

# Here's why the 2017 flu season was so bad

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Vaccines for the flu offer mediocre coverage compared with those for other diseases. Credit: PLRANG ART/Sutterstock

Australia has recorded 221,853 flu infections so far in 2017, <u>more than</u> <u>any other year</u>. As the flu season comes to <u>an end</u>, we're beginning to understand why it was so bad. And it wasn't because of increased, or more sensitive, testing.

A better vaccine could have reduced the rates, but not the high-dose Fluzone vaccine <u>doctors were touting</u> at the start of the week.



#### This year's flu viruses

Multiple flu viruses circulate each year and are broadly grouped into two types: A and B.

Influenza B viruses have two main strains, while the influenza A viruses are more variable. The influenza As you get each year are usually A/H3N2 – the main player so far this season – or A/H1N1, which lingers on from its 2009 "swine flu" pandemic.

This was a busy flu year with H1N1 strains and different strains of flu B all circulating, sometimes in the same person at the same time.

#### How vaccines are formulated

The formulation for a vaccine to immunise against all four flu viruses was decided back in <u>September 2016</u> and the final product released seven months later.

Commonly, each <u>virus</u> is amplified by injecting well-growing "seed" virus into vast numbers of fertilised (embryonated) hens' eggs. Fluid containing lots of new virus is removed, the virus is <u>inactivated</u> and the new vaccine is manufactured.

Today's flu vaccines remain <u>mediocre</u> compared to those for <u>measles</u> or <u>human papilloma virus</u>, which offer 97% and 90% protection respectively. It's normal for less than 60% of those vaccinated against the flu to develop a protective immune response.

In <u>one study of the 2017 flu season</u> a paltry 27% of Australians were vaccinated (73% weren't), including just 6% of children. Among those vaccinated, 33% were effectively protected, though rates differed between the strains. The vaccine was 5-19% effective at protecting



against H3N2 and 37% effective at protecting against H1N1 or flu B infections.

Flu vaccination doesn't produce the same degree of immunity to flu viruses that wild infection does. But vaccine protection is much safer than getting the flu, and vaccination cannot give you the flu.

#### What went wrong in 2017

It's difficult to predict what strains will dominate months later. Viruses within each flu type also change over time so H3N2 from years ago differs from H3N2 in 2017. These mutations <u>reduce the effectiveness</u> of flu vaccines.

Over the past year, H3N2 mutated <u>after it was chosen</u> as a vaccine strain. Additionally, the egg-grown H3N2 vaccine virus strain <u>changed</u> during vaccine production. For both these reasons, the vaccine no longer matched what we faced.

### Would Fluzone have helped?

This week, media stories implied that if Australia had purchased a different vaccine, Sanofi Pasteur's egg-grown inactivated Fluzone, the massive flu epidemic would not have happened. Rubbish.

The Fluzone product being discussed is a high-dose <u>flu vaccine</u> licensed in the United States (2009) and Canada (2016) for those aged 65 years or older, but not for other age groups. It's not available in Australia because Sanofi <u>hasn't applied</u> to register it here.

This vaccine contains four times more antibody-inducing active ingredient, hemagglutinin ( $60\mu g$  rather than the usual  $15\mu g$ ) than a standard-dose vaccine. But it remains susceptible to problems discussed



earlier – though this hasn't been well studied.

High-dose Fluzone is a trivalent formulation (meaning it protects against three strains of flu), not quadrivalent (four-strain protection) as <u>recommended in Australia</u> this year.

No current vaccine could have prevented 2017's flu epidemic.

## How to better protect the elderly

H3N2 viruses cause more harm among the elderly than the young, whereas flu B strains tend to impact children more.

Flu virus infection directly damages cells and paves the way for bacterial infections. The elderly are particularly affected by the flu because an older immune system struggles to defend against infections; specialised immune cells are less effective, less able to respond to new viruses and prefer reminiscing about past viral battles which confuse new skirmishes.

An ageing immune system also loses vaccine-induced antibody protection <u>faster</u> than a younger one.

To better protect the elderly, some rapid response options include moving immunisation closer to the start of <u>flu season</u>, adding an additional substance to boost the immune response to the vaccine, or increasing the amount of the virus active ingredient to produce a stronger defence against infection.

Is a high-dose vaccine safe and more effective than the standard-dose vaccines in the elderly?

Sanofi's own research found its more expensive vaccine better protected



older people from lab-confirmed flu, producing higher antibody levels than a standard-dose vaccine. High-dose Fluzone <u>showed benefit</u> over standard-dose vaccines across two seasons.

Other researchers found the elderly had <u>fewer doctor visits and</u> <u>hospitalisations</u> and that <u>deaths were prevented</u>.

But temporary, mild injection-site reactions were <u>more common</u> after high-dose vaccines because more active ingredient was injected.

However, absolute effectiveness is not clear as studies to date usually compare Fluzone to another vaccine.

## **Stopping a repeat in 2018**

The Fluzone high-dose vaccines can reduce disease and death in the elderly but won't halt spread of an efficiently transmitting flu virus.

Stopping <u>flu viruses</u> requires vaccines that are effective in those who most efficiently carry and spread the virus – <u>pre-school</u> and <u>school-aged</u> children, in whom rates are highest. It also needs higher uptake by the community.

The promise of a "universal flu <u>vaccine</u>" has been dangled since at least <u>1980</u>. We need focused research to commercialise and license vaccines that effectively protect us from seasonal influenza.

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