



Influenza vaccines – Frequently Asked Questions

This fact sheet provides answers to common questions about influenza viruses and available vaccines, including the new influenza vaccine programs in 2023. More detailed information can be found in the NCIRS fact sheet Influenza vaccines for Australians.

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Questions about influenza virus and influenza vaccines

Q1. What's the difference between influenza and the common cold?

Influenza is a respiratory illness that occurs after an infection with influenza viruses. Influenza is often referred to as 'the flu'. Sometimes the term 'the flu' is used incorrectly to describe the common cold, other respiratory viruses or even gastrointestinal illnesses. This is because their symptoms can be similar to those caused by influenza. There are many different viruses and some bacteria that can cause these symptoms. The influenza vaccine will only protect you from the influenza virus. The following table compares symptoms from the common cold and influenza. Usually influenza is more severe and lasts longer than a cold or other viral respiratory illness.

Cold	Symptom	Influenza
8	Fever	888
8	Headache	888
88	General aches and pains	888
88	Tired and weak	888
8	Extreme fatigue	888
888	Runny, stuffy nose	88
888	Sneezing	88
888	Sore throat	888
88	Chest discomfort, coughing	888

 $\alpha = \text{rarely}; \ \alpha \alpha = \text{sometimes}; \ \alpha \alpha \alpha = \text{often}$

Table adapted from: Immunize Canada - Is it a cold or influenza? Available from: www.immunize.ca/sites/default/files/resources/176e.pdf (Accessed February 2023)

Q2. Is it worth getting the influenza vaccine? I'm a healthy person and have heard that influenza isn't serious.

Yes, it is important to get the influenza vaccine even if you are healthy. Most Australians who get influenza are quite sick for a few days with fever, aches and pains, and sore throat, and then recover without lasting effects (see Q1). However, influenza can cause serious illness in some people; this can lead to hospitalisation or even death.

It is not possible to predict who will be severely affected by influenza. Each year, previously healthy people are hospitalised and die from the virus. Although around 100 deaths and 5,100 hospitalisations due to influenza are reported each year,³ many cases don't get identified, so the true impact of influenza is much greater.

Apart from becoming ill, having influenza can be a big inconvenience. People can miss time from work or school because either they are too sick to attend or they have to take time off to care for a sick child. One study has shown that parents of children under 3 years of age missed an average of 3 days of work to stay home and care for their sick child.⁴ The estimated cost to the Australian healthcare system for GP visits and hospitalisations was \$115 million per year for each year between April 2000 and March 2006.⁵

In addition to protecting you from influenza, vaccination also protects people around you. If you don't catch influenza, then you can't spread the infection. It is particularly important to protect vulnerable people who can't get the vaccine themselves or don't respond strongly to it, such as young babies under 6 months of age and people who have low immunity.^{6,7}

The influenza vaccine is not completely protective because the circulating influenza strain may change and people's individual response to the vaccine may differ. However, it is our best protection against infection and potentially serious complications, including death.

Given SARS-CoV-2 (the virus that causes COVID-19) continues to circulate, it is even more important to get an influenza vaccine to stay healthy, to protect others from getting influenza and to reduce impact on our health systems.

Q3. Why do I need the influenza vaccine if there has not been any influenza in Australia?

Influenza is a highly contagious infection of the airways. It is often a mild disease but it can cause serious illness in otherwise healthy people. It can affect people of all ages and in some people, it can cause hospitalisation and death.

Although during the COVID-19 pandemic there has been reduced circulation of influenza virus and lower levels of influenza vaccine coverage than in previous years, in 2022, with border protection measures easing and international travel resuming, Australia observed a resurgence of influenza with the influenza season commencing earlier than in previous years. During 2023, influenza epidemiology may be atypical, particularly in the context of COVID-19.

Vaccination is a safe and effective way to protect yourself from serious disease caused by influenza. By getting vaccinated against influenza, you can also help protect other people, especially people who are too sick or too young to be vaccinated. The more people who are vaccinated in your community, the less likely the disease will spread.

Yearly influenza vaccination is recommended for people aged 6 months and older. Anyone who wants to protect themselves against influenza can talk to their immunisation provider about getting immunised (see Q6).

Q4. Why do healthy young children need an influenza vaccine?

Compared with older children and adults, infants and children under 5 years of age, including those without pre-existing medical conditions, are more likely to get severe influenza infection, resulting in hospitalisation.^{3,8} In 2017, approximately 1 in 400 children were diagnosed with laboratory-confirmed influenza. Previously healthy children can be severely ill and suffer from influenza-associated complications such as pneumonia and encephalitis.⁹⁻¹¹

Q5. Why do children 6 months to <9 years of age who have never received influenza vaccine need two doses?

Children 6 months to <9 years of age receiving influenza vaccine for the first time need two doses, given at least 4 weeks apart to maximise the immune response to the vaccine strains.

Children who received one or more doses of influenza vaccine in a previous season only need one dose in the current and future seasons.

Q6. If the influenza vaccine is recommended for everyone then why can only certain people get it for free?

The influenza vaccine is funded by the government under the National Immunisation Program (NIP) for certain groups of people who are at the highest risk of severe influenza or are more likely to get complications from influenza than the general population. ¹² These groups include:

- children aged 6 months to under 5 years
- all Aboriginal and Torres Strait Islander people older than 6 months
- all adults aged 65 years and older
- pregnant women
- people with certain underlying medical conditions.

Since 2005, decisions on what vaccines are provided for free, and for whom, are made following a process that involves the Pharmaceutical Benefits Advisory Committee.¹³ This ensures any government spending on a health intervention is cost-effective. This is important as there is a limited amount of money that is available for healthcare in Australia and these funds need to be used for the greatest benefit for the whole population.

However, people who are not eligible for influenza vaccine on the NIP will still benefit from vaccination (see Q2). The <u>Australian Immunisation Handbook</u> (the national clinical guideline advising on the safest and most effective use of vaccines in Australia) recommends the influenza vaccine from 6 months of age. ¹⁴ A range of providers can administer private influenza vaccines (e.g. GPs, pharmacists). Influenza vaccines can be purchased for around \$20–\$30 each.

Q7. How do I protect myself from influenza if travelling to the northern hemisphere or returning to Australia?

Travellers may be exposed to the influenza virus at any time throughout the year regardless of their destination. The influenza season in the southern hemisphere is mostly from April to September; in the northern hemisphere, it is from October to May. Influenza activity in the tropics has been reported throughout the year.

People travelling in large tourist groups or those travelling in confined spaces for days to weeks, such as on a cruise ship, are at particular risk of influenza.^{15,16} People can get infection either before departure or from travel to areas of the world where influenza may be circulating at that time.

Influenza vaccination is recommended if travelling, especially if it is known before travel that influenza is circulating in the destination region. Some brands of current southern hemisphere

influenza vaccine are available from March through February the following year (when the vaccine expires). A northern hemisphere formulation of influenza vaccine may be preferred if travelling in the northern hemisphere during their influenza season, but is generally unavailable in Australia (see Q9) The southern hemisphere formulation is considered as an acceptable alternative; a second dose late in the season may be given even if the person has received this vaccine earlier in the current season (see Q8).

If you have returned to Australia from living or travelling overseas, consider getting vaccinated with the currently available local vaccine for optimal protection if you have not had it before, regardless of the time of the year.

Q8. Is a second dose of influenza vaccine recommended before travel to the northern hemisphere?

Yes. Individuals who have received a current southern hemisphere influenza vaccine and are travelling to the northern hemisphere during their influenza season (usually October to May) may receive a second dose of influenza vaccine before travel (ideally 2 weeks before departure). Further information about influenza vaccination and travellers is available in the <u>Australian Immunisation Handbook</u>.

Q9. Why are there different influenza vaccines in the northern hemisphere and the southern hemisphere?

The influenza vaccine is designed to protect against the four influenza viruses that are most likely to spread and cause illness among people during the upcoming influenza season.

The circulating strains of influenza virus vary each year in both the northern and southern hemispheres. Therefore, the influenza vaccine composition is reviewed each year based on which influenza viruses are making people sick, the extent to which those viruses are spreading and how well the previous season's vaccine protects against those viruses.

Annual influenza immunisation is hence strongly recommended to have the best possible protection against influenza disease and its complications.

Q10. How effective is the influenza vaccine in preventing influenza illness?.

Extensive research has shown the influenza vaccine is effective. It takes 2 weeks after the influenza vaccination for you to develop immunity.

How well the influenza vaccine works can vary among different people and in different years, as it depends on several factors, such as age and health of the person receiving the vaccine and the match between the vaccine strains and those circulating in the community.

Influenza vaccination can prevent illness in about 50–60% of young children and healthy adults under the age of 65 years, although this figure varies year by year. 17,18

People with an underlying medical condition, such as those with low immunity, or the elderly, may not respond as well to the influenza vaccine as healthy adults do and so the level of protection they get from the vaccine may be less. However, among high-risk individuals such as nursing home residents, the vaccine prevents pneumonia and hospitalisation due to influenza. Because of the higher risk of severe influenza in the elderly, any protection provided by vaccination against influenza is worthwhile.

As the vaccine is not 100% effective, it means a small proportion of people may catch the influenza virus after getting the vaccine. People sometimes think they have caught influenza after getting vaccination but that is not the case. Influenza vaccine cannot give you influenza because

it does not contain a live virus. Sometimes people catch influenza before getting the influenza vaccine but their symptoms don't appear until shortly after being vaccinated. This makes them think the vaccine didn't work or even (mistakenly) that the vaccine made them sick.

Similarly, a person who is vaccinated against influenza may catch a different virus that is mistaken for influenza (see Q1). For instance, respiratory syncytial virus (RSV) and parainfluenza are viruses that cause symptoms similar to those of influenza, spread in the community at the same time as influenza and can cause severe illness and complications just like influenza.²⁰

Q11. What is an 'enhanced' influenza vaccine and how is it different from other influenza vaccines?

An enhanced influenza vaccine is one that is specifically designed to increase the immune system's response to the vaccine. It can contain the standard amount of antigen but with an adjuvant (a compound that stimulates a higher immune response to a vaccine) or contain a higher amount of antigen but with no adjuvant.

The highest disease burden from influenza occurs in the elderly in terms of serious complications and death rates.³ The elderly do not respond as well to the influenza vaccine as younger adults do, as the immune system weakens with age. The level of protection they get from the influenza vaccine is usually less than that of a younger person. This underpins the need for 'enhanced' influenza vaccines for people 65 years of age and older to better protect them from influenza infection.

Two 'enhanced' influenza vaccines are available for older people in 2023. Fluad Quad (adjuvanted) is available for people aged ≥65 years and Fluzone High-Dose Quadrivalent is available for people aged ≥60 years.

Fluzone High-Dose Quadrivalent, an inactivated split-virus influenza vaccine, became available in Australia in 2022. It contains 240 micrograms haemagglutinin (60 µg for each strain), four times the amount in standard-dose influenza vaccines. It has a similar safety and superior effectiveness profile as the standard-dose influenza vaccines and a similar safety and effectiveness profile as the adjuvanted influenza vaccine. ²¹⁻²⁴ Both Fluzone High-Dose Quadrivalent and Fluad Quad are preferentially recommended over standard-dose influenza vaccine. There is no preference between Fluad Quad and Fluzone High-Dose Quadrivalent. Both vaccines are latex-free and safe in people with latex allergy.

While both enhanced vaccines are recommended in preference over standard-dose influenza vaccines for people aged ≥65 years, in 2023, only Fluad Quad is available free of charge via the National Immunisation Program (NIP).

Q12. Are there any precautions associated with the Fluzone High-Dose Quadrivalent influenza vaccine?

Precautions to be taken when administering high-dose influenza vaccines are comparable to those for standard-dose and adjuvanted influenza vaccines. Fluzone High-Dose Quadrivalent is contraindicated in individuals with a history of severe allergic reaction after receiving any influenza vaccine and to any component of the vaccine. Although this is an egg-based vaccine, people with a history of anaphylaxis to egg can still receive it.

There are currently no specific data on the co-administration of Fluzone High-Dose Quadrivalent with other vaccines. However, people can receive it with other vaccines, including COVID-19 vaccines.

In addition, adverse events after vaccination with Fluzone High-Dose Quadrivalent are slightly more common than those with standard-dose influenza vaccines, but these are mostly mild and

temporary.²⁵ Adverse events following vaccination with high-dose influenza vaccines are similar to those following vaccination with adjuvanted influenza vaccines.²⁶⁻²⁸

Q13. When should I get the influenza vaccine and when is it too late in the season to get it?

It is recommended you get annual influenza vaccine before the influenza season starts. Immunisation providers should start giving vaccinations as soon as vaccine is available.

The peak of influenza activity in Australia can vary each year. Typically it occurs between June and September, but infections can occur year round.

It is never too late in the year to get influenza vaccine. Vaccination should be available throughout the influenza season (note: most years vaccine brands don't expire until end of February the following year). Evidence suggests optimal protection occurs in the 3–4 months following vaccination; ideally vaccination before the expected winter peak is advisable.^{29,30}

For pregnant women, influenza vaccine is recommended in every pregnancy and it is safe for the mother and the unborn baby to receive the vaccine at any stage of pregnancy. Women who are in their first trimester in the first quarter of the year may wish to wait until the current year's seasonal influenza vaccine becomes available, rather than receiving the previous year's influenza vaccine. Influenza vaccine can safely be given at the same time as pertussis vaccine.

Q14. Can I get an influenza vaccine at the same time as other vaccines, including childhood and COVID-19 vaccines?

Yes, you can receive influenza vaccine at any time before or after, or with, most other vaccines, including COVID-19 vaccines.. Studies show that co-administration of COVID-19 and influenza vaccines is safe and produces a good immune response.

The COVID-19 vaccine does not protect against influenza, so you should still have your annual influenza vaccine.

For more information, refer to <u>ATAGI advice on influenza and COVID-19 vaccines</u>.

Q15. I had influenza infection recently, do I still need to get the influenza vaccine?

Yes, even if you have had influenza infection recently, you should get the influenza vaccine as it can lower your risk of becoming ill from other strains of the influenza virus. Yearly influenza vaccination is recommended for people aged 6 months and older.

Q16. How long after COVID-19 infection can I have the influenza vaccine?

There is no set timeframe to wait between having a COVID-19 infection and then having the influenza vaccine. Once you are feeling well and have no fever, you may receive an influenza vaccine.

Q17. Who needs to get more than one dose of influenza vaccine within a year?

People recommended to receive a second dose of influenza vaccine within a year include:

- Children 9 years of age and younger receiving their influenza vaccine for the first time. Two doses 4 weeks apart are required for an adequate immune response.
- People who have had a haematopoietic stem cell transplant or solid organ transplant and are receiving influenza vaccine for the first time after transplant.
- Pregnant women, who may be vaccinated with the next season's influenza vaccine if it becomes available in the latter part of their pregnancy, even if they were vaccinated with the previous season's vaccine prior to or earlier in pregnancy (see Q26).

 Overseas travellers, who may benefit from a second dose of this season's influenza vaccine if going to the northern hemisphere winter and receiving the northern hemisphere formulation there is not feasible (see Q7, Q8 and Q9).

There is not enough evidence to routinely support a second dose in the general population at this time, even if the influenza vaccine was given early in the season. The few studies of antibody responses (an indirect measure of protection) following a second dose of influenza vaccine in the same season in adults have not shown consistent results. For example, two studies in the elderly of a second vaccine dose at either 1 month or 3 months after the first dose did not show higher antibody levels, 31-33 while a third study suggested a better antibody response after a booster dose at 3 months in frail elderly patients. 34

Q18. Is the influenza vaccine available all year round?

The influenza vaccine is usually available from late March through February the following year (when the vaccine expires). This means there may be a gap of about 1–2 month where no influenza vaccine may be available.

Q19. What type of vaccine is the influenza vaccine?

All influenza vaccines available in Australia are either split virion (inactivated, 'split virus') or subunit vaccines (inactivated, fragments of the 'shell' of the virus). These vaccines do not contain live virus and cannot cause influenza illness.

The purified inactivated influenza virus used in vaccine preparation is cultivated in embryonated hens' eggs (standard influenza vaccines and adjuvanted influenza vaccine (with an additive to enhance immune response)) or propagated in mammalian cell lines (cell-based influenza vaccine). See Q21.

Q20. What are the ingredients of the influenza vaccine?

All influenza vaccines available in Australia are quadrivalent as they contain 4 influenza virus strains – 2 influenza A subtypes and 2 influenza B lineages.

Standard (egg-based) influenza vaccines contain 15 µg of haemagglutinin per strain per dose with no adjuvant. Similarly, the available cell-based influenza vaccine contains 15 µg of haemagglutinin per strain per dose with no adjuvant.

The available adjuvanted influenza vaccine is egg-based, and contains the standard 15 µg of haemagglutinin per strain per dose, with MF59 as the adjuvant. The adjuvanted influenza vaccine is formulated to induce a greater immune response than standard influenza vaccines.

High-dose (egg-based) influenza vaccines contain 60 µg of haemagglutinin per strain per dose with no adjuvant. The high-dose influenza vaccine is also formulated to induce a greater immune response than standard influenza vaccines.

Although egg-based vaccines may contain traces of egg-derived protein (ovalbumin), they contain less than 1 μ g of ovalbumin per dose.

Q21. What is a cell-based influenza vaccine?

Cell-based influenza vaccines are prepared in Madin-Darby canine kidney (MDCK) cells. This method is different from the traditional method of producing standard influenza vaccines, which involves hens' eggs. In recent years, there has been concern that the replication of influenza viruses in eggs during vaccine production results in antigenic changes that could make them less closely related to the circulating strains. This 'antigenic mismatch' has been a particular problem for influenza A/H3N2 subtypes and is thought to have contributed to lower influenza vaccine effectiveness in some years (particularly when A/H3N2 was the predominant strain). 35,36

Also, there is a logistical concern that supply of influenza vaccines could be compromised if there was a worldwide shortage of eggs, or if rapid increases in vaccines were needed (e.g. in an influenza pandemic). Cell-based influenza vaccine production diversifies the supply lines and theoretically mitigates the 'antigenic mismatch' issue of egg-based vaccine production.

Flucelvax Quad is currently the only cell-based inactivated influenza vaccine that has been registered for use in Australia. It is approved for use in people from 2 years of age. This vaccine is currently not funded under the NIP for any population groups.

Q22. Is the cell-based influenza vaccine preferred over standard (egg-based) influenza vaccines?

Although there are some theoretical advantages of cell-based influenza vaccines (see Q21), studies have shown that the cell-based influenza vaccine has a similar efficacy and safety profile to standard influenza vaccines.³⁷⁻⁴¹ There is no preferential recommendation for its use over the standard influenza vaccines.

Standard influenza vaccines are currently preferred for use in pregnancy because a large body of evidence supports their safety in pregnant women. While the use of cell-based influenza vaccines in pregnancy has not been assessed, there are no theoretical concerns regarding their safety in pregnant women.

The remaining eligible population can receive either standard influenza vaccine or cell-based influenza vaccine.

Questions about the safety of influenza vaccines

Q23. Can the influenza vaccine cause influenza?

No. It is not possible for the influenza vaccine to give you influenza. This is because all influenza vaccines in use in Australia are 'inactivated' (the virus is killed) and the vaccine is either split into components or made up only of fragments of the outside shell of the influenza virus, and therefore, it is not able to function like a whole live virus.⁴²

Sometimes the normal responses the body experiences after getting the vaccine (i.e. side effects) are similar to the early signs of influenza. This can make people think they have gotten influenza from the vaccine. For example, the expected side effects of the vaccine are swelling, redness and pain at the injection site but also fever, tiredness and muscle aches which also occur when you get influenza (see Q1, Q10). However, these side effects are a sign that the vaccine is triggering an immune response, which is what it is designed to do. The symptoms can start within a few hours of being vaccinated, last 1–2 days, and are generally much milder than an actual influenza infection. These symptoms go away on their own once your body has successfully made an immune response to the vaccine, which will protect you from influenza virus.⁴³

Q24. How do we know that influenza vaccines are safe?

The safety of vaccines is reviewed and monitored at all stages of the vaccine development process, from initial lab-based research, vaccine registration, recommendations on the use of the vaccine to ongoing surveillance once the vaccine is being used in the population.

In Australia, the Therapeutic Goods Administration (TGA) is responsible for registering vaccines for use in the country. The TGA uses the latest research and testing information available to evaluate and ensure the safety and efficacy of vaccines. Independent medical and scientific

advice on the safety, quality and efficacy of vaccines is provided by experts who make up the Advisory Committee on Vaccines (ACV).⁵⁴

Once vaccines are registered and in use, the TGA continues to monitor their safety and effectiveness through a national monitoring system. If the TGA receives information that there are safety concerns about a vaccine, the issue is investigated immediately.⁵⁵

Australia also conducts active safety monitoring of vaccines using a system called AusVaxSafety. AusVaxSafety collects adverse events data directly from patients via an SMS-based survey after vaccination at certain GPs and immunisation providers. Data are collected in near real-time and analysed for safety concerns. The latest safety data are available on the AusVaxSafety website.

Another important body is the Australian Technical Advisory Group on Immunisation (ATAGI).⁵⁶ This group advises the government on existing, new and emerging vaccines in relation to their effectiveness and use in Australian populations. ATAGI produces the <u>Australian Immunisation Handbook</u>, the national clinical guideline advising on the safest and most effective use of vaccines in Australia. ATAGI and the ACV work together with other bodies on the implementation of immunisation policies, procedures and vaccine safety.

Q25. Are there any side effects from influenza vaccines?

Common side effects

Approximately 10 percent of children (1 out of every 10) who have an influenza vaccine experience swelling, redness or pain at the injection site. Between 1 percent and 10 per cent of people (1–10 out of every 100) who have an influenza vaccine experience a fever, headache, malaise (tiredness or lack of energy) or myalgia (muscle aches). These symptoms can last for 1-2 days. 43,57,58

Rare side effects

Rarely, approximately 1 in every 20,000 children who receive the influenza vaccine have a type of seizure known as a febrile convulsion.⁴⁹ This can happen when a baby or child's temperature (fever) goes up suddenly. Influenza infection itself can also cause fever and results in many more febrile seizures than vaccination.⁴⁷ In one study, 6% (6 out of every 100) of children hospitalised with influenza suffered a febrile seizure.⁴⁸

In Australia in 2010, higher than expected numbers of fever and febrile seizures following influenza vaccination were detected in children under 5 years of age, particularly in children under 3 years of age.⁵⁰ Upon investigation, the reports were linked to only one manufacturer's influenza vaccine and the issue was likely caused by the manufacturing process for that particular vaccine.⁵¹⁻⁵³ This vaccine is no longer available in Australia.

Enhanced safety monitoring systems for influenza vaccines introduced in recent years, such as <u>AusVaxSafety</u> (see <u>Q24</u>), have confirmed that influenza vaccine is safe in children under 5 years, reporting low rates of fever and medical attendance after vaccination.⁴⁴

Very rare side effects

More serious side effects following influenza vaccination are very rare. Anaphylaxis occurs in approximately 1 per 1 million people who receive the influenza vaccine.⁵⁹ While Guillain-Barré syndrome (GBS) has shown a very low rate of less than 1 in 1 million doses of influenza vaccine.⁶⁰ Studies have also shown that a person is more likely to get GBS from infection with the influenza virus than from the influenza vaccine⁶¹ (see Q29).

Q26. I've been recommended to get the influenza vaccine when pregnant to protect me and my baby. Is this safe?

Influenza vaccine is safe during pregnancy. The vaccine is recommended with every pregnancy and at any stage of pregnancy to protect both the mother and her unborn child against complications from influenza.

Influenza can cause severe disease in pregnant women and young babies. Getting sick with influenza while pregnant can lead to complications such as premature delivery and even perinatal death. For young children, especially those younger than 6 months, are more likely to be hospitalised or die from influenza than older children.

During pregnancy, protective antibodies are transferred through the placenta from the mother to the baby. Babies born to women vaccinated against influenza while pregnant are less likely to be born prematurely or have a low birth weight. 63,64

Maternal vaccination is estimated to reduce the risk of influenza in infants <6 months of age by 48%. 65-67 However, the protection wears off as babies get to 6 months of age, at which time babies can start to receive the vaccine themselves. 68

Multiple safety studies have found no increase in foetal death, spontaneous abortion or congenital malformation after maternal influenza vaccination in pregnancy.⁶³ The frequency of expected adverse events after vaccination, like injection site reactions, is the same in both pregnant and non-pregnant women. Influenza vaccine is also safe when given to mothers who are breastfeeding and can provide protection to the baby through antibodies that are transferred to the baby in breastmilk.⁶⁹

For pregnant women who received an influenza vaccine in 2022, revaccinate if the 2023 influenza vaccine becomes available before the end of pregnancy. For women who received an influenza vaccine before becoming pregnant, revaccinate during pregnancy to protect the unborn infant. Influenza vaccine can safely be given at the same time as pertussis vaccine.

Q27. I'm a breastfeeding mother. Will influenza vaccine reduce my milk supply?

Influenza vaccine is safe for pregnant and breastfeeding women. Although there is little published data specifically examining the impact of influenza vaccine on breastmilk production, influenza vaccines have been used in women who are breastfeeding for many years and decreased milk supply has not been identified as a <u>safety signal</u> (i.e. an adverse event that has been detected and is flagged for further monitoring) in any vaccine safety surveillance systems, nor <u>listed as having an association with the influenza vaccine</u>.

A review from the Netherlands on COVID-19 vaccines suggests that decreased milk supply after vaccination could be related to stress, which can happen as part of the body's immune response to vaccination, and it is theoretically possible that this could occur after other vaccines. Some people will feel unwell for a day or two after receiving the influenza vaccine. This is a sign that the vaccine has triggered an immune response. However, being infected with the influenza virus can make people sick for much longer, and can <u>reduce breastmilk supply in some women</u>. The benefits of vaccination greatly outweigh any risks associated with vaccination.

If you have any concerns about receiving the influenza vaccine while you are pregnant or breastfeeding you can speak to your regular healthcare professional or call <u>Tresilian</u>, the <u>Australian Breastfeeding Association</u> or <u>Karitane</u> for advice.

Q28. Can I get the influenza vaccine if I have an egg or a latex allergy?

Egg allergy

People with egg allergy can safely receive the influenza vaccine (which usually contain less than 1 microgram of egg protein per dose).

Reactions such as hives, angioedema (a skin reaction with swelling similar to hives) or anaphylaxis (severe allergic reaction) are rare side effects following influenza vaccination. They can be due to an allergic response to something in the vaccines, such as egg protein.

Recent studies have shown that people with egg allergy, including egg-induced anaphylaxis, have safely received the influenza vaccine.^{70,71}

The Australasian Society of Clinical Immunology and Allergy (ASCIA) guidelines should be referred to for additional information on influenza vaccination of individuals with an allergy to eggs, including risk, dosage and observation period.⁷²

Latex allergy

Influenza vaccines used in Australia are latex-free and safe for use by people with a latex allergy or sensitivity. Although the product information for Fluarix Tetra and Fluad Quad state that some presentations of the vaccine cannot be considered latex-free, these presentations are not supplied in Australia.

Q29. Can the influenza vaccine be given to someone who has had Guillain-Barré syndrome?

People with a history of GBS whose first episode was not after vaccination have an extremely low risk of recurrence of GBS after vaccination.⁷³⁻⁷⁵ Vaccination is recommended for these people.

Vaccination is generally not recommended for people with a history of GBS whose first episode occurred within 6 weeks of influenza vaccination. There are limited data in people where the first episode occurred within 6 weeks of influenza vaccination (i.e. the first episode was possibly triggered by the vaccine). In these people, discuss the potential for GBS recurrence if vaccinated, the potential for exacerbation following influenza infection, and other protective strategies (e.g. vaccination of household members). Vaccination can be considered in special circumstances.

Guillain-Barré syndrome (GBS) is a rare disorder in which the immune system damages nerve cells, causing muscle weakness and sometimes paralysis. The symptoms usually last for a few weeks followed by a full or partial recovery. In very rare cases people have died of GBS. The risk of the syndrome increases with age and is greatest for those aged 50 years or older. Diagnosis of GBS is complex and must be made by a doctor.

A small increased risk of GBS was found in people given a specific influenza vaccine in the United States in 1976.⁷⁶ Since then, close monitoring has shown that GBS has occurred at a very low rate of less than 1 in 1 million doses of influenza vaccine.⁶⁰ Studies suggest that a person is up to 15 times more likely to get GBS from infection with the influenza virus than from the influenza vaccine.⁶¹

Q30. Can the influenza vaccine be given to someone taking immune checkpoint inhibitors?

Immune checkpoint inhibitors are a class of monoclonal antibodies currently used in the treatment of a number of cancers, including metastatic melanoma, renal clear cell carcinoma, non-Hodgkin lymphoma, non-small cell lung cancer and other solid organ tumours.

Checkpoint inhibitors include:

- CTLA-4 inhibitors (such as ipilimumab)
- PD-1 and PD-L1 inhibitors (such as nivolumab or pembrolizumab)

People taking checkpoint inhibitors may have a higher risk of immune-related adverse events following immunisation with influenza vaccine.⁷⁷ Consult the person's treating oncologist about the risks and benefits of influenza vaccination in people taking treatments.

Additional resources for primary medical care/vaccination providers

- NCIRS Influenza vaccines for Australians fact sheet
- Australian Technical Advisory Group on Immunisation (ATAGI) advice on seasonal influenza vaccines in 2023
- Australian Government Department of Health and Aged Care immunisation website
- National Immunisation Program schedule
- Sharing Knowledge About Immunisation (SKAI)

References

- 1. Eccles R. Understanding the symptoms of the common cold and influenza. *The Lancet Infectious Diseases* 2005;5:718-25.
- 2. Nichol KL, Lind A, Margolis KL, et al. The effectiveness of vaccination against influenza in healthy, working adults. *New England Journal of Medicine* 1995;333:889-93.
- 3. Li-Kim-Moy J, Yin JK, Patel C, et al. Australian vaccine preventable disease epidemiological review series: Influenza 2006 to 2015. *Communicable Diseases Intelligence* 2016;40:E482-95.
- 4. Heikkinen T, Silvennoinen H, Peltola V, et al. Burden of influenza in children in the community. *Journal of Infectious Diseases* 2004;190:1369-73.
- 5. Newall AT, Scuffham PA. Influenza-related disease: the cost to the Australian healthcare system. *Vaccine* 2008;26:6818-23.
- 6. Mertz D, Kim TH, Johnstone J, et al. Populations at risk for severe or complicated influenza illness: systematic review and meta-analysis. *BMJ* 2013;347:f5061.
- 7. Rasmussen SA, Jamieson DJ, Uyeki TM. Effects of influenza on pregnant women and infants. *American Journal of Obstetrics and Gynecology* 2012;207(3 Suppl):S3-8.
- 8. Izurieta HS, Thompson WW, Kramarz P, et al. Influenza and the rates of hospitalization for respiratory disease among infants and young children. *New England Journal of Medicine* 2000;342:232-9.
- Coffin SE, Zaoutis TE, Rosenquist AB, et al. Incidence, complications, and risk factors for prolonged stay in children hospitalized with community-acquired influenza. *Pediatrics* 2007;119:740-8.

- 10. Britton PN, Blyth CC, Macartney K, et al. The spectrum and burden of influenza-associated neurological disease in children: combined encephalitis and influenza sentinel site surveillance from Australia, 2013-2015. *Clinical Infectious Diseases* 2017;65:653-60.
- 11. Britton PN, Dale RC, Blyth CC, et al. Influenza-associated encephalitis/encephalopathy identified by the Australian Childhood Encephalitis Study 2013-2015. *Pediatric Infectious Disease Journal* 2017;36:1021-6.
- 12. Australian Technical Advisory Group on Immunisation (ATAGI). Australian Technical Advisory Group on Immunisation (ATAGI) advice for immunisation providers regarding the administration of seasonal influenza vaccines in 2018. Canberra: Australian Technical Advisory Group on Immunisation (ATAGI); 2018. Available from:

 https://beta.health.gov.au/resources/publications/atagi-advice-on-seasonal-influenza-vaccines-in-2018 (Accessed 5 April 2018).
- 13. Nolan TM. The Australian model of immunization advice and vaccine funding. *Vaccine* 2010;28 Suppl 1:A76-83.
- 14. Australian Technical Advisory Group on Immunisation (ATAGI). The Australian Immunisation Handbook 10th ed. Canberra: The Australian Government Department of Health and Ageing; 2018. Available from: http://www.immunise.health.gov.au/internet/immunise/publishing.nsf/Content/Handbook10-home~handbook10part4~handbook10-4-7 (Accessed 6/4/2018).
- 15. Steffen R. Influenza in travelers: epidemiology, risk, prevention, and control issues. *Curr Infect Dis Rep* 2010;12:181-5.
- 16. Marti F, Steffen R, Mutsch M. Influenza vaccine: a travelers' vaccine? *Expert Review of Vaccines* 2008;7:679-87.
- 17. Osterholm MT, Kelley NS, Sommer A, Belongia EA. Efficacy and effectiveness of influenza vaccines: a systematic review and meta-analysis. *The Lancet Infectious Diseases* 2012;12:36-44.
- 18. Blyth CC, Jacoby P, Effler PV, et al. Effectiveness of trivalent flu vaccine in healthy young children. *Pediatrics* 2014;133:e1218-25.
- Jefferson T, Di Pietrantonj C, Al-Ansary LA, et al. Vaccines for preventing influenza in the elderly. Cochrane Database of Systematic Reviews 2010;(2):CD004876. doi:10.1002/14651858.CD004876.pub3.
- 20. Zambon MC, Stockton JD, Clewley JP, Fleming DM. Contribution of influenza and respiratory syncytial virus to community cases of influenza-like illness: an observational study. *The Lancet* 2001;358:1410-6.
- 21. Shay DK, Chillarige Y, Kelman J, et al. Comparative effectiveness of high-dose versus standard-dose influenza vaccines among US Medicare beneficiaries in preventing postinfluenza deaths during 2012-2013 and 2013-2014. *Journal of Infectious Diseases* 2017;215:510-7.
- 22. Young-Xu Y, Thornton Snider J, Mahmud SM, et al. High-dose influenza vaccination and mortality among predominantly male, white, senior veterans, United States, 2012/13 to 2014/15. *Euro Surveillance* 2020:25.
- 23. Izurieta HS, Chillarige Y, Kelman J, et al. Relative effectiveness of influenza vaccines among the United States elderly, 2018-2019. *Journal of Infectious Diseases* 2020;222:278-87.

- 24. Izurieta HS, Lu M, Kelman J, et al. Comparative effectiveness of influenza vaccines among US Medicare beneficiaries ages 65 years and older during the 2019-2020 season. *Clinical Infectious Diseases* 2021;73:e4251-e9.
- 25. Pepin S, Nicolas JF, Szymanski H, et al. Immunogenicity and safety of a quadrivalent high-dose inactivated influenza vaccine compared with a standard-dose quadrivalent influenza vaccine in healthy people aged 60 years or older: a randomized Phase III trial. *Human Vaccine Immunotherapy* 2021;17:5475-86.
- 26. Cowling BJ, Thompson MG, Ng TWY, et al. Comparative reactogenicity of enhanced influenza vaccines in older adults. *Journal of Infectious Diseases* 2020;222:1383-91.
- 27. Pillsbury AJ, Fathima P, Quinn HE, et al. Comparative postmarket safety profile of adjuvanted and high-dose influenza vaccines in individuals 65 years or older. *JAMA Network Open* 2020;3:e204079.
- 28. Schmader KE, Liu CK, Harrington T, et al. Safety, reactogenicity, and health-related quality of life after trivalent adjuvanted vs trivalent high-dose inactivated influenza vaccines in older adults: A randomized clinical trial. *JAMA Network Open* 2021;4:e2031266.
- 29. Belongia EA, Sundaram ME, McClure DL, et al. Waning vaccine protection against influenza A (H3N2) illness in children and older adults during a single season. *Vaccine* 2015;33:246-51.
- 30. Sullivan SG, Komadina N, Grant K, et al. Influenza vaccine effectiveness during the 2012 influenza season in Victoria, Australia: influences of waning immunity and vaccine match. *Journal of Medical Virology* 2014;86:1017-25.
- 31. Buxton JA, Skowronski DM, Ng H, et al. Influenza revaccination of elderly travelers: antibody response to single influenza vaccination and revaccination at 12 weeks. *Journal of Infectious Diseases* 2001;184:188-91.
- 32. Gross PA, Weksler ME, Quinnan GV, Jr., et al. Immunization of elderly people with two doses of influenza vaccine. *Journal of Clinical Microbiology* 1987;25:1763-5.
- 33. Nichol KL. Booster doses of influenza vaccine. JAMA 1996;276:1857.
- 34. Roos-van Eijndhoven DG, Cools HJM, Westendorp RGJ, et al. Randomized controlled trial of seroresponses to double dose and booster influenza vaccination in frail elderly subjects. *Journal of Medical Virology* 2001;63:293-8.
- 35. Wu NC, Zost SJ, Thompson AJ, et al. A structural explanation for the low effectiveness of the seasonal influenza H3N2 vaccine. *PLoS Pathog* 2017;13:e1006682.
- 36. Zost SJ, Parkhouse K, Gumina ME, et al. Contemporary H3N2 influenza viruses have a glycosylation site that alters binding of antibodies elicited by egg-adapted vaccine strains. *Proceedings of the National Academy of Sciences of the United States of America* 2017;114:12578-83.
- 37. Bruxvoort KJ, Luo Y, Ackerson B, et al. Comparison of vaccine effectiveness against influenza hospitalization of cell-based and egg-based influenza vaccines, 2017-2018. *Vaccine* 2019;37:5807-11.
- 38. Divino V, Krishnarajah G, Pelton SI, et al. A real-world study evaluating the relative vaccine effectiveness of a cell-based quadrivalent influenza vaccine compared to egg-based quadrivalent influenza vaccine in the US during the 2017-18 influenza season. *Vaccine* 2020;38:6334-43.

- 39. Krishnarajah G, Divino V, Postma MJ, et al. Clinical and Economic Outcomes Associated with Cell-Based Quadrivalent Influenza Vaccine vs. Standard-Dose Egg-Based Quadrivalent Influenza Vaccines during the 2018-19 Influenza Season in the United States. *Vaccines* (*Basel*) 2021;9.
- 40. Nolan T, Chotpitayasunondh T, Capeding MR, et al. Safety and tolerability of a cell culture derived trivalent subunit inactivated influenza vaccine administered to healthy children and adolescents: A Phase III, randomized, multicenter, observer-blind study. *Vaccine* 2016;34:230-6.
- 41. Ambrozaitis A, Groth N, Bugarini R, et al. A novel mammalian cell-culture technique for consistent production of a well-tolerated and immunogenic trivalent subunit influenza vaccine. *Vaccine* 2009;27:6022-9.
- 42. Gross PA, Ennis FA, Gaerlan PF, et al. A controlled double-blind comparison of reactogenicity, immunogenicity, and protective efficacy of whole-virus and split-product influenza vaccines in children. *Journal of Infectious Diseases* 1977;136:623-32.
- 43. Mahajan D, Roomiani I, Gold MS, et al. Annual report: surveillance of adverse events following immunisation in Australia, 2009. *Communicable Diseases Intelligence* 2010;34:259-76.
- 44. Pillsbury A, Quinn H, Cashman P, Leeb A, Macartney K. Active SMS-based influenza vaccine safety surveillance in Australian children. *Vaccine* 2017;35:7101-6.
- 45. Patel N, Ram D, Swiderska N, et al. Febrile seizures. BMJ 2015;351:h4240.
- 46. Waruiru C, Appleton R. Febrile seizures: an update. *Archives of Disease in Childhood* 2004;89:751-6.
- 47. Francis JR, Richmond P, Robins C, et al. An observational study of febrile seizures: the importance of viral infection and immunization. *BMC Pediatr* 2016;16:202.
- 48. Li-Kim-Moy J, Yin JK, Blyth CC, et al. Influenza hospitalizations in Australian children. *Epidemiology and Infection* 2017;145:1451-60.
- 49. Tse A, Tseng HF, Greene SK, Vellozzi C, Lee GM. Signal identification and evaluation for risk of febrile seizures in children following trivalent inactivated influenza vaccine in the Vaccine Safety Datalink Project, 2010–2011. *Vaccine* 2012;30:2024-31.
- 50. Armstrong PK, Dowse GK, Effler PV, et al. Epidemiological study of severe febrile reactions in young children in Western Australia caused by a 2010 trivalent inactivated influenza vaccine. *BMJ Open* 2011:1:e000016.
- 51. Australian Government Department of Health, Therapeutic Goods Administration. Seasonal flu vaccine: Investigation into febrile reactions in young children following 2010 seasonal trivalent influenza vaccination. Status report as at 2 July 2010 (updated 24 September 2010). Available from: http://www.tga.gov.au/alert/seasonal-flu-vaccine-investigation-febrile-reactions-young-children-following-2010-seasonal-trivalent-influenza-vaccination (Accessed 20 March 2019).
- 52. Rockman S, Becher D, Dyson A, et al. Role of viral RNA and lipid in the adverse events associated with the 2010 Southern Hemisphere trivalent influenza vaccine. *Vaccine* 2014;32:3869-76.

- 53. Rockman S, Dyson A, Koernig S, et al. Evaluation of the bioactivity of influenza vaccine strains in vitro suggests that the introduction of new strains in the 2010 Southern Hemisphere trivalent influenza vaccine is associated with adverse events. *Vaccine* 2014;32:3861-8.
- 54. Therapeutic Goods Administration (TGA). Advisory Committee on Vaccines (ACV) Available from: http://www.tga.gov.au/committee/advisory-committee-vaccines-acv (Accessed 9 March 2018).
- 55. Australian Government Department of Health and Ageing. Are vaccines safe. Canberra: 2018. Available from: https://beta.health.gov.au/topics/immunisation/getting-started/are-vaccines-safe (Accessed 9 March 2018).
- 56. Australian Government Department of Health and Ageing. Australian Technical Advisory Group on Immunisation (ATAGI). Available from: https://beta.health.gov.au/committees-and-groups/australian-technical-advisory-group-on-immunisation (Accessed 9 March 2018).
- 57. Wood NJ, Blyth CC, Willis GA, et al. The safety of seasonal influenza vaccines in Australian children in 2013. *Medical Journal of Australia* 2014;201:596-600.
- 58. Mahajan D, Menzies R, Cook J, Macartney K, McIntyre P. Supplementary report: surveillance of adverse events following immunisation among children aged less than 7 years in Australia, 1 January to 30 June 2010. *Communicable Diseases Intelligence* 2011;35:21-8.
- 59. Australasian Society of Clinical Immunology and Allergy Limited (ASCIA). Guidelines Vaccination of the egg-allergic individual. 2017. Available from:

 https://www.allergy.org.au/hp/papers/vaccination-of-the-egg-allergic-individual (Accessed 1 July 2019).
- 60. Nelson KE. Invited commentary: influenza vaccine and Guillain-Barré syndrome is there a risk? *American Journal of Epidemiology* 2012;175:1129-32.
- 61. Kwong JC, Vasa PP, Campitelli MA, et al. Risk of Guillain-Barre syndrome after seasonal influenza vaccination and influenza health-care encounters: a self-controlled study. *The Lancet Infectious Diseases* 2013;13:769-76.
- 62. McMillan M, Porritt K, Kralik D, Costi L, Marshall H. Influenza vaccination during pregnancy: a systematic review of fetal death, spontaneous abortion, and congenital malformation safety outcomes. *Vaccine* 2015;33:2108-17.
- 63. Fell DB, Dodds L, MacDonald NE, Allen VM, McNeil S. Influenza vaccination and fetal and neonatal outcomes. *Expert Review of Vaccines* 2013;12:1417-30.
- 64. Legge A, Dodds L, MacDonald NE, Scott J, McNeil S. Rates and determinants of seasonal influenza vaccination in pregnancy and association with neonatal outcomes. *Canadian Medical Association Journal* 2014;186:E157-64.
- 65. Benowitz I, Esposito DB, Gracey KD, Shapiro ED, Vázquez M. Influenza vaccine given to pregnant women reduces hospitalization due to influenza in their infants. *Clinical Infectious Diseases* 2010;51:1355-61.
- 66. Nunes MC, Madhi SA. Influenza vaccination during pregnancy for prevention of influenza confirmed illness in the infants: a systematic review and meta-analysis. *Human vaccines & immunotherapeutics* 2017:1-9.
- 67. Zaman K, Roy E, Arifeen SE, et al. Effectiveness of maternal influenza immunization in mothers and infants. *New England Journal of Medicine* 2008;359:1555-64.

- 68. Nunes MC, Cutland CL, Jones S, et al. Duration of infant protection against influenza illness conferred by maternal immunization: secondary analysis of a randomized clinical trial. *JAMA Pediatrics* 2016;170:840-7.
- 69. Brady RC, Jackson LA, Frey SE, et al. Randomized trial comparing the safety and antibody responses to live attenuated versus inactivated influenza vaccine when administered to breastfeeding women. *Vaccine* 2018;36:4663-71.
- 70. Des Roches A, Paradis L, Gagnon R, et al. Egg-allergic patients can be safely vaccinated against influenza. *Journal of Allergy and Clinical Immunology* 2012;130:1213-6.e1.
- 71. Greenhawt MJ, Li JT, Bernstein DI, et al. Administering influenza vaccine to egg allergic recipients: a focused practice parameter update. *Annals of Allergy, Asthma and Immunology* 2011;106:11-6.
- 72. Australian Society of Clinical Immunology and Allergy (ASCIA). Guidelines: vaccination of the egg-allergic individual. ASCIA; 2017. Available from:

 https://www.allergy.org.au/images/stories/pospapers/ASCIA_Guidelines_vaccination_egg_allergic_individual_2017.pdf (Accessed 9 March 2018).
- 73. Wijdicks EF, Fletcher DD, Lawn ND. Influenza vaccine and the risk of relapse of Guillain-Barre syndrome. *Neurology* 2000;55:452-3.
- 74. Baxter R, Lewis N, Bakshi N, Vellozzi C, Klein NP. Recurrent Guillain-Barre syndrome following vaccination. *Clinical Infectious Diseases* 2012;54:800-4.
- 75. Kuitwaard K, Bos-Eyssen ME, Blomkwist-Markens PH, van Doorn PA. Recurrences, vaccinations and long-term symptoms in GBS and CIDP. *Journal of Peripheral Nervous System* 2009;14:310-5.
- 76. Haber P, Sejvar J, Mikaeloff Y, DeStefano F. Vaccines and Guillain-Barre syndrome. *Drug Safety* 2009:32:309-23.
- 77. Laubli H, Balmelli C, Kaufmann L, et al. Influenza vaccination of cancer patients during PD-1 blockade induces serological protection but may raise the risk for immune-related adverse events. *J Immunother Cancer* 2018;6:40.