

Immunity to pertussis transmission and disease among household contacts of infected children (PITA study)

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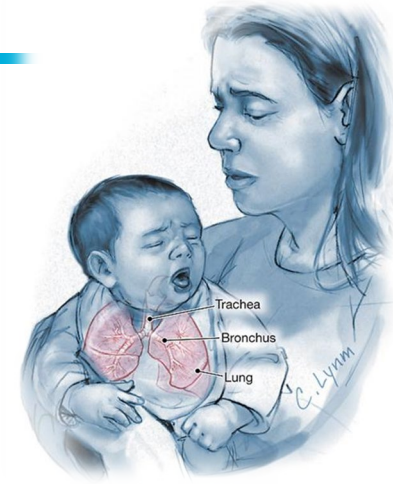
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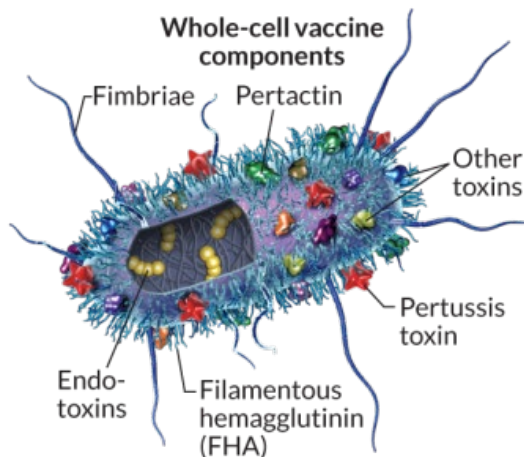
Radboudumc

Pertussis

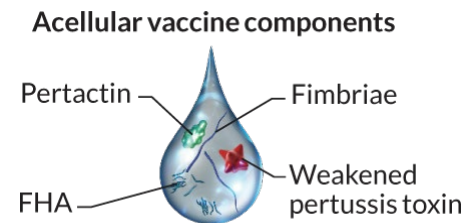
- *Bordetella pertussis*
- Uncontrollable coughing
- $R_0 \approx 10-17$
- Vaccine-preventable



jamanetwork.com

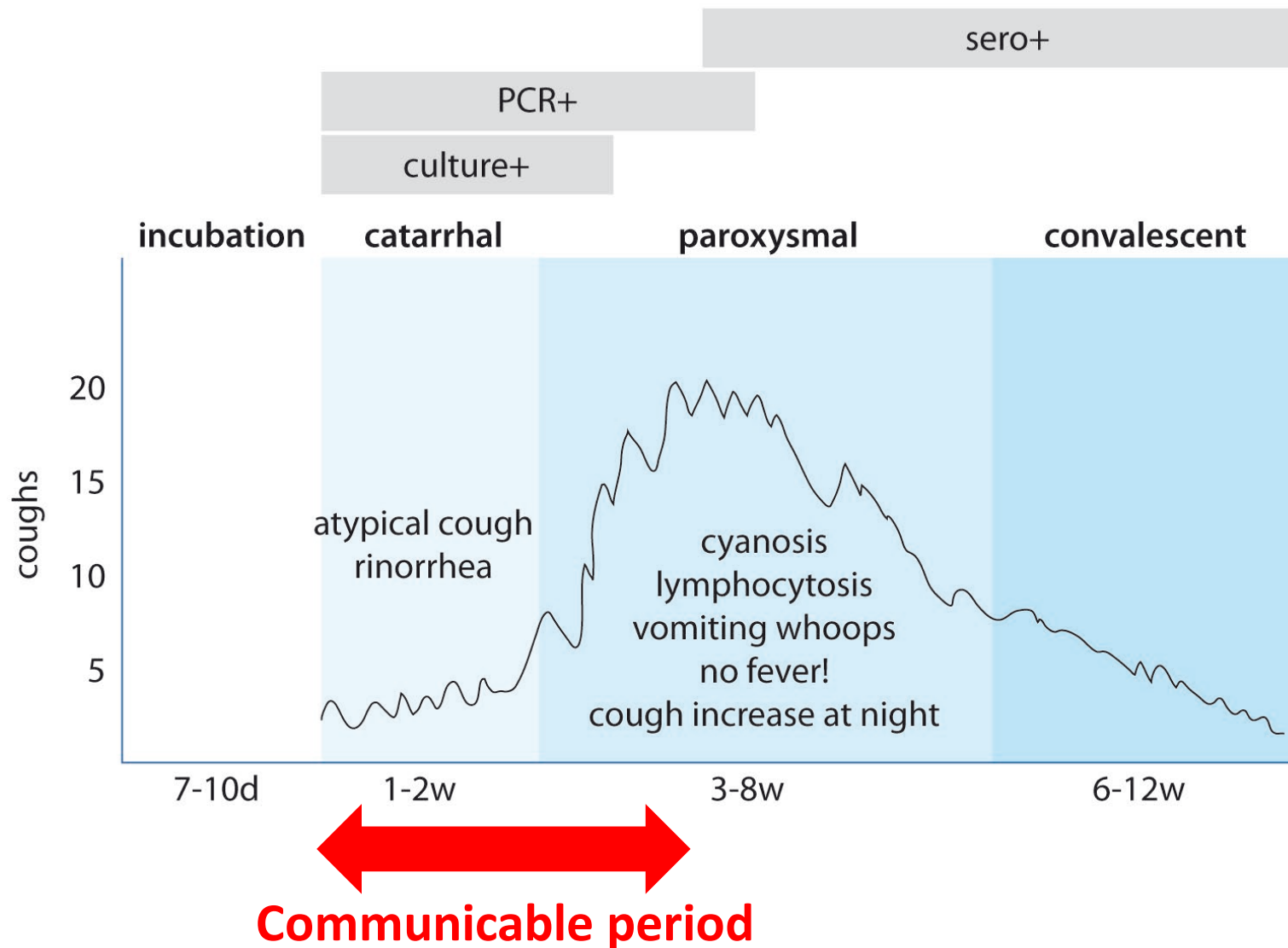


1957

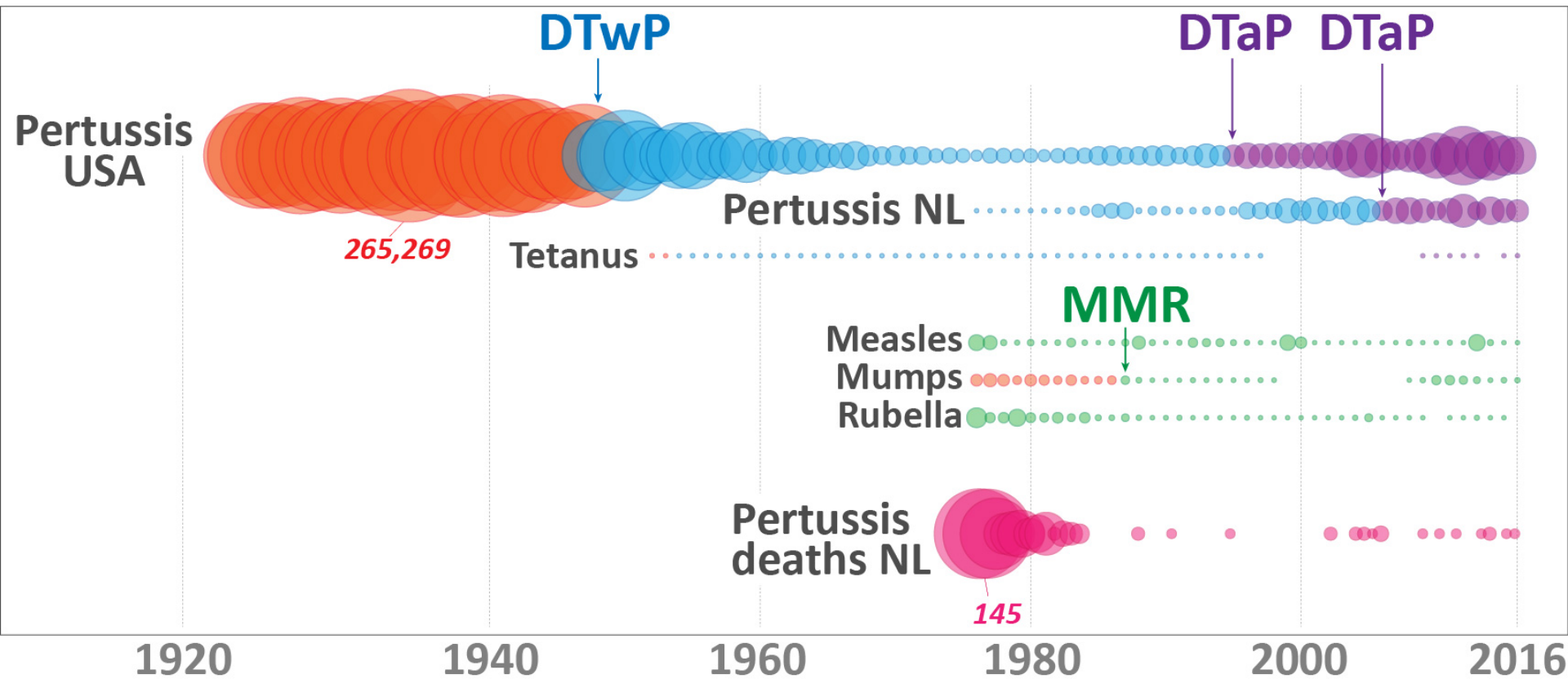


2005

Classical pertussis



Vaccination history



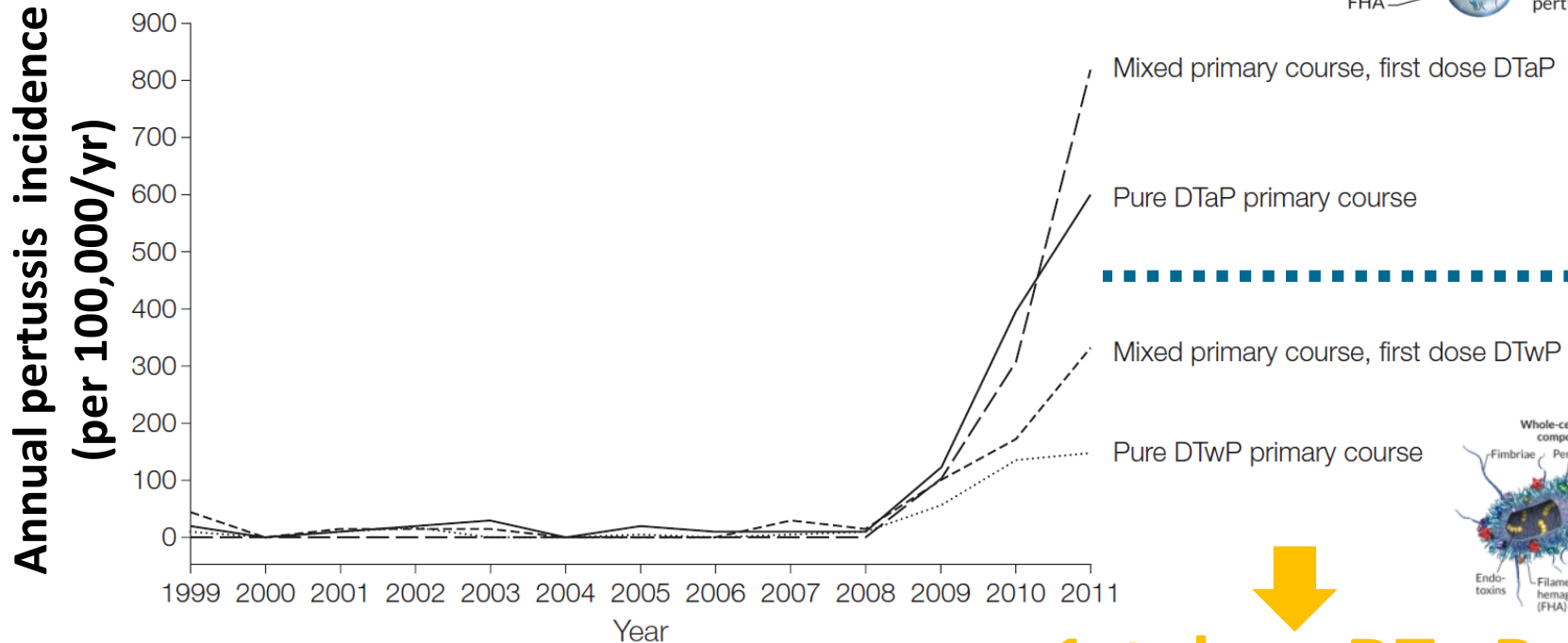
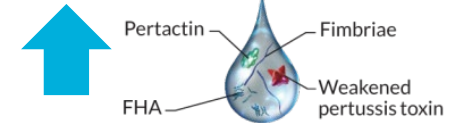
Explanations for the resurgence

1. Improved awareness and molecular diagnostics
2. Pathogen adaptation
3. End-of-honeymoon
4. Switch to acellular vaccines
 - a) Herd immunity – asymptomatic transmission
 - b) Duration of protection

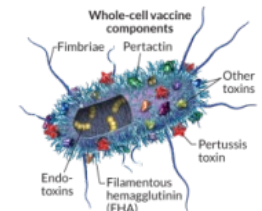
Impact of priming

1st dose DTaP

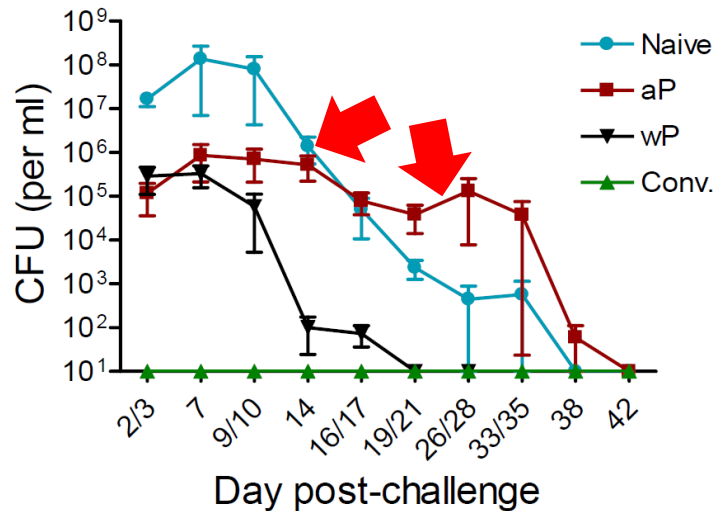
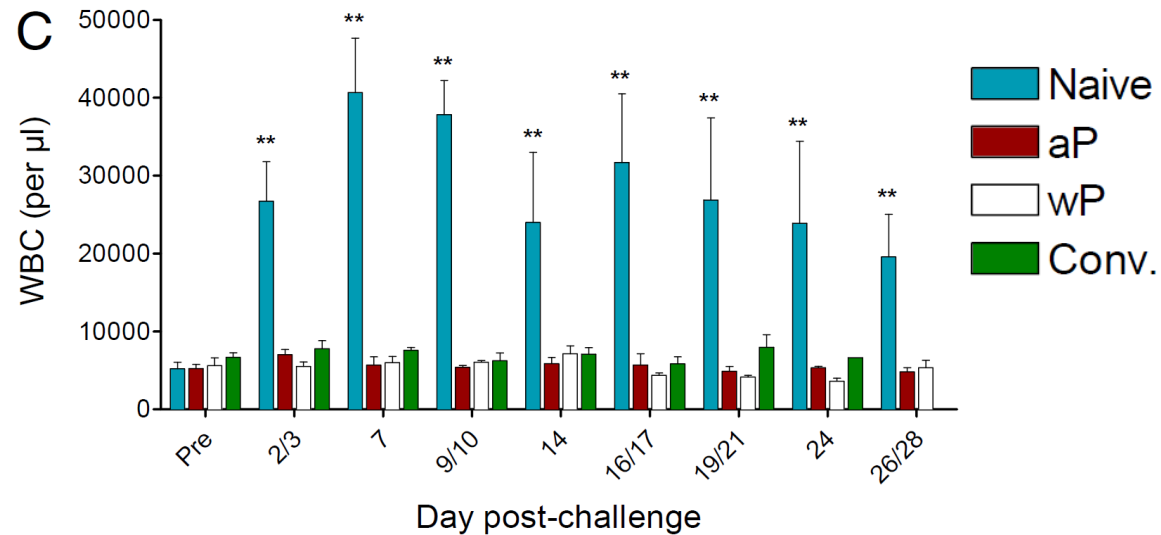
Acellular vaccine components



1st dose DTwP



Effects of priming on disease & colonisation



➡ Asymptomatic infection

Transmission of pertussis

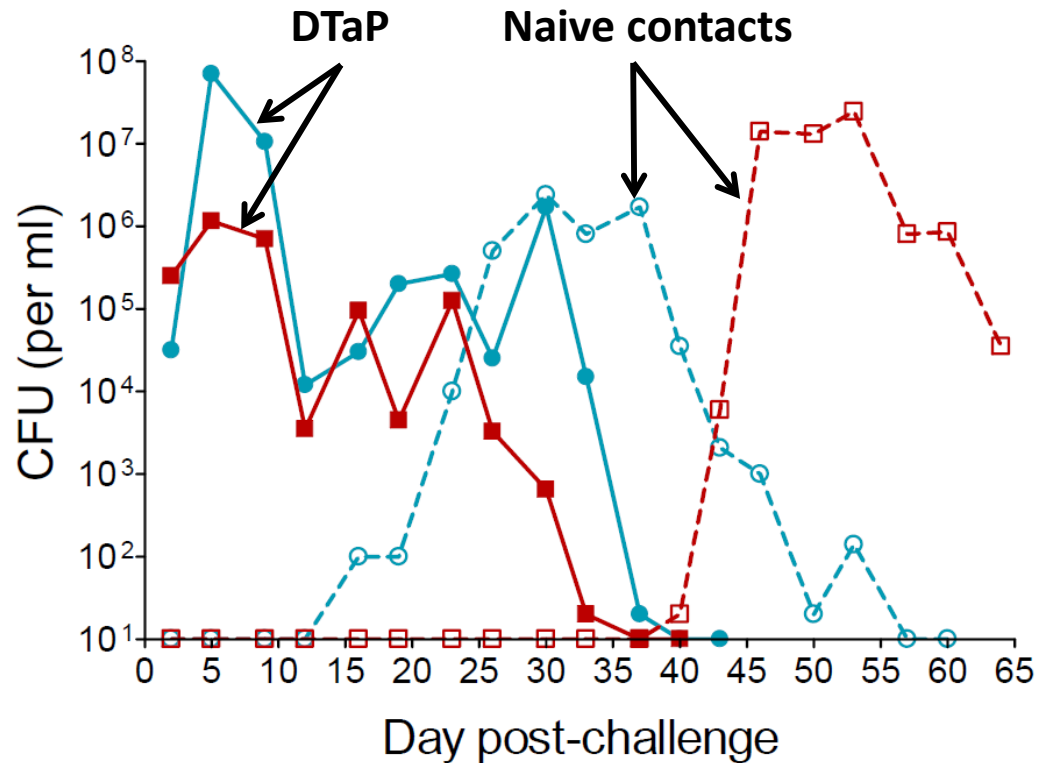
Acellular vaccine components
Pertactin Fimbriae
FHA Weakened pertussis toxin



DTaP
+ infection



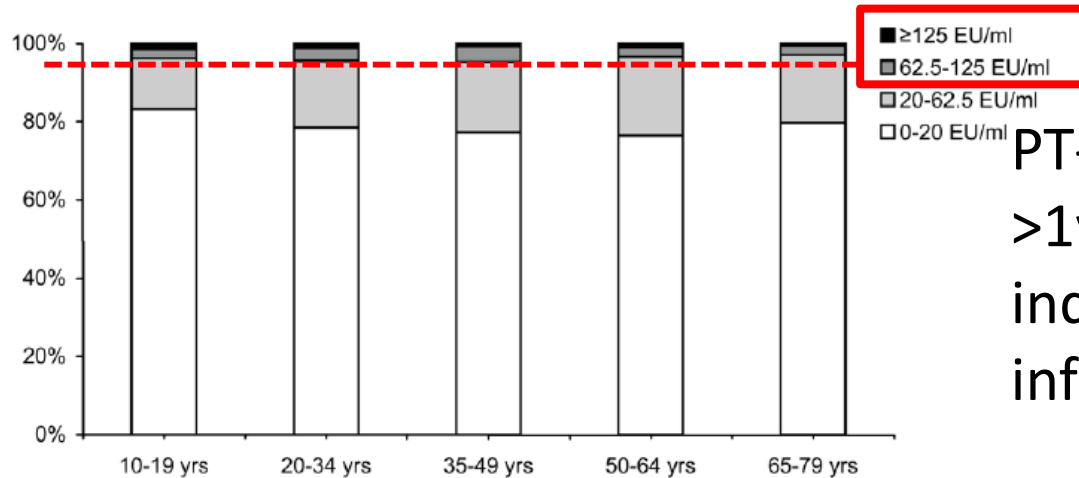
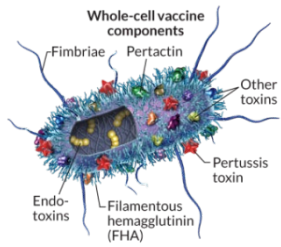
Naive
co-housed



Increased circulation in the Netherlands

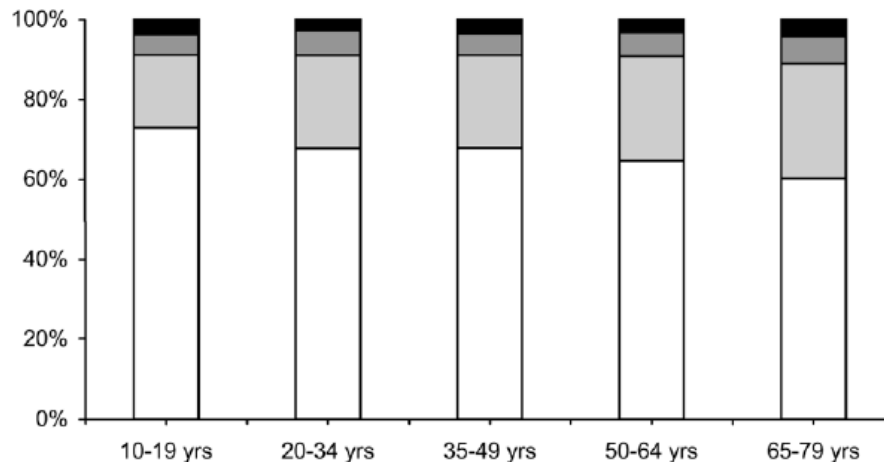
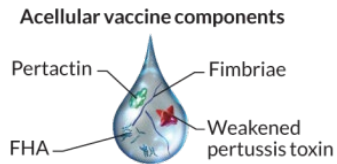
NL:

1995-1996



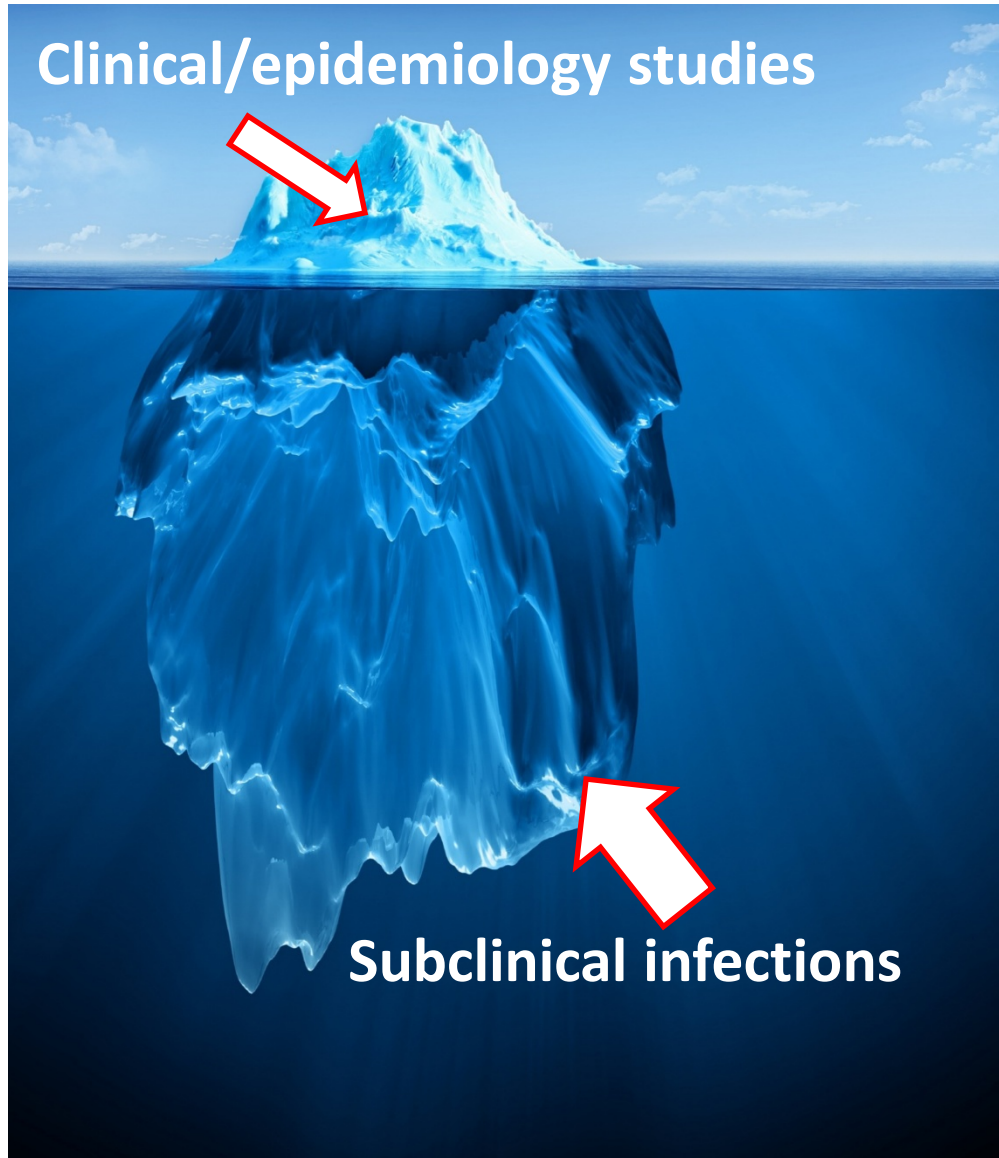
PT-IgG >62.5 &
>1y since last vax
indicates recent
infection

2006-2007



9% of >9y
exposed/infected
each year!

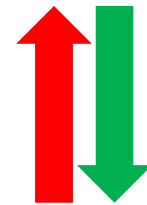
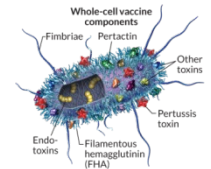
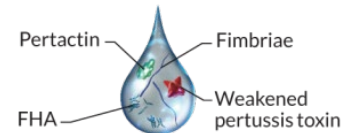
Correlates of protection?



Disease: lungs

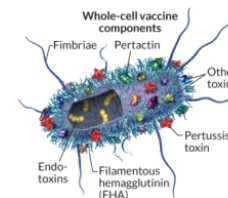
Mechanisms of protection

Acellular vaccine components



**Colonization/
transmission**

Mechanisms of protection?





What is the PERISCOPE consortium ?

Public Partners (20)

RUMC (C)	UNIBAS
UOXF	UB
PHE	LUMC
TCD	USAL
UTU	ULB
CEA	CHUV
ICL	US
RIVM	QB
IMIC	MRC
	EURICE

EFPIA Partners (2)

Sanofi-Pasteur
GlaxoSmithKline

FINANCING

BMGF	€	7M
EFPIA in-kind	€	7M
IMI	€	14M
Total	€	28M

Strategic objectives

- 01** Foster expertise and increase capacity in Europe to evaluate new pertussis vaccines in (pre)clinical models
- 02** Identify (early) biomarkers of long-lasting protective immunity to pertussis in humans. This will accelerate and de-risk clinical development of next generation pertussis vaccines
- 03** Investigate the impact of maternal vaccination on the infant response to pertussis vaccination



Start: March '16
Duration: 60m

Transmission in humans

- Pertussis is very infectious in close settings like families and in schools
- Infection rate in families: **>80%**
 - **46%** of infected family members (vaccinated during childhood) **remain asymptomatic**
 - Attack rates highest in mothers and sibling 9-13y old

PITA study

Objective

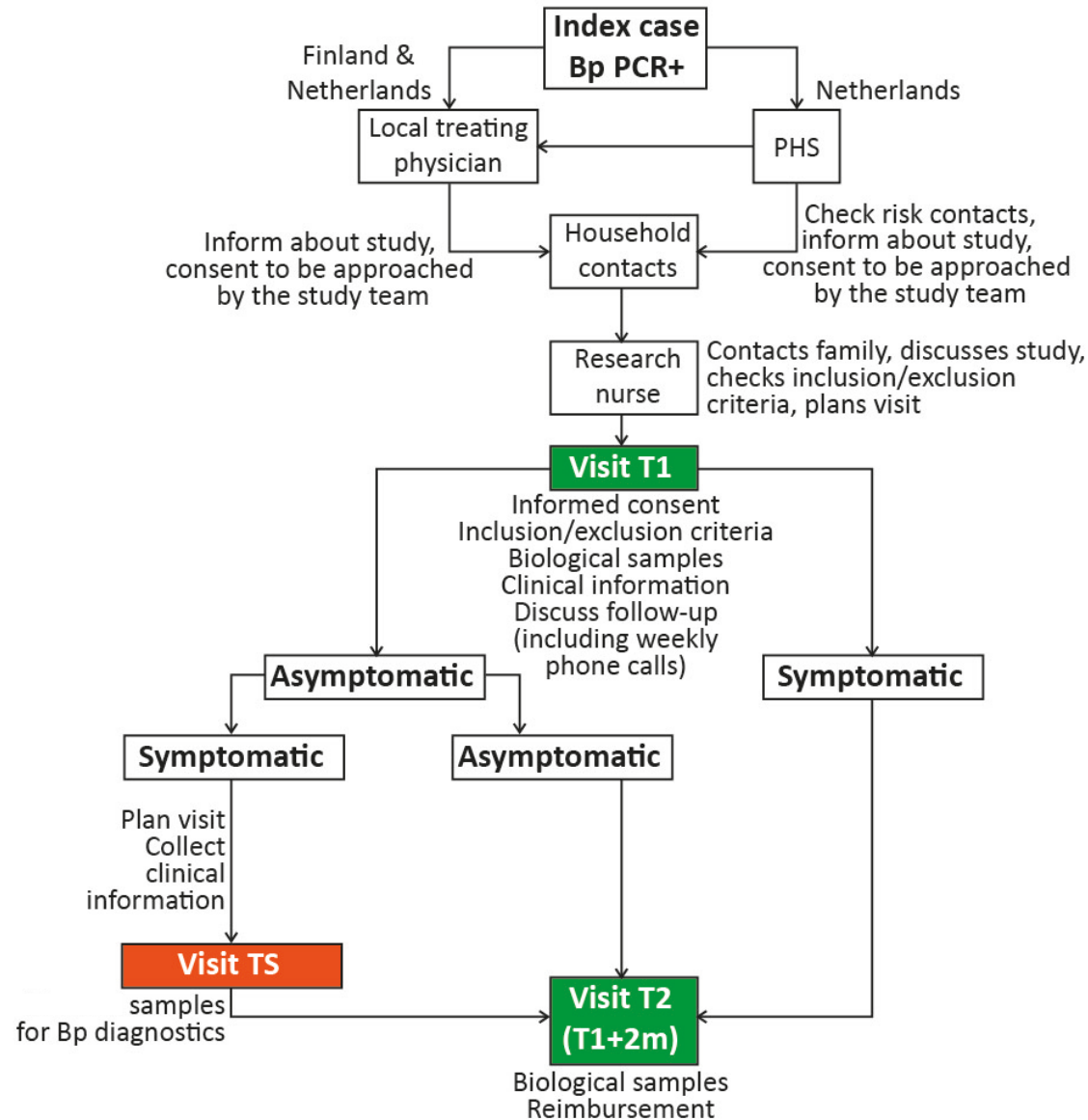
To study the **spread of *B. pertussis*** and the **development of symptoms** in relation to **immunological biomarkers** in a family setting with high potential exposure to *B. pertussis*.

Study design

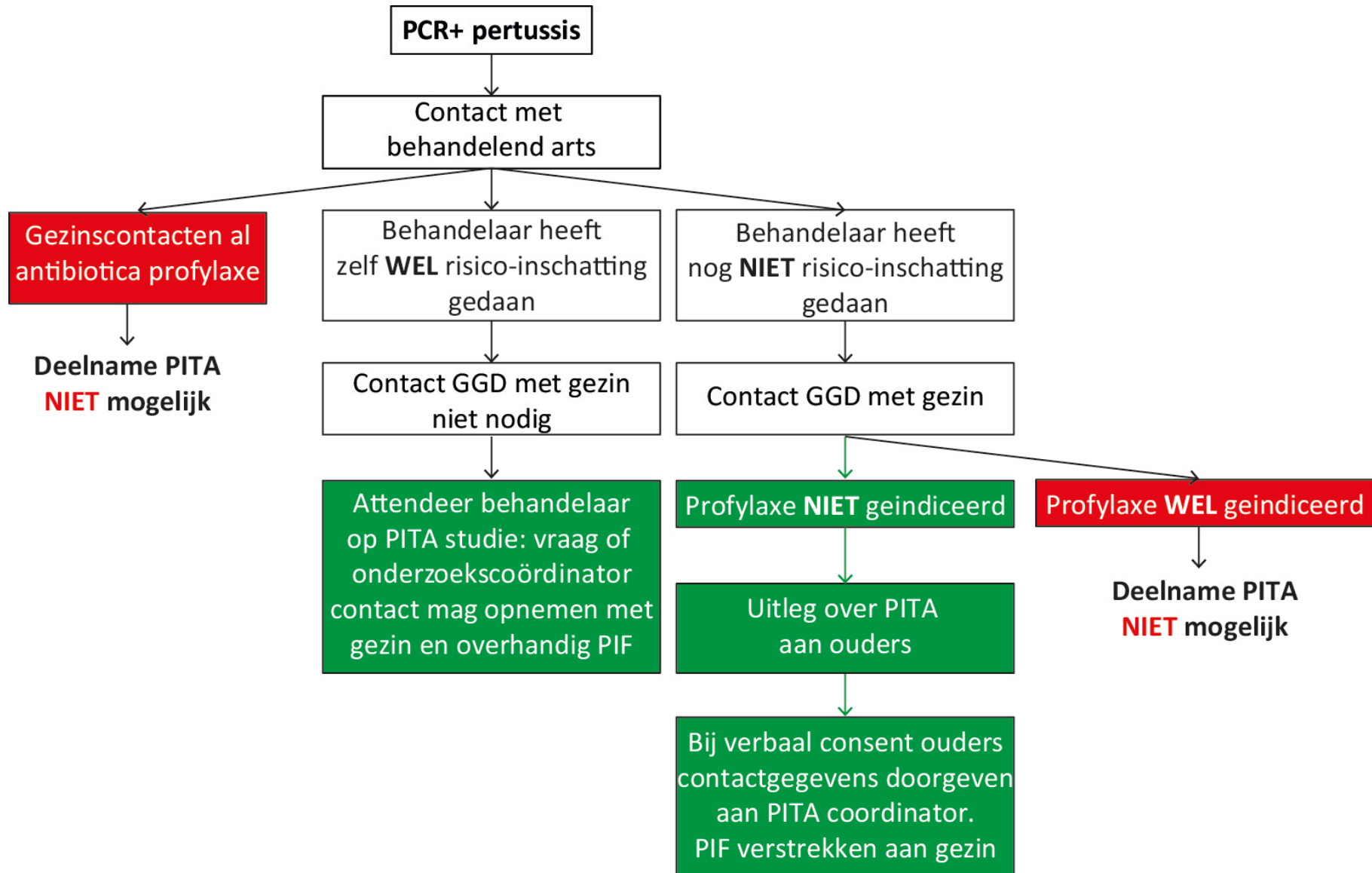
Household contact study in Finland and in the Netherlands

Study population

- **Index cases:** Children <16 diagnosed with pertussis (PCR)
- **Contacts:** All household contacts can be included (50 families in total)



Contact index-casus



Screening

Inclusion criteria:

- index case <16 years old in the family with **PCR-proven pertussis (start with <1y old index cases)**
- **≥ 2 asymptomatic household** contacts who consent to participate at T1

Exclusion criteria:

- Families with risk contacts → **indication for post exposition prophylactic treatment**
- Families for which it is **not possible to organize visit T1 within 5 days** after PHS or the treating physician has informed them about the study
- **Incapacitated household contacts**

DE PITA STUDIE

VERSPREIDING VAN KINKHOEST IN HUISELIJKE KRING

WE BESTUDEREN DE VERSPREIDING VAN KINKHOEST EN DE ONTWIKKELING VAN KINKHOEST KLACHTEN BINNEN HUISELIJKE KRING. BIJ GEZINSLEDEN BESTUDEREN WE DE AFWEER TEGEN KINKHOEST OM HET HUIDIGE VACCIN TE VERBETEREN.



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innovative
medicines
initiative



BILL & MELINDA
GATES foundation



Turun yliopisto
University of Turku



Rijksinstituut voor Volksgezondheid
en Milieu
Ministerie van Volksgezondheid,
Welzijn en Sport