

From Evidence to Action: The 2nd International HIV Treatment as Prevention Workshop

> April 22-25, 2012 www.treatmentaspreventionworkshop.org

Julio Montaner MD, DSc(hon), FRCPC, FCCP, FRSC, OBC

Director, BC-Centre for Excellence in HIV/AIDS, Providence Health Care Professor of Medicine and Head, Division of AIDS, University of British Columbia Past-President, International AIDS Society



BRITISH COLUMBIA CENTRE for EXCELLENCE in HIV/AIDS



a place of mindTHE UNIVERSITY OF BRITISH COLUMBIA





Scientific Advisory Committee

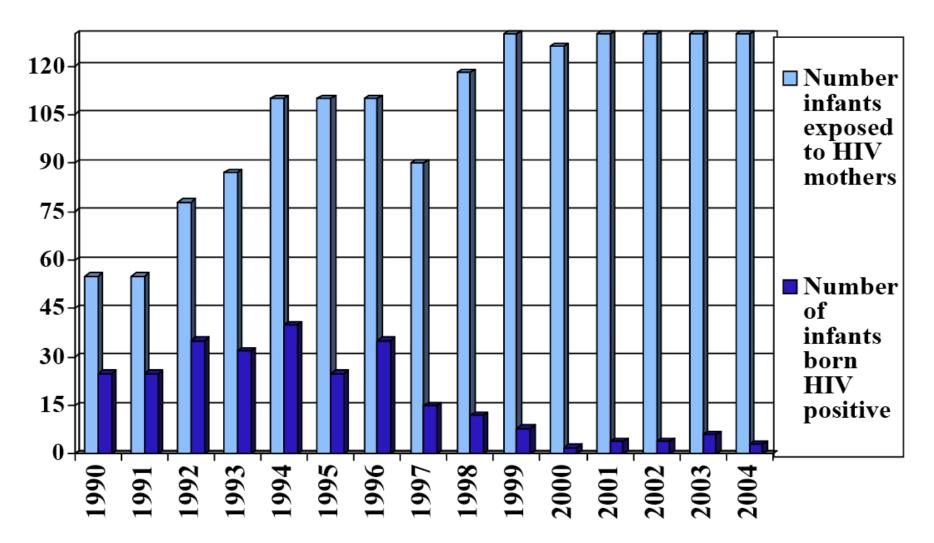
Julio S.G. Montaner, Chair Elly Katabira, Co-Chair **Bertrand Audoin** Curt Beckwith Pedro Cahn Yvette Calderon Chinazo Cunningham Carlos Del Rio Wafaa El-Sadr Alison End Geoff Garnett **Reuben Granich**

Mark Harrington **Gregory Kirk** Ann Kurth Viviane Dias Lima Jacqueline Ying-Ru Lo Rob Lyerla Jean-Michel Molina **Jacques Normand** Mariangela Simao Papa Salif Sow **Brian Williams** Zunyou Wu

7⁸⁹!'#' 1(# 0' "+ # 05!- "3 ! ' (10' 2 1#' !0 &!*(131#' "+ # !:' ; 8* < = + (>& + * 3 &4!56789.--!

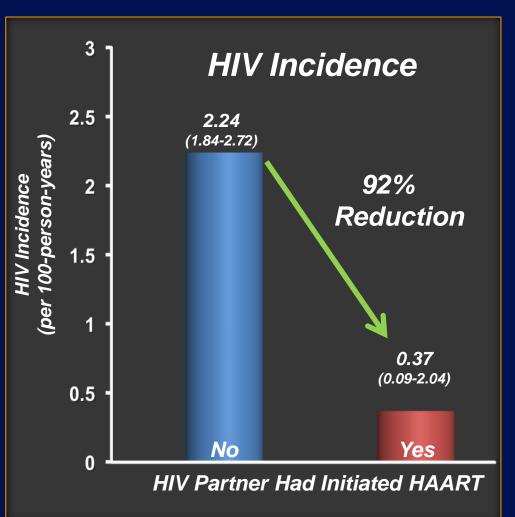


Canada: Infants Exposed to HIV and Born HIV Positive



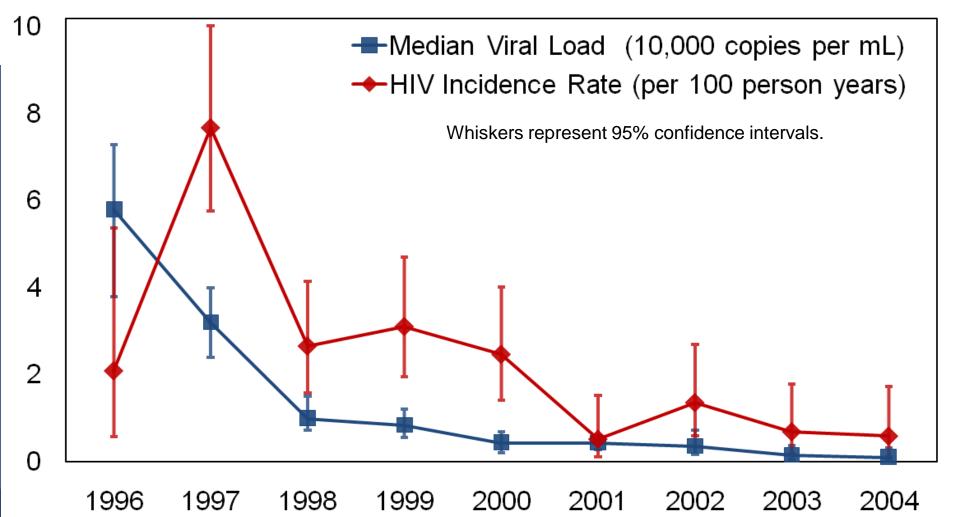
Partners for the Prevention of HSV/HIV Transmission Study: Discordant Couples

- Prospective cohort analysis (n=3408 heterosexual HIV discordant couples
- 7 African countries
- 349 HIV+ initiated HAART
- 103 Linked HIV transmissions Only 1 while on HAART. Adjusted incidence rate ratio 0.08% (0.00-0.57; P=0.004)
- Sexual risk behaviors decreased after HAART 6.2% vs 3.7% (P=0.03)
- No change in sexual frequency

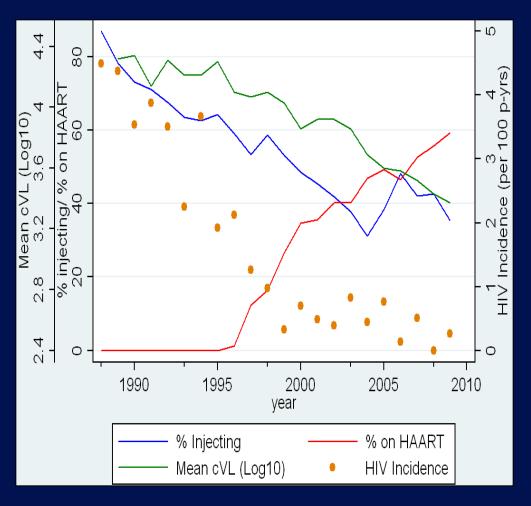


Longitudinal community plasma HIV-1 RNA concentrations and incidence of HIV-1 among injecting drug users: prospective cohort study

BMJ | 16 MAY 2009 | VOLUME 338



HAART Reduces HIV incidence in IDUs

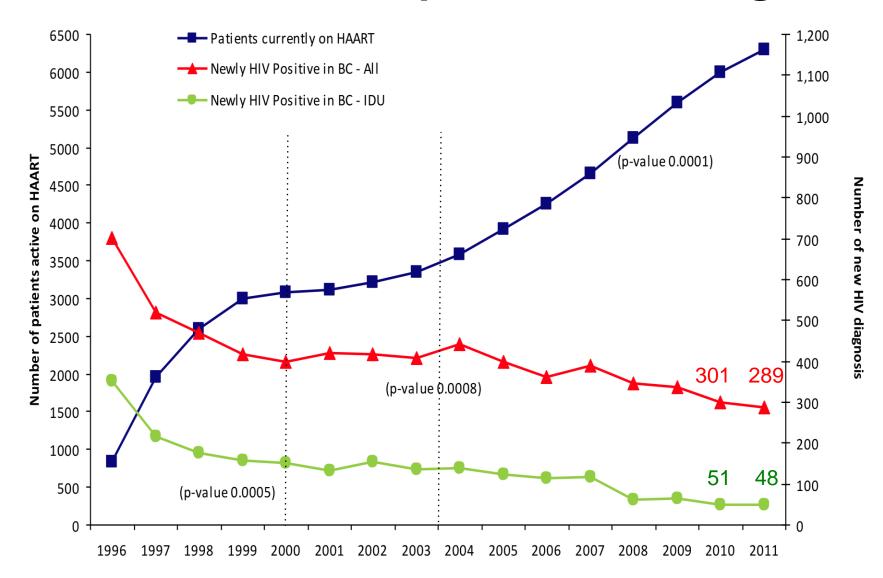


From 1997, HIV incidence
decreased by 74% for each log
decline in community HIV viral
load

In a separate model, HIV
incidence decreased by 5% for
each 1% increase in HAART
coverage

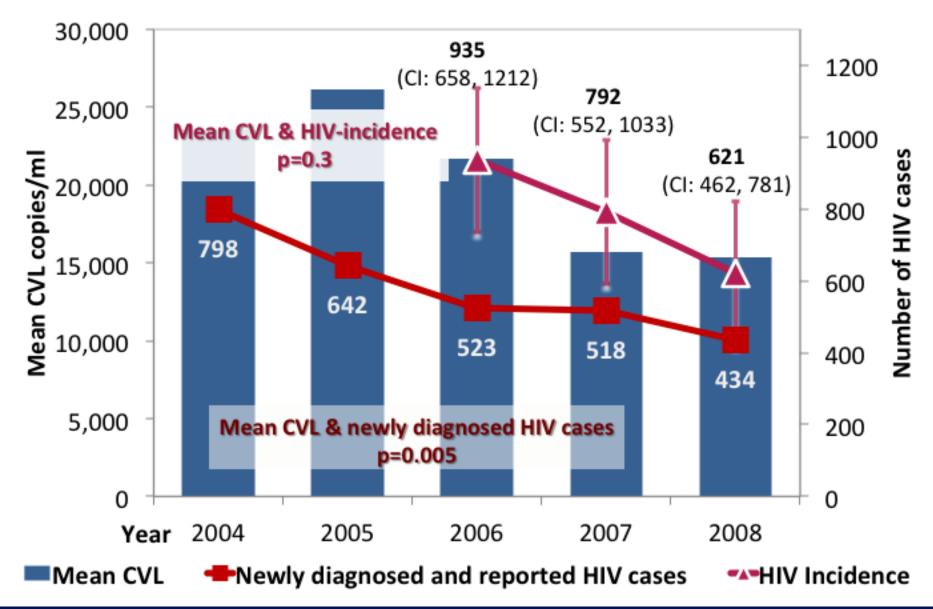
Modified from G Kirk et al, for the Alive Cohort, CROI 2011

Increasing HAART Coverage <u>within</u> Evolving Guidelines in BC - Impact on New Diagnoses

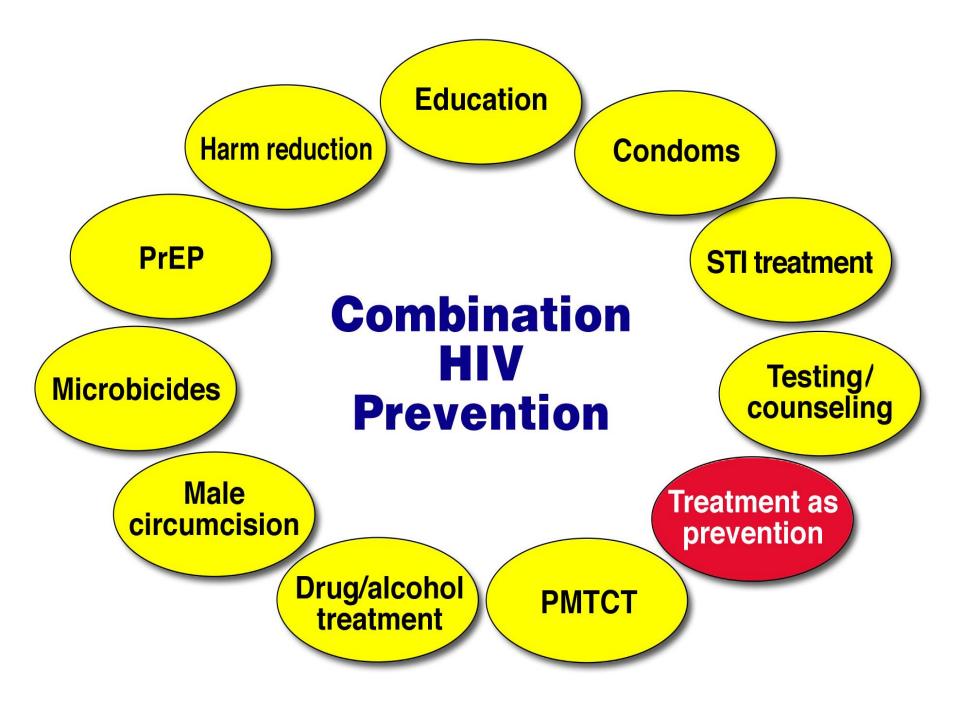


TasP Workshop 2012 (Updated from Montaner et al, Lancet, 2010)

Mean CVL and New HIV Infections, 2004-08



Moupali Das et al, TasP Workshop 2011



More evidence on the way: TasP Relevant Studies



Current HIV Research, 2011, 9, 355-366



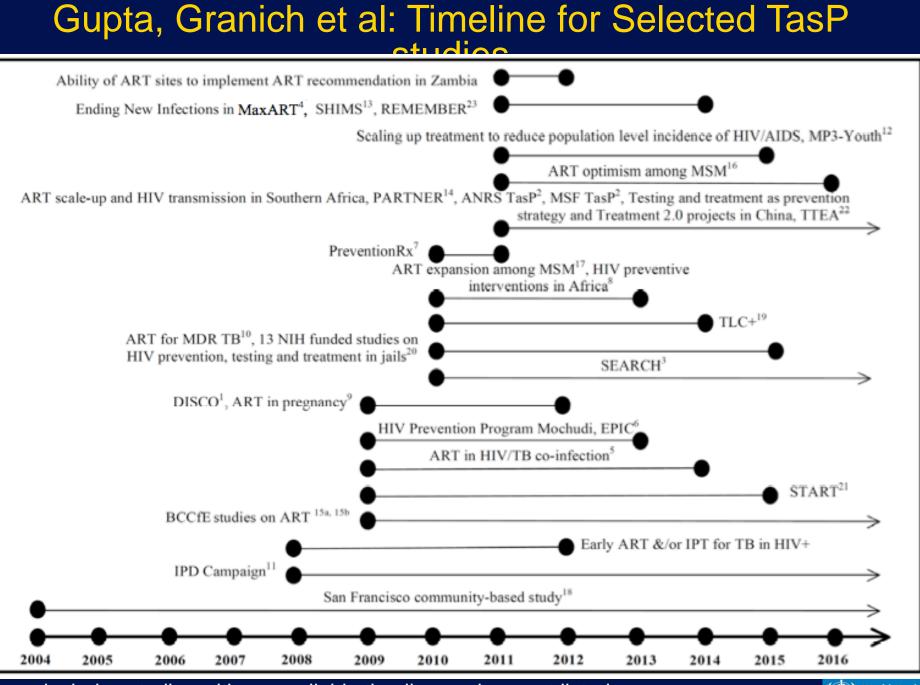
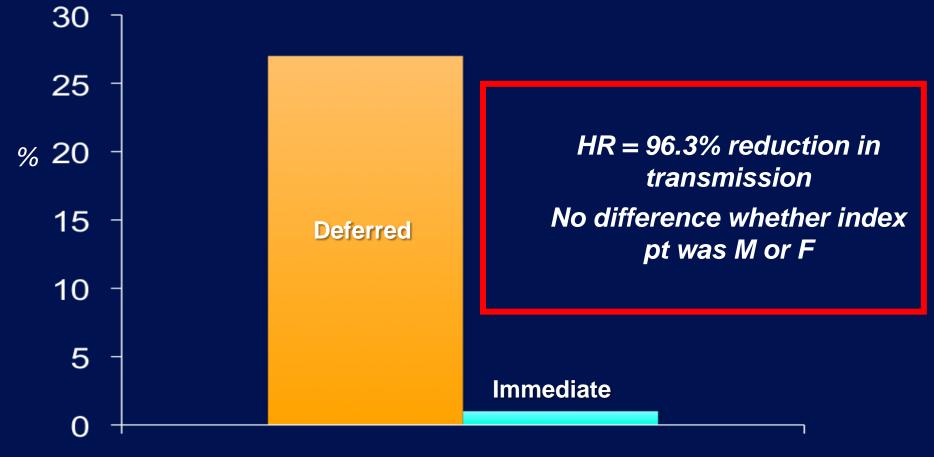


Figure includes studies with an available timeline—others not listed.

World Health Organization

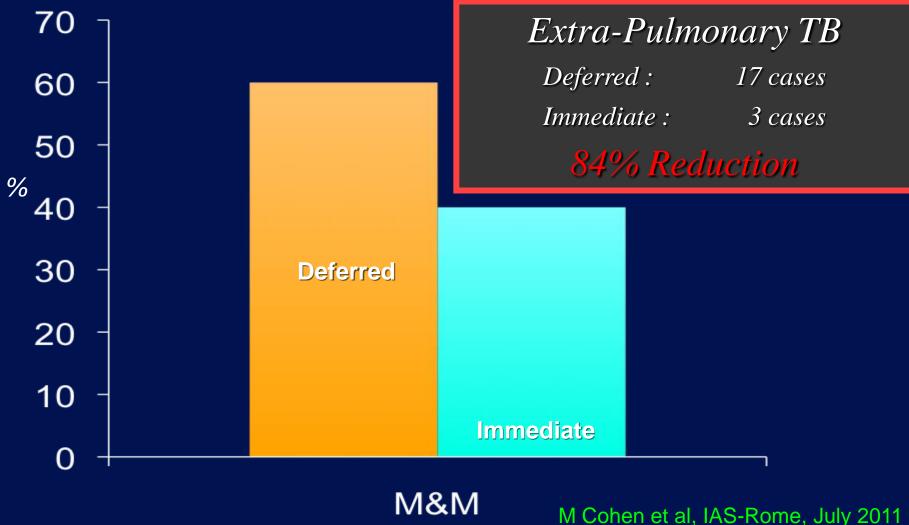
HPTN 052: Immediate vs Delayed ART in Sero-discordant Couples



Sero-Conversions

Cohen MS, et al. IAS 2011. Abst MOAX0102 Cohen MS, et al. N Engl J Med. 2011

HPTN 052: Immediate vs Delayed ART in Sero-discordant Couples



M Cohen, NEJM 2011

The Economist

30HE 47H-10TH 2011

The trap for Turkey Wall Street's plumbing problem Lady Gaga, Mother Teresa and profits Brazil's boiling economy The farce that is FIFA

The end of AIDS?

Economist.com

How 5 million lives have been saved, and a plague could now be defeated

Aids - is the end really in sight?

US Secretary of State Hillary Clinton has offered to lead the endstage of the battle against Aids - but will the money be forthcoming to finish the job and deliver an "Aids-free generation"?



Three Key Pillars:

MC MTCT TasP

The Guardian, UK Nov 8th 2011

US secretary of state Hillary Clinton has urged the United States and other nations to increase their funding for HIV/Aids prevention and treatment. Photograph: Win McNamee/Getty Images



Few could have imagined that we'd be talking about the real possibility of an AIDS-free generation. But that's what we're talking about...make no mistake, we are going to win this fight.

President Obama, December 1, 2011

By the end of 2013, PEPFAR will directly support more than 6 M people on HAART– 2M more than previously targeted.

Ambassador Eric Goosby, TasP Workshop 2012

BREAKTHROUGH

HIV Treatment as Prevention

On 1 December, George Washington University in Washington, D.C., hosted "The Beginning of the End of AIDS," a splashy World AIDS Day event that featured three U.S. presidents, business magnates, and rock stars. The catalyst that brought them together was something Anthony Fauci, the top U.S. government HIV/AIDS scientist, told the crowd even 1 year ago would have seemed "wishful thinking": a clinical trial dubbed HPTN 052 and its "astounding" result.

HIV/AIDS researchers have long debated whether antiretroviral drugs (ARVs)



The researchers planned to compare the groups until 2015. But on 28 April, an independent monitoring board that periodically

e data stunned Cohen and his when it recommended that the e trial be made public as soon Of the 28 people who become n HIV that genetically matched n their long-term partners, only e early treatment group—which nced 41% fewer serious health roblems associated with HIV. nfected people in the delayed arm f the study were offered ARVs nmediately. The HPTN 052 results and

BREAKTHROUGH OF THE YEAR

HIV Treatment as Prevention

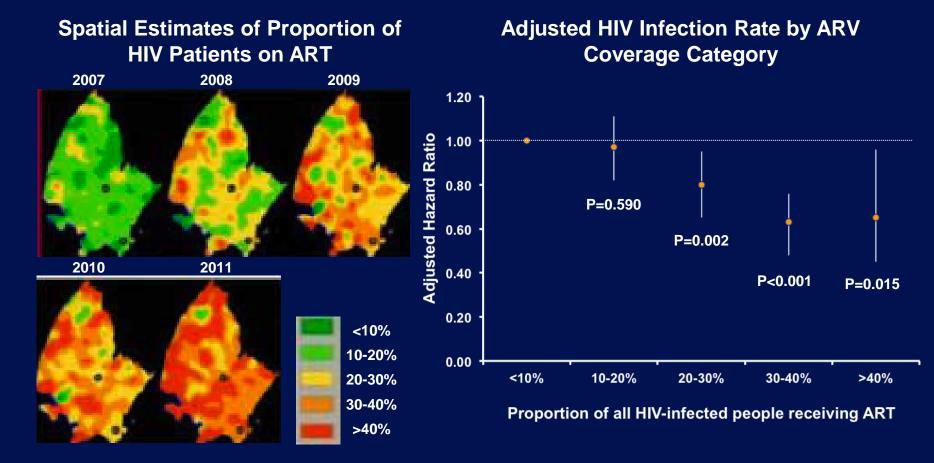
Treatment Discordant Couples in 2011

	ART	No. couples	Person-year observed	No. HIV new infections	(%)
	Yes	18,371	13366.2	151	1.13
10 - 9 -	No	16,158	12195.9	305	2.50
8- 7-	Total	35,016	25562.1	456	1.78
6 5 4 3 2 1 0	■非治疗组 ■ 治疗组		ART reduce 55%		
	0.1 0.2 0.3 0.4 0.5 0.6 0.7 0.8 0.9 1.0		transmis 20 ⁴		
		观察人年	Z Wu, TasP Workshop 2012		

阳转概率(%)

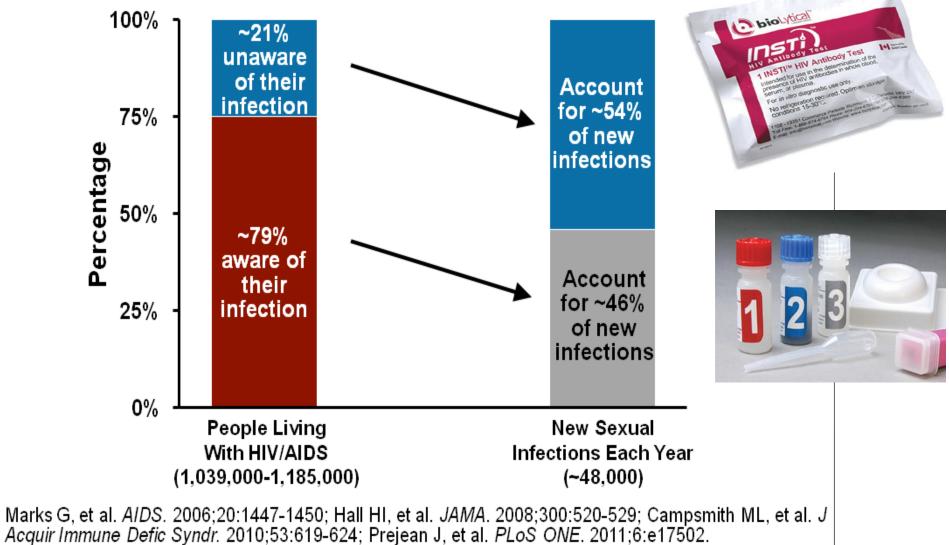
Treatment as Prevention: Effect of ART Coverage on HIV Incidence in Rural South Africa

- Annual population based HIV surveillance in rural KwaZulu-Natal 2004 to 2011
- 1395 HIV seroconversions among 16,588 HIV negative adults ≥15 years of age

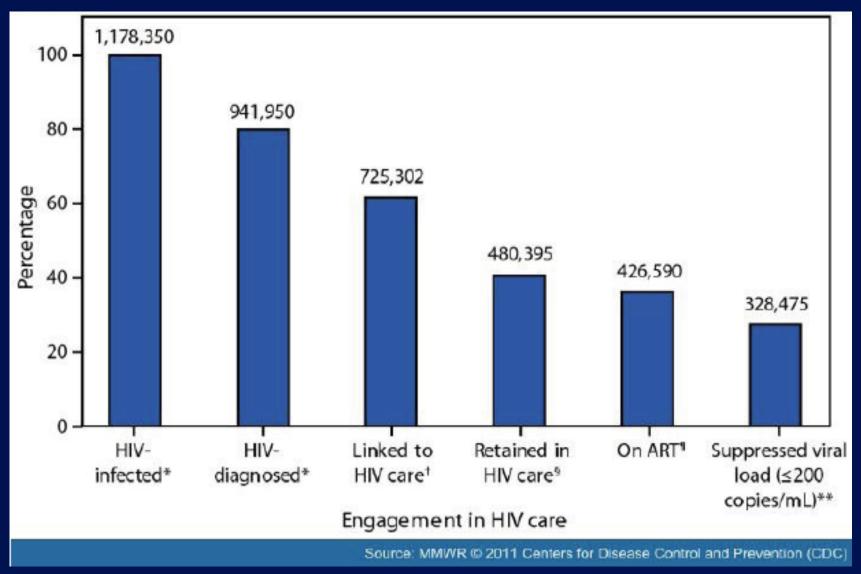


Tanser F, et al. 19th CROI, 2012. Abst. 136LB

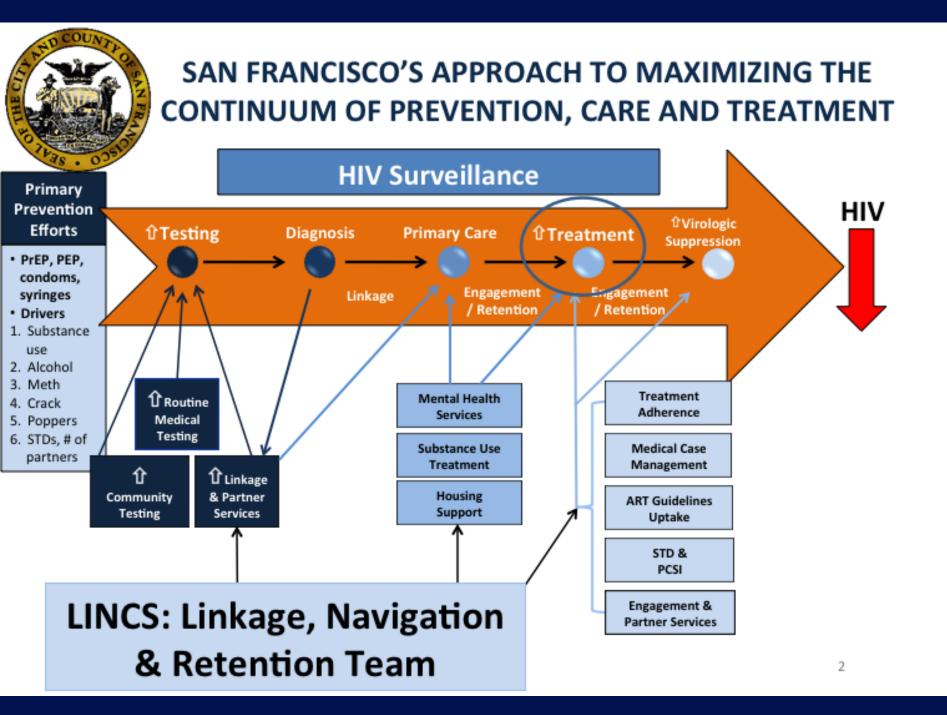
Majority of HIV Transmissions From **People Unaware of Their Infection**



Spectrum of Engagement in care - USA



CDC-USA



IAS-USA Guidelines: When to Start

CD4+ Cell Count	Recommendation
< 350 cells/mm ³	 Start ART (AI)
 350-500 cells/mm³ 	 Start ART (AII)
> 500 cells/mm ³	 Start ART (BIII)

Clinical Conditions Favoring Initiation of Therapy Regardless of CD4+ Cell Count

- History of AIDS-defining illness (AI)
- Pregnancy (AI)
- HIV-associated nephropathy (AII)
- HBV co-infection (AII)
- Patients at risk of transmitting HIV to sexual partners (AI, heterosexuals; AIII, others)
- HCV co-infection* (BII)
- Patients > 50 years of age (BIII)

Offer ART to all HIV infected patients unless patient is elite controller or has stable CD4+ count and low HIV-1 RNA in absence of ART.

Thompson MA, et al. JAMA. 2010;304;321-333.

DHHS, 2012: When to Start

ART recommended for <u>all</u> HIV-infected patients

strength of recommendation varies according to CD4+ cell count

CD4+ Cell Count	Recommendation
< 350 cells/mm ³	 Start ART (AI)
 350-500 cells/mm³ 	 Start ART (AII)
> 500 cells/mm ³	 Start ART (BIII)

Clinical Conditions Favoring Initiation of Therapy Regardless of CD4+ Cell Count

- History of AIDS-defining illness (AI)
- Pregnancy (AI)
- HIV-associated nephropathy (AII)
- HBV co-infection (AII)
- Patients at risk of transmitting HIV to sexual partners (AI, heterosexuals; AIII, others)
- HCV co-infection* (BII)
- Patients > 50 years of age (BIII)

*Including those with high CD4 count and/or with cirrhosis. If CD4> 500/mm³ may elect to defer ART until after HCV therapy

• *Rating of Recommendations:* A = Strong; B = Moderate; C = Optional

• Rating of Evidence: I = data from RCTs; II = data from well-designed nonrandomized trials or cohort studies with long-term clinical outcomes; III = expert opinion

DHHS Guidelines for Antiretroviral Therapy in Adults and Adolescents. March 27, 2012.

WHO Guidance on couples HIV testing and counselling, including antiretroviral therapy for treatment and prevention in serodiscordant couples

RECOMMENDATIONS

- 1. Couples and partners should be offered voluntary HIV testing and counselling with support for mutual disclosure. *Strong recommendation, low-quality evidence.*
- 2. Couples and partners in antenatal care settings should be offered voluntary HIV testing and counselling with support for mutual disclosure. *Strong recommendation, low-quality evidence.*
- 3. Couples and partner voluntary HIV testing and counselling with support for mutual disclosure should be offered to individuals with known HIV status and their partners. Strong recommendation, low-quality evidence for all people with HIV in all epidemic settings / Conditional recommendation, low-quality evidence for HIV-negative people depending on countryspecific HIV prevalence.
- 4. People with HIV in serodiscordant couples and who are started on antiretroviral therapy (ART) for their own health should be advised that ART is also recommended to reduce HIV transmission to the uninfected partner. *Strong recommendation, high-quality evidence.*
- 5. HIV-positive partners with >350 CD4 cells/µL in serodiscordant couple should be offered ART to reduce HIV transmission to uninfected partners. *Strong recommendation, high-quality evidence.*

5. HIV-Positive partners with >350 CD4 cells/uL in serodiscordant couples should be offered ART to reduce HIV transmission to uninfected partners.

Strong recommendation, high quality evidence.

Released April 18th 2012 at http://www.who.int/hiv/pub/guidelines/9789241501972/en/index.html

Antiretroviral prophylaxis: a defining moment in HIV control

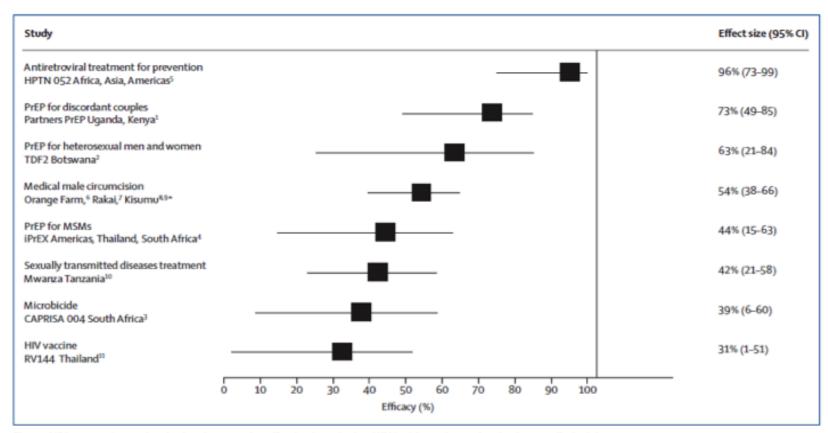
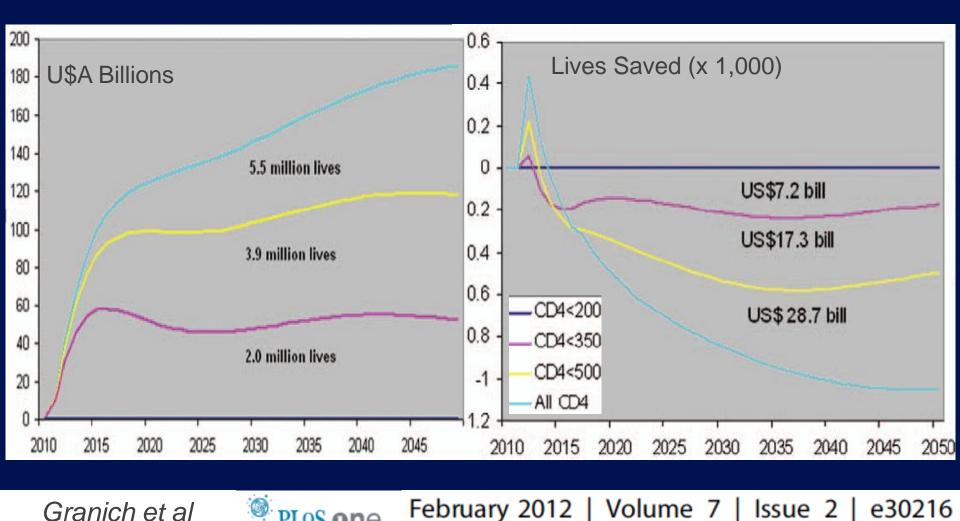


Figure: HIV prevention technologies shown to be effective in reducing HIV incidence in randomised controlled trials¹⁻³³ PrEP=Pre-exposure prophylaxis. *Meta-analysis of circumcision trials.

Published Online July 15, 2011 DOI:10.1016/S0140-6736(11)61136-7

Expanding ART for Treatment and Prevention of HIV in South Africa: Estimated Cost and Cost-Effectiveness 2011-2050



PLos one

Granich et al

Medscape Connect AIDScan TasP Works, So Let's Start Planning for Implementation

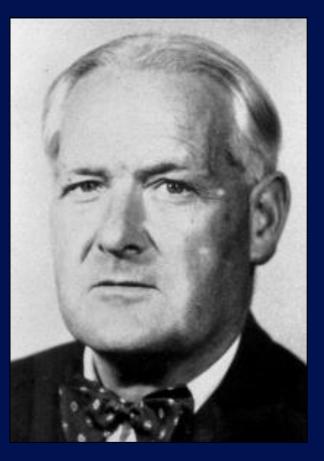
Jose M Zuniga, Other, 10:01AM May 2, 2012

The workshop's bottom-line messages:

- TasP works in combination with other prevention interventions
- Additional research is needed, without which some will have grounds to dispute TasP
- While research continues, planning for TasP implementation must start now
- HIV testing uptake and linkages to TasP and treatment as treatment must improve
- The health workforce requires training and support to implement TasP
- TasP must count on community acceptance and requires community ownership
- HIV-positive patients require treatment literacy and should not be coerced into treatment without truly informed consent

In less than three months, 30,000-plus global AIDS warriors will gather in Washington, DC, for the XIX International AIDS Conference. Several TasP studies will report out at this year's conference – which will infuse the conference with the same spirit of optimism that permeated Vancouver almost two decades ago. Our challenge is to turn optimism into action!

"The Environment and Disease: Association or Causation: A Case for Action"



All Scientific work is incomplete – whether it be observational or experimental. All scientific work is liable to be upset or modified by advancing knowledge. That does not confer upon us a freedom to ignore the knowledge we already have, or to postpone the action that it appears to demand at a given time.

Austin Bradford Hill (1897-1991) British epidemiologist best known for his research with Richard Doll, which linked smoking with lung cancer. He was also widely acknowledged as the world's leading medical statistician and a pioneer in the use of RCTs.