### **Biomedical Agents for Prevention:**

### HIV Viral Load Suppression as a Pathway to Epidemic Control

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### HPTN Annual Meeting







Defining HIV/AIDS Epidemic Control

 Viral load suppression as pathway to decrease in HIV incidence

Future directions

### What is HIV/AIDS epidemic control?





### Goal:

- Absolute rate of HIV incidence less than one per 10,000 adults per year
- 90% reduction in new HIV infections by 2030 from 2010 (UNAIDS)

# Population viral load suppression as pathway to epidemic control

- Focus on reduction of new HIV infections by achieving 95:95:95 targets
  - First 95 (Diagnosed): 95% of PLHIV know or are aware of their HIV+ status
  - Second (On Treatment): 95% of those aware of their PLHIV status are on antiretroviral therapy
  - Third 95 (With VLS): 95% of those on ART with viral load suppression



#### Viral Load Level and HIV Sexual Transmission: Observational Evidence





- Zero transmissions from PLHIV with VL <1,500 copies/mL
- Dose-response between PLHIV VL and HIV transmission
- PLHIV VL >50,000cp/mL associated with x23 higher transmission rates

Quinn TC, Wawer MJ, Sewankambo N, Serwadda D, et al (2000). Viral load and heterosexual transmission of Human Immunodeficiency Virus type 1. NEJM, Vol 342,#13.921-929

## Early ART with viral load suppression minimizes sexual transmission: Evidence from HPTN 052





- No linked infections when participants were stably virally suppressed.
- Linked were infections associated with high VL
- A substantial number of *unlinked* transmissions (36%): suggesting multiple sexual partners: *need to ensure diagnosis, treatment and VLS for all PLHIV.*

## ART, viral load suppression and sexual transmission: Evidence from Partners Study



- 1,593 couple-years of follow-up from 782 couples
- PLHIV were on ART
  - Few missed pills for 4+ days
  - 99% virally suppressed <200 copies/mL
- Zero within couple transmissions through condomless sex



Figure 1: Rate of within-couple HIV transmission through condomless sex according to sexual behaviour reported by the HIV-negative partner

#### **New HIV infections: Where are we now?**





Source: IN DANGER: UNAIDS Global AIDS Update 2022.

#### Population viral load suppression: Where do we need to be?



#### **UNAIDS** Targets for VLS at Population Level



#### **Population HIV Impact Assessments** (PHIA)



- Nationally representative household surveys funded by PEPFAR through CDC
- Conducted since 2011 in 15 countries in SSA and Haiti by Ministries of Health with ICAP support
  - Several countries with 2 PHIAs in the past ~5 years
  - Eswatini with 3 PHIAs since 2011
- Standardized methodology with random selection of sampling units and households
- All individuals in selected households 15 years and older are surveyed
- All receive HIV testing with viral load testing for those HIV-positive (irrespective of awareness of HIV)
- All HIV-positive persons with VL<1000 copies per ml have ARV levels measured in blood
- 95-95-95 are ARV adjusted



# 2015-2021: 95-95-95 Achievements among individuals 15+ years in 5 SSA Countries

Eswatini





# **2015-2021:** 95-95-95 Achievements among individuals 15+ years in 5 SSA Countries



 All 5 countries did better on all 95's in a ~5-year period

 First 95 (HIV testing, awareness) is a persistent gap, yet a critical gateway to ART and population VLS

### How are achievements in 95-95-95 translating to population VLS among ALL individuals 15+ years?





PHIA Population VLS in 2016 vs 2020

- There is progress in the past~ 5 years among all countries
- >20% of the PLHIV still capable of HIV transmission in many countries

### Eswatini: 95-95-95 and Population VLS among adults 18-49 years (2011-2021)



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13

## Eswatini: 95-95-95 and Population VLS among adults 18-49 years (2011-2021)





Nkambule R, Philip NM, Reid G, Mnisi Z, Nuwagaba-Biribonwoha H, Ao TT, et al. (2021) HIV incidence, viremia, and the national response in Eswatini: Two sequential population-based surveys. PLoS ONE 16(12): e0260892. https://doi.org/10.1371/journal.pone.0260892

### Eswatini: HIV incidence among adults 18-49 years (2011-2021)





Significant decline in HIV incidence among men and women 18-49 years

# Eswatini National HIV/AIDS Program 2010-2021



Combination HIV Prevention	<ul> <li>Programs at scale among individuals 15 years and older</li> <li>Community and key stakeholder engagement</li> <li>Differentiated service delivery, multi-month dispensing</li> </ul>	
	>500,000	<ul> <li>Condoms distributed annually to target KP</li> </ul>
	>300,000	<ul> <li>Average annual HIV tests 2014-2021, test and start strategy for all populations, index testing</li> </ul>
	>200,000	<ul> <li>PLHIV on ART, average 16,000 new PLHIV on ART annually 2011-2021</li> </ul>
	>150,000	<ul> <li>Average #VL tests conducted annually 2017-2021</li> </ul>
HIV diagnosis, treatment, VLS	>24,000	<ul> <li># enrolled in HIV recent infection surveillance program since 2019, 15 hotspot response activities, including community dialogues</li> </ul>
	~10,000	Average number of men receiving VMMC annually 2010-2021
	~10,000	Average # adults initiating oral PrEP annually 2019- 2021

Source: Eswatini Ministry of Health Annual Program Reports: 2010-2021

#### Future Directions: Address Program Gaps Men lag behind women in awareness of HIV-positive status



### Gaps: Youth as a vulnerable population





#### Older Population (25+)



### **Gaps: Key populations**





- 2020 ZIMPHIA data in general population vs 2020 Zim BBS data
- 95's almost achieved in general population, <50% of KP know their HIV status
- Population VLS among KP lower than general population: continued potential of HIV transmission
- <50% KP had disclosed their PLHIV status to their partner

#### Source:

Zimbambwe HIV Population HIV Impact Assessment Survey (ZimPHIA 2020). Summary Sheet. Dec. 2020

HIV and STI biobehavioral survey among men who have sex with men, transgender women, and genderqueer individuals in Zimbabwe: Final Report, Aug 2020

### Conclusions



- There has been substantial progress towards achieving 95-95-95 targets in several African countries, and tangible
  decline in HIV incidence where these targets have been achieved
  - Yet gaps remain among youth, men, key populations
  - These gaps mean a substantial proportion of the PLHIV population is still capable of HIV transmission
- VLS is not a one-time goal, it must be maintained over a lifetime. PLHIV need comprehensive support over time and differentiated service delivery
  - Ensuring commodities are available to deliver 95-95-95 at scale: HIV test kits, ART, viral load testing
  - Navigating ART fatigue and side effects: long-acting injectable ART offers new opportunities
  - Rapid diagnosis of virological failure and ART resistance, and prompt switching of ART regimens
  - Prevention, diagnosis and management of co-morbidities: hypertension, diabetes, cancers, mental health
- Primary HIV prevention is important
  - Long-acting PrEP offers important new option
  - Continued scale up of VMMC
  - Continuing need for comprehensive STI management (including point of care testing) and wholistic reproductive health services

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