

Strengthening the HIV cascade to ensure an effective future ART response in sub-Saharan Africa

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Timely linkage to antiretroviral therapy (ART) care is critical for reducing HIV-related morbidity, mortality and transmission. Studies investigating interventions to improve linkage to, and retention in, pre-ART care in sub-Saharan Africa were reviewed. Certain interventions used to overcome economic barriers for ART-patients (i.e. integration of services, medical and food incentives, intensified counselling and peer support) have also shown favourable results in the pre-ART period. A combined package of interventions found to be effective in the pre-ART and ART period might be effective for reducing attrition in both periods. Further operational research in this area is needed to identify local solutions.

Keywords: ART, Attrition, HIV, Loss to follow-up, Pre-ART, Retention

Sub-Saharan Africa is home to the largest HIV cohorts in the world, with antiretroviral therapy (ART) programmes now nearly a decade old, serving over 30 000 HIV-infected individuals.^{1,2} As ART services have been brought to scale several programmatic challenges have emerged, including suboptimal rates of HIV testing, ART adherence and retention in ART care.^{3,4} Currently, HIV programmes are faced with an additional challenge of delayed linkage to HIV and ART care and high pre-ART attrition among HIV-infected individuals, which is hindering further scale-up of these programmes and attainment of universal coverage.⁵⁻⁷

Timely linkage to ART care is critical for reducing HIV-related morbidity and mortality. In addition, there is increasing interest in treating people earlier in their disease progression as a way to reduce HIV transmission. This strategy of 'treatment as prevention' is primarily aimed at increasing the proportion of patients on ART who are virologically suppressed and at a negligible risk of transmitting the virus on to others.⁸ The success of this strategy hinges on the ability to treat HIV-infected individuals before their HIV-RNA viral loads increase, ensure timely linkage to ART care, and maintain patients on effective lifelong treatment.^{9,10} However, ART-eligible individuals often present late to clinical services and despite the expanding ART-eligibility threshold from <200 to <350 cells/ μ l (and when the 2013 WHO guidelines are implemented, to <500 cells/ μ l),¹¹ the median CD4 count at ART initiation remains low (81–120 cells/ μ l),¹² with one in four

individuals starting ART with a CD4 count <100 cells/ μ l.¹¹ Thus, currently most individuals in sub-Saharan Africa are commencing ART after a period of sustained high viral load. While much of the focus on treatment as prevention has been on earlier ART initiation, the success of this approach in reducing HIV transmission is also dependent on long-term retention in care, yet retention in care has been estimated to be 60% at 2 years, after treatment initiation, in ART programmes in sub-Saharan Africa,⁵ with long-term retention of only 38% at 7 years in an urban clinic in South Africa.¹³

Three systematic reviews have documented the high attrition occurring in the period between HIV diagnosis and ART initiation.⁵⁻⁷ Similarities in the barriers to linkage to, and retention in, care exist for both ART-ineligible and -eligible individuals. Getting to the clinic and ensuring regular clinic attendance is a common problem encountered by both groups of individuals as it is challenged by high transport costs, long distances and other competing priorities.¹⁴ Certain interventions used to overcome these economic barriers for patients on ART (i.e. integration of care, 15,16 treatment supporters, 17 nutrition support $^{18-20}$) have also been evaluated for the pre-ART care period and have shown favourable results. Integration of health services reduces the number of trips an HIV-infected individual has to make to the clinic. A study conducted in Zambia which integrated ART and antenatal-care found a twofold-increase in the risk of enrolling in HIV care and ART initiation.²¹ Similarly, increase in linkage

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COMMENTARY

to ART care was also observed in a study conducted in Cambodia, in which the impact of an improved referral, communication and teaching intervention among co-infected TB patients was assessed.²² Intensified post-test counselling and enhanced peersupport in Uganda was also found to increase the proportion of HIV-infected individuals enrolling into pre-ART care, following diagnosis, by 29%.²³ In addition, two studies from Kenya have shown that regular home visits by a patient navigator increases linkage to HIV care,²⁴ and provision of cotrimoxazole to encourage regular pre-ART clinic attendance doubled pre-ART retention. Of note, food incentives increased regular attendance in children²⁶ and decreased time to ART initiation among drug users in India,²⁷ although these interventions have yet to be assessed in sub-Saharan Africa. Interventions to overcome the main health system barriers such as an efficient appointment and patientmonitoring system, providing after-hours and weekend clinics need further investigation, as this could potentially overcome obstacles faced by both ART-ineligible and -eligible individuals.

Strengthening the existing HIV cascade through improving linkage to, and retention in, pre-ART and ART care is, therefore, critical to ensuring successful future ART scale-up and to maximise the benefits of long-term ART, including reduced HIV transmission. Integration of services, enhanced counselling, peer support and referral as well as medical and food incentives have been found to be effective in improving both linkage to, and retention in, pre-ART and ART care. These interventions could be implemented simultaneously to target both periods of care, although 'one size' is unlikely to 'fit all' and a combined package of interventions might be more effective. Further operational research in this area is urgently needed in order to identify local solutions that work to support the future effectiveness of the ART response.

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References

- 1 Garone DB, Hilderbrand K, Boulle AM et al. Khayelitsha 2001–2011: 10 years of primary care HIV and TB programmes. Southern African Journal of HIV Medicine 2011;12:33–8.
- 2 Fox MP, Maskew M, MacPhail AP et al. Cohort Profile: The Themba Lethu Clinical Cohort, Johannesburg, South Africa. Int. J. Epidemiol 2013;42:430–9.
- 3 Mills EJ, Nachega JB, Buchan I et al. Adherence to antiretroviral therapy in sub-Saharan Africa and North America. JAMA 2006;296:679–90.
- 4 Rosen S, Fox MP, Gill CJ. Patient retention in antiretroviral therapy programs in sub-Saharan Africa: a systematic review. PLoS Med 2007;4:e298.

- 5 Rosen S, Fox MP. Retention in HIV Care between Testing and Treatment in Sub-Saharan Africa: A Systematic Review. PLoS Med 2011;8:e1001056.
- 6 Mugglin C, Estill J, Wandeler G et al. Loss to programme between HIV diagnosis and initiation of antiretroviral therapy in sub-Saharan Africa: systematic review and meta-analysis. Trop Med Int Health 2012;17:1509–20.
- 7 Kranzer K, Govindasamy D, Ford N et al. Quantifying and addressing losses along the continuum of care for people living with HIV infection in sub-Saharan Africa: a systematic review. J Int AIDS Soc 2012;15:17383.
- 8 Granich RM, Gilks CF, Dye C et al. Universal voluntary HIV testing with immediate antiretroviral therapy as a strategy for elimination of HIV transmission: a mathematical model. Lancet 2009;373:48–57.
- 9 Gardner EM, McLees MP, Steiner JF et al. The spectrum of engagement in HIV care and its relevance to test-and-treat strategies for prevention of HIV infection. Clin Infect Dis 2011;52:793–800.
- 10 Kulkarni SP, Shah KR, Sarma KV, Mahajan AP. Clinical Uncertainties, Health Service Challenges, and Ethical Complexities of HIV "Test-and-Treat": A Systematic Review. Am J Public Health 2013;103:e14–23.
- 11 Global update on HIV treatment 2013: results, impact and opportunities. 2013. Geneva: World Health Organization. http://www.who.int/hiv/pub/ progressreports/update2013/en/index.html [accessed 04 November 2013].
- 12 Lahuerta M, Ue F, Hoffman S et al. The problem of late ART initiation in sub-Saharan Africa: a transient aspect of scale-up or a long-term phenomenon? J Health Care Poor Underserved 2013;24:359–83.
- 13 Fox MP, Shearer K, Maskew M et al. Treatment outcomes after 7 years of public-sector HIV treatment. AIDS 2012;26:1823–8.
- 14 Govindasamy D, Ford N, Kranzer K. Risk factors, barriers and facilitators for linkage to antiretroviral therapy care: a systematic review. AIDS 2011;26:2059–67.
- 15 Suthar AB, Hoos D, Beqiri A et al. Integrating antiretroviral therapy into antenatal care and maternal and child health settings: a systematic review and meta-analysis. Bull World Health Organ 2013;91:46–56.
- 16 Uyei J, Coetzee D, Macinko J, Guttmacher S. Integrated delivery of HIV and tuberculosis services in sub-Saharan Africa: a systematic review. Lancet Infect Dis 2011;11:855–67.
- 17 Franke MF, Kaigamba F, Socci AR et al. Improved retention associated with community-based accompaniment for antiretroviral therapy delivery in rural Rwanda. Clin Infect Dis 2013;56:1319–26.
- 18 Cantrell RA, Sinkala M, Megazinni K et al. A pilot study of food supplementation to improve adherence to antiretroviral therapy among food-insecure adults in Lusaka, Zambia. J Acquir Immune Defic Syndr 2008;49:190–5.
- 19 Posse M, Tirivayi N, Saha U, Baltussen R. The effect of Food Assistance on Adherence to Antiretroviral Therapy among HIV/AIDS Patients in Sofala Province, in Mozambique: A Retrospective Study. J AIDS Clin Res 2013;4:2.
- 20 Ivers LC, Chang Y, Gregory Jerome J, Freedberg KA. Food assistance is associated with improved body mass index, food security and attendance at clinic in an HIV program in central Haiti: A prospective observational cohort study. AIDS Res Ther 2010;7:33.
- 21 Killam WP, Tambatamba BC, Chintu N et al. Antiretroviral therapy in antenatal care to increase treatment initiation in HIV-infected pregnant women: a stepped-wedge evaluation. AIDS 2010;24:85–91.
- 22 Choun K, Pe R, Thai S et al. Timing of antiretroviral therapy in Cambodian hospital after diagnosis of tuberculosis: impact of revised WHO guidelines. Bull World Health Organ 2013;91:195-206.
- 23 Muhamadi L, Tumwesigye NM, Kadobera D et al. A single-blind randomized controlled trial to evaluate the effect of extended

counseling on uptake of pre-antiretroviral care in Eastern Uganda. Trials 2011;12:187.

- 24 Hatcher AM, Turan JM, Leslie HH et al. Predictors of linkage to care following community-based HIV counseling and testing in rural Kenya. AIDS Behav 2012;16:1295–307.
- 25 Kohler PK, Chung MH, McGrath CJ et al. Implementation of free cotrimoxazole prophylaxis improves clinic retention among antiretroviral therapy-ineligible clients in Kenya. AIDS 2011;25:1657–61.
- 26 Kundu CK, Samanta M, Sarkar M et al. Food supplementation as an incentive to improve pre-antiretroviral therapy clinic adherence in HIV-positive children-experience from eastern India. J Trop Pediatr 2012;58:31–7.
- 27 Solomon S, Srikrishnan AK, Vasudevan C et al. The impact of voucher incentives on linkage to care and ART initiation among drug users: Chennai, India; 19th Conference on Retroviruses and Opportunistic Infections 2012; Washington, USA. Poster number: 1145.