S1 Supplementary Material: Interview guides

The interviews were guided by a series of questions that were sent to Experts prior to the call. Questions differed slightly for DOH/MOH staff versus WHO/ECDC staff. Calls did not strictly follow the questions and interviewees were encouraged to elaborate and raise other issues, as appropriate.

Proposed questions for interviews with jurisdictions/country MOH staff:

Past pandemic plans:

- 1. What was the strategic objective of vaccine use prior to/during the 2009 pandemic? ie transmission reduction, mitigation, protect the health workforce (or multiple aims?)
- 2. Given (1), was there a prioritisation schedule prior to/during the 2009 pandemic? If yes,
	- a. Was it followed?
		- i. If not, why not? E.g. change in strategy needed in real time, issues with availability, lack of public/practitioner engagement
	- b. Was prioritization ranked?
	- c. Was prioritization dependent on any factors associated with risk; e.g. if determined that elderly would not be affected, were the no longer a priority group?
	- d. Was there a rationale for prioritization?
	- e. Was it evaluated?
		- i. Did that result in specific changes at the time, or in subsequent preparedness planning?
		- ii. What worked best?
		- iii. What didn't work?

Current plans:

- 3. Is there a new prioritization plan? Or is one being planned?
	- a. Is it a ranked list?
	- b. Is there a rationale for the ranking?
	- c. Were practical considerations taken into account?
	- d. Were these recommendations evidence based? i.e. based on published references, simulations, ranking of the materials, modelling, etc.
	- e. Does the plan include a definition for essential personnel?
	- f. Who were the key stakeholders that contributed to the list?
	- g. Has a human ethics committee been involved in the development of the list?
- 4. Has the country ever prioritised delivery of seasonal vaccine?
	- a. E.g. by age, or risk group?
- 5. Are there any specific documents published by your DOH that you can share or we should be aware of?
- 6. Availability for a follow-up call with ADOH

Proposed questions for interviews with ECDC/WHO:

Pre 2009 preparedness planning:

- 1. Prior to 2009, did WHO/ECDC have a suggested prioritisation schedule for vaccination?
	- a. How did it relate to the strategic objectives of vaccination?
	- b. Was prioritization ranked?
	- c. Was there a rationale for prioritization?

Evolution of recommendations during pandemic response:

- 2. How did evolving experience of the 2009 pandemic influence these recommendations?
	- a. Did the strategic objective of immunisation change?
	- b. Did the priority groups change?
	- c. What committees/stakeholders were involved in reviewing evidence and making recommendations?
	- d. Was the prioritization evaluated? (formally or informally)
		- i. What were the outcomes?

Current pandemic plan:

- 3. Is there a new set of recommendations for prioritising groups for vaccination? Or is one being planned?
	- a. What are the strategic objectives of immunisation as part of the response?
	- b. Is there a ranked list of groups for immunisation?
	- c. Is there a rationale for the ranking?
	- d. Were practical considerations taken into account?
	- e. Were these recommendations evidence based? i.e. based on published references, simulations, ranking of the materials, etc.
	- f. Does the plan include a definition for essential personnel?
	- g. Who were the key stakeholders that contributed to the list?
	- h. Has a human ethics committee been involved in the development of the list?
- 4. Are there any specific documents that we should be aware of?
- 5. Availability for a follow-up call with ADOH

S2 Supplementary material: mathematical model

S2.1 Model of influenza transmission

The transmission model is based on a classic susceptible-exposed-infectious-recovered (SEIR) paradigm. All individuals are assumed fully susceptible (S) at the outset of the epidemic, and vulnerable to acquiring infection (E) upon contact with an infectious (I) case. Once recovered (R), individuals are assumed to be fully resistant to reinfection. At the time the simulations commence, there are already 100 prevalent infections in the population. There are 50 exposed individuals and 50 infectious individuals, distributed across the population strata in proportion to each stratum's population size.

The model incorporates a dynamic "contact" label, applied to a fixed number of individuals drawn from the whole population each time a new infectious case appears. We define these contacts, based on the findings of sociological studies, as those people who have been sufficiently close to an infected individual to conceivable contract infection. Only contacts of an infectious case may proceed to the exposed and infectious classes, however the majority of contacts escape unscathed, returning to their original state within 72 hours of exposure.

This model has been described in detail in previous publications[[7,](#page-10-0) [8\]](#page-10-1); here we describe the key modifications and features that were developed and/or used in this study.

S2.1.1 Population stratification and mixing

The Australian population is stratified into 23 distinct sub-groups, agreed in consultation with the Office of Health Protection, with sizes based on June 2014 statistics from the Australian Bureau of Statistics. Population sub-groups are defined in terms of age groups and by Indigenous status, with further sub-categories defined in terms of location (urban/remote), risk factors (pregnant women and adults with comorbidities), and essential workforce (see [Table S1\)](#page-3-0).

Each sub-group can be targeted independently for vaccination; subsequent seroconversion (two weeks post-vaccination) reduces susceptibility to infection, but does not modify the clinical course of disease in the case that a vaccinated individual becomes infected.

In previous iterations of this model, the entire population has mixed homogeneously; population sub-groups were defined in order to support differential risk factors, targeted interventions, and reporting of burden in sub-populations. In this version of the model we incorporated inhomogeneous mixing to facilitate (a) intensive mixing between young children; and (b) intensive mixing between indigenous groups. We assumed that 67% of the contacts of primary-school children were also primary-school children, and that 80% of the contacts of an Indigenous person were also Indigenous.

S2.1.2 Differential risk factors

We have previously defined the risks of ICU admission and death for hospitalised cases *in the absence of treatment*; these values are shown in [Table S2](#page-3-1). These values were obtained

Age:	$0 - 4$	$5 - 9$	$10 - 12$	$13 - 18$	$19 - 65$	$66+$
Indigenous:						
Urban						
Remote						
Non-Indigenous:						
General						
Pregnant						
Co-morbidities						
Healthcare						
Emergency Services						
Infrastructure						
Low mixing			Moderate mixing		High mixing	
Low risk of hospitalisation				High risk		

Table S1: Population strata, showing variation in mixing and in risk of severe disease.

	High-Risk Children Others		
ICU admission	39.5%	14.4\%	14.4\%
Deaths (in ICU)	94.9%	46.1%	46.1%

Table S2: Risks of ICU admission and death, given hospital admission.

by using risk ratios for total influenza-related complications — 0.74 for otherwise healthy patients and 0.37 for high-risk patients $[2]$ $[2]$ $[2]$ — and calculating the counter-factual risks of ICU admission and death.

We have assumed that (a) all Indigenous strata have the same risks as the "High-Risk" group in [Table S2](#page-3-1); (b) pregnant women and other adults with co-morbidities have the same risks as the "High-Risk" group in [Table S2](#page-3-1); and (c) that all other strata (non-Indigenous, no co-morbidities) have the same risks as the "Children" and "Others" groups in [Table S2](#page-3-1).

We have also assumed that all Indigenous strata, pregnant women, and other adults with co-morbidities, are five times more likely to require hospitalisation, given infection, than the general population.

S2.1.3 Vaccination

The model simulations consider a vaccine that becomes available at 6 weeks after the start of the pandemic, and that batches of 3.4 million doses are delivered at 6-week intervals starting from this time. Consistent with previous studies, we assume that:

- 5% of vaccine doses are "wasted" and cannot be used to immunise individuals.
- 30% of vaccinated individuals do not seroconvert and receive no benefit from vaccination.
- The maximum rate of vaccine provision is 750,00 doses per week.

In this model, either one dose or two doses are required to confer full protection (i.e., to transition to the S_V compartment) and this requirement can differ for each stratum. In order to represent individuals with partial protection, by virtue having received one dose when two doses are required for full protection, we introduced a new compartment S_{PV} . To confer complete protection to such individuals, two doses are simultaneously consumed from the vaccine stockpile and the individual is moved to the S_{PV} compartment, which has a mean residence time of 2 weeks (after which they transition to S_V).

S2.2 Targeted vaccination strategies

We investigated the impact of two vaccination strategies that prioritised different subgroups for vaccination. The **Direct Protection** strategy prioritised the provision of vaccine to sub-groups with a disproportionate risk of severe disease, and involved:

- 1. Provision of vaccine to 300,000 essential services workers, distributed evenly across healthcare, emergency services, and infrastructure.
- 2. Provision of vaccine to all High-Risk groups, until an intermediate coverage target of 25% was achieved in these groups.
- 3. Provision of vaccine to all groups, until an ultimate coverage target of 50% in High-Risk groups and 25% in all other groups was achieved.

The **Indirect Protection** strategy prioritised the provision of vaccine to sub-groups that drive the transmission, and involved:

- 1. Provision of vaccine to 300,000 essential services workers, distributed evenly across healthcare, emergency services, and infrastructure.
- 2. Provision of vaccine to primary-school children (aged 5–12), until a coverage target of 50% was achieved in these groups.
- 3. Provision of vaccine to all groups, until an ultimate coverage target of 50% in High-Risk groups and 25% in all other groups was achieved.

Table S3: The capacity constraints for each healthcare setting.

S2.3 Measures of impact

Model outputs include the rate of infection, disease, severe disease, and death, for each population sub-group. By comparing these results against the same scenarios, but in the absence of any vaccine, we reported the following measures of population impact:

- Reduction in infections;
- Reduction in clinical presentations:
- Reduction in hospital (general ward) admissions;
- Reduction in ICU admissions; and
- Reduction in observed deaths (in ICUs);

The transmission model described above accounts for the spread of pandemic influenza in the Australian population and the population-level effects of vaccination. But it does not model the clinical pathways and inpatient capacities of the Australian healthcare system. These aspects of the pandemic influenza scenarios are instead captured by a separate model, which uses daily incidence of mild and severe presentations (as generated by the transmission model) to determine the available capabilities for inpatient admission, outpatient consultation, and patient treatment, and reports outcomes such as the peak and excess burden on each healthcare setting [\[7\]](#page-10-0).

For each pandemic scenario, the transmission model reports the daily number of mild and severe presentations, where "severe cases" are those that will require hospitalisation. All mild presentations are assumed to occur in outpatient settings (i.e., general practice clinics and hospital emergency departments). In addition, we assume that some fraction of the severe cases presents to an outpatient facility prior to requiring hospitalisation.

Patients are admitted to general wards with a mean length of stay of 5 days, and are admitted to ICUs with a mean length of stay of 10 days. Therefore, it is the prevalence of cases requiring hospitalisation that determines the available ward and ICU bed capacities for new admissions. The capacity constraints for each healthcare setting are shown in [Table S3.](#page-5-0)

Transmission R_0		Clinical Severity η		Mean α_m Mean CAR
Low Medium High	$1.05 - 1.20$ High $1.40 - 1.70$ High	$1.20-1.40$ Moderate	$10^{-2} - 10^{-1}$ 29.8% $10^{-3} - 10^{-2}$ 11.6\% $10^{-2} - 10^{-1}$ 29.8%	5.7% 4.8% 17.1%

Table S4: Pandemic influenza scenarios; note that low-transmissibility represents lowlevel epidemic activity rather than "sporadic cases". The mean presenting proportion (α_m) and the mean Clinical Attack Rate (CAR) summarise some of the key differences between these scenarios.

S2.4 Pandemic influenza scenarios

As per previous consultancies for the Commonwealth Office of Health Protection [[7](#page-10-0)] and as used in the Australian Health Management Plan for Pandemic Influenza [[1](#page-9-1)], pandemic scenarios were classified by pandemic impact. This comprises both transmission (R_0) and clinical severity $(\eta, \, \text{the proportion of (adult) infections that are sufficiently severe})$ to require hospitalisation), as summarised in [Table S4](#page-6-0).

We allowed the proportion of adult infections that require hospitalisation to vary from 1 in 1,000 ($\eta = 10^{-3}$, moderate clinical severity) to 1 in 10 ($\eta = 10^{-1}$, high clinical severity). Within each scenario, uncertainty on the time-course of the epidemic, effectiveness of vaccination, and other biological and epidemiological co-variates was accounted for through the use of Latin Hypercube Sampling, in which 10,000 simulations of the model were run, each with randomly assigned parameterisations[[7\]](#page-10-0). For example, the proportion α_m of all infections that "present" (i.e., are visible in the health system) was determined through a combination of the transmissibility (R_0) and the clinical severity (η) , and varied from 5% to 75%.

S2.5 Supplementary figures

Figure S1: The effect of vaccine availability on vaccine impact. The greatest impact is observed when vaccine is immediately available ("0 weeks"), but the relative impact of a 6-week and a 12-week delay vary across these pandemic scenarios. Note that the y-axis scale for the Low Transmission / High Severity scenario is markedly different from that of the other scenarios.

Figure S2: The hospital admissions as a result of each vaccination strategy, shown for population sub-groups as median estimates (points) and the 5th and 95th percentiles (lines). Note that the y-axis scale for the Low Transmission / High Severity scenario is markedly different from that of the other scenarios.

S2.6 Modifications to suit COVID-19

The model described here, which incorporated a number assumptions regarding pandemic influenza, was subsequently tailored to COVID-19 to inform the COVID-19 pandemic response in Australia [\[9](#page-10-2)]. This involved the following modifications:

- 1. Increasing the latent period to 3.2 days, assuming 2 days of pre-symptomatic transmission before completion of incubation period, based on estimates from Ganyani et al. [\[3](#page-9-2)] and Tindal et al. [\[11\]](#page-10-3).
- 2. Increasing the infectious period to 9.68 days, based on a doubling time of 6.4 days in Wuhan, China [\[12\]](#page-10-4) and an incubation period of 5.2 days [[4](#page-9-3), [6](#page-9-4)].
- 3. A higher value for R_0 than the high severity scenario used here $(1.4 \text{ to } 1.7)$, based on the latent and infectious periods above, and a doubling time of 6.4 days.

Further modifications to this model are required to consider COVID-19 vaccination:

- 1. The age-specific hospitalisation and mortality rates should be altered to reflect country-specific values for COVID-19.
- 2. Where relevant to the local context, contact rates should be reduced to reflect population compliance with physical distancing measures [[5,](#page-9-5) [10](#page-10-5)].
- 3. The vaccine effectiveness estimate would need to be updated to reflect candidate COVID-19 vaccine estimates. Recently-published estimates for several candidate vaccines range from 70% to 95%, and it appears likely that 2 doses will be required.
- 4. Our pandemic scenarios considered a vaccine that became available soon after the onset of an epidemic wave in a fully-susceptible population. For countries that have successfully limited local transmission of COVID-19 to date, these timelines remain relevant because the majority of the population in these countries remain susceptible to COVID-19. For countries that have experienced substantial local transmission of COVID-19, and where a substantial proportion of the population are likely to have some degree of protective immunity, the initial model population state should be adapted accordingly.

S2.7 References

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S Supplementary material: Experience of pandemic vaccine use in 2009

Multinational Level

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