# HIV Prevention in People who Inject Drugs (PWID)

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**HPTN Annual Meeting 2023** 





### Disclosures

- Grants, products and honoraria from Gilead Sciences
- Grants, products and honoraria from Abbott Laboratories

### Why People who Inject Drugs?

#### The New Hork Times

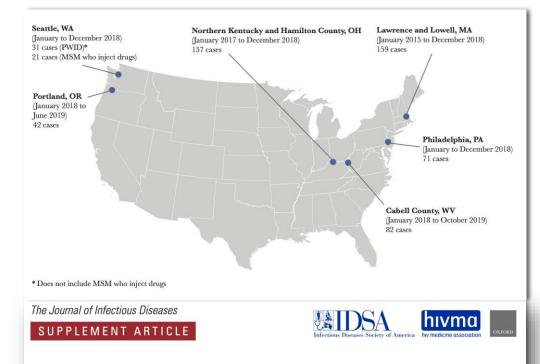
#### **Overdose Deaths Reached Record** High as the Pandemic Spread

More than 100,000 Americans died from drug overdoses in the yearlong period ending in April, government researchers said.

Give this article L\_ 847



A memorial service in Baltimore last year for a man who died of an overdose.



Responding to Outbreaks of Human Immunodeficiency Virus Among Persons Who Inject Drugs—United States, 2016-2019: Perspectives on Recent Experience and Lessons Learned Sheryl B. Lyss,<sup>1,2</sup> Kate Buchacz,<sup>1</sup> R. Paul McClung,<sup>1,2</sup> Alice Asher,<sup>3</sup> and Alexandra M. Oster<sup>1,2</sup>

Clinical Infectious Diseases MAJOR ARTICLE



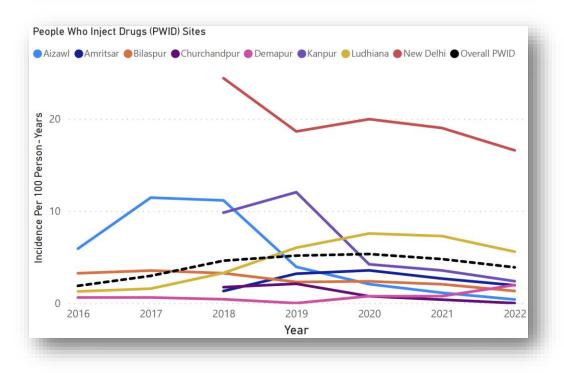
Estimated Number of People Who Inject Drugs in the **United States** 

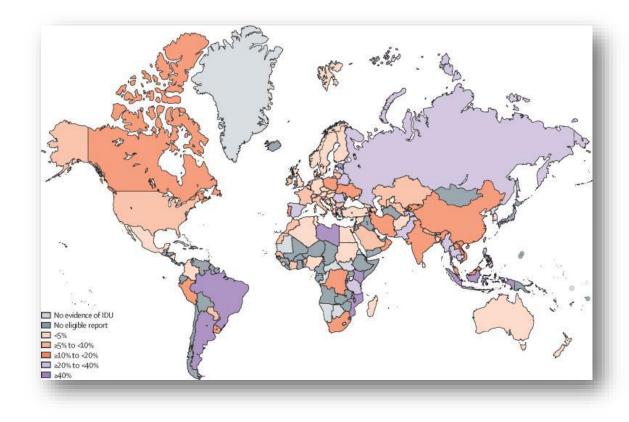
Heather Bradley,<sup>1</sup> Eric W. Hall,<sup>2</sup> Alice Asher,<sup>3</sup> Nathan W. Furukawa,<sup>3</sup> Christopher M. Jones,<sup>4</sup> Jalissa Shealey,<sup>1</sup> Kate Buchacz,<sup>3</sup> Senad Handanagic,<sup>3</sup> Nicole Crepaz,<sup>3</sup> and Eli S. Rosenberg<sup>5,6</sup>

# And not just in the US...

#### HIV incidence in a community-based cohort in New Delhi (2017-2020)

Number HIV negative	Person years of	Number of incident	Incidence rate
with ≥1 follow-up	follow-up	infections	(95% Cl)
782	747	159	21.3 (18.2 – 24.9)





#### Prevalence Estimates:

- South Africa (21% 35%)
- Kenya (15 21%)
- Tanzania (34%)

Clipman, Sci Adv 2022; McFall, CROI 2023; Degenhardt, Lancet Glob Health 2017; Scheibe, Harm Red J 20119; Zanoni, PLoS One 2023; Kurth, JAIDS 2015; Kawambwa, BMC Pun Health 2020

### My HPTN "all-time" favorites!!!

VOL. 365 NO. 6

#### The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

AUGUST 11, 2011

#### Prevention of HIV-1 Infection with Early Antiretroviral Therapy

Myron S. Cohen, M.D., Ying Q. Chen, Ph.D., Marybeth McCauley, M.P.H., Theresa Gamble, Ph.D., Mina C. Hosseinipour, M.D., Nagalingeswaran Kumarasamy, M.B., B.S., James G. Hakim, M.D., Johnstone Kumwenda, F.R.C.P., Beatriz Grinsztejn, M.D., Jose H.S. Pilotto, M.D., Sheela V. Godbole, M.D., Sanjay Mehendale, M.D., Suwat Chariyalertsak, M.D., Breno R. Santos, M.D., Kenneth H. Mayer, M.D., Irving F. Hoffman, P.A., Susan H. Eshleman, M.D., Estelle Piwowar-Manning, M.T., Lei Wang, Ph.D., Joseph Makhema, F.R.C.P., Lisa A. Mills, M.D., Guy de Bruyn, M.B., B.Ch., Ian Sanne, M.B., B.Ch., Joseph Eron, M.D., Joel Gallant, M.D., Diane Havlir, M.D., Susan Swindells, M.B., B.S., Heather Ribaudo, Ph.D., Vanessa Elharrar, M.D., David Burns, M.D., Taha E. Taha, M.B., B.S., Karin Nielsen-Saines, M.D., David Celentano, Sc.D., Max Essex, D.V.M., and Thomas R. Fleming, Ph.D., for the HPTN 052 Study Team\*

Couple	Index Case (HIV-infected)
<ul> <li>Reports a history of injection drug use within the last five years.</li> <li>Previous and/or current participant in an HIV vaccine study.</li> <li>Any condition that, in the opinion of the site investigator, would make participation in the study unsafe, complicate interpretation of study outcome data, or otherwise interfere with achieving the study objectives.</li> <li>Incarceration in a correctional facility, prison, or jail; and involuntary incarceration in a medical facility for psychiatric or physical (<i>e.g.</i> infectious disease) illness</li> </ul>	<ul> <li>Current or previous AIDS-defining illness</li> <li>Current or previous use of any ART drugs (some exceptions apply)</li> <li>Documented or suspected acute hepatitis within 30 days prior to enrollment, irrespective of AST (SGOT) and ALT (SGPT) values.</li> <li>Acute therapy for serious medical illnesses, in the opinion of the site investigator, within 14 days prior to enrollment. Candidates with chronic, acute, or recurrent infections that are serious, in the opinion of the site investigator, who must continue with chronic (maintenance) therapy (e.g., TB), must have completed at least 14 days of therapy prior to study entry and be clinically stable.</li> </ul>

#### HPTN 052 Study Exclusion Criteria

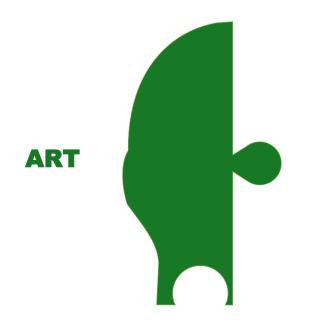
#### The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

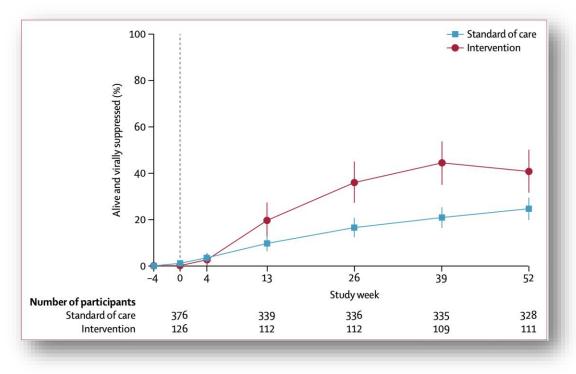
#### Cabotegravir for HIV Prevention in Cisgender Men and Transgender Women

R.J. Landovitz, D. Donnell, M.E. Clement, B. Hanscom, L. Cottle, L. Coelho,
R. Cabello, S. Chariyalertsak, E.F. Dunne, I. Frank, J.A. Gallardo-Cartagena,
A.H. Gaur, P. Gonzales, H.V. Tran, J.C. Hinojosa, E.G. Kallas, C.F. Kelley,
M.H. Losso, J.V. Madruga, K. Middelkoop, N. Phanuphak, B. Santos, O. Sued,
J. Valencia Huamaní, E.T. Overton, S. Swaminathan, C. del Rio, R.M. Gulick,
P. Richardson, P. Sullivan, E. Piwowar-Manning, M. Marzinke, C. Hendrix, M. Li,
Z. Wang, J. Marrazzo, E. Daar, A. Asmelash, T.T. Brown, P. Anderson, S.H. Eshleman,
M. Bryan, C. Blanchette, J. Lucas, C. Psaros, S. Safren, J. Sugarman, H. Scott, J.J. Eron,
S.D. Fields, N.D. Sista, K. Gomez-Feliciano, A. Jennings, R.M. Kofron, T.H. Holtz,
K. Shin, J.F. Rooney, K.Y. Smith, W. Spreen, D. Margolis, A. Rinehart, A. Adeyeye,
M.S. Cohen, M. McCauley, and B. Grinsztejn, for the HPTN 083 Study Team\*

minute.<sup>16</sup> Cisgender MSM and transgender women who have sex with men who were recruited for the trial were at high risk for HIV infection, as defined in the protocol. Key exclusion criteria were the use of illicit intravenous drugs within 90 days before enrollment, previous participation in the active treatment group of an HIV vaccine trial, coagulopathy, buttock implants or fillers, a



## Viral suppression among PWID



#### **HPTN 074** Peer navigation + Counseling + ART

#### Viral suppression over time

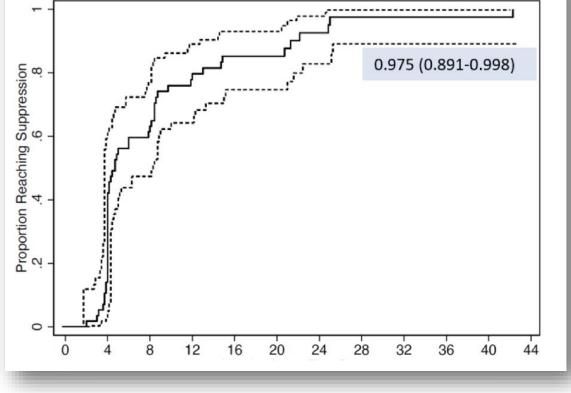


ICC+ (Lucas/Mehta) Outreach workers + contingency management

#### Miller, Lancet 2018; Solomon, CROI 2022

### **Role of LA ARVs**

**Figure**: KM curve of probability of reaching virologic suppression (VL <30) on LA ART (n=57); dotted lines 95% CI



Oh, no, no, no, no, no. The only one that sticks me with needles is me and my tattoo artist. [LAUGH] Oh no, I'll stick with the pill Providence, cisgender man

So, see them then, get injected or get the shot, and go on and not worry about it for a couple months. That's awesome. That's better than taking something every day. Providence, cisgender woman

## **Resistance: Baseline vs. follow-up**



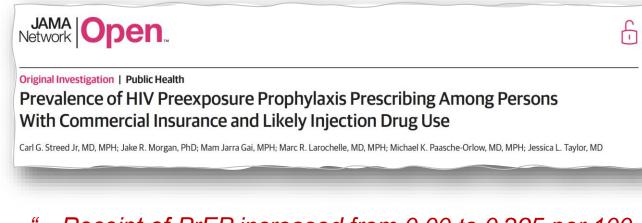
- On-treatment resistance:
  - Cyclical patterns of injection, cessation, relapse
  - Long tail of injectable ARVs
- Transmitted drug resistance?

#### Trials with long-term follow-up



# **PrEP uptake in PWID**

- Ideal population for PrEP
  - Low viral suppression
  - High incidence
  - Challenge with daily oral adherence
    - LA PrEP

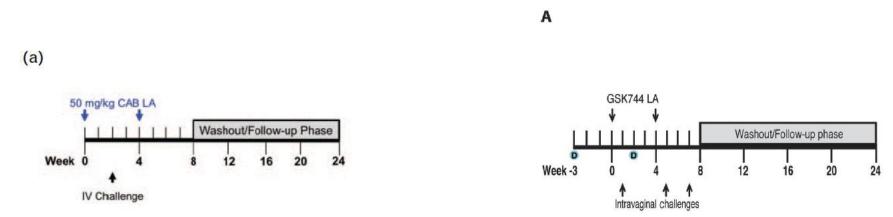


"...Receipt of PrEP increased from 0.00 to 0.295 per 100 person-years between 2010 and 2019 among PWID...."





### **Evidence for LA PrEP in PWID**



**Results:** CAB long acting was highly protective with 21 of the 24 CAB long-actingtreated macaques remaining aviremic, resulting in 88% protection. The plasma CAB concentration at the time of virus challenge appeared to be more important for protection than sustaining therapeutic plasma concentrations with the second CAB long acting injection.

**Conclusion:** These results support the clinical investigation of CAB long acting as PrEP in people who inject drugs. Copyright © 2017 Wolters Kluwer Health, Inc. All rights reserved.

Intravenous Challenge

**Intravaginal Challenge** 

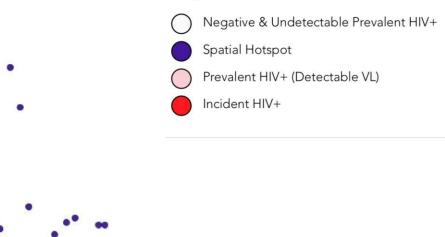
# LA PrEP in PWID: Way forward...

- Drug-drug interaction data
  - Commons substances of misuse
  - Medications for opioid use disorder
- Efficacy data in humans
  - Phase III trials (?unbalanced allocation)
    - Incidence still high among PWID in many settings globally
- Delivery of LA PrEP
  - MOUD/OTP centers
  - Field/van based
    - Could we target high transmission venues?

#### Sociospatial Network

Interact with this figure online: <u>Tiny.cc/sociospatial</u>

#### Legend



**113 (71%)** of incident infections reported injecting at venue #40

For every increased step in path to venue #40, risk of HIV acquisition reduced by 23% (IRR: 0.77)

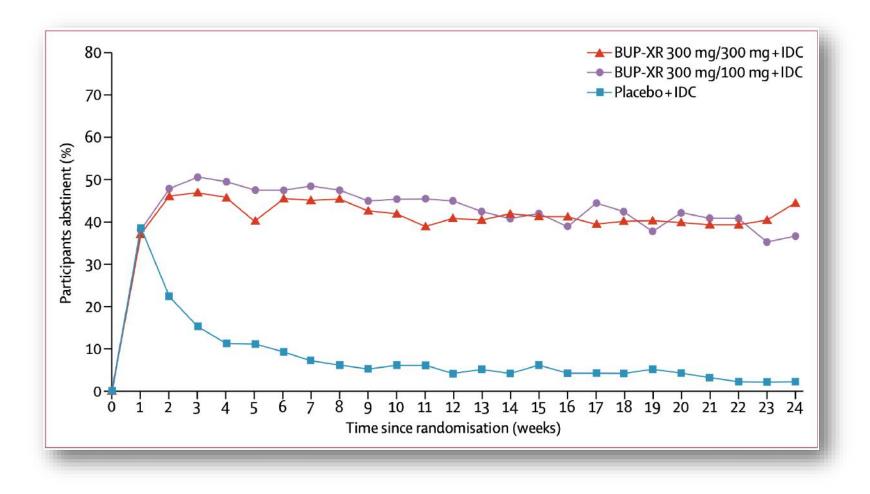
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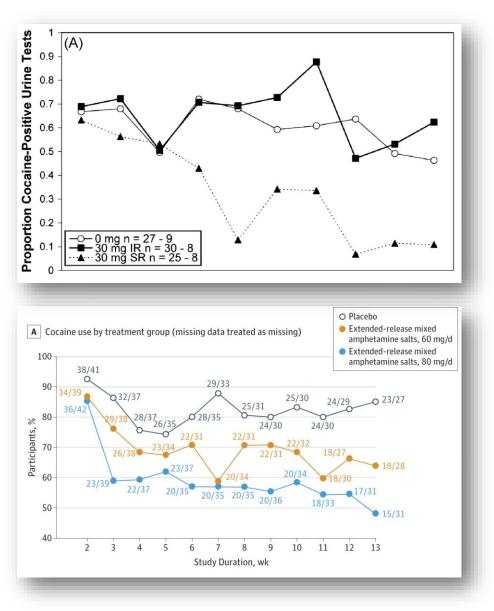
### **Management of substance use**

- Syringe service programs
- Fentanyl test strips
- Access to naloxone (Narcan)
- Medically supervised injection centers/sites
- Cognitive behavioral therapy
- Pharmacotherapy:
  - Buprenorphine
  - Methadone
  - No FDA approved agents for stimulants

#### **Long-acting buprenorphine**

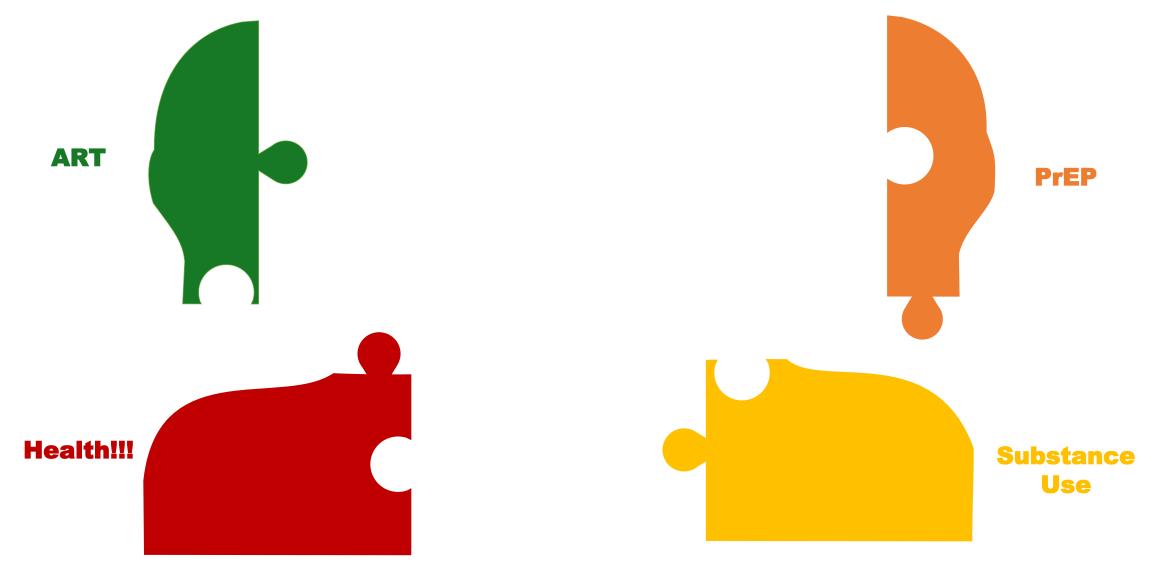


### **Treating stimulants with stimulants?**

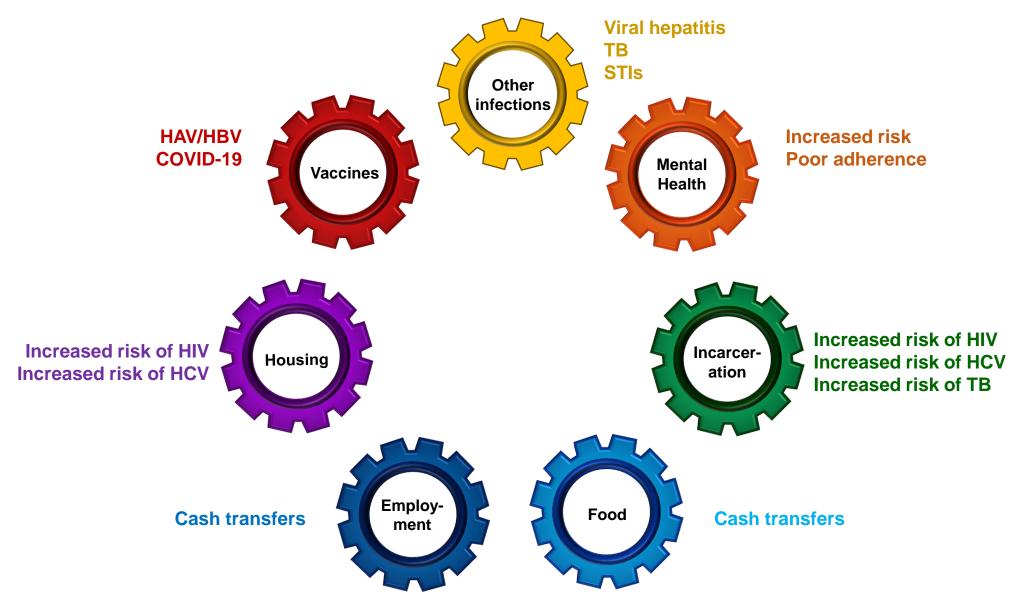


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+ Community Reinforcement Approach	i	3.92 (1.94, 1
		3.63 (2.01, 6
		4.84 (0.75, 3
Contingency Management		1.82 (0.37, 8 1.99 (0.48, 8
+ 12-Step Programme		2.47 (0.72, 8
+ 12-Step Programme		1.97 (0.63, 6
		3.27 (0.49, 2
Community Reinforcement Approach		1.25 (0.31, 5
		1.23 (0.33, 4
+ Non-Contingent Rewards	i i i i i i i i i i i i i i i i i i i	4.32 (1.47, 1 1.61 (0.63, 4
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contingency management	i	1.39 (1.09, 1
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		0.83 (0.22, 3
+ Non-Contingent Rewards		1.04 (0.34, 3 0.79 (0.27, 2
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and the second second and second and second at the		1.37 (0.48, 3 1.10 (0.23, 5
Meditation Based Therapies		0.86 (0.42, 1
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		1.17 (0.79, 1
Cognitive Behavioural Therapy		1.06 (0.75, 1
		1.42 (1.05, 1 1.47 (1.08, 2
		0.89 (0.53, 1
		0.86 (0.52, 1
Non-Contingent Rewards	!	0.59 (0.40, 0
	·	1.93 (1.27, 2
		1.76 (1.16, 2
		0.63 (0.24, 1
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		stinence at the longest follow
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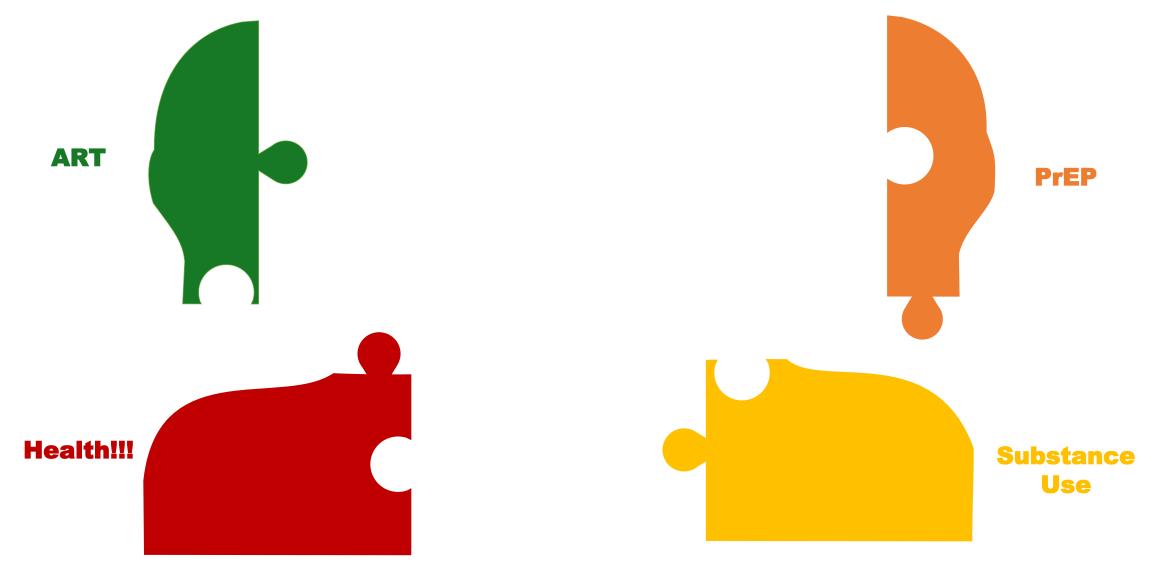
Mooney, DAD 2009; Levin, JAMA Psych 2015; Brandt, Addiction 2020; De Crescenzo, PLoS Med 2018



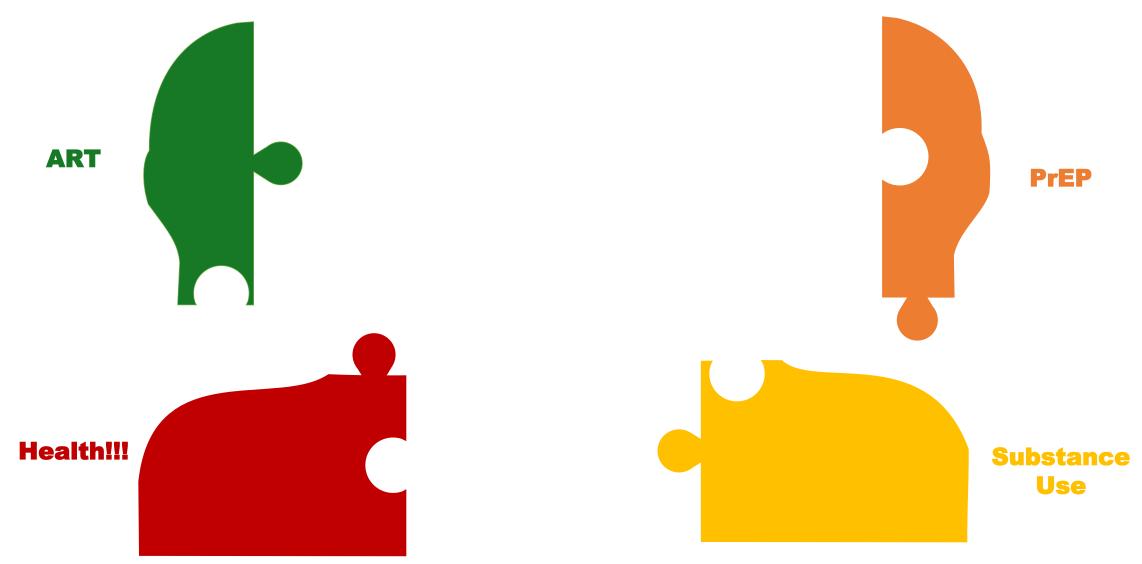
### More to health than HIV...



Altice, Lancet 2016; Stone, Lancet Public Health 2022; Richetrman, Nature 2023;



#### "Status-Neutral Whole Person"



# Acknowledgements

- People who graciously participate in research studies globally
- Johns Hopkins University
  - Shruti Mehta, Gregory Lucas, Steven Clipman, Carl Latkin, Allison McFall, David Celentano
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  - Abbott Laboratories
  - Gilead Sciences
  - Elton John AIDS Foundation
  - USAID/PEPFAR

# THANK YOU