National Center for Immunization and Respiratory Diseases



Principles of Vaccination

Pink Book Web-on-Demand Series July 2, 2024

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- Describe the fundamental principles of the immune response.
- Describe immunization best practices.
- Describe an emerging immunization issue.
- For each vaccine-preventable disease, identify those for whom routine immunization is recommended.
- For each vaccine-preventable disease, describe characteristics of the vaccine used to prevent the disease.
- Locate current immunization resources to increase knowledge of team's role in program implementation for improved team performance.

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The findings and conclusions in this presentation are those of the authors and do not necessarily represent the official position of the Centers for Disease Control and Prevention.

Immunity

Human Immune System

 Complex network of interacting cells and proteins whose purpose is to identify, and eliminate, foreign substances

American Academy of Pediatrics. Active and passive immunization. In: Kimberlin D, Brady M, Jackson M, et al., eds. Red Book: 2018 Report of the Committee on Infectious Diseases. 31st ed. Itasca, IL: American Academy of Pediatrics;2018:13–64 Pinkbook | Principles of Vaccination | Epidemiology of VPDs | CDC

Plotkin S. Vaccines, vaccination, and vaccinology. J Infect Dis 2003;187:1347–59.

Siegrist C. Vaccine immunology. In: Plotkin S, Orenstein W, Offit P, et al., eds. Plotkin's Vaccines. 7th ed. Elsevier;2018:16–34



Immunity

- The ability of the human body to:
 - Tolerate the presence of material indigenous to the body
 - Eliminate foreign substances
- Self vs. "non-self"

American Academy of Pediatrics. Active and passive immunization. In: Kimberlin D, Brady M, Jackson M, et al., eds. Red Book: 2018 Report of the Committee on Infectious Diseases. 31st ed. Itasca, IL: American Academy of Pediatrics;2018:13–64.

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Immunity, cont.

- Most organisms recognized as foreign.
 - Virus, bacteria, fungi
 - Immune system provides protection from infectious diseases.
- Immunity is generally specific to a single organism.
 - E.g., immunity against a specific bacteria will not protect against a specific virus.



American Academy of Pediatrics. Active and passive immunization. In: Kimberlin D, Brady M, Jackson M, et al., eds. Red Book: 2018 Report of the Committee on Infectious Diseases. 31st ed. Itasca, IL: American Academy of Pediatrics;2018:13–64.

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Antigen

- Live or inactivated substances (e.g., viruses, bacteria, toxins)
 - Capable of stimulating an immune response
- Antigen = antibody generator



Antibody

- Protein molecules (immunoglobulins)
- Help infection-fighting cells recognize and kill foreign organisms, including viruses
- Antibodies are produced by the body



Arms of the Immune System

- Humoral
- Cell-mediated

Antigen

Arms of the Immune System, cont.

- Humoral
 - Antibodies attach to invading organisms and interfere with its ability to produce more invading organisms.
 - Antibodies are produced by B-cells (lymphocytes) to bind to a corresponding antigen, such as a virus (lock and key mechanism).

Antibodies attaching to antigens

Antibody

Antigen

Antibody

Arms of the Immune System, cont.

- Cell-mediated
 - Involves the activation of T-cells, macrophages, and other substances that eliminate the antigen.
 - T-cells mature in the thymus gland.





Immune System



Which of the following helps infectionfighting cells recognize and kill a microorganism?

A. Antigen B. Antibody





Which of the following helps infection-fighting cells recognize and kill a microorganism?

A. Antigen



Types of Immunity: Passive and Active

Types of Immunity

- Passive immunity
- Active immunity

Passive Immunity

- Passive immunity is the transfer of antibody produced by one human, animal, or recombinant technology, to another.
- It is temporary protection that wanes with time.
- There are different sources.

Sources of Passive Immunity (1)

- Transfer of antibody from the mother through placenta to the infant.
- These antibodies can help protect the baby from some diseases during the first few months of life.



Sources of Passive Immunity (2)

- Many types of blood or blood products
- Homologous pooled human antibody (immune globulin or IG)
 - IgG antibody from the blood of thousands of adult donors
 - Hepatitis A and measles post-exposure prophylaxis (PEP)



Sources of Passive Immunity (3)

- Homologous human hyperimmune globulin
- From donors with high concentrations of a specific antibody
 - Hepatitis B Immune Globulin (HBIG), Human Rabies Immune Globulin (HRIG), Tetanus Immune globulin (TIG), Varicella Zoster Immune Globulin (Human) (VariZIG), Vaccinia Immune Globulin Intravenous (Human) (VIGIV)
- Heterologous hyperimmune serum
 - Antitoxin (e.g., diphtheria antitoxin)
 - Serum sickness

Sources of Passive Immunity (4)

- Monoclonal antibodies are derived from a single type, or clone, of antibody-producing cells (B-cells) or through recombinant technology.
 - Immune globulin from human sources is polyclonal (contains many kinds of antibodies).
 - Monoclonal antibody is specific to a single antigen or closely related group of antigens.
 - Used for diagnosis of and therapy for certain cancers and autoimmune and infectious diseases (e.g., COVID-19)
 - Used for prevention of certain infectious diseases and transplant rejection.
 - Monoclonal-antibody-derived drugs end in -mab.

Antibody for Prevention of RSV

- Nirsevimab (Beyfortus)
 - Monoclonal
 - Contains only RSV antibody
 - Will not interfere with the response to a live-virus vaccine

Active Immunity

- Protection produced by a person's own immune system
- The process of developing antibodies takes time.
 - It can take up to a few weeks.
- Lasts for many years, often lifetime



Sources of Active Immunity

• Infection with disease-causing form of organism



• Vaccination



Vaccination

- Active immunity produced by vaccine
 - Vaccine delivers an attenuated (weakened, nonpathogenic) or non-live (dead or disrupted) form of the pathogen.
- Vaccination primes the body to make antibodies when exposed to the specific organism the vaccine was designed to protect against.
 - Immunologic memory allows for an anamnestic response after the primary immune response so that antibody reappears when the antigen is introduced.





Which type of immunity lasts longer?

A. Passive immunity

B. Active immunity





Which type of immunity lasts longer?

A. Passive immunity

B. Active Immunity

Principles of Vaccination

5

Principles of Vaccination

 The more similar the antibody [or immune response] produced by vaccination is to antibody produced by natural disease (infection), the better the protection provided by the vaccine.

Factors that Affect Immune Response to Vaccines

- Presence of maternal antibodies
- Nature and amount of antigen in vaccine
- Route of administration
- Presence of an adjuvant (ingredient that promotes a stronger immune response)
- Storage and handling of vaccine
- Vaccine recipient
 - Age
 - Nutritional status
 - Genetics
 - Coexisting disease
 - Immune suppression

Community Immunity



Community Immunity, cont.



Community Immunity, cont.


Classification of Vaccines

Classification of Vaccines

- Live
 - Most live vaccines used in the United States are "live, attenuated," meaning that the microbe in the vaccine is alive but has been weakened (attenuated).

- Non-live
 - A non-live vaccine contains inactivated or dead virus or bacterium, or a fraction of the organism.
 - Non-live vaccines are sometimes referred to as inactivated vaccines.

Classification of Vaccines (2)

- Live
 - Viral or bacterial
- Non-live
 - Viral or bacterial

Classification of Vaccines (3)

- Live, attenuated
- Non-live
 - Whole-cell
 - Subunit
 - Toxoid
 - Recombinant
 - mRNA

Live, Attenuated Vaccines

- "Wild" virus or bacterium weakened by repeated passage in culture media
- Must replicate to produce an immune response
- Immune response similar to natural infection
- Usually produce long-lasting immunity with 1-2 doses
 - Except those administered orally or intranasally

Live, Attenuated Vaccines (2)

- Severe reactions possible in persons with immune compromise.
- Interference from circulating antibody, including:
 - Maternal antibody
 - Other sources of passive immunity, i.e., blood products
- Fragile must be stored and handled carefully

Live, Attenuated Vaccines

Viral

- Live Attenuated Influenza Vaccine (LAIV), Measles Mumps Rubella (MMR), Measles Mumps Rubella Varicella (MMRV), Mpox (Mpox)[‡], Varicella (VAR), Rotavirus (RV1,RV5), Dengue (DEN4CYD), Chikungunya, Yellow fever (YF), oral adenovirus^{*}, oral polio[†], Ebola
- Bacterial

Bacille Calmette-Guérin (BCG)[§],
 oral typhoid, oral cholera

[§]Not used in the United States for routine TB protection

^{*}Live, but not attenuated. Limited use in the United States

⁺Not used in the United States

[‡]Jynneos vaccine does not replicate and behaves like non-live vaccines

Individual Response to a Hypothetical Live, Attenuated Vaccine



Population Response to a Hypothetical Live, Attenuated Vaccine



Non-live Vaccines

- Cannot replicate
- May be less affected by circulating antibody than live vaccines.
- May require multiple doses
- Antibody titer diminishes with time
- May include an adjuvant

Non-live Vaccines

- Whole-cell, inactivated
 - Polio, hepatitis A, rabies
- Subunit
 - Antigens can be protein, polysaccharide, or combination of polysaccharide and protein molecule (i.e., conjugate vaccine)
- Toxoid
 - Diphtheria, tetanus
- Recombinant
 - Hepatitis B, HPV
- mRNA
 - COVID-19

Adjuvants

- An adjuvant is an ingredient used in some vaccines that helps create a stronger immune response.
- Adjuvants have been used safely in vaccines for decades.



Several Different Adjuvants are Used in U.S. Vaccines

Vaccines
DTaP (Daptacel, Infanrix), DTaP-HepB-IPV (Pediarix), DTaP-IPV (Kinrix, Quadracel), DTaP-IPV/Hib (Pentacel), DTaP-IPV-Hib-HepB (VAXELIS), HepA (Havrix, Vaqta), HepB (Engerix-B, PREHEVBRIO, Recombivax), HepA/HepB (Twinrix), HIB (PedvaxHIB), HPV (Gardasil 9), MenB (Bexsero, Trumenba), Pneumococcal (Prevnar 20, VAXNEUVANCE), Td (Tenivac, no trade name), Tdap (Adacel, Boostrix),
RZV (Shingrix)
HepB (Heplisav-B)
COVID-19 (Novavax)
Influenza (Fluad and Fluad Quadrivalent)

Individual Response to Hypothetical Non-live Vaccine



Population Response to Hypothetical Non-live Vaccine





Which type of vaccine must replicate to generate an immune response?

A. Live, attenuated vaccinesB. Non-live vaccines





Which type of vaccine must replicate to generate an immune response?

A. Live, attenuated vaccines

B. Non-live vaccines

Capsular Polysaccharide



Pure Polysaccharide Vaccines

- Immune response typically T-cell-independent
- Not consistently immunogenic in children younger than 2 years of age
- No booster response
- Antibody with less functional activity (IgM rather than IgG)
- Immunogenicity improved by conjugation
 - i.e., combined with a protein

Polysaccharide Vaccines

• Pure polysaccharide

- Pneumococcal (PPSV23)
- Salmonella typhi (Vi)

• Conjugate polysaccharide

- Haemophilus influenzae type b (Hib)
- Pneumococcal (PCV13, PCV15, PCV20)
- Meningococcal ACWY





Which type of polysaccharide vaccine has improved immunogenicity?

A. Pure polysaccharide vaccineB. Conjugated polysaccharide vaccine



Which type of polysaccharide vaccine has improved immunogenicity?

A. Pure polysaccharide vaccine

B. Conjugated polysaccharide vaccine

ACIP Immunization Schedules

Recommended Child and Adolescent Immunization Schedule for Ages 18 Years or Younger, United States, 2024

Vaccine	Birth	1 mo	2 mos	4 mos	6 mos	9 mos	12 mos	15 mos	18 mos	19–23 mos	2–3 yrs	4–6 yrs	7-10 yrs	11–12 yrs	13–15 yrs	16 yrs	17–18 yrs		
Hepatitis B (HepB)	1# dose	4 2 nd (dose>				3 rd dose		>										
Rotavirus (RV): RV1 (2-dose series), RV5 (3-dose series)			1 st dose	2 nd dose	See Notes														
Diphtheria, tetanus, acellular pertussis (DTaP <7 yrs)			1 st dose	2 nd dose	3 ^{el} dose			∢ 4 th d	lose ►			5 th dose							
Haemophilus influenzae type b (Hib)			1 st dose	2 nd dose	See Notes		<a>3rd or 4 See 1	th dose		8									
Pneumococcal conjugate (PCV13)			1 st dose	2 nd dose	3 rd dose		∢ 4 th c	dose>											
Inactivated poliovirus (IPV <18 yrs)			1ª dose	2 nd dose	•		3ª dose					4 th dose							
Influenza (IIV4)						Annual vaccination 1 or 2 doses							Annual vaccination 1 dose only						
Influenza (LAIV4)											Annua 1 o	l vaccination r 2 doses	n	Annua	lvaccinatior	n 1 dose on	ly		
Measles, mumps, rubella (MMR)					See Notes							2 nd dose	se						
Varicella (VAR)							< 1= dose>					2 nd dose							
Hepatitis A (HepA)					See 1	Notes		2-dose serie	s, See Note	s									
Tetanus, diphtheria, acellular pertussis (Tdap ≥7 yrs)														1 dose					
Human papillomavirus (HPV)														See Notes					
Meningococcal (MenACWY-D ≥9 mos, MenACWY-CRM ≥2 mos, MenACWY-TT ≥2years)								See Notes						1 [#] dose		2 nd dose			
Meningococcal B (MenB-4C, MenB- FHbp)															See No	ites			
Pneumococcal polysaccharide (PPSV23)														See Notes					
Dengue (DEN4CYD; 9-16 yrs)													Se	ropositive i (S	n endemic a ee Notes)	reas only			
Range of recommended ages for all children	Range of r for catch-u	ecommend p vaccinati	ed ages on	Rar	nge of recon	nmended a n-risk group	ges s	Recomr can beg	mended vac in in this ag	cination le group	Re	commende shared clin	ed vaccinatio	m based making	No	recomment ot applicabl	ndation/ e		

Child and Adolescent Immunization Schedule – Healthcare Providers | CDC

Recommended Catch-up Immunization Schedule for Children and Adolescents Who

Start Late or Who Are More than 1 Month Behind, United States, 2024

 Table 2
 Recommended Catch-up Immunization Schedule for Children and Adolescents Who Start Late or Who Are More than 1 Month Behind, United States, 2024

The table below provides catch-up schedules and minimum intervals between doses for children whose vaccinations have been delayed. A vaccine series does not need to be restarted, regardless of the time that has elapsed between doses. Use the section appropriate for the child's age. Always use this table in conjunction with Table 1 and the Notes that follow.

raccine	Minimum Age for		Minimum Interval Between Doses		
	Dose 1	Dose 1 to Dose 2	Dose 2 to Dose 3	Dose 3 to Dose 4	Dose 4 to Dose 5
Hepatitis B	Birth	4 weeks	8 weeks and at least 16 weeks after first dose minimum age for the final dose is 24 weeks		00101000000
lotavirus	6 weeks Maximum age for first dose is 14 weeks, 6 days.	4 weeks	4 weeks maximum age for final dose is 8 months, 0 days		
Diphtheria, tetanus, and cellular pertussis	6 weeks	4 weeks	4 weeks	6 months	6 months A fifth dose is not ne if the fourth dose wa administered at age older and at least 6 m after dose 3
iaemophilus influenzae ype b	6 weeks	No further does needed iffnet does was administered at age 15 months or older. 4 weeks iffnet does was administered before the 1* birthday. 8 weeks (as final dose) if first does was administered at age 12 through 14 months.	No further does needed if previous does was administered at age 15 months or older 4 weeks 16 current age is younger than 12 months and first dose was administered at younger than age 7 months and at least 1 previous dose was PPP-1 (Actilie) ⁶ Pentacel ⁶ , Hieberk ⁶) Xaeell ⁶ or unknown 8 weeks and age 12 through 59 months (as final dose 8 weeks and age 12 through 59 months (as final dose was administered at age 7 through 11 months; OR 16 current age is younger than 12 months and first dose was administered at age 7 through 11 months; OR 16 current age is through 59 months: and first dose was administered before the ¹ bitthday and second dose was administered at younger than 15 months; OR 16 both doss was even eadministered before the 1 st bitthday	B weeks (as final dose) This dose only necessary for children age 12 through 59 months who received 3 doses before the 1° birthday.	
neumococcal conjugate	6 weeks	No further doese needed for healthy children if first does was administered at age 24 months or older 4 weeks if first dose was administered before the 1* birthdgy 8 weeks (as final does for healthy children) if first dose was administered at the 1* birthdgy or after	No further doses needed for healthy children if previous dose was administered at age 24 months or older devices age surgers than 12 months and previous dose was administered at <7 months old surgers (as final dose for healthy children) previous dose as administered between 7-11 months (wait until at least 12 months old: OR if current age is 12 months or older and at least 1 dose was administered before age 12 months	8 weeks (as final dose) This dose is only necessary for children age 12 through 59 months regardless of risk, or age 60 through 71 months with any risk, who received 3 doses before age 12 months.	
nactivated poliovirus	6 weeks	4 weeks	4 weeks if current age is <4 years 6 months (as final dose) if urrent age is 4 years or older	6 months (minimum age 4 years for final dose)	
Measles, mumps, rubella	12 months	4 weeks			
/aricella	12 months	3 months			
lepatitis A	12 months	6 months			
Meningococcal ACWY	2 months MenACWY-CRM 2 years MenACWY-TT	8 weeks	See Notes	See Notes	
			Children and adolescents age 7 through 18 years		
Meningococcal ACWY	Not applicable (N/A)	8 weeks			
fetanus, diphtheria; etanus, diphtheria, and icellular pertussis	7 years	4 weeks	4 weeks if first dose of DTaP/DT was administered before the 1* birthday 6 months (as final dose) if first dose of DTaP/DT or Tdap/Td was administered at or after the 1* birthday	6 months if first dose of DTaP/DT was administered before the 1 st birthday	
Human papillomavirus	9 years	Routine dosing intervals are recommended.			
lepatitis A	N/A	6 months			
Hepatitis B	N/A	4 weeks	8 weeks and at least 16 weeks after first dose		
nactivated poliovirus	N/A	4 weeks	6 months A fourth does is not necessary if the third dose was administered at age 4 years or older and at least 6 months after the previous dose.	A fourth dose of IPV is indicated if all previous doses were administered at <4 years OR if the third dose was administered <6 months after the second dose.	
Measles, mumps, rubella	N/A	4 weeks			
/aricella	N/A	3 months if younger than age 13 years.			
		- meens mage to years of order			

Table 3 Recommended Child and Adolescent Immunization Schedule by Medical Indication, United States, 2024

Always use this table in conjunction with Table 1 and the Notes that follow. Medical conditions are often not mutually exclusive. If multiple conditions are present, refer to guidance in all relevant columns. See Notes for medical conditions not listed.

Vaccine			HIV infect percentage	tion CD4 and count ^a	CCE lask av	Asplenia or persistent		Kidney failure,				
and other immunizing agents	Pregnancy	(excluding HIV infection)	<15% or <200mm	≥15% and ≥200mm	CSF leak or cochlear implant	complement component deficiencies	Heart disease or chronic lung disease	End-stage renal disease or on Dialysis	Chronic liver disease	Diabetes		
RSV-mAb (nirsevimab)		2nd RSV seasor	n	1 d RSV	ose depending on vaccination status,	maternal See Notes	2nd RSV season for chronic 1 dose depending on maternal lung disease (See Notes) RSV vaccination status, See Note					
Hepatitis B												
Rotavirus		SCID ^b										
DTaP/Tdap	DTaP Tdap: 1 dose each pregnancy											
Hib		HSCT: 3 doses	See Not	es		See Notes						
Pneumococcal												
IPV												
COVID-19		See N	lotes									
IIV4												
LAIV4							Asthma, wheezing: 2–4 years ^c					
MMR	•											
VAR	•											
Hepatitis A												
HPV	•	3 dose series	s. See Notes									
MenACWY												
MenB												
RSV (Abrysvo)	Seasonal administration, See Notes											
Dengue												
Мрох	See Notes											
Recommend eligible child documentati vaccination s	ed for all age- ren who lack on of a complete eries or	t recommended for all children t is recommended for some ildren based on increased risk fo severe outcomes from disease	or	Recomme children, a necessary or other in	nded for all age-eligit nd additional doses n based on medical cor dications. See Notes.	ole nay be ndition	Precaution: Might be indicated if benefit of protection outweighs risk of adverse reaction	Contraindicated recommended "Vaccinate after if indicated	or not pregnancy,	No Guidance/ Not Applicable		

Child and Adolescent Immunization Schedule – Healthcare Providers | CDC

Recommended Adult Immunization Schedule by Age Group, United States, 2024

Table 1 Recommended Adult Immunization Schedule by Age Group, United States, 2024

Vaccine	19-26 years	27–49 years		50–64 years		≥65 years							
COVID-19		1 or more doses of updated (2023–2024 Formula) vaccine (See Notes)											
Influenza inactivated (IIV4) or Influenza recombinant (RIV4)		1 dose annually											
Influenza live, attenuated (LAIV4)	1 dose a												
Respiratory Syncytial Virus (RSV)	Seasonal administration d	≥60 years											
Tetanus, diphtheria, pertussis		1 dose Tdap each pregnancy; 1 dose Td/Tdap for wound management (see notes)											
(ldap or ld)		1 dose Tdap, then T	d or Tda	ap booster every 10 years									
Measles, mumps, rubella (MMR)		For healthcare personnel, see notes											
Varicella (VAR)	2 dose (if born in 1980	s) or later)			2 doses								
Zoster recombinant (RZV)	2 doses for immunocompro	mising conditions (see notes)		2 doses									
Human papillomavirus (HPV)	2 or 3 doses depending on age at initial vaccination or condition	27 through 45 years											
Pneumococcal (PCV15, PCV20, PPSV23)						See Notes See Notes							
Hepatitis A (HepA)		2, 3, or 4 doses depending on vaccine											
Hepatitis B (HepB)		2, 3, or 4 doses depending on vaccine or condition											
Meningococcal A, C, W, Y (MenACWY)		1 or 2 doses depending on indication, see notes for booster recommendations											
Meningococcal B (MenB)	19 through 23 years 2 or 3 doses depending on vaccine and indication, see notes for booster recommendations												
Haemophilus influenzae type b (Hib)	1 or 3 doses depending on indication												
Мрох													
Recommended vaccination for adults lack documentation of vaccination, or	who meet age requirement, lack evidence of immunity	h an n	Recommended vaccinati clinical decision-making	on based on	shared No recommendation/ Not applicable								

Adult Immunization Schedule by Age | CDC

Table 2 Recommended Adult Immunization Schedule by Medical Condition or Other Indication, United States, 2024

Always use this table in conjunction with Table 1 and the Notes that follow. Medical conditions or indications are often not mutually exclusive. If multiple medical conditions or indications are present, refer to guidance in all relevant columns. See Notes for medical conditions or indications not listed.



Schedule Addendums



A Schedules Home

Child Immunization Schedule Addendum

Best Practice: Immunization Schedules

Recommende for ages 19 years o	ed Adu rolder	ult Immunizatio	on Sche	dule		2	024	1							
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Verlag	Althorn detires in	Trade purperiol	How to us	se the adu	ult immun	ization so	chedule								
COVD-19 vaccine	WOWNES	Continuity/ThereBoltTech COVD-19Vectine Spikews/SWadews COVD-19Vectine Receive COVD-19Vectine	1 Deservice recommended socializes	Assessmed two-different recommended	3 Review vacalies representers and	4 Faster contractions and percent test	5 Fasterine Activities	w 1							
//bemophilasintiaerape type b vaccine	185	Activitien Hilbarbon Pedvasivitien	by ape (Table 1)	vectivetions by readical condition or other indication	internals, and consideration alion special stations. (Meters)	for vectine type (Appendia)	i Okddendu	*							
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Liepatitis A and hepatitis Evacuine	Hep4-Hep3	Twints"													
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Meningococcol seragroups A, C, W, Y vaccine	MeetACWY CRM	Merwer" Merchaelle	Questions or co	mments											
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Mpaxwarrine	Mpos	Jymeos*	THE MANAGEMENT			11									024
Presimococcel conjugate recoine	PO/15 PO/20	Vakineuvence" Previou 20"	Helpful informa Complete Advisory	rtion Committee on inc	nunkost on Practices	(ACIP) recommen	dations		chedule*	Harrison		ahild an a			
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2024

Use the QR code located on the first page to access the online schedule to check for updates!



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Immunization Schedule Best Practices





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Know your patient's complete immunization history

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ONIPINFO@cdc.gov

Thank You from Atlanta!

For more information, contact CDC 1-800-CDC-INFO (232-4636) TTY: 1-888-232-6348 www.cdc.gov

The findings and conclusions in this report are those of the authors and do not necessarily represent the official position of the Centers for Disease Control and Prevention.



