

The Changing Face of Pertussis

Colin D Marchant MD

**Boston University School of Medicine
Boston Medical Center**

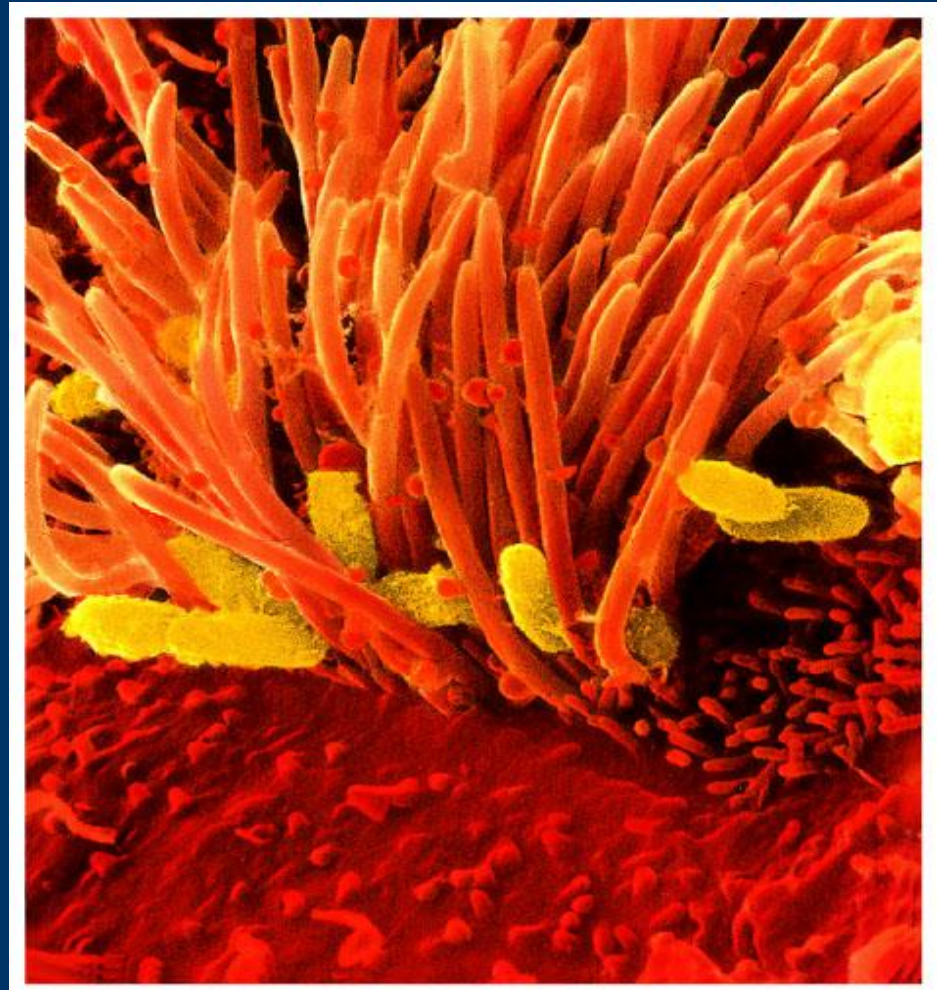
**Tufts University and Tufts Medical Center
Boston, Massachusetts**

Dr. Marchant has the following past, present or intended financial relationships:

- Employment at Boston University
- Medical Practice at: Boston Medical Center, Tufts-New England Medical Center, and Franciscan Hospital for Children
- Research support from the Centers for Disease Control
- Research support, consultant, and/or speakers bureau for the following pharmaceutical companies: Abbott, GlaxoSmithKline, Johnson & Johnson, Merck, MedImmune, Novartis, Pfizer, Replidyne, Sanofi-Pasteur,.
- No patents for drugs, vaccines or medical devices
- No direct stock ownership in any pharmaceutical or healthcare company

Bordetella pertussis – Whooping Cough

- Fastidious gram-negative coccobacillus
- Antigenic and biologically active components:
 - pertussis toxin (PT)
 - filamentous hemagglutinin (FHA)
 - agglutinogens
 - adenylate cyclase
 - pertactin
 - tracheal cytotoxin
- Reservoir: humans only
- Transmission: via aerosols, highly contagious



Morbidity of Pertussis in Adolescents

- Coughing 100%
- Paroxysms of cough 74%
- Vomiting 56%
- Weight loss 33%
- Problems sleeping 77%

- Still coughing at 108 days 38%
- Missed school 83%
 - Mean number of missed days: 5.5
 - Range: 0.4 to 32 days

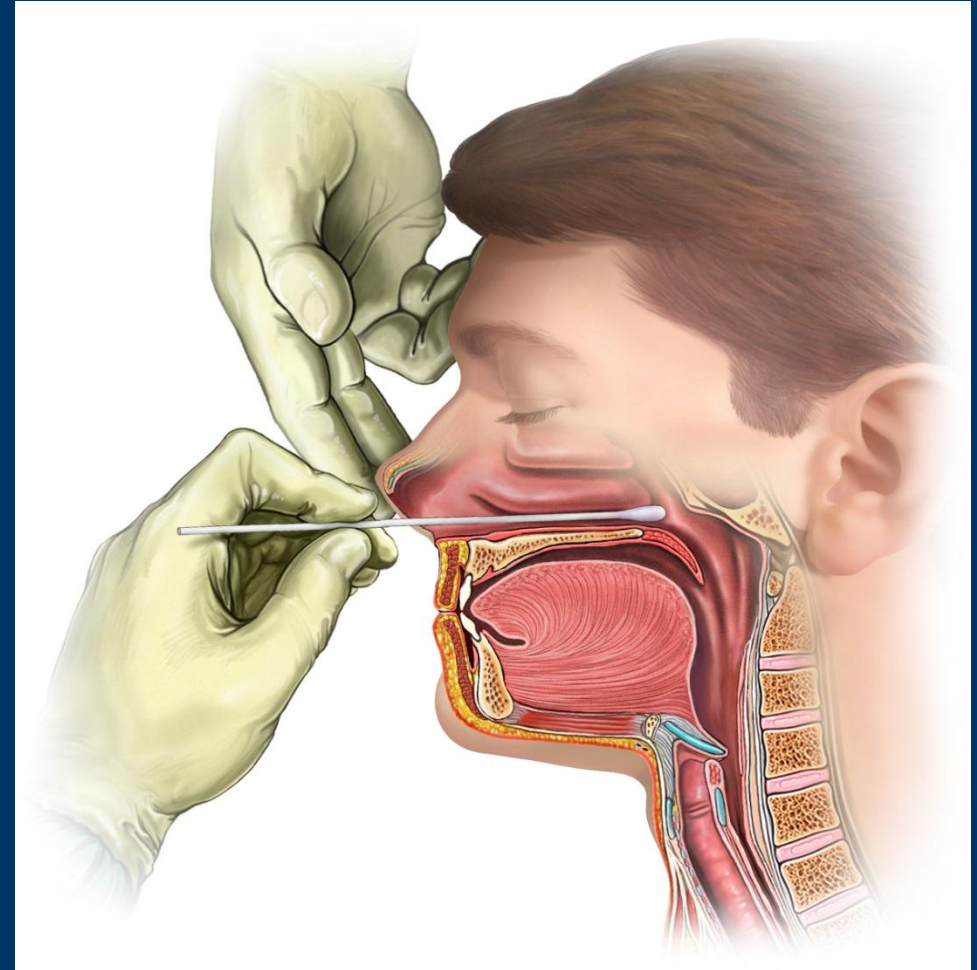
Morbidity of Pertussis in Adults

- Coughing 100%
- Paroxysms of cough 84%
- Vomiting 54%
- Weight loss 33%
- Problems sleeping 84%

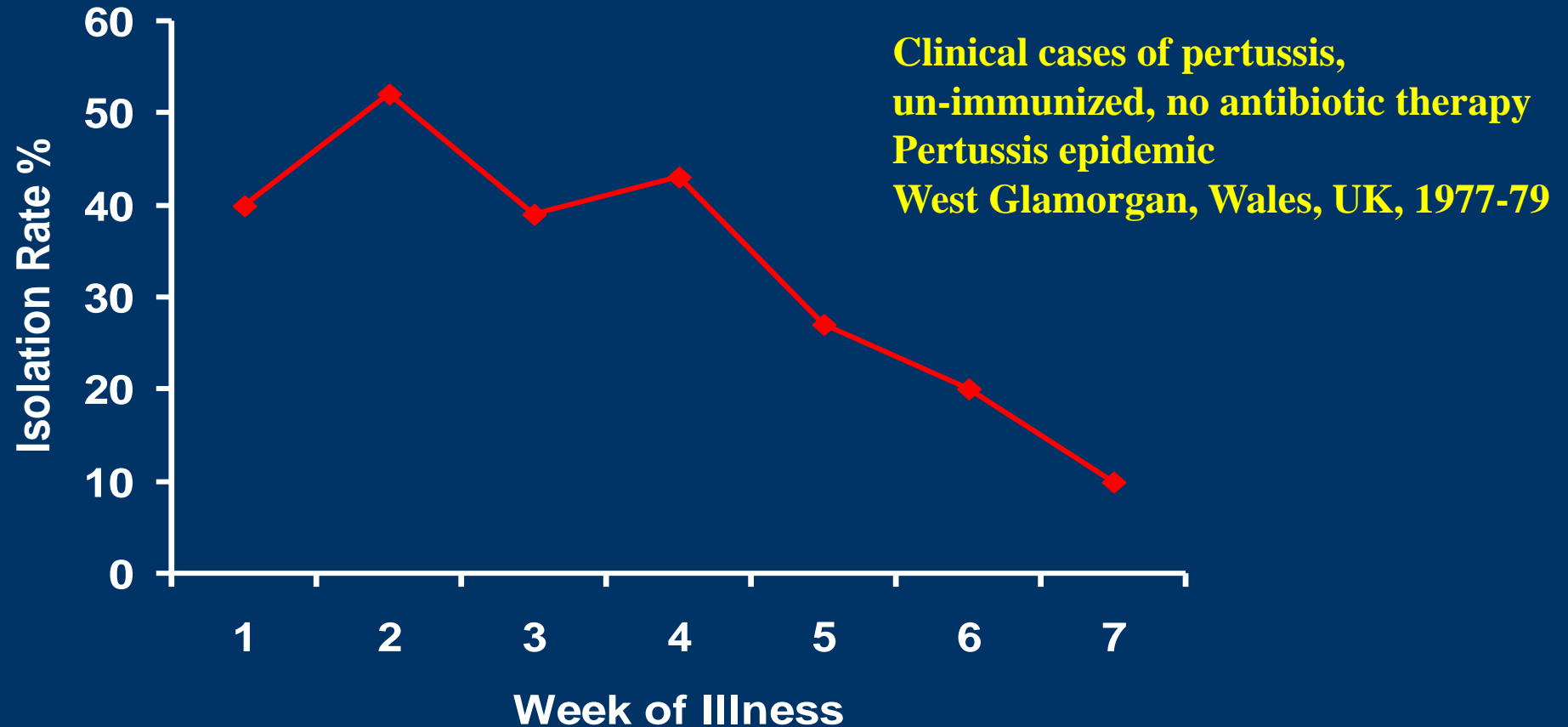
- Still coughing at 94 days 61%
- Missed work 61%
 - Mean number of missed days: 9.8
 - Range: 0.1 to 180 days

Diagnostic Tests for Pertussis

- Nasopharyngeal (NP) culture on special media (Regan-Lowe, Bordet-Gengou)
- Polymerase chain reaction (PCR)
- Serologic tests
- Increased white blood cell (WBC) count with absolute lymphocytosis
- Direct fluorescent antibody (DFA)—variable sensitivity/specificity



Isolation of *Bordetella pertussis* by week of illness



from Kwantes et al J Hyg Camb 1983;90:149-58

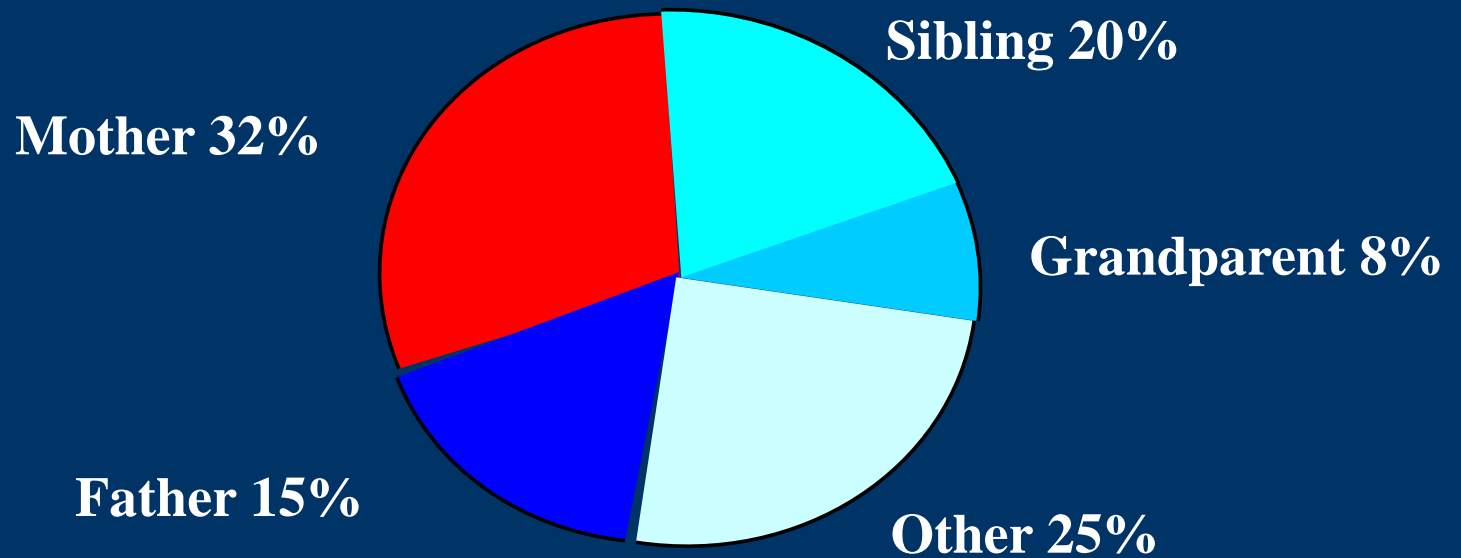
Intrafamilial spread of pertussis

- 21 families (97 individuals) of patients with whooping cough diagnosed by culture or by ELISA serology
- Follow-up (average 6 months): 83% infected (ELISA)
- 46% of secondary cases asymptomatic, mostly adults or vaccinated children.
- Unvaccinated infants had classic whooping cough
- Classic symptoms of pertussis decreased with age; atypical pertussis was usually culture negative (diagnosed by ELISA)

Mertsola J, et al J Pediatr. 1983 ;103:359-63;
see also Long SS et al J Infect Diis 1990; 161:480-86

Infant Pertussis: Who Is the Source?

- 774 infant cases from 4 states (GA, IL, MA, and MN)
 - 616 families interviewed
- 264 cases had known or suspected source identified
- Sources:

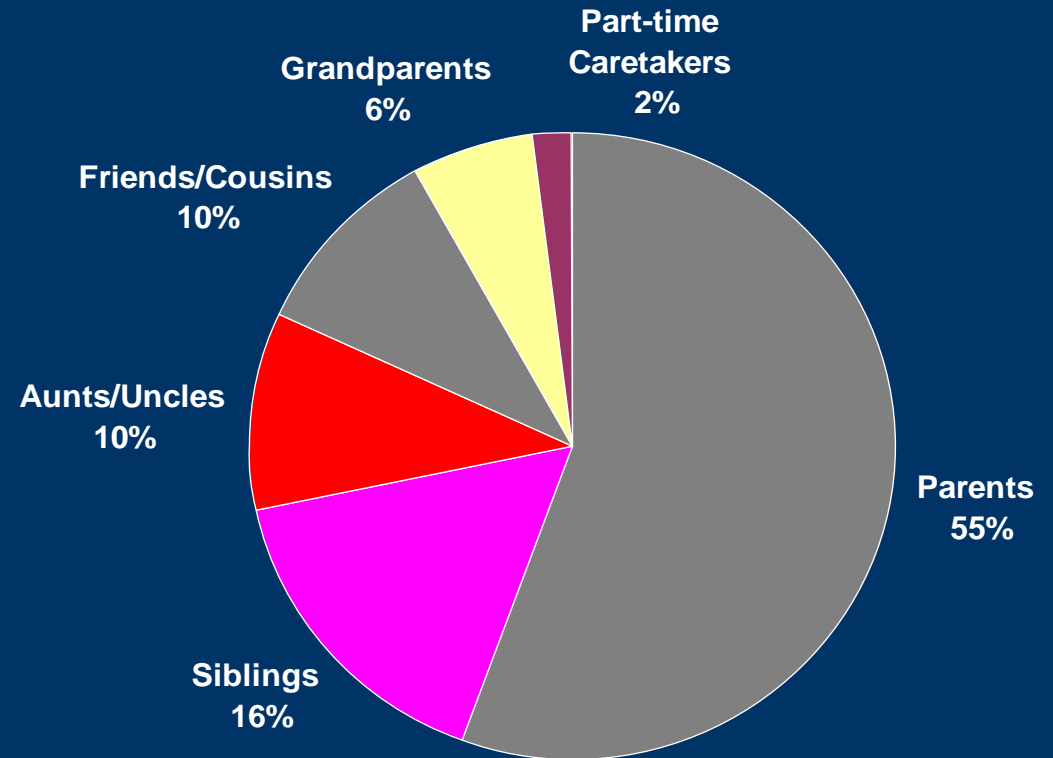


Reference:

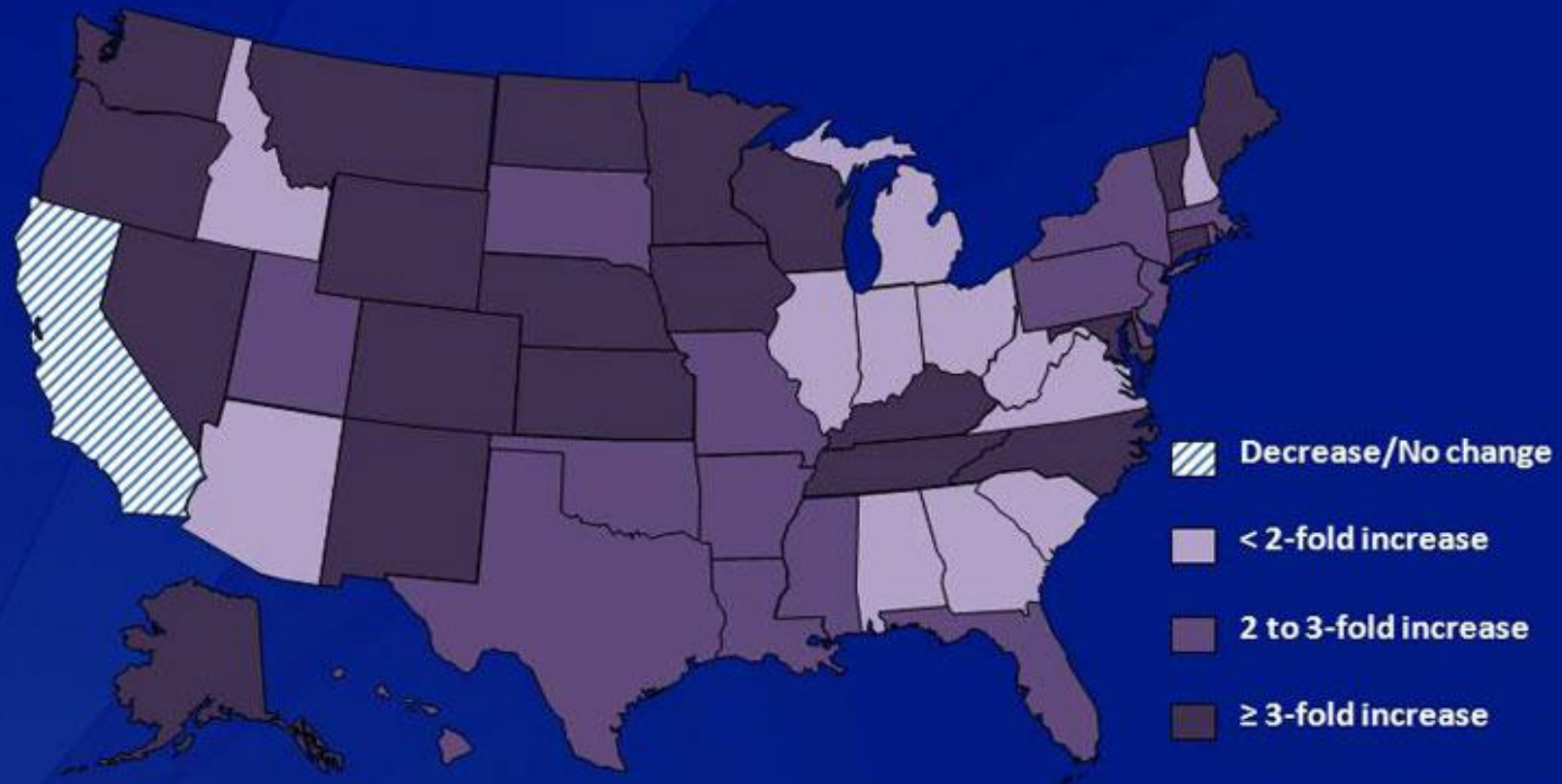
1. Bisgard KM, et al. *Pediatr Infect Dis J.* 2004;23:985-989.

Infant Pertussis: Who Is the Source?

- Multicenter study in France, Germany, Canada, and the United States
- 95 infants ≤ 6 months of age with laboratory-confirmed pertussis and 404 household and close non-household contacts
- Household members responsible for 76%-83% of transmission to infants



Changes in Pertussis Reporting by State from 2011 to 2012* †



*Data for 2012 are provisional.

†Cases reported through Week 52 in 2011 were compared with cases reported through Week 52 in 2012; fold-changes were calculated for each state.



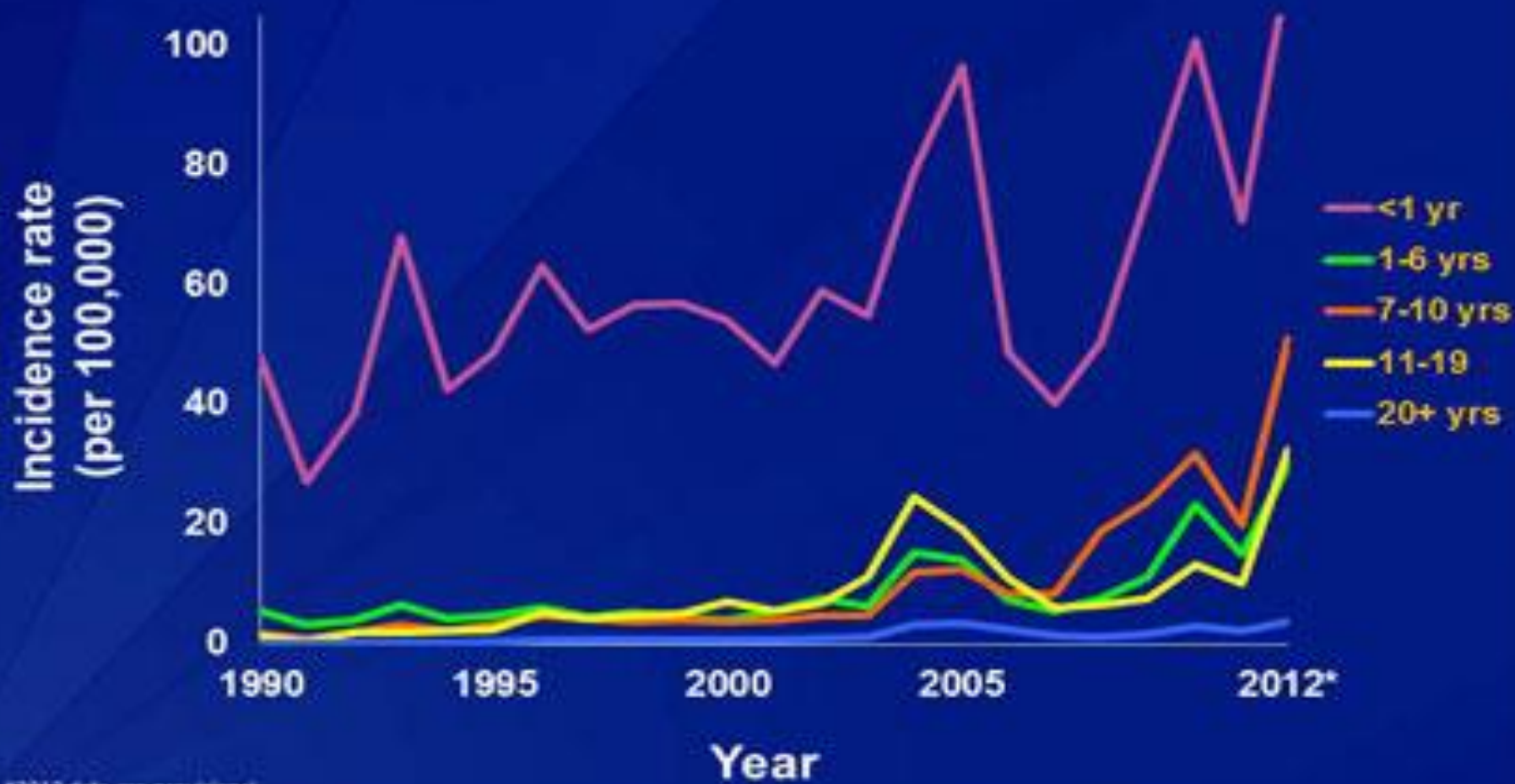
Reported NNDSS pertussis cases: 1922-2012*



*2012 data are provisional.

SOURCE: CDC, National Notifiable Diseases Surveillance System and Supplemental Pertussis Surveillance System and 1922-1949, passive reports to the Public Health Service

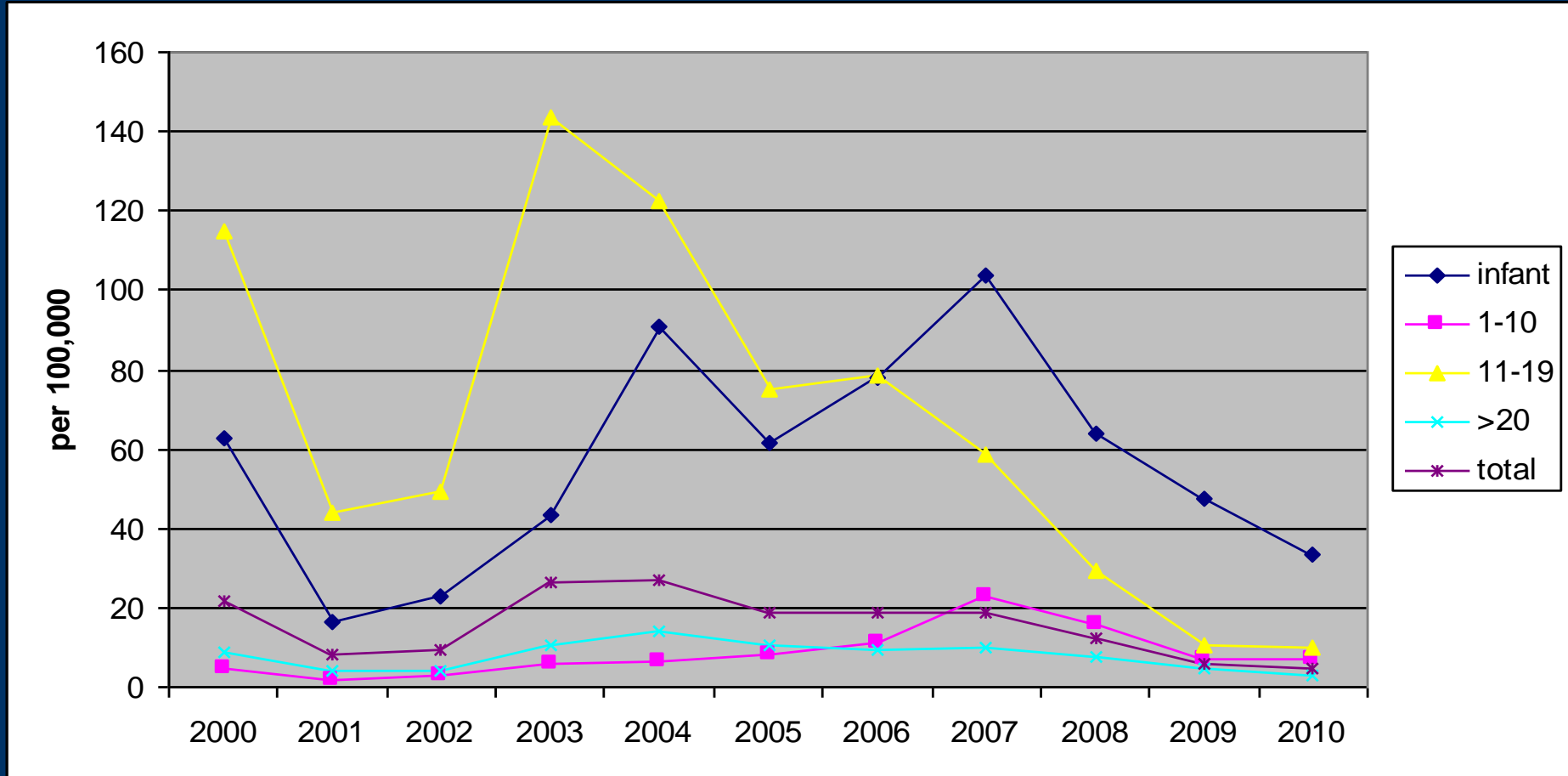
Reported pertussis incidence by age group: 1990-2012*



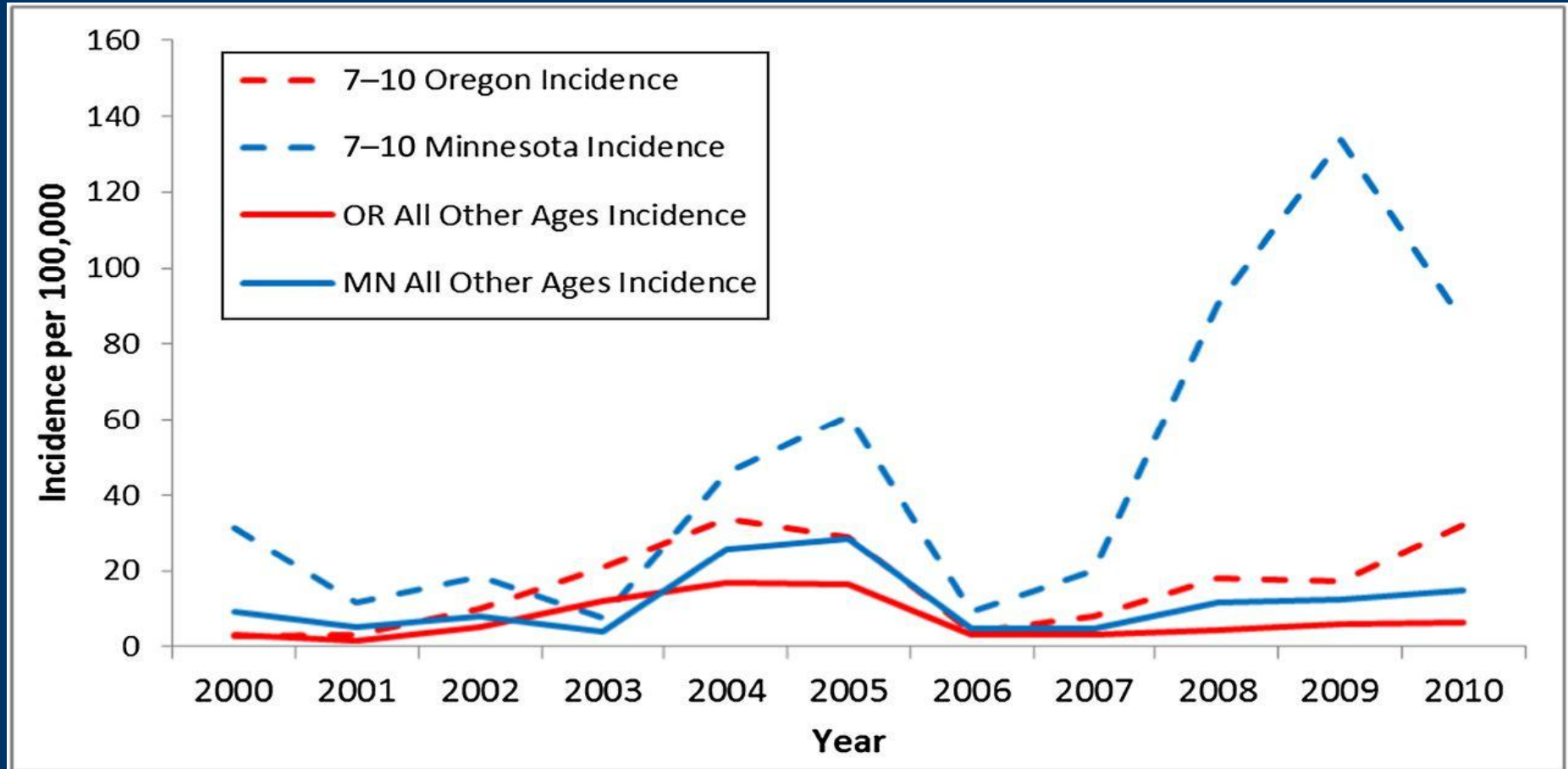
*2012 data are provisional.

SOURCE: CDC National Notifiable Diseases Surveillance System and Supplemental Pertussis Surveillance System.

Pertussis Incidence by Age Massachusetts 2000 -2010

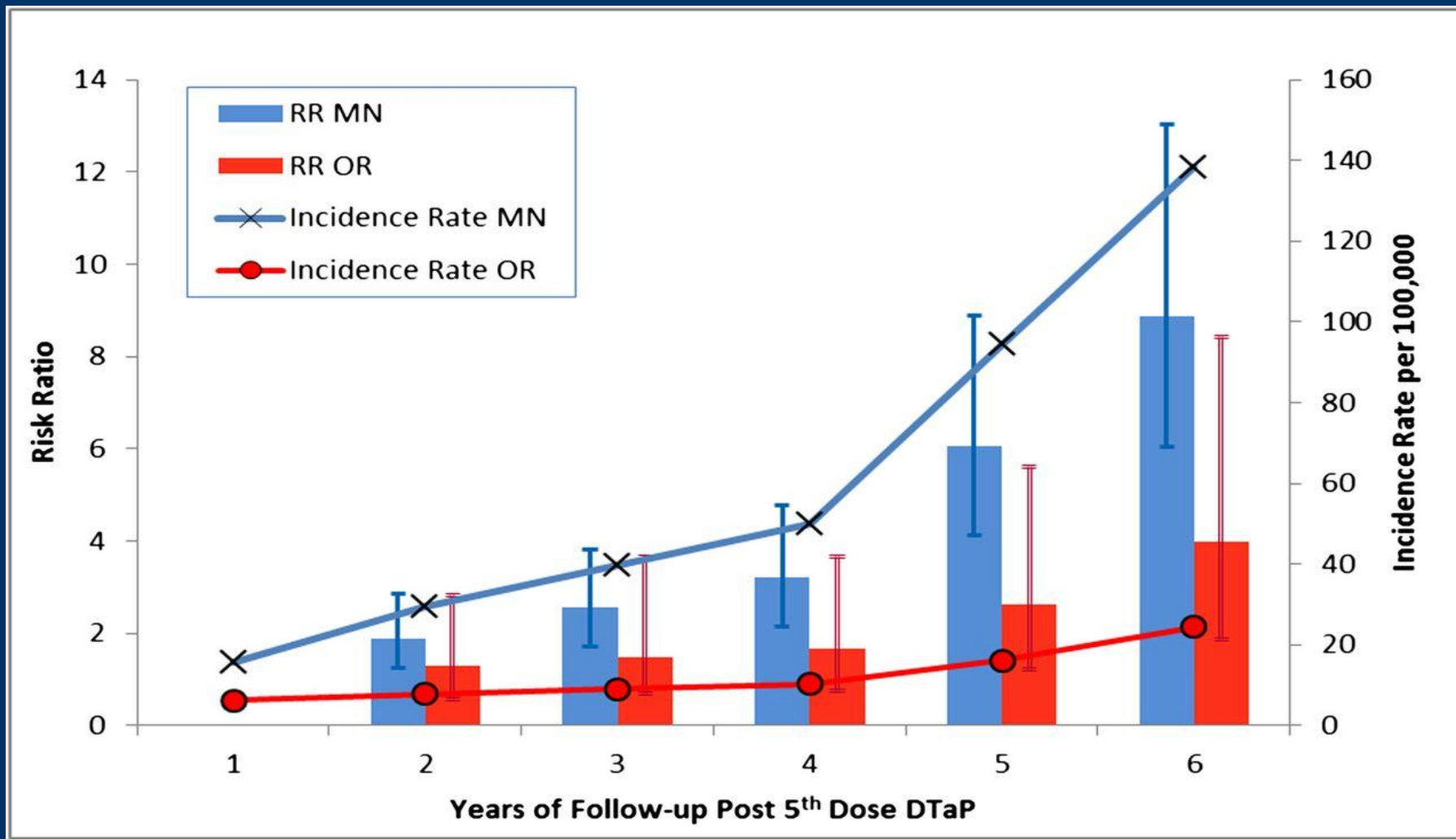


Statewide pertussis incidence and incidence of cases in 7- to 10-year-olds, Oregon (OR) and Minnesota (MN), 2000–2010.



Tartof S Y et al. Pediatrics 2013;131:e1047-e1052

Risk ratios and incidence rates for pertussis by year of follow-up post fifth-dose DTaP, Minnesota (MN) and Oregon (OR), 2010.



Tartof S Y et al. Pediatrics 2013;131:e1047-e1052

Waning Vaccine Effectiveness after the 5th Dose of Dtap Vaccine, California 2010

Months Since 5 th Dose	Cases No.	Controls No.	Vaccine Effectiveness [95% CI]
0 doses (reference)	53	19	1 (reference)
<12	19	354	98.1% [96.1, 99.1]
12-23	51	391	95.3 % [91.2, 97.5]
24-35	79	366	92.3 % [86.6, 95.5]
36-47	108	304	87.3 % [76.2, 93.2]
48-59	141	294	82.8 % [68.7, 90.6]
≥ 60	231	288	71.2 % [45.8, 84.8]

Immunization of Adults with Tetanus-diphtheria-acellular pertussis vaccine (TdaP)

ACIP Recommendation

- All adults should receive a single dose of TdaP, followed by doses of Td every 7 to 10 years

Immunization of Pregnant Women with Tetanus-diphtheria-acellular pertussis vaccine (TdaP)

2011 ACIP Recommendations

- Reviewed safety data from manufacturers' pregnancy registries and the VAERS data: no evidence for safety problems when TdaP was given to pregnant women
- Reviewed evidence for the effect of maternal TdaP on infant immune responses to routine (2,4,6 month) DTaP vaccine: infants born to mothers who had received TdaP during pregnancy did develop immune responses to active immunization, but the responses were quantitatively lower – these differences were not judged to be important

Immunization of Pregnant Women with Tetanus-diphtheria-acellular pertussis vaccine (TdaP)

2011 ACIP Recommendation

- Immunize pregnant women who have not previously received a dose of TdaP after 20 weeks gestation

Benefits:

- Mothers will be protected from pertussis and are thus less likely to infect their infants
- Infants will receive anti-pertussis IgG antibodies from their mothers during the 3rd trimester and will likely have protection from pertussis for several months

Immunization of Pregnant Women with Tetanus-diphtheria-acellular pertussis vaccine (Tdap)

2013 ACIP Recommendation

- Immunize women with Tdap during each pregnancy
- Optimal timing for Tdap administration is between 27 and 36 weeks gestation although Tdap may be given at any time during pregnancy
- “Cocooning” still recommended: adolescents and adults (e.g., parents, siblings, grandparents, child-care providers, and health-care personnel) who anticipate having close contact with an infant aged <12 months should receive a single dose of Tdap if they have not received Tdap previously.

Tdap Immunization Rates, US, 2010

	Sample size	%	(95% CI)	Difference from 2009
Total	14,824	8.2	(7.6–8.8)	1.6
White	7,830	9.1	(8.3–9.9)	1.7
Black	2,441	7.4	(6.1–8.8)	1.6
Hispanic or Latino	3,183	4.8	(3.9–5.9)	0.1
Asian	1,058	9.2	(6.9–12.1)	4.8
Living with an infant aged <1 yr	624	10.6	(7.9–14.2)	0.3
Not living with an infant aged <1 yr	14,200	8.1	(7.5–8.7)	1.7