Response to Screening of diabetes mellitus among people living with HIV – a comment on “Diabetes mellitus burden among people living with HIV from the Asia-Pacific region” (Han et al. 2019)

Win M Han1§, Awachana Jiamsakul2, Sasisopin Kiertiburanakul3, Jeremy Ross4 and Anchalee Avihingsanon1,5

§Corresponding author: Win M Han, HIV-NAT, Thai Red Cross AIDS Research Centre, 104 Ratchadamri Rd., Pathumwan, Bangkok, Thailand 10330. Tel: 662 652 3040. (win.m@hivnat.org)

Keywords: diabetes; HIV; screening; viral suppression; Asia-Pacific

Dear Editor,

We appreciate the opportunity to respond to a Letter to the Editor about our recently published article entitled “Diabetes mellitus burden among people living with HIV from the Asia-Pacific region” [1]. The letter notes that the sensitivity of the screening tests used in our study differed substantially, and the author of the letter favoured haemoglobin A1c (HbA1c) and the two-hour plasma glucose level after an oral glucose tolerance test (OGTT) over fasting blood glucose (FBG). The author also highlighted the discrepancies in the proportions of patients diagnosed with diabetes mellitus (DM) by each of the screening tests used in our study and that most of the DM patients were diagnosed by FBG.

While we fully acknowledge the differing sensitivities of these screening tests in diagnosing DM, we want to emphasize that the data analyzed in our observational cohort study were collected from routine clinical care in participating sites. Not all patients included in our analysis would have undergone all mentioned DM screening tests, as test availability and use have varied across the sites in the cohort over time. FBG has been the most widely accessible test, so it would be expected that FBG data availability was greater than that of HbA1c and OGTT in our cohort. The differences in the number of participants diagnosed by FBG compared to HbA1c and OGTT may reflect the later inclusion of HbA1c and OGTT into our cohort data collection as well as differential costs associated with those tests in the context of routine care. Notably, study investigators did not select which tests would be used a priori, and we acknowledge that this as a limitation of the study analysis. However, we believe this does not alter our recommendation for improved monitoring and routine screening of non-communicable diseases including DM among people living with HIV (PLHIV) on long-term ART using available laboratory testing.

The letter stated that HbA1c screening identified more cases of DM than FBG in studies in Vietnam [2] and Taiwan [3], we would note that participants from both of these studies were from HIV-negative populations. More studies are recommending the use of FBG testing as the primary screening tool to diagnose DM among PLHIV due to the potential risk of HbA1c underestimating glycaemia in some PLHIV (e.g. on certain ART regimens, who have a higher mean corpuscular volume of red blood cells and lower CD4 counts) [4-7]. We do agree with the author of the letter that testing beyond FBG would be useful for DM screening among PLHIV who are receiving chronic HIV care, and will monitor test availability and reliability in future studies of cardiometabolic outcomes in our cohort.

AUTHORS’ AFFILIATIONS
1HIV-NAT/Thai Red Cross AIDS Research Centre, Bangkok, Thailand; 2The Kirby Institute, UNSW Sydney, Sydney, Australia; 3Faculty of Medicine Ramathibodi Hospital, Mahidol University, Bangkok, Thailand; 4TREAT Asia, amfAR - The Foundation for AIDS Research, Bangkok, Thailand; 5Faculty of Medicine, Chulalongkorn University, Bangkok, Thailand

COMPETING INTERESTS
The authors have no conflicts of interest to declare.

AUTHORS’ CONTRIBUTIONS
WMH and AJ drafted the response. WMH, AJ, SK, JR and AA reviewed and provided revisions to the draft prior, before WMH circulated the letter for approval from all authors from the original article.

ACKNOWLEDGEMENTS
We would like to thank the journal editors and the author of the original letter for giving us the opportunity to discuss this important topic. We particularly thank Dr Annette Sohn for helpful discussions and comments in preparing this response.
REFERENCES

1. Han WM, Jiamsakul A, Kiertiburanakul S, Ng OT, Sim BLH, Sun LP, et al. Diabetes mellitus burden among people living with HIV from the Asia-Pacific region. J Int AIDS Soc. 2019;22(1):e25236.

2. Ho-Pham LT, Nguyen UDT, Tran TX, Nguyen TV. Discordance in the diagnosis of diabetes: comparison between HbA1c and fasting plasma glucose. PLoS ONE. 2017;12(8):e0182192.

3. Wang JS, Lee IT, Lee WJ, Lin SY, Fu CP, Lee WL, et al. Comparing HbA1c, fasting and 2-h plasma glucose for screening for abnormal glucose regulation in patients undergoing coronary angiography. Clin Chem Lab Med. 2015;53(9):1441–9.

4. Kim PS, Woods C, Georgoff P, Crum D, Rosenberg A, Smith M, et al. A1C underestimates glycemia in HIV infection. Diabetes Care. 2009;32(9):1591–3.

5. Slama L, Palella FJ Jr, Abraham AG, Li X, Vignal C, Pialoux G, et al. Inaccuracy of haemoglobin A1c among HIV-infected men: effects of CD4 cell count, antiretroviral therapies and haematological parameters. J Antimicrob Chemother. 2014;69(12):3360–7.

6. Monroe AK, Glesby MJ, Brown TT. Diagnosing and managing diabetes in HIV-infected patients: current concepts. Clin Infect Dis. 2014;60(3):453–62.

7. Coelho AR, Moreira FA, Santos AC, Silva-Pinto A, Sarmento A, Carvalho D, et al. Diabetes mellitus in HIV-infected patients: fasting glucose, A1c, or oral glucose tolerance test - which method to choose for the diagnosis? BMC Infect Dis. 2018;18(1):309.