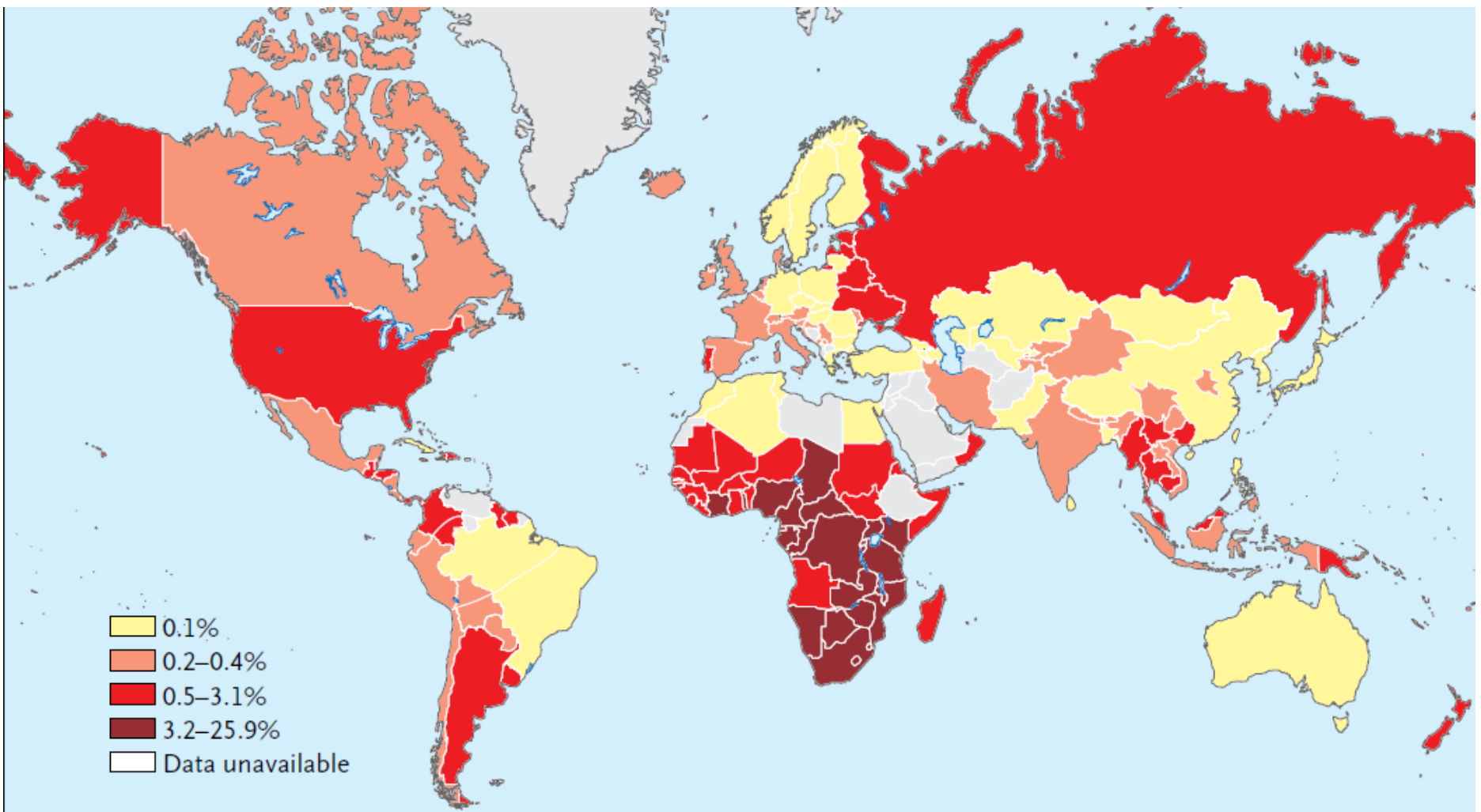


DETECTION OF PRIMARY OR EARLY HIV-1 INFECTIONS  
IN PRETORIA  
– (preliminary results)

*S. Mayaphi<sup>1</sup>, T. Rossouw<sup>2</sup>, S.A.S. Olorunju<sup>3</sup>, D. Martin<sup>1</sup>*

*1 – Clinical virologist, 2 – HIV specialist, 3 – Biostatistician*

# GLOBAL HIV Prevalence (Population Age 15 – 49, 2009)

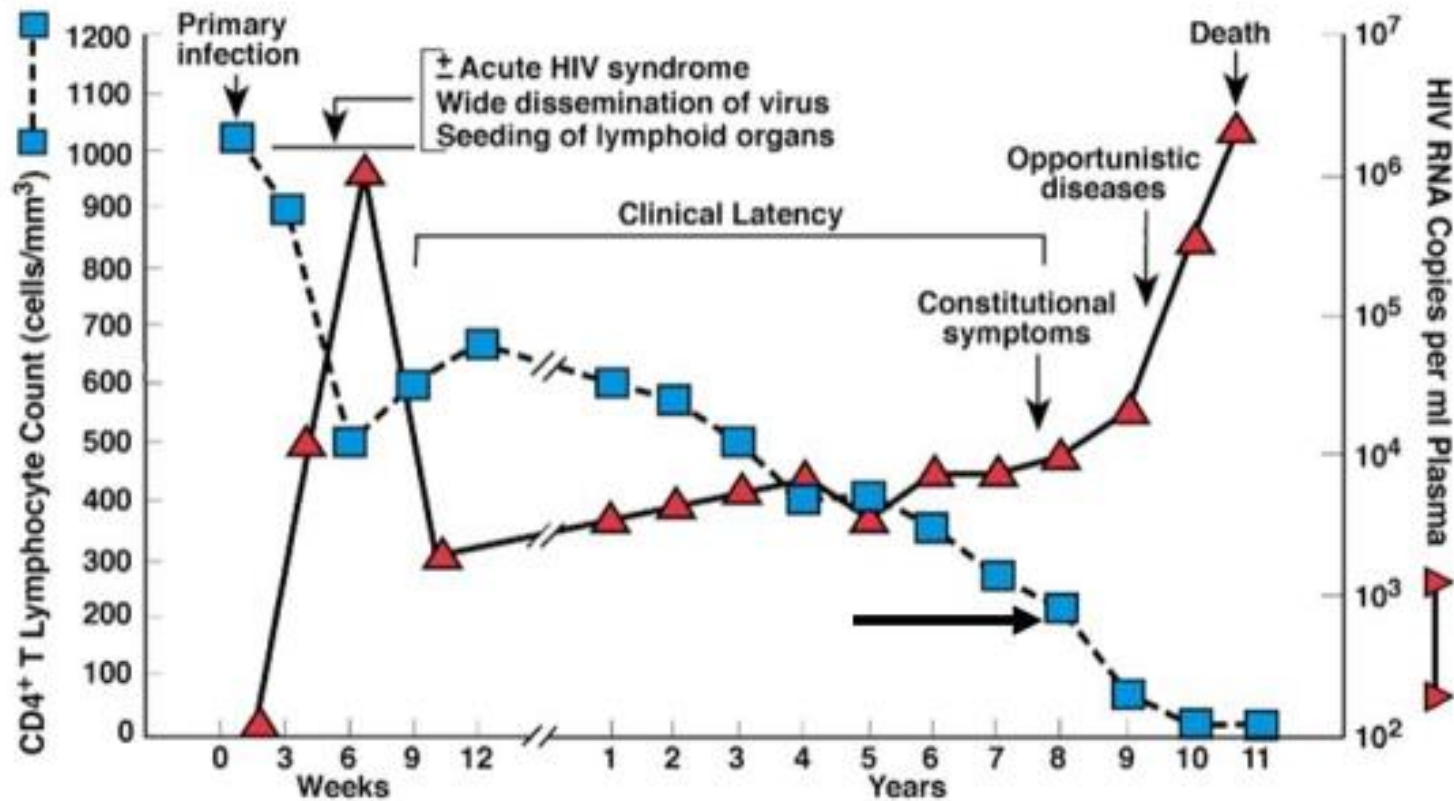


Kourtis AP, et al. N Engl J Med 2012; 366 (19)1749 – 52.

# PRIMARY (ACUTE) HIV INFECTION

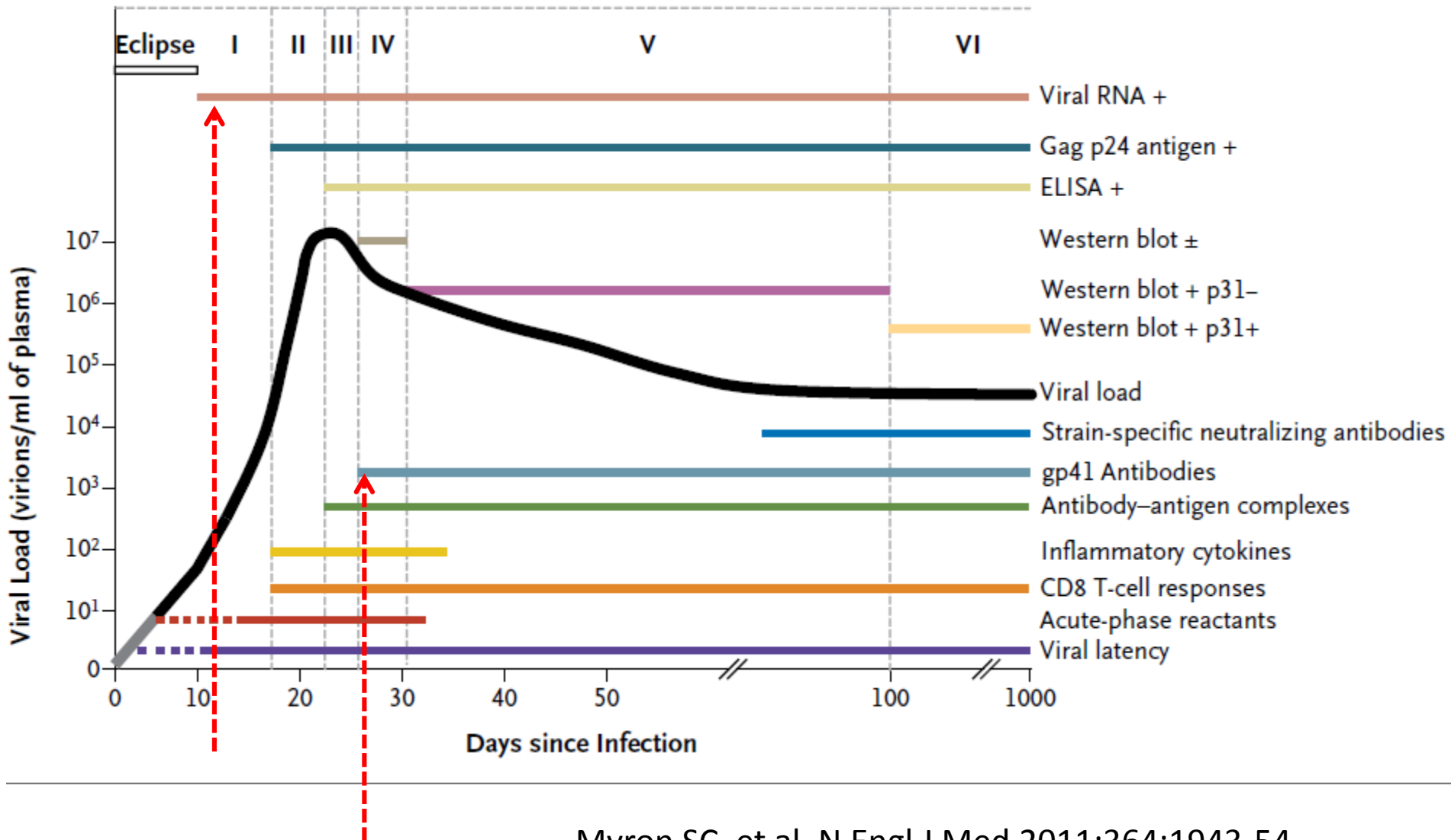
- Primary HIV infection (PHI)
  - is defined as the interval between the time of infection with HIV and that of detectable antibodies (~3 - 12 weeks)
  - extremely high levels of infectious virus are detectable in serum and genital secretions and persist for 10 – 12 weeks
  - the rate of transmission during PHI is **~26 times** as high as that during established HIV infection
  - can account for 10%–50% of all new HIV infections, especially in the context of high sexual partner concurrency or high rates of partner change

# Typical Course of HIV Infection



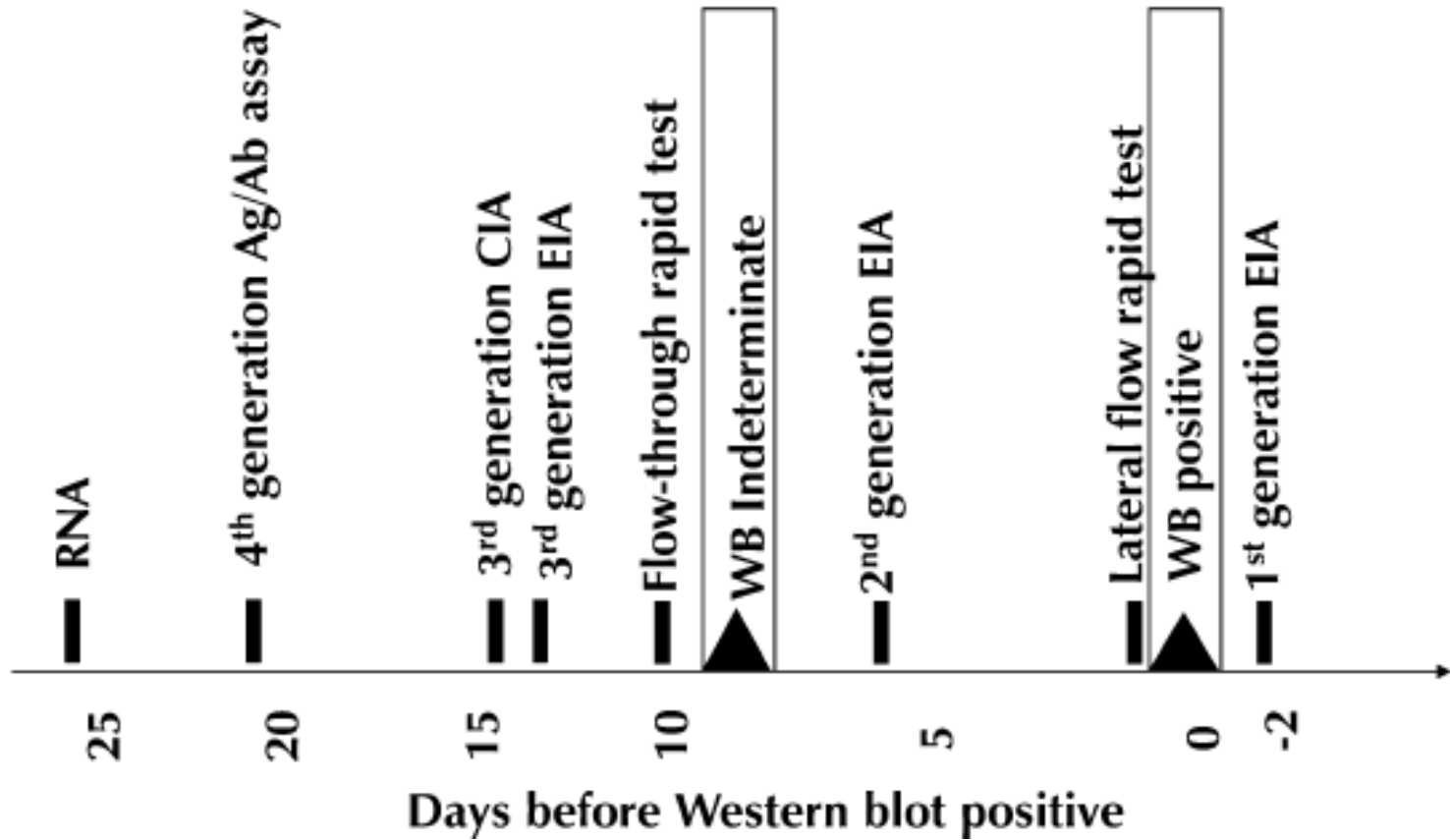
Modified From: Fauci, A.S., et al, *Ann. Intern. Med.*, 124:654, 1996

# FIEBIG'S STAGING OF PHI



Myron SC, et al. N Engl J Med 2011;364:1943-54.

# HIV ASSAYS vs WESTERN BLOT



# STUDY AIM

- To assess the burden of primary HIV infections in VCT clinics around Pretoria.

## **Objectives**

- To use pooled nucleic acid testing (pNAT) to detect the presence of PHI in individuals who test negative on rapid HIV tests.
- To subtype detected PHIs and check their ARV resistance profile.
- To assess if a questionnaire tool that captures HIV risk behaviour can be used to predict PHIs

# MATERIALS AND METHODS

## Study design and sample size:

- This is a cross-sectional study that will enroll about 4000 ✓ Condom use

## Study duration:

- This study is expected to last for a period of 3 years (from 2012 – 2014).

## Study sites:

- Tshwane district hospital VCT clinic
- FF Ribiero clinic
- Skinner clinic

## Study documents:

- Consent form and questionnaire (HIV risk behaviour)

- 
- ✓ History of unprotected sex
  - ✓ Number of sexual partners
  - ✓ Drug abuse
  - ..and more...



HIV Rapid test

Negative

Positive

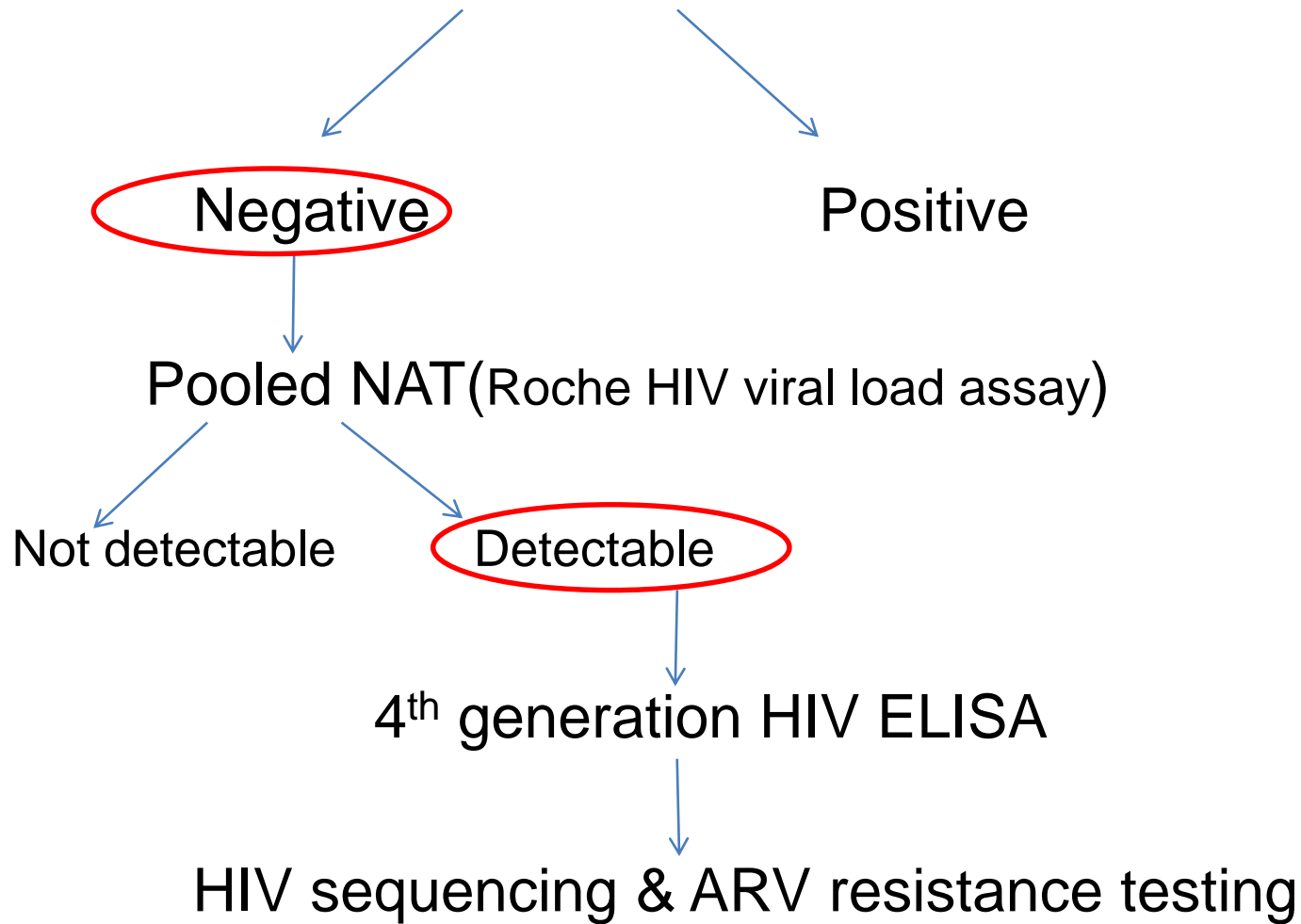
Pooled NAT (Roche HIV viral load assay)

Not detectable

Detectable

4<sup>th</sup> generation HIV ELISA

HIV sequencing & ARV resistance testing



# POOLED NUCLEIC ACID TESTING (pNAT)



200µl plasma  
from each  
tube

# LOWER DETECTION LIMITS OF HIV MOLECULAR ASSAYS (in plasma) USED IN NHLS LABORATORIES IN **2012**

**Qualitative HIV PCR (Roche CAP-CTM ): 514 copies/mL**

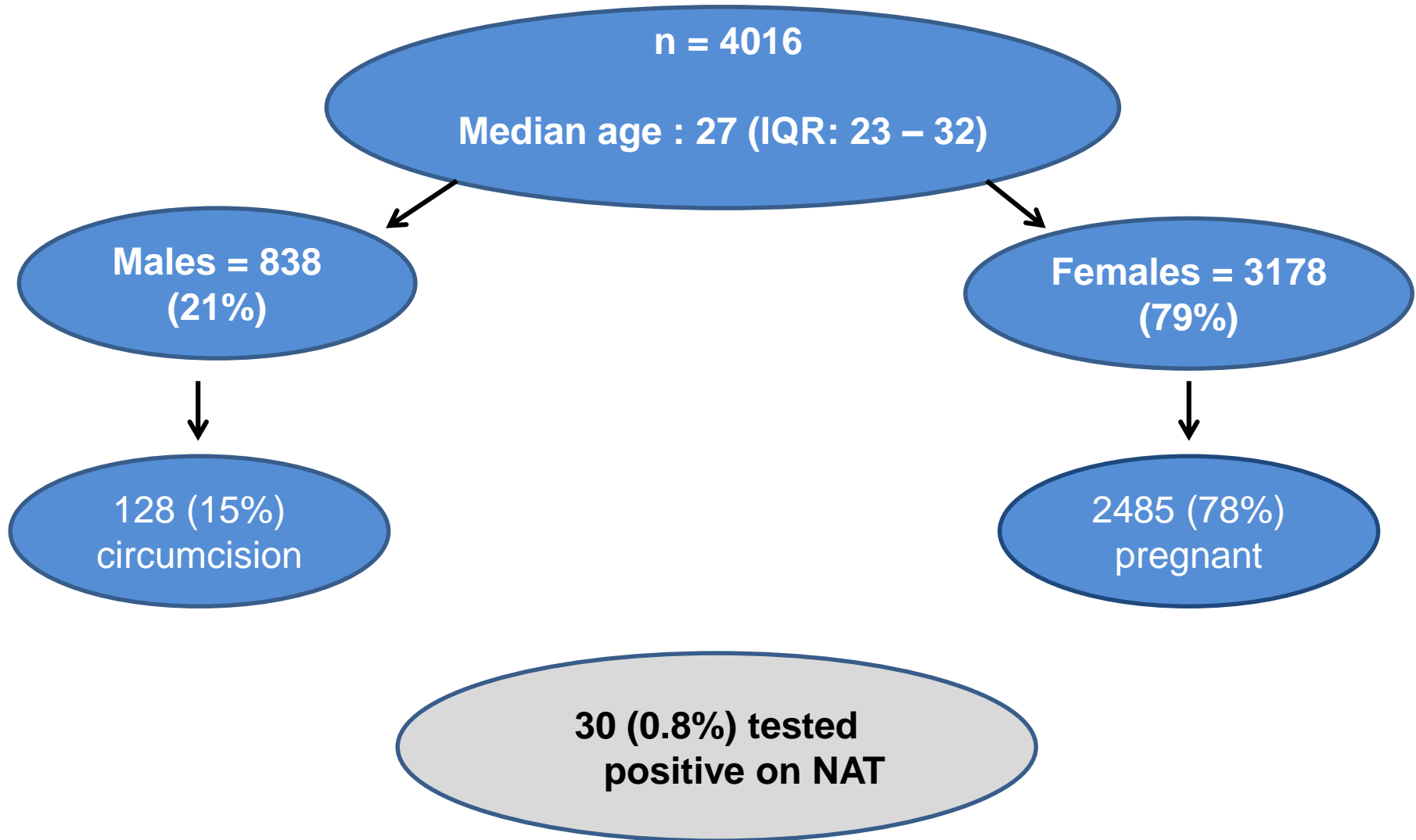
**Abbott HIV viral load assay (m2000): 40 copies/mL**

**Roche HIV viral load assay (CAP-CTM v2): 20 copies/mL**

1. Stevens W, et al. J Clin Microbiol 2008; 46 (12) 3941–3945.
2. Roche and Abbot HIV viral loads packages inserts.

# PRELIMINARY RESULTS

(March 2012 – mid Sep 2014)



# INCIDENT HIV INFECTIONS IN STUDY SUBGROUPS

- **30 (0.8%) – overall incidence**

**Circumcision (n - 128)  
= 0%**

**Non-pregnant group  
(n - 1531)  
= 0.7%**

**Pregnant women  
(n - 2485)  
= 0.8%**

# SUMMARY OF POSITIVE PARTICIPANTS (n = 30)

- All participants had **negative HIV rapid tests** at enrolment

Pregnant women	Non-pregnant group
19	11 (7 females)

HIV viral load levels	$\leq 10^2$	$10^3$	$10^4$	$10^5$	$10^6$	$>10^7$
n	3	7	13	4	1	2

- 4<sup>th</sup> generation HIV ELISA performed – first 15 participants
  - all tested positive except for one
  - HIV antibody and p24 antigen will be tested separately later

## TIME INTERVAL TO POSITIVE RAPID TEST

- 21 participants had follow up rapid HIV test
  - all tested positive except for one

2 – 6 weeks

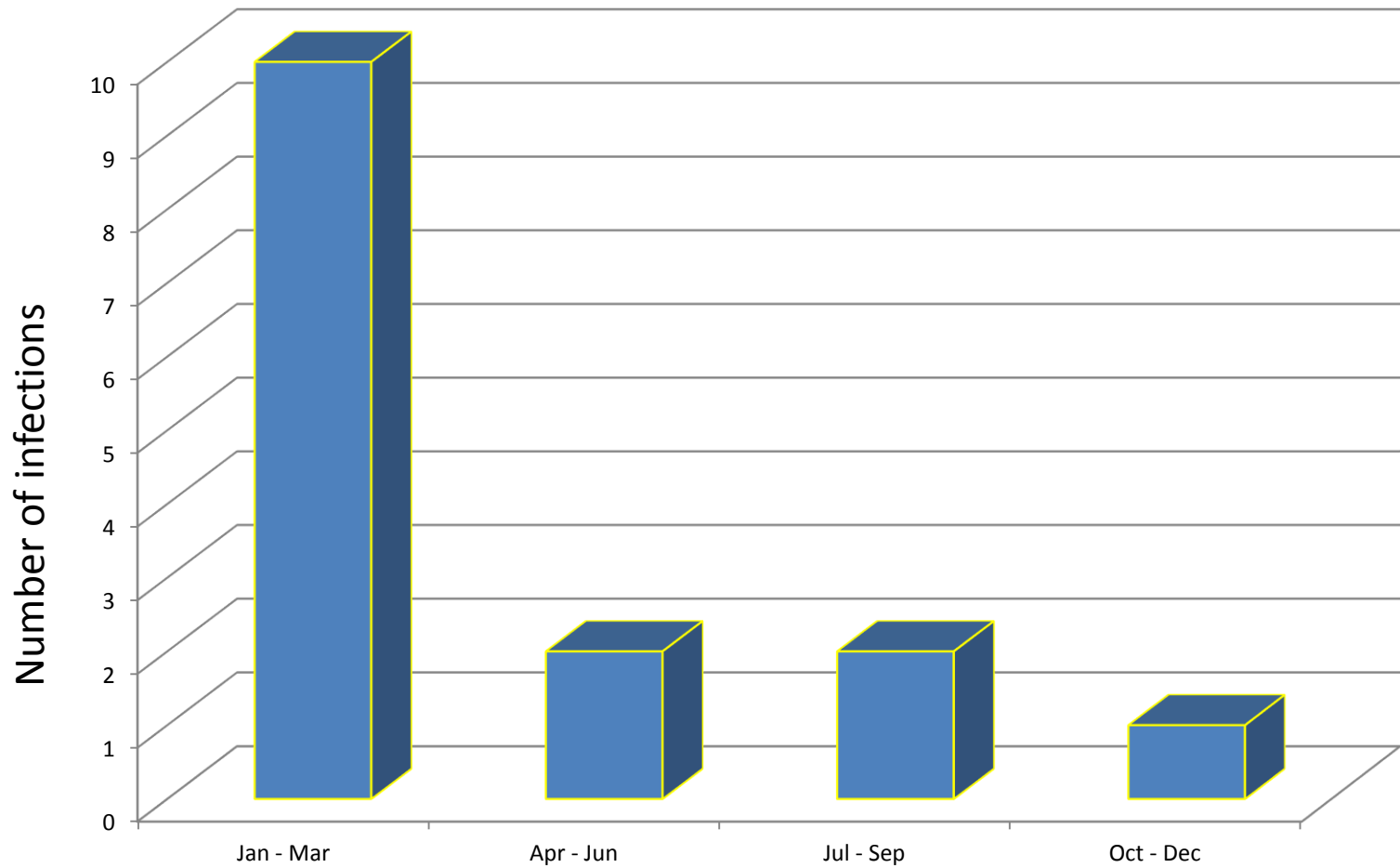
n = 15

7 - 14 weeks

n = 6\*

\* = 1 tested negative at 10 week follow up

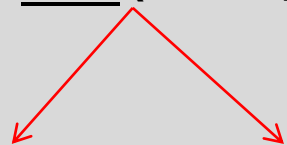
# SEASONS AND PHI INFECTIONS (March 2012 - Feb 2014)





# FREQUENCY OF CONDOM USE (n = 4014)

Never (n = 1857, 46%)

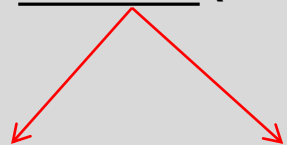


Gender: M - 353 F - 1504

Marital status distribution

S - 818 M - 1028 D - 11

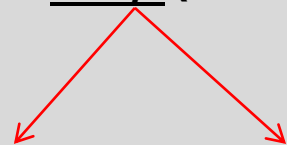
Sometimes (n = 1936, 48%)



M - 371 F - 1565

S - 1690 M - 236 D - 10

Always (n = 221, 6%)



M - 114 F - 107

S - 195 M - 19 D - 7

Incident HIV infections: 13 17 0

Marital status and infections:

Single: 23 Married: 6 Divorced: 1

# PRIMARY HIV INFECTION INCIDENCE (measured by nucleic acid tests)

Publications	Country	Acute HIV incidence	Sample size (rapid HIV - negative)
Pilcher CD, et al. <b>2005</b> N Engl J Med;352:1873-83.	USA	0.02%	108667
Shepard CW, et al. <b>2008</b> MMWR; CDC.	USA	0.08%	21241
Stekler JD, et al. <b>2009</b> CID; 49:444–53.	USA	0.3%	13677

# CONCLUSIONS

- Feasibility of incident HIV detection in SA through the use of pNAT
- Other options of detecting these infections are:
  - HIV ELISA
  - Repeat rapid test at 6 weeks later
- A questionnaire tool can be used for prediction of incident HIV infections
- Innovative ideas are needed for promotion of condom use in SA
- Detection of incident HIV infections missed by the rapid tests has a huge potential of reducing HIV spread and prevalence



I'm ~ 95% effective!!!



# ACKNOWLEDGEMENTS

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- NHLS Research Trust grant
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- University of Pretoria Research assistant grant
- Virology department staff – University of Pretoria



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