses (12 months)	2
2 doses (6 months)	69
Not yet introduced	44
Unknown schedule	2



Last update:

8/22/2025 10:12:27
AM

77/149
(51.3%)
one dose

And 83 /150
(55.3%)
Both sex

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So where are we with HPV vaccine....



- Extremely effective and safe prophylactic vaccines in use since 2006
- Over 500 million doses, proven to prevent cervical cancer
- Supply constraints easing, one dose schedule in use
- Primary pillar of the global cervical cancer elimination strategy
 - Very cost effective in almost all settings
- Can be successfully implemented at scale at any income level
 - School based delivery, with systems for out of school girls, and clear eligibility criteria
 - Micro plan, involve teachers and multisector collaborations
 - Proactive, early community mobilisation essential
 - Messaging focused on cancer prevention, govt endorsement, safety and efficacy, when and where delivered



Vaccination of young adolescents against HPV is safe and prevents cervical cancer

Human Papillomavirus (HPV) is the cause of cervical cancer and is the most common sexually transmitted infection.

Tsu et al Prev Med 2021, Gallagher PlosOne 2017, Howard PVR 2017, Kabakama BMC Public Health 2016, Soi et al Implementation Science 2018, Gallagher PVR 2017, LaMontagne IntJGyneObst 2017, WHO Elimination Strategy 2020

Invasive meningococcal disease in adolescents

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Professor of Vaccinology and NHMRC Fellow, The University of Adelaide



THE UNIVERSITY
of ADELAIDE

Invasive meningococcal disease in adolescents

- Second highest incidence to infants
- ~10% case fatality rate in adolescents
- sequelae – blindness, deafness, limb amputation
- Most cases (85%) of meningococcal disease in adolescents are due to the B strain (A, B, C, W, Y strains)
- 10% of adolescents carry the meningococcus in their oropharynx
- MenACWY conjugate vaccine has been funded through the National Immunisation Program(NIP) for adolescents
- MenB protein-based vaccines are not included on the NIP for adolescents and are available for private purchase.
- SA, NT, Queensland, Tasmania (to commence) have state funded programs

Adelaide teenager Jack Klemich dies of meningococcal disease

JILL PENGELLEY AND JOANNA VAUGHAN THE ADVERTISER MAY 26, 2009 9:00PM

SHARE    

 SAVE THIS STORY

JACK Klemich was popular, talented and fortunate - he had it all.

But three days after developing meningococcal disease, the vibrant, athletic son of real estate identity Oren Klemich was gone.

The savage speed of the disease has stunned mates who saw Jack, 18, play football for his St Peter's College team on Saturday. Twenty-four hours later, he was fighting for his life and early today, he quietly slipped away.

Despite emergency treatment with antibiotics and being put on life support, the fit sportsman could not be saved.



AMEND (Adolescent MENingococcal Disease) study; the lived experience

I drove home ..., which is about 20 minutes, and I was crying the whole time because I knew something wasn't right, and by the time I got home I couldn't get the key in the backdoor - P15 Female, 16.

So normally when I'm sick and I don't go to work, mum just leaves me, shuts my door, doesn't even come in, but for some reason she came in my room that day. I was covered in vomit, my eyes were rolling back and she rang the ambulance. - P11, Female, 20

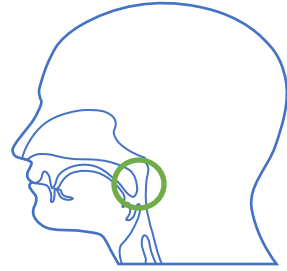
[The] doctor saw me and she said oh, it looks like you're coming down with the flu, just go home, take some Nurofen, you'll be fine – P25, Female, 18

And then by the time the home doctor comes, I can't really use my legs anymore. So I'm kind of like crawling or like, you know, holding on to things drag myself through the door to let him in. He said that he thought I might have a bladder infection. So to wait until tomorrow and get some antibiotics. So [he] gave me a prescription and left. – P6, Female 23

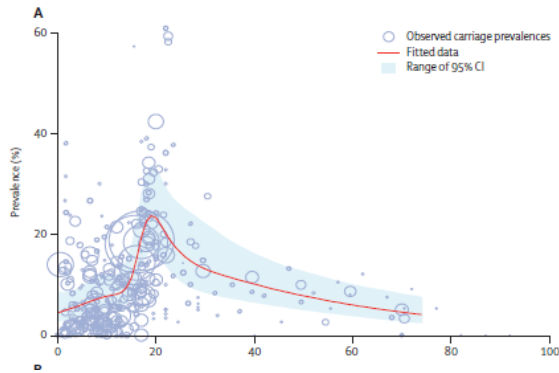
My mum's probably like the worst affected by it. She can't really even talk about it. She just starts crying. – P18, female, 19

There's still like a limit on how much I can do in the day without just crashing for a week afterwards. Placement was really hard this year. – P4, female, 16

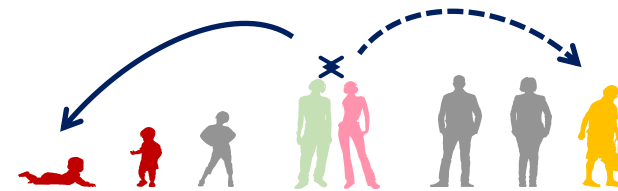
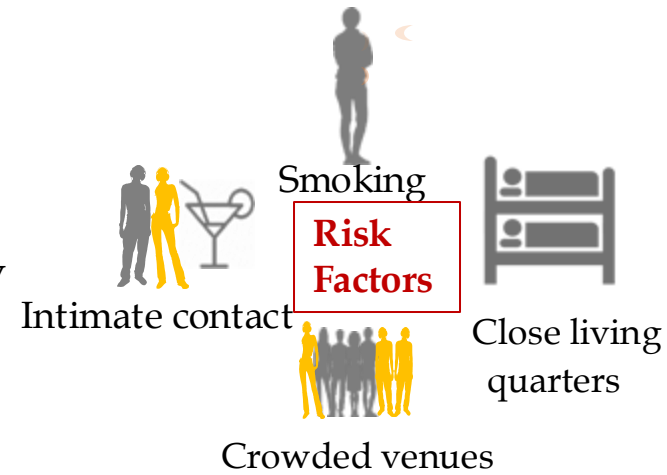
Oro-pharyngeal carriage of *Neisseria meningitidis*



N. Meningitidis colonises the nasopharynx of healthy individuals^{1,2}



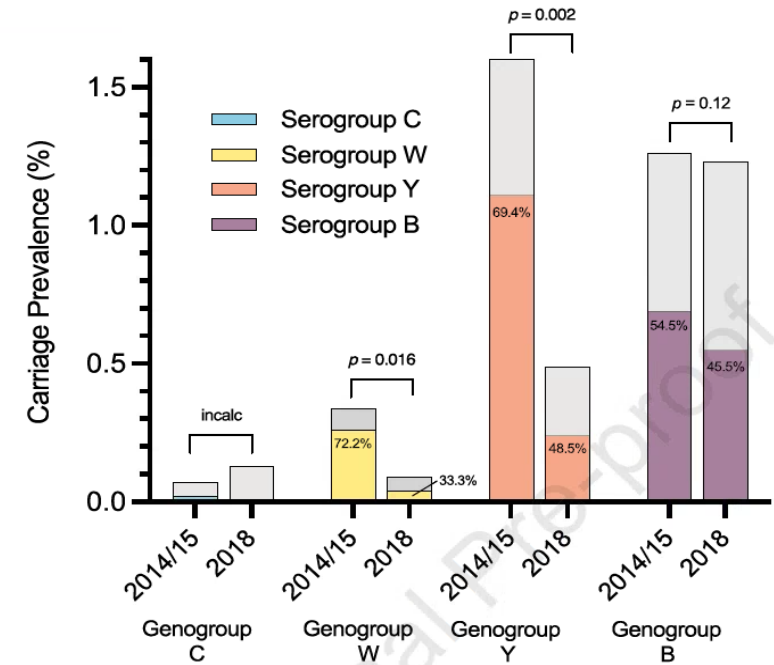
~10% of the general population are asymptomatic carriers rates are highest in older adolescents and young adults^{1,2}



Asymptomatic carriers are a major source of transmission⁴

Meningococcal ACWY vaccines are safe and effective against IMD and have a herd immunity impact

Vaccine type (IMD outcome)	Country	Age		Odds Ratio (95% CI)	% Weight
Author, year					
MCC vaccines (Group C IMD)					
Bose, 2003	England	15 to 19 yrs		0.07 (0.01, 0.61)	7.86
Cardoso, 2015	Brazil	< 5 & 10 to 24 yrs		0.03 (0.00, 0.21)	8.74
DeWals, 2011	Canada	2 mths to 20 yrs		0.13 (0.06, 0.25)	47.64
Pezzotti, 2018	Italy	1 to 22 yrs		0.20 (0.08, 0.46)	35.76
Subtotal (I-squared = 13.6%, p = 0.324)				0.13 (0.07, 0.23)	100.00
MenACWY vaccines (Group ACWY IMD)					
Cohn, 2017	United States	11 to 19 yrs		0.31 (0.20, 0.49)	100.00



Meningococcal B vaccine effectiveness in adolescents against IMD and carriage

'B Part of It' study

Impact of 4CMenB vaccine on carriage of *Neisseria meningitidis* in adolescents

Professor Helen Marshall MBBS MD MPH
National Health and Medical Research Council Practitioner Fellow
Adelaide Medical School and Deputy Director, RRI, The University of Adelaide
Senior Medical Practitioner, Women's and Children's Health Network, South Australia

www.bpartofit.com.au

On behalf of the Investigators and Scientific Advisory Committee: Ann Koehler, Andrew Lawrence, Tom Sullivan, Shamez Ladhani, Adam Finn, Ray Borrow, Martin Maiden, Jenny Maclennan, Matthew Snape, Caroline Trotter, Mary Ramsey, Charlene Kahler, Peter Richmond



Randomised, controlled trial to assess the impact of 4CMenB on carriage of *Neisseria meningitidis*

The 'B Part of It' study administered 4CMenB to high-school students in South Australia, 2017/18¹



62% of all South Australian students in Years 10 and 11 (ages 15–17 years) enrolled

91% of South Australian schools participated in the school-based programme (237/260)



Intervention group



Control group

Key:  Oropharyngeal swab

 BEXSERO dose

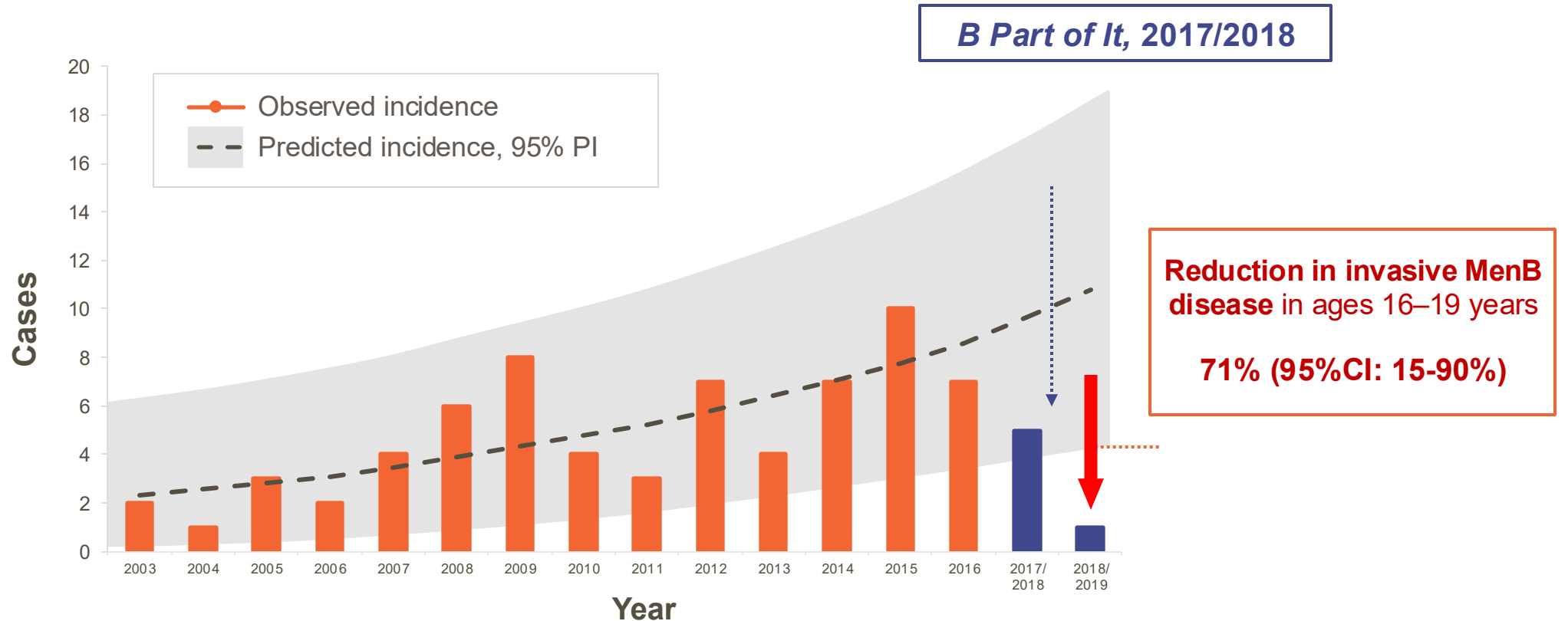
Total enrolment 34,489 students

No impact of 4CMenB on carriage of meningococcal strains associated with disease (capsule) but reduction in carriage of unencapsulated strains

<i>N. meningitidis</i>	Vaccinated		Unvaccinated		aOR (95% CI)	ap-value
All	4.3 %	547/12746	4.9%	561/11523	0.85 (0.70, 1.04)	0.117
Disease causing	2.6%	326/12746	2.5%	291/11523	1.02 (0.8, 1.31)	0.845
Secondary analysis carriage	Vaccinated		Control		aOdds ratio	ap-value
	%	n	%	n	95% CI	
Non-groupable (unencapsulated)	1.65	179/10841	2.23	229/10285	0.71 (0.54, 0.91)	0.008
Group W	0.16%	17/10841	0.18%	18/10285	0.89 (0.43, 1.85)	0.751
Group C	0.11%	12/10841	0.07%	7/10285	1.87 (0.63, 5.55)	0.260
Group X	0.07%	8/10841	0.01%	1/10285	-	-

<i>Acquisition N. meningitidis</i>	vaccinated		unvaccinated		aOR (95% CI)	ap-value
All groups	3.38 %	430/12746	3.7%	427/11523	0.91 (0.73, 1.13)	0.298
Disease causing groups	2.13%	272/12746	2.07%	238/11523	1.03 (0.79, 1.34)	0.954

Reduction in meningococcal B disease, following the state-wide study, evidence for direct protection



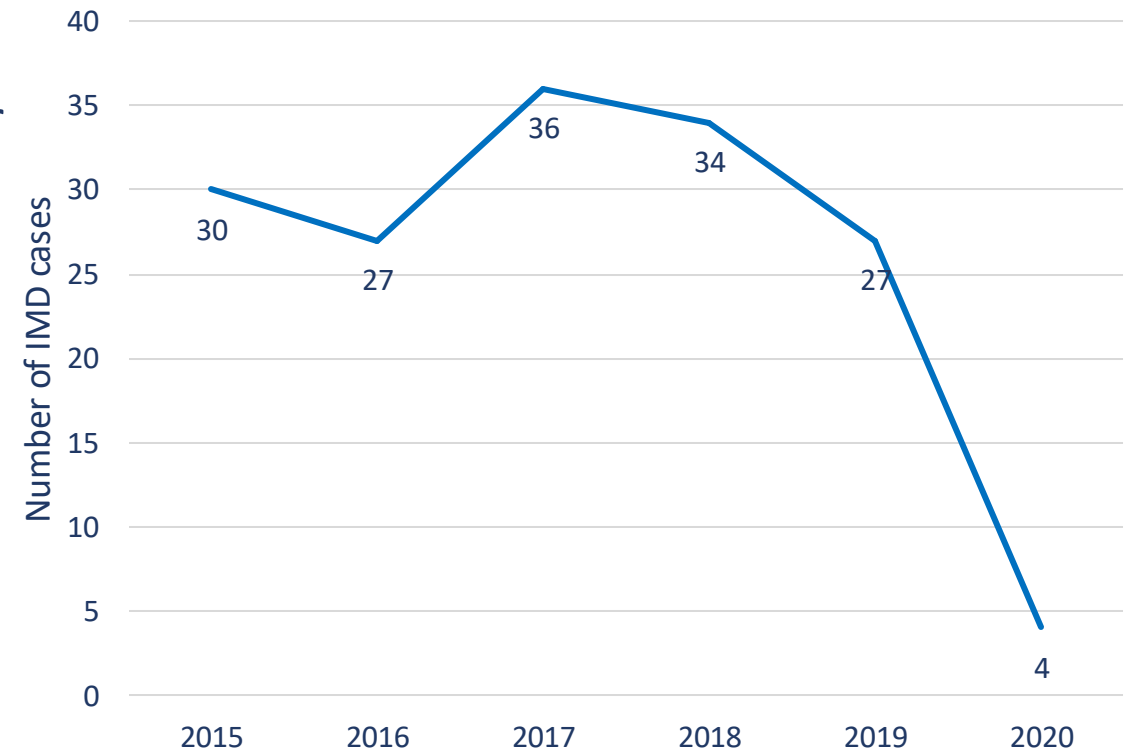
Vaccine effectiveness two years after introduction of the funded 4CMenB vaccine program in SA 2018–2020

Vaccine effectiveness against meningococcal B disease in infants

- 94.7% (95% CI: 40.3–99.5, $P=0.025$) for two doses

Vaccine effectiveness against meningococcal B disease in adolescents

- ~100% for two doses (Variance and confidence intervals for VE cannot be calculated because of the lack of vaccinated cases)



IMD in South Australia following introduction of 4CMenB (state funded) and MenACWY (NIP)

Moderate effectiveness of 4CMenB against gonorrhoea at 2 years post 4CMenB vaccine introduction

Previous evidence of VE against gonorrhoea with OMV vaccines

- MenB-OMV vaccines showed a post vaccine program reduction in gonorrhoea in Cuba
- MeNZB vaccine showed VE=31%¹

Cohort and case control study in SA assessing effect of 4CMenB against gonorrhoea ²

- Population: 15-20 yr olds in SA, 2 years post-introduction of 4CMenB
- VI=24% (aIRR=0.76 (0.48, 1.22) reduction in gonorrhoea incidence
- VE=33% (8.3, 50.6) against gonorrhoea

There is currently no indication for 4CMenB to be used for protection against gonorrhoea

Effectiveness and impact of the 4CMenB vaccine against invasive serogroup B meningococcal disease and gonorrhoea in an infant, child, and adolescent programme: an observational cohort and case-control study



Bing Wang, Lynne Giles, Prabha Andraweera, Mark McMillan, Sara Almond, Rebecca Beazley, Janine Mitchell, Noel Lally, Michele Ahoure, Emma Denshy, Ann Koehler, Louise Flood, Helen Marshall

Summary

Background A programme of vaccination with the four-component serogroup B meningococcal (4CMenB) vaccine was introduced in South Australia for infants and children aged 0–3 years on Oct 1, 2018, and for senior school students in school years 10 and 11 (aged 15–16 years) and young adults aged 17–20 years on Feb 1, 2019. We aimed to evaluate vaccine effectiveness and impact on serogroup B meningococcal disease and gonorrhoea 2 years after implementation of the programme.

Lancet Infect Dis 2022; 22: 1011–20
April 12, 2022
https://doi.org/10.1016/S1473-3099(21)0054-4

Methods We did a cohort and case-control study among those targeted by the South Australia 4CMenB vaccination programme. We obtained disease notification data from SA Health, Government of South Australia, and vaccine coverage data from the South Australian records of the Australian Immunisation Register. Vaccine effectiveness was estimated as the reduction in the odds of infection using the screening and case-control methods. Vaccine impact was estimated as incidence rate ratios (IRRs), obtained by comparing case numbers in each year following the start of the vaccination programme with cases in the equivalent age cohort during the pre-vaccination programme years. We used Poisson or negative binomial models, as appropriate, with adjustment for changes in the incidence of serogroup B meningococcal disease in age cohorts not eligible for vaccination through the state programme.

See Comment page 919
Vaccinology and Immunology Research Trials Unit, Women's and Children's Health Network, Adelaide, SA, Australia
(B Wang PhD, P Andraweera PhD, M McMillan PhD, Prof H Marshall MD), Robinson Research Institute (B Wang, L Giles PhD, P Andraweera, M McMillan, Prof H Marshall MD), Adelaide Medical School (B Wang, P Andraweera, M McMillan, Prof H Marshall), and School of Public Health (L Giles), University of Adelaide, Adelaide, SA, Australia; Communicable Disease Control Branch, SA Health, Adelaide, SA, Australia (S Almond GradCert/INMP, R Beazley MPH, J Mitchell MPH, N Lally MPH, M Ahoure GradCert/INMP, E Denshy MPH, A Koehler FRCPA, L Flood FRPHM)

Findings 4CMenB vaccine coverage 2 years after introduction of the childhood vaccination programme was 94·9% (33 357 of 35 144 eligible individuals) for one dose, 91·4% (26 443 of 28 922) for two doses, and 79·4% (15 440 of 19 436) for three doses in infants. The one-dose (77·1%, 16 422 of 21 305) and two-dose (69·0%, 14 704 of 21 305) coverage was highest in adolescents born in 2003 (approximately year 10 students). 2 years after implementation of the childhood vaccination programme, incidence of serogroup B meningococcal disease was significantly reduced compared with before programme implementation in infants aged 12 weeks to 11 months [adjusted IRR (aIRR) 0·40 [95% CI 0·23–0·69], p=0·0011], but not in those aged 1 year (0·79 [0·16–3·87], p=0·77), 2 years (0·75 [0·18–3·14], p=0·70), or 4 years (3·00 [0·47–18·79], p=0·24). aIRRs were not calculable in those aged 3 or 5 years because of no cases occurring after programme implementation. aIRR for serogroup B meningococcal disease was 0·27 (0·06–1·16, p=0·078) in adolescents aged 15–18 years 2 years after implementation of the adolescent and young adult programme, and 1·20 (0·70–2·06, p=0·51) in those aged 19–21 years in the first year. Two-dose vaccine effectiveness against serogroup B meningococcal disease was estimated to be 94·2% (95% CI 36·6–99·5) using the screening method and 94·7% (40·3–99·5) using the case-control method in children, and 100% in adolescents and young adults (no cases reported after implementation). Estimated two-dose vaccine effectiveness against gonorrhoea in adolescents and young adults was 32·7% (8·3–50·6) based on the case-control method using age-matched individuals with chlamydia infection as controls.

Interpretation 4CMenB vaccine shows sustained effectiveness against serogroup B meningococcal disease 2 years after introduction in infants and adolescents, and moderate effectiveness against gonorrhoea in adolescents. The high vaccine effectiveness against serogroup B meningococcal disease is likely due to high coverage in the target age

Correspondence to:
Prof Helen Marshall, Vaccinology and Immunology Research Trials Unit, Women's and Children's Hospital, Adelaide, SA 5006, Australia
helen.marshall@adelaide.edu.au

Gonococcal infection burden of disease

- 82 million new infections annually¹, increasing globally, despite WHO's global strategy for 90% reduction by 2030
- **Burden of disease affects mostly women and children**; pelvic inflammatory disease - infertility, ophthalmia neonatorum – blindness
 - Up to 80% of gonococcal infections in women are asymptomatic
 - 50% risk of infertility if ≥ 3 infections²
- **Increasing development of antibiotic resistance** with potential for gonorrhoea to become untreatable³

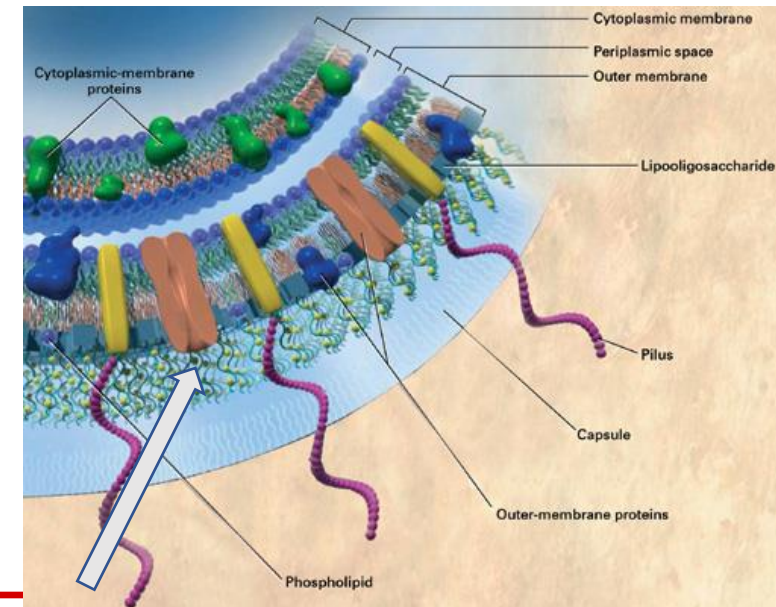


High risk populations: MSM, sex workers, people in prison, people living with HIV, adolescents and young adults, Indigenous populations (complex social and cultural determinants, barriers and access to healthcare)

- **x24 higher risk for Aboriginal and Torres Strait Islander people, Australia**
- 8.2% prevalence in adolescent girls and young women, East Africa

Mechanism for 4CMenB vaccine protection against gonorrhoea

- ~ 90% genetic homology, both mucosal pathogens
- 4CMenB vaccine induced antibodies have been shown to recognise gonococcal proteins²
- reduction in carriage of unencapsulated meningococci in the vaccinated group¹
 - vaccine-type outer membrane proteins are potentially more exposed without a capsule
 - the gonococcus is an unencapsulated Neisseria bacteria proteins targeted by 4CMenB vaccine expressed on *N. meningitidis* as well as *N. gonorrhoeae* (OMV, NHBA) and have high homology²



Gonorrhoea notifications are highest in 15-19 year olds in the NT which is also a high risk age group for invasive meningococcal disease

Table D.1 Gonorrhoea notifications in the NT by age

Age group	Prior 5-year mean for Jan-Jun (2016-2020)	
	Number	Proportion (%)
<10	0.4	0.0
10-14	39.2	4.6
15-19	230.6	27.2
20-24	178.6	21.1
25-29	134.2	15.9
30-34	101.0	11.9
35-39	65.0	7.7
40-44	42.4	5.0
45-49	26.8	3.2
50-54	13.0	1.5
55-59	6.2	0.7

Surveillance Update for Notifiable STI and BBV in the Northern Territory Vol. 24, No. 2 July to December 2023

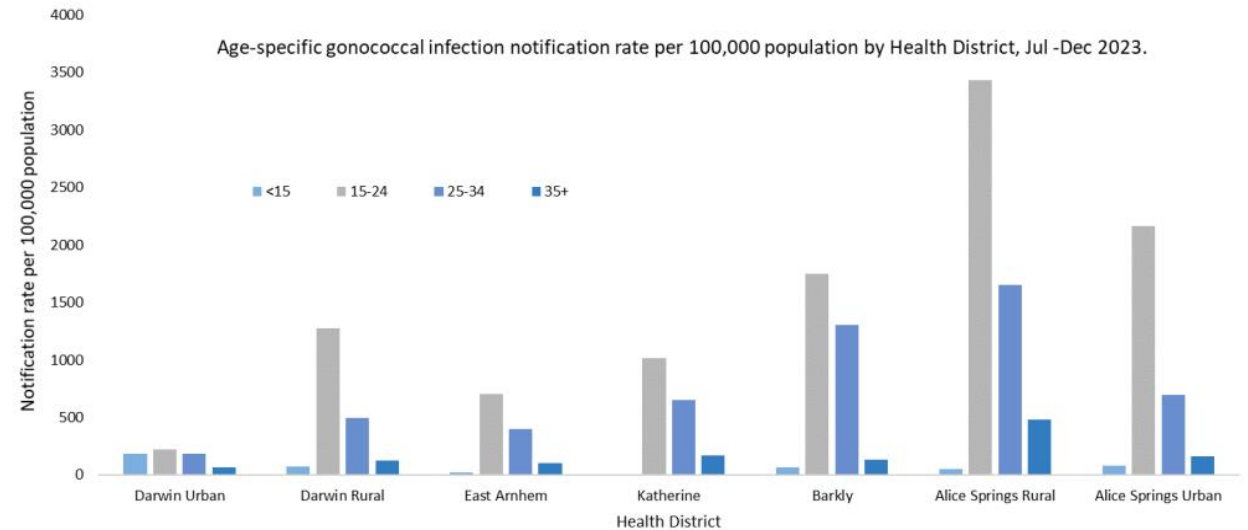


Figure C1: Age-specific gonococcal infection notification rate per 100,000 population by Health District, Jul-Dec 2023

B Part Of It NT study

Population: 14-19 year olds in urban, rural and remote regions of the Northern Territory

Aims: Assess impact of 4CMenB vaccine on

1. *Neisseria gonorrhoeae*
2. Oro-pharyngeal carriage of *Neisseria meningitidis*

Intervention: 4CMenB vaccine offered to all eligible adolescents

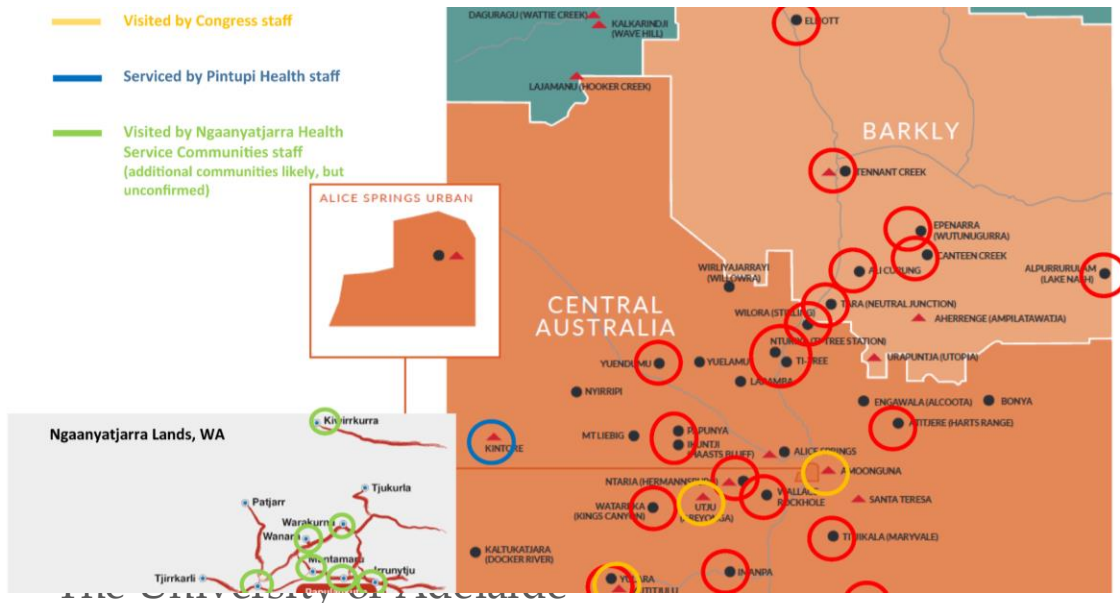
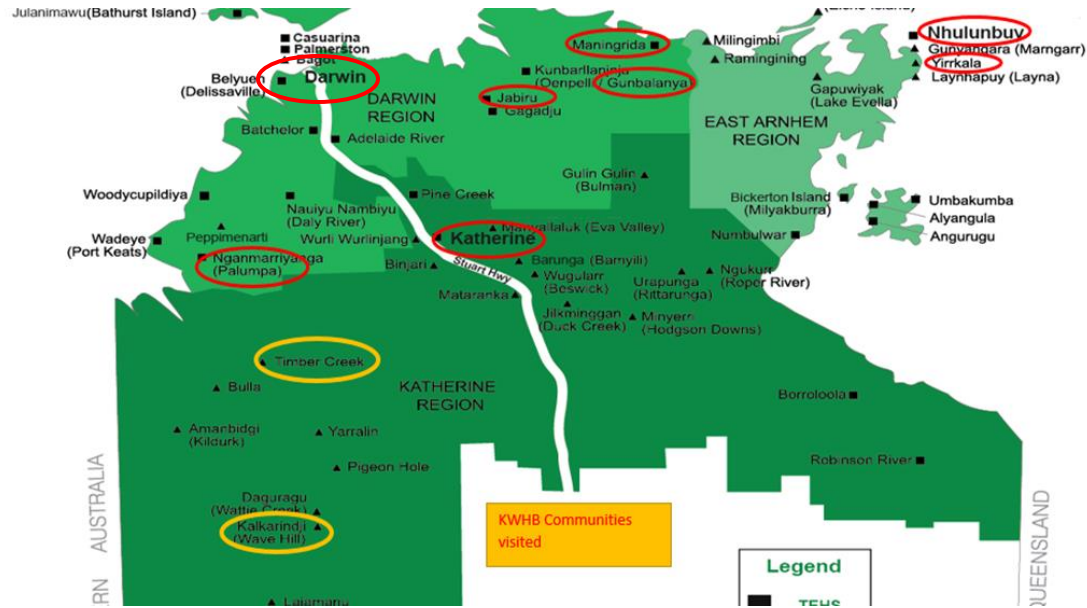
Outcome: Time to gonococcal notification

Time to event analysis: VE was assessed using a Cox proportional hazards model



B Part of it NT study results

- Of 1,324 individuals diagnosed with gonorrhoea, 97.5% were Indigenous young people, 59.4% were females
- Overall vaccine uptake in 14-19 year old cohort was 8.5% (up to 18% in 15 year olds)
- **VE =38.1% (95%CI: 18.1-53.4%)** against gonorrhoea



A 4CMenB vaccine program has now been introduced in the NT for infants and adolescents with ongoing evaluation in a high uptake program

Long term vaccine effectiveness and impact against IMD and gonorrhoea in adolescents 5 years post 4CMenB program introduction

IMD

Vaccine Effectiveness

- VE=92.6% (95%CI 37.5, 99.1; p=0.017)

Vaccine impact

- 76.2% relative reduction in IMD in adolescents 15-18 years of age

IMD cases =19 in adolescents

- 14-24 years of age
- 63% male
- **84% unvaccinated**
- 10% First Nations young people

Gonorrhoea

Vaccine Effectiveness:

- 40.1 % (95%CI: 31.3-46.0) p<0.001

Vaccine Impact:

- 35.5% relative reduction in incidence of gonococcal infection

Vaccine effectiveness results (chlamydia controls) subgroup analysis suggest waning by 5 years

Two-dose 4CMenB vs unvaccinated	Adjusted OR (95%CI)	Vaccine effectiveness (%) (95%CI)
Duration of protection		
3-60 months since vaccination	0.582 (0.513-0.660) p<0.001	41.8 (34.0 to 48.7)
>60 months since vaccination	1.063 (0.782-1.445) p=0.695	-6.3 (-44.5 to 21.8)
Sex		
Female	0.596 (0.507-0.699) p<0.001	40.4 (30.1 to 49.3)
Male	0.630 (0.525-0.755) p<0.001	37.0 (24.5 to 47.5)
Co-infection		
Cases co-infected, chlamydia controls	0.525 (0.430-0.643) p<0.001	47.5 (35.7-57.0)
Gonorrhoea only, chlamydia controls	0.653 (0.568-0.751) p<0.001	34.7 (24.9-43.2)

Impact of 4CMenB on repeat gonococcal infection

Survival analysis using a multivariable Cox proportional hazards regression model

- Adjusted hazard ratio for repeat gonococcal infections in vaccinated vs unvaccinated gonococcal cases (adjusted for sex, SES with stratification by age)

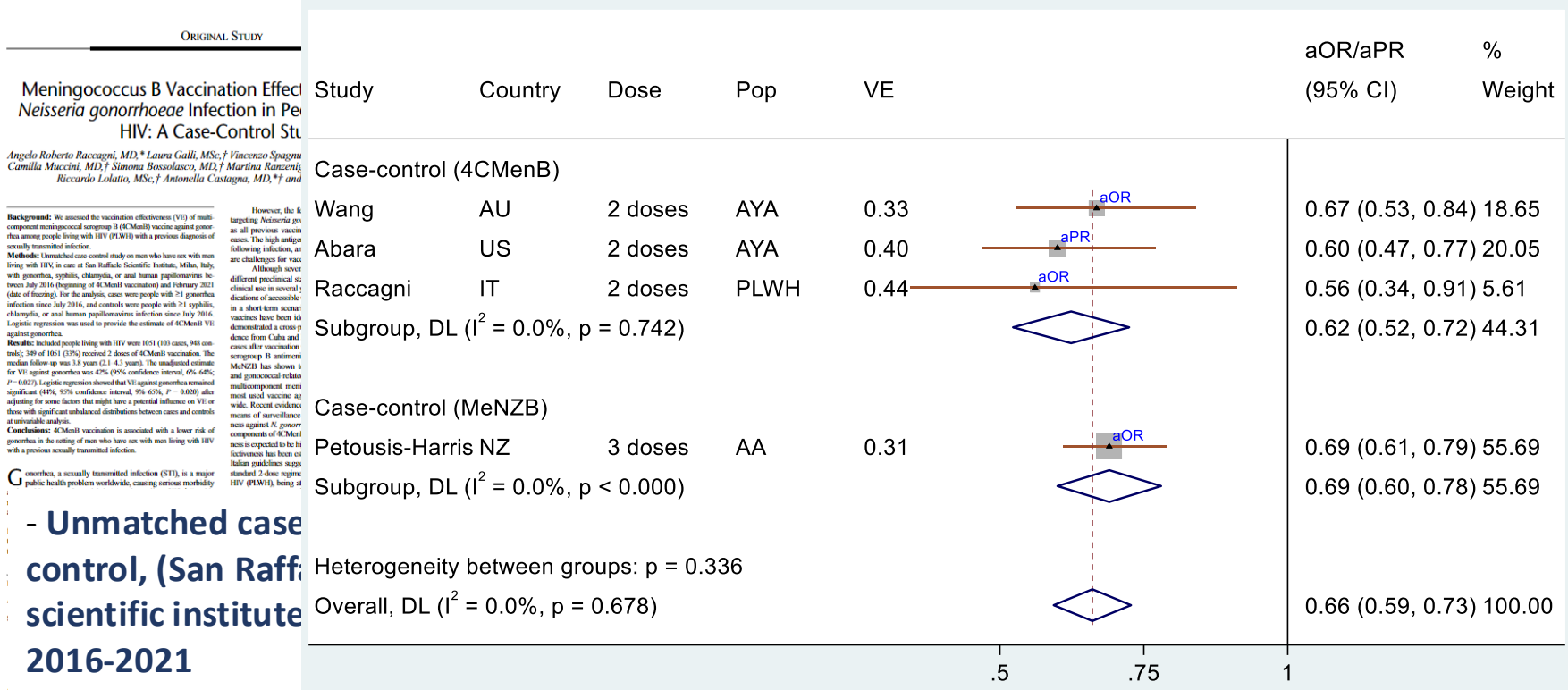
The risk of a second episode of gonococcal infection was lower in gonococcal cases who were vaccinated with two doses

- **VE=29.1% aHR=0.730 (95%CI 0.540, 0.988) p=0.042**

No difference was observed for multiple repeat gonococcal infections

- **VE=7.8% (aHR= 0.922; 95% CI, 0.643–1.320; p=0.656)**

Mounting global real world evidence for a moderate effectiveness of 4CMenB against gonorrhoea



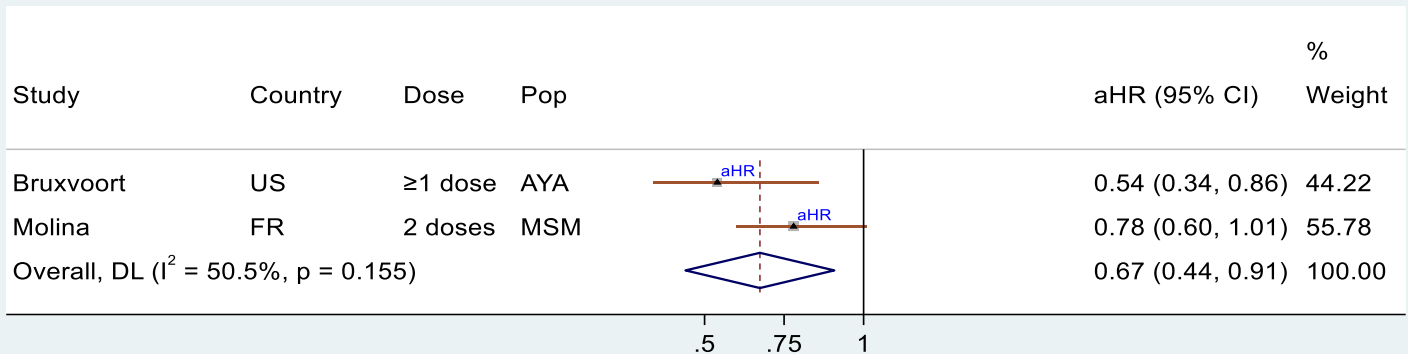
- Unmatched case control, (San Raffaele scientific institute 2016-2021)

- 103 cases in 105 living with HIV

- 37-51 yrs

- VE=44% (95%CI)

- Italy



Effectiveness of a serogroup B outer membrane vesicle meningococcal vaccine against gonorrhoea: a retrospective observational study

Winston E Abara, Kyle T Bernstein, Felicia M T Lewis, Julia A Schillinger, Kristen Feemster, Preeti Pathela, Susan Hariri, Aras Islam, Michael Eberhart, Iris Cheng, Alexandra Terrier, Jennifer Sanderson Slutsker, Sarah Mbaayi, Robbie Madera, Robert D Kirkcaldy

Summary
Background Declining antimicrobial susceptibility to current gonorrhoea antibiotic treatment and inadequate treatment options have raised the possibility of untreatable gonorrhoea. New prevention approaches, such as vaccination, are needed. Outer membrane vesicle meningococcal serogroup B vaccines might be protective against gonorrhoea. We evaluated the effectiveness of a serogroup B meningococcal outer membrane vesicle vaccine (MenB-4C) against gonorrhoea in individuals aged 16–23 years in two US cities.

Methods We identified laboratory-confirmed gonorrhoea and chlamydia infections among individuals aged 16–23 years from sexually transmitted infection surveillance records in New York City and Philadelphia from 2016 to 2018. We linked gonorrhoea and chlamydia case records to immunisation registry records to determine MenB-4C vaccination status at infection, defined as complete vaccination (two MenB-4C doses administered 30–180 days apart), partial vaccination (single MenB-4C vaccine dose), or no vaccination (serogroup B meningococcal vaccine naive). Using log-binomial regression with generalised estimating equations to account for correlations between multiple infections per patient, we calculated adjusted prevalence ratios (aPR) and 95% CIs to determine if

-Retrospective case-control study (CDC), 2016-2018

-167,706 gonorrhoea infections, 16-23 yrs

-VE=40%(23, 53) against gonorrhoea (2 dose)

VE=26% (12, 37) (1 dose)

- Philadelphia, USA

1. Raccagni et al. Sexually transmitted diseases 2023;50(5):247-251. 2. Molina J-M et al, CROI conference, March 2024. 3. Bruxvoort KJ et al. CID 2023;76:e1341-49. 4. Abara WE et al. Lancet ID 2022;22(7):1021-9. 5. Wang B et al. Journal of Infection 2024; 106225

Policy recommendations for use of 4CMenB to reduce the risk of gonorrhoea in high risk groups, UK

JCVI, UK recommendation, 10 November 2023 and funded program launched on 22nd May 2025.

Recommendation for a targeted programme using 4CMenB to prevent gonorrhoea among individuals at higher risk of infection attending sexual health services.

These risk criteria may include but not be limited to:

- a recent history of gonorrhoea or other bacterial STI diagnosis, individuals should also be offered vaccination
- after a gonorrhoea diagnosis (whether symptomatic or asymptomatic)
- reporting high-risk sexual behaviours with multiple partners during sexual health screening and assessment

Selectively vaccinating high risk groups was the most cost-effective intervention.

In conclusion

- Although uncommon, IMD is a life-threatening disease associated with long term disability in adolescents
- Meningococcal vaccines are available to reduce the risk of IMD
- Population programs of meningococcal ACWY and B vaccine are highly effective
- Evidence suggest cross protection against gonorrhoea
- Policy considerations - 4CMenB for groups at higher risk of both diseases including Aboriginal and Torres Strait Islander young people



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disease control

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AIM is to determine the most effective immunisation program for prevention of meningococcal disease and gonorrhoea in Australia and globally

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CRE** Centre for Research
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disease control

By 2030, the NEIS CRE will deliver a roadmap to eliminate meningitis and drive down gonorrhoea globally

<https://neiscre.com.au>

Safety of adolescent vaccines

- School immunization programs have protocols in place to ensure safety of adolescents vaccinated in the school setting (eg to avoid mass reactions)
- Passive TGA surveillance
- Active surveillance through AusVaxSafety
- Incident diseases in adolescence differ from those in infancy early childhood
- Signal investigation often around diseases with new onset or frequent occurrence in adolescence eg autoimmune diseases, migraine



Vaccines given at 14–16 years*

Vaccine	Protects against
NIMENRIX OR MENQUADFI*	Meningococcal disease (types A, C, W and Y)

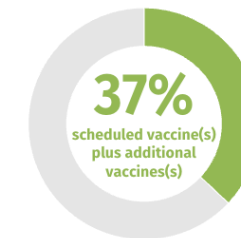
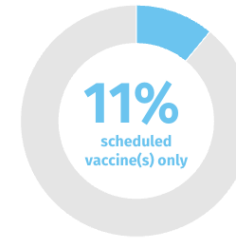
*Includes individuals aged up to 19 years who received a MenACWY vaccine as per recommendations in the Australian Immunisation Handbook

Vaccine safety surveys completed

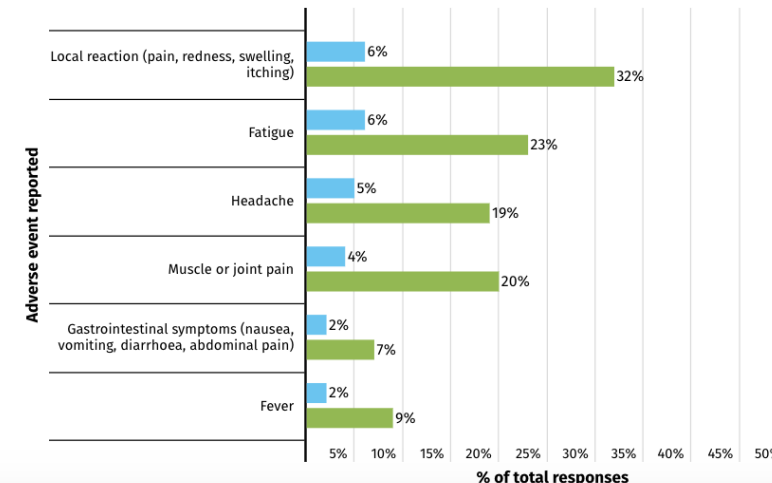
81,249
scheduled vaccine(s) only

3,082
scheduled vaccine(s) plus
additional vaccines(s)

Reported at least one adverse event after vaccination



Commonly reported adverse events after vaccination



Ongoing HPV vaccine safety evaluations



- Over 9 million doses to December 2017
- Purposeful enhanced surveillance 2013 and 2014 – increased rates of syncope noted

“Over an 11-year period, reporting rates of AE following 4vHPV administration in Australia were consistent with data from similar surveillance systems internationally and did not reveal any new or concerning safety issues.” Phillips et al, Vaccine 2020; 38:6038–46

- Ongoing passive national surveillance
- State based clinical support services
- Active real time surveillance via text

<https://ausvaxsafety.org.au/national-immunisation-program-schedule-vaccines/12-13-years-schedule-point>

See also: Macartney K et al, Drug Safety 2013
Phillips et al, Drug Safety 2018

12-13
years

Schedule point

Medical attendance

1 April 2022 - 30 June 2022

HPV (1st dose) and dTpa vaccines together

Less than 1 in 100 people reported taking their child to a doctor or emergency department in the days after vaccination.



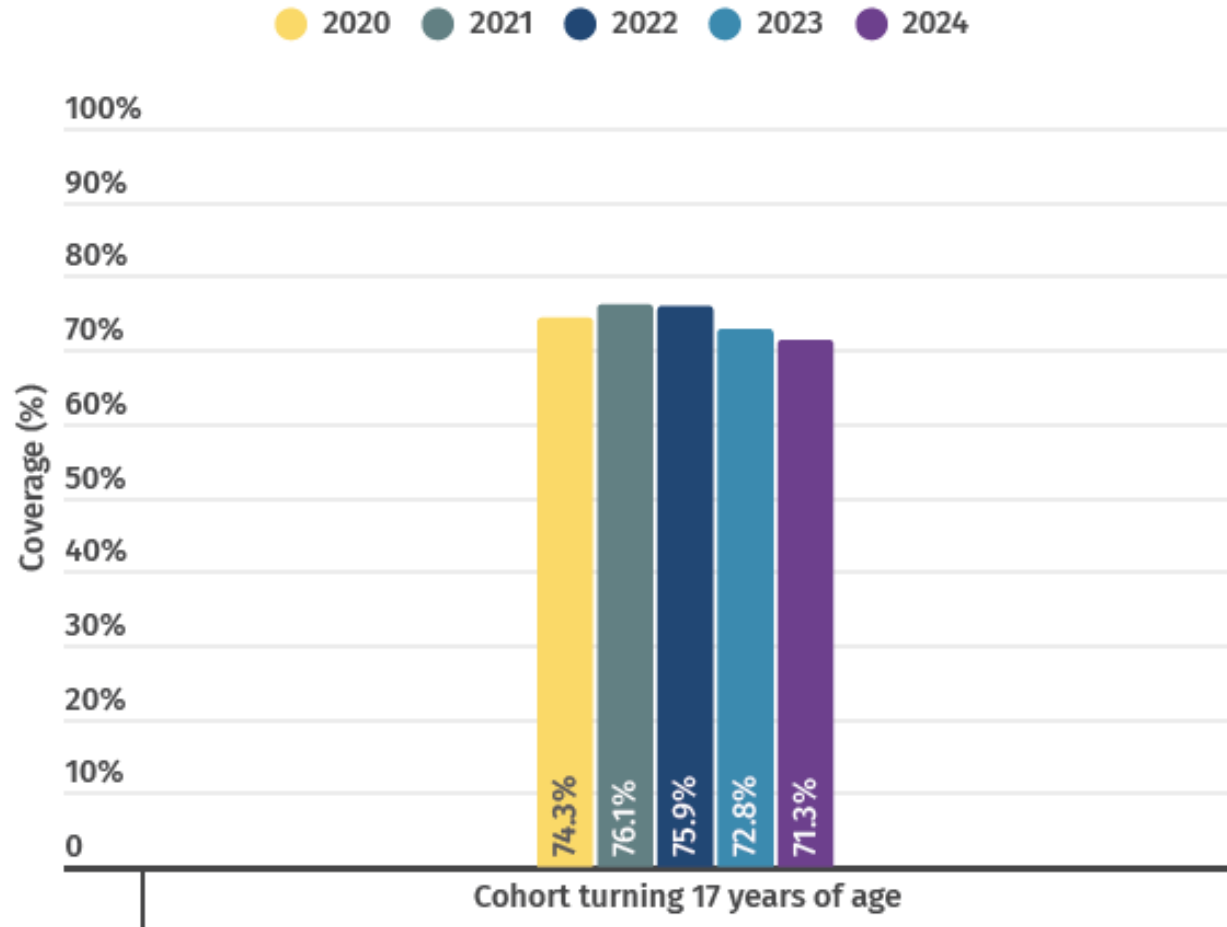
HPV vaccine (2nd dose) alone

Less than 1 in 100 people reported taking their child to a doctor or emergency department in the days after vaccination.





COVERAGE IN ADOLESCENCE



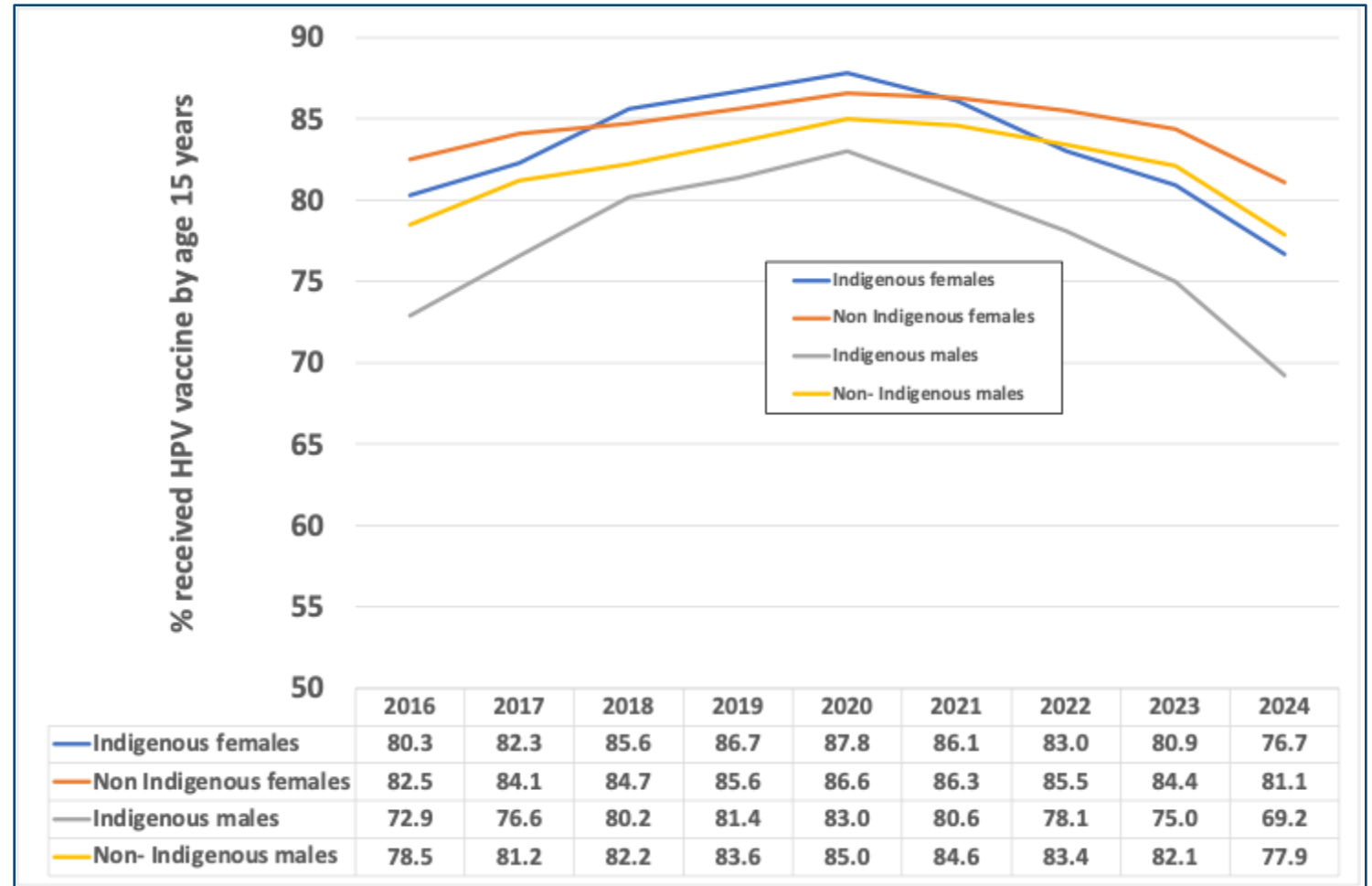
Meningococcal ACWY vaccine

Coverage of adolescent meningococcal ACWY vaccine in adolescents turning 17 years of age has also decreased in recent years and was lower than for other adolescent vaccines.



HPV vaccine rates are trending down

- HPV vaccine coverage peaked in Australia in 2020
- Ongoing small declines across childhood and adolescent vaccine schedule
- Also seen in similar countries post pandemic – USA, Canada, UK

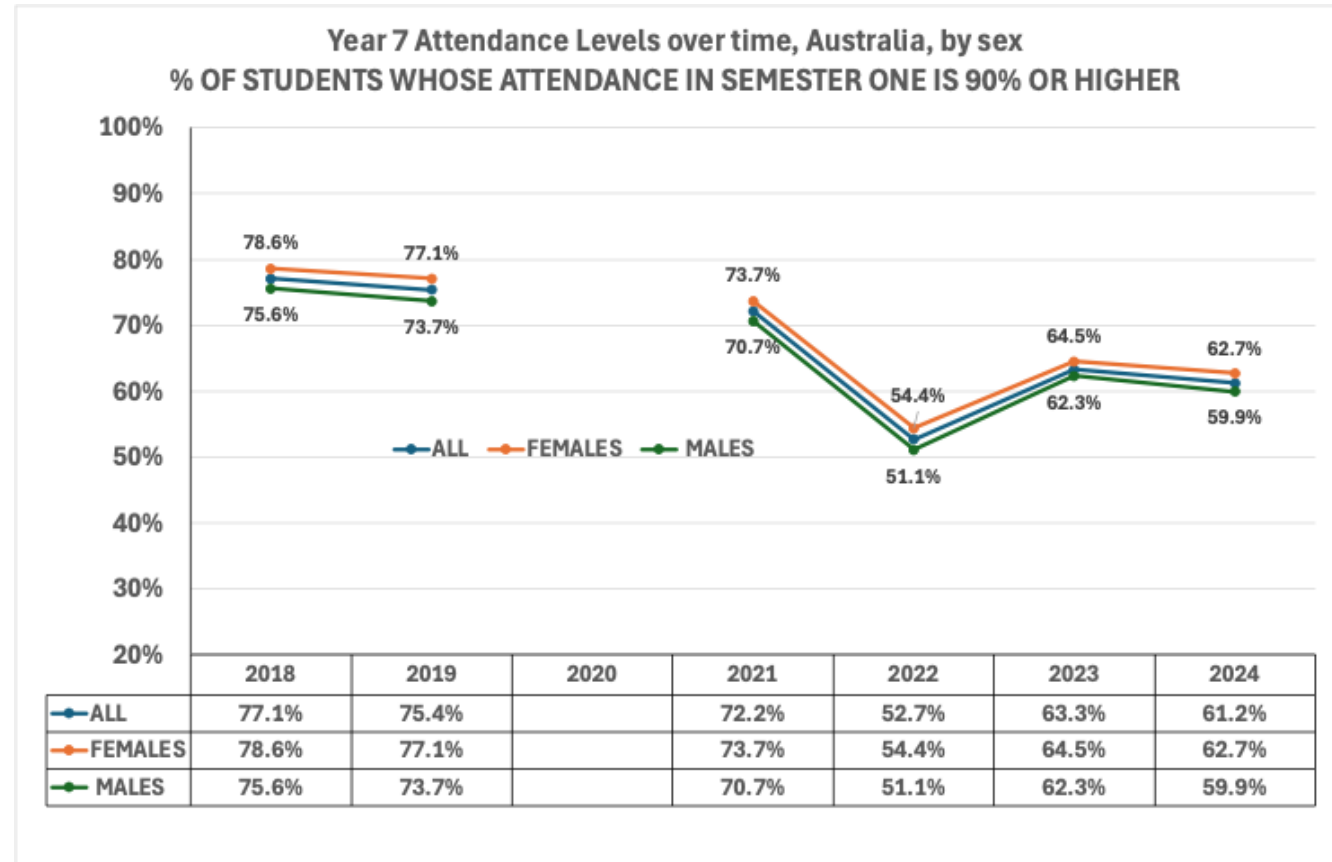


Data from Brotherton J, Machalek D, Smith M et al., 2024 Cervical Cancer Elimination Progress Report: Australia's progress towards the elimination of cervical cancer as a public health problem. Published online 14/03/2025, Melbourne, Australia, at <https://www.cervicalcancercontrol.org.au>. Updated with NCIRS interim report data for 2024



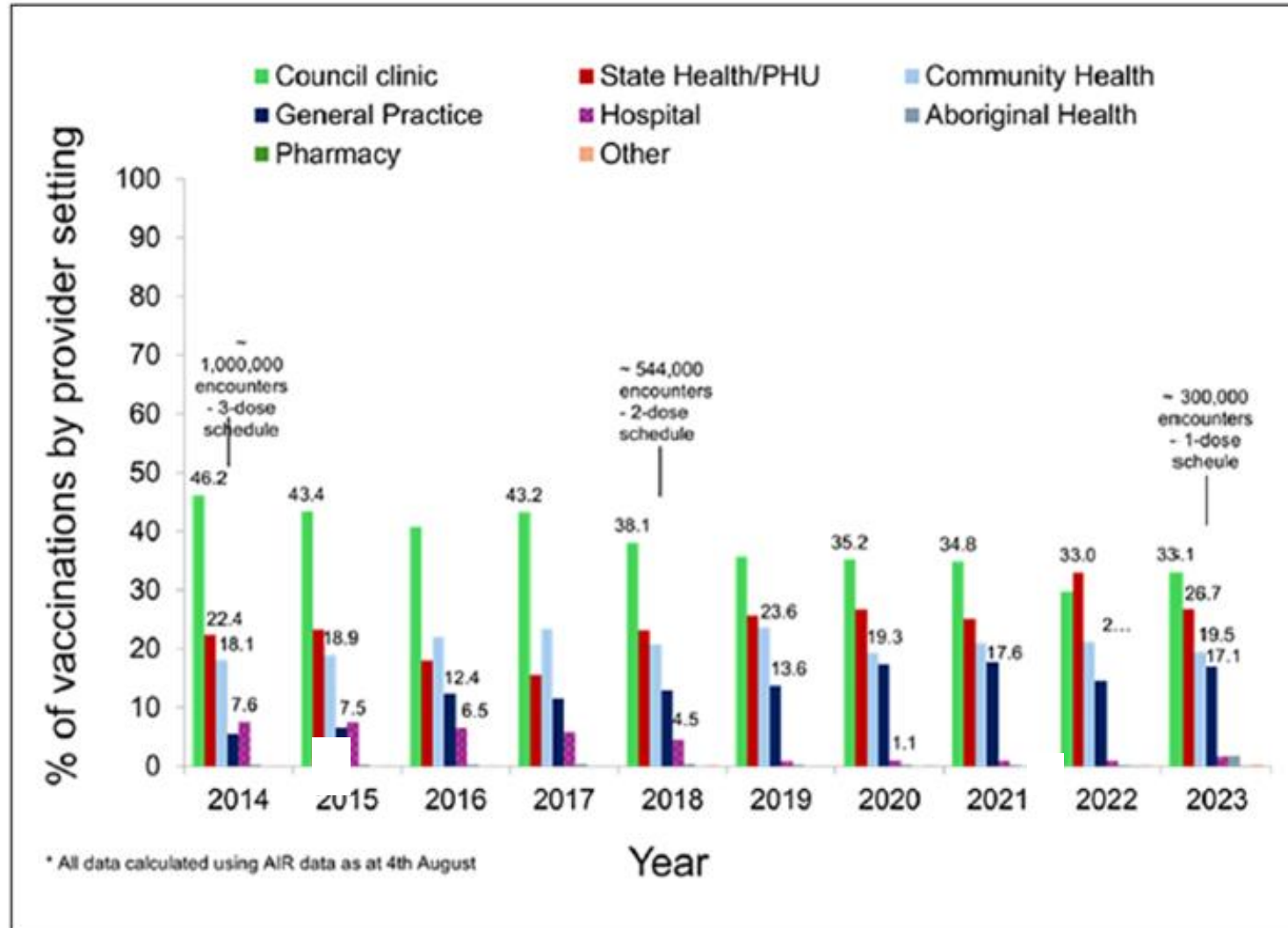
Hypotheses

- Kids at school less, more anxious
- Consent processes are challenging (mix of electronic and paper)
- Education/schools overwhelmed – staffing challenges and hands full. Limited resources/engagement
- Loss of/difficulty maintaining second visit in Year 7
- Parents ?hesitancy – overwhelmed, disengaged from vaccination, antigovernment



Source: ACARA data; Steffens et al, manuscript in preparation; McIndoe L, et al. Exploring Communication Barriers and Facilitators in School Vaccination: A Case Study in South Eastern Sydney, Australia. Vaccines (Basel). 2024 Oct 31;12(11):1243.

Proportion of HPV vaccine by provider, adolescents 10-19, 2014-2023, Australia



Reference: Hull BP, Hendry A, Beard F, Dey A. The Australian Immunisation Register (AIR): Insights from working with AIR data. Health Information Management Journal, 2025

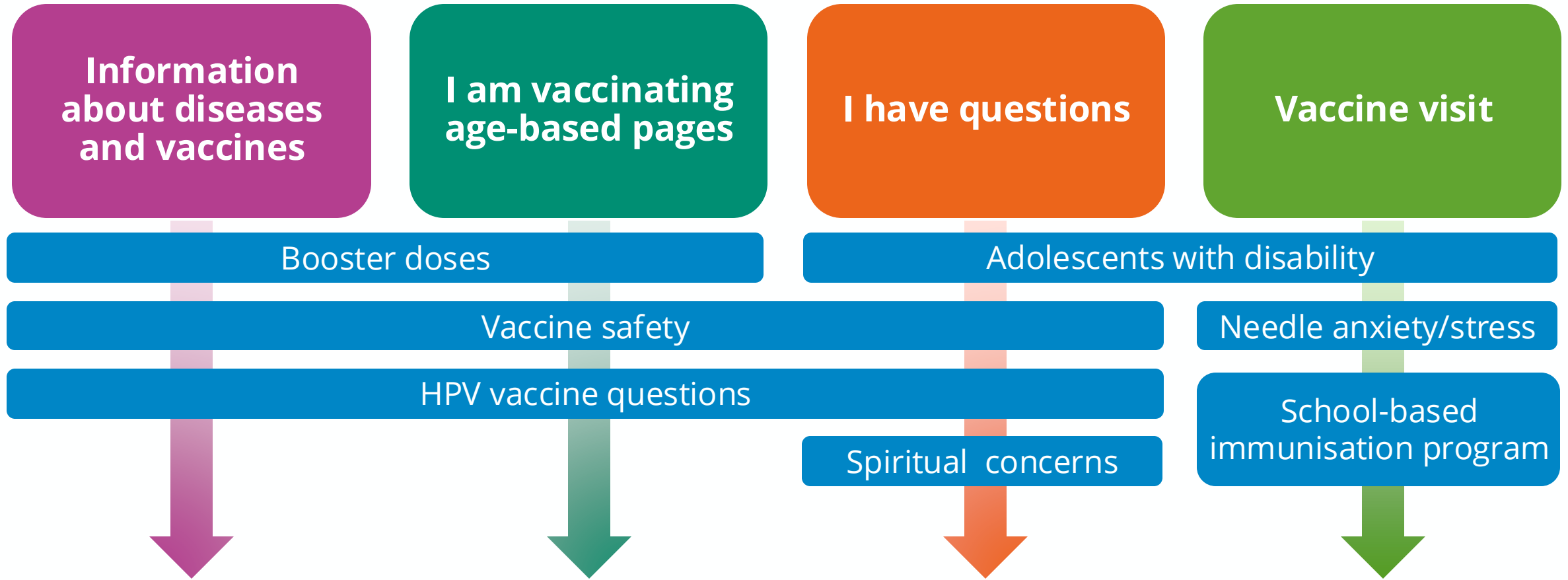
SKAI Adolescent: empowering conversations about adolescent immunisation



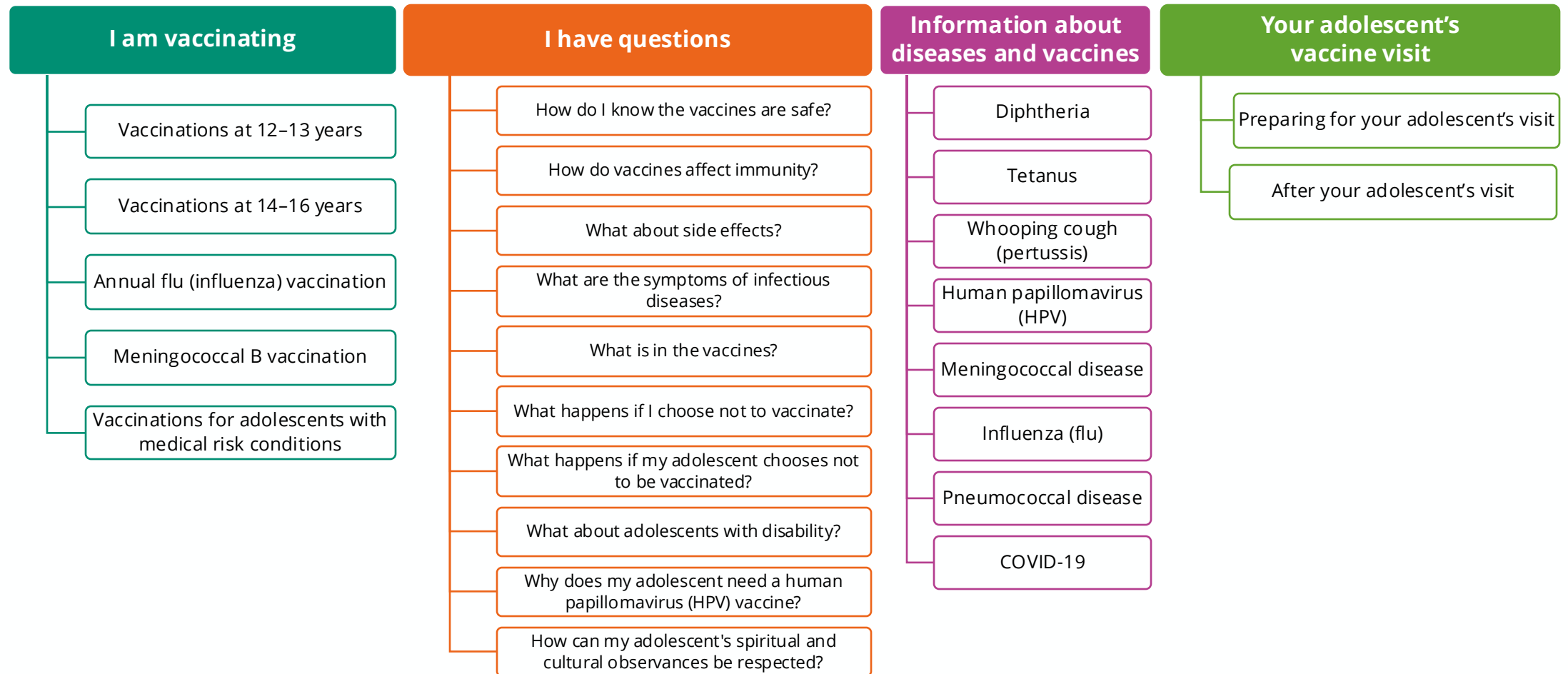
Providing practical guidance and answers to common questions related to:

- human papillomavirus (HPV) vaccination
- diphtheria-tetanus-pertussis (dTpa)m booster vaccination
- meningococcal B and meningococcal ACWY vaccination
- influenza vaccination
- vaccination for adolescents with medical risk factors
- vaccination for adolescents with disability

SKAI Adolescent: meeting the need for quality vaccination information



SKAI Adolescent: website content map



Novel strategies to improve uptake of vaccines in young people



THE UNIVERSITY
of ADELAIDE

NEW YORK TIMES BESTSELLER



MORE THAN
750,000
COPIES SOLD

Nudge

Improving Decisions About
Health, Wealth, and Happiness

Richard H. Thaler and Cass R. Sunstein

Revised and Expanded Edition

"One of the few books I've read recently that fundamentally changes the way
I think about the world." —Steven D. Levitt, coauthor of *Freakonomics*



Richard Thaler
Professor of Economics
University of Chicago
Behavioural Insight Team, UK
Cabinet Office



Cass Sunstein
Professor of Law
Harvard University
Regulatory Czar, Obama
Administration

Nudge is “any aspect of the choice architecture that *alters people’s behaviour in a predictable way* without forbidding any options or significantly changing their economic incentives”.

Choice architecture is the environments within which people make choices.

Consider this...



Schiphol Airport,
Amsterdam
80% decline
in 'spillage'

And this...



Lake Shore Drive
Chicago
36% fewer crashes

CHICAGO



GSB

The University of Chicago Graduate School of Business



The 9 general MINDSPACE principles of human behaviour change allow us to influence people



Messenger

We are heavily influenced by who communicates information to us



Incentives

Our responses to incentives are shaped by predictable mental shortcuts e.g. strongly avoiding losses



Norms

We are strongly influenced by what others do



Defaults

We go with the flow of pre-set options



Salience

Our attention is drawn to what is novel and what seems relevant to us



Priming

Our acts are often influenced by subconscious cues



Affect

Our emotional associations can powerfully shape our actions



Commitment

We seek to be consistent with our public promises, and reciprocate acts



Ego

We act in ways that make us feel better about ourselves

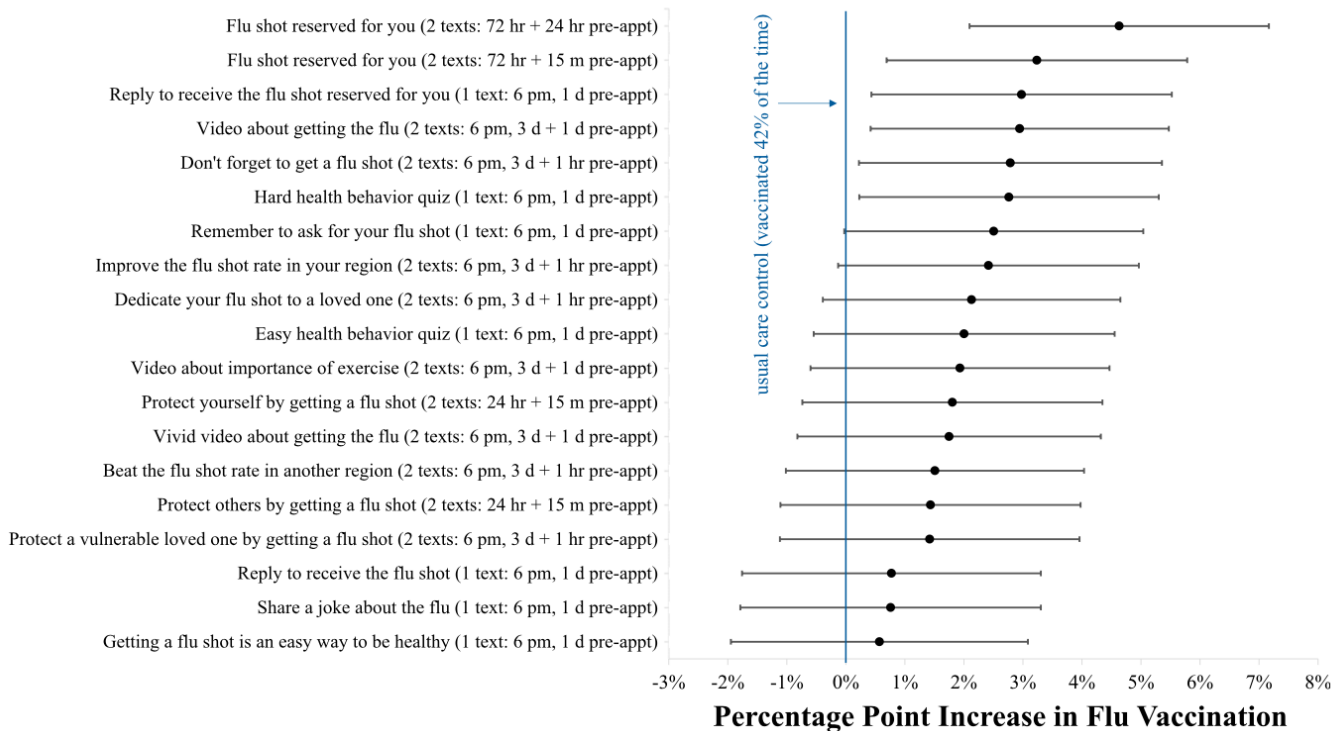
Incentives



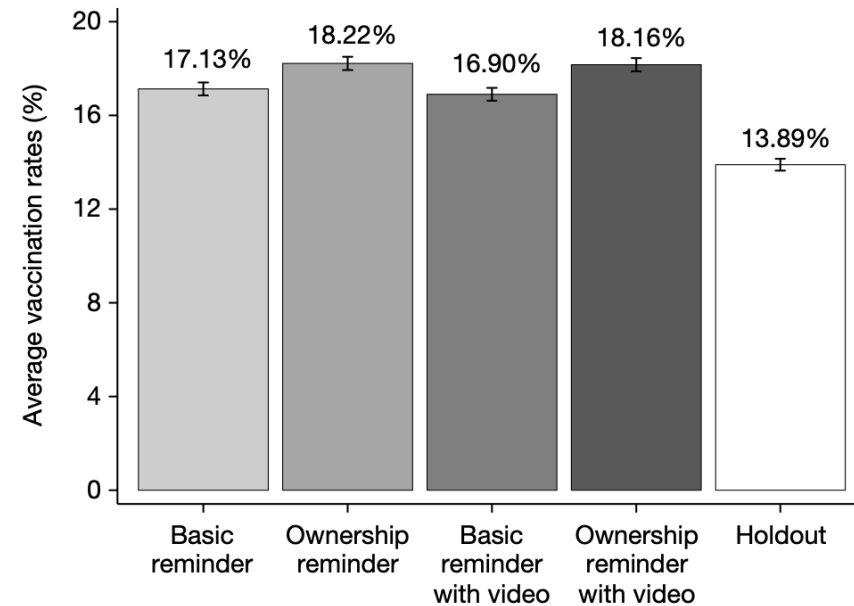
The value of an incentive depends on where we see it from – its reference point

Psychological Ownership nudges increase flu and COVID-19 vaccinations

N=47,306, 19 different text messages



UCLA Health: John Smith, a COVID-19 vaccine has just been made available to you at UCLA Health. Claim your dose today by making a vaccination appointment here: uclahealth.org/schedule vs you can get the COVID vaccine at UCLA health



FluText4U nudge study

- Assess whether a SMS intervention could increase influenza vaccine uptake in children with chronic medical conditions
- SMS sent 1 week prior to WCH appointment, 1 week after appointment and 2 weeks after the 2nd me

Influenza may be serious for children with medical conditions. The Women's and Children's Hospital recommends the influenza vaccine for all medically at-risk children.

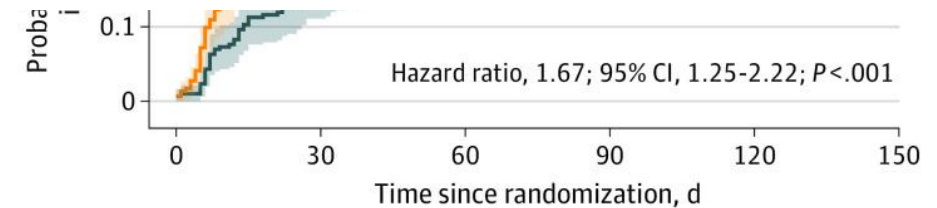
[name, lastname] is recommended to receive the influenza vaccine.

Table 2a Effect of intervention on receipt of COVID-19 vaccine within 3 months of randomization.

Variable	Intervention	Standard care	Odds ratio (95% CI)	P-value	Risk ratio (95% CI)	Absolute risk difference (95% CI)
COVID-19 vaccine received within 3 months of randomization, N (%)	8/554 (1.4)	9/551 (1.6)	0.89 (0.34, 2.35)	.82	0.90 (0.35, 2.29)	-0.17 (-1.61, 1.27)

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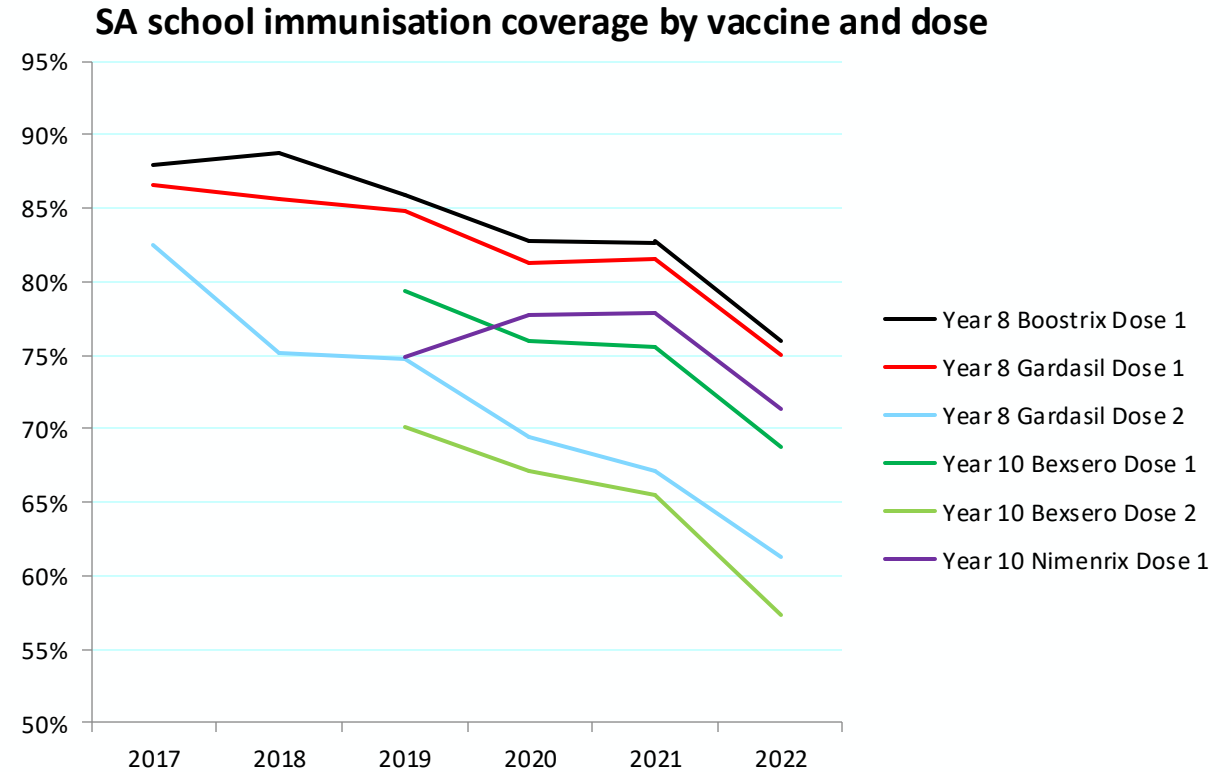
- A greater proportion receiving the flutext-4U were vaccinated during the optimal period (April-June) [48% vs 34%; aOR=1.97 (95%CI 1.28, 3.06 P<0.001)]



No. at risk	0	30	60	90	120	150
Control	302	256	216	163	101	54
SMS intervention	293	209	184	127	79	33

Nudges to increase uptake of vaccines delivered to young people

- Co-design a nudge intervention with students at “low uptake” schools
- Implement the nudge in a cluster randomised controlled trial in schools



Slide courtesy of Noel Lally, Executive Director, CDC, SA Health

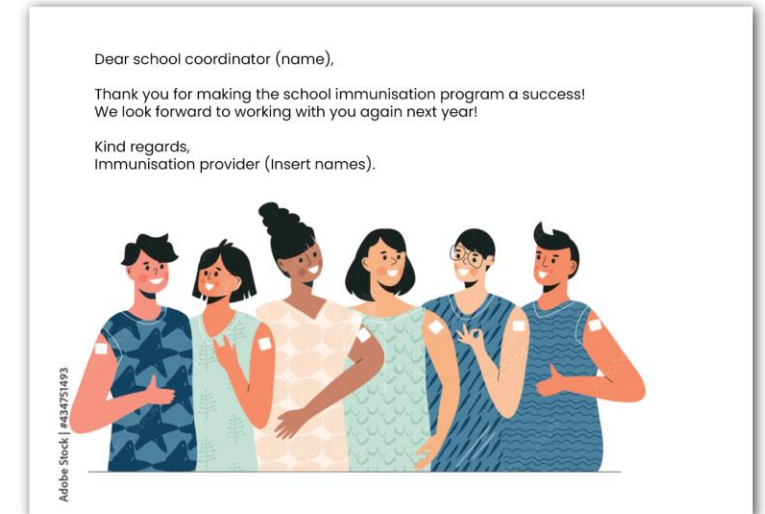
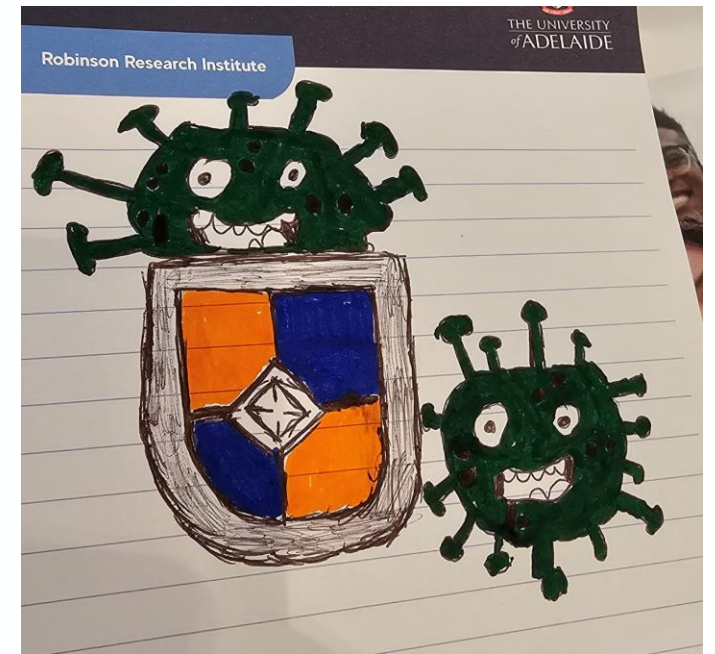
Co-designed nudge solutions



- ✓ Nudgeathon Workshop (Dec 2024)
- ✓ Framework: MINDSPACE principles
- ✓ 58 Participants:
 - Consumers: Year 7 and 10 students and their parents, representing a range of ethnicities and culturally and linguistically diverse backgrounds.
 - Key stakeholders: staff members from SA Health, Department for Education, private vaccination and health service providers, local councils and school
 - academia
 - psychologist
 - graph designer
- ✓ Evaluation: APEASE criteria¹ (Affordability, Practicability, Effectiveness/Cost-Effectiveness, Acceptability, Side-Effects/Safety, and Equity)

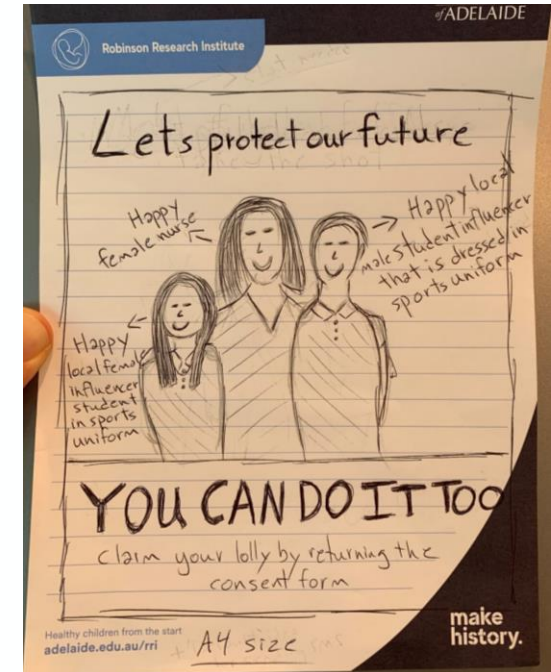
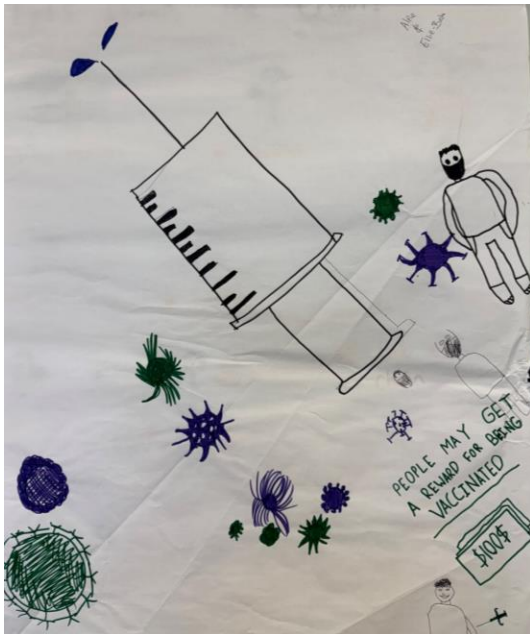
Teacher-Focused Nudges

- Checklist & Commitment:
 - ✓ Teachers receive a letter and checklist, encouraging them to support consent form distribution and collection.
 - ✓ Commitment signing and recognition (e.g. school-wide prize for highest return rate).
- Posters & Reminders:
 - ✓ Visual prompts in staff rooms
 - ✓ SMS/email reminders
- Comparison Data:
 - ✓ Share return rates across schools to stimulate motivation through friendly competition.
- Ego Appeal:
 - ✓ “Thank you” e-cards to acknowledge teacher efforts.

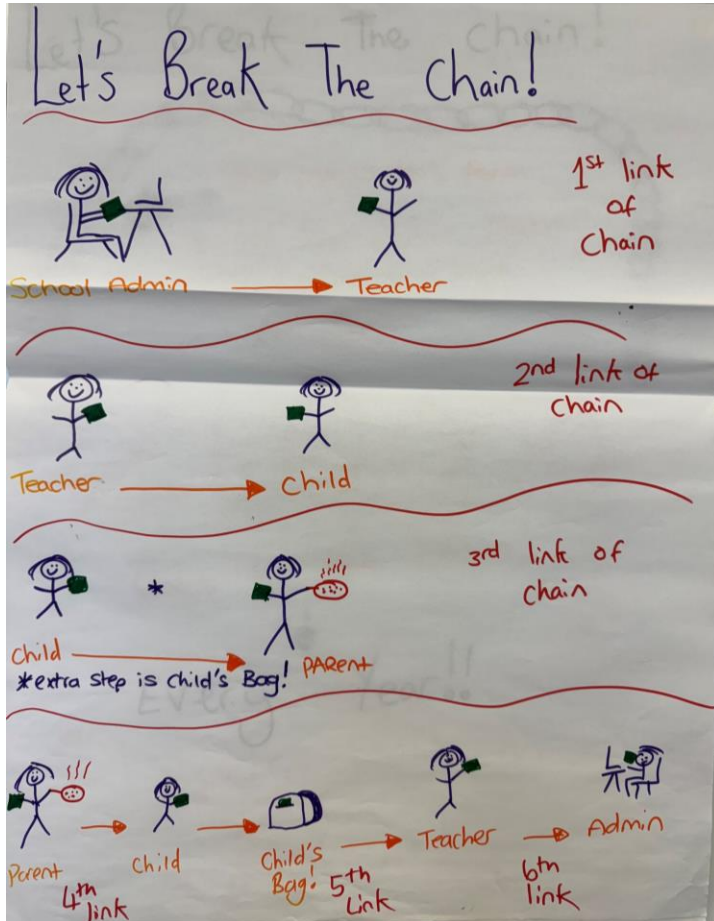


Student-Focused Nudges

- Immunisation Day = Positive Day:
 - ✓ Make vaccination day engaging (casual clothes, games, treats, competitions, etc.)
- Peer Norms & Social Proof:
 - ✓ Showcase videos with peer messengers and highlight group success (e.g. class pizza party if >85% vaccinated)
- Reminders:
 - ✓ Bright flyers, school app messages, and SMS reminders to parents
- Student-friendly forms:
 - ✓ visuals, slogans like "Protect Your Future"



System-Level Nudges



➤ Streamlined Consent Process:

- ✓ Normalise consent as a routine school activity
- ✓ Include immunisation consent forms in school enrolment packs
- ✓ Send reminders or updates before year 7 & 10 vaccinations based on prior consent.
- ✓ Explore default consent models requiring legal review & transparency
- ✓ Priority scheduling: block immunisation days in calendars

➤ Curriculum Integration:

- ✓ Include immunisation education in health or science classes

➤ Centralised Coordination:

- ✓ Improve school-provider coordination through calendar reminders, clearer timelines, and reduced form handling.



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By 2030, the NEIS CRE will deliver a roadmap to eliminate meningitis and drive down gonorrhoea globally