

# Vaccination Guidelines

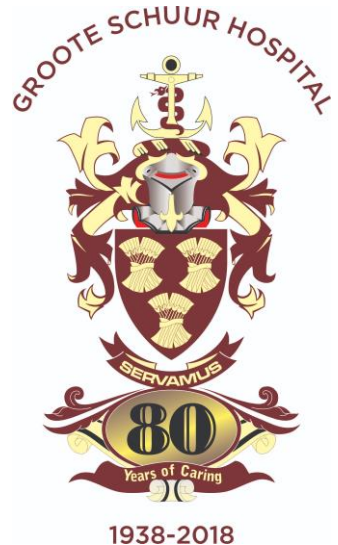
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**4<sup>th</sup> Southern African HIV Clinicians Society Conference / 24-27 October 2018**  
**Gallagher Convention Centre, Midrand**



# Calculating immunisation's return on investment

- In Gavi-supported countries, 2011–2020

**48x**  
(full income approach)

**18x**  
(cost of illness)

**3x**



Public infrastructure



**7x**



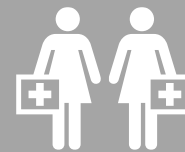
Pre-school education



**9x**



Community health workers



Immunisation



# Whole Life Approach to Immunization

- Importance of vaccination
  - Prevention to avert health spending
  - Prevention is a “best buy”
  - Vaccines seen as a solution for national & economic security
  - Dual function of vaccines

# Vaccination essential element for promoting

- Health equity
- Economic equity (reducing medical & non-medical costs)
- Social equity –access to the health care system
- Vertical equity intervention- vaccines for diseases of poverty

# WORLD IMMUNIZATION WEEK 2017 #VACCINESWORK

When **immunization rates are high**, the wider community is **protected** including:

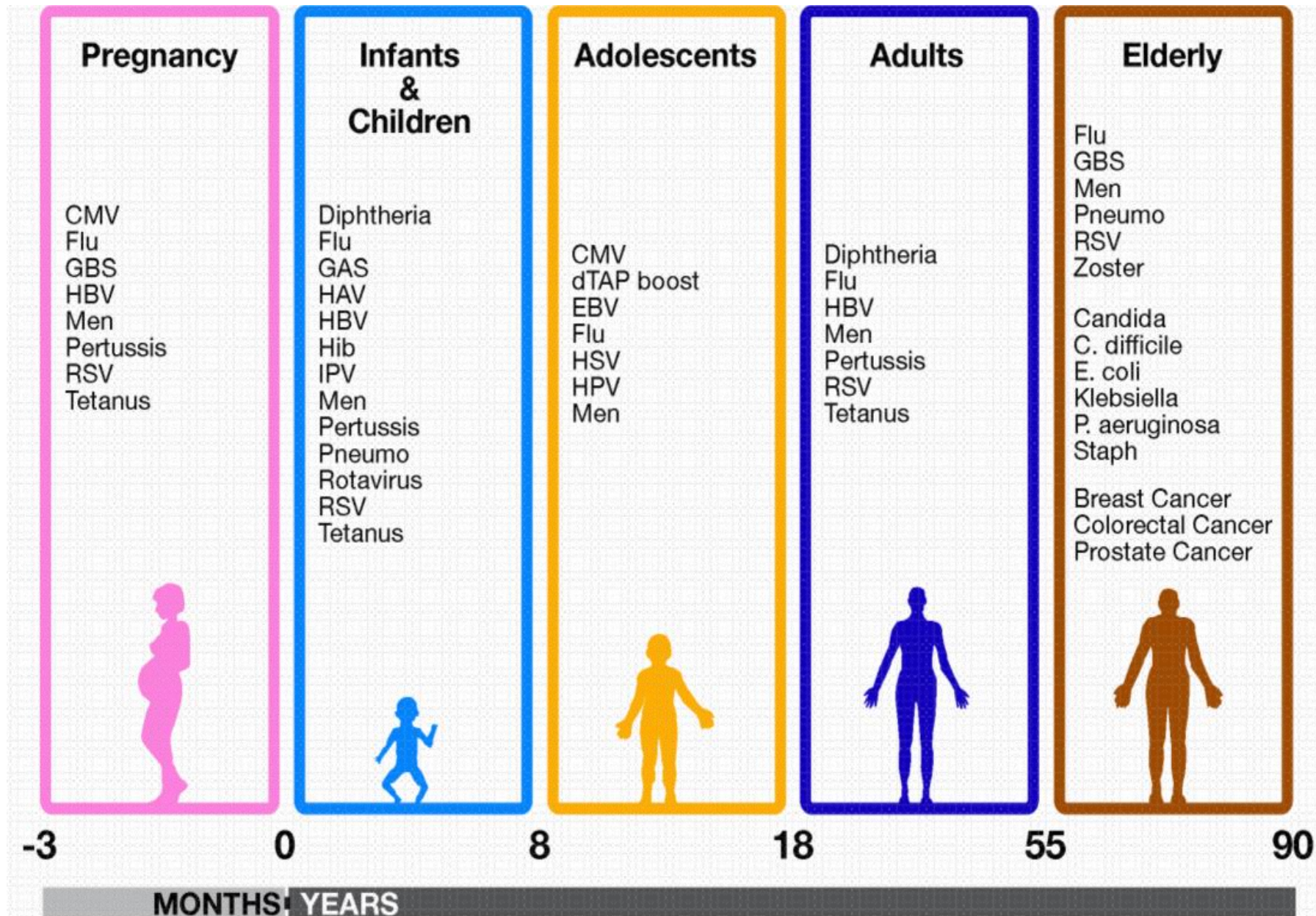
**Infants who are too young** to receive their vaccines.



**Older adults** at risk of serious diseases.

**People who take medication** that lowers their immune systems.





# Guidelines for the vaccination of HIV-infected adolescents and adults in South Africa



## Authors:









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TABLE 1: Vaccination guidelines for HIV-infected adolescents and adults.

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MMR, measles, mumps, and rubella; R, recommended; RS, recommended in selected individuals; NR, not recommended; VL, viral load; HBsAb, hepatitis B surface antibody; MSM, men who have sex with men.

# Guidelines for the vaccination of HIV-infected adolescents and adults in South Africa



- Recommendations made on the basis:
  - **Vaccines with strong local evidence for use**
    - Influenza
    - Pneumococcal vaccination
    - Hepatitis B, Tetanus-diphtheria
  - **Vaccines recommended but either local data lacking or warranted in selected cases**
    - Pertussis
    - Meningococcal, hepatitis A
  - **Vaccines with no recommendation (NR) OR recommended in selected individuals (RS)**
    - Varicella
    - Herpes Zoster
    - Measles, mumps & rubella



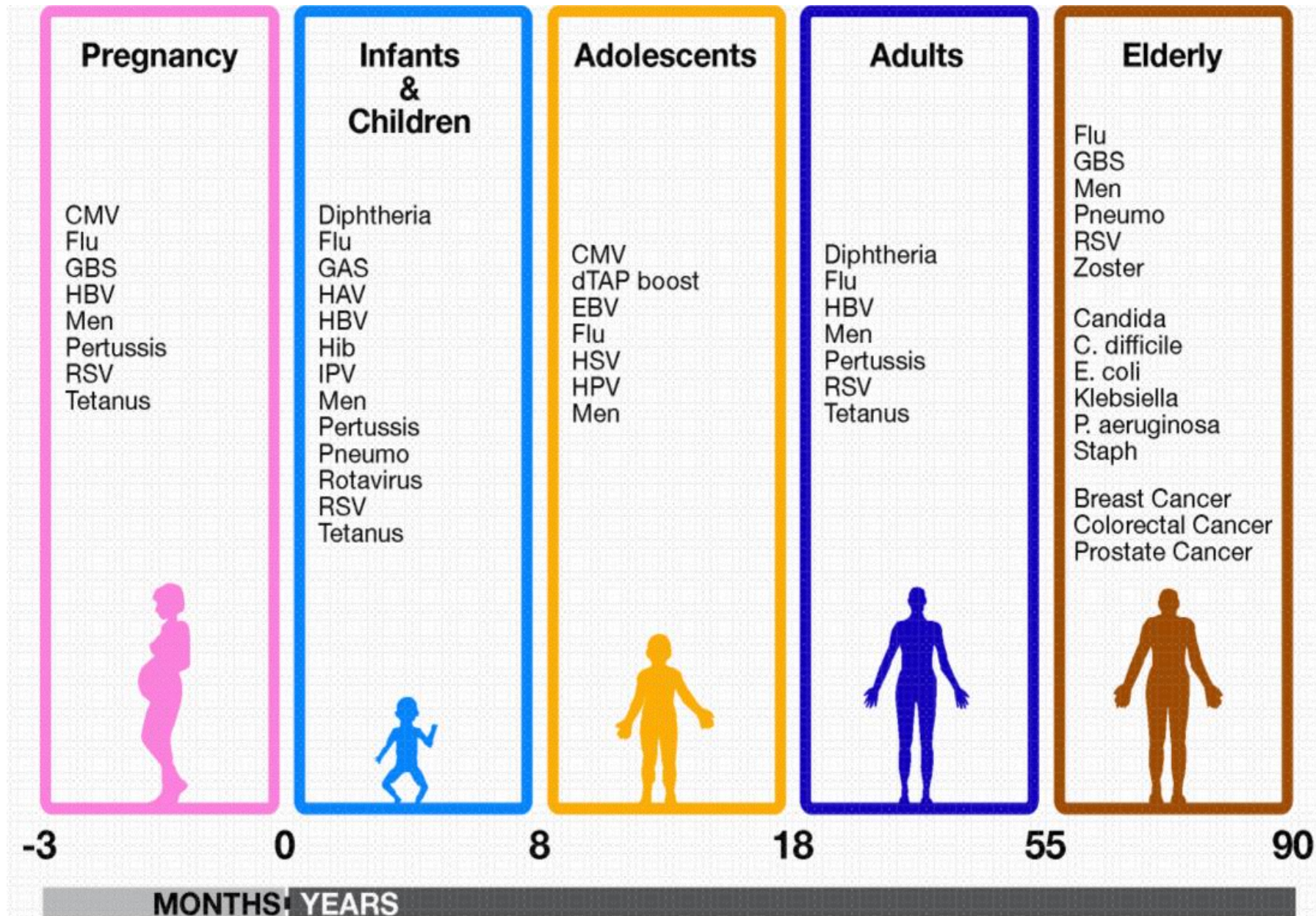
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# Ms XB

- 24 Year old Nursing student
  - HIV positive CD4<sup>+</sup> Count of 450 cells/ $\mu$ L
  - On ART for the past 3 years
    - Regimen of FDC (TDF/FTC/EFV)
    - Last viral load –LDL 6 months ago
  - 14 weeks pregnant
  - No other medical conditions & clinically well
  - **What vaccines does she require?**



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Viral Load

15 000 copies/ml



# Guidelines for the vaccination of HIV-infected adolescents and adults in South Africa



- **Pertussis**

- Emerging epidemiological data on burden of pertussis in HIV endemic countries
- Only pregnant women regardless of CD4+ count or viral load
- Recommend acellular vaccine

# Guidelines for the vaccination of HIV-infected adolescents and adults in South Africa



- **Influenza**

- 1 dose yearly
- Irrespective of CD4+ cell count, HIV viral load or pregnancy status

# NK

- 38-yr-old
- Diagnosed HIV +ve 2004
  - CD4<sup>+</sup> 11 (nadir)
  - ART initiation 03/2004
    - D4T/3TC/EFV
  - 06/2008
    - Diabetes Mellitus
  - Virologic failure 09/2010
    - Put on second line regimen
    - TDF/FTC/LPV/r
    - VL- <50 cps/mL
- Previous OI's
  - Cryptococcal Meningitis 2004
  - PTB 2011 (treated for 8/12)
- Social history
  - Employed as a senior clerk
  - Alcohol on week-ends

Do you need to vaccinate and what vaccines?

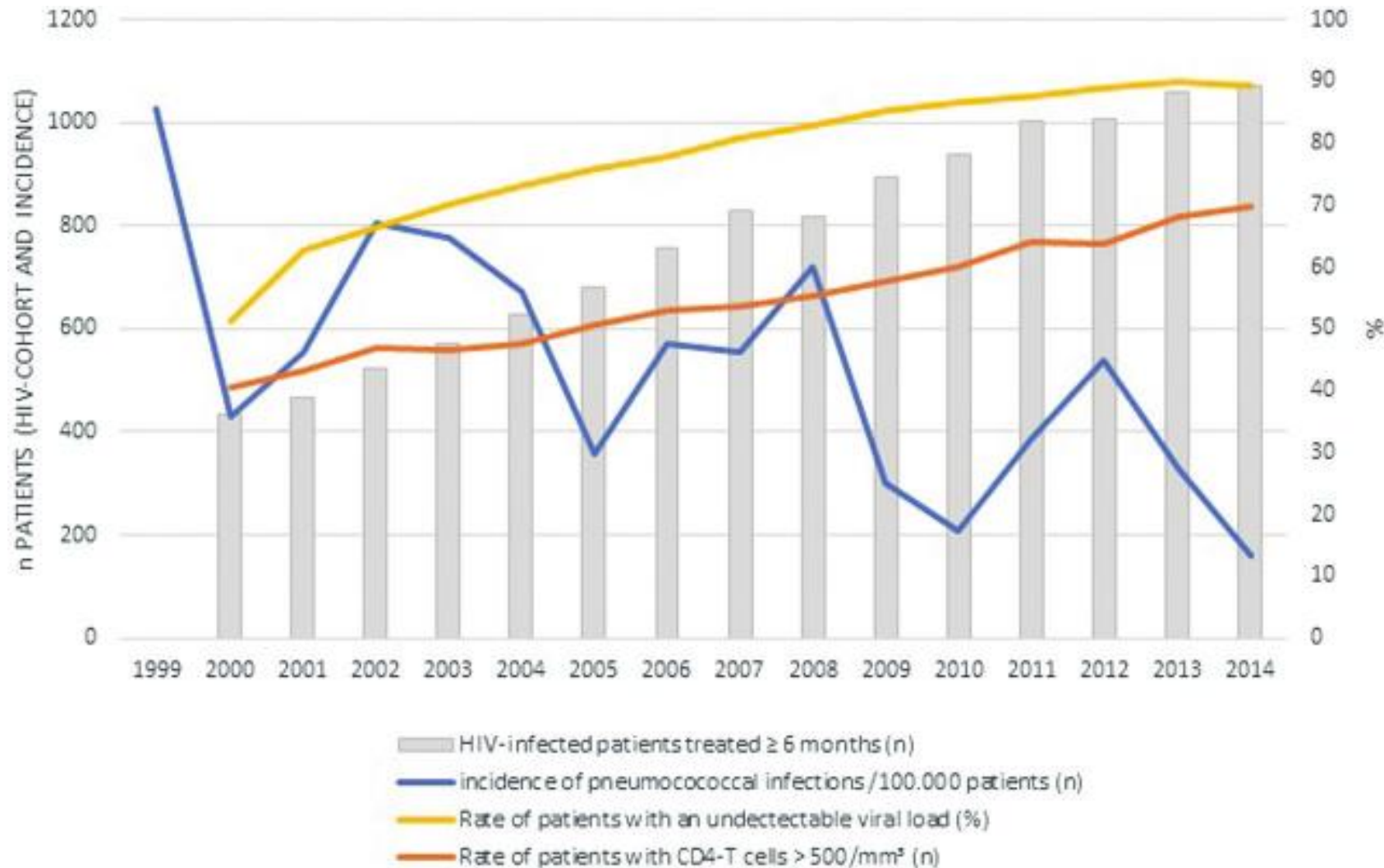
**Table 1** Factors associated with an increased risk of pneumococcal diseases

Age	Host factors		External factors	Behavioural
	Immunocompetent	Immunocompromised		
<2 years	Underlying medical conditions	HIV	Socioeconomic	Smoking
≥50 years	<ul style="list-style-type: none"> <li>• CCVD</li> <li>• CPD</li> <li>• Diabetes</li> <li>• Alcoholism</li> <li>• CLD</li> <li>• Cerebrospinal fluid leaks</li> </ul>	<ul style="list-style-type: none"> <li>CRF, nephrotic syndrome</li> <li>Cancer (solid, haematological)</li> <li>Organ and bone marrow transplant</li> <li>Auto-immune diseases</li> <li>Immunosuppressive therapy, corticosteroids</li> <li>Primary immunodeficiencies</li> <li>Functional and anatomical asplenia</li> </ul>	<ul style="list-style-type: none"> <li>Environmental</li> <li>• Preceding viral respiratory infection</li> <li>• Residence in an institution (e.g. nursing home)</li> <li>• Frequent contact with children</li> </ul>	Heavy alcohol use

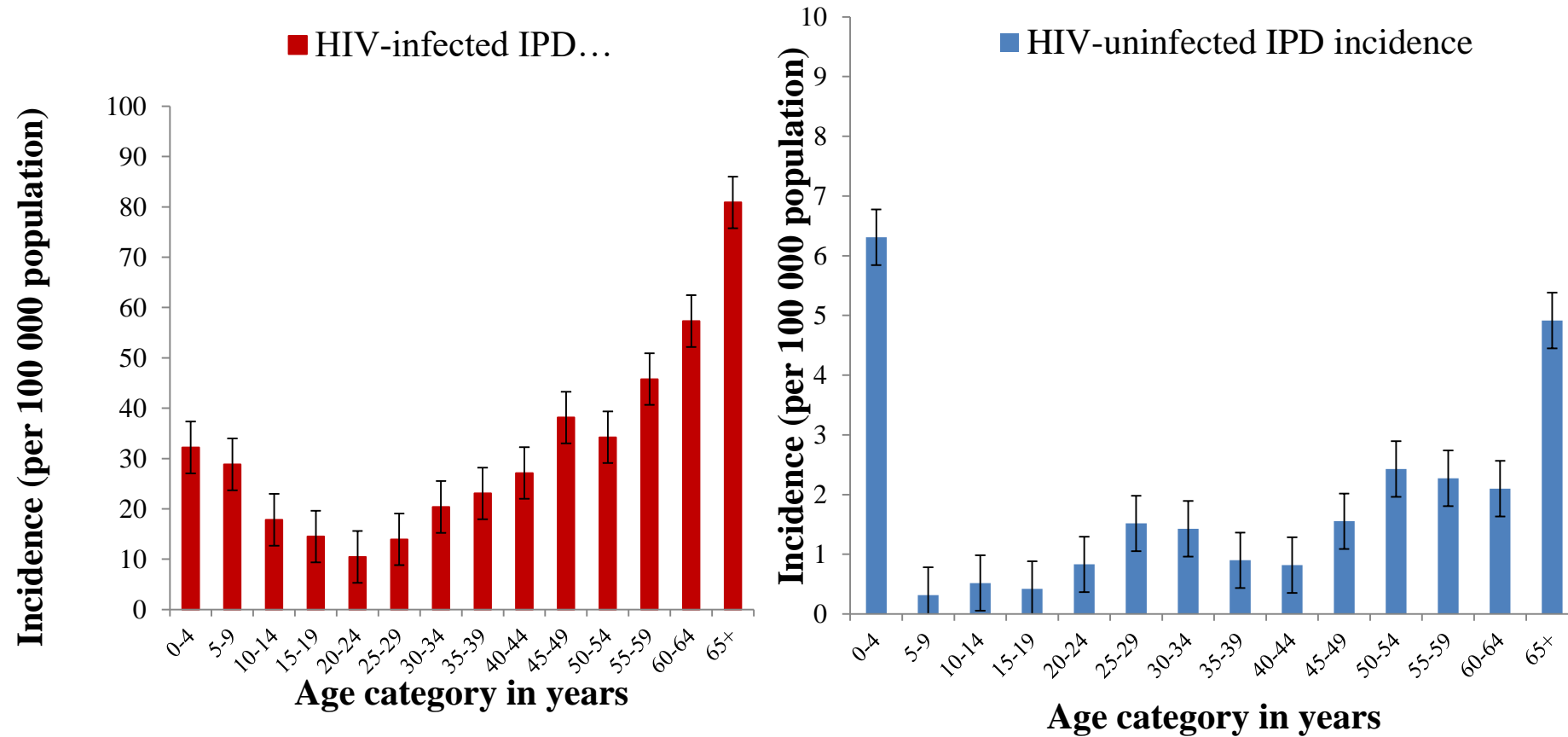
CCVD: cardiovascular and cerebrovascular disease; CPD: chronic pulmonary disease, CLD: chronic liver disease; CRF: chronic renal failure; HIV: human immunodeficiency virus



# Antiretroviral Therapy as Prevention of ... Pneumococcal Infections?



## Estimated incidence of invasive pneumococcal disease amongst HIV-infected and -uninfected persons by age category, South Africa, 2017



GERMS-SA, unpublished data

Slide courtesy – Susan Meiring

# Guidelines for the vaccination of HIV-infected adolescents and adults in South Africa



- **Pneumococcal**

- All HIV-infected regardless of CD4+ with suppressed viral load
- Prime-boost approach
- PCV13 followed by PPV23 eight weeks later
- PCV13 alone is sufficient

# DPH

- 21-yr-old female
  - HIV positive current CD4+ 218 on FDC (TFE)
  - Previously treated Disseminated Kaposi Sarcoma
    - Pulmonary involvement
- previously vaccinated with polysaccharide vaccine
- What vaccines are necessary?



# Immunogenicity and Safety of 13-Valent Pneumococcal Conjugate Vaccine in HIV-Infected Adults Previously Vaccinated with Pneumococcal Polysaccharide Vaccine

Marshall J. Glesby,<sup>1</sup> Wendy Watson,<sup>3</sup> Cynthia Brinson,<sup>4</sup> Richard N. Greenberg,<sup>5</sup> Jacob P. Lalezari,<sup>6</sup> Daniel Skiest,<sup>7</sup> Vani Sundaraiyer,<sup>8</sup> Robert Natuk,<sup>2</sup> Alejandra Gurtman,<sup>2</sup> Daniel A. Scott,<sup>2</sup> Emilio A. Emini,<sup>2</sup> William C. Gruber,<sup>2</sup> and Beate Schmoele-Thoma<sup>9</sup> *The Journal of Infectious Diseases*® 2015;212:18–27

- Immunogenicity studies with single-dose or combination regimens of PCV7 and PPSV23 in HIV – yielded variable results
- Open-label, single-arm study
  - Safety & immunogenicity of 3 doses of PCV13
  - Given at 6 monthly intervals
- **Conclusion:**
  - Vaccination with PCV13 induces anticapsular immunoglobulin G and opsonophagocytic anti-body responses in HIV-infected adults with prior PPSV23 vaccination and CD4 cell count  $\geq 200$  cells/mm<sup>3</sup>. The observations support the use of PCV13 in this population.

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- **Pneumococcal**

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# South African Guidelines for Pneumococcal Vaccination



[www.sahivsoc.org](http://www.sahivsoc.org)

New guidelines released May 2018



[www.pulmonology.co.za](http://www.pulmonology.co.za)

A updated guideline released in 2017

Guidelines for the vaccination of HIV-infected adolescents and adults  
In South Africa presented to the National Advisory Group on Immunisation  
(NAGI) of South Africa in May 2018



**Table 1** Summary of advisory group recommendations for inactivated vaccines with broad indications for HIV-infected adults

Vaccine	WHO [6, 7]	UK [8]	Europe [12]	France [14]	US [9–11, 13]
Pneumococcal	Not recommended in resource-limited settings	<p>Recommended for all patients. <u>Use PCV-13 (one dose)</u> regardless of HIV control.</p> <p>PPV recommended only for those with additional risk factors which include:</p> <ul style="list-style-type: none"> <li>·Age &gt;65 years old</li> <li>·Younger adults with concurrent comorbidity (e.g., asplenia) based on national program recommendations</li> </ul> <p>Dosed as 1 dose of PPV-23 with PPV-23 given <math>\geq 3</math> months after PCV-13</p> <p>No repeat doses of PPV-23 or PCV-13 are advised</p>	<p>Recommended for all patients. Use <u>PCV-13 (one dose)</u></p> <p>No repeat dosing advised</p>	<p>Recommended for all patients. Use PCV-13 and PPV-23</p> <p>Previously unvaccinated: 1 dose of PCV-13 followed by PPV-23 at <math>\geq 2</math> months later</p> <p>Previously vaccinated with PPV-23: 1 dose of PCV-13 at <math>\geq 3</math> years followed 2 months later with 1 dose of PPV-23</p>	<p>Recommended for all patients. Use PCV-13 and PPV-23</p> <p>Previously unvaccinated: 1 dose of PCV-13 followed by 1 dose of PPV-23 at <math>\geq 8</math> weeks later (preferably when CD4 count <math>\geq 200</math> cells/mm<sup>3</sup>). Repeat PPV-23 dose 5 years later</p> <p>Previously vaccinated with PPV-23, give PCV-13 at <math>\geq 1</math> year later followed by PPV-23 at 5 years later</p>

# PCD

- 14-year-old female HIV +ve patient on ART- TLD
  - CD4+ count 450 cells/L
  - VL <50 cps/mL
  - No other medical conditions
- **Which vaccines are indicated in this patient?**
- **How many vaccines can be given at the same time?**

# Guidelines for the vaccination of HIV-infected adolescents and adults in South Africa



- **Tetanus-diphtheria (Td)**
  - Vaccinated irrespective of CD4+ count
  - Booster vaccine every 10 years (until more data available)

# Guidelines for the vaccination of HIV-infected adolescents and adults in South Africa



PREVENT CERVICAL CANCER

Human Papillomavirus (HPV) vaccine is now available for **grade 4** girls who are 9 years and older.

Protecting young girls, future women of South Africa against cervical cancer.



- **Human papilloma virus**

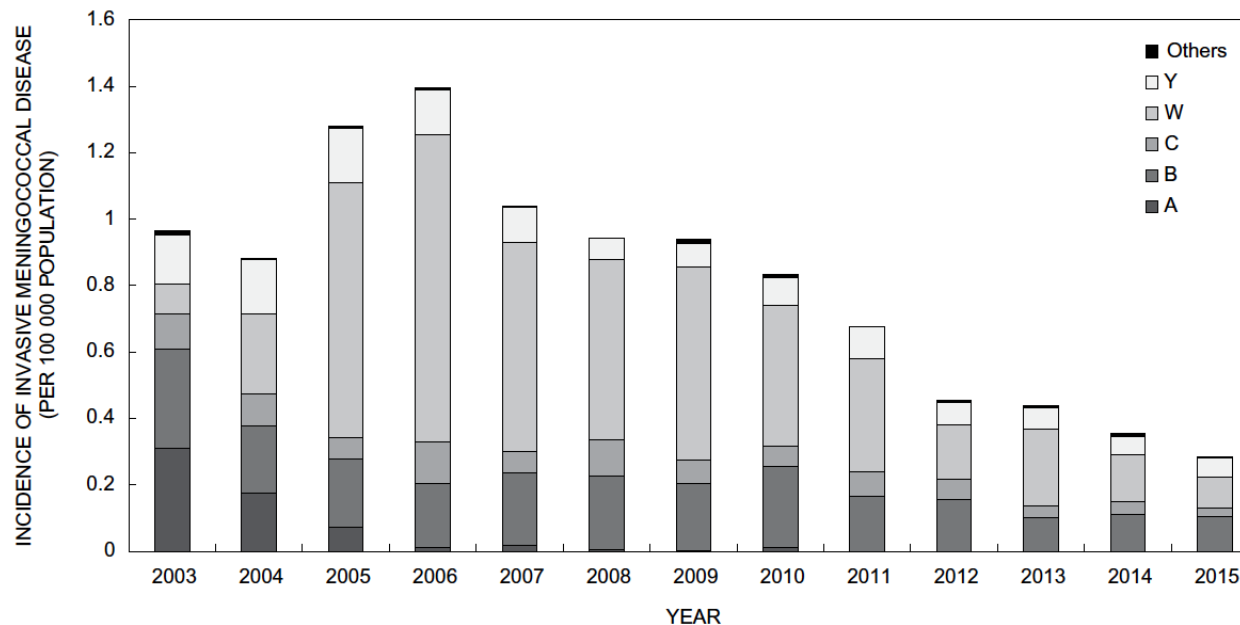
- In SA HPV- preteen girls 9-13 yrs- regardless of HIV status
- Recommended for all HIV-infected adult men (**MSM**) & **women**,
- Can be given regardless of CD4+ count, ART use or viral load

# Guidelines for the vaccination of HIV-infected adolescents and adults in South Africa



## • Meningococcal

- Should be considered
- 2 dose schedule (12 weeks apart)
- Booster every 5 years







Safety and Immunogenicity of an Adjuvanted Herpes Zoster Subunit Candidate Vaccine in HIV-Infected Adults: A Phase 1/2a Randomized, Placebo-Controlled Study



**SHINGRIX**  
(ZOSTER VACCINE  
RECOMBINANT, ADJUVANTED)

**The Journal of Infectious Diseases® 2015;211:1279–87**

**ZOSTAVAX®**  
Zoster Vaccine Live

**Zostavax® is Generally Safe and Immunogenic in HIV-Infected Adults with CD4 Counts  $\geq 200$  Cells/ $\mu$ L Virologically Suppressed on ART: Results of a Phase 2, Randomized, Placebo-Controlled Trial**

CA Benson, L Hua, JW Anderson, JH Jiang, DR Bozzolo, K Bergstrom, PW Annunziato, SW Read, R Pollard, D Rusin, J Lennox

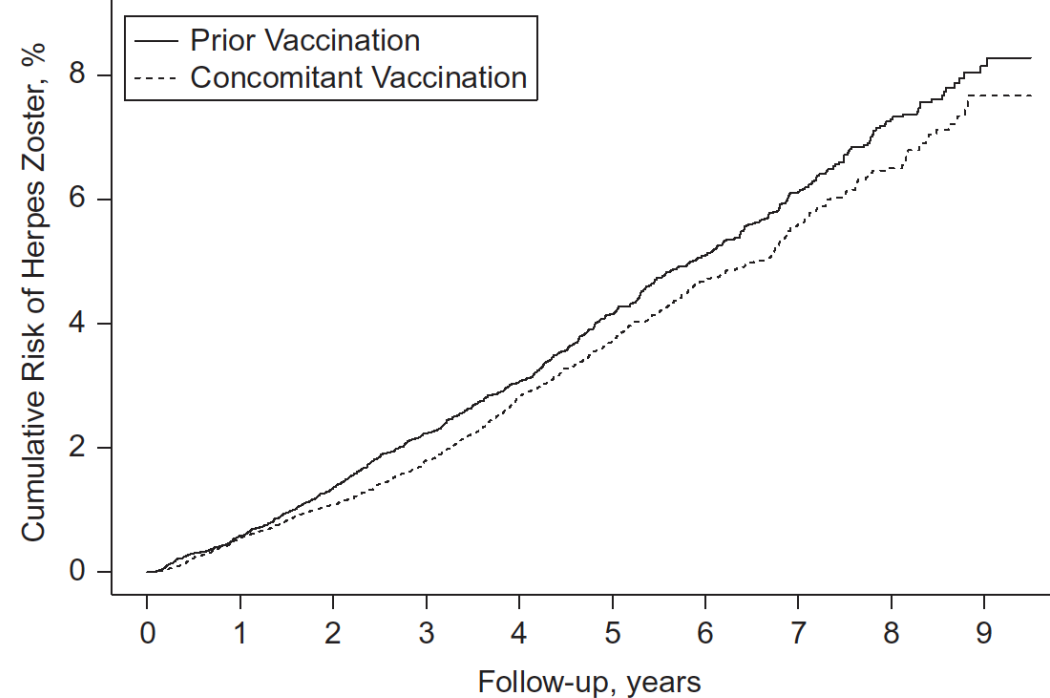
for the ACTG A5247 Team

Abstract #96

- Benefits of zoster vaccine
  - Reduce incidence of shingles
  - Reduce severity of disease
  - Reduce occurrence of post-herpetic neuralgia
- Concerns that remain
  - Lack of data on ideal dosing schedule
  - Safety & efficacy



# Real-World Evidence for Regulatory Decisions: Concomitant Administration of Zoster Vaccine Live and Pneumococcal Polysaccharide Vaccine



- Results from this study suggest no evidence of a diminished VZV immune response following concomitant administration of ZVL with PPSV23.
- Concomitant vaccination is recommended to minimize barriers to patients & providers & improve vaccination coverage

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Protected Together

#VACCINESWORK

WORLD IMMUNIZATION WEEK 2018



# Vaccine advocacy

- Advocacy is all about change:
  - Changed attitudes
  - Behaviour
  - Policies
  - Practices
  - Bridging the gap from problem to solution
- **What we need to ask in South Africa is:**
  - What **CHANGE** is needed?
  - Who can make the **CHANGE** happen?
  - How do we persuade for **CHANGE**?

Protected Together

#VACCINESWORK

WORLD IMMUNIZATION WEEK 2018



# Intervention strategies to increase adult vaccination rates

- **Pillar 1:** Convenient vaccination services
- **Pillar 2:** Communication with patients about importance of vaccination and the availability of vaccines
- **Pillar 3:** Enhanced office systems to facilitate adult vaccination
- **Pillar 4:** Motivation through an office immunization champion



# Conclusion

- Are opportunities to expand immunization for HIV-infected Adolescents & Adults
- Vaccinate during stable disease
- Communicate with patients about the importance of vaccination and the availability of vaccines
- Vaccination is the most cost effective intervention of 21<sup>st</sup> century

**THANK YOU!**

