

Supported by an educational grant from Merck & Co., Inc.



Activity Description

Target Audience

This activity is designed as a comprehensive approach to address the practice needs of primary care providers, including primary care physicians, doctors of osteopathy, physician assistants, nurse practitioners, and allied healthcare professionals, who are at the forefront of caring for adult patients eligible for immunizations and/or at risk for vaccine-preventable diseases.

Learning Objectives

At the conclusion of the educational activity, the learner should be able to:

- Evaluate the latest clinical research on the impact of HPV vaccination in the prevention of various types of cancer in men and women
- Identify strategies to adhere to ACIP HPV vaccination recommendations and overcome barriers by both healthcare providers and patients to vaccinate younger adults
- Describe the clinical consequences of pneumococcal disease and its associated complications among older adults
- State the latest ACIP recommendations for pneumococcal vaccination among older adults

Faculty and Disclosure

Michael J. Donnelly, MD, FACP, FAAP
Professor of Medicine and Pediatrics
The Philip Calcagno/Jeane and James Dixon Chair in Pediatrics
Georgetown University
Pediatrician-in-Chief
MedStar Georgetown University Hospital
Washington, DC

Dr. Michael Donnelly does not have any relevant financial relationships to disclose. *Dr. Donnelly does not intend to discuss off-label uses of products.*

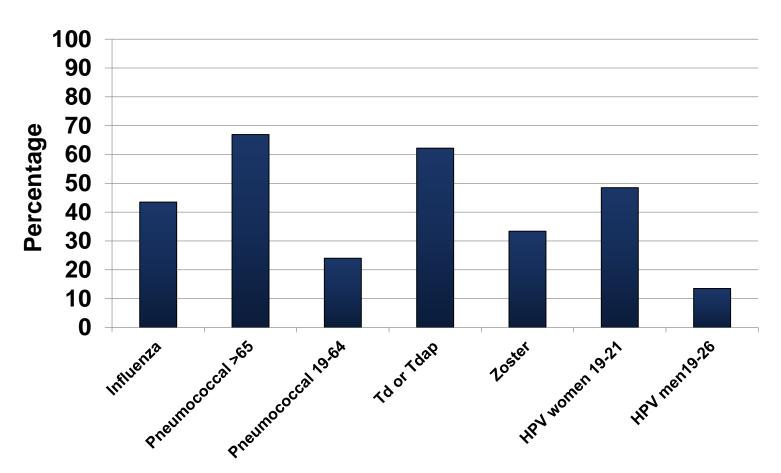
No (other) speakers, authors, planners or content reviewers have any relevant financial relationships to disclose.

Content review confirmed that the content was developed in a fair, balanced manner free from commercial bias. Disclosure of a relationship is not intended to suggest or condone commercial bias in any presentation, but it is made to provide participants with information that might be of potential importance to their evaluation of a presentation.

Burden of Vaccine-Preventable Disease

- Influenza
 - Since 2010
 - 140,000–710,000 flu-related hospitalizations per year
 - Between 12,000 and 56,000 deaths yearly
- Hepatitis B
 - 700k–1.4M people live with chronic hepatitis B
- We will take a close look at pneumococcal disease and HPV today

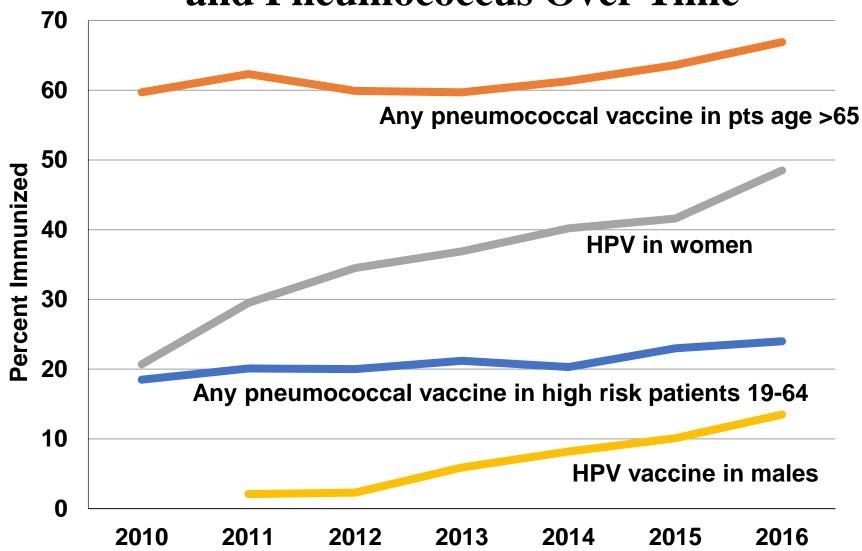
Vaccination Rates 2016



CDC. National Health Interview Survey, 2016. https://www.cdc.gov/vaccines/imz-managers/coverage/adultvaxview/pubs-resources/NHIS-2016.html.

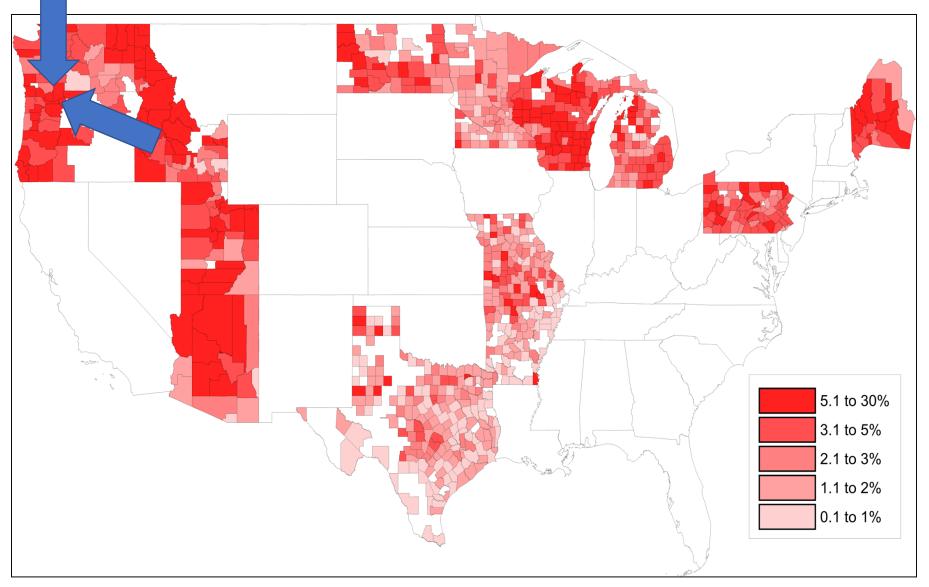


Adult Vaccination Rate for HPV and Pneumococcus Over Time



CDC. National Health Interview Survey, 2016. https://www.cdc.gov/vaccines/imz-managers/coverage/adultvaxview/pubs-resources/NHIS-2016.html.

Vaccine Refusal



The state of the antivaccine movement in the United States: A focused examination of nonmedical exemptions in states and counties. Jacqueline K, et al. *PLOS Medicine*. June 12, 2018. https://doi.org/10.1371/journal.pmed.1002578



Friends!!! I am desperate here: I am a newly non vaccinator and I want to write about vaccination dangers for my final thesis in college: however. I

Superbugs and anti-vaxxers make WHO's list of 10 global health threats

By Holly Yan, CNN

Updated 12:58 AM ET, Mon January 21, 2019



Thank you all!!! ***



The fact that you cannot find any journals to back up your viewpoint MAY suggest you need to reassess your viewpoint...

US Adults Attitudes Toward Vaccines

American Osteopathic Association

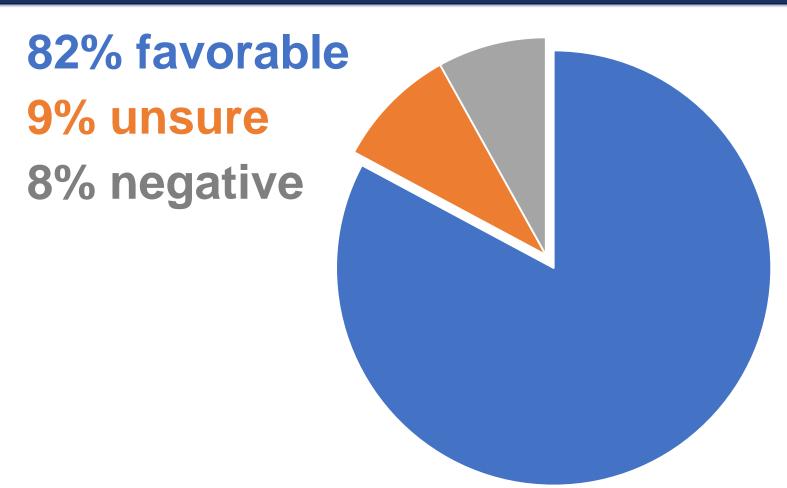
- The Harris Poll
- >2,000 US adults
- May 2019

45% of American Adults Doubt Vaccine Safety

Which of the following have caused you to doubt the safety of vaccines?

Nothing—I don't doubt the safety of vaccines 55%

Safety and Efficacy



Sources of Information

Top 3:

16% said Online Articles

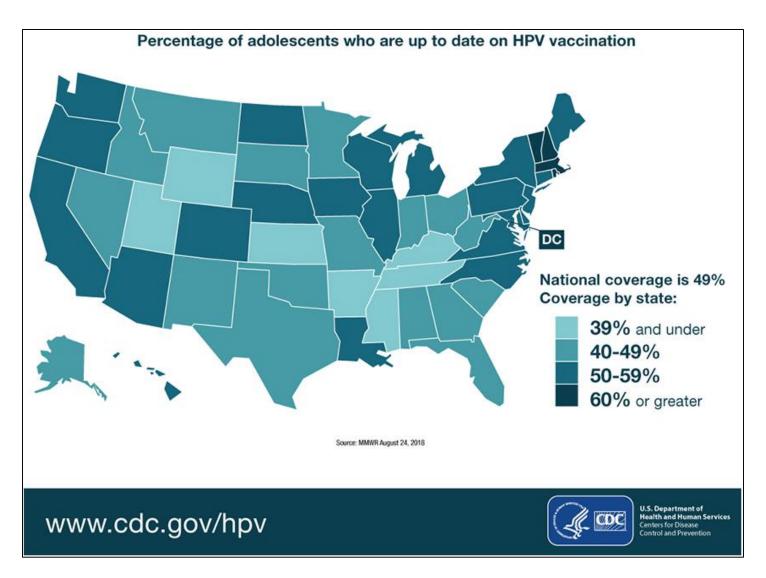
12% past wrongdoing by industry

11% info from Medical Experts

HPV vaccine it's not just warts



Kansas Vaccination Rates



Walker TY et al. Natl, Reg, State, and Selected Area Vaccination Coverage Among Adolescents Aged 13–17 Years — United States, 2018. MMWR 2019;68:718–723.

HPV-Related Cancer Incidence

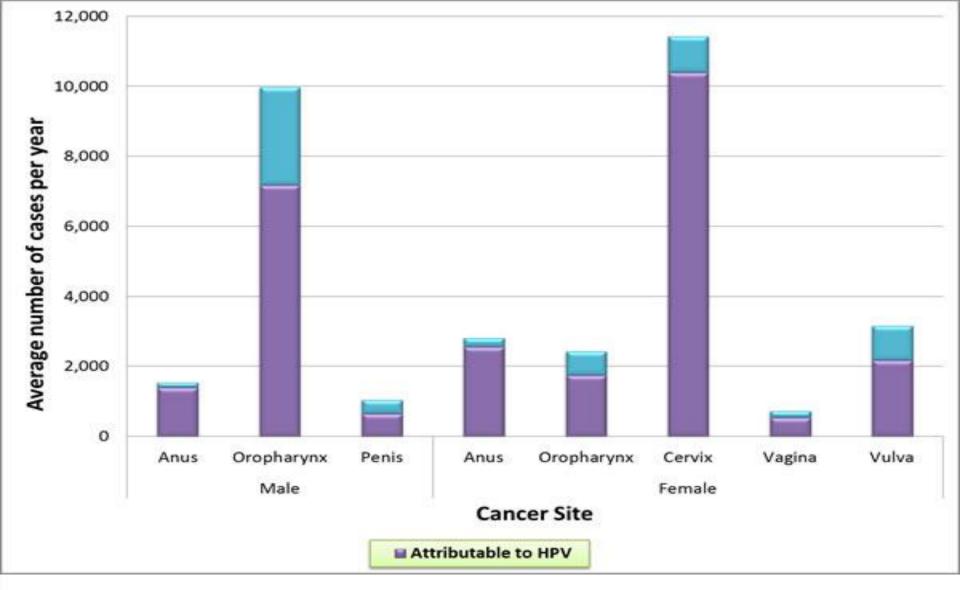
In 2015 alone-

CDC estimate- 43,371 HPV-related cancers

- From 1999–2015
 - HPV-associated cancer incidence
 - rose by 0.9%
 - cervical cancer dropped by 1.6%
- Estimated to result in ~18,000 deaths²

¹Van Dyne EA, et al. *MMWR Weekly*. 2018;67(33);918–924.

²American Cancer Society. Cancer Statistics Center. https://cancerstatisticscenter.cancer.org/#!/.



Over 90% of Anal and Cervical Cancers and 60% of Penile Cancers are HPV+

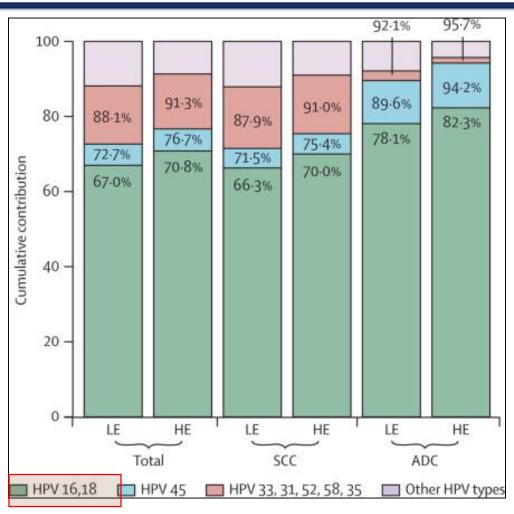
Supplement: Assessing the Burden of HPV-Associated Cancers in the United States. Cancer. 2008;113(S10):2837–3057.

Quiz

Which serotype of HPV is most carcinogenic?

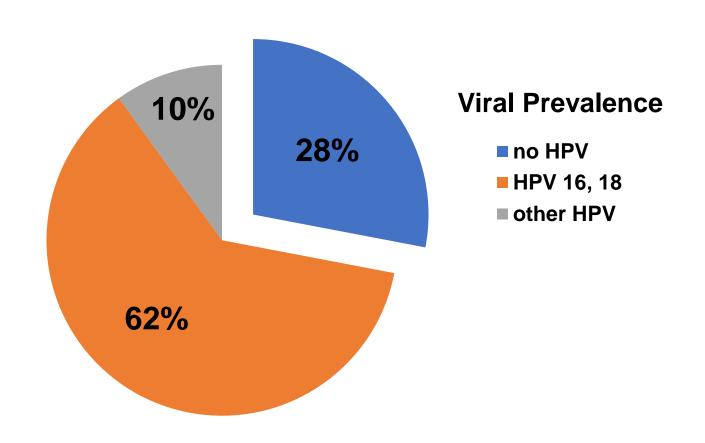
- A)6
- B) 11
- C)16
- D)18
- E)31

Genotype Attribution for Cervical Cancer

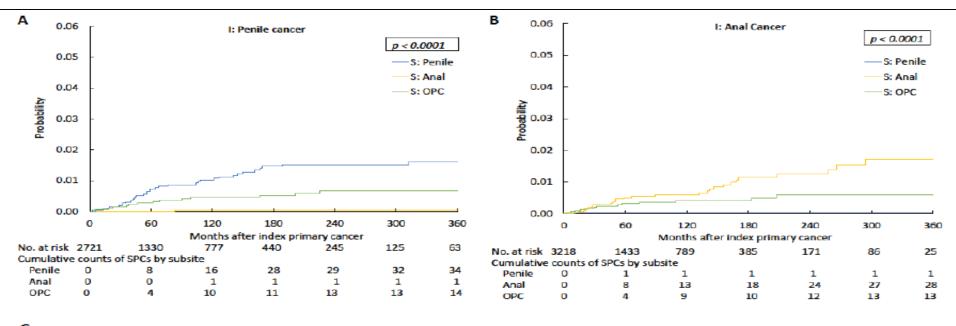


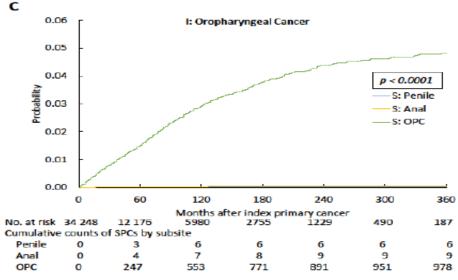
de Sanjose S, et al. Lancet Oncol. 2010;11:1048-1056.

Genotype Attribution: Oropharyngeal Cancer



Second Cancers Occur in Both Genders





Cumulative incidence of 2nd HPV-associated primary cancers among men

Suk R, et al. JAMA Network Open. 2018;1(5):e181999. doi:10.1001/jamanetworkopen.2018.

Vaccine is Effective for HPV 16 & 18

Reduces Cervical Infection

 Reduced incidentally detected 6- and 12-month infection by 94% and 91%¹

From 2003–06 to 2009–2012, 4vHPV-type prevalence decreased:

- 64% in 14–19-yr-olds
- 34% in 20–24-yr-olds³

Reduces Oropharyngeal Infection

• Estimated vaccine efficacy: 93%²

- 1. Paavonen J, et al. Lancet. 2009;374(9686):301-14.
- 2. Herrero R, et al. PLoS ONE. 2013;8:e68329.
- 3. Markowitz LE, et al. Pediatrics. 2016;137(2):e20151968.

Vaccine Rates in Teens are Increasing

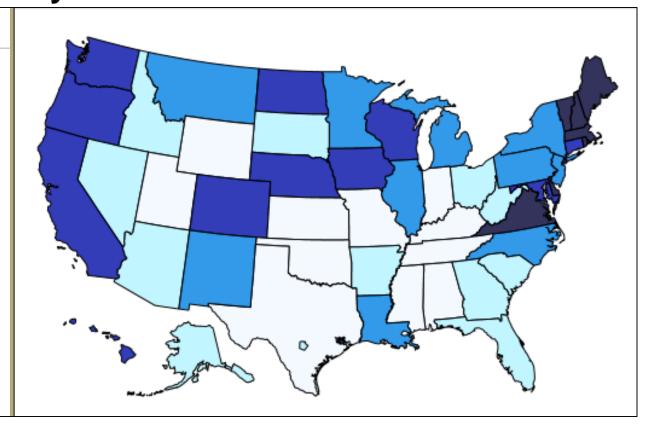
NIS-Teen 2017 rates

2017	2016
------	------

• Teen Girls 53.1% 49.5%

• Teen Boys 44.3% 37.5%

Legend (%) 46.9 - 59.3 59.4 - 65.0 65.1 - 69.1 69.2 - 73.0 73.1 - 91.9 NA

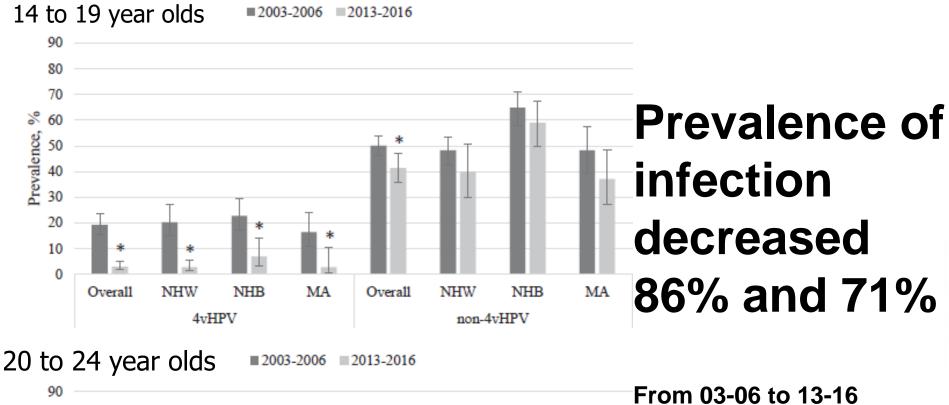


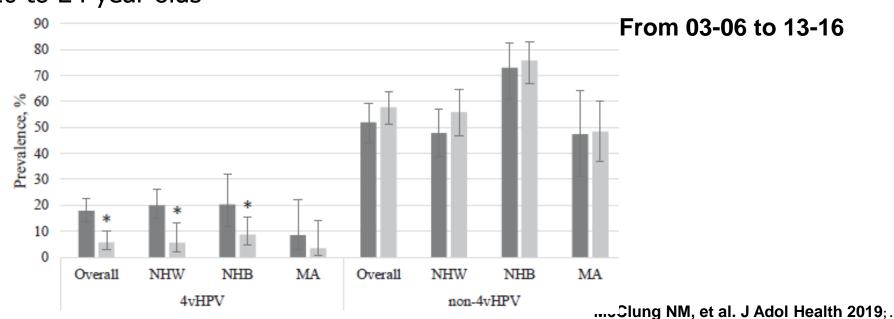
CDC. TeenVaxView. https://www.cdc.gov/vaccines/imz-managers/coverage/teenvaxview/data-reports/hpv/trend/index.html.

Quiz

HPV vaccine was initially recommended in the US in 2009 in women. Since that time, which of the following have been shown regarding the incidence of vaccine type HPV?

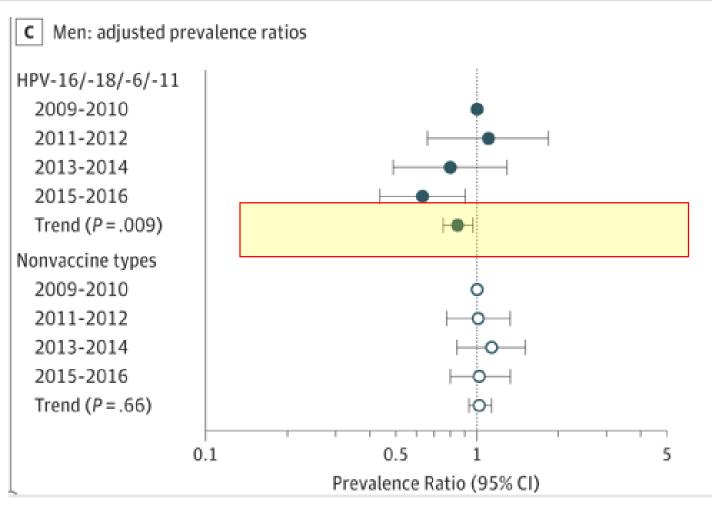
- A) Cervical infection has decreased in women who have been unvaccinated.
- B) Cervical infection is unchanged in women who are unvaccinated.
- C) Oropharyngeal infection in males has decreased.
- D) Both A and C are correct.





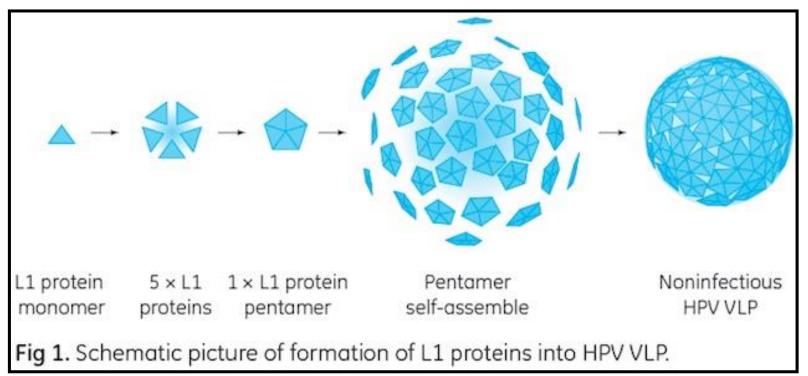
https://www.clinicalkey.com/#!/content/journal/1-s2.0-S1054139X19303532

Herd Immunity-Oral HPV Prevalence in Men



Chaturvedi AK, et al. JAMA 2019;322(10):977-979.

How Do We Make It??





http://biopharma-asia.com/technical-papers/use-capto-core-700-capto-q-impres-purification-human-papilloma-virus-like-particles/

ACIP Recommendations for HPV 9

- Earliest: age 9 years
- Recommended: age 11–12 years
- 2 doses: 0 then 6–12 months if <15 years otherwise 3 doses
- Catchup
 - Males: 13–21 years, "may" vaccinate up to 26 years
 - Females: age 13 up to 26 years

FDA Expands Indication to age 45 in October 2018

- 10-year efficacy in women 27–45 years using quadrivalent HPV, not 9vHPV
 - (Merck study done in Columbia)

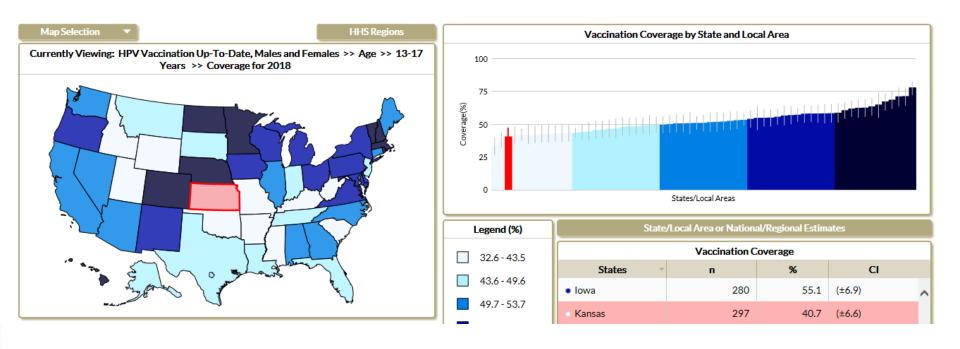
ACIP June 2019

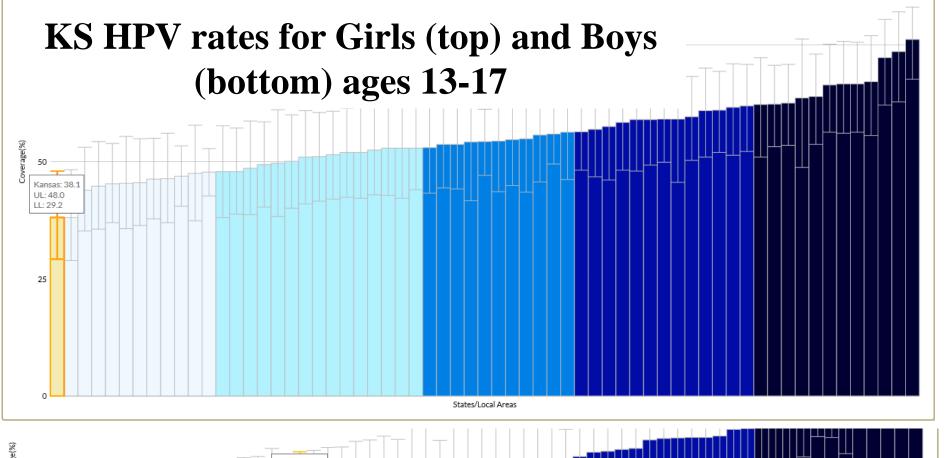
2 votes:

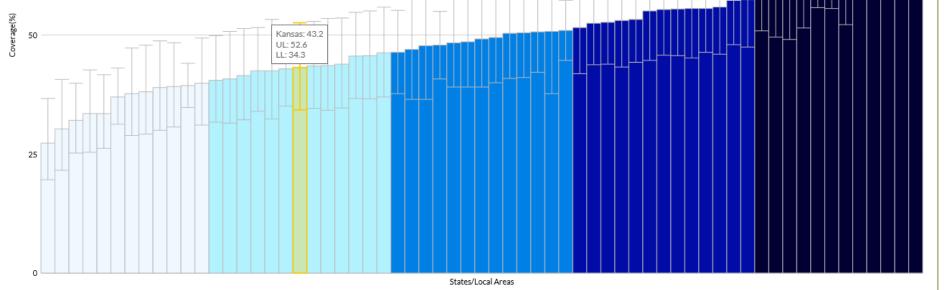
- HPV vaccination recommended for both males and females through age 26 years
- For ages 27 to 45 years, decision to vaccinate based on "shared clinical decision-making"

Recommendations based on June 2019 ACIP meeting. www.cdc.gov/vaccines/acip/meetings/downloads/agenda-archive/agenda-2019-06-508.pdf. Accessed July 23, 2019.

HPV Vaccination Coverage Among Adolescents 13-17 years by State, HHS Region, and the US, National Immunization Survey-Teen (NIS-TEEN), 2018







HPV Over the Ages

HPV DNA prevalence and report of at least 1 new sex partner — United States, 2013-2014

100

90

80

70

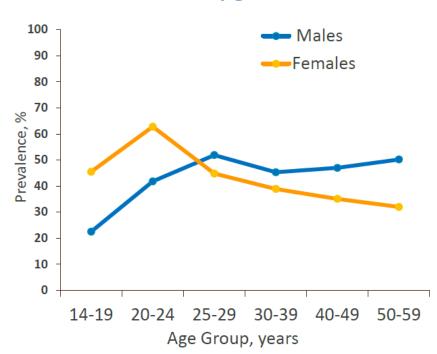
50 40

30

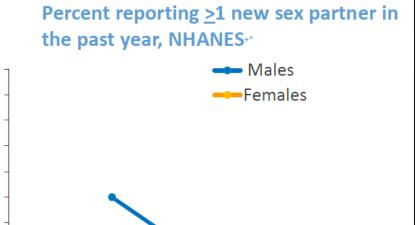
20

reporting ≥1 new partner in past year

Prevalence of any genital HPV, NHANES*



Lewis et al, JID 2018; Gargano et al, JID 2017 NHANES, National Health and Nutrition Examination Survey





12

*among sexually experienced persons

†CDC, unpublished data

Shared Decision Making Framework HPV Vaccine in 27-45 year olds

LEANS toward VACCINATE

- Having a new/multiple sex partners is a risk
- Vaccine efficacy is high if you haven't been exposed to that serotype

LEANS toward DON'T VACCINATE

- Monogamous long term relationships
- Not likely ever to be sexually active

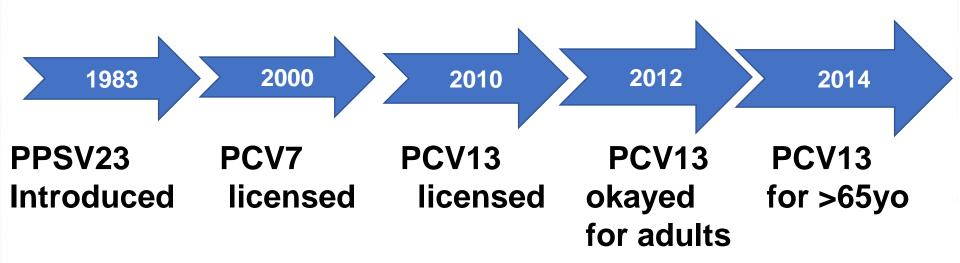
Vaccine Hesitancy

- Hesitancy has been increasing among patients and parents
- It is a spectrum: many are neither pro nor anti-vaccine but are in the middle.
- Provider introduction and recommendation is very important.

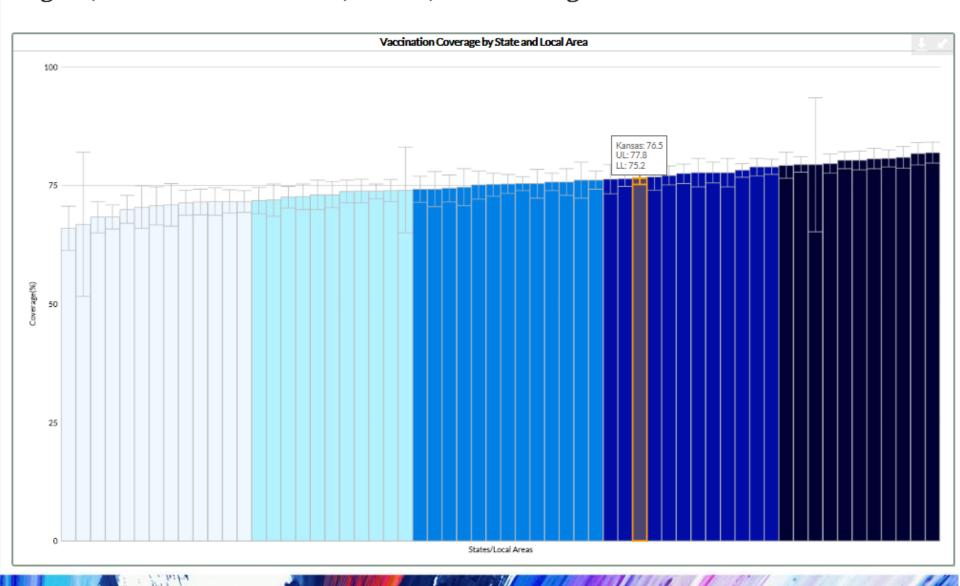
Hesitancy

- Presumptive style of communication
 - (continue discussion about why the vaccine is important vs deferring vaccination)
- Strong, direct communication
 - Even when parents verbally assertively expressed hesitancy, 33% were vaccinated same day.

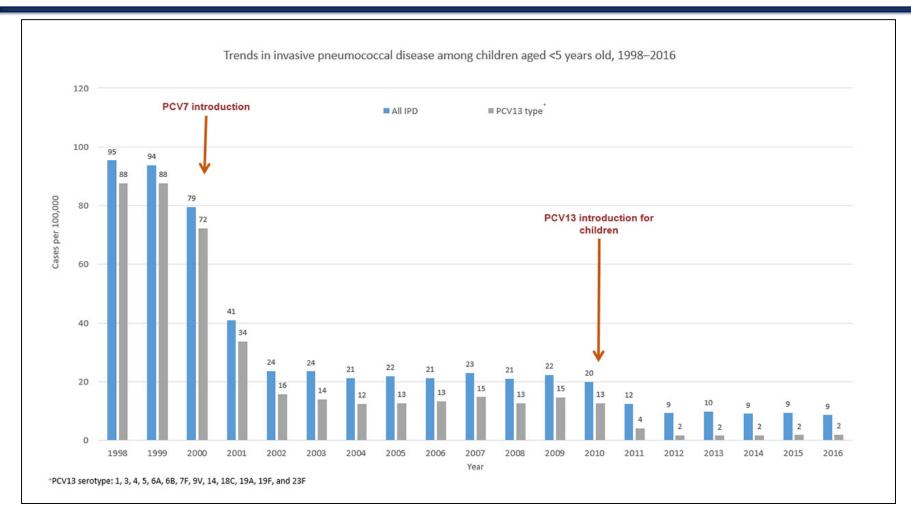
Pneumococcal Vaccines Remember there are 2 of them



Pneumococcal vaccination coverage among adults 18–64 years at increased risk and ≥65 years, Td and Tdap vaccination coverage among adults ≥18 years, and shingles vaccination coverage among adults ≥60 years by selected local area, state, HHS Region, and the United States, BRFSS, 2008 through 2017



Pneumococcal Vaccines Effects in Children



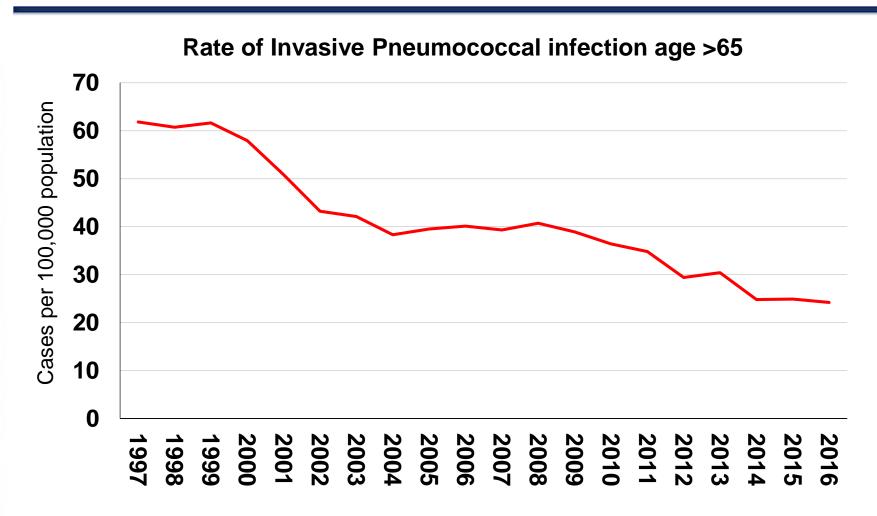
CDC. Active Bacterial Core surveillance. https://www.cdc.gov/abcs/reports-findings/survreports/spneu-types.html.

Quiz

Which of the following had the largest impact on the incidence of IPD in US adults?

- A) PPSV23 Vaccination for 65 year olds
- B) PCV7 Vaccination in children
- C) PCV7/13 Vaccination of High Risk 19-64 yr olds
- D) PPSV23 Vaccination of High Risk 19-64 year olds

Trends in Adults



CDC. Pneumococcal Disease: Surveillance and Reporting. https://www.cdc.gov/pneumococcal/surveillance.html.

Polysaccharide vs. Conjugate

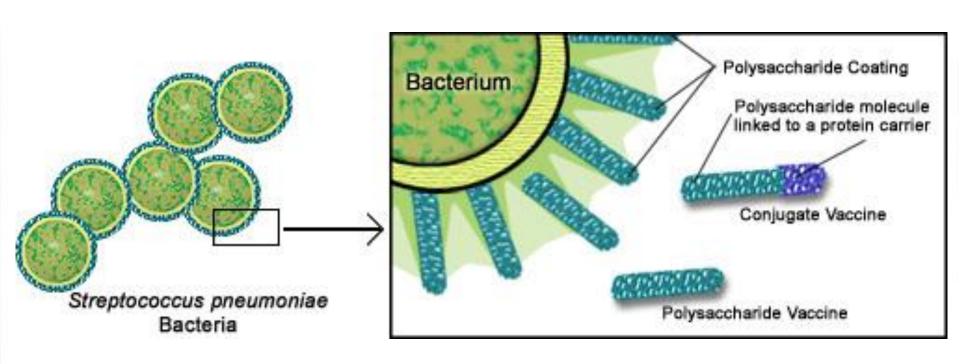


Image Source: https://www2a.cdc.gov/nip/isd/ycts/mod1/courses/genrec/10351.asp?seg=H.

Vaccine Effectiveness

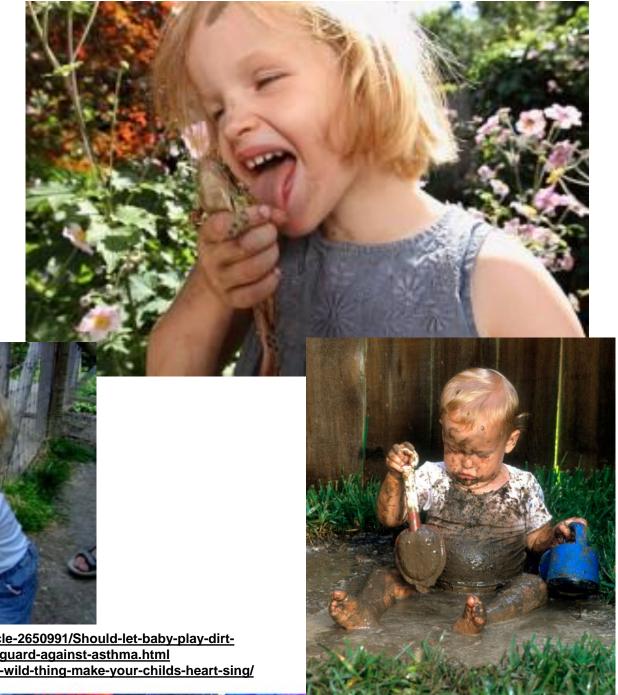
- PPSV23 is only modestly effective protecting against invasive disease (~50% reduction)
- Poor against non-bacteremic pneumonia (NPP) (4–17%)
- NPP is ~10 times more common than bacteremic pneumonia in adults

Conjugates vs Polysaccharides

- Induces herd immunity by decreasing naso-pharyngeal carriage.
 (Seen in Hib previously)
- Long-term immunity via T-cell activation also may decrease need for booster doses.



So Who was Getting the Adults Sick?



http://www.dailymail.co.uk/sciencetech/article-2650991/Should-let-baby-play-dirt-researchers-say-help-guard-boost-immune-guard-against-asthma.html
http://news.hugofox.com/2013/10/07/project-wild-thing-make-your-childs-heart-sing/

Whom to Vaccinate: ACIP 2012 Immunocompetent Patients

Underlying medical condition	PCV13	PPSV23
Chronic heart/lung/liver disease		~
Diabetics/smokers/alcoholism		✓
CSF leak	✓	✓
Immunocompromised Persons	PCV13	PPSV23 + rpt in 5yrs
Renal failure/nephrotic	✓	✓
Generalized or heme malignancy	✓	✓
latrogenic immunosuppression/transplant	✓	✓
Congenital or acquired asplenia Sickle cell/hemoglobinopathy	~	✓

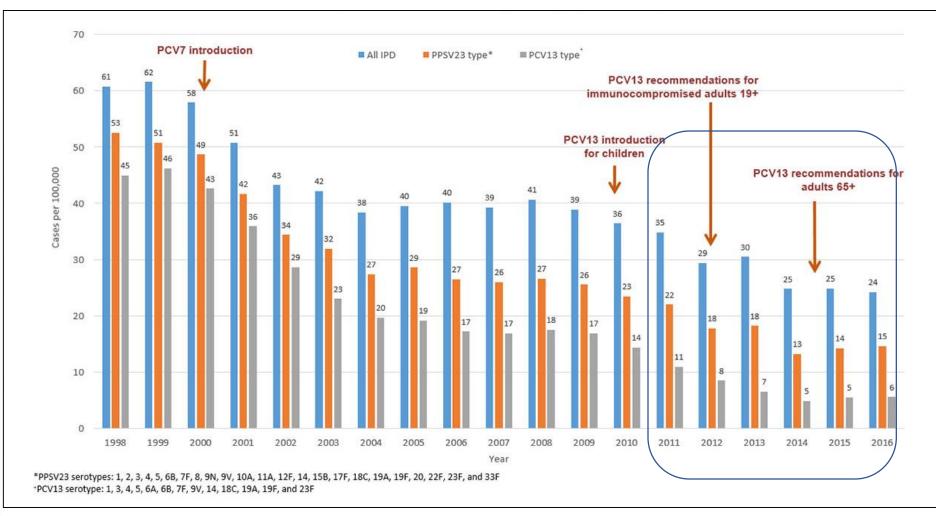
ACIP Meeting June 2019

For immunocompetent adults ≥65 years and no history of pneumococcal vaccine:

- Remove the recommendation for a single dose of PCV13
- "Shared clinical decision making" can be used to decide whom to vaccinate with PCV13
- A single dose of PPSV23 vaccine should continue to be administered

Recommendations based on June 2019 ACIP meeting. www.cdc.gov/vaccines/acip/meetings/downloads/agenda-archive/agenda-2019-06-508.pdf. Accessed July 23, 2019.

Trends in IPD Among Adults ≥65 Years, 1998–2016



CDC. Pneumococcal Disease: Surveillance and Reporting. https://www.cdc.gov/pneumococcal/surveillance.html.

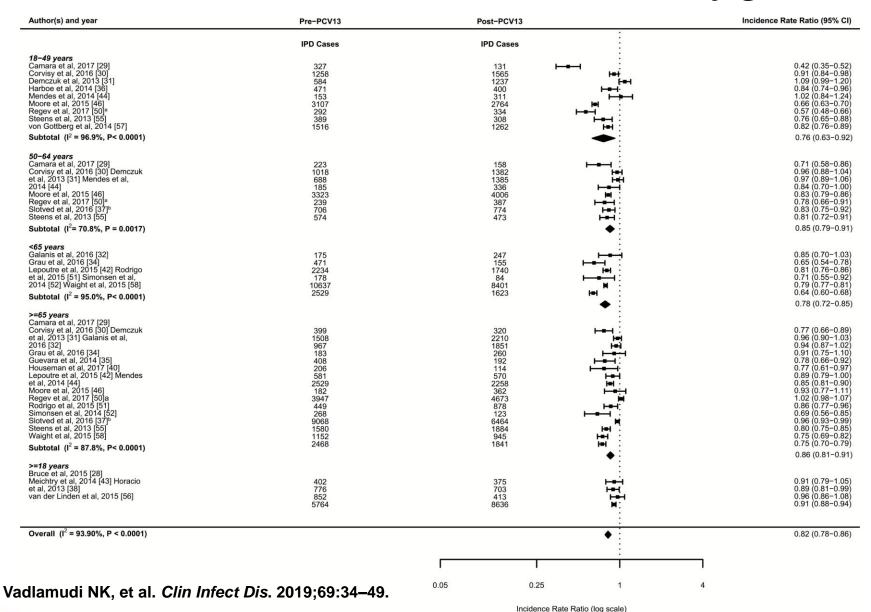
Indirect Effect of Pediatric Vaccines

- 9-fold reduction in IPD in adults >65 years
 In PCV13 types alone: 3-fold reduction
- Steady rate since 2014
- ~ Same as Europe
 - 77% reduction in PCV7 types
 - 38% reduction in PCV13-non7 types

Active Bacterial Core Surveillance, https://www.cdc.gov/abcs/reports-findings/surv-reports.html, comparing 2000 to 2014. Hanquet G, et al. *Thorax*. 2019;74:473-82.

ACIP meeting February 2019. https://www.cdc.gov/vaccines/acip/meetings/downloads/slides-2019-02/Pneumococcal-2-Pilishvili-508.pdf

16 Country Risk of Invasive Pneumococcal Disease, Before and After the Introduction of 13-valent Pneumococcal Conjugate ...



Annual Number Needed to Vaccinate (NNV) among Adults ≥65 Years Old*

Outcome	Incidence per 100,000	Vaccine Effectiveness (VE)	(95%CI)	NNV	(95%CI)
PCV13-type IPD	5ª	76% ^b	(48, 89)	26,300	(22,500, 41,700)
PCV13-type pneumonia,					
inpatient	17°-76d	43% ^e	(12, 63)	3,000–14,000	(2,100, 50,200)
PCV13-type pneumonia,					
_outpatient	88 ^f	43% ^e	(12, 63)	2,600	(1,800, 9,500)

^{*}Calculation: NNV= 1/(incidence rate*VE)

Matonack A. ACIP presentation, February 2019. https://www.cdc.gov/vaccines/acip/meetings/downloads/slides-2019-02/Pneumococcal-5-Matanock-508.pdf.

a Unpublished ABCs data [3]

b Bonten [1]*

^cGierke [11], estimated by applying the %PCV13-type IPD to the NIPP incidence estimate

d Swerdlow [10]*

e Webber [6]*

^fNelson et al. 2008, estimated as 5.1% of all-cause outpatient pneumonia is PCV13-type

While reasonable to remove the recommendation to vaccinate with PCV13, what does that mean for reimbursement, etc.?

In conclusion, tremendous advances in science have given us new vaccines

Which one of the following is NOT a method of developing a currently licensed vaccine?

- A) use entire genome's open reading frames to synthesize proteins, inject them in a rodent to test immunogenicity, find one that evokes a response, then purify it.
- B) Make envelope protein from pentamers, get it to self-assemble to a larger particle.
- C) Get a harmless recombinant insect virus to infect a moth ovary and then secrete a target protein.
- D) Get a target protein, add HbSag to it, then also add AS01 adjuvant.
- E) Take a part of a polysaccharide capsule, and connect it to tetanus toxoid.

The End

