

Predictors of return to HIV care

Patterns and Predictors of Incident Return to HIV Care Among Traced, Disengaged Patients in  
Zambia: Analysis of a prospective cohort

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Conflicts of Interest and Sources of Funding: No authors have conflicts of interest to declare.

**d) Address for reprints:** same as that of corresponding author

**e) Meetings at which parts of the data were presented:** Conference on Retroviruses and Opportunistic Infections, 2019

Research reported in this publication was supported by the National Institute of Mental Health of the National Institutes of Health under Award Numbers F31MH109378. The content is solely the responsibility of the authors and does not necessarily represent the official views of the National Institutes of Health. This research was also supported by the Bill and Melinda Gates Foundation

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grant number OPP1105071 and the Johns Hopkins University Center for AIDS Research P30AI094189.

## Abstract

**Background:** Dynamic movement of patients in and out of HIV care is prevalent, but there is limited information on patterns of patient re-engagement or predictors of return to guide HIV programs to better support patient engagement.

**Methods:** From a probability-based sample of lost to follow-up, adult patients traced by peer educators from 31 Zambian health facilities, we prospectively followed disengaged HIV patients for return clinic visits. We estimated cumulative incidence of return and time to return using Kaplan Meier methods. We used univariate and multivariable Cox proportional hazards regression to conduct a risk factor analysis identifying predictors of incident return across a social ecological framework.

**Results:** Of the 556 disengaged patients, 73.0% (95% CI: 61.0-83.8) returned to HIV care. Median follow-up time from disengagement was 32.3 months (IQR: 23.6-38.9). The rate of return decreased with time post-disengagement. Independent predictors of incident return included a prior gap in care (aHR: 1.95, 95%CI: 1.23-3.09) and confronting a stigmatizer once in the past year (aHR: 2.14, 95%CI: 1.25-3.65). Compared to a rural facility, patients were less likely to return if they sought care from an urban facility (aHR: 0.68, 95%CI: 0.48-0.96) or hospital (aHR: 0.52, 95%CI: 0.33-0.82).

**Conclusions:** Interventions are needed to hasten re-engagement in HIV care. Early and differential interventions by time since disengagement may improve intervention effectiveness.

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Patients in urban and tertiary care settings may need additional support. Improving patient resilience, outreach after a care gap, and community stigma reduction may facilitate return. Future re-engagement research should include causal evaluation of identified factors.

**Key words:** HIV, Zambia, retention, antiretroviral therapy

## **Introduction**

Re-engagement in care is a critical but poorly understood step in the HIV care cascade globally<sup>1-4</sup>. Dynamic movement of patients in and out of care is prevalent<sup>2,4,5</sup>, making care interruptions part of the natural history of HIV treatment<sup>4,6</sup>. These interruptions put patients at risk of poor health outcomes<sup>7-9</sup> and onward transmission of HIV<sup>10-12</sup>. They threaten achievement of the global 95-95-95 targets<sup>13</sup>. However, return to care is a positive patient behavior which has the potential to improve treatment outcomes. Especially as the burden of undiagnosed disease continues to diminish, and time on treatment for the average patient increases, understanding how quickly disengaged patients return to care, what factors facilitate return, and ways to encourage more rapid return represents an important scientific agenda with a potentially significant magnitude of effect and public health relevance<sup>14</sup>.

HIV policy, service delivery and monitoring must recognize and account for dynamic patient movement in the HIV care cascade<sup>3,6,15,16</sup>. However, most extant literature focuses on the traditional, linear steps including testing, linkage, ART initiation and viral suppression<sup>17</sup>. Much less is known about patient re-engagement after a care-seeking absence. To date, the few return-to-care studies have been primarily retrospective and examined demographic and clinical

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characteristics only. These studies suggest that between one third and one half of patients with a gap in care have a return visit<sup>3,18</sup>, while studies including patient tracing observe return ranging from 20-70%<sup>19-21</sup>. Factors associated with return in studies from east and southern Africa include older age<sup>22</sup>, lower CD4 count<sup>19,22</sup>, female gender<sup>20,22</sup>, health facility outreach<sup>19,20,22</sup>, ART use<sup>19,22</sup>, and latent patient factors related to poverty and poor care quality<sup>21</sup>. Several additional factors were identified in a north American context, but not explored in studies conducted in African countries, including mental health concerns, secure housing and substance use<sup>18</sup>. Several qualitative studies have explored patients experiences, identifying factors such as reduced stigma and social support as important for care engagement<sup>21,23-25</sup>, but few studies measure these factors quantitatively to examine their association with return. There is a lack of prospective analyses of re-engagement that assess the effect of a comprehensive set of potential patient-related, clinical, and social influences on return.

To improve the understanding of re-engagement in HIV care and treatment in sub-Saharan Africa, our study prospectively identified incident return to HIV care and time to return among a representative sample of traced, lost to follow-up (LTFU) patients confirmed to be disengaged from care from 31 facilities across four provinces in Zambia. We conducted a risk factor analysis identifying predictors of return from a range of factors at the individual, social and facility levels. This analysis can inform future research and intervention development through patient re-engagement risk stratification and hypothesis generation around re-engagement support opportunities.

## **Methods**

### **Study Background and Procedures**

This analysis is nested within a larger study, 'Better Information for Health in Zambia' (BetterInfo)<sup>26,27</sup>. BetterInfo enumerated all LTFU adult patients at 31 sampled study facilities who had at least one HIV care visit between 1<sup>st</sup> August 2013 - 31<sup>st</sup> July 2015. Patients were determined to be LTFU if they were >90 days late for their last scheduled appointment and had a subsequent unknown care status. Approximately 10% of LTFU patients were then randomly sampled for BetterInfo study tracing. As described elsewhere<sup>26,27</sup>, sampled patients were traced by a peer educator using paper medical record review, phone calls and in-person visits to ascertain if the patient was: 1) deceased, 2) alive and in-care or 3) alive and out of care. All contacted, disengaged patients were verbally encouraged to return to care, and, while not systematically applied, in some cases tracers accompanied the returning patient to the facility or met them for their return visit. Upon in-person patient contact, tracers obtained voluntary written informed consent and used tablet computers to administer a survey recording care status, demographic, social, behavioral and household characteristics, and reported barriers to care engagement. No medical care was administered during the tracing interaction. BetterInfo study surveys were administered in Nyanja, Bemba, Tonga or English based on patient preference. Our nested study then extracted approximately two and a half years of follow-up HIV visit data after the cohort closed using electronic medical records (EMR) linked through unique patient identifiers. (Supplemental Figure 1, <http://links.lww.com/QAI/B562>)

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### Study Population

Our analysis included all out of care patients identified through BetterInfo tracing who: a) confirmed that they did not have an HIV care visit since the last one identified in their medical record, b) completed the study survey at time of tracing, and c) were interviewed after their estimated date of disengagement (>90 days from last scheduled appointment based on paper medical record review, as recorded in the study database).

### Measurements

Our study outcome, return to care, was obtained from facility visit dates in the EMR follow-up data extraction. Potential predictors of return (Supplemental Figure 2, <http://links.lww.com/QAI/B562>) including clinical characteristics at the time of LTFU (e.g. CD4 count, time in HIV care, facility-type) and gaps in care of >90 days prior to the BetterInfo study-identified gap were gathered from the patient's EMR at LTFU. All demographic (e.g. age, marital status), social (e.g. HIV status disclosure, stigma), behavioral (e.g. alcohol use, travel) and household (e.g. wealth, violence tolerance) factors potentially predictive of return were taken from the patient survey administered by the tracer. Missing survey items were taken from the EMR, if available (e.g. age, marital status). Most potential predictors were measured using closed-ended yes/no, multi-choice or Likert scale questions. To capture patient reasons for disengagement, changes needed to return, and return intentions, however, tracers asked the open-ended questions, 'Why did you stop going to any clinic for HIV care?' and 'What would have to happen for you to come back to care at any clinic?', listened to the response, and recorded tick marks in as many pre-defined response options as were consistent with the patient's reply. Pre-

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defined sub-categories developed through prior research<sup>28</sup> included 'structural' (e.g. transport, work issues), 'psychosocial' (e.g. need encouragement, family, disclosure issues), 'clinic' (e.g. poor care quality, wait too long), and 'medical' (e.g. felt well, too many pills) options, each of which had 4-13 detailed response options. The response category 'other' captured responses that did not fit under the pre-defined options.

## Analysis

### *Potential predictors of return*

Analysis of possible predictors of return to care was guided by an adapted social ecological conceptual framework<sup>29</sup> of incident patient return to HIV care developed using extant literature<sup>2,18-25,28,30-32</sup> and contextual knowledge (Supplemental Figure 2, <http://links.lww.com/QAI/B562>). To model potential predictor variables, we first assessed the distribution of categorical variables, excluding variables where  $\geq 97\%$  of responses were the same. We assessed the relationship between continuous variables and return (on the log odds scale) using LOWESS plots. Time from enrollment to disengagement was dichotomized at 18 months based on the LOWESS plot. From our 18 stigma questions developed to be consistent with draft and final HPTN 071<sup>33</sup> stigma questions, we used exploratory and confirmatory factor analysis to identify four stigma sub-scales with adequate internal consistency: internalized (Cronbach  $\alpha=0.70$ ), anticipated (Cronbach  $\alpha=0.87$ ), experienced (Cronbach  $\alpha=0.72$ ) and resilience (single question: 'I confronted, challenged, or educated someone stigmatizing and/or discriminating against me'). Stigma sub-scale scores were summed from item responses. For internalized stigma, patient responses were dichotomized as low versus high at the median scale



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score. Experienced stigma was dichotomized as none versus any, and anticipated stigma was broken into approximate tertiles. Household wealth was estimated from ownership of 14 possible household items using the Demographic and Health Survey wealth index approach<sup>34</sup> and broken into tertiles. Household violence tolerance scores were summed, with one point for each positive response to the two, yes/no questions previously used in HIV research in Zambia, 'If someone in the household misuses money is it acceptable to beat him/her?' and 'In my household if a wife comes home late without permission of the husband, she will be beaten.'<sup>35</sup> Alcohol use was analyzed using the AUDIT-C<sup>36</sup> binge drinking question. For 'patient reasons for disengagement' and 'needs to return', participants were analyzed as 'yes' for a particular sub-category of 'reason for stopping' or 'need to return' if  $\geq 1$  detailed response option was selected for that patient under the specified sub-category. Sub-categories were not mutually exclusive<sup>28</sup>.

We used descriptive statistics to assess missingness. If a participant was missing data on binge drinking but replied that they drank ' $\geq 5$ -6 drinks on a typical day' on a separate AUDIT-C question<sup>36</sup>, their binge value was set to 'yes'. For stigma sub-scales, we imputed the mean of available sub-scale items for a missing sub-scale item if at least two sub-scale items were available. We used multiple imputation with chained equations and 10 imputed data sets to account for remaining missing predictor data in the multivariable model.

## *Disengaged Patient characteristics*

We described the disengaged study population by potential predictors of return and used Kaplan-Meier methods to estimate cumulative incidence of and time to return.

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### *Return to care*

We used Cox proportional hazards regression to estimate incident return to care. The time origin was the date of disengagement from care (90 days from last appointment or 180 days from last visit if the appointment date missing). The time scale was days since disengagement. Study entry was the date of in-person tracer contact, the point at which a patient was confirmed to be out of care. The event, incident return to care, is defined as the first HIV visit date of any type (i.e. clinical, pharmacy or laboratory) on or after the date of in-person tracing contact. Patients were censored at database closure.

We first examined the complete case, univariate association of each potential predictor with return to care. The final multivariable model was informed by theory (Supplemental Figure 2, <http://links.lww.com/QAI/B562>), including the following variables: gender, age, CD4 count at last visit, time in HIV care, past care gaps, past facility outreach for return, facility type, mobility (having to travel for >1 month in the past year), and having a psychosocial reason for disengagement or psychosocial need to return. We additionally included factors with a univariate association significance of  $p < 0.05$ . We examined variance inflation factors to assess multicollinearity and examined Schoenfeld residuals and adjusted log-log plots for each covariate to assess the proportional hazards assumption.

### *Supplemental Analyses*

To better understand disengaged patients, we descriptively compared LTFU patients successfully traced and determined to be out of care to those found to be in-care.

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To better understand return within a shorter time period, we conducted a supplemental analysis for incident return to HIV care within one year of disengagement, following the same analytic approach outlined above. To support a smaller model (more appropriate for fewer outcomes), the final supplemental multivariable model included only sex, age and variables with a univariate association significance of  $p < 0.05$ . Acknowledging the important role of theory in a risk factor analysis, we also ran a multivariable model for return by one year with the theory-driven variables described above as a sensitivity analysis.

Analyses were conducted using Stata 15.1 IC (StataCorp, 2018) and Mplus 8.2 (Muthen & Muthen, 2018).

## *Ethical Review*

This study was approved by the University of Zambia Research Ethics Committee, the Zambian Ministry of Health, and the University of Alabama at Birmingham Institutional Review Board (UAB IRB). The Johns Hopkins University and University of California at San Francisco had reliance agreements with the UAB IRB.

## **Results**

### *Disengaged Patient Characteristics*

There were 556 patients identified through tracing as disengaged and included in our study sample (Supplemental Figure 3, <http://links.lww.com/QAI/B562>). Disengaged traced patients were 41.7% male, had a median age at disengagement of 33.6 years (IQR: 28.4-39.9,

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min: 18.5, max: 80.3) and median time in care prior to disengagement of 0.9 years (IQR: 0.4-2.6, min: 0.3, max: 10.7) (Table 1). The first supplemental analysis showed that, compared to LTFU patients successfully traced and determined to be in-care, disengaged traced patients were more likely to be male, younger, never married, to have had a higher CD4 count at last visit, not yet initiated ART and have been lost from a facility in Lusaka Province (Supplemental Table 1, <http://links.lww.com/QAI/B562>). Traditional healer contact was dropped from further analysis due to >97% of responses being the same.

### *Patterns of Return to Care*

Most disengaged traced patients, 73.0% (95% CI: 61.0-83.8) had a return HIV visit. Median follow-up time was 32.3 months (IQR: 23.6-38.9). The cumulative proportion of patients returning were 23.4% (95% CI: 6.5-65.7) by 90 days, 33.7% (95% CI: 14.2-66.7) by 180 days, and 51.4% (95% CI: 33.2-72.5) by 365 days (Figure 1A). The overall incidence rate of return is 0.73 per 1,000 person years (95% CI: 0.64-0.84), declining with time since disengagement and no additional returns after 3.5 years post-disengagement (Figure 1B). Among returners, the median time spent out of care was 19.1 months (IQR: 13.9-25.4).

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Univariate analyses indicate that disengaged, traced patients were significantly (p-value <0.05) more likely to return to care if they had been contacted more than three times by the facility after past missed visits and if they had challenged, confronted or educated someone stigmatizing them once in the past year. Patients were significantly less likely to return if they

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sought care from an urban health center or hospital, compared to a rural health center or were from the richest wealth tertile (Table 2).

Independent predictors of incident return to HIV care from the multivariable model with p-values at or below 0.01 level included having had a prior gap in care (aHR: 1.95, 95%CI: 1.23-3.09) and the patient having challenged, educated or confronted someone stigmatizing them once in the past year (aHR: 2.14, 95%CI: 1.25-3.65; more than once aHR: 0.65, 95%CI: 0.33-1.27). Patients were less likely to return to care if they sought care from an urban health center (aHR: 0.68, 95%CI: 0.48-0.96) or a hospital (aHR: 0.52, 95%CI: 0.33-0.82) compared to a rural health center (Table 2). While the overall p-value of the wealth tertile was 0.01, the hazard ratio estimates and confidence intervals did not show a consistent direction of association between increased wealth and return. (wealthiest aHR: 0.71, 95%CI: 0.47-1.08, middle tertile aHR: 1.27, 95%CI: 0.89-1.80) (Table 2).

#### *Supplemental Analysis: Predictors of return by one year*

An estimated 51.4% (95%CI: 33.2-72.5) of participants returned by one-year post-disengagement. In the multivariable model built based on significant predictors from univariate analyses, statistically significant (at the 0.01 level) independent predictors of incident return within 1-year of disengagement included being 45 years or older and having used herbal remedies in the past 6 months (Supplemental Table 2, <http://links.lww.com/QAI/B562>). Patients were less likely to return by 1-year if they reported a psychosocial or clinic-related reason for stopping care (Supplemental Table 2, <http://links.lww.com/QAI/B562>). The sensitivity analysis using the theory-driven model showed consistent results for the age and psychosocial reasons

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variables and identified no other significant predictors of return. Estimate precision was poor in these models due to limited events.

## **Discussion**

With sufficient follow-up time, a high proportion of disengaged, traced patients, 73%, return to care across four provinces in Zambia. More needs to be done, however, to hasten return. Among those patients returning to care, median time spent disengaged was 19 months. Our data show that the rate of return is higher soon after disengagement. Earlier efforts to facilitate return may be more effective. Indeed, retrospective analysis of patient outreach in Kenya demonstrated improve return with more rapid tracing<sup>22</sup>. However, more rapid return soon after disengagement may also indicate that patients who do not return quickly may require targeted support to come back to care.

Interventions to support patient resilience to stigma and to limit stigma in the social environment may facilitate increased re-engagement. Our data indicate that, compared to not confronting stigma at all, confronting stigma once in the past year facilitates re-engagement. This is consistent with existing literature on the relationship between coping, resilience and improved health outcomes<sup>37,38</sup>. However, we do not see a traditional dose-response relationship, as challenging stigmatizers multiple times does not further increase return. We theorize that repeat confrontation of stigmatizers may represent a more hostile social environment or chronic stress, limiting any positive effect the ability to respond to a stigmatizer may bring. Research has shown that the effect of HIV stigma on health is worse in the context of low perceived community support<sup>39</sup> and that the pathways through which resilience to stigma operates in the context of

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chronic stress are complex<sup>40</sup>. Future re-engagement research should include stigma and resilience measures and test effectiveness of resilience interventions to improve return to care<sup>41,42</sup>.

Despite tracing, 27% of disengaged patients did not return to care by the end of study follow-up. Our data suggest that disengaged patients from urban health centers and hospitals are at higher risk than rural patients of remaining disengaged and may require targeted interventions. Greater likelihood of return among those at rural health centers may be consistent with the more personal relationship-based care often available in rural, compared to urban and tertiary care centers. Existing research supports the importance of health care worker-patient relationships in patient engagement<sup>43,44</sup>. Additionally, urban versus rural patients may have different needs driving engagement. Past research has shown differences, for example, in which differentiated service delivery models (DSDs) for HIV treatment access are preferred between urban and rural patients<sup>45</sup>. More research is needed to understand the mechanisms underlying facility-level difference in re-engagement and how to best address them to support return.

The finding that prior care gaps predict incident re-engagement adds additional urgency to the need to conceptualize care engagement as a dynamic process<sup>4,5,10,14</sup>, and the need for effective interventions to support continuity of care. While complex factors are likely associated with both having a prior care gap and a patient's subsequent re-engagement, our findings suggest that investment in supporting patient return after one care gap may pay future re-engagement dividends. The greater than 2.5 fold increase in the hazard of return among disengaged patients who were repeatedly contacted by the clinic beyond the standard of care is consistent with this suggestion and other retention literature<sup>46,47</sup>. Together these results warrant further investigation

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into the mechanisms through which extended outreach may support return, such as relationship development, and outreach effectiveness evaluation.

Our analysis suggests that factors predictive of return by one-year post-disengagement are more proximal to the patient care experience than predictors within the full study period. This suggests that effective interventions early on may need to target different mechanisms than interventions for people who remain disengaged for a longer time. In addition to older age and the use of herbal remedies in the six months prior to the survey, independent predictors of return by one year included not reporting a clinic-related complaint (e.g. poor quality of care, lack of respect, spending too much time at the facility) or a psychosocial reason (e.g. clinic attendance creating conflicts, risking disclosure, being told to stop by someone influential, depression, forgetting or seeking alternative care), for stopping care. While self-treatment with herbal remedies may indicate illness-driven care-seeking, finding ways to reduce clinic and psychosocial barriers, such as improving patient clinic experiences<sup>43,44,48</sup> and engaging social support<sup>49,50</sup> may be important to encourage return sooner after disengagement.

## *Limitations*

Despite intensive tracing efforts, we were not able to obtain an updated vital or care status on 25% of the sampled patients. If disengaged patients not successfully traced are systematically different from those found, the estimates may be biased. Using EMR data to compare, patients we found were more likely to be from rural health centers and from provinces other than Lusaka, indicating that our estimates may over-represent rural experiences. These two groups were similar on other demographics (data not shown). Our study was only able to identify



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return among patients whose return care visit was documented using the same unique patient number in the four study provinces. It is possible that patients returned as a 'new' patient under a new unique patient number, or to a facility outside of the study area, potentially underestimating return. Patients in urban or tertiary care settings may have more health facility options due to higher facility density, which may make them more likely to have an undocumented return under a new patient number. Predictors were largely collected using survey responses, which are subject to self-report error, recall, and social desirability biases. As study observation began after disengagement, we assume that survey-measured predictors are time invariant in the interim. Due to poor documentation of mortality in the EMR we were unable to look at the competing risk of death.

### Conclusions

The most appropriate models of HIV care engagement show dynamic engagement patterns that demand multifaceted flexibility and support for retention, as is true for many chronic diseases<sup>51,52</sup>. Return to care after disengagement is a critical, yet under-researched step of the HIV care cascade. Our findings suggest that patients in urban and tertiary care settings may need additional return support, and that efforts to improve patient resilience and outreach after any care gap may facilitate return. Other important re-engagement influences may include positive patient experience at the clinic, having a supportive psychosocial environment, not being in the wealthiest population tertile, and older age. Future re-engagement research should include measures of these predictors to investigate their mechanisms of effect and evaluate their causal effect on return to care.

### **Acknowledgements**

We gratefully acknowledge the contributions of the Better Information for Health in Zambia study participants and research team, without whom this work would not have been possible. We thank and remember Nancy Czaicki for her contributions. We appreciate the data access support provided by Paul Kaumba, Jacob Mutale and Mwansa Lumpa at the Centre for Infectious Disease Research in Zambia. Thank you to Drs. David Celentano and Caitlin Kennedy for their feedback on the analysis.

### **References**

1. Kranzer K, Govindasamy D, Ford N, Johnston V, Lawn SD. 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(2):17383.
2. Mugavero MJ, Amico KR, Horn T, Thompson MA. The state of engagement in HIV care in the United States: from cascade to continuum to control. *Clinical infectious diseases : an official publication of the Infectious Diseases Society of America.* 2013;57(8):1164-1171.
3. Lee H, Wu XK, Genberg BL, et al. Beyond binary retention in HIV care: predictors of the dynamic processes of patient engagement, disengagement, and re-entry into care in a US clinical cohort. *AIDS (London, England).* 2018;32(15):2217-2225.

#### Predictors of return to HIV care

4. Lee H, Hogan JW, Genberg BL, et al. A state transition framework for patient-level modeling of engagement and retention in HIV care using longitudinal cohort data. *Statistics in medicine*. 2018;37(2):302-319.
5. Hallett TB, Eaton JW. A side door into care cascade for HIV-infected patients? *Journal of acquired immune deficiency syndromes (1999)*. 2013;63 Suppl 2:S228-232.
6. Powers KA, Miller WC. Critical Review: Building on the HIV Cascade: A Complementary "HIV States and Transitions" Framework for Describing HIV Diagnosis, Care, and Treatment at the Population Level. *Journal of acquired immune deficiency syndromes (1999)*. 2015;69(3):341-347.
7. Ahonkhai AA, Noubary F, Munro A, et al. Not all are lost: interrupted laboratory monitoring, early death, and loss to follow-up (LTFU) in a large South African treatment program. *PloS one*. 2012;7(3):e32993.
8. Alamo ST, Colebunders R, Ouma J, et al. Return to normal life after AIDS as a reason for lost to follow-up in a community-based antiretroviral treatment program. *Journal of acquired immune deficiency syndromes (1999)*. 2012;60(2):e36-45.
9. Crawford TN, Sanderson WT, Thornton A. Impact of poor retention in HIV medical care on time to viral load suppression. *Journal of the International Association of Providers of AIDS Care*. 2014;13(3):242-249.
10. Krentz HB, Vu Q, Gill MJ. The Impact of "Churn" on Plasma HIV Burden Within a Population Under Care. *Open forum infectious diseases*. 2019;6(6):ofz203.
11. Gardner EM, McLees MP, Steiner JF, Del Rio C, Burman WJ. The spectrum of engagement in HIV care and its relevance to test-and-treat strategies for prevention of

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- HIV infection. *Clinical infectious diseases : an official publication of the Infectious Diseases Society of America*. 2011;52(6):793-800.
12. Skarbinski J, Rosenberg E, Paz-Bailey G, et al. Human immunodeficiency virus transmission at each step of the care continuum in the United States. *JAMA internal medicine*. 2015;175(4):588-596.
  13. UNAIDS. *Fast-Track: ending the AIDS Epidemic by 2030*. Geneva, Switzerland: UNAIDS;2014.
  14. Nsanzimana S, Binagwaho A, Kanters S, Mills EJ. Churning in and out of HIV care. *The lancet HIV*. 2014;1(2):e58-59.
  15. Yehia BR, Stephens-Shields AJ, Fleishman JA, et al. The HIV Care Continuum: Changes over Time in Retention in Care and Viral Suppression. *PloS one*. 2015;10(6):e0129376.
  16. Haber N, Pillay D, Porter K, Barnighausen T. Constructing the cascade of HIV care: methods for measurement. *Current opinion in HIV and AIDS*. 2016;11(1):102-108.
  17. Gueler A, Vanobberghen F, Rice B, Egger M, Muglin C. The HIV Care Cascade from HIV diagnosis to viral suppression in sub-Saharan Africa: a systematic review and meta-regression analysis protocol. *Systematic reviews*. 2017;6(1):172.
  18. Cunningham CO, Buck J, Shaw FM, Spiegel LS, Heo M, Agins BD. Factors associated with returning to HIV care after a gap in care in New York State. *Journal of acquired immune deficiency syndromes (1999)*. 2014;66(4):419-427.
  19. Krebs DW, Chi BH, Mulenga Y, et al. Community-based follow-up for late patients enrolled in a district-wide programme for antiretroviral therapy in Lusaka, Zambia. *AIDS care*. 2008;20(3):311-317.

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20. Nabaggala MS, Parkes-Ratanshi R, Kasirye R, et al. Re-engagement in HIV care following a missed visit in rural Uganda. *BMC research notes*. 2018;11(1):762.
21. Camlin CS, Neilands TB, Odeny TA, et al. Patient-reported factors associated with reengagement among HIV-infected patients disengaged from care in East Africa. *AIDS (London, England)*. 2016;30(3):495-502.
22. Rebeiro PF, Bakoyannis G, Musick BS, et al. Observational Study of the Effect of Patient Outreach on Return to Care: The Earlier the Better. *Journal of acquired immune deficiency syndromes (1999)*. 2017;76(2):141-148.
23. Pecoraro A, Royer-Malvestuto C, Rosenwasser B, et al. Factors contributing to dropping out from and returning to HIV treatment in an inner city primary care HIV clinic in the United States. *AIDS care*. 2013;25(11):1399-1406.
24. Gill VC, Krentz HB. Patient Perspectives on Leaving, Disengaging, and Returning to HIV Care. *AIDS Patient Care STDS*. 2015;29(7):400-407.
25. Layer EH, Brahmhatt H, Beckham SW, et al. "I pray that they accept me without scolding:" experiences with disengagement and re-engagement in HIV care and treatment services in Tanzania. *AIDS Patient Care STDS*. 2014;28(9):483-488.
26. Holmes CB. Estimated mortality on HIV treatment among active patients and patients lost to follow-up in 4 provinces of Zambia: Findings from a multistage sampling-based survey. 2018;15(1).
27. Sikazwe I, Eshun-Wilson I, Sikombe K, et al. Retention and viral suppression in a cohort of HIV patients on antiretroviral therapy in Zambia: Regionally representative estimates using a multistage-sampling-based approach. *PLoS medicine*. 2019;16(5):e1002811.

Predictors of return to HIV care

28. Geng EH, Odeny TA, Lyamuya R, et al. Retention in Care and Patient-Reported Reasons for Undocumented Transfer or Stopping Care Among HIV-Infected Patients on Antiretroviral Therapy in Eastern Africa: Application of a Sampling-Based Approach. *Clinical infectious diseases : an official publication of the Infectious Diseases Society of America*. 2016;62(7):935-944.
29. Sweat MD, Denison JA. Reducing HIV incidence in developing countries with structural and environmental interventions. *AIDS (London, England)*. 1995;9 Suppl A:S251-257.
30. Ahonkhai AA, Adeola J, Banigbe B, et al. Impact of Unplanned Care Interruption on CD4 Response Early After ART Initiation in a Nigerian Cohort. *Journal of the International Association of Providers of AIDS Care*. 2017;16(1):98-104.
31. Ware NC, Wyatt MA, Geng EH, et al. Toward an understanding of disengagement from HIV treatment and care in sub-Saharan Africa: a qualitative study. *PLoS medicine*. 2013;10(1):e1001369; discussion e1001369.
32. Nakiwogga-Muwanga A, Musaaazi J, Katabira E, Worodria W, Talisuna SA, Colebunders R. Patients who return to care after tracking remain at high risk of attrition: experience from a large HIV clinic, Uganda. *International journal of STD & AIDS*. 2015;26(1):42-47.
33. Hargreaves JR, Stangl A, Bond V, et al. HIV-related stigma and universal testing and treatment for HIV prevention and care: design of an implementation science evaluation nested in the HPTN 071 (PopART) cluster-randomized trial in Zambia and South Africa. *Health policy and planning*. 2016;31(10):1342-1354.
34. Rutstein SO, Johnson K. *The DHS Wealth Index*. Calverton, Maryland: ORC Macro;2004.

Predictors of return to HIV care

35. Gari S, Malungo JR, Martin-Hilber A, Musheke M, Schindler C, Merten S. HIV testing and tolerance to gender based violence: a cross-sectional study in Zambia. *PloS one*. 2013;8(8):e71922.
36. Bush K, Kivlahan DR, McDonell MB, Fihn SD, Bradley KA. The AUDIT alcohol consumption questions (AUDIT-C): an effective brief screening test for problem drinking. Ambulatory Care Quality Improvement Project (ACQUIP). Alcohol Use Disorders Identification Test. *Archives of internal medicine*. 1998;158(16):1789-1795.
37. Varni SE, Miller CT, McCuin T, Solomon SE. Disengagement and Engagement Coping with HIV/AIDS Stigma and Psychological Well-Being of People with HIV/AIDS. *Journal of social and clinical psychology*. 2012;31(2):123-150.
38. Zhang L, Li X, Qiao S, et al. The mediating role of individual resilience resources in stigma-health relationship among people living with HIV in Guangxi, China. *AIDS care*. 2015;27(10):1317-1325.
39. Earnshaw VA, Lang SM, Lippitt M, Jin H, Chaudoir SR. HIV stigma and physical health symptoms: do social support, adaptive coping, and/or identity centrality act as resilience resources? *AIDS and behavior*. 2015;19(1):41-49.
40. Logie C, James L, Tharao W, Loutfy M. Associations between HIV-related stigma, racial discrimination, gender discrimination, and depression among HIV-positive African, Caribbean, and Black women in Ontario, Canada. *AIDS Patient Care STDS*. 2013;27(2):114-122.
41. Mak WWS, Mo PKH, Ma GYK, Lam MYY. Meta-analysis and systematic review of studies on the effectiveness of HIV stigma reduction programs. *Social science & medicine (1982)*. 2017;188:30-40.

Predictors of return to HIV care

42. Stangl AL, Lloyd JK, Brady LM, Holland CE, Baral S. A systematic review of interventions to reduce HIV-related stigma and discrimination from 2002 to 2013: how far have we come? *J Int AIDS Soc.* 2013;16(3 Suppl 2):18734.
43. Flickinger TE, Saha S, Moore RD, Beach MC. Higher quality communication and relationships are associated with improved patient engagement in HIV care. *Journal of acquired immune deficiency syndromes (1999).* 2013;63(3):362-366.
44. Beach MC, Keruly J, Moore RD. Is the quality of the patient-provider relationship associated with better adherence and health outcomes for patients with HIV? *J Gen Intern Med.* 2006;21(6):661-665.
45. Eshun-Wilson I, Mukumbwa-Mwenechanya M, Kim HY, et al. Differentiated Care Preferences of Stable Patients on Antiretroviral Therapy in Zambia: A Discrete Choice Experiment. *Journal of acquired immune deficiency syndromes (1999).* 2019;81(5):540-546.
46. Smith LR, Fisher JD, Cunningham CO, Amico KR. Understanding the behavioral determinants of retention in HIV care: a qualitative evaluation of a situated information, motivation, behavioral skills model of care initiation and maintenance. *AIDS patient care and STDs.* 2012;26(6):344-355.
47. Broaddus MR, Hanna CR, Schumann C, Meier A. "She makes me feel that I'm not alone": linkage to Care Specialists provide social support to people living with HIV. *AIDS care.* 2015;27(9):1104-1107.
48. Man JD, Mayega RW, Sarkar N, et al. Patient-Centered Care and People-Centered Health Systems in Sub-Saharan Africa: Why So Little of Something So Badly Needed? *The International Journal of Patient Centered Medicine.* 2016;6(3).



Predictors of return to HIV care

49. Sariah A, Rugemalila J, Protas J, et al. Why did I stop? And why did I restart? Perspectives of women lost to follow-up in option B+ HIV care in Dar es Salaam, Tanzania. *BMC Public Health*. 2019;19(1):1172.
50. Nixon SA, Bond V, Solomon P, et al. Optimism alongside new challenges: using a rehabilitation framework to explore experiences of a qualitative longitudinal cohort of people living with HIV on antiretroviral treatment in Lusaka, Zambia. *AIDS care*. 2018;30(3):312-317.
51. Frost J, Garside R, Cooper C, Britten N. A qualitative synthesis of diabetes self-management strategies for long term medical outcomes and quality of life in the UK. *BMC health services research*. 2014;14(1):348.
52. Bodenheimer T, Wagner EH, Grumbach K. Improving Primary Care for Patients With Chronic Illness. *Jama*. 2002;288(14):1775-1779.

Figure 1. (A) Cumulative Incidence of Re-engagement in Care (n=556) (B) Hazard for Returning to Care based on Time since Disengagement

Table 1. Disengaged Patient Characteristics (n=556)

	Total	
	n	%
<b>Potential predictors of return</b>		
<b>Sex</b>		
Male	232	41.7
Female	324	58.3
<b>Age at disengagement (years)</b>		
18-24	77	13.9
25-34	228	41.0
35-44	182	32.7
45+	69	12.4
<b>Marital status</b>		
Single, Never Married	115	20.7
Married	290	52.1
Separated, Divorced, Widowed	151	27.2
<b>Education</b>		
No formal education	40	7.2
Primary	247	44.4
Secondary	218	39.2
Tertiary	51	9.2
<b>Religion<sup>†</sup></b>		
Pentecostal	105	19.0
Universal Church of Zambia (UCZ)	37	6.7
7th Day Adventist	102	18.5
New Apostolic	79	14.3
Catholic	92	16.7
Other	137	24.8
<b>Province</b>		
Lusaka	238	42.8
Eastern	108	19.4
Southern	117	21.1
Western	93	16.7
<b>Facility type</b>		
Rural Health Center	131	23.6
Urban Health Center	301	54.1
Hospital	124	22.3
<b>Last CD4 count (cells/<math>\mu</math>mol) prior to loss<sup>°</sup></b>		
<350	155	35.6
351-500	100	23.0
>500	180	41.4
<b>Ill at enrollment, WHO Stage III or IV or enrollment CD4&lt;200<sup>§</sup></b>		
	162	30.9
<b>Time from HIV care enrollment to disengagement</b>		
$\leq$ 18 months	343	61.7
>18 months	213	38.3
<b>Prior gap in HIV care before study LTFU</b>		
	203	36.5
<b>Initiated ART</b>		
	247	44.4
<b>HIV status disclosure to someone</b>		
	479	86.2
<b>Patient ever contacted by facility in past when missed a visit prior to study<sup>¶</sup></b>		
No	472	86.1
Contacted as per standard of care (up to 3 times)	67	12.2
Contacted beyond standard of care (>3 times)	9	1.7
<b>Travel time from usual residence to facility</b>		
Less than 1 hour	239	43.0
1 to under 2 hours	139	25.0
2 hours or more	178	32.0
<b>Did not spend &gt;1 month away from usual residence in past year<sup>^</sup></b>		
	255	46.7
<b>Relationship to head of household<sup>†</sup></b>		
Head	274	49.4
Wife or husband of head	155	27.9
Other	126	22.7

<b>Used herbal remedies in past 6 months<sup>°</sup></b>	29	5.3
<b>No binge alcohol use<sup>°</sup></b>	367	67.8
<b>Wealth tertile<sup>α</sup></b>		
Poorest	184	33.6
Middle	193	35.3
Richest	170	31.1
<b>Tolerance of household violence<sup>ε</sup></b>		
No tolerance	426	80.2
Some tolerance	54	10.2
High tolerance	51	9.6
<b>High internalized stigma v. Low<sup>^</sup></b>	218	39.9
<b>Anticipated stigma<sup>α</sup></b>		
Anticipated, low	189	34.6
Anticipated, medium	144	26.3
Anticipated, high	214	39.1
<b>Experienced stigma in past 12 months<sup>°</sup></b>	133	24.5
<b>Challenged, educated or confronted stigmatizer in past 12 months<sup>†</sup></b>		
No	470	86.9
One time	29	5.3
More than once	42	7.8
<b>Patient reported reasons for disengagement<sup>#</sup></b>		
Any structural reason for stopping care	241	43.6
Any psychosocial reason for stopping care	227	41.1
Any clinic reason for stopping care	191	34.5
Any medical reason for stopping care	147	26.6
<b>Patient reported needs for return to care<sup>~</sup></b>		
Any structural barrier to return to care	96	17.7
Any psychosocial barrier to return to care	140	25.7
Any clinic barrier to return to care	229	42.1
Patient reported already planning to return	295	54.2

-n=552, °n=435, §n=524, ¶n=548, ^n=546, †n=555, °n=543, \*n=541, αn=547, εn=531, †n=541, #n=553 – categories are not mutually exclusive, ~n=544 – categories are not mutually exclusive

Table 2. Crude and adjusted predictors of return to care among disengaged patients

Predictors of return	Crude (univariate, complete case analysis)				Adjusted* (n=556)			
	Hazard Ratio	95% CI		p-value	Hazard Ratio	95% CI		p-value
<b>Male sex v. Female</b>	1.02	0.77	1.35	0.90	0.91	0.65	1.26	0.57
<b>Age at disengagement (years)</b>				0.55				0.65
18-24	1.00				1.00			
25-34	0.97	0.62	1.52		0.89	0.56	1.42	
35-44	1.16	0.74	1.82		1.01	0.63	1.64	
45+	1.28	0.74	2.20		1.21	0.68	2.17	
<b>Marital status</b>				0.64	//			
Single, Never Married	1.00							
Married	1.17	0.81	1.70					
Separated, Divorced, Widowed	1.20	0.80	1.81					
<b>Education</b>				0.07	//			
No formal education	1.00							
Primary	1.29	0.73	2.25					
Secondary	0.95	0.54	1.69					
Tertiary	0.67	0.31	1.44					
<b>Religion<sup>†</sup></b>				0.85	//			
Pentecostal	1.00							
Universal Church of Zambia (UCZ)	1.37	0.75	2.51					
7th Day Adventist	1.24	0.78	1.97					
New Apostolic	1.16	0.70	1.92					
Catholic	1.31	0.82	2.08					
Other	1.10	0.71	1.71					
<b>Province</b>				0.58	//			
Lusaka	1.00							
Eastern	0.96	0.65	1.42					
Southern	0.88	0.60	1.29					
Western	1.21	0.82	1.78					
<b>Facility type</b>				<0.01*				0.01
Rural Health Center	1.00				1.00			
Urban Health Center	0.63	0.46	0.87		0.68	0.48	0.96	
Hospital	0.43	0.28	0.66		0.52	0.33	0.82	
<b>Last CD4 count (cells/μmol) prior to loss<sup>°</sup></b>				0.38				0.63
<350	1.00				1.00			
351-500	0.75	0.49	1.15		0.85	0.55	1.32	
>500	0.85	0.59	1.20		0.83	0.55	1.25	
<b>Ill at enrollment (WHO Stage III or IV or enrollment CD4&lt;200) v. Not<sup>§</sup></b>	1.23	0.90	1.67	0.19	//			
<b>&gt;18 months from HIV care enrollment to disengagement v. ≤18 months</b>	0.96	0.72	1.29	0.81	0.58	0.36	0.94	0.03
<b>Prior gap in care v. no gap prior to study LTFU</b>	1.25	0.94	1.66	0.13	1.95	1.23	3.09	<0.01
<b>Initiated ART v. No ART</b>	0.88	0.66	1.16	0.36	//			
<b>HIV status disclosure to someone v. No disclosure</b>	1.21	0.78	1.87	0.40	//			
<b>Patient ever contacted by facility in past when missed a visit prior to study<sup>¶</sup></b>				0.04*				0.11
No	1.00				1.00			
Contacted as per standard of care (up to 3 times)	0.91	0.58	1.44		1.09	0.68	1.75	
Contacted beyond standard of care (>3 times)	2.84	1.26	6.43		2.65	1.04	6.73	

<b>Travel time from usual residence to facility</b>				0.32	//			
Less than 1 hour	1.00							
1 to under 2 hours	0.98	0.70	1.38					
2 hours or more	0.78	0.56	1.10					
<b>Did not spend &gt;1 month away from usual residence in past year v. Did<sup>^</sup></b>	1.26	0.95	1.67	0.11	1.05	0.78	1.41	0.74
<b>Relationship to head of household<sup>l</sup></b>				0.57	//			
Head	1.00							
Wife or husband of head	1.16	0.83	1.62					
Other	1.17	0.83	1.65					
<b>Used herbal remedies in past 6 mo v. Did not<sup>o</sup></b>	1.38	0.77	2.48	0.28	//			
<b>No binge alcohol use v. Binge alcohol use<sup>y</sup></b>	0.98	0.73	1.32	0.89	//			
<b>Wealth tertile<sup>α</sup></b>				<b>&lt;0.01*</b>				<b>0.01</b>
Poorest	1.00				1.00			
Middle	1.17	0.84	1.62		1.27	0.89	1.80	
Richest	0.64	0.44	0.92		0.71	0.47	1.08	
<b>Tolerance of household violence<sup>c</sup></b>				0.75	//			
No tolerance	1.00							
Some tolerance	0.83	0.51	1.35					
High tolerance	0.99	0.59	1.66					
<b>High internalised stigma v. Low<sup>^</sup></b>	1.20	0.90	1.60	0.21	//			
<b>Anticipated stigma<sup>α</sup></b>				0.40	//			
Anticipated, low	1.00							
Anticipated, medium	1.17	0.81	1.68					
Anticipated, high	1.26	0.90	1.76					
<b>Experienced stigma v. Did not experience stigma in past 12 months<sup>o</sup></b>	1.05	0.76	1.45	0.78	//			
<b>Challenged, educated or confronted stigmatizer in past 12 months<sup>†</sup></b>				<b>0.01*</b>				<b>&lt;0.01</b>
No	1.00				1.00			
One time	1.90	1.15	3.14		2.14	1.25	3.65	
More than once	0.63	0.33	1.19		0.65	0.33	1.27	
<b>Patient reported reasons for disengagement<sup>#</sup></b>								
Any structural reason for stopping care v. no structural	1.05	0.79	1.39	0.74	//			
Any psychosocial reason for stopping care v. no psychosocial	0.80	0.60	1.07	0.13	0.94	0.68	1.29	0.68
Any clinic reason for stopping care v. no clinic	1.11	0.83	1.49	0.47	//			
Any medical reason for stopping care v. no medical	1.13	0.83	0.45	0.52	//			
<b>Patient reported needs for return to care<sup>~</sup></b>								
Any structural barrier to return to care v. no structural barrier	1.04	0.72	1.50	0.84	//			
Any psychosocial barrier to return to care v. no psychosocial barrier	0.71	0.50	1.01	0.06	0.71	0.48	1.06	0.10
Any clinic barrier to return to care v. no clinic barrier	1.07	0.80	1.42	0.66	//			
Patient reported already planning to return v. not	1.05	0.79	1.40	0.72	//			
-n=552, °n=435, §n=524, ¶n=548, ^n=546, †n=555, °n=543, ¥n=541, αn=547, çn=531, †n=541, #n=553, ~n=544					*adjusted based on theory and 0.05 univariate significance: sex, age, last CD4 count, time since enrollment, past care gaps, past facility contact after loss, facility type, mobility, psychosocial barriers to care			

Figure 1. (A) Cumulative Incidence of Re-engagement in Care (n=556) (B) Hazard for Returning to Care based on Time since Disengagement

