Please note: This <u>report</u> has been corrected. An <u>erratum</u> has been published.



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Prevention and Control of Seasonal Influenza with Vaccines: Recommendations of the Advisory Committee on Immunization Practices — United States, 2023–24 Influenza Season



**U.S. Department of Health and Human Services** Centers for Disease Control and Prevention

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#### CDC Adoption of ACIP Recommendations for MMWR Recommendations and Reports, MMWR Policy Notes, and Immunization Schedules (Child/Adolescent, Adult)

Recommendations for routine use of vaccines in children, adolescents, and adults are developed by the Advisory Committee on Immunization Practices (ACIP). ACIP is chartered as a Federal Advisory Committee to provide expert external advice and guidance to the Director of CDC on use of vaccines and related agents for the control of vaccine preventable diseases in the civilian population of the United States. Recommendations for routine use of vaccines in children and adolescents are harmonized to the greatest extent possible with recommendations made by the American Academy of Pediatrics (AAP), the American Academy of Family Physicians (AAFP), and the American College of Obstetricians and Gynecologists (ACOG). Recommendations for routine use of vaccinations in adults are harmonized with recommendations of AAFP, ACOG, and the American College of Physicians (ACP). ACIP recommendations are forwarded to CDC's Director and once adopted become official CDC policy. These recommendations are then published in CDC's Morbidity and Mortality Weekly Report (MMWR). Additional information is available at https://www.cdc.gov/vaccines/acip.

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# Prevention and Control of Seasonal Influenza with Vaccines: Recommendations of the Advisory Committee on Immunization Practices — United States, 2023–24 Influenza Season

Lisa A. Grohskopf, MD<sup>1</sup>; Lenee H. Blanton, MPH<sup>1</sup>; Jill M. Ferdinands, PhD<sup>1</sup>; Jessie R. Chung, MPH<sup>1</sup>, Karen R. Broder, MD<sup>2</sup>; H. Keipp Talbot, MD<sup>3</sup>

<sup>1</sup>Influenza Division, National Center for Immunization and Respiratory Diseases, CDC; <sup>2</sup>Immunization Safety Office, National Center for Emerging and Zoonotic Infectious Diseases, CDC; <sup>3</sup>Division of Infectious Diseases, Vanderbilt University Medical Center, Nashville, Tennessee

#### Summary

This report updates the 2022–23 recommendations of the Advisory Committee on Immunization Practices (ACIP) concerning the use of seasonal influenza vaccines in the United States (MMWR Recomm Rep 2022;71[No. RR-1]:1–28). Routine annual influenza vaccination is recommended for all persons aged  $\geq 6$  months who do not have contraindications. All seasonal influenza vaccines expected to be available in the United States for the 2023–24 season are quadrivalent, containing hemagglutinin (HA) derived from one influenza A(H1N1)pdm09 virus, one influenza A(H3N2) virus, one influenza B/Victoria lineage virus, and one influenza B/Yamagata lineage virus. Inactivated influenza vaccines (IIV4s), recombinant influenza vaccine (RIV4), and live attenuated influenza vaccine (LAIV4) are expected to be available.

For most persons who need only 1 dose of influenza vaccine for the season, vaccination should ideally be offered during September or October. However, vaccination should continue after October and throughout the season as long as influenza viruses are circulating and unexpired vaccine is available. Influenza vaccines might be available as early as July or August, but for most adults (particularly adults aged  $\geq 65$  years) and for pregnant persons in the first or second trimester, vaccination during July and August should be avoided unless there is concern that vaccination later in the season might not be possible. Certain children aged 6 months through 8 years need 2 doses; these children should receive the first dose as soon as possible after vaccine is available, including during July and August. Vaccination during July and August can be considered for children of any age who need only 1 dose for the season and for pregnant persons who are in the third trimester during these months if vaccine is available.

ACIP recommends that all persons aged  $\geq 6$  months who do not have contraindications receive a licensed and age-appropriate seasonal influenza vaccine. With the exception of vaccination for adults aged  $\geq 65$  years, ACIP makes no preferential recommendation for a specific vaccine when more than one licensed, recommended, and age-appropriate vaccine is available. ACIP recommends that adults aged  $\geq 65$  years preferentially receive any one of the following higher dose or adjuvanted influenza vaccines: quadrivalent high-dose inactivated influenza vaccine (HD-IIV4), quadrivalent recombinant influenza vaccine (RIV4), or quadrivalent adjuvanted inactivated influenza vaccine (aIIV4). If none of these three vaccines is available at an opportunity for vaccine administration, then any other age-appropriate influenza vaccine should be used.

Primary updates to this report include the following two topics: 1) the composition of 2023-24 U.S. seasonal influenza vaccines and 2) updated recommendations regarding influenza vaccination of persons with egg allergy. First, the composition of 2023-24U.S. influenza vaccines includes an update to the influenza A(H1N1)pdm09 component. U.S.-licensed influenza vaccines will contain HA derived from 1) an influenza A/Victoria/4897/2022 (H1N1)pdm09-like virus (for egg-based vaccines) or an influenza A/Wisconsin/67/2022 (H1N1)pdm09-like virus (for cell culture-based and recombinant vaccines); 2) an influenza A/Darwin/9/2021 (H3N2)-like virus (for egg-based vaccines) or an influenza A/Darwin/6/2021 (H3N2)-like virus (for cell culture-based and recombinant vaccines); 3) an influenza B/Austria/1359417/2021 (Victoria lineage)-like virus; and 4) an influenza B/Phuket/3073/2013 (Yamagata lineage)-like virus. Second, ACIP recommends that all persons aged  $\geq 6$  months with egg allergy should receive influenza vaccine. Any influenza vaccine (egg based or nonegg based) that is otherwise appropriate for the recipient's age and health status can be used. It is no longer recommended that persons who have had an allergic reaction to egg involving symptoms other than urticaria should be vaccinated in an inpatient or outpatient medical setting supervised by a health care provider who is able to recognize and manage severe allergic reactions if an egg-based vaccine is used. Egg allergy alone necessitates no additional safety measures for influenza vaccination beyond those recommended for any recipient of any vaccine,

**Corresponding author:** Lisa A. Grohskopf, Influenza Division, National Center for Immunization and Respiratory Diseases, CDC. Telephone: 404-639-2552; Email: lgrohskopf@cdc.gov. regardless of severity of previous reaction to egg. All vaccines should be administered in settings in which personnel and equipment needed for rapid recognition and treatment of acute hypersensitivity reactions are available. This report focuses on recommendations for the use of vaccines for the prevention and control of seasonal influenza during the 2023–24 influenza season in the United States. A brief summary of the recommendations and a link to the most recent Background Document containing additional information are available at https://www.cdc.gov/vaccines/hcp/acip-recs/vacc-specific/flu.html. These recommendations apply to U.S.-licensed influenza vaccines used according to Food and Drug Administration–licensed indications. Updates and other information are available from CDC's influenza website (https://www.cdc.gov/flu). Vaccination and health care providers should check this site periodically for additional information.

## Introduction

Influenza viruses typically circulate annually in the United States, most commonly from the late fall through the early spring. Most persons who become ill after influenza virus infection recover without serious complications or sequelae. However, influenza can be associated with serious illnesses, hospitalizations, and deaths, particularly among older adults, very young children, pregnant persons, and persons of all ages with certain chronic medical conditions (1-7). Influenza also is an important cause of missed work and school (8-10).

Routine annual influenza vaccination for all persons aged ≥6 months who do not have contraindications has been recommended by CDC and the Advisory Committee on Immunization Practices (ACIP) since 2010 (11). Vaccination provides important protection from influenza illness and its potential complications. The effectiveness of influenza vaccination varies depending on multiple factors, such as the age and health of the recipient; the type of vaccine administered; the types, subtypes (for influenza A), and lineages (for influenza B) of circulating influenza viruses; and the degree of similarity between circulating viruses and those included in the vaccine (12). During each of the six influenza seasons from 2010-11 through 2015-16, influenza vaccination prevented an estimated 1.6-6.7 million illnesses, 790,000-3.1 million outpatient medical visits, 39,000-87,000 hospitalizations, and 3,000-10,000 respiratory and circulatory deaths each season in the United States (13). During the severe 2017–18 season, notable for an unusually long duration of widespread high influenza activity throughout the United States and higher rates of outpatient visits and hospitalizations compared with recent seasons, vaccination prevented an estimated 7.1 million illnesses, 3.7 million medical visits, 109,000 hospitalizations, and 8,000 deaths (14), despite an overall estimated vaccine effectiveness of 38% (62% against influenza A[H1N1]pdm09 viruses, 22% against influenza A[H3N2] viruses, and 50% against influenza B viruses) (14).

Influenza circulated at historically low levels in the United States and globally during the 2020–21 influenza season, coincident with widespread implementation of nonpharmaceutical interventions (e.g., masking, social distancing, and suspension of in-person work and school) intended to prevent transmission of SARS-CoV-2 (the virus that causes COVID-19) (15). The 2021–22 influenza season saw increased activity compared with 2020–21, with influenza activity remaining elevated later into the spring than any previous season for which data are available (16). The 2022–23 season was marked by early influenza activity peaking in late November to early December (17). The timing, intensity, and severity of the 2023–24 influenza season cannot be predicted. Influenza vaccination remains an important tool for the prevention of potentially severe respiratory illness.

This report updates the 2022–23 ACIP recommendations regarding the use of seasonal influenza vaccines (*18*) and provides recommendations and guidance for vaccination providers regarding the use of influenza vaccines in the United States for the 2023–24 season. Various formulations of influenza vaccines are available (Table 1). Contraindications and precautions for the use of influenza vaccines are summarized (Tables 2 and 3). Abbreviations are used in this report to denote the various types of vaccines (Box). A summary of these recommendations and a Background Document containing additional information on influenza, influenza-associated illness, and influenza vaccines are available at https://www.cdc.gov/vaccines/hcp/acip-recs/vacc-specific/flu.html.

## **Methods**

ACIP provides annual recommendations for the use of influenza vaccines for the prevention and control of seasonal influenza in the United States. The ACIP Influenza Work Group meets by teleconference once to twice per month throughout the year. Work Group membership includes multiple voting members of ACIP, representatives of ACIP liaison organizations, and consultants. Discussions include topics such as influenza surveillance, vaccine effectiveness and safety, vaccination coverage, program feasibility, cost effectiveness, and vaccine supply. Presentations are requested from invited experts and published and unpublished data are discussed.

The Background Document that supplements this report contains literature related to recommendations made in previous seasons. The information included in the Background Document for such topics is not a systematic review; it is intended to provide an overview of background literature

#### TABLE 1. Influenza vaccines — United States, 2023-24 influenza season\*

Trade name (manufacturer)	Presentation	Age indication	μg HA (IIV4s and RIV4) or virus count (LAIV4) for each vaccine virus (per dose)	Route	Mercury (from thimerosal, if present) µg/0.5 mL
IIV4 (standard-dose, egg-based vac	cines <sup>†</sup> )				
Afluria Quadrivalent	0.5-mL PFS <sup>§</sup>	≥3 yrs <sup>§</sup>	15 μg/0.5 mL	IM¶	**
(Seqirus)	5.0-mL MDV <sup>§</sup>	≥6 mos <sup>§</sup> (needle and syringe) 18 through 64 yrs (jet injector)	7.5 μg/0.25 mL 15 μg/0.5 mL	IM¶	24.5
Fluarix Quadrivalent (GlaxoSmithKline)	0.5-mL PFS	≥6 mos	15 μg/0.5 mL	IM¶	—
FluLaval Quadrivalent (GlaxoSmithKline)	0.5-mL PFS	≥6 mos	15 μg/0.5 mL	IM¶	_
Fluzone Quadrivalent	0.5-mL PFS <sup>++</sup>	≥6 mos <sup>††</sup>	15 μg/0.5 mL	IM¶	_
(Sanofi Pasteur)	0.5-mL SDV <sup>††</sup>	≥6 mos <sup>††</sup>	$15 \mu g/0.5 m L$	IM¶	_
	5.0-mL MDV <sup>††</sup>	≥6 mos <sup>††</sup>	7.5 μg/0.25 mL 15 μg/0.5 mL	IM¶	25.0
ccIIV4 (standard-dose, cell culture-	based vaccine)				
Flucelvax Quadrivalent	0.5-mL PFS	≥6 mos	15 μg/0.5 mL	IM¶	_
(Seqirus)	5.0-mL MDV	≥6 mos	15 μg/0.5 mL	IM¶	25.0
HD-IIV4 (high-dose, egg-based vac	cine <sup>†</sup> )				
Fluzone High-Dose Quadrivalent (Sanofi Pasteur)	0.7-mL PFS	≥65 yrs	60 µg/0.7 mL	IM¶	_
allV4 (standard-dose, egg-based va	ccine <sup>†</sup> with MF59 adjuvant)				
Fluad Quadrivalent (Seqirus)	0.5-mL PFS	≥65 yrs	15 μg/0.5 mL	IM¶	—
RIV4 (recombinant HA vaccine) Flublok Quadrivalent (Sanofi Pasteur)	0.5-mL PFS	≥18 yrs	45 μg/0.5 mL	IM¶	_
LAIV4 (egg-based vaccine <sup>†</sup> ) FluMist Quadrivalent (AstraZeneca)	0.2-mL prefilled single-use intranasal sprayer	2 through 49 yrs	10 <sup>6.5–7.5</sup> fluorescent focus units/0.2 mL	NAS	_

Abbreviations: ACIP = Advisory Committee on Immunization Practices; HA = hemagglutinin; IIV4 = inactivated influenza vaccine, quadrivalent; IM = intramuscular; LAIV4 = live attenuated influenza vaccine, quadrivalent; MDV = multidose vial; PFS = prefilled syringe; RIV4 = recombinant influenza vaccine, quadrivalent; SDV = single-dose vial.

\* Manufacturer package inserts and updated CDC and ACIP guidance should be consulted for additional information concerning, but not limited to, indications, contraindications, warnings, and precautions. Package inserts for U.S.-licensed vaccines are available at https://www.fda.gov/vaccines-blood-biologics/vaccines/vaccines-licensed-use-united-states. Availability and characteristics of specific products and presentations might change or differ from what is described in this table and in the text of this report.

<sup>†</sup> Although a history of severe allergic reaction (e.g., anaphylaxis) to egg is a labeled contraindication to the use of egg-based IIV4s and LAIV4, ACIP recommends that all persons aged ≥6 months with egg allergy should receive influenza vaccine and that any influenza vaccine (egg based or nonegg based) that is otherwise appropriate for the recipient's age and health status can be used (see Persons with a History of Egg Allergy).

<sup>5</sup> The approved dose volume for Afluria Quadrivalent is 0.25 mL for children aged 6 through 35 months and 0.5 mL for persons aged ≥3 years. However, 0.25-mL prefilled syringes are no longer available. For children aged 6 through 35 months, a 0.25-mL dose must be obtained from a multidose vial.

<sup>1</sup> IM-administered influenza vaccines should be administered by needle and syringe only, with the exception of the MDV presentation of Afluria Quadrivalent, which may alternatively be given by the PharmaJet Stratis jet injector for persons aged 18 through 64 years only. For older children and adults, the recommended site for IM influenza vaccination is the deltoid muscle. The preferred site for infants and young children is the anterolateral aspect of the thigh. Additional specific guidance regarding site selection and needle length for IM administration is available in the General Best Practice Guidelines for Immunization available at https://www.cdc.gov/vaccines/hcp/acip-recs/general-recs/index.html.

\*\* Not applicable.

<sup>++</sup> Fluzone Quadrivalent is approved for children aged 6 through 35 months at either 0.25 mL or 0.5 mL per dose; however, 0.25-mL prefilled syringes are no longer available. If a prefilled syringe of Fluzone Quadrivalent is used for a child in this age group, the dose volume will be 0.5 mL per dose.

and is periodically updated with articles being identified primarily through a broad search for English-language articles on influenza and influenza vaccines. In general, longstanding recommendations in this document that were made in previous seasons reflect expert opinion, and systematic review and assessment of evidence was not performed. Systematic review and evidence assessment are not performed for minor wording changes to existing recommendations, changes in the Food and Drug Administration (FDA)–recommended viral antigen composition of seasonal influenza vaccines, and minor changes in guidance for the use of influenza vaccines (e.g., guidance for timing of vaccination and other programmatic issues, guidance for dosage in specific populations, guidance for selection of vaccines for specific populations that are already recommended for vaccination, and changes that reflect use that is consistent with FDA-licensed indications and prescribing information).

Typically, systematic review and evaluation of evidence using the Grading of Recommendations Assessment, Development

Vaccine type	Contraindications	Precautions
Egg-based IIV4s	<ul> <li>History of severe allergic reaction (e.g., anaphylaxis) to any component of the vaccine<sup>†</sup> or to a previous dose of any influenza vaccine (i.e., any egg-based IIV, ccIIV, RIV, or LAIV)<sup>§</sup></li> </ul>	<ul> <li>Moderate or severe acute illness with or without fever</li> <li>History of Guillain-Barré syndrome within 6 weeks of receipt of influenza vaccine</li> </ul>
cclIV4	<ul> <li>History of severe allergic reaction (e.g., anaphylaxis) to a previous dose of any cclIV or any component of cclIV4<sup>§</sup></li> </ul>	<ul> <li>Moderate or severe acute illness with or without fever</li> <li>History of Guillain-Barré syndrome within 6 weeks of receipt of influenza vaccine</li> <li>History of severe allergic reaction to a previous dose of any other influenza vaccine (i.e., any egg-based IIV, RIV, or LAIV)<sup>¶</sup></li> </ul>
RIV4	<ul> <li>History of severe allergic reaction (e.g., anaphylaxis) to a previous dose of any RIV or any component of RIV4<sup>§</sup></li> </ul>	<ul> <li>Moderate or severe acute illness with or without fever</li> <li>History of Guillain-Barré syndrome within 6 weeks of receipt of influenza vaccine</li> <li>History of severe allergic reaction to a previous dose of any other influenza vaccine (i.e., any egg-based IIV, ccIIV, or LAIV)<sup>¶</sup></li> </ul>
LAIV4	<ul> <li>History of severe allergic reaction (e.g., anaphylaxis) to any component of the vaccine<sup>†</sup> or to a previous dose of any influenza vaccine (i.e., any egg-based IIV, ccIIV, RIV, or LAIV)<sup>§</sup></li> <li>Concomitant aspirin- or salicylate-containing therapy in children and adolescents<sup>§</sup></li> <li>Children aged 2 through 4 years who have received a diagnosis of asthma or whose parents or caregivers report that a health care provider has told them during the preceding 12 months that their child had wheezing or asthma or whose medical record indicates a wheezing episode has occurred during the preceding 12 months</li> <li>Children and adults who are immunocompromised due to any cause, including but not limited to immunosuppression caused by medications, congenital or acquired immunodeficiency states, HIV infection, anatomic asplenia, or functional asplenia (e.g., due to sickle cell anemia)</li> <li>Close contacts and caregivers of severely immunosuppressed persons who require a protected environment</li> <li>Pregnancy</li> <li>Persons with active communication between the CSF and the oropharynx, nasopharynx, nose, or ear or any other cranial CSF leak</li> <li>Persons with cochlear implants**</li> <li>Receipt of influenza antiviral medication within the previous 48 hours for oseltamivir and zanamivir, previous 5 days for peramivir, and previous 17 days for baloxavir<sup>††</sup></li> </ul>	<ul> <li>Moderate or severe acute illness with or without fever</li> <li>History of Guillain-Barré syndrome within 6 weeks of receipt of influenza vaccine</li> <li>Asthma in persons aged ≥5 years</li> <li>Other underlying medical conditions that might predispose to complications after wild-type influenza infection (e.g., chronic pulmonary, cardiovascular [except isolated hypertension], renal, hepatic, neurologic, hematologic, or metabolic disorders [including diabetes mellitus])</li> </ul>

Abbreviations: ACIP = Advisory Committee on Immunization Practices; ccIIV = cell culture–based inactivated influenza vaccine (any valency); ccIIV4 = cell culture–based inactivated influenza vaccine, quadrivalent; CSF = cerebrospinal fluid; FDA = Food and Drug Administration; IIV = inactivated influenza vaccine (any valency); IIV4 = inactivated influenza vaccine, quadrivalent; LAIV = live attenuated influenza vaccine (any valency); RIV4 = recombinant influenza vaccine, quadrivalent; RIV = recombinant influenza vaccine (any valency); RIV4 = recombinant influenza vaccine, quadrivalent.

- \* Manufacturer package inserts and updated CDC and ACIP guidance should be consulted for additional information concerning, but not limited to, indications, contraindications, warnings, and precautions. When a contraindication is present, a vaccine should not be administered. When a precaution is present, vaccination should generally be deferred but might be indicated if the benefit of protection from the vaccine outweighs the risk for an adverse reaction (see the General Best Practice Guidelines for Immunization, available at https://www.cdc.gov/vaccines/hcp/acip-recs/general-recs/index.html). Package inserts for U.S.-licensed vaccines are available at https://www.fda.gov/vaccines-blood-biologics/vaccines/ucensed-use-united-states.
- <sup>†</sup> Although a history of severe allergic reaction (e.g., anaphylaxis) to egg is a labeled contraindication to the use of egg-based IIV4s and LAIV4, ACIP recommends that all persons aged ≥6 months with egg allergy should receive influenza vaccine, and that any influenza vaccine (egg based or nonegg based) that is otherwise appropriate for the recipient's age and health status can be used (see Persons with a History of Egg Allergy).
- <sup>§</sup> Labeled contraindication noted in package insert.
- <sup>1</sup> If administered, vaccination should occur in a medical setting and should be supervised by a health care provider who can recognize and manage severe allergic reactions. Providers can consider consultation with an allergist in such cases to assist in identification of the component responsible for the allergic reaction.
- \*\* Age-appropriate injectable vaccines are recommended for persons with cochlear implant because of the potential for CSF leak, which might exist for a period after implantation. Providers might consider consultation with a specialist concerning risk for persistent CSF leak if an age-appropriate inactivated or recombinant vaccine cannot be used.
- <sup>++</sup> Use of LAIV4 in context of influenza antivirals has not been studied; however, interference with activity of LAIV4 is biologically plausible, and this possibility is noted in the package insert for LAIV4. In the absence of data supporting an adequate minimum interval between influenza antiviral use and LAIV4 administration, the intervals provided are based on the half-life of each antiviral. The interval between influenza antiviral receipt and LAIV4 for which interference might potentially occur might be further prolonged in the presence of medical conditions that delay medication clearance (e.g., renal insufficiency). Influenza antivirals might also interfere with LAIV4 if initiated within 2 weeks after vaccination. Persons who receive antivirals during the period starting with the specified time before receipt of LAIV4 through 2 weeks after receipt of LAIV4 should be revaccinated with an age-appropriate IIV or RIV4.

TABLE 3. Influenza vaccine contraindications and precautions for persons with a history of severe allergic reaction to a previous dose of an
influenza vaccine* — United States, 2023–24 influenza season

Vaccine (of any valency) associated with	Available 2023–24 influenza vaccines			
previous severe allergic reaction (e.g., anaphylaxis)	Egg-based IIV4s and LAIV4	ccIIV4	RIV4	
Any egg-based IIV or LAIV	Contraindication <sup>†</sup>	Precaution <sup>§</sup>	Precaution <sup>§</sup>	
Any ccllV	Contraindication <sup>†</sup>	Contraindication <sup>†</sup>	Precaution§	
Any RIV	Contraindication <sup>†</sup>	Precaution <sup>§</sup>	Contraindication <sup>†</sup>	
Unknown influenza vaccine	Allergist consultation recommended			

Abbreviations: ACIP = Advisory Committee on Immunization Practices; ccIIV = cell culture–based inactivated influenza vaccine (any valency); ccIIV4 = cell culture–based inactivated influenza vaccine, quadrivalent; IIV = inactivated influenza vaccine (any valency); IIV4 = inactivated influenza vaccine, quadrivalent; LAIV = live attenuated influenza vaccine, quadrivalent; RIV = recombinant influenza vaccine (any valency); RIV4 = recombinant influenza vaccine (any valency); RIV4 = recombinant influenza vaccine, quadrivalent.

\* Manufacturer package inserts and updated CDC and ACIP guidance should be consulted for additional information, including, but not limited to indications, contraindications, warnings, and precautions. Package inserts for U.S.-licensed vaccines are available at https://www.fda.gov/vaccines-blood-biologics/vaccines/vaccines/licensed-use-united-states.

<sup>†</sup> When a contraindication is present, a vaccine should not be administered, consistent with the General Best Practice Guidelines for Immunization (Source: Kroger A, Bahta L, Hunter P. General best practice guidelines for immunization; https://www.cdc.gov/vaccines/hcp/acip-recs/general-recs/index.html). In addition to the contraindications based on history of severe allergic reaction to influenza vaccines that are noted in the table, each individual influenza vaccine is contraindicated for persons who have had a severe allergic reaction (e.g., anaphylaxis) to any component of that vaccine. Vaccine components can be found in package inserts. Although a history of severe allergic reaction (e.g., anaphylaxis) to egg is a labeled contraindication to the use of egg-based IIV4s and LAIV4, ACIP recommends that all persons aged ≥6 months with egg allergy should receive influenza vaccine, and that any influenza vaccine (egg based or nonegg based) that is otherwise appropriate for the recipient's age and health status can be used (see Persons with a History of Egg Allergy).

<sup>§</sup> When a precaution is present, vaccination should generally be deferred but might be indicated if the benefit of protection from the vaccine outweighs the risk for an adverse reaction, consistent with the General Best Practice Guidelines for Immunization (Source: Kroger A, Bahta L, Hunter P. General best practice guidelines for immunization; https://www.cdc.gov/vaccines/hcp/acip-recs/general-recs/index.html). Providers can consider using the following vaccines in these instances; however, vaccination should occur in an inpatient or outpatient medical setting with supervision by a health care provider who is able to recognize and manage severe allergic reactions: 1) for persons with a history of severe allergic reaction (e.g., anaphylaxis) to any egg-based IIV or LAIV of any valency, the provider can consider administering RIV4; and 3) for persons with a history of severe allergic reaction (e.g., anaphylaxis) to any ccIIV of any valency, the provider can consider administering RIV4; and 3) for persons with a history of severe allergic reaction (e.g., anaphylaxis) to any RIV of any valency, the provider can consider administering RIV4; and 3) for persons with a niltergist to help determine which vaccine component is responsible for the allergic reaction.

and Evaluation (GRADE) approach (19) are performed for new recommendations or substantial changes in the current recommendations (e.g., expansion of the recommendation for influenza vaccination to new populations not previously recommended for vaccination or potential preferential recommendations for specific vaccines).

Evidence is reviewed with the ACIP influenza Work Group, and Work Group considerations are included within the ACIP Evidence to Recommendations framework (EtR) (20) to inform the development of recommendations that are proposed for vote by the ACIP. Systematic review, GRADE, and the ACIP EtR framework were used in the development of the updated recommendations for influenza vaccination of persons with egg allergy discussed in this report.

# **Primary Changes and Updates**

Primary changes and updates to the recommendations described in this report include 1) the composition of 2023–24 U.S. seasonal influenza vaccines; 2) updated recommendations regarding influenza vaccination of persons with egg allergy. Information relevant to these changes includes the following:

• The composition of the 2023–24 U.S. seasonal influenza vaccines includes an update to the influenza A(H1N1)pdm09 component. For the 2023–24 season, U.S.-licensed influenza vaccines will contain hemagglutinin (HA)

derived from 1) an influenza A/Victoria/4897/2022 (H1N1)pdm09-like virus (for egg-based vaccines) or an influenza A/Wisconsin/67/2022 (H1N1)pdm09-like virus (for cell culture-based and recombinant vaccines); 2) an influenza A/Darwin/9/2021 (H3N2)-like virus (for egg-based vaccines) or an influenza A/Darwin/6/2021 (H3N2)-like virus (for cell culture-based and recombinant vaccines); 3) an influenza B/Austria/1359417/2021 (Victoria lineage)-like virus; and 4) an influenza B/Phuket/3073/2013 (Yamagata lineage)-like virus.

Recommendations for the composition of Northern Hemisphere influenza vaccines are made by the World Health Organization (WHO), which organizes a consultation, usually in February of each year. Surveillance data are reviewed and candidate vaccine viruses are discussed. Information about the WHO meeting of February 2023 for selection of the 2023-24 Northern Hemisphere influenza vaccine composition is available at https://www.who.int/publications/m/item/recommendedcomposition-of-influenza-virus-vaccines-for-use-inthe-2023-2024-northern-hemisphere-influenza-season. Subsequently, FDA, which has regulatory authority over vaccines in the United States, convenes a meeting of its Vaccines and Related Biological Products Advisory Committee (VRBPAC). This committee considers the recommendations of WHO, reviews and discusses similar

# BOX. Abbreviation conventions for influenza vaccines discussed in this report

- Main influenza vaccine types:
  - IIV = inactivated influenza vaccine
  - RIV = recombinant influenza vaccine
  - LAIV = live attenuated influenza vaccine
- Numerals following letter abbreviations indicate valency (the number of influenza virus hemagglutinin antigens represented in the vaccine):
  - **4** for quadrivalent vaccines: one A(H1N1), one A(H3N2), and two B viruses (one from each lineage)
  - **3** for trivalent vaccines: one A(H1N1), one A(H3N2), and one B virus (from one lineage)
- All influenza vaccines expected to be available in the United States for the 2023–24 season are quadrivalent vaccines. However, abbreviations for trivalent vaccines (e.g., IIV3) might be used in this report when discussing information specific to trivalent vaccines.
- Abbreviations for general vaccine categories (e.g., IIV) might be used when discussing information that is not specific to valency or to a specific vaccine in that category.
- Prefixes are used when necessary to refer to certain specific IIVs:
  - **a** for MF59-adjuvanted inactivated influenza vaccine (e.g., aIIV3 and aIIV4)
  - **cc** for cell culture–based inactivated influenza vaccine (e.g., ccIIV3 and ccIIV4)
  - **HD** for high-dose inactivated influenza vaccine (e.g., HD-IIV3 and HD-IIV4)
  - **SD** for standard-dose inactivated influenza vaccine (e.g., SD-IIV3 and SD-IIV4)

data, and makes a final decision regarding the composition of influenza vaccines licensed and marketed in the United States. Materials from the VRBPAC discussion on March 7, 2023, during which the composition of the 2023–24 U.S. influenza vaccines was discussed, are available at https://www.fda.gov/advisory-committees/ advisory-committee-calendar/vaccines-and-relatedbiological-products-advisory-committee-march-7-2023meeting-announcement.

• Regarding influenza vaccination of persons with egg allergy, ACIP recommends that all persons aged ≥6 months with egg allergy should receive influenza vaccine. Any influenza vaccine (egg based or nonegg based) that is otherwise appropriate for the recipient's age and health status can be used. It is no longer recommended that

persons who have had an allergic reaction to egg involving symptoms other than urticaria should be vaccinated in an inpatient or outpatient medical setting supervised by a health care provider who is able to recognize and manage severe allergic reactions if an egg-based vaccine is used. Egg allergy alone necessitates no additional safety measures for influenza vaccination beyond those recommended for any recipient of any vaccine, regardless of severity of previous reaction to egg. All vaccines should be administered in settings in which personnel and equipment needed for rapid recognition and treatment of acute hypersensitivity reactions are available.

To inform this recommendation, a systematic review and GRADE of evidence concerning safety of influenza vaccination for egg-allergic persons was conducted. A summary of this review and the GRADE evidence tables is available at https://www.cdc.gov/vaccines/acip/recs/grade/ influenza-egg-allergy.html. A summary of the ACIP EtR framework is available at https://www.cdc.gov/vaccines/ acip/recs/grade/influenza-egg-allergy-etr.html.

# Recommendations for the Use of Influenza Vaccines, 2023–24

## **Groups Recommended for Vaccination**

Routine annual influenza vaccination of all persons aged  $\geq 6$  months who do not have contraindications continues to be recommended. Influenza vaccines expected to be available for the 2023–24 season, their age indications, and their presentations are described (Table 1). Available influenza vaccines and age indications are expected to be similar to those of the 2022–23 season. Recommendations regarding timing of vaccination, considerations for specific populations, the use of specific vaccines, and contraindications and precautions are summarized in the sections that follow.

## **Timing of Vaccination**

Because timing of the onset, peak, and decline of influenza activity varies, the ideal time to start vaccinating cannot be predicted each season. Decisions about timing necessitate balancing considerations regarding this unpredictability of the influenza season, possible waning of vaccine-induced immunity over the course of a season, and programmatic considerations. For most persons who need only 1 dose of influenza vaccine for the season, vaccination should ideally be offered during September or October. However, vaccination should continue after October and throughout the influenza season as long as influenza viruses are circulating and unexpired vaccine is available. Influenza vaccines might be available as early as July or August; however, vaccination during these months is not recommended for most groups because of the possible waning of immunity over the course of the influenza season (21–37). However, vaccination of such persons during July and August can be considered in instances where there is concern that the recipient will not be vaccinated at a later date. Considerations for timing of vaccination include the following:

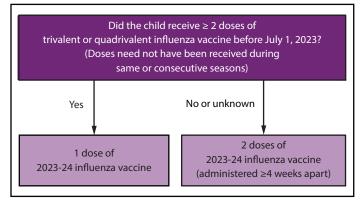
- For most adults (particularly adults aged ≥65 years) and for pregnant persons in the first or second trimester: Vaccination during July and August should be avoided unless there is concern that vaccination later in the season might not be possible.
- Children who require 2 doses: Certain children aged 6 months through 8 years require 2 doses of influenza vaccine for the season (see Children Aged 6 Months Through 8 Years: Number of Influenza Vaccine Doses) (Figure). These children should receive their first dose as soon as possible (including during July and August, if vaccine is available) to allow the second dose (which must be administered ≥4 weeks later) to be received, ideally, by the end of October.
- Children who require only 1 dose: Vaccination during July and August can be considered for children of any age who need only 1 dose of influenza vaccine for the season. Although waning of immunity after vaccination over the course of the season has been observed among all age groups (21–37), there are fewer published studies reporting results specifically among children (21,30,32,33,37). Moreover, children in this group might visit health care providers during the late summer months for medical examinations before the start of school. Vaccination can be considered at this time because it represents a vaccination opportunity.
- **Pregnant persons in the third trimester:** Vaccination during July and August can be considered for pregnant persons who are in the third trimester during these months because vaccination has been associated in multiple studies with reduced risk for influenza illness in their infants during the first months after birth, when they are too young to receive influenza vaccine (38–41). For pregnant persons in the first or second trimester during July and August, waiting to vaccinate until September or October is preferable, unless there is concern that later vaccination might not be possible.

Community vaccination programs should balance maximizing the likelihood of persistence of vaccine-induced protection through the season with avoiding missed opportunities to vaccinate or vaccinating after onset of influenza circulation occurs. Efforts should be structured to optimize vaccination coverage before influenza activity in the community begins. Vaccination should continue to be offered as long as influenza viruses are circulating and unexpired vaccine is available. To avoid missed opportunities for vaccination, providers should offer vaccination during routine health care visits and hospitalizations. No recommendation is made for revaccination (i.e., providing a booster dose) later in the season of persons who have been fully vaccinated for the season, regardless of when the current season vaccine was received.

Optimally, vaccination should occur before onset of influenza activity in the community. However, because timing of the onset, peak, and decline of influenza activity varies, the ideal time to start vaccinating cannot be predicted each season. Moreover, more than one outbreak might occur in a community in a single season. In the United States, localized outbreaks indicating the start of seasonal influenza activity can occur as early as October. However, in 30 (77%) of 39 influenza seasons from 1982–83 through 2021–22, peak influenza activity did not occur until January or later, and in 24 (62%) seasons, the peak was in February or later (42). Activity peaked in February in 17 (44%) of these seasons (42).

An increasing number of observational studies (21-37) have reported decreases in vaccine effectiveness with increasing time postvaccination within a single influenza season. Waning effects have not been observed consistently across age groups, influenza viruses (types, subtypes, and lineages), or seasons. Certain studies suggest waning occurs to a greater degree against influenza A(H3N2) viruses than against influenza A(H1N1) or influenza B viruses (25,31,35). This effect also might vary with recipient age; in certain studies, waning was more pronounced among older adults (21,22,24,31,34) and younger children (21). Relatively fewer reports include results specific to children (21,30,32,33,37); findings suggestive of waning have been reported in certain studies (21,32,33,37) but not others (30). Rates of decline in vaccine effectiveness also varied. A multiseason (2011-12 through 2014-15) analysis from the U.S. Influenza Vaccine Effectiveness (U.S. Flu VE) Network found that vaccine effectiveness decreased by approximately 7% per month for influenza A(H3N2) and influenza B and 6%-11% per month for influenza A(H1N1)pdm09 (23). A prospective test-negative case control study of the 2015-16 through 2019-20 seasons observed an average decline in vaccine effectiveness against influenza-associated hospitalizations of 1.3% per month among children aged ≤8 years and 4.7% per month among those aged 9 through 17 years (37). In the Hospitalized Adult Influenza Vaccine Effectiveness Network (HAIVEN) during the 2015–16 through 2018–19 seasons, vaccine effectiveness against influenza-associated hospitalizations declined by approximately 8%-9% per month for all adults and

FIGURE. Influenza vaccine dosing algorithm for children aged 6 months through 8 years\* — Advisory Committee on Immunization Practices, United States, 2023–24 influenza season



\* Children aged 6 months through 8 years who require 2 doses of influenza vaccine should receive their first dose as soon as possible (including during July and August, if vaccine is available) to allow the second dose (which must be administered ≥4 weeks later) to be received, ideally, by the end of October. For children aged 8 years who require 2 doses of vaccine, both doses should be administered even if the child turns age 9 years between receipt of dose 1 and dose 2.

approximately 10%–11% per month for those aged  $\geq 65$  years (24). An analysis of the 2010-11 through 2013-14 seasons noted estimated effectiveness ranging from 54% to 67% during days 0-180 postvaccination; estimated vaccine effectiveness was not significant during the period between days 181 and 365 (32). A multiseason analysis (2010–11 through 2014–15) conducted in Europe noted a decline in vaccine effectiveness to 0% at 111 days postvaccination against influenza A(H3N2) viruses. Vaccine effectiveness against influenza B viruses decreased more slowly, and vaccine effectiveness against influenza A(H1N1)pdm09 viruses remained roughly stable at 50%-55% throughout the influenza season (25). A metaanalysis of 14 studies examining waning of influenza vaccine effectiveness using the test-negative design found a significant decline in effectiveness within 180 days after vaccination against influenza A(H3N2) and influenza B but not against influenza A(H1N1) (43). In addition to the factors observed to be associated with waning immunity across studies, observed decreases in protection might be at least in part attributable to bias, unmeasured confounding, or the late-season emergence of antigenic drift variants of influenza viruses that are less wellmatched to the vaccine viruses.

Varying data concerning the presence and rate of waning immunity after influenza vaccination, coupled with the unpredictable timing of the influenza season each year, prevent determination of an optimal time to vaccinate each season. Programmatic issues also are a consideration. Although delaying vaccination might result in greater immunity later in the season, deferral also might result in missed opportunities to vaccinate as well as difficulties in vaccinating a population within a more constrained period. The potential contributions of these factors among persons aged  $\geq 65$  years have been assessed using a simulated mathematical model examining various scenarios of vaccination timing, timing of onset of the influenza season, vaccine effectiveness, and rate of waning (44). In this model, during an influenza season beginning in October and peaking in January, delaying vaccination until October resulted in more hospitalizations if >14% of persons aged  $\geq 65$  years who would have been vaccinated in August or September did not get vaccinated. However, these predictions varied considerably with assumed timing of season onset, rate of waning immunity, and vaccine effectiveness.

Vaccination efforts should continue throughout the season because the duration of the influenza season varies, and influenza activity might not occur in certain communities until February, March, or later (17). Providers should offer influenza vaccine at healthcare visits to those not yet vaccinated, and organized vaccination campaigns should continue throughout the influenza season, including after influenza activity has begun in the community. Although vaccination by the end of October is recommended, vaccine administered in December or later, even if influenza activity has already begun, might be beneficial in most influenza seasons. Providers should offer influenza vaccination to unvaccinated persons who have already become ill with influenza during the season because the vaccine might protect them against other circulating influenza viruses.

## Guidance for Influenza Vaccination in Specific Populations and Situations

## Populations at Higher Risk for Medical Complications Attributable to Severe Influenza

All persons aged  $\geq 6$  months who do not have contraindications should be vaccinated annually. However, vaccination to prevent influenza is particularly important for persons who are at increased risk for severe illness and complications from influenza and for influenza-related outpatient, emergency department, or hospital visits. When vaccine supply is limited, vaccination efforts should focus on vaccination of persons at higher risk for medical complications attributable to severe influenza who do not have contraindications. These persons include the following (order of listing does not imply hierarchy or prioritization among these populations):

- All children aged 6 through 59 months.
- All persons aged  $\geq 50$  years.
- Adults and children who have chronic pulmonary (including asthma), cardiovascular (excluding isolated hypertension), renal, hepatic, neurologic, hematologic, or metabolic disorders (including diabetes mellitus).

- Persons who are immunocompromised due to any cause (including but not limited to immunosuppression caused by medications or HIV infection).
- Persons who are or will be pregnant during the influenza season.
- Children and adolescents (aged 6 months through 18 years) who are receiving aspirin- or salicylate-containing medications and who might be at risk for experiencing Reye syndrome after influenza virus infection.
- Residents of nursing homes and other long-term care facilities.
- American Indian or Alaska Native persons.
- Persons who are extremely obese (body mass index ≥40 for adults).

An age-appropriate IIV4 or RIV4 is suitable for all persons recommended for vaccination, including those in the risk groups listed. LAIV4 is not recommended for certain populations, including certain of these listed groups. Contraindications and precautions for the use of LAIV4 are noted (Table 2).

## Persons Who Live with or Care for Persons at Higher Risk for Influenza-Related Complications

All persons aged  $\geq 6$  months without contraindications should be vaccinated annually. However, emphasis also should be placed on vaccination of persons who live with or care for those who are at increased risk for medical complications attributable to severe influenza. When vaccine supply is limited, vaccination efforts should focus on administering vaccination to persons at higher risk for influenza-related complications as well as persons who live with or care for such persons, including the following:

- Health care personnel, including all paid and unpaid persons working in health care settings who have the potential for exposure to patients or to infectious materials. These personnel might include but are not limited to physicians, nurses, nursing assistants, nurse practitioners, physician assistants, therapists, technicians, emergency medical service personnel, dental personnel, pharmacists, laboratory personnel, autopsy personnel, students and trainees, contractual staff members, and others not directly involved in patient care but who might be exposed to infectious agents (e.g., clerical, dietary, housekeeping, laundry, security, maintenance, administrative, and billing staff members and volunteers). ACIP guidance for vaccination of health care personnel has been published previously (45).
- Household contacts (including children aged ≥6 months) and caregivers of children aged ≤59 months (<5 years) and adults aged ≥50 years, particularly contacts of children aged <6 months.</li>

• Household contacts (including children aged ≥6 months) and caregivers of persons with medical conditions that put them at higher risk for severe complications from influenza.

Health care personnel and persons who are contacts of persons in these groups (with the exception of contacts of severely immunocompromised persons who require a protected environment) may receive any influenza vaccine that is otherwise indicated. Persons who care for severely immunocompromised persons requiring a protected environment should not receive LAIV4. ACIP and the Healthcare Infection Control Practices Advisory Committee (HICPAC) have previously recommended that health care personnel who receive LAIV should avoid providing care for severely immunocompromised persons requiring a protected environment for 7 days after vaccination and that hospital visitors who have received LAIV should avoid contact with such persons for 7 days after vaccination (46). However, such persons need not be restricted from caring for or visiting less severely immunocompromised persons.

## Influenza Vaccination of Persons with COVID-19

Specific data concerning the optimal timing of influenza vaccination of persons with COVID-19 illness are not available. For those who have moderate or severe COVID-19, vaccination should usually be deferred until they have recovered from the acute illness, consistent with General Best Practice Guidelines for Immunization (47). For those with mild or asymptomatic COVID-19, further deferral might be considered to avoid confusing COVID-19 symptoms with potential postvaccination reactions. Other considerations for determination of when to vaccinate include current local influenza activity, the recipient's individual risk for severe influenza illness, current or recent use of immunosuppressive therapeutic agents that might blunt immune response to vaccines, and risk for exposing others in the vaccination setting to COVID-19. Information concerning precautions for persons with COVID-19 is available at https://www.cdc. gov/coronavirus/2019-ncov/hcp/duration-isolation.html.

## Children Aged 6 Through 35 Months: Influenza Vaccine Dose Volumes

Five IIV4s are approved for children aged  $\geq 6$  months (Table 1). Four of these vaccines are egg based (Afluria Quadrivalent, Fluarix Quadrivalent, FluLaval Quadrivalent, and Fluzone Quadrivalent), and one is cell culture-based (Flucelvax Quadrivalent). For these vaccines, the approved dose volumes for children aged 6 through 35 months are as follows (Table 4):

• Afluria Quadrivalent: 0.25 mL per dose. However, 0.25-mL prefilled syringes are no longer available. For

children aged 6 through 35 months, a 0.25-mL dose must be obtained from a multidose vial (*48*).

- Fluarix Quadrivalent: 0.5 mL per dose (49).
- Flucelvax Quadrivalent: 0.5 mL per dose (50).
- FluLaval Quadrivalent: 0.5 mL per dose (51).
- Fluzone Quadrivalent: Either 0.25 mL or 0.5 mL per dose. Per the package insert, each dose may be given at either volume (*52*); however, 0.25-mL prefilled syringes are no longer available.

For all of these IIV4s, persons aged  $\geq 36$  months ( $\geq 3$  years) should receive 0.5 mL per dose. Alternatively, healthy children aged  $\geq 24$  months ( $\geq 2$  years) can receive LAIV4, 0.2 mL intranasally (0.1 mL in each nostril) (53). LAIV4 is not recommended for certain populations and is not approved for children aged <2 years or adults >49 years (see Contraindications and Precautions for the Use of LAIV4) (Table 2). RIV4 is not approved for children aged <18 years (54). High-dose inactivated influenza vaccine (HD-IIV4) (55) and adjuvanted inactivated influenza vaccine (aIIV4) (56) are not approved for persons aged <65 years.

Care should be taken to administer an age-appropriate vaccine at the appropriate volume for each dose. For IIV4s, the recommended volume may be administered from a prefilled syringe containing the appropriate volume (as supplied by the manufacturer), a single-dose vial, or a multidose vial. Singledose vials should be used for only 1 dose, and multidose vials should be used only for the maximum number of doses specified in the package insert. Any vaccine remaining in a vial after the maximum number of doses has been removed should be discarded, regardless of the volume of the doses obtained or any remaining volume in the vial.

## Children Aged 6 Months Through 8 Years: Number of Influenza Vaccine Doses

Children aged 6 months through 8 years require 2 doses of influenza vaccine administered a minimum of 4 weeks apart during their first season of vaccination for optimal protection (57-60). Determination of the number of doses needed is based on 1) the child's age at the time of the first dose of 2023–24 influenza vaccine and 2) the number of doses of influenza vaccine received in previous influenza seasons.

- For children aged 6 months through 8 years, the number of doses of influenza vaccine needed for the 2023–24 influenza season is determined as follows (Figure):
  - Those who have previously received ≥2 total doses of trivalent or quadrivalent influenza vaccine ≥4 weeks apart before July 1, 2023, require only 1 dose for the 2023–24 season. The previous 2 doses of influenza vaccine do not need to have been received in the same season or consecutive seasons.

- Those who have not previously received ≥2 doses of trivalent or quadrivalent influenza vaccine ≥4 weeks apart before July 1, 2023, or whose previous influenza vaccination history is unknown, require 2 doses for the 2023–24 season. The interval between the 2 doses should be ≥4 weeks. Children aged 6 months through 8 years who require 2 doses of influenza vaccine should receive their first dose as soon as possible (including during July and August, if vaccine is available) to allow the second dose (which must be administered ≥4 weeks later) to be received, ideally, by the end of October. For children aged 8 years who require 2 doses of vaccine, both doses should be administered even if the child turns age 9 years between receipt of dose 1 and dose 2.
- Adults and children aged ≥9 years need only 1 dose of influenza vaccine for the 2023–24 season.

## **Pregnant Persons**

Pregnant and postpartum persons have been observed to be at higher risk for severe illness and complications from influenza, particularly during the second and third trimesters. Influenza vaccination during pregnancy is associated with reduced risk for respiratory illness and influenza among pregnant and postpartum persons as well as infants during the first months of life (38-41,61). ACIP and the American College of Obstetricians and Gynecologists recommend that persons who are pregnant or who might be pregnant or postpartum during the influenza season receive influenza vaccine (62). Any licensed, recommended, and age-appropriate IIV4 or RIV4 may be used. LAIV4 should not be used during pregnancy but can be used postpartum. Influenza vaccine can be administered at any time during pregnancy (i.e., during any trimester), before and during the influenza season. Early vaccination (i.e., during July and August) can be considered for persons who are in the third trimester during these months if vaccine is available because this can provide protection for the infant during the first months of life when they are too young to be vaccinated (38-41,61).

Although experience with the use of IIVs during pregnancy is substantial, data specifically reflecting administration of influenza vaccines during the first trimester are limited. Most studies have not noted an association between influenza vaccination and adverse pregnancy outcomes, including spontaneous abortion (miscarriage) (63–73). One observational Vaccine Safety Datalink (VSD) study conducted during the 2010–11 and 2011–12 seasons noted an association between receipt of IIV containing influenza A(H1N1)pdm09 and risk for miscarriage in the 28 days after receipt of IIV, when an H1N1pdm09-containing vaccine also had been received the previous season (74). However, in a larger VSD

TABLE 4. Dose volumes for inactivated influenza vaccines approved for children aged 6 through 35 months<sup>\*</sup> — United States, 2023–24 influenza season

Trade name (manufacturer)	Dose volume for children aged 6 through 35 mos (µg HA per vaccine virus)
Afluria Quadrivalent (Seqirus)	0.25 mL (7.5 μg) <sup>†</sup>
Fluarix Quadrivalent (GlaxoSmithKline)	0.5 mL (15 μg)
Flucelvax Quadrivalent (Segirus)	0.5 mL (15 μg)
FluLaval Quadrivalent (GlaxoSmithKline)	0.5 mL (15 μg)
Fluzone Quadrivalent (Sanofi Pasteur)	0.5 mL (15 μg) <sup>§</sup>

Abbreviation: HA = hemagglutinin.

\* For persons aged ≥36 months (≥3 years), the dose volume is 0.5 mL per dose for all inactivated influenza vaccines with the exception of Fluzone High-Dose Quadrivalent (HD-IIV4), which is licensed for persons aged ≥65 years and for which the dose volume is 0.7 mL per dose.

<sup>+</sup> The approved dose volume for Afluria Quadrivalent is 0.25 mL for children aged 6 through 35 months and 0.5 mL for persons aged ≥3 years. However, 0.25-mL prefilled syringes are no longer available. For children aged 6 through 35 months, a 0.25-mL dose must be obtained from a multidose vial.

<sup>§</sup> Per the package insert, Fluzone Quadrivalent is approved for children aged 6 through 35 months at either 0.25 mL or 0.5 mL per dose; however, 0.25-mL prefilled syringes are no longer available. If a prefilled syringe of Fluzone Quadrivalent is used for a child in this age group, the dose volume will be 0.5 mL per dose. The 0.5-mL single-dose vials should be accessed for only 1 dose and multidose vials for only 10 doses, regardless of the volume of the doses obtained or any remaining volume in the vial. Any vaccine remaining in a vial after the maximum number of doses has been removed should be discarded.

follow-up study, IIV was not associated with an increased risk for miscarriage during the 2012–13, 2013–14, and 2014–15 seasons, regardless of previous season vaccination (*75*).

There is less experience with the use of more recently licensed influenza vaccines (e.g., quadrivalent, cell culture-based, and recombinant vaccines) during pregnancy compared with previously available products. For ccIIV, a review of Vaccine Adverse Event Reporting System (VAERS) reports from 2013 through 2020 (76) and a prospective cohort study conducted from 2017 through 2020 (77) did not reveal unexpected safety events among pregnant persons. Data from a randomized clinical trial conducted at Clinical Immunization Safety Assessment (CISA) Project sites comparing the safety of RIV4 versus IIV4 in 382 pregnant persons supported the safety of RIV4 in pregnancy (78). Pregnancy registries and surveillance studies exist for certain products, for which information can be found in package inserts.

#### **Older Adults**

ACIP recommends that adults aged ≥65 years preferentially receive any one of the following higher dose or adjuvanted influenza vaccines: quadrivalent high-dose inactivated influenza vaccine (HD-IIV4), quadrivalent recombinant influenza vaccine (RIV4), or quadrivalent adjuvanted inactivated influenza vaccine (aIIV4). If none of these three vaccines

is available at an opportunity for vaccine administration, then any other age-appropriate influenza vaccine should be administered (*79,80*).

Older adults (aged  $\geq 65$  years) are at increased risk for severe influenza-associated illness, hospitalization, and death compared with younger persons (4,17,81). Influenza vaccines are often less effective in this population (82). HD-IIV, RIV, and aIIV have been evaluated in comparison with nonadjuvanted SD-IIVs in this age group. Two of these vaccines, HD-IIV and RIV, are higher dose vaccines, which contain an increased dose of HA antigen per virus compared with nonadjuvanted SD-IIVs (60 µg for HD-IIV4 and 45 µg for RIV4, compared with 15 µg for standard-dose inactivated vaccines). The adjuvanted vaccine contains 15 µg of HA per virus, similarly to nonadjuvanted SD-IIVs, but contains the adjuvant MF59.

HD-IIV, RIV, and aIIV have shown relative benefit compared with SD-IIVs in certain studies, with the most evidence available for HD-IIV3. Randomized efficacy studies comparing these vaccines with nonadjuvanted SD-IIVs against laboratory-confirmed influenza outcomes are few in number (83-85) and cover few influenza seasons. Observational studies, predominantly retrospective cohort studies using diagnostic code-defined (rather than laboratory-confirmed) influenza outcomes, are more numerous and include more influenza seasons (86-96). Certain observational studies have reported relative benefit for HD-IIV, RIV, and aIIV in comparison with nonadjuvanted SD-IIVs, particularly in prevention of influenza-associated hospitalizations. The size of this relative benefit has varied from season to season and is not observed in all studies in all seasons, making it difficult to generalize the findings to all or most seasons. Studies directly comparing HD-IIV, RIV, and aIIV with one another are few and do not support a conclusion that any one of these vaccines is consistently superior to the others across seasons (86-88,91,97,98).

During the 2020–21 season, quadrivalent formulations of high-dose (HD-IIV4) and adjuvanted (aIIV4) influenza vaccines were introduced. Trivalent formulations of these vaccines are no longer available. Data summarizing comparisons of these newer quadrivalent formulations relative to nonadjuvanted SD-IIV4s are limited. In a pragmatic randomized open-label feasibility study of HD-IIV4 compared with SD-IIV4 conducted in Denmark among persons aged 65 through 79 years during the 2021–22 influenza season that collected data from health registries, HD-IIV4 was associated with lower risk for diagnostic code–defined pneumonia and influenza hospitalizations (relative vaccine effectiveness 64.4; 95% CI = 24.4–84.6) (*99*).

#### Immunocompromised Persons

ACIP recommends that persons with compromised immunity (including but not limited to persons with congenital and acquired immunodeficiency states, persons who are immunocompromised due to medications, and persons with anatomic and functional asplenia) should receive an age-appropriate IIV4 or RIV4. ACIP recommends that LAIV4 not be used for these groups because of the uncertain but biologically plausible risk for disease attributable to the live vaccine virus. Use of LAIV4 in persons with these and other conditions is discussed in more detail (see Dosage, Administration, Contraindications, and Precautions) (Table 2).

Immunocompromised states comprise a heterogeneous range of conditions with varying risks for severe infections. In many instances, limited data are available regarding the effectiveness of influenza vaccines in the setting of specific immunocompromised states (100). Timing of vaccination might be a consideration (e.g., vaccinating during a period either before or after an immunocompromising intervention). The Infectious Diseases Society of America has published detailed guidance for the selection and timing of vaccines for persons with specific immunocompromising conditions (101). Immune response to influenza vaccines might be blunted in persons with certain conditions, such as congenital immune deficiencies, and in persons receiving cancer chemotherapy, posttransplant regimens, or immunosuppressive medications.

## Persons with a History of Guillain-Barré Syndrome After Influenza Vaccination

A history of Guillain-Barré syndrome (GBS) within 6 weeks of a previous dose of any type of influenza vaccine is considered a precaution for influenza vaccination (Table 2). Persons who are not at higher risk for severe influenza complications (see Populations at Higher Risk for Medical Complications Attributable to Severe Influenza) and who are known to have experienced GBS within 6 weeks of a previous influenza vaccination typically should not be vaccinated. As an alternative to vaccination, providers might consider using influenza antiviral chemoprophylaxis for these persons (*102*). However, the benefits of influenza vaccination might outweigh the possible risks for certain persons who have a history of GBS within 6 weeks after receipt of influenza vaccine and who also are at higher risk for severe complications from influenza.

#### Persons with a History of Egg Allergy

ACIP recommends that all persons aged ≥6 months with egg allergy should receive influenza vaccine. Any influenza vaccine (egg based or nonegg based) that is otherwise appropriate for the recipient's age and health status can be used (https://www. cdc.gov/vaccines/acip/recs/grade/influenza-egg-allergy.html; https://www.cdc.gov/vaccines/acip/recs/grade/influenza-eggallergy-etr.html). It is no longer recommended that persons who have had an allergic reaction to egg involving symptoms other than urticaria should be vaccinated in an inpatient or outpatient medical setting supervised by a health care provider who is able to recognize and manage severe allergic reactions if an egg-based vaccine is used. Egg allergy alone necessitates no additional safety measures for influenza vaccination beyond those recommended for any recipient of any vaccine, regardless of severity of previous reaction to egg. All vaccines should be administered in settings in which personnel and equipment needed for rapid recognition and treatment of acute hypersensitivity reactions are available.

Most available influenza vaccines, with the exceptions of RIV4 (Flublok Quadrivalent, licensed for persons aged  $\geq 18$  years) and ccIIV4 (Flucelvax Quadrivalent, licensed for persons aged  $\geq 6$  months), are prepared by propagation of virus in embryonated eggs and might contain trace amounts of egg proteins, such as ovalbumin. Among those U.S.-licensed influenza vaccines for which ovalbumin content is reported, quantities are generally small ( $\leq 1 \mu g/0.5 mL$  dose) (Table 1).

The Joint Task Force on Practice Parameters of the American Academy of Allergy, Asthma & Immunology (AAAAI) and the American College of Allergy, Asthma, & Immunology (ACAAI) have recommended that administration of influenza vaccine to egg allergic persons requires no additional precautions other than those recommended for administration of any vaccine to any individual (103). Since the 2016–17 influenza season, the American Academy of Pediatrics (AAP) has recommended that no additional precautionary measures are needed when administering influenza vaccine to egg-allergic persons (104). A review of 20 studies (16 of IIVs, one of virosomal influenza vaccine, and three of LAIV) that examined reactions after administration of seasonal influenza vaccines to egg-allergic persons via either full single-dose or split-dose administration protocols (of which 13 reported inclusion of persons with a history of severe reaction or anaphylaxis to egg) included no reports of anaphylaxis (certainty level: very low) (105-122). Less severe reactions not described as anaphylaxis but involving cardiovascular symptoms, respiratory symptoms, angioedema, or generalized urticaria, or which involved treatment with medications or outpatient or emergency department attention occurred with low frequency (<1%). A similar profile was noted among 13 studies of monovalent H1N1pdm09 influenza vaccine, with no reported instances of anaphylaxis; frequency of reactions involving cardiovascular symptoms, respiratory symptoms, angioedema, or generalized urticaria events of <1%, and of events involving treatment with medications or outpatient or emergency department attention of approximately 1.5% (certainty level: very low) (114,115,123-132). One

instance of anaphylaxis meeting a surveillance case definition (i.e., Brighton Level 1 criteria) in a person with possible egg allergy was noted in a summary of VAERS reports after administration of monovalent H1N1pdm09 influenza vaccine during the 2009-10 influenza season; no denominator of doses administered was available but it was noted that approximately 127 million doses of monovalent IIV were distributed in the United States that season (133). Of note, severe allergic reactions after administration of the egg-free vaccine RIV to egg-allergic persons have been noted in VAERS reports (134-136). These reports highlight both the possibility that observed reactions after egg-based influenza vaccines might be caused by substances other than egg proteins and the importance of being prepared to recognize and manage serious hypersensitivity reactions when administering any vaccine to any recipient (regardless of allergy history).

Severe and life-threatening reactions to vaccines can rarely occur with any vaccine and in any vaccine recipient, regardless of allergy history. Providers are reminded that all vaccines should be administered in settings in which personnel and equipment needed for rapid recognition and treatment of acute hypersensitivity reactions are available. All vaccination providers should be familiar with their office emergency plan and be certified in cardiopulmonary resuscitation (47). No postvaccination observation period is recommended specifically for egg-allergic persons. However, ACIP recommends that vaccination providers consider observing patients (seated or supine) for 15 minutes after administration of any vaccine to decrease the risk for injury should syncope occur (47).

Although egg allergy is neither a contraindication nor precaution to the use of any influenza vaccine, there are contraindications and precautions related to allergies to vaccine components other than egg and to previous allergic reactions to influenza vaccines (see Persons with Previous Allergic Reactions to Influenza Vaccines and Dosage, Administration, Contraindications, and Precautions) (Tables 2 and 3).

# Persons with Previous Allergic Reactions to Influenza Vaccines

As is the case for all vaccines, influenza vaccines contain various components that might cause allergic and anaphylactic reactions. Most influenza vaccine package inserts list among contraindications to their use a history of previous severe allergic reaction (e.g., anaphylaxis) to any component of the vaccine or to a previous dose of any influenza vaccine (48,49,51-53,55,56). For ccIIV4 and RIV4, a history of a severe allergic reaction to any vaccine component is listed as a contraindication; no labeled contraindication is specified for a history of allergic reaction to any other influenza vaccine (50,54). However, severe allergic reactions, although rare, can occur after influenza vaccination, even among persons with no previous reactions or known allergies. Vaccine components and excipients can be found in package inserts. However, identifying the causative agent without further evaluation (i.e., through evaluation and testing for specific allergies) can be difficult. Severe allergic reactions after vaccination with an RIV have been reported to VAERS, certain of which have occurred among persons reporting previous allergic reactions to egg or to influenza vaccines and that might represent a predisposition to development of allergic manifestations in affected persons (134–136). Because these rare but severe allergic reactions can occur, ACIP recommends the following for persons with a history of severe allergic reaction to a previous dose of an influenza vaccine (Table 3):

- For egg-based IIV4s and LAIV4:
  - A history of severe allergic reaction (e.g., anaphylaxis) to any influenza vaccine (i.e., any egg-based IIV, ccIIV, RIV, or LAIV of any valency) is a contraindication to future receipt of all egg-based IIV4s and LAIV4. Each individual egg-based IIV4 and LAIV4 is also contraindicated for persons who have had a severe allergic reaction (e.g., anaphylaxis) to any component of that vaccine (excluding egg; see Persons with a History of Egg Allergy).
- For ccIIV4:
  - A history of a severe allergic reaction (e.g., anaphylaxis) to any egg-based IIV, RIV, or LAIV of any valency is a precaution for the use of ccIIV4. If ccIIV4 is administered in such instances, vaccination should occur in an inpatient or outpatient medical setting and should be supervised by a health care provider who is able to recognize and manage severe allergic reactions. Providers also can consider consultation with an allergist to help determine the vaccine component responsible for the allergic reaction.
  - A history of a severe allergic reaction (e.g., anaphylaxis) to any ccIIV of any valency or to any component of ccIIV4 is a contraindication to future receipt of ccIIV4.
- For RIV4:
  - A history of a severe allergic reaction (e.g., anaphylaxis) to any egg-based IIV, ccIIV, or LAIV of any valency is a precaution for the use of RIV4. If RIV4 is administered in such instances, vaccination should occur in an inpatient or outpatient medical setting and should be supervised by a health care provider who is able to recognize and manage severe allergic reactions. Providers can also consider consultation with an allergist to help determine the vaccine component responsible for the allergic reaction.

• A history of a severe allergic reaction (e.g., anaphylaxis) to any RIV of any valency or to any component of RIV4 is a contraindication to future receipt of RIV4.

## **Vaccination Issues for Travelers**

In temperate climate regions of the Northern and Southern Hemispheres, influenza activity is seasonal, occurring during approximately October–May in the Northern Hemisphere and April–September in the Southern Hemisphere. In the tropics, influenza might occur throughout the year (137). The timing of influenza activity and predominant types and subtypes of influenza viruses in circulation vary by geographic region (138). Travelers can be exposed to influenza when traveling to an area where influenza is circulating or when traveling as part of large tourist groups (e.g., on cruise ships) that include persons from areas of the world where influenza viruses are circulating (139–142).

Travelers who want to reduce their risk for influenza should consider influenza vaccination, preferably at least 2 weeks before departure. In particular, persons who live in the United States and are at higher risk for influenza complications and who were not vaccinated with influenza vaccine during the previous Northern Hemisphere fall or winter should consider receiving influenza vaccination before departure if they plan to travel to the tropics, to the Southern Hemisphere during the Southern Hemisphere influenza season (April–September), or with organized tourist groups or on cruise ships to any location. Persons at higher risk who received the previous season's influenza vaccine before travel should consult with their health care provider to discuss the risk for influenza and other travel-related diseases before embarking on travel during the summer. All persons (regardless of risk status) who are vaccinated in preparation for travel before the upcoming influenza season's vaccine is available, or who received the immediately preceding Southern Hemisphere influenza vaccine, should receive the current U.S. seasonal influenza vaccine the following fall or winter.

Influenza vaccine formulated for the Southern Hemisphere might differ in viral composition from the Northern Hemisphere vaccine. For persons traveling to the Southern Hemisphere during the Southern Hemisphere influenza season, receipt of a current U.S.-licensed Southern Hemisphere influenza vaccine formulation before departure might be reasonable but might not be feasible because of limited access to or unavailability of Southern Hemisphere formulations in the United States. Most Southern Hemisphere influenza vaccine formulations are not licensed in the United States, and they are typically not commercially available. More information on influenza vaccines and travel is available at https://wwwnc.cdc. gov/travel/diseases/influenza-seasonal-zoonotic-and-pandemic. Additional information on global influenza surveillance by region is available at https://www.who.int/tools/flunet.

## **Use of Influenza Antiviral Medications**

Administration of any IIV4 or RIV4 to persons receiving influenza antiviral medications for treatment or chemoprophylaxis of influenza is acceptable. Data concerning vaccination with LAIV4 in the setting of influenza antiviral use are not available. However, influenza antiviral medications might interfere with the action of LAIV4 because this vaccine contains live influenza viruses.

The package insert for LAIV4 notes that influenza antiviral agents might reduce the effectiveness of the vaccine if administered within the interval from 48 hours before to 14 days after vaccination (53). However, the newer influenza antivirals peramivir and baloxavir have longer half-lives than oseltamivir and zanamivir, approximately 20 hours for peramivir (143) and 79 hours for baloxavir (144), and could interfere with the replication of LAIV4, if administered >48 hours before vaccination. Potential interactions between influenza antivirals and LAIV4 have not been studied, and the ideal intervals between administration of these medications and LAIV4 are not known. Assuming a period of at least 5 half-lives for substantial decrease in drug levels (145), a reasonable assumption is that that peramivir might interfere with the mechanism of LAIV4 if administered from 5 days before through 2 weeks after vaccination and baloxavir might interfere if administered from 17 days before through 2 weeks after vaccination. The interval between influenza antiviral receipt and LAIV4 during which interference might occur could be further prolonged in the presence of medical conditions that delay medication clearance (e.g., renal insufficiency). Persons who receive these medications during these periods before or after receipt of LAIV4 should be revaccinated with another appropriate influenza vaccine (e.g., IIV4 or RIV4).

## Administration of Influenza Vaccines with Other Vaccines

IIV4s and RIV4 can be administered simultaneously or sequentially with other inactivated vaccines or live vaccines. Injectable vaccines that are given concomitantly should be administered at separate anatomic sites. COVID-19 vaccines that are administered at the same time as influenza vaccines that might be more likely to be associated with local injection site reactions (e.g., HD-IIV4 and aIIV4) should be given in different limbs, if possible. LAIV4 can be administered simultaneously with other live or inactivated vaccines. However, if two live vaccines are not given simultaneously, at least 4 weeks should pass after administration of one live vaccine (such as LAIV4) before another live vaccine is administered (47).

In recent years, multiple vaccines containing nonaluminum adjuvants have been licensed for use in the United States for the prevention of various infectious diseases. Examples include AS01<sub>B</sub> (in Shingrix, recombinant zoster subunit vaccine) (146),  $AS01_{\rm F}$  (in Arexy, respiratory syncytial virus vaccine) (147) MF59 (in Fluad Quadrivalent [aIIV4]) (56), and cytosine phosphoguanine oligodeoxynucleotide (in Heplisav-B, a recombinant hepatitis B surface antigen vaccine) (148). Data are limited regarding coadministration of these vaccines with other adjuvanted or nonadjuvanted vaccines, including COVID-19 vaccines. Coadministration of Shingrix with nonadjuvanted IIV4 has been studied, and no evidence of decreased immunogenicity or safety concerns was noted (149). Data on the immunogenicity and safety of simultaneous or sequential administration of two nonaluminum adjuvantcontaining vaccines are limited, and the ideal interval between such vaccines when given sequentially is not known. In the study of Shingrix and IIV4 (149), most reactogenicity symptoms resolved within 4 days. Because of the limited data on the safety of simultaneous administration of two or more vaccines containing nonaluminum adjuvants and the availability of nonadjuvanted influenza vaccine options, selection of a nonadjuvanted influenza vaccine may be considered in situations in which influenza vaccine and another vaccine containing a nonaluminum adjuvant are to be administered concomitantly. However, influenza vaccination should not be delayed if a specific vaccine is not available. As recommended for all vaccines, vaccines with nonaluminum adjuvants should be administered at separate anatomic sites from other vaccines that are given concomitantly (47).

For more recently introduced and new vaccines (e.g., respiratory syncytial virus [RSV] vaccine) data informing simultaneous administration with influenza vaccines might be limited or evolving. Providers should consult current CDC/ACIP recommendations and guidance for up-to-date information.

# Influenza Vaccine Composition and Available Vaccines

## Influenza Vaccine Composition for the 2023–24 Season

All influenza vaccines licensed in the United States will contain components derived from influenza viruses antigenically similar to those recommended by FDA (https://www.fda.gov/ advisory-committees/advisory-committee-calendar/vaccinesand-related-biological-products-advisory-committee-march-7-2023-meeting-announcement). All influenza vaccines expected to be available in the United States for the 2023–24 season will be quadrivalent vaccines. For the 2023–24 season, U.S. egg-based influenza vaccines (i.e., vaccines other than ccIIV4 and RIV4) will contain HA derived from

- an influenza A/Victoria/4897/2022 (H1N1)pdm09-like virus,
- an influenza A/Darwin/9/2021 (H3N2)-like virus,
- an influenza B/Austria/1359417/2021 (Victoria lineage)like virus, and
- an influenza B/Phuket/3073/2013 (Yamagata lineage)-like virus.

For the 2023–24 season, U.S. cell culture–based inactivated (ccIIV4) and recombinant (RIV4) influenza vaccines will contain HA derived from

- an influenza A/Wisconsin/67/2022 (H1N1)pdm09-like virus,
- an influenza A/Darwin/6/2021 (H3N2)-like virus,
- an influenza B/Austria/1359417/2021 (Victoria lineage)-like virus, and
- an influenza B/Phuket/3073/2013 (Yamagata lineage)-like virus.

## Vaccines Available for the 2023–24 Season

Availability of specific types and brands of licensed seasonal influenza vaccines in the United States is determined by the manufacturers of the vaccines. Information presented concerning vaccines expected to be available and their approved indications and usage reflects current knowledge and is subject to change.

Various influenza vaccines will be available for the 2023–24 season (Table 1). For many vaccine recipients, more than one type or brand of vaccine might be appropriate within approved indications and ACIP recommendations. A licensed influenza vaccine that is appropriate for the recipient's age and health status should be used. Specific age indications for licensed influenza vaccines are summarized (Table 1). Current prescribing information should be consulted for authoritative, up-to-date information. Contraindications and precautions for the different types of influenza vaccines are summarized (Table 2 and 3), as are dose volumes (Table 4).

Not all influenza vaccines are likely to be uniformly available in any specific practice setting or geographic locality. Vaccination should not be delayed to obtain a specific product when an appropriate one is available. Within these guidelines and approved indications, ACIP makes no preferential recommendation for the use of any one influenza vaccine over another when more than one licensed, recommended, and age-appropriate vaccine is available, with the exception of selection of influenza vaccines for persons aged  $\geq 65$  years (see Older Adults).

## Dosage, Administration, Contraindications, and Precautions

## Quadrivalent Inactivated Influenza Vaccines (IIV4s)

Available Vaccines. As in recent seasons, various inactivated influenza vaccines (IIVs) are expected to be available for 2023–24 (Table 1); all are expected to be quadrivalent (IIV4s). Standard-dose, nonadjuvanted IIV4s are licensed for persons aged as young as 6 months. However, for certain IIV4s, the approved dose volume for children aged 6 through 35 months differs from that for older children and adults (Table 4). Two IIV4s, the MF59-adjuvanted IIV4 (aIIV4) and the highdose IIV4 (HD-IIV4), are approved only for persons aged  $\geq$ 65 years. Care should be taken to administer the appropriate dose volume of an age-appropriate vaccine to each recipient.

Standard-dose, nonadjuvanted IIV4s contain 15 µg of HA per vaccine virus in a 0.5-mL dose (7.5  $\mu$ g of HA per vaccine virus in a 0.25-mL dose). For 2023-24, this category is expected to include five different vaccines (Table 1). Four of these are egg-based vaccines, and one is a cell culture-based vaccine (Flucelvax Quadrivalent [ccIIV4]). All are approved for persons aged  $\geq 6$  months. Egg-based and cell culture–based vaccines differ in the substrate in which reference vaccine viruses supplied to the manufacturer are propagated in quantities sufficient to produce the needed number of doses of vaccine. For the IIV4s Afluria Quadrivalent (48), Fluarix Quadrivalent (49), FluLaval Quadrivalent (51), and Fluzone Quadrivalent (52), reference vaccine viruses are propagated in eggs. For Flucelvax Quadrivalent (ccIIV4), reference vaccine viruses are propagated in Madin-Darby canine kidney cells instead of eggs (50).

Two additional IIV4s that will be available for the 2023–24 season are approved only for persons aged  $\geq$ 65 years. These vaccines are egg based. Quadrivalent high-dose inactivated influenza vaccine (Fluzone High-Dose Quadrivalent; HD-IIV4) contains 60 µg of HA per vaccine virus (240 µg total) in a 0.7-mL dose (55). Quadrivalent adjuvanted inactivated influenza vaccine (Fluad Quadrivalent; aIIV4) contains 15 µg of HA per vaccine virus (60 µg total) and MF59 adjuvant (56).

**Dosage and Administration.** Standard-dose nonadjuvanted IIV4s are approved for children aged as young as 6 months. Certain of these IIV4s are approved at different dose volumes for very young children than for older children and adults. Care should be taken to administer an age-appropriate vaccine at the approved dose volume for each needed dose (see Children Aged 6 Through 35 Months: Influenza Vaccine Dose Volumes) (Tables 1 and 4):

- Afluria Quadrivalent: The approved dose volume for children aged 6 through 35 months is 0.25 mL per dose. Persons aged ≥36 months (≥3 years) should receive 0.5 mL per dose (48).
- Fluarix Quadrivalent: The approved dose volume is 0.5 mL per dose for all persons aged ≥6 months (*49*).
- Flucelvax Quadrivalent: The approved dose volume is 0.5 mL per dose for all persons aged ≥6 months (*50*).
- FluLaval Quadrivalent: The approved dose volume is 0.5 mL per dose for all persons aged ≥6 months (51).
- Fluzone Quadrivalent: The approved dose volume for children aged 6 through 35 months is either 0.25 mL or 0.5 mL per dose. Persons aged ≥36 months (≥3 years) should receive 0.5 mL per dose (52).

If prefilled syringes are not available, the appropriate volume can be administered from a single-dose or multidose vial. If a 0.5-mL single-dose vial is used for a 0.25-mL dose for a child aged 6 through 35 months, only one half of the vial volume should be administered, and the remaining one half should be discarded. Of note, dose volume is distinct from the number of doses. Children in this age group who require 2 doses for 2023–24 need 2 separate doses administered ≥4 weeks apart, regardless of the specific IIV4 used and volume given for each dose (see Children Aged 6 Months Through 8 Years: Number of Influenza Vaccine Doses) (Figure).

For children aged 36 months (3 years) through 17 years and adults aged  $\geq 18$  years, the dose volume for IIV4s is 0.5 mL per dose, with the exception of Fluzone High-Dose Quadrivalent (HD-IIV4, licensed for persons aged  $\geq 65$  years), for which the correct volume is 0.7 mL per dose. If a smaller vaccine dose (e.g., 0.25 mL) is inadvertently administered to a person aged  $\geq 36$  months, the remaining volume needed to make a full dose should be administered during the same vaccination visit or, if measuring the needed remaining volume is a challenge, administering a repeat dose at the full volume is acceptable. If the error is discovered later (after the recipient has left the vaccination setting), a full dose should be administered as soon as the recipient can return. Vaccination with a formulation approved for adult use should be counted as a single dose if inadvertently administered to a child.

IIV4s are administered intramuscularly (IM). For adults and older children, the deltoid muscle is the preferred site. Infants and younger children should be vaccinated in the anterolateral thigh. Additional specific guidance regarding site selection and needle length for IM injection is provided in the General Best Practice Guidelines for Immunization (47). One IIV4, Afluria Quadrivalent, is licensed for IM injection via the PharmaJet Stratis jet injector for persons aged 18 through 64 years (48). Persons in this age group may receive Afluria Quadrivalent via either needle and syringe or this specific jet injection device.

Children aged 6 months through 17 years and adults aged  $\geq$ 65 years should receive this vaccine by needle and syringe only. No other IIV4s are licensed for administration by jet injector.

Contraindications and Precautions for the Use of IIV4s. Manufacturer package inserts and updated CDC and ACIP guidance should be consulted for information on contraindications and precautions for individual influenza vaccines. Each IIV, whether egg based or cell culture based, has a labeled contraindication for persons with a history of a severe allergic reaction to any component of that vaccine (Tables 2 and 3). However, although egg is a component of all IIV4s other than ccIIV4, ACIP makes specific recommendations for the use of influenza vaccine for persons with egg allergy (see Persons with a History of Egg Allergy). All egg-based IIV4s are contraindicated in persons who have had a severe allergic reaction (e.g., anaphylaxis) to a previous dose of any influenza vaccine (any egg-based IIV, ccIIV, RIV, or LAIV of any valency). Use of ccIIV4 is contraindicated in persons who have had a severe allergic reaction (e.g., anaphylaxis) to any ccIIV of any valency. A history of severe allergic reaction (e.g., anaphylaxis) to any other influenza vaccine (i.e., any eggbased IIV, RIV, or LAIV of any valency) is a precaution for the use of ccIIV4 (see Persons with Previous Allergic Reactions to Influenza Vaccines) (Tables 2 and 3). If ccIIV4 is administered in such an instance, vaccination should occur in an inpatient or outpatient medical setting and should be supervised by a health care provider who is able to recognize and manage severe allergic reactions. Providers can also consider consultation with an allergist to help identify the vaccine component responsible for the reaction. Information about vaccine components can be found in the package inserts for each vaccine. Prophylactic use of antiviral agents is an option that can be considered for preventing influenza among persons who cannot receive vaccine, particularly for those who are at higher risk for medical complications attributable to severe influenza (102).

Moderate or severe acute illness with or without fever is a general precaution for vaccination (47). A history of GBS within 6 weeks after receipt of a previous dose of influenza vaccine is considered a precaution for the use of all influenza vaccines (Table 2).

#### Quadrivalent Recombinant Influenza Vaccine (RIV4)

**Available Vaccine.** One recombinant influenza vaccine, Flublok Quadrivalent (RIV4), is expected to be available during the 2023–24 influenza season. RIV4 is approved for persons aged  $\geq$ 18 years. This vaccine contains recombinant HA produced in an insect cell line using genetic sequences from cell-derived influenza viruses and is manufactured without the use of influenza viruses or eggs (*54*). **Dosage and Administration**. RIV4 is administered by IM injection via needle and syringe. A 0.5-mL dose contains 45  $\mu$ g of HA derived from each vaccine virus (180  $\mu$ g total).

Contraindications and Precautions for the Use of RIV4. Manufacturer package inserts and updated CDC and ACIP guidance should be consulted for information on contraindications and precautions for individual influenza vaccines. RIV4 is contraindicated in persons who have had a severe allergic reaction (e.g., anaphylaxis) to a previous dose of any RIV of any valency or to any component of RIV4. A history of a severe allergic reaction (e.g., anaphylaxis) to any other influenza vaccine (i.e., any egg-based IIV, ccIIV, or LAIV of any valency) is a precaution for the use of RIV4. If RIV4 is administered in such an instance, vaccination should occur in an inpatient or outpatient medical setting and should be supervised by a health care provider who is able to recognize and manage severe allergic reactions. Providers can also consider consulting with an allergist to help identify the vaccine component responsible for the reaction (Tables 2 and 3).

Moderate or severe acute illness with or without fever is a general precaution for vaccination (47). A history of GBS within 6 weeks after receipt of a previous dose of influenza vaccine is considered a precaution for the use of all influenza vaccines (Table 2). RIV4 is not licensed for children aged <18 years.

## Quadrivalent Live Attenuated Influenza Vaccine (LAIV4)

**Available Vaccine.** One live attenuated influenza vaccine, FluMist Quadrivalent (LAIV4), is expected to be available during the 2023–24 influenza season. LAIV4 is approved for persons aged 2 through 49 years. LAIV4 contains live attenuated influenza viruses that are propagated in eggs. These viruses are cold adapted (so that they replicate efficiently at 25°C [77°F]) and temperature sensitive (so that their replication is restricted at higher temperatures, 39°C [102.2°F] for influenza A viruses and 37°C [98.6°] for influenza B viruses). The live attenuated vaccine viruses replicate in the nasopharynx, which is necessary to promote an immune response (*53*). No preference is expressed for LAIV4 versus other influenza vaccines used within specified indications.

**Dosage and Administration.** LAIV4 is administered intranasally using the supplied prefilled, single-use sprayer containing 0.2 mL of vaccine. Approximately 0.1 mL (i.e., one half of the total sprayer contents) is sprayed into the first nostril while the recipient is in the upright position. An attached dose-divider clip is removed from the sprayer to permit administration of the second half of the dose into the other nostril. Sniffing of the dose is not necessary. If the recipient sneezes immediately after administration, the dose should

not be repeated. However, if nasal congestion is present that might impede delivery of the vaccine to the nasopharyngeal mucosa, deferral of administration should be considered until resolution of the illness, or another appropriate vaccine should be administered instead. Each total dose of 0.2 mL contains  $10^{6.5-7.5}$  fluorescent focus units of each vaccine virus (*53*).

**Contraindications and Precautions for the Use of LAIV4.** Manufacturer package inserts and updated CDC and ACIP guidance should be consulted for information on contraindications and precautions for individual influenza vaccines. Conditions considered by ACIP to be contraindications and precautions for the use of LAIV4 are summarized (Table 2). These include two labeled contraindications that appear in the package insert (53) and other conditions for which there is either uncertain but biologically plausible potential risk associated with live viruses or limited data for use of LAIV. Contraindications to use of LAIV4 include the following (Tables 2 and 3):

- Severe allergic reaction (e.g., anaphylaxis) to any component of the vaccine or to a previous dose of any influenza vaccine (i.e., any egg-based IIV, ccIIV, RIV, or LAIV of any valency; a labeled contraindication noted in the package insert). However, although egg is a component of LAIV4, ACIP makes specific recommendations for the use of influenza vaccine for persons with egg allergy (see Persons with a History of Egg Allergy).
- Children and adolescents receiving concomitant aspirin- or salicylate-containing medications, because of the potential risk for Reye syndrome (a labeled contraindication noted in the package insert).
- Children aged 2 through 4 years who have received a diagnosis of asthma or whose parents or caregivers report that a health care provider has told them during the preceding 12 months that their child had wheezing or asthma or whose medical record indicates a wheezing episode has occurred during the preceding 12 months.
- Children and adults who are immunocompromised due to any cause, including but not limited to immunosuppression caused by medications, congenital or acquired immunodeficiency states, HIV infection, anatomic asplenia, or functional asplenia (such as that due to sickle cell anemia).
- Close contacts and caregivers of severely immunosuppressed persons who require a protected environment.
- Pregnancy.
- Persons with active communication between the cerebrospinal fluid (CSF) and the oropharynx, nasopharynx, nose, or ear or any other cranial CSF leak.
- Persons with cochlear implants, because of the potential for CSF leak that might exist for a period after implantation

(providers might consider consultation with a specialist concerning the risk for persistent CSF leak if an age-appropriate inactivated or recombinant vaccine cannot be used).

• Receipt of influenza antiviral medication within the previous 48 hours for oseltamivir and zanamivir, previous 5 days for peramivir, and previous 17 days for baloxavir. The interval between influenza antiviral receipt and LAIV4 during which interference might potentially occur might be further prolonged in the presence of medical conditions that delay medication clearance (e.g., renal insufficiency).

Precautions to the use of LAIV4 include the following (Tables 2 and 3):

- Moderate or severe acute illness with or without fever.
- History of GBS within 6 weeks after receipt of any influenza vaccine.
- Asthma in persons aged  $\geq 5$  years.
- Other underlying medical condition (other than those listed under contraindications) that might predispose to complications after wild-type influenza virus infection (e.g., chronic pulmonary, cardiovascular [except isolated hypertension], renal, hepatic, neurologic, hematologic, or metabolic disorders [including diabetes mellitus]).

# Storage and Handling of Influenza Vaccines

In all instances, approved manufacturer packaging information should be consulted for authoritative guidance concerning storage and handling of specific influenza vaccines. Typically, influenza vaccines should be protected from light and stored at temperatures that are recommended in the package insert. Recommended storage temperatures are typically 36°F–46°F (2°C–8°C) and should be maintained at all times with adequate refrigeration and temperature monitoring. Vaccine that has frozen should be discarded. Specific recommendations for appropriate refrigerators and temperature monitoring equipment can be found in the Vaccine Storage and Handling Toolkit, available at https://www.cdc.gov/vaccines/ hcp/admin/storage/toolkit/index.html.

Vaccines should not be used beyond the expiration date on the label. In addition to the expiration date, multidose vials also might have a beyond-use date (BUD), which specifies the number of days the vaccine can be kept once first accessed. After being accessed for the first dose, multidose vials should not be used after the BUD. If no BUD is provided, then the listed expiration date is to be used. Multidose vials should be returned to recommended storage conditions between uses. Package information might also specify a maximum number of doses contained in multidose vials (regardless of remaining volume). No more than the specified number of doses should be removed, and any remainder should be discarded. Single-dose vials should not be accessed for more than 1 dose. Providers should contact the manufacturer for information on permissible temperature excursions and other departures from recommended storage and handling conditions that are not discussed in the package labeling.

# Additional Sources of Information Regarding Influenza and Influenza Vaccines

## Influenza Surveillance, Prevention, and Control

Updated information regarding influenza surveillance, detection, prevention, and control is available at https://www.cdc. gov/flu. U.S. surveillance data are updated weekly throughout the year on FluView (https://www.cdc.gov/flu/weekly) and can be viewed in FluView Interactive (https://www.cdc.gov/flu/weekly/fluviewinteractive.htm). In addition, periodic updates regarding influenza are published in MMWR (https://www. cdc.gov/mmwr/index.html). Additional information regarding influenza and influenza vaccines can be obtained from CDCINFO by calling 1-800-232-4636. State and local health departments should be consulted about availability of influenza vaccines, access to vaccination programs, information related to state or local influenza activity, reporting of influenza outbreaks and influenza-related pediatric deaths, and advice concerning outbreak control.

## Vaccine Adverse Event Reporting System (VAERS)

The National Childhood Vaccine Injury Act of 1986 requires health care providers to report any adverse event listed by the vaccine manufacturer as a contraindication to future doses of the vaccine or any adverse event listed in the VAERS Table of Reportable Events Following Vaccination (https://vaers.hhs. gov/docs/VAERS\_Table\_of\_Reportable\_Events\_Following\_ Vaccination.pdf) that occurs within the specified period after vaccination. In addition to mandated reporting, health care providers are encouraged to report any clinically significant adverse event after vaccination to VAERS. Information on how to report a vaccine adverse event is available at https://vaers. hhs.gov/index.html.

# National Vaccine Injury Compensation Program (VICP)

The National Vaccine Injury Compensation Program (VICP), established by the National Childhood Vaccine Injury Act of 1986, as amended, is a no-fault alternative to the traditional tort system. It provides compensation to people found to be injured by certain vaccines. VICP covers most vaccines routinely given in the United States. The Vaccine Injury Table (https://www.hrsa.gov/sites/default/files/hrsa/ vicp/vaccine-injury-table-01-03-2022.pdf) lists the vaccines covered by VICP and the associated injuries and conditions that might receive a legal presumption of causation. If the injury or condition is not in the table or does not meet the requirements in the table, persons must prove that the vaccine caused the injury or condition. Claims must be filed within specified time frames. Persons of all ages who receive a VICPcovered vaccine might be eligible to file a claim. Additional information is available at https://www.hrsa.gov/vaccinecompensation or by calling 1-800-338-2382.

## **Additional Resources**

## **ACIP Statements**

- Recommended Adult Immunization Schedule for Ages 19 Years or Older, United States: https://www.cdc.gov/ vaccines/schedules/hcp/imz/adult.html
- Recommended Child and Adolescent Immunization Schedule for Ages 18 Years or Younger, United States: https://www.cdc.gov/vaccines/schedules/hcp/imz/childadolescent.html
- Immunization of Health Care Personnel: Recommendations of the Advisory Committee on Immunization Practices (ACIP), 2011. MMWR Recomm Rep 2011;60(No.RR-7):1–45: https://www.cdc.gov/ mmwr/preview/mmwrhtml/rr6007a1.htm

## General Best Practices Guidelines for Immunization:

• General Best Practice Guidelines for Immunization: https://www.cdc.gov/vaccines/hcp/acip-recs/general-recs/ index.html

#### **COVID-19 Vaccine Recommendations and Guidance**

• ACIP recommendations for the use of COVID-19 vaccines: https://www.cdc.gov/vaccines/hcp/acip-recs/vacc-specific/covid-19.html

- Clinical Care Considerations for COVID-19 Vaccination: https://www.cdc.gov/vaccines/covid-19/clinicalconsiderations/index.html
- Use of COVID-19 Vaccines in the United States Interim Clinical Considerations: https://www.cdc.gov/ vaccines/covid-19/clinical-considerations/covid-19vaccines-us.html
- FDA COVID-19 Vaccines page: https://www.fda.gov/ emergency-preparedness-and-response/coronavirusdisease-2019-covid-19/covid-19-vaccines

#### **Vaccine Information Sheets**

- IIV4 and RIV4: https://www.cdc.gov/vaccines/hcp/vis/ vis-statements/flu.pdf
- LAIV4: https://www.cdc.gov/vaccines/hcp/vis/visstatements/flulive.pdf

## Influenza Vaccine Package Inserts

 https://www.fda.gov/vaccines-blood-biologics/vaccines/ vaccines-licensed-use-united-states

## **CDC Influenza Antiviral Guidance**

• Influenza Antiviral Medications: Summary for Clinicians: https://www.cdc.gov/flu/professionals/antivirals/ summary-clinicians.htm

## Infectious Diseases Society of America Influenza Antiviral Guidance

• Clinical Practice Guidelines by the Infectious Diseases Society of America: 2018 Update on Diagnosis, Treatment, Chemoprophylaxis, and Institutional Outbreak Management of Seasonal Influenza: https://academic.oup. com/cid/article/68/6/e1/5251935

#### **American Academy of Pediatrics Guidance**

• American Academy of Pediatrics Recommendations for Prevention and Control of Influenza in Children (Red Book Online): https://publications.aap.org/redbook

## Infectious Diseases Society of America Guidance for Vaccination of Immunocompromised Hosts

• 2013 IDSA Clinical Practice Guideline for Vaccination of the Immunocompromised Host: https://academic.oup. com/cid/article/58/3/e44/336537

## American College of Obstetricians and Gynecologists

 Influenza Vaccination During Pregnancy, ACOG Committee Opinion No. 732: https://www.acog. org/clinical/clinical-guidance/committee-opinion/ articles/2018/04/influenza-vaccination-during-pregnancy

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Voting members of the Advisory Committee on Immunization Practices: Grace M. Lee, MD, Stanford University School of Medicine, Stanford, California (Chair); Lynn Bahta, MPH, Minnesota Department of Health, St. Paul, Minnesota; Beth P. Bell, MD, University of Washington, Seattle, Washington; Oliver Brooks, MD, Watts HealthCare Corporation, Los Angeles, California; Wilbur H. Chen, MD, University of Maryland School of Medicine, Baltimore, Maryland; Sybil Cineas, MD, Warren Alpert Medical School of Brown University, Providence, Rhode Island; Matthew F. Daley, MD, Kaiser Permanente Colorado, Aurora, Colorado; Camille Nelson Kotton, MD, Harvard Medical School, Boston, Massachusetts; Jamie Loehr, MD, Cayuga Family Medicine, Ithaca, New York; Sarah S. Long, MD, Drexel University College of Medicine, Philadelphia, Pennsylvania; Veronica V. McNally, JD, Franny Strong Foundation, West Bloomfield, Michigan; Katherine A. Poehling, MD, Wake Forest School of Medicine, Winston-Salem, North Carolina; Pablo J. Sánchez, MD, The Research Institute at Nationwide Children's Hospital, Columbus, Ohio; Helen Keipp Talbot, MD, Vanderbilt University, Nashville, Tennessee.

Donna S. Hummell, MD, Vanderbilt University, Nashville, Tennessee. Alicia Budd, MPH; Sascha Ellington, PhD; Brendan Flannery, PhD; Andrew Kroger, MD; Samantha Olson, MPH; Tom Shimabukuro, MD; Mark Tenforde, MD; Lindsay Trujillo, MPH, CDC.

#### **ACIP Influenza Vaccine Work Group**

H. Keipp Talbot, MD, Nashville, Tennessee (Chair); Robert Atmar, MD, Houston, Texas; Kevin Ault, MD, Kalamazoo, Michigan; Edward Belongia, MD, Marshfield, Wisconsin; Henry Bernstein, DO, Hempstead, New York; Thomas Boyce, MD, Marshfield, Wisconsin; Timothy Brennan, MD, Silver Spring, Maryland; Kristina Angel Bryant, MD, Louisville, Kentucky; Doug Campos-Outcalt, MD, Phoenix, Arizona; Uzo Chukwuma, PhD, Rockville, Maryland; Sarah Coles, MD, Phoenix, Arizona; Alicia Fry, MD, Atlanta, Georgia; Sandra Adamson Fryhofer, MD, Atlanta, Georgia; Denise Jamieson, MD, Atlanta, Georgia; Jeffrey Kelman, MD, Baltimore, Maryland; Krissy Moehling Geffel, PhD, Pittsburgh, Pennsylvania; Wendy Keitel, MD, Houston, Texas; Camille Kotton, MD, Boston, Massachusetts; Marie-Michèle Léger, MPH, Alexandria, Virginia; Jamie Loehr, MD, Ithaca, New York; Valerie Marshall, MPH, Washington, DC; Zackary Moore, MD, Raleigh, North Carolina; Rebecca L. Morgan, PhD, Cleveland, Ohio; Kathleen M. Neuzil, MD, Baltimore, Maryland; Cynthia Nolletti, MD, Silver Spring, Maryland; Chris Roberts, PhD; Rockville, Maryland; William Schaffner, MD, Nashville, Tennessee; Robert Schechter, MD, Richmond, California; Kenneth Schmader, MD, Durham, North Carolina; Tamara Sheffield, MD, Salt Lake City, Utah; Patricia A. Stinchfield, MS, St. Paul, Minnesota; Peter Szilagyi, MD, Los Angeles, California; Matthew Zahn, MD, Santa Ana, California.

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#### References

- Barker WH. Excess pneumonia and influenza associated hospitalization during influenza epidemics in the United States, 1970-78. Am J Public Health 1986;76:761–5. PMID:3717461 https://doi.org/10.2105/ AJPH.76.7.761
- Barker WH, Mullooly JP. Impact of epidemic type A influenza in a defined adult population. Am J Epidemiol 1980;112:798–813. PMID:7457471 https://doi.org/10.1093/oxfordjournals.aje.a113052
- Coleman BL, Fadel SA, Fitzpatrick T, Thomas SM. Risk factors for serious outcomes associated with influenza illness in high-versus lowand middle-income countries: systematic literature review and metaanalysis. Influenza Other Respir Viruses 2018;12:22–9. PMID:29197154 https://doi.org/10.1111/irv.12504
- Mullooly JP, Bridges CB, Thompson WW, et al.; Vaccine Safety Datalink Adult Working Group. Influenza- and RSV-associated hospitalizations among adults. Vaccine 2007;25:846–55. PMID:17074423 https://doi.org/10.1016/j.vaccine.2006.09.041
- Poehling KA, Edwards KM, Griffin MR, et al. The burden of influenza in young children, 2004-2009. Pediatrics 2013;131:207–16. PMID:23296444 https://doi.org/10.1542/peds.2012-1255
- Poehling KA, Edwards KM, Weinberg GA, et al.; New Vaccine Surveillance Network. The underrecognized burden of influenza in young children. N Engl J Med 2006;355:31–40. PMID:16822994 https://doi.org/10.1056/NEJMoa054869
- Siston AM, Rasmussen SA, Honein MA, et al.; Pandemic H1N1 Influenza in Pregnancy Working Group. Pandemic 2009 influenza A(H1N1) virus illness among pregnant women in the United States. JAMA 2010;303:1517–25. PMID:20407061 https://doi.org/10.1001/ jama.2010.479
- Fragaszy EB, Warren-Gash C, White PJ, et al.; Flu Watch Group. Effects of seasonal and pandemic influenza on health-related quality of life, work and school absence in England: results from the Flu Watch cohort study. Influenza Other Respir Viruses 2018;12:171–82. PMID:28991409 https://doi.org/10.1111/irv.12506
- Van Wormer JJ, King JP, Gajewski A, McLean HQ, Belongia EA. Influenza and workplace productivity loss in working adults. J Occup Environ Med 2017;59:1135–9. PMID:28759481 https://doi. org/10.1097/JOM.00000000001120
- Willis GA, Preen DB, Richmond PC, et al.; WAIVE Study Team. The impact of influenza infection on young children, their family and the health care system. Influenza Other Respir Viruses 2019;13:18–27. PMID:30137663 https://doi.org/10.1111/irv.12604
- Fiore AE, Uyeki TM, Broder K, et al.; Centers for Disease Control and Prevention (CDC). Prevention and control of influenza with vaccines: recommendations of the Advisory Committee on Immunization Practices (ACIP), 2010. MMWR Recomm Rep 2010;59(No. RR-8):1–62. PMID:20689501

- 12. CDC. How flu vaccine effectiveness and efficacy are measured: questions and answers. Atlanta, GA: US Department of Health and Human Services, CDC; 2023.
- Rolfes MA, Foppa IM, Garg S, et al. Annual estimates of the burden of seasonal influenza in the United States: a tool for strengthening influenza surveillance and preparedness. Influenza Other Respir Viruses 2018;12:132–7. PMID:29446233 https://doi.org/10.1111/irv.12486
- 14. Rolfes MA, Flannery B, Chung JR, et al.; US Influenza Vaccine Effectiveness (Flu VE) Network, the Influenza Hospitalization Surveillance Network, and the Assessment Branch, Immunization Services Division, CDC. Effects of Influenza Vaccination in the United States During the 2017-2018 Influenza Season. Clin Infect Dis 2019;69:1845–53. PMID:30715278 https://doi.org/10.1093/cid/ciz075
- 15. CDC. 2020–21 flu season summary FAQ. Atlanta, GA: US Department of Health and Human Services, CDC; 2021. https://www. cdc.gov/flu/season/faq-flu-season-2020-2021.htm
- CDC. 2021–22 flu season summary. Atlanta, GA: US Department of Health and Human Services, CDC; 2023. https://www.cdc.gov/flu/ season/faq-flu-season-2021-2022.htm
- 17. CDC. Fluview. Weekly influenza surveillance report. Atlanta, GA: US Department of Health and Human Services, CDC; 2023. https://www. cdc.gov/flu/weekly/index.htm
- Grohskopf LA, Blanton LH, Ferdinands JM, et al. Prevention and control of seasonal influenza with vaccines: recommendations of the Advisory Committee on Immunization Practices—United States, 2022–23 influenza season. MMWR Recomm Rep 2022;71(No. RR-1):1–28. PMID:36006864 https://doi.org/10.15585/mmwr.rr7101a1
- Ahmed F. US Advisory Committee on Immunization Practices (ACIP) handbook for developing evidence-based recommendations, version 1.2. Atlanta, GA: US Department of Health and Human Services, CDC; 2013.
- 20. CDC. Advisory Committee on Immunization Practices (ACIP): evidence-based recommendations—GRADE. Atlanta, GA: US Department of Health and Human Services, CDC; 2023. https://www. cdc.gov/vaccines/acip/recs/grade/about-grade.html
- Belongia EA, Sundaram ME, McClure DL, Meece JK, Ferdinands J, VanWormer JJ. Waning vaccine protection against influenza A (H3N2) illness in children and older adults during a single season. Vaccine 2015;33:246–51. PMID:24962752 https://doi.org/10.1016/j. vaccine.2014.06.052
- 22. Castilla J, Martínez-Baz I, Martínez-Artola V, et al.; Primary Health Care Sentinel Network; Network for Influenza Surveillance in Hospitals of Navarre. Decline in influenza vaccine effectiveness with time after vaccination, Navarre, Spain, season 2011/12. Euro Surveill 2013;18:20388. PMID:23399423 https://doi.org/10.2807/ ese.18.05.20388-en
- Ferdinands JM, Fry AM, Reynolds S, et al. Intraseason waning of influenza vaccine protection: evidence from the US Influenza Vaccine Effectiveness Network, 2011-12 through 2014-15. Clin Infect Dis 2017;64:544–50. PMID:28039340
- Ferdinands JM, Gaglani M, Martin ET, et al. Waning vaccine effectiveness against influenza-associated hospitalizations among adults, 2015–2016 to 2018–2019, United States Hospitalized Adult Influenza Vaccine Effectiveness Network. Clin Infect Dis 2021;73:726–9. PMID:33462610 https://doi.org/10.1093/cid/ciab045
- 25. Kissling E, Nunes B, Robertson C, et al.; I-MOVE case–control study team. I-MOVE multicentre case-control study 2010/11 to 2014/15: Is there within-season waning of influenza type/subtype vaccine effectiveness with increasing time since vaccination? Euro Surveill 2016;21:1–12. PMID:27124420 https://doi.org/10.2807/1560-7917. ES.2016.21.16.30201

- 26. Kissling E, Valenciano M, Larrauri A, et al. Low and decreasing vaccine effectiveness against influenza A(H3) in 2011/12 among vaccination target groups in Europe: results from the I-MOVE multicentre casecontrol study. Euro Surveill 2013;18:20390. PMID:23399425 https://doi.org/10.2807/ese.18.05.20390-en
- Mira-Iglesias A, López-Labrador FX, García-Rubio J, et al. Influenza vaccine effectiveness and waning effect in hospitalized older adults. Valencia region, Spain, 2018/2019 season. Int J Environ Res Public Health 2021;18:1129. PMID:33514058 https://doi.org/10.3390/ ijerph18031129
- Ng Y, Nandar K, Chua LAV, et al. Evaluating the effectiveness of the influenza vaccine during respiratory outbreaks in Singapore's long term care facilities, 2017. Vaccine 2019;37:3925–31. PMID:31160102 https://doi.org/10.1016/j.vaccine.2019.03.054
- Pebody R, Andrews N, McMenamin J, et al. Vaccine effectiveness of 2011/12 trivalent seasonal influenza vaccine in preventing laboratoryconfirmed influenza in primary care in the United Kingdom: evidence of waning intra-seasonal protection. Euro Surveill 2013;18:20389. PMID:23399424 https://doi.org/10.2807/ese.18.05.20389-en
- Powell LN, Bégué RE. Influenza vaccine effectiveness among children for the 2017–2018 season. J Pediatric Infect Dis Soc 2020;9:468–73. PMID:31774120 https://doi.org/10.1093/jpids/piz077
- Puig-Barberà J, Mira-Iglesias A, Tortajada-Girbés M, et al.; Valencia Hospital Network for the Study of Influenza and other Respiratory Viruses (VAHNSI, Spain). Waning protection of influenza vaccination during four influenza seasons, 2011/2012 to 2014/2015. Vaccine 2017;35:5799–807. PMID:28941618 https://doi.org/10.1016/j. vaccine.2017.09.035
- 32. Radin JM, Hawksworth AW, Myers CA, Ricketts MN, Hansen EA, Brice GT. Influenza vaccine effectiveness: maintained protection throughout the duration of influenza seasons 2010-2011 through 2013-2014. Vaccine 2016;34:3907–12. PMID:27265447 https://doi. org/10.1016/j.vaccine.2016.05.034
- 33. Ray GT, Lewis N, Klein NP, et al. Intraseason waning of influenza vaccine effectiveness. Clin Infect Dis 2019;68:1623–30. PMID:30204855 https://doi.org/10.1093/cid/ciy770
- 34. Young BE, Mak TM, Ang LW, et al. Influenza vaccine failure in the tropics: a retrospective cohort study of waning effectiveness. Epidemiol Infect 2020;148:e299. PMID:33261680 https://doi.org/10.1017/ S0950268820002952
- 35. Hu W, Sjoberg PA, Fries AC, DeMarcus LS, Robbins AS. Waning vaccine protection against influenza among department of defense adult beneficiaries in the United States, 2016–2017 through 2019–2020 influenza seasons. Vaccines (Basel) 2022;10:888. PMID:35746496 https://doi.org/10.3390/vaccines10060888
- 36. Regan AK, Fielding JE, Chilver MB, et al. Intraseason decline in influenza vaccine effectiveness during the 2016 southern hemisphere influenza season: a test-negative design study and phylogenetic assessment. Vaccine 2019;37:2634–41. PMID:30952499 https://doi. org/10.1016/j.vaccine.2019.02.027
- 37. Sahni LC, Naioti EA, Olson SM, et al. Sustained within-season vaccine effectiveness against influenza-associated hospitalization in children: evidence from the New Vaccine Surveillance Network, 2015–2016 through 2019–2020. Clin Infect Dis 2023;76:e1031–9. PMID:35867698 https://doi.org/10.1093/cid/ciac577
- Madhi SA, Cutland CL, Kuwanda L, et al.; Maternal Flu Trial (Matflu) Team. Influenza vaccination of pregnant women and protection of their infants. N Engl J Med 2014;371:918–31. PMID:25184864 https://doi.org/10.1056/NEJMoa1401480
- 39. Steinhoff MC, Katz J, Englund JA, et al. Year-round influenza immunisation during pregnancy in Nepal: a phase 4, randomised, placebo-controlled trial. Lancet Infect Dis 2017;17:981–9. PMID:28522338 https://doi.org/10.1016/S1473-3099(17)30252-9

- 40. Tapia MD, Sow SO, Tamboura B, et al. Maternal immunisation with trivalent inactivated influenza vaccine for prevention of influenza in infants in Mali: a prospective, active-controlled, observer-blind, randomised phase 4 trial. Lancet Infect Dis 2016;16:1026–35. PMID:27261067 https://doi.org/10.1016/S1473-3099(16)30054-8
- 41. Zaman K, Roy E, Arifeen SE, et al. Effectiveness of maternal influenza immunization in mothers and infants. N Engl J Med 2008;359:1555–64. PMID:18799552 https://doi.org/10.1056/NEJMoa0708630
- 42. CDC. Flu season. Atlanta, GA: US Department of Health and Human Services, CDC; 2023. https://www.cdc.gov/flu/about/season
- 43. Young B, Sadarangani S, Jiang L, Wilder-Smith A, Chen MI. Duration of influenza vaccine effectiveness: a systematic review, meta-analysis, and metaregression of test-negative design case-control studies. J Infect Dis 2018;217:731–41. PMID:29220496 https://doi.org/10.1093/infdis/jix632
- 44. Ferdinands JM, Alyanak E, Reed C, Fry AM. Waning of influenza vaccine protection: exploring the trade-offs of changes in vaccination timing among older adults. Clin Infect Dis 2020;70:1550–9. PMID:31257422 https://doi.org/10.1093/cid/ciz452
- 45. Advisory Committee on Immunization Practices; CDC. Immunization of health-care personnel: recommendations of the Advisory Committee on Immunization Practices (ACIP). MMWR Recomm Rep 2011;60(No. RR-7):1–45. PMID:22108587
- 46. Pearson ML, Bridges CB, Harper SA; Healthcare Infection Control Practices Advisory Committee (HICPAC); Advisory Committee on Immunization Practices (ACIP). Influenza vaccination of health-care personnel: recommendations of the Healthcare Infection Control Practices Advisory Committee (HICPAC) and the Advisory Committee on Immunization Practices (ACIP). MMWR Recomm Rep 2006;55(No. RR-2):1–16. PMID:16498385
- 47. Kroger A, Bahta L, Long S, Sanchez P. General best practice guidelines for immunization. Atlanta, GA: US Department of Health and Human Services, CDC; 2023. https://www.cdc.gov/vaccines/hcp/acip-recs/ general-recs/index.html
- 48. Afluria Quadrivalent [Package Insert]. Parkville, Victoria, Australia: Seqirus; 2023.
- 49. Fluarix Quadrivalent [Package Insert]. Dresden, Germany: GlaxoSmithKline; 2023.
- 50. Flucelvax Quadrivalent [Package Insert]. Holly Springs, NC: Seqirus; 2023.
- 51. FluLaval Quadrivalent [Package Insert]. Quebec City, QC, Canada: ID Biomedical Corporation of Quebec; 2023.
- 52. Fluzone Quadrivalent [Package Insert]. Swiftwater, PA: Sanofi Pasteur; 2023.
- 53. FluMist Quadrivalent [Package Insert]. Gaithersburg, MD: MedImmune; 2023.
- 54. Flublok Quadrivalent [Package Insert]. Meriden, CT: Protein Sciences; 2023.
- 55. Fluzone High-Dose Quadrivalent [Package Insert]. Swiftwater, PA: Sanofi Pasteur; 2023.
- 56. Fluad Quadrivalent [Package Insert]. Holly Springs, NC: Seqirus; 2023.
- Allison MA, Daley MF, Crane LA, et al. Influenza vaccine effectiveness in healthy 6- to 21-month-old children during the 2003–2004 season. J Pediatr 2006;149:755–62e1. PMID:17137887 https://doi. org/10.1016/j.jpeds.2006.06.036
- 58. Eisenberg KW, Szilagyi PG, Fairbrother G, et al.; New Vaccine Surveillance Network. Vaccine effectiveness against laboratoryconfirmed influenza in children 6 to 59 months of age during the 2003-2004 and 2004-2005 influenza seasons. Pediatrics 2008;122:911–9. PMID:18977968 https://doi.org/10.1542/ peds.2007-3304
- 59. Neuzil KM, Jackson LA, Nelson J, et al. Immunogenicity and reactogenicity of 1 versus 2 doses of trivalent inactivated influenza vaccine in vaccine-naive 5-8-year-old children. J Infect Dis 2006;194:1032–9. PMID:16991077 https://doi.org/10.1086/507309
- 60. Ritzwoller DP, Bridges CB, Shetterly S, Yamasaki K, Kolczak M, France EK. Effectiveness of the 2003-2004 influenza vaccine among children 6 months to 8 years of age, with 1 vs 2 doses. Pediatrics 2005;116:153–9. PMID:15995046 https://doi.org/10.1542/peds.2005-0049

- 61. Eick AA, Uyeki TM, Klimov A, et al. Maternal influenza vaccination and effect on influenza virus infection in young infants. Arch Pediatr Adolesc Med 2011;165:104–11. PMID:20921345 https://doi. org/10.1001/archpediatrics.2010.192
- 62. ACOG Committee on Obstetric Practice. ACOG Committee opinion no. 732: influenza vaccination during pregnancy. Obstet Gynecol 2018;131:e109–14. PMID:29578985 https://doi.org/10.1097/ AOG.000000000002588
- 63. Bratton KN, Wardle MT, Orenstein WA, Omer SB. Maternal influenza immunization and birth outcomes of stillbirth and spontaneous abortion: a systematic review and meta-analysis. Clin Infect Dis 2015;60:e11–9. PMID:25409473 https://doi.org/10.1093/cid/ciu915
- 64. Chambers CD, Johnson D, Xu R, et al.; OTIS Collaborative Research Group. Risks and safety of pandemic H1N1 influenza vaccine in pregnancy: birth defects, spontaneous abortion, preterm delivery, and small for gestational age infants. Vaccine 2013;31:5026–32. PMID:24016809 https://doi.org/10.1016/j.vaccine.2013.08.097
- 65. Chambers CD, Johnson DL, Xu R, et al.; OTIS Collaborative Research Group. Safety of the 2010-11, 2011-12, 2012-13, and 2013-14 seasonal influenza vaccines in pregnancy: birth defects, spontaneous abortion, preterm delivery, and small for gestational age infants, a study from the cohort arm of VAMPSS. Vaccine 2016;34:4443–9. PMID:27449682 https://doi.org/10.1016/j.vaccine.2016.06.054
- 66. Heikkinen T, Young J, van Beek E, et al. Safety of MF59-adjuvanted A/H1N1 influenza vaccine in pregnancy: a comparative cohort study. Am J Obstet Gynecol 2012;207:177.e1–8. PMID:22939717 https://doi.org/10.1016/j.ajog.2012.07.007
- 67. Huang WT, Tang FW, Yang SE, Chih YC, Chuang JH. Safety of inactivated monovalent pandemic (H1N1) 2009 vaccination during pregnancy: a population-based study in Taiwan. Vaccine 2014;32:6463–8. PMID:25285884 https://doi.org/10.1016/j. vaccine.2014.09.054
- Irving SA, Kieke BA, Donahue JG, et al.; Vaccine Safety Datalink. Trivalent inactivated influenza vaccine and spontaneous abortion. Obstet Gynecol 2013;121:159–65. PMID:23262941 https://doi. org/10.1097/AOG.0b013e318279f56f
- 69. Ma F, Zhang L, Jiang R, et al. Prospective cohort study of the safety of an influenza A(H1N1) vaccine in pregnant Chinese women. Clin Vaccine Immunol 2014;21:1282–7. PMID:24990911 https://doi. org/10.1128/CVI.00375-14
- 70. 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. PMID:25758932 https://doi.org/10.1016/j. vaccine.2015.02.068
- 71. Pasternak B, Svanström H, Mølgaard-Nielsen D, et al. Vaccination against pandemic A/H1N1 2009 influenza in pregnancy and risk of fetal death: cohort study in Denmark. BMJ 2012;344:e2794. PMID:22551713 https://doi.org/10.1136/bmj.e2794
- 72. Sammon CJ, Snowball J, McGrogan A, de Vries CS. Evaluating the hazard of foetal death following H1N1 influenza vaccination; a population based cohort study in the UK GPRD. PLoS One 2012;7:e51734. PMID:23341865 https://doi.org/10.1371/journal. pone.0051734
- 73. Oppermann M, Fritzsche J, Weber-Schoendorfer C, et al. A(H1N1) v2009: a controlled observational prospective cohort study on vaccine safety in pregnancy. Vaccine 2012;30:4445–52. PMID:22564554 https://doi.org/10.1016/j.vaccine.2012.04.081
- 74. Donahue JG, Kieke BA, King JP, et al. Association of spontaneous abortion with receipt of inactivated influenza vaccine containing H1N1pdm09 in 2010-11 and 2011-12. Vaccine 2017;35:5314–22. PMID:28917295 https://doi.org/10.1016/j.vaccine.2017.06.069

- 75. Donahue JG, Kieke BA, King JP, et al. Inactivated influenza vaccine and spontaneous abortion in the Vaccine Safety Datalink in 2012-13, 2013-14, and 2014-15. Vaccine 2019;37:6673–81. PMID:31540812 https://doi.org/10.1016/j.vaccine.2019.09.035
- 76. Moro PL, Marquez P. Reports of cell-based influenza vaccine administered during pregnancy in the Vaccine Adverse Event Reporting System (VAERS), 2013-2020. Vaccine 2021;39:678–81. PMID:33358703 https://doi.org/10.1016/j.vaccine.2020.12.045
- 77. Robinson C, Van Boxmeer J, Tilson H, et al. Outcomes in pregnant persons immunized with a cell-based quadrivalent inactivated influenza vaccine: a prospective observational cohort study. Vaccines (Basel) 2022;10:1600. PMID:36298465 https://doi.org/10.3390/ vaccines10101600
- 78. Swamy GK. Clinical Trial to compare safety of Recombinant Influenza Vaccine (RIV4) versus Quadrivalent Inactivated Influenza Vaccine (IIV4) in pregnancy. Presented at the Advisory Committee on Immunization Practices meeting, Atlanta, GA; October 20, 2022.
- 79. CDC. Advisory Committee on Immunization Practices: GRADE: higher dose and adjuvanted influenza vaccines for persons aged ≥65 Years. Atlanta, GA: US Department of Health and Human Services, CDC; 2023. https://www.cdc.gov/vaccines/acip/recs/grade/influenzaolder-adults.html
- 80. CDC. Advisory Committee on Immunization Practices: evidence to recommendations (EtR) framework: higher dose and adjuvanted influenza vaccines for persons aged ≥65 years. Atlanta, GA: US Department of Health and Human Services, CDC; 2023. https://www.cdc.gov/vaccines/acip/recs/grade/influenza-older-adults-etr.html
- Thompson WW, Shay DK, Weintraub E, et al. Mortality associated with influenza and respiratory syncytial virus in the United States. JAMA 2003;289:179–86. PMID:12517228 https://doi.org/10.1001/ jama.289.2.179
- 82. CDC. Past seasons' vaccine effectiveness estimates. Atlanta, GA: US Department of Health and Human Services, CDC; 2023. https://www. cdc.gov/flu/vaccines-work/past-seasons-estimates.html
- DiazGranados CA, Dunning AJ, Kimmel M, et al. Efficacy of high-dose versus standard-dose influenza vaccine in older adults. N Engl J Med 2014;371:635–45. PMID:25119609 https://doi.org/10.1056/ NEJM0a1315727
- 84. Dunkle LM, Izikson R, Patriarca P, et al.; PSC12 Study Team. Efficacy of recombinant influenza vaccine in adults 50 years of age or older. N Engl J Med 2017;376:2427–36. PMID:28636855 https://doi. org/10.1056/NEJMoa1608862
- Keitel WA, Treanor JJ, El Sahly HM, et al. Comparative immunogenicity of recombinant influenza hemagglutinin (rHA) and trivalent inactivated vaccine (TIV) among persons > or =65 years old. Vaccine 2009;28:379–85. PMID:19879222 https://doi.org/10.1016/j. vaccine.2009.10.037
- 86. Izurieta HS, Chillarige Y, Kelman J, et al. Relative effectiveness of cell-cultured and egg-based influenza vaccines among elderly persons in the United States, 2017–2018. J Infect Dis 2019;220:1255–64. PMID:30561688 https://doi.org/10.1093/infdis/jiy716
- Izurieta HS, Chillarige Y, Kelman J, et al. Relative effectiveness of influenza vaccines among the United States elderly, 2018–2019. J Infect Dis 2020;222:278–87. PMID:32100009 https://doi.org/10.1093/ infdis/jiaa080
- 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. Clin Infect Dis 2021;73:e4251–9. PMID:33211809 https://doi.org/10.1093/cid/ciaa1727
- 89. Izurieta HS, Thadani N, Shay DK, et al. Comparative effectiveness of high-dose versus standard-dose influenza vaccines in US residents aged 65 years and older from 2012 to 2013 using Medicare data: a retrospective cohort analysis. Lancet Infect Dis 2015;15:293–300. PMID:25672568 https://doi.org/10.1016/S1473-3099(14)71087-4

- 90. Lu Y, Chillarige Y, Izurieta HS, et al. Effect of age on relative effectiveness of high-dose versus standard-dose influenza vaccines among US Medicare beneficiaries aged ≥65 years. J Infect Dis 2019;220:1511–20. PMID:31290553 https://doi.org/10.1093/infdis/jiz360
- 91. Pelton SI, Divino V, Shah D, et al. Evaluating the relative vaccine effectiveness of adjuvanted trivalent influenza vaccine compared to high-dose trivalent and other egg-based influenza vaccines among older adults in the US during the 2017–2018 influenza season. Vaccines (Basel) 2020;8:446. PMID:32784684 https://doi.org/10.3390/ vaccines8030446
- 92. Richardson DM, Medvedeva EL, Roberts CB, Linkin DR; Centers for Disease Control and Prevention Epicenter Program. Comparative effectiveness of high-dose versus standard-dose influenza vaccination in community-dwelling veterans. Clin Infect Dis 2015;61:171–6. PMID:25829001 https://doi.org/10.1093/cid/civ261
- 93. 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. J Infect Dis 2017;215:510–7. PMID:28329311 https://doi.org/10.1093/infdis/jiw641
- 94. Young-Xu Y, Snider JT, van Aalst R, et al. Analysis of relative effectiveness of high-dose versus standard-dose influenza vaccines using an instrumental variable method. Vaccine 2019;37:1484–90. PMID:30745146 https://doi.org/10.1016/j.vaccine.2019.01.063
- 95. 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 Surveill 2020;25:1900401. PMID:32431290 https://doi.org/10.2807/1560-7917. ES.2020.25.19.1900401
- 96. Young-Xu Y, Van Aalst R, Mahmud SM, et al. Relative vaccine effectiveness of high-dose versus standard-dose influenza vaccines among Veterans Health Administration patients. J Infect Dis 2018;217:1718–27. PMID:29452380 https://doi.org/10.1093/infdis/jiy088
- 97. Pelton SI, Divino V, Postma MJ, et al. A retrospective cohort study assessing relative effectiveness of adjuvanted versus high-dose trivalent influenza vaccines among older adults in the United States during the 2018-19 influenza season. Vaccine 2021;39:2396–407. PMID:33810903 https://doi.org/10.1016/j.vaccine.2021.03.054
- 98. van Aalst R, Gravenstein S, Mor V, et al. Comparative effectiveness of high dose versus adjuvanted influenza vaccine: a retrospective cohort study. Vaccine 2020;38:372–9. PMID:31606249 https://doi. org/10.1016/j.vaccine.2019.09.105
- Johansen ND, Modin D, Nealon J, et al. A pragmatic randomized feasibility trial of influenza vaccine. N Engl J Med Evid 2023;2:1–11. https://doi.org/10.1056/EVIDoa2200206
- 100. Bosaeed M, Kumar D. Seasonal influenza vaccine in immunocompromised persons. Hum Vaccin Immunother 2018;14:1311–22. PMID:29485353 https://doi.org/10.1080/21645 515.2018.1445446
- 101. Rubin LG, Levin MJ, Ljungman P, et al.; Infectious Diseases Society of America. 2013 IDSA clinical practice guideline for vaccination of the immunocompromised host. Clin Infect Dis 2014;58:e44–100. PMID:24311479 https://doi.org/10.1093/cid/cit684
- 102. CDC. Influenza antiviral medications: summary for clinicians. Atlanta, GA: US Department of Health and Human Services, CDC; 2023. https://www.cdc.gov/flu/professionals/antivirals/summary-clinicians.htm
- 103. Greenhawt M, Turner PJ, Kelso JM. Administration of influenza vaccines to egg allergic recipients: a practice parameter update 2017. Ann Allergy Asthma Immunol 2018;120:49–52. PMID:29273128 https://doi.org/10.1016/j.anai.2017.10.020

- 104. Byington CL, Maldonado YA, Barnett ED, et al.; Committee on Infectious Diseases. Committee on Infectious Diseases. Recommendations for prevention and control of influenza in children, 2016–2017. Pediatrics 2016;138:e20162527. PMID:27600320 https://doi.org/10.1542/peds.2016-2527
- 105. Anvari S, Scurlock AM, Jones SM, Perry TT, Hall PH. Influenza vaccination can be safely administered to most egg-allergic patients. J Ark Med Soc 2011;108:103–4. PMID:23252020
- 106. Chung EY, Huang L, Schneider L. Safety of influenza vaccine administration in egg-allergic patients. Pediatrics 2010;125:e1024–30. PMID:20368312 https://doi.org/10.1542/peds.2009-2512
- 107. Comeau J, Orr K, Mehr S, Wood N. ASCIA-P6: why is Australian policy still cautious with influenza vaccination in severe egg allergic children? Intern Med J 2016;46(Suppl 4):6. https://doi.org/10.1111/imj.6\_13197
- 108. Des Roches A, Paradis L, Gagnon R, et al.; PCIRN (Public Health Agency of Canada/Canadian Institutes of Health Research Influenza Research Network). Egg-allergic patients can be safely vaccinated against influenza. J Allergy Clin Immunol 2012;130:1213–1216.e1. PMID:23021881 https://doi.org/10.1016/j.jaci.2012.07.046
- 109. Erlewyn-Lajeunesse M, Keeton D, King R, et al. Multicentre audit of influenza immunisation in children allergic to egg. United Kingdom: Allergy: European Journal of Allergy and Clinical Immunology; 2010. https://onlinelibrary.wiley.com/doi/ epdf/10.1111/j.1398-9995.2010.02393.x
- 110. Esposito S, Gasparini C, Martelli A, et al. Safe administration of an inactivated virosomal adjuvanted influenza vaccine in asthmatic children with egg allergy. Vaccine 2008;26:4664–8. PMID:18639601 https://doi.org/10.1016/j.vaccine.2008.06.095
- 111. Greenhawt MJ, Spergel JM, Rank MA, et al. Safe administration of the seasonal trivalent influenza vaccine to children with severe egg allergy. Ann Allergy Asthma Immunol 2012;109:426–30. PMID:23176882 https://doi.org/10.1016/j.anai.2012.09.011
- 112. Howe LE, Conlon ASC, Greenhawt MJ, Sanders GM. Safe administration of seasonal influenza vaccine to children with egg allergy of all severities. Ann Allergy Asthma Immunol 2011;106:446–7. PMID:21530882 https://doi.org/10.1016/j.anai.2011.01.024
- 113. James JM, Zeiger RS, Lester MR, et al. Safe administration of influenza vaccine to patients with egg allergy. J Pediatr 1998;133:624–8. PMID:9821418 https://doi.org/10.1016/S0022-3476(98)70101-5
- 114. Leo SH, Dean J, Chan ES. Safety of H1N1 and seasonal influenza vaccines in egg allergic patients in British Columbia. All Asth Clin Immun 2010;6(S2):P4. https://doi.org/10.1186/1710-1492-6-S2-P4
- 115. Paschall VL, Siles RI, Bhatti H, Nolder PJ, Melton AL. Do egg-specific IgE levels predict reactions to seasonal influenza or H1N1 vaccination? J Allergy Clin Immunol 2011;127:AB182. https://doi.org/10.1016/j. jaci.2010.12.725
- 116. Shimizu M, Imai T, Yamazaki S, et al. Safety of influenza vaccination in children with severe allergy to hen's eggs: a prospective case series study. Arerugi 2016;65:128–33. PMID:27086959
- 117. Thanik ES, Cox AL, Sampson HA. Administration of a low eggcontaining influenza vaccine [Fluarix] in an egg-allergic pediatric population. J Allergy Clin Immunol 2010;125(Suppl 1):AB25. https:// doi.org/10.1016/j.jaci.2009.12.130
- 118. Hotte SL, Lejtenyi C, Primeau M. A 6-year experience with influenza vaccination in egg allergic patients. J Allergy Clin Immunol 2008;121:S239. https://doi.org/10.1016/j.jaci.2007.12.947
- 119. Park AY, Pien GC, Stinson R, Brown-Whitehorn TF, Spergel JM. Administration of influenza vaccine to patients with egg allergy. J Allergy Clin Immunol 2008;121:S240. https://doi.org/10.1016/j. jaci.2007.12.949

- 120. Des Roches A, Samaan K, Graham F, et al. Safe vaccination of patients with egg allergy by using live attenuated influenza vaccine. J Allergy Clin Immunol Pract 2015;3:138–9. PMID:25577640 https://doi. org/10.1016/j.jaip.2014.08.008
- 121. Turner PJ, Southern J, Andrews NJ, Miller E, Erlewyn-Lajeunesse M; SNIFFLE-2 Study Investigators. Safety of live attenuated influenza vaccine in young people with egg allergy: multicentre prospective cohort study. BMJ 2015;351:h6291. PMID:26645895 https://doi. org/10.1136/bmj.h6291
- 122. Turner PJ, Southern J, Andrews NJ, et al.; SNIFFLE Study Investigators. Safety of live attenuated influenza vaccine in atopic children with egg allergy. J Allergy Clin Immunol 2015;136:376–81. PMID:25684279 https://doi.org/10.1016/j.jaci.2014.12.1925
- 123. Upton JEM, Hummel DB, Kasprzak A, Atkinson AR. No systemic reactions to influenza vaccination in egg-sensitized tertiary-care pediatric patients. Allergy Asthma Clin Immunol 2012;8:2. PMID:22385977 https://doi.org/10.1186/1710-1492-8-2
- 124. Spiegel W, Anolik R. Administration of H1N1 vaccine in an egg allergic population. Ann Allergy Asthma Immunol 2010;105:A4. https://doi. org/10.1016/j.anai.2010.09.019
- 125. Siret-Alatrista A, Bouali F, Demoly M, Bousquet PJ, Demoly P. The 2009-2010 H1N1 vaccination campaign for patients with egg allergy in a region of France. Allergy 2011;66:298–9. PMID:20716317 https://doi.org/10.1111/j.1398-9995.2010.02461.x
- 126. Pitt T, Kalicinsky C, Warrington R, Cisneros N. Assessment of epicutaneous testing of a monovalent Influenza A (H1N1) 2009 vaccine in egg allergic patients. Allergy Asthma Clin Immunol 2011;7:3. PMID:21314925 https://doi.org/10.1186/1710-1492-7-3
- 127. Pien GC, LeBenger KS, Carotenuto DR, et al. Coordination of multidisciplinary resources for vaccination of egg-allergic individuals during an H1N1 (novel) influenza pandemic. Allergy Asthma Proc 2010;31:507–10. PMID:20943012 https://doi.org/10.2500/ aap.2010.31.3396
- 128. Greenhawt MJ, Chernin AS, Howe L, Li JT, Sanders G. The safety of the H1N1 influenza A vaccine in egg allergic individuals. Ann Allergy Asthma Immunol 2010;105:387–93. PMID:21055666 https://doi. org/10.1016/j.anai.2010.08.015
- 129. Gagnon R, Primeau MN, Des Roches A, et al.; PHAC-CIHR Influenza Research Network. Safe vaccination of patients with egg allergy with an adjuvanted pandemic H1N1 vaccine. J Allergy Clin Immunol 2010;126:317–23. PMID:20579720 https://doi.org/10.1016/j. jaci.2010.05.037
- 130. Forsdahl BA. Reactions of Norwegian children with severe egg allergy to an egg-containing influenza A (H1N1) vaccine: a retrospective audit. BMJ Open 2012;2:e000186. PMID:22223837 https://doi. org/10.1136/bmjopen-2011-000186
- 131. Didenko I, Ferreira F, Ferrao A, et al. Influenza A (H1N1) vaccination in individuals with egg allergy. Lisbon, Portugal: Revista Portuguesa de Imunoalergoloica; 2010. https://www.spaic.pt/client\_files/rpia\_ artigos/vacinacao-para-a-influenza-a-(h1n1)-em-individuos-comalergia-ao-ovo.pdf
- 132. Schuler JE, King WJ, Dayneka NL, et al. Administration of the adjuvanted pH1N1 vaccine in egg-allergic children at high risk for influenza A/H1N1 disease. Can J Public Health 2011;102:196–9. PMID:21714318 https://doi.org/10.1007/BF03404895

- 133. Halsey NA, Griffioen M, Dreskin SC, et al.; Hypersensitivity Working Group of the Clinical Immunization Safety Assessment Network. Immediate hypersensitivity reactions following monovalent 2009 pandemic influenza A (H1N1) vaccines: reports to VAERS. Vaccine 2013;31:6107–12. PMID:24120547 https://doi.org/10.1016/j. vaccine.2013.09.066
- 134. Woo EJ. Allergic reactions after egg-free recombinant influenza vaccine: reports to the US Vaccine Adverse Event Reporting System. Clin Infect Dis 2015;60:777–80. PMID:25428412 https://doi.org/10.1093/cid/ciu948
- 135. Woo EJ, Moro PL, Cano M, Jankosky C. Postmarketing safety surveillance of trivalent recombinant influenza vaccine: reports to the Vaccine Adverse Event Reporting System. Vaccine 2017;35:5618–21. PMID:28886946 https://doi.org/10.1016/j.vaccine.2017.08.047
- 136. Woo EJ, Moro PL. Postmarketing safety surveillance of quadrivalent recombinant influenza vaccine: Reports to the vaccine adverse event reporting system. Vaccine 2021;39:1812–7. PMID:33678452 https://doi.org/10.1016/j.vaccine.2021.02.052
- 137. CDC. Influenza prevention: information for travelers. Atlanta, GA: US Department of Health and Human Services, CDC; 2023. https://www.cdc.gov/flu/school-business/travelersfacts.htm
- 138. World Health Organization. Global influenza programme: FluNet. Geneva, Switzerland: World Health Organization; 2023. https://www. who.int/tools/flunet
- 139. Millman AJ, Kornylo Duong K, Lafond K, Green NM, Lippold SA, Jhung MA. Influenza outbreaks among passengers and crew on two cruise ships: a recent account of preparedness and response to an everpresent challenge. J Travel Med 2015;22:306–11. PMID:26031322 https://doi.org/10.1111/jtm.12215
- 140. Mutsch M, Tavernini M, Marx A, et al. Influenza virus infection in travelers to tropical and subtropical countries. Clin Infect Dis 2005;40:1282–7. PMID:15825030 https://doi.org/10.1086/429243
- 141. Ratnam I, Black J, Leder K, et al. Incidence and risk factors for acute respiratory illnesses and influenza virus infections in Australian travellers to Asia. J Clin Virol 2013;57:54–8. PMID:23380660 https://doi. org/10.1016/j.jcv.2013.01.008
- 142. Uyeki TM, Zane SB, Bodnar UR, et al.; Alaska/Yukon Territory Respiratory Outbreak Investigation Team. Large summertime influenza A outbreak among tourists in Alaska and the Yukon Territory. Clin Infect Dis 2003;36:1095–102. PMID:12715302 https://doi. org/10.1086/374053
- 143. Rapivab (peramivir injection) [Package Insert]. Durham, NC: BioCryst; 2021.
- 144. Xofluza (baloxavir marboxil) [Package Insert]. South San Francisco, CA: Genentech; 2022. https://www.accessdata.fda.gov/drugsatfda\_ docs/label/2018/210854s000lbl.pdf
- 145. Food and Drug Administration. Guidance for industry: bioavailability and bioequivalence studies for orally administered drug products: general considerations. Silver Spring, MD: US Department of Health and Human Services, Food and Drug Administration; 2003.
- 146. Shingrix [Package Insert]. Durham, NC: GlaxoSmithKline; 2023.
- 147. Arexy [Package insert]. Durham, NC: GlaxoSmithKline; 2023.
- 148. Heplisav-B [Package Insert]. Emeryville, CA: Dynavax; 2023.
- 149. Levin MJ, Buchwald UK, Gardner J, et al. Immunogenicity and safety of zoster vaccine live administered with quadrivalent influenza virus vaccine. Vaccine 2018;36:179–85. PMID:28830693 https://doi. org/10.1016/j.vaccine.2017.08.029

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