TOWARDS HIV ELIMINATION

The XII IAS (International AIDS Society) conference on HIV science VII 2023, Brisbane

(translated from Latvian)

"We are acknowledging the traditional owners of the lands on which we are meeting and paying our respect to the elders past, present and emerging" was a usual start to each of the sessions in Brisbane.

This bi-annual conference gathered more than <u>5000 scientists</u>, clinicians, public health experts, as well as community educators from all the continents.

Maybe <u>Australia</u> was chosen as it **has reached the 90-90-90** (90% of PWH aware of their HIV positive status, 90% of those on ART, and 90% of those on ART are virally suppressed) targets in 2020 already and is on track to achieve the UNAIDS 95-95-95 targets by 2025.

Anecdotally, some HIV specialists in major Australian urban areas say they rarely see new diagnoses!

But not Australia alone. <u>Close to reaching the 95-95-95</u> global targets <u>are Botswana, Eswatini, Rwanda, Tanzania, Zimbabwe, and 16 other countries</u> as well!

The only region in the world with a <u>rapidly growing HIV epidemic</u> is E Europe and C Asia. There was a special Gilead satellite on Monday: "**E Europe** and C Asia **in crisis**".

At the conference, **WHO** released the **new scientific** and normative **guidance** on HIV. Among others, it states that **viral load (VL) is**

- <u>Undetectable</u> = no measurable virus = 0 risk of transmission, minimal risk of vertical transmission:
- <u>Suppressed</u> = ≤1000 copies/ml = almost 0 or negligible risk of transmission;
- <u>Unsuppressed</u> = >1000 ,, = increased vulnerability of becoming ill and transmitting HIV to sexual partner(s) and/or children.

ART

<u>DOR + ISL</u> (the approved NNRTI doravirine + investigational NRTTI islatravir) are non-inferior to BIC/FTC/TAF (bictegravir/ emtricitabine/tenofovir alafenamide) for initial treatment of HIV-1 and was generally well-tolerated (oral LBX0102).

Switching to <u>injectable long-acting CAB+RPV</u> was associated with improved treatment satisfaction, while also providing relief from the fear of disclosure and anxiety surrounding adherence (*poster exhibition TUPEB06*).

<u>Long-acting lenacapavir</u> (*LEN*) is approved for multidrug-resistant HIV-1 in combination with other antiretrovirals for heavily treatment-experienced individuals (*e-poster EPB0230*).

A POTENTIAL HIV CURE

"Geneva Patient" was diagnosed with HIV decades ago, but has been in HIV remission (i.e., still undetectable) for 20 months without ARV following a stem cell transplant – and whose donor lacked the rare stem cell mutation (i.e., had normal or "wild type" stem cells!) that has been linked to all known HIV cure cases (total of 5) to date (track A, LB5819).

There have been several children maintaining undetectable VL for months/years off therapy: a Mississippi baby, S African boy, and cases from Texas and France.

A study of 281 mother-child pairs monitored from delivery following in utero HIV transmission was conducted in S Africa, KwaZulu-Natal. Infants received cART prior to birth via placental transfer and received ART at birth. Exceptionally, **five boys** were identified in whom <u>aviremia was maintained</u> (for 3-19 months) despite persistent cART non/bad adherence (oral OALBX0104).

Another novel class of drugs for the potential development of an HIV cure strategy is DPP9 inhibitors. Besides HIV-1 infected cells, **the DPP9 inhibitor Val-boroPro** (*VbP*) was also able to significantly increase the clearance of HIV-1 latent reservoirs from PWH, **eliminating 75% of latent reservoirs**. This strategy has the capacity to clear HIV-1 infected cells in vitro, ex vivo, and in vivo (poster exhibition TUPEA11).

STATIN BENEFITS!

The biggest story of the conference was the first large-scale clinical study to test a primary cardiovascular prevention strategy in PWH (having low to moderate risk of cardiovascular disease), REPRIEVE.

The study's participants who took a daily statin (in this study: pitavastatin calcium: its patent ends in 2023) lowered their risk of major adverse cardiovascular events by 35% compared with those receiving a placebo. Adverse drug events observed in the study were like those in the general population taking statins. In short, statin therapy prevents major CV events in PWH with low/moderate CV risk and normal-range LDL cholesterol (symposium SY0601).

FRAILTY TRANSITION

A study among virologically suppressed PWH aged ≥50 years in Bangkok used 5 frailty criteria (weight loss, low physical activity, exhaustion, weak grip strength, and slow gait speed). Weak grip strength was the predominant frailty phenotype characteristic at year 5. Excessive alcohol consumption, longer duration of ART, and NNRTI regimen at baseline (compared to PI or INSTI) were associated with worsening of the frailty stage. Within 5 years frailty of ~1/4 of PWH worsened. However, the majority of frail PWH at baseline showed improvements in severity, suggesting frailty can be reversible (e-poster EPB0203).

STIs

While STIs occur frequently in MSM/HIV+, guidelines' recommendations do not include screening for asymptomatic people. A study in Mexico investigated the **prevalence of asymptomatic STIs** in these patients. At least one microorganism was detected in 23% of urethral samples, and more than one – in 12% of patients. The primary reason that gonorrhea and chlamydia infections are untreated is that infected people never have symptoms. Regular screening should be considered in PWH at least every 6-12 months: *EPB0142*

PREVENTION

A novel HIV prevention method for women: the **Dopivirine Vaginal Ring** is the <u>first long-acting</u>, woman-controlled HIV prevention product recommended by the WHO (*e-poster EPD0601*).

Tenofovir douche was well tolerated in young males and showed no adverse events. **TFV douche as PREP** to prevent HIV acquisition via receptive anal intercourse would fill a critical gap in pre-exposure prophylaxis product availability (*e-poster EPC0360*).

COVID-19 VACCINATION

WHO Global Clinical Platform (>800.000 children & adults) showed that PWH had a

- **54% higher risk of death** during the pre-Delta variant wave,
- 56% during Delta variant wave, and
- 142% during Omicron variant wave

compared to the HIV-negative population, with the risk being higher among those with CD4=200.

PWH with at least one dose of COVID-19 vaccination had

- 39% lower risk of death during the Delta variant wave and
- 38% during the Omicron variant wave

compared to the unvaccinated.

While the mortality risk among HIV-negative people decreased in the Omicron wave, only a modest reduction was observed in PWH. This highlights the need to implement **WHO guidelines recommending booster vaccines for** populations most at risk of severe COVID-19 outcomes, including **PWH** (oral OALBC0604).

Undisentangeably yours – A. Kalnins, AGIHAS