

Influenza vaccines

INFLUENZA VACCINES FOR AUSTRALIANS: INFORMATION FOR IMMUNISATION PROVIDERS

This fact sheet provides information for immunisation providers on seasonal influenza vaccines that are available in Australia in 2018. It can be used in conjunction with the NCIRS fact sheet [Influenza vaccines – frequently asked questions](#) which provides responses to common questions about influenza viruses and seasonal influenza vaccines.

- Influenza remains a common cause of hospitalisation and death in Australia.
- Annual influenza vaccination is recommended for all people aged ≥ 6 months.
- Annual influenza vaccination is funded under the National Immunisation Program (NIP) for people aged ≥ 6 months who are at increased risk of severe influenza, including:
 - Aboriginal and/or Torres Strait Islander people aged 6 months to < 5 years and ≥ 15 years
 - All adults aged ≥ 65 years
 - People with specified medical conditions (refer to [Table 1](#))
 - Pregnant women (during any stage of pregnancy)
- Most states and territories (WA, QLD, NSW, ACT, VIC and TAS) also provide free influenza vaccine for all children aged 6 months to < 5 years.
- The strains used in seasonal influenza vaccines can change from year to year depending on which viruses are predicted to circulate in each upcoming season.
- In 2018, in addition to the quadrivalent influenza vaccines (QIVs), two higher-immunogenicity trivalent influenza vaccines ([TIVs]; one 'high-dose' vaccine and another containing an adjuvant) are funded under the NIP for adults aged ≥ 65 years.
- The 'high-dose' TIV is estimated to be approximately 24% more effective against laboratory-confirmed influenza than the standard TIV in adults aged ≥ 65 years. The adjuvanted TIV is estimated to be approximately 25% more effective against hospitalisation for influenza or pneumonia than the standard TIV.
- Providers are reminded that influenza vaccinations given to people of all ages should be reported to the Australian Immunisation Register (AIR).
- Providers are also reminded to vaccinate pregnant women at any time of the year and any stage of pregnancy. Vaccination of pregnant women provides protection to mothers and their newborn infants.

The Australian Technical Advisory Group on Immunisation (ATAGI) publishes annual advice on the use of influenza vaccines in Australia:

<https://beta.health.gov.au/resources/publications/atagi-advice-on-seasonal-influenza-vaccines-in-2018>

The disease

Influenza or ‘the flu’ is an acute viral illness that mainly affects the respiratory system.

Causative agent

Influenza is caused by influenza viruses which are classified as type A, B or C.¹ Only influenza A and B viruses are included in seasonal influenza vaccines as they cause the majority of disease in humans. Type A influenza viruses are further categorised into subtypes according to two kinds of proteins on their surface: haemagglutinin (H) and neuraminidase (N).¹ Type B influenza viruses are categorised into two lineages (Yamagata and Victoria). Both influenza A and B can be further broken down into different strains.

The genes for the H and N proteins on the virus surface mutate frequently, which results in constant change to influenza viruses. These minor changes to the H and N proteins of both influenza A and B are referred to as ‘antigenic drift’ and result in new virus strains. Antibody cross-protection against drifted strains is likely to be reduced. If a major change happens in the H or N protein of influenza A, it is called ‘antigenic shift’. Previous immunity is usually not adequate against disease from a ‘shifted’ strain. This creates the potential for a pandemic.

Transmission

Influenza is spread easily, mainly through large particle droplets produced by sneezing and coughing.^{1,2} Droplets containing the influenza virus also settle onto surfaces, and can then pass from hands to the nose, mouth or eyes. People with influenza can be infectious to others from 24 hours before symptoms start until 1 week after the start of symptoms. In previously healthy individuals, symptoms typically subside within 5–8 days.

People of all ages are susceptible to influenza. The percentage of people in the general community affected by flu each year is typically 5–10%, but may be up to 20% in some years. This percentage is higher for children, with 10–40% infected each year.^{1–3} Influenza is more easily spread where large numbers of people gather together.¹ As such, infection rates may be 2–3 times higher in closed populations (e.g. childcare centres, aged care facilities and households).^{4,5}

Clinical features

Influenza symptoms usually have a sudden onset. The most common symptoms are fever, dry non-productive cough, nasal congestion, headache, sore throat and constitutional complaints such as myalgia, malaise and fatigue. The elderly may present with atypical symptoms such as malaise and confusion, and more often develop pulmonary complications. Non-respiratory symptoms

such as gastrointestinal complaints and calf muscle pain occur more frequently in children than in adults.^{1,3}

Although most influenza infections are symptomatically worse and more severe than other viral upper respiratory tract infections, some may be mild.^{1–3} Serious complications from influenza occur in a small proportion of people who are infected.^{1–3} Complications include pneumonia, myocarditis and neurologic complications, which can lead to hospitalisation and death. People at highest risk of complications from influenza include those with pre-existing medical conditions. However, previously healthy people can also have severe complications.

Diagnosis

Laboratory tests are required to confirm an influenza infection. The virus can be detected in a nose or throat swab by rapid antigen-based tests, viral culture or more commonly by molecular methods, such as polymerase chain reaction (PCR). Serological diagnosis can be established by measuring antibodies in acute and convalescent blood specimens.

Treatment

Treatment of influenza, including bed rest, pain relief such as aspirin/paracetamol and fluid intake, generally aims to prevent or minimise symptoms.^{1–3} Children <16 years of age must not be given aspirin or aspirin-containing medications while sick with influenza because of the increased risk of developing Reye syndrome, a condition that causes swelling in the liver and brain.

Antiviral medication, which requires a prescription, can help reduce the severity and duration of symptoms of influenza. To be most effective, it needs to be administered within 48 hours of symptom onset.¹

Prevention

Vaccination is the only way to specifically prevent influenza infection and its complications (refer to [Who should be vaccinated](#)).

Practising cough etiquette (such as covering the nose and mouth with a tissue when coughing or sneezing) and washing hands before eating can help reduce the likelihood of transmitting and contracting the influenza virus. Anyone who is unwell with influenza should stay home from work, school and social gatherings to prevent close contact with other people which could lead to transmission of the virus.^{1–3,6}

Who should be vaccinated

Annual influenza vaccination is recommended for all people aged ≥ 6 months unless contraindicated (refer to [Contraindications](#)).

There are a number of groups who are at increased risk of influenza and its complications and so annual influenza vaccination is strongly recommended for these groups. For some of these groups, seasonal influenza vaccination is provided free of charge through the National Immunisation Program (NIP) on the basis of demonstrated cost-effectiveness as a public health intervention.⁷ However, annual influenza vaccine should be actively promoted for all individuals at increased risk of severe complications from influenza, regardless of eligibility for a free vaccine.

Influenza vaccination is strongly recommended and funded on the NIP for the following groups:

- Aboriginal and/or Torres Strait Islander people aged 6 months to <5 years or ≥15 years
- All adults aged ≥65 years
- All people aged ≥6 months with a medical condition that increases the risk of influenza complications
- Pregnant women (during any stage of pregnancy)

The medical conditions that are associated with an increased risk of influenza complications are summarised in [Table 1](#) for easy reference.

Influenza vaccination is strongly recommended but not funded on the NIP for the following groups:

- Children aged 6 months to <5 years – NOTE, most states and territories (WA, QLD, NSW, ACT, VIC and TAS) now provide vaccine for free for all children in this age group
- Aboriginal and/or Torres Strait Islander children aged 5 years to <15 years
- Individuals with certain medical conditions (in addition to those funded on the NIP):
 - Down syndrome
 - class III obesity (body mass index ≥40 kg/m²)
 - chronic liver disease
- Residents and staff (including volunteers) of aged care and long-term residential care facilities
- Homeless people
- Carers and household contacts of those in high-risk groups
- Commercial poultry or pork industry workers
- Essential services providers
- Travellers

Detailed information on influenza vaccine recommendations is provided in *The Australian Immunisation Handbook* (refer to [Additional resources for primary medical care/vaccination providers](#)).

Contraindications

The only absolute contraindications to influenza vaccines are:

- anaphylaxis after a previous dose of any influenza vaccine
- anaphylaxis after any vaccine component
- people allergic or sensitive to latex should not receive Flud. All other influenza vaccines used in Australia are safe for people who are allergic or sensitive to latex.

Note: Egg allergy is not a contraindication to influenza vaccine. Persons with egg allergy, including anaphylaxis, can be safely vaccinated with influenza vaccines. Persons with a history of anaphylaxis to egg can be vaccinated with a full vaccine dose in medical facilities with staff experienced in recognising and treating anaphylaxis.

Vaccines

The 2018 southern hemisphere seasonal influenza vaccines contain:

- A (H1N1) – an A/Michigan/45/2015 (H1N1)pdm09-like virus
 - A (H3N2) – an A/Singapore/INFIMH-16-0019/2016 (H3N2)-like virus
 - B (Yamagata lineage) – a B/Phuket/3073/2013-like virus
 - B (Victoria lineage) – a B/Brisbane/60/2008-like virus (not included in the trivalent influenza vaccines [TIVs])
- Vaccines are registered on the basis of evidence of their effectiveness and safety (refer to [Supplementary information](#)). Multiple registered influenza vaccine products are available each year. The age group(s) in which each vaccine can be used and their NIP availability vary.

The available seasonal influenza vaccines by brand, recommended age and NIP availability are summarised in [Table 2](#) for easy reference.

More detailed information on seasonal influenza vaccines is provided in the *ATAGI advice for immunisation providers regarding the administration of seasonal influenza vaccines in 2018* (refer to [Additional resources for primary medical care/vaccination providers](#)).

Dosage and administration

The preferred route of administration for influenza vaccines is by intramuscular injection; however, they may also be given by the subcutaneous route. The recommended vaccine dose volume varies by age, and the number of vaccine doses varies by age and immune status of the vaccine recipient.

The recommended brand and dose of seasonal influenza vaccine by age group is summarised in [Table 3](#)

Although protection provided by influenza vaccine is generally expected to last for the whole season, optimal protection occurs within the first 3 to 4 months after vaccination.^{8,9} While influenza continues to circulate, it is never too late to vaccinate.

All influenza vaccines available in Australia may be co-administered with any other vaccine (refer to [Supplementary information, Safety in infants and children](#)). Detailed information on the administration of influenza vaccines, including co-administration and vaccine interchangeability, is provided in *The Australian Immunisation Handbook* (refer to [Additional resources for primary medical care/vaccination providers](#)).

Supplementary information

Epidemiology

Influenza is a seasonal disease in temperate regions. Most cases in Australia occur during the winter months between June and September.¹⁰ In the northern hemisphere, influenza usually occurs between December and April, whereas in the tropics, influenza can occur all year round.

Annual influenza epidemics are most often due to a single virus subtype or lineage. However, the circulating subtypes/lineages can vary year to year and different subtypes/lineages may appear sequentially or simultaneously in the same season.¹¹

Influenza is an important cause of morbidity and mortality. The number of affected people varies considerably from year to year depending on the characteristics of the circulating virus strains and the level of immunity in the population. It has long been recognised that the impact of influenza is often substantially under-estimated.^{6,12} Between 2006 and 2013 (excluding the 2009 pandemic year), an average of 100 deaths and approximately 5,100 hospitalisations due to influenza occurred annually in Australia.¹³ In the 2017 influenza season, the highest levels of activity since the 2009 pandemic year were recorded. Around 750 deaths were reported nationally among notified cases of laboratory-confirmed influenza.¹⁴ However, a mathematical modelling study estimated that influenza is actually associated with more than 3,000 deaths and 13,500 hospitalisations each year in Australia, just in people aged over 50 years.¹²

There are a number of groups who are at a higher risk of influenza and its complications and who experience increased morbidity and mortality associated with influenza compared to the rest of the population. The highest rates of influenza notifications and hospitalisations are seen in the elderly and children <5 years of age.¹³ Aboriginal and Torres Strait Islander people experience a greater disease burden from influenza than non-Indigenous Australians across all age groups.^{13,15} In addition, people with certain underlying medical conditions such as chronic heart, lung and neuromuscular disease, among others, are also at increased risk of severe influenza complications compared with otherwise healthy individuals.¹⁶ Pregnant women are more likely than other women to be hospitalised with influenza, and infants born to mothers who contract influenza during pregnancy are at risk of preterm birth and low birth weight.¹⁷

Vaccine effectiveness

Influenza vaccine provides varying levels of protection against influenza virus, depending on age, whether a person is immunocompromised and how good the match is between influenza strains in the vaccine and those circulating in the community.⁶ In a clinical trial among adults aged ≥ 65 years, the 'high-dose' TIV was estimated to be approximately 24% more effective against laboratory-confirmed influenza compared to standard TIV.¹⁸ In a large post-licensure study of community-dwelling adults aged ≥ 65 years, the adjuvanted TIV was estimated to be approximately 25% more effective against hospitalisation for influenza or pneumonia compared to standard TIV.¹⁹

A systematic review estimated the overall efficacy of standard TIV against laboratory-confirmed influenza in healthy adults <65 years of age to be 59%, although efficacy varied by influenza season.²⁰ Similar levels of protection have been achieved in young children, with an estimated vaccine effectiveness of 65% against laboratory-confirmed influenza in those aged 6 to 59 months.²¹⁻²³

Clinical trials of quadrivalent influenza vaccine (QIV), demonstrated equivalent antibody levels (an accepted surrogate for protection against influenza) to standard TIV for the shared strains in adults and children aged >6 months and added protection against the additional B strain.²⁴⁻²⁷

Vaccine safety

The common symptoms after influenza vaccination can mimic influenza infection, but are due to the vaccine's interaction with the immune system. The influenza vaccines currently registered do not contain live virus, so they cannot cause influenza.

Fever, headache, arthralgia and myalgia occur in <15% of those who receive influenza vaccine. Injection site reactions such as swelling, redness and pain are not uncommon. A higher rate of injection site reactions has been observed in clinical trials with the high-dose and adjuvanted TIVs registered for use in adults ≥ 65 years compared to that with standard TIVs.^{28,29} Around 30% of high-dose recipients reported injection site reactions compared to around 20% of standard dose recipients.²⁸ The majority of reactions were mild.²⁸ More injection site reactions in the week after vaccination were also seen among adjuvanted TIV recipients than those in non-adjuvanted TIV recipients (around 35% versus 18%).²⁹ Less than 1% of local reactions following either adjuvanted TIV or standard TIV were severe.²⁹ These side effects may commence within a few hours of vaccination and can last for 1–2 days.⁶

Surveillance of influenza vaccine safety through active enhanced surveillance systems such as AusVaxSafety showed that in 2017 across all ages low rates of any adverse event (6.6%) and medical attendance (0.4%) were reported after vaccination. In 2018, active surveillance by AusVaxSafety will be conducted in persons of all ages and data will be made available on a weekly basis at www.ausvaxsafety.org.au.

More severe immediate adverse effects such as hives, angioedema or anaphylaxis are rare consequences of influenza vaccination.^{6,30,31}

A small increased risk of Guillain-Barré syndrome (GBS) was associated historically with one influenza vaccine in the United States in 1976. But since then, close surveillance has shown that GBS has occurred at a very low rate of less than 1 in 1 million doses of influenza vaccine, if at all.^{32,33}

Safety in infants and children

Surveillance of influenza vaccine safety in young children through AusVaxSafety has shown that influenza vaccine is safe in children aged 6 months to <5 years, with low rates of fever (approximately 4%) and medical attendance (1%) reported after vaccination.^{34,35}

In young children, febrile convulsions are a relatively common response to fever of any cause, most often occurring among children 12–23 months of age.³⁶ Febrile convulsions related to influenza vaccination are uncommon, occurring at a rate of 1 per 1,000 or less in vaccinated individuals.^{37,38}

A slightly higher risk of fever and febrile convulsions in children aged 6 months to <5 years (especially those aged 12–24 months) has been reported following the concurrent administration of inactivated trivalent influenza vaccine and the 13-valent pneumococcal conjugate vaccine.³⁷ These vaccines are not routinely administered together in Australia and more recent data have not demonstrated the same association with febrile convulsions.³⁹ It is acceptable to administer these vaccines concurrently when both vaccines are indicated.⁴⁰

Safety in pregnant women

All influenza vaccines in Australia are inactivated vaccines, which can be safely given to pregnant women at any stage during pregnancy. The rate of adverse events after vaccination in pregnant women is no different from the rate in women who are not pregnant. In addition, studies of mother–baby pairs have shown that receiving the influenza vaccine while pregnant does not increase maternal or fetal complications during pregnancy.⁴¹

A number of high-quality studies have demonstrated that influenza vaccination during pregnancy provides protection not only to the mother but also to her newborn in the first few months of life when they are most vulnerable.^{42–46}

For further information, refer to the NCIRS fact sheet on [Vaccinations during pregnancy](#).

Additional resources for primary medical care/vaccination providers

- *The Australian Immunisation Handbook*, 10th edition – the most up-to-date clinical recommendations are contained in the online version of the *Handbook*
www.immunise.health.gov.au/internet/immunise/publishing.nsf/Content/Handbook10-home
- Immunise Australia website
www.immunise.health.gov.au
- Australian Technical Advisory Group on Immunisation (ATAGI) advice for immunisation providers regarding the administration of seasonal influenza vaccines in 2018
<https://beta.health.gov.au/resources/publications/atagi-advice-on-seasonal-influenza-vaccines-in-2018>

Table 1: Medical conditions that are associated with an increased risk of influenza complications and for which individuals are eligible for vaccination under the NIP*

(from ATAGI advice for immunisation providers regarding the administration of seasonal influenza vaccines in 2018 – refer to [Additional resources for primary medical care/vaccination providers](#))

| Category | Vaccination strongly recommended for (but not limited to) individuals with the following clinical conditions |
|---|---|
| Cardiac disease | Cyanotic congenital heart disease Congestive heart failure Coronary artery disease |
| Chronic respiratory conditions [†] | Severe asthma (for which frequent medical consultations or the use of multiple medications is required) Cystic fibrosis Bronchiectasis Suppurative lung disease Chronic obstructive pulmonary disease (COPD) Chronic emphysema |
| Chronic neurological conditions [†] | Hereditary and degenerative CNS diseases [†] (including multiple sclerosis) Seizure disorders Spinal cord injuries Neuromuscular disorders |
| Immunocompromising conditions [‡] | Immunocompromised due to disease or treatment (e.g. malignancy, transplantation and/or chronic steroid use) Asplenia or splenic dysfunction HIV infection |
| Diabetes and other metabolic disorders | Type 1 diabetes Type 2 diabetes Chronic metabolic disorders |
| Renal disease | Chronic renal failure |
| Haematological disorders | Haemoglobinopathies |
| Long-term aspirin therapy in children aged 6 months to 10 years | These children are at increased risk of Reye syndrome following influenza infection |

* **Note:** ATAGI also strongly recommends influenza vaccination for persons who have the following conditions (but vaccination is **not funded** under the NIP for such persons unless they also fall under one of the categories in the table above):

- Down syndrome
- obesity (class III), defined as body mass index (BMI) ≥ 40 kg/m²
- chronic liver disease (defined as histological evidence of fibrosis or cirrhosis, or clinical evidence of chronic liver disease).

Further details are provided in *The Australian Immunisation Handbook* (refer to [Additional resources for primary medical care/vaccination providers](#)).

[†] Persons who have any condition that compromises the management of respiratory secretions or is associated with an increased risk of aspiration should be vaccinated.

[‡] Persons with certain immunocompromising conditions (i.e. haematopoietic stem cell transplant, solid organ transplant) receiving influenza vaccine for the first time post transplant are recommended to receive 2 vaccine doses at least 4 weeks apart (irrespective of age) and 1 dose annually thereafter.

Table 2: Seasonal influenza vaccines available for use in Australia in the 2018 influenza season, by brand and recommended age

(from ATAGI advice for immunisation providers regarding the administration of seasonal influenza vaccines in 2018 [Table 1] – refer to [Additional resources for primary medical care/vaccination providers](#))

| Vaccine | Quadrivalent | | | | | Trivalent (for age ≥ 65 years only) | |
|---------------------------|---|----------------------------------|-----------------------------------|--------------------------------------|--------------------------------------|---|-------------------------------|
| | FluQuadri Junior 0.25 mL (Sanofi) | FluQuadri 0.50 mL (Sanofi) | Fluarix Tetra 0.50 mL (GSK) | Afluria Quad 0.50 mL (Seqirus) | Influvac Tetra 0.50 mL (Mylan) | Fluzone High-Dose 0.50 mL (Sanofi) | Fluad 0.50 mL (Seqirus) |
| <6 months | X | | | | | | |
| 6 to 35 months (<3 years) | ✓ | x | x | x | x | x | x |
| ≥ 3 to 17 years | x | ✓ | ✓ | x | x | x | x |

| | | | | | | | |
|-----------|---|---|---|---|---|---|---|
| ≥18 years | x | ✓ | ✓ | ✓ | ✓ | x | x |
| ≥65 years | x | ✓ | ✓ | ✓ | ✓ | ✓ | ✓ |

Table 3: Recommended doses of influenza vaccine by age

(adapted from the current online version of *The Australian Immunisation Handbook, 10th edition*— refer to [Additional resources for primary medical care/vaccination providers](#))

| Age | Dose (volume per dose) | Number of doses required | |
|-----------------------|------------------------|--|--|
| | | In the first year of influenza vaccination | If previously received any prior dose of influenza vaccine |
| 6 months to <3 years* | 0.25 mL [†] | 2 | 1 |
| ≥3 to <9 years* | 0.5 mL [‡] | 2 | 1 |
| ≥9 years | 0.5 mL [‡] | 1 [§] | 1 |

* Children aged 6 months to <9 years receiving influenza vaccine for the first time require 2 doses, at least 4 weeks apart, to maximise the immune response to the vaccine strains.

† If a child aged 6 months to <3 years inadvertently receives a 0.5 mL dose of influenza vaccine, no immediate action is necessary, and any additional dose(s) required that season or in future seasons should be given following standard recommendations. There is some evidence that a 0.5 mL dose of inactivated influenza vaccine is immunogenic and safe in children <3 years of age.^{25,47}

‡ If a child aged ≥3 years or an adult inadvertently receives a 0.25 mL dose of influenza vaccine, an age-appropriate dose (0.5 mL) should be repeated. Any additional dose(s) required that season or in future seasons should then be given following standard recommendations.

§ Two doses, at least 4 weeks apart, are recommended for persons with certain immunocompromising conditions (i.e. haematopoietic stem cell transplant or solid organ transplant) receiving influenza vaccine for the first time post transplant (irrespective of their age).

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