

# Linkage to HIV care after home-based HIV counselling and testing in sub-Saharan Africa: A systematic review

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## Abstract

**Background:** Home-based HIV counselling and testing (HBHCT) has the potential to increase HIV testing uptake in sub-Saharan Africa (SSA) but data on linkage to HIV care after HBHCT are limited. We conducted a systematic review of linkage to care after HBHCT in SSA.

**Methods:** Five databases were searched for studies published between 1<sup>st</sup> January 2000 and 19<sup>th</sup> August 2016 that reported on linkage to care among adults newly identified with HIV infection through HBHCT. Eligible studies were reviewed, assessed for risk of bias and findings summarised using the PRISMA guidelines.

**Results:** Fourteen studies from six countries met the eligibility criteria; nine used specific strategies (point-of-care CD4 count testing, follow-up counselling, provision of transport funds to clinic, and counsellor facilitation of HIV clinic visit) in addition to routine referral to facilitate linkage to care. Time intervals for ascertaining linkage ranged from one week to twelve months post-HBHCT. Linkage ranged from 8.2% [95% confidence interval (CI), 6.8%-9.8%] to 99.1% (95% CI, 96.9%-99.9%). Linkage was generally lower (<33%) if HBHCT was followed by referral only, and higher (>80%) if additional strategies were used. Only one study assessed linkage by means of a randomised trial. Five studies had data on cotrimoxazole (CTX) prophylaxis and twelve on ART eligibility and initiation. CTX uptake among those eligible ranged from 0% to 100%. The proportion of persons eligible for ART

ranged from 16.5% (95% CI, 12.1-21.8) to 77.8% (95% CI, 40.0-97.2). ART initiation among those eligible ranged from 14.3% (95% CI, 0.36%-57.9%) to 94.9% (95% CI, 91.3%-97.4%). Additional linkage strategies, whilst seeming to increase linkage, were not associated with higher uptake of CTX and/or ART. Most of the studies were susceptible to risk of outcome ascertainment bias. A pooled analysis was not performed because of heterogeneity across studies with regard to design, setting, and the key variable definitions.

**Conclusion:** Only few studies from SSA investigated linkage to care among adults newly diagnosed with HIV through HBHCT. Linkage was often low after routine referral but higher if additional interventions were used to facilitate it. The effectiveness of linkage strategies should be confirmed through randomised controlled trials.

## **Introduction**

Access to antiretroviral therapy (ART) in sub-Saharan Africa (SSA) has expanded considerably but AIDS-related mortality remains high [1]. A major cause of this mortality is the late presentation of patients for treatment [2]. Early ART initiation is dependent on early HIV diagnosis, and prompt linkage to and retention in care [3]. HIV counselling and testing (HCT) is essential in expanding HIV prevention and treatment services [4]. However, HCT uptake in SSA remains low [5]. For instance, the proportion of HIV-positive adults in SSA who are aware of their HIV status has been estimated to be only 60% [6]. In order to expand access to HIV testing in settings with generalised HIV epidemics, the World Health Organisation (WHO) recommends community-based HCT with linkage to prevention, care and treatment services, in addition to facility-based HCT [7].

In the community-based HCT model, services are delivered through mobile, workplace-, school-, and home-based approaches thus removing structural, logistical, and social barriers to HCT [8]. Community-based HCT may also be delivered as part of multi-disease campaigns that involve intensive community mobilization lasting 1-2 weeks followed by mobile HIV testing, often coupled with other preventive medical services (campaign HCT) [9].

Under the home-based HIV counselling and testing (HBHCT) approach, HCT services are conducted by trained HCT service providers in the client's home [10]. HBHCT may be provided to everyone in a community through a door-to-door approach or to household members of known tuberculosis (TB) or HIV-positive patients [4, 10]. According to WHO, there were at least 39 HBHCT programmes in 10 SSA countries by early 2011 [10]. A systematic review published the following year found 21 studies that had reported on the uptake of HBHCT in SSA [11]. Since then, several studies have reported successful implementation of HBHCT in rural [12-24] and urban [19, 23] populations in SSA. HBHCT is highly acceptable and has the potential to substantially increase HIV testing uptake in SSA [11]. It is cost-effective at reaching previously untested persons compared with other HCT models in settings with high HIV prevalence [25, 26], promotes equitable access of services [27] and may help to promote HCT among couples [28] and prevention of mother-to-child HIV transmission [29]. Importantly, HBHCT facilitates early HIV diagnosis and therefore provides an opportunity for early linkage to care [29]. These attributes highlight the potential

of HBHCT as an effective platform for HIV prevention and population-based test-and-treat strategies.

Despite these advantages, few data are available on linkage to care after HBHCT particularly among newly identified HIV-positive persons or on the effectiveness of strategies to increase linkage after HCT [30]. In the absence of interventions to facilitate linkage to care, individuals that test HIV positive through HBHCT may find it more challenging to enter care compared to those identified in facility-based HCT [31]. This is because HBHCT is more likely to reach socio-economically disadvantaged populations that have difficulty accessing healthcare services compared to facility-based HCT [32]. In order to identify effective linkage strategies, data are specifically required on linkage to care soon after HBHCT. The reasons for this include the current WHO recommendation to initiate ART among all HIV-positive adults regardless of WHO clinical stage and at any CD4 count [33]; increasing use of HBHCT in Africa; and growing importance of early treatment for improved clinical outcomes [34-36] and HIV prevention [37]. A recent systematic review on linkage to care following community- and facility-based HCT [9] included 10 HBHCT studies, but did not distinguish linkage outcomes between HBHCT and campaign HCT, between newly and previously diagnosed HIV-positive individuals, or between children/adolescents and adults. Individuals who previously tested HIV-positive and have not yet linked to care are likely to differ from newly identified patients with regard to barriers that may prevent service uptake [12]. Similarly linkage to care among children/adolescents may be influenced by factors that are unique to this population [38, 39].

The specific objectives of our review were to: estimate the proportion of individuals in SSA linking to care within 12 months among those who were newly diagnosed with HIV; the proportion initiating daily cotrimoxazole (CTX) prophylaxis (i.e. the people who initiated daily CTX prophylaxis among those who linked to care and were eligible for CTX); and the proportion initiating ART (i.e. the people who initiated ART among those who linked to care and were eligible for ART); and to summarise data on the strategies that have been used to increase linkage to care after HBHCT.

## **Methods**

### **Search strategy**

We searched five databases (Medline, Embase, Global Health, Web of Science, and Africa-Wide information) for studies published between 1<sup>st</sup> January 2000 (time at which roll-out of ART programmes began in SSA [11]) and 19<sup>th</sup> August 2016. The following key terms were used: (HIV diagnosis OR HIV voluntary counselling and testing OR HIV testing and counselling OR HIV counselling and testing) AND (home based OR mobile OR community OR household OR door-to-door OR survey) AND (linkage OR access OR uptake OR enrolment OR non-enrolment OR retention OR loss to follow-up OR loss to care OR care OR treatment OR pre- antiretroviral therapy) AND (Africa OR individual names of countries in SSA). No language restriction was applied to the literature search. Identified articles were exported using Endnote reference management software and duplicate articles removed. Two authors (ER and SB) independently screened titles and abstracts of articles to identify eligible publications, discussed inconsistencies, and reached a consensus on their eligibility. Studies were eligible if they were conducted in SSA, and had original data on linkage to care among adults ( $\geq 18$  years) newly identified with HIV infection through HBHCT, defined as HCT services offered in an individual's home. Studies whose study populations included persons  $< 18$  years were eligible but only data for participants aged  $\geq 18$  years were utilised to estimate linkage and other outcomes. Studies for which the required information was not published but might have been collected were identified and the corresponding authors approached with requests for additional data. Where two or more eligible articles reported on similar or overlapping populations, the article with the most complete data was included. Review articles were excluded but their bibliographies as well as those of the identified articles were manually checked to identify any additional studies. Conference abstracts were excluded. All potentially eligible papers were then subjected to full text screening.

### **Data extraction and synthesis**

A data extraction form was used to collect the following information from each eligible article: first author's name, publication year, country and setting where study was conducted, study population, sample size, study design, definition of linkage to care, strategies used to promote and time for evaluation of linkage to care. We also obtained the number of HIV-positive adults who were newly diagnosed, and, among those, the numbers who linked to care (as defined in the respective studies), and were eligible for, and initiated, CTX prophylaxis

and ART (based on national guidelines that were in use at the time of the respective studies). Risk of bias in the included studies was assessed using a component approach, similar to the Cochrane Collaboration's [40], and based on three items: selection bias, outcome ascertainment, and attrition. The results were summarised using the PRISMA guidelines [41].

We used the reported data to calculate the proportions [and their 95% confidence intervals (CI), using the Clopper–Pearson method] who linked to care, initiated CTX prophylaxis, were eligible for and initiated ART. The denominator for linkage was all newly diagnosed HIV-positive adults ( $\geq 18$  years) who had a potential minimum follow-up period corresponding to the time point when linkage was assessed (i.e. including those who out-migrated, died, or were lost to follow-up, but excluding those who entered the study at a later date so had a shorter potential follow-up period). We did not stratify our linkage estimates by individual-level factors that may potentially influence referral uptake, such as HIV disease stage or CD4 count at the time of HIV diagnosis, as these data were not available for most of the studies. However, we compared linkage estimates between studies conducted under different CD4 ART eligibility thresholds i.e.  $\leq 250$  cells/ $\mu\text{L}$  versus  $\leq 350$  cells/ $\mu\text{L}$ . The denominator for ART eligibility was all individuals who linked within the specified time period, and those for initiation of CTX prophylaxis and ART were all individuals who linked and were eligible for CTX and ART respectively. We did not perform a meta-analysis because the identified studies varied widely with regard to design, setting, definition of linkage to care, the time points of and method for ascertaining linkage, and with regard to the strategies used to facilitate linkage.

## Results

### *Summary of search results*

The search identified 5,905 articles of which 61 were subjected to full text screening (Figure 1). Of those screened, 21 were eligible for detailed review; two were excluded on the basis of reporting on overlapping study populations [20, 42]. Of the remaining 19 articles, one [19] had all the required data. Additional data were obtained for 13 [12-14, 18, 21-24, 43-47] of the remaining 18 articles after contacting the respective corresponding authors. Thus, 14 studies were included in the review. A summary description of the included studies is presented in Table 1. The studies were conducted in six countries i.e. Uganda [13, 14, 43,

44], South Africa [13, 14, 18, 23, 24, 45], Kenya [12, 19], Malawi [46, 47], Lesotho [22], and Swaziland [21] between 2005 and 2015. Most (92%) studies were based in rural [12-14, 18, 21, 22, 24, 43, 45, 47] or semirural settings [44, 46] settings; two were conducted in both rural and urban [19, 23] populations. The number of newly identified HIV-positive adults varied widely across studies (range: 15–1637).

#### *Summary of study objectives and populations*

The included studies aimed to assess acceptability of HBHCT [18, 19, 21-24, 46, 47], and HIV prevalence [19], linkage to care [12-14, 18, 19, 21-24, 43-45], uptake of ART [13, 14, 18, 24, 44, 47], and viral suppression [13, 14, 18], in the context of HBHCT. Two studies compared HBHCT and mobile HCT with regard to uptake and cost of HCT, HIV positivity rates and linkage to care [21, 22]. Three observational studies [13, 18, 44] and one randomised trial [14] were designed to evaluate the effect of specific interventions on linkage to HIV care, uptake of ART, and other outcomes. Except for one study in which HBHCT was provided to household contacts of TB index patients [23], all the other studies used the door-to-door HBHCT approach. Study participants mostly comprised individuals aged  $\geq 13$  years. However, two studies also recruited children ( $< 13$  years) [19, 21]. With the exception of two studies that recruited newly identified HIV-positive individuals [22, 23], participants in other studies were a mixture of previously and newly identified HIV-positive patients.

#### *Risk of bias*

Only two [13, 18] studies had a low risk of bias for all assessed items (Table 2). Risk of selection bias was low ( $\geq 80\%$  HBHCT coverage) in four studies [13, 18, 19, 44]. Risk of attrition bias was low ( $\geq 80\%$  participant retention) in eight [13, 14, 18, 23, 24, 44-46] of ten studies in which participants were followed. Self-reported linkage to care was confirmed by tracking referrals and review of records at the referral clinic in only two studies [43, 45]. In the first study [43], no information was reported on the proportion of participants for whom clinic records were not found. In the second study [45], clinic records were found for only 71% of the tracked referrals and self-reported data was used to ascertain linkage for the rest of the participants. Self-reported linkage to care was confirmed by review of documentation issued to patients by HIV clinics (e.g. clinic cards) in three studies [13, 14, 18]; linkage was not verified with the HIV clinics. In five studies [12, 21, 22, 24, 47], ascertainment of linkage to care was based on data from HIV clinics in the areas where the studies were conducted;

participants who may have linked to HIV clinics outside of the study areas were not tracked. Only self-reported data was used in the rest of the studies [19, 23, 44, 46].

### *Linkage to HIV care*

In all studies, persons who tested HIV positive were referred for care. Additional strategies to facilitate linkage to care were used in nine (64%) [13, 14, 18, 22, 24, 43-46] of the studies (Table 1). These strategies included: provision of funds for transport to the HIV clinic [46]; follow-up counselling [13, 14, 18, 24, 43-45]; lay counsellor facilitation of the initial HIV clinic visits (the counsellor met the HIV-positive participant at the clinic and explained the clinic processes and the benefits of ART) [14]; point-of-care (POC) CD4 count [13, 14, 18, 22] and home-based collection of samples for viral load [13] testing and provision of results. Linkage to care was ascertained within 3 months of HBHCT in 50% of studies [12, 19, 22, 23, 44-46]. Ascertainment of linkage in the remaining studies was done >3 to 6 [18, 21, 43] and >6 to 12 [13, 14, 24, 47] months after HBHCT.

Linkage to care ranged from 8.2% (95% CI, 6.8%-9.8%) [12] to 85.4% (95% CI, 75.8%-92.2%) [14] when only referral was offered, and 24.3% (95% CI, 11.8%-41.2%) [22] to 99.1% (95% CI, 96.9%-99.9%) [13] when referral plus additional interventions to facilitate linkage were offered (Table 1). In general, linkage to care was lower (<33%) in the studies that offered referral only [12, 19, 21, 23, 47] and higher (>80%) in those that used a combination of additional linkage strategies [13, 14, 18, 44].

Among seven studies that were conducted in the context of a CD4 ART eligibility threshold of  $\leq 250$  cells/ $\mu$ L [12, 19, 43-47], linkage to care ranged from 8.2% (95% CI, 6.8%-9.8%) [12] to 81.8% (95% CI, 71.4-89.7) [44]. Linkage was lower (<30.0%) in the studies that offered referral only [12, 19, 47] and higher (>50%) in those that used at least one additional linkage strategy [43-46].

Among four studies conducted in the context of a CD4 ART eligibility threshold of  $\leq 350$  cells/ $\mu$ L [13, 21-23], linkage to care ranged from 24.3% (95% CI, 11.8-41.2) [22] to 99.1% (95% CI 96.9-99.9) [13]. Linkage was <33% in the two studies that offered referral only [21, 23]. Linkage was also low (24.3%) in one study that used referral and POC CD4 count testing



[22], but very high (99.1%) in a study that offered referral, POC CD4 testing and additional linkage strategies [13].

#### *Uptake of CTX prophylaxis and ART*

Five studies conducted in Kenya [19], Uganda [43, 44], and Uganda and South Africa [13, 14] had data on initiation of CTX prophylaxis among eligible individuals who linked to care. CTX prophylaxis is only recommended for patients with CD4 count  $\leq 200$  cells/ $\mu\text{L}$ , WHO stage 3 or 4 or HIV/TB co-infection in South Africa [48] but is routinely provided to all HIV-positive persons irrespective of CD4 count or WHO disease stage in Uganda [49] and Kenya [50]. Of the studies conducted in Uganda and Kenya, additional interventions to facilitate referral uptake were offered in all except one. CTX uptake in these studies ranged from 78.2% (95% CI, 69.3-85.5%) [13] to 100% [14, 43, 44] (Table 1). CTX uptake was also high [90.6% (95% CI, 87.3%-93.2%)] in the study that offered referral only [19]. Interventions to facilitate linkage were offered in the two studies that were conducted in South Africa. Uptake of CTX among patients with CD4 count  $\leq 200$  cells/ $\mu\text{L}$  in these studies ranged from 0% [14] to 33.3% (95% CI, 4.3-77.7) [14].

Twelve studies [12-14, 18, 19, 21, 22, 24, 43-45, 47] had data on ART eligibility and ART initiation among patients who linked to care. The proportion of individuals eligible for ART initiation ranged from 16.5% (95% CI, 12.1-21.8) [45] to 77.8% (95% CI, 40.0-97.2) [22] (Table 1). ART uptake among those who linked to care ranged from 33.0% (95% CI, 24.2%-41.7%) [19] to 94.0% (95% CI, 85.4%-98.3%) [47] in the studies that provided referral only. A similar range i.e. 14.3% (95% CI, 0.36%-57.9%) [22] to 94.9% (95% CI, 91.3%-97.4%) [43] was observed in the studies that provided referral plus additional linkage interventions. ART initiation rates were highest ( $\geq 90\%$ ) in the two studies in which HIV care services were provided through community-based research clinics [43, 47].

## **Discussion**

HBHCT is increasingly being used in SSA, and an effectively conducted HBHCT strategy would be a key precondition for HIV control programmes that propagate a test-and-treat approach for HIV prevention [45]. The success of such programmes will partly depend on their capacity to achieve high levels of linkage to care following HIV diagnosis [16]. Hence, it is necessary to identify and set up strategies that will effectively link persons identified

with HIV through HBHCT to care and treatment. As observed in this review however, only a few studies have investigated linkage to care among adults newly identified with HIV through HBHCT in SSA. Linkage to care was below 33% in five of six studies where participants were only referred for care with no further interventions to facilitate referral [12, 19, 21, 23, 47], a figure that is lower than those for client-initiated facility-based HCT (61%) and provider-initiated facility-based HCT (55%) [9]. With the exception of two studies [22, 24], studies that used additional linkage strategies recorded moderate (>50% to <80%) [43, 45, 46] to high ( $\geq 80\%$ ) [13, 14, 18, 44] levels of linkage. These trends remained irrespective of the CD4 ART eligibility threshold at the time of the studies. In general, linkage to care was highest when participants were offered POC CD4 count testing and follow-up counselling [13, 14, 18]. These findings suggest that HBHCT coupled with interventions to facilitate referral uptake may achieve similar or even higher linkage compared to facility-based HCT.

WHO recommends CTX prophylaxis for all adults with WHO stage 3 and 4 HIV disease and/or with a CD4 count  $\leq 350$  cells/mm<sup>3</sup>, and regardless of CD4 count in settings where malaria and/or severe bacterial infections are highly prevalent [33]. For this reason, CTX prophylaxis is an essential component of HIV care in many settings in SSA, and its uptake may be used as an indicator of access to HIV care [19, 51]. However, only a small number of studies included in this review had data on the uptake of CTX. In Kenya and Uganda where routine CTX prophylaxis is recommended irrespective of CD4 count or clinical stage [49, 50], uptake was high (>70%) irrespective of whether or not additional interventions were applied. This may be because CTX is widely available, inexpensive, and simple to use [52]. In contrast and in spite of facilitated linkage, uptake of CTX prophylaxis among those eligible (based on guidelines that were in use at the time of the studies) in South Africa was low ( $\leq 33\%$ ). The reasons for this are not clear. However previous studies have found irregular supply and lack of stocks of CTX, lack of awareness among health care workers, and perceived low priority of CTX prophylaxis due to the absence of a reporting requirement, to be some of the barriers to implementation of CTX prophylaxis policies [53].

Consistent with previous findings [11], significant proportions of HIV-positive persons identified through HBHCT were still ineligible for ART (based on national guidelines that were in use at the time of the respective studies). The finding that ART uptake was highest in the studies where services were provided through community-based research clinics may be

attributed to such clinics being more accessible and less prone to limitations that characterise many public sector ART care programmes in SSA, including the requirement for several visits to prepare patients for ART [14], crowded, busy and unwelcoming clinics [54], non-functioning laboratories [54], and inadequate supply or lack of antiretroviral drugs [55]. Additionally, some research clinics are likely to have close and long standing relationships with communities in which they are located.

HBHCT studies with facilitated linkage have been shown to achieve higher ART initiation rates among participants who link to care compared to those without facilitated linkage [9]. However, ART initiation rates were high in some but not all studies with facilitated linkage described in this review. Additionally, some studies without facilitated linkage achieved higher ART initiation rates [47] than those with facilitated linkage [13, 14, 18, 22, 24, 44, 45]. It is likely that clinic level factors such as those mentioned above as well as individual-level confounding factors such as HIV disease stage may be more important in influencing events after linkage to care. Also, people who link to care in the absence of facilitated linkage may be more motivated to receive care.

This review has some limitations. The review was limited to SSA. Even so, the number of relevant studies found is small and represents only six countries, four of which are in Southern Africa. This may limit the generalizability of the findings. The methodologies used in the identified studies varied widely, making it impractical to combine findings from individual studies into a pooled analysis. Outcome assessments in most of the included studies were based solely on self-reports or records in HIV clinics within the respective study areas; hence linkage to care may have been overestimated in the case of self-reports or underestimated if some individuals linked to clinics outside of their communities.

Importantly, most of the studies that included interventions to facilitate linkage to HIV care were observational, determining intervention effects without control groups. A major limitation of observational studies is that it is difficult to account for the effects of confounding factors such as HIV disease stage, fear of stigma, healthcare seeking behaviour, and familiarity with the health care services. Moreover, in some of the studies, two or more interventions were delivered concurrently, making it difficult to distinguish the effects of each. Randomised trials represent the gold-standard methodology in the evaluation of an

intervention including its likely effect size [56]. It is therefore desirable to confirm the impact and establish the cost effectiveness of these interventions in randomised controlled trials. Indeed, a number of trials have been designed to investigate the effect of different interventions on linkage to care among persons that test HIV positive through HBHCT [57-61]. Findings from these trials are expected in the near future and may provide more robust data on the impact of interventions on linkage to care after HBHCT.

Except for one study in which some of the newly identified HIV-positive individuals were offered immediate ART irrespective of CD4 count [24], all the other studies were conducted in the context of old ART eligibility criteria i.e. CD4 counts of  $\leq 200$  cells/ $\mu$ L,  $\leq 250$  cells/ $\mu$ L,  $\leq 350$  cells/ $\mu$ L or  $\leq 500$  cells/ $\mu$ L and by implication, prolonged pre-ART care periods. Therefore, the extent to which the findings of this review are relevant to settings in which the new WHO recommendation of immediate ART initiation irrespective of CD4 count [33] has been adopted is not clear. A future review of the evidence on linkage to care after HBHCT under the new treatment guidelines will be needed.

In conclusion, we found that only few published studies investigated linkage to HIV care among adults newly diagnosed with HIV through HBHCT in SSA. In general, HBHCT without additional intervention strategies to increase service uptake achieved inadequate linkage while HBHCT combined with some kind of additional strategy seemed to achieve higher linkage. There is a need to confirm the impact of the most promising linkage strategies through randomised controlled trials before they can be recommended for large scale adoption. Moreover, it will be important to demonstrate the effectiveness of linkage strategies under the new WHO treat-all policy.

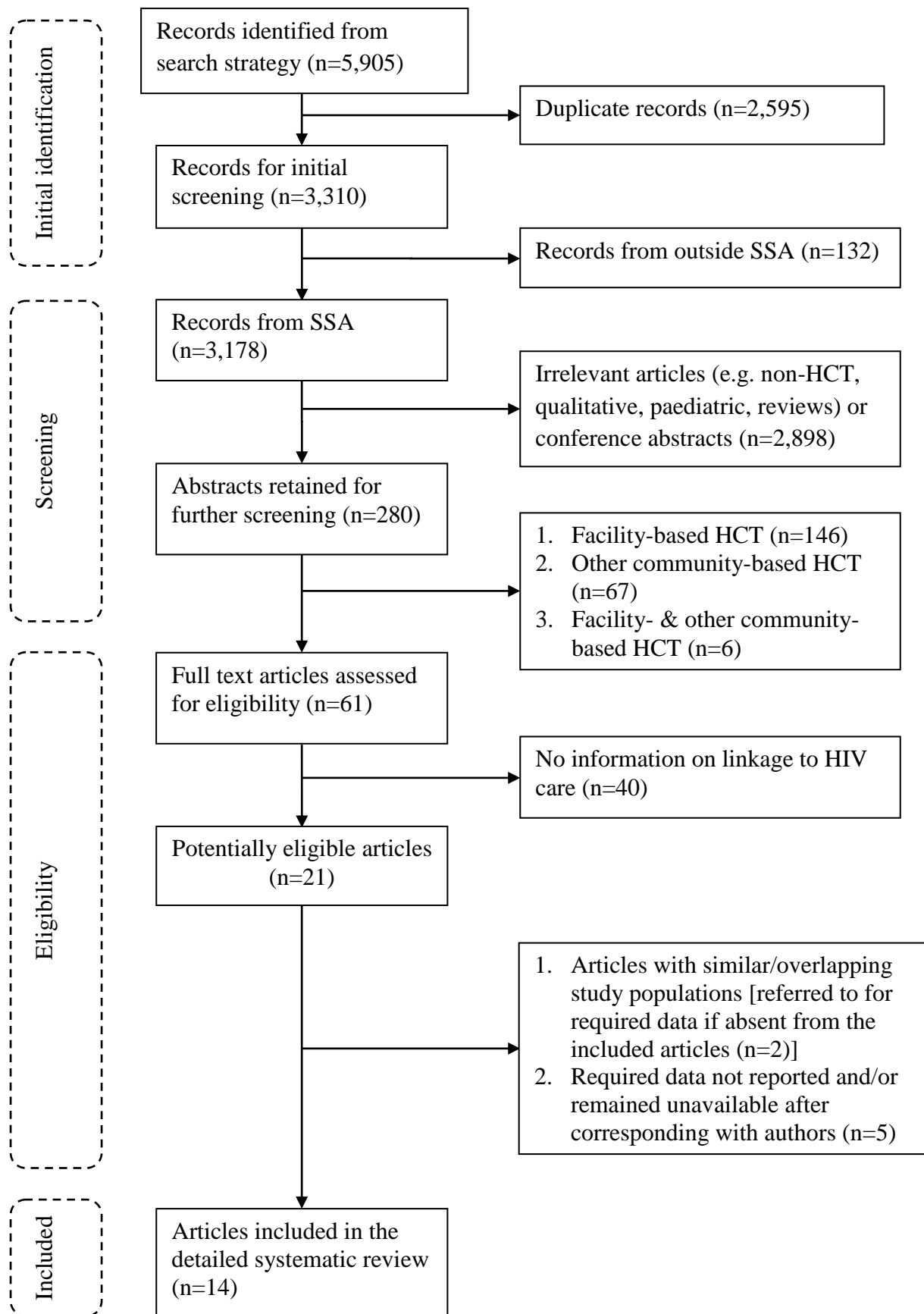
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**Figure 1: Systematic search flow diagram**



**Table 1: Description of studies included in the systematic review**

Author, year	Country, setting	Study period	Study design	Age eligibility (years)	HIV care provider	CD4 ART eligibility threshold (cells/ $\mu$ L)	Linkage to care definition	Linkage assessment time (months)	Linkage strategies	Number in analysis (Number of HIV-positive persons in study) <sup>a</sup>	Linked n (%), 95% CI	Initiated CTX <sup>b</sup> n/N (%), 95% CI	Eligible for ART <sup>c</sup> n (%), 95% CI	Initiated ART n (%), 95% CI
Genberg, 2015 [12]	Kenya, rural	2009-2011	Retrospective population-based cohort study of HBHCT among participants enrolled in a clinical care program	$\geq 13$	Public clinic	$\leq 250$	Registration at clinic	3	Referral only	1329 (3482)	109 (8.2, 6.8-9.8)	Not reported	41 (37.6, 28.5-47.4)	23 (56.1, 39.7-71.5)
Dalal, 2013 [19]	Kenya, rural & urban	2008	Prospective population-based cohort study of HBHCT among participants enrolled in a disease surveillance program	All ages <sup>d</sup>	Public clinic	$\leq 250$	Registration at clinic	1	Referral only	1637 (2759)	414 (25.3, 23.2-27.5)	375/414 (90.6, 87.3-93.2)	120 (29.0, 24.7-33.6)	39 (33.0, 24.2-41.7)
Wringe, 2012 [47]	Malawi, rural	2008-2010	Retrospective population-based cohort study of HBHCT among participants enrolled in a demographic surveillance program	$\geq 15$	Research clinic	$\leq 250$	Screening for ART eligibility	12	Referral only	431 (473)	126 (29.2, 24.9-33.8)	Not reported	67 (53.2, 44.1-62.1)	63 (94.0, 85.4-98.3)
Parker, 2015 [21]	Swaziland, rural	2013	Prospective population-based intervention of HBHCT versus mobile HCT	$\geq 18$ months	Public clinic	$\leq 350$	Registration at clinic	6	Referral only	142 (170)	45 (31.7, 24.1-40.0)	Not reported	17 (37.8, 23.8-53.5)	9 (52.9, 27.8-77.0)

Author, year	Country, setting	Study period	Study design	Age eligibility (years)	HIV care provider	CD4 ART eligibility threshold (cells/ $\mu$ L)	Linkage to care definition	Linkage assessment time (months)	Linkage strategies	Number in analysis (Number of HIV-positive persons in study) <sup>a</sup>	Linked n (%; 95% CI)	Initiated CTX <sup>b</sup> n/N (%; 95% CI)	Eligible for ART <sup>c</sup> n (%; 95% CI)	Initiated ART n (%; 95% CI)
Velen, 2016 [23]	South Africa, rural & urban	2013-2014	Prospective cohort study of HBHCT among household contacts of TB index patients enrolled in a TB contact tracing trial	$\geq 14$	Public clinic	$\leq 350$	Registration at clinic	3	Referral only	25 (26)	8 (32.0, 14.9-53.5)	Not reported	Not reported	Not reported
Iwuji, 2016 [24]	South Africa, rural	2012-2014	Community-based cluster randomised trial of immediate ART initiation versus ART initiation according to national guidelines following HBHCT	$\geq 16$	Public & research clinics	Any CD4 count (intervention arm); $\leq 350$ (standard-of-care arm)	Registration at clinic	12	Referral & follow-up counselling <sup>e</sup>	358 (2569)	162 (45.3, 40.0-50.6)	Not reported	101 (71.6, 63.4-78.9) <sup>f</sup>	81 (80.2, 71.1-87.5)
Naik, 2015 [45]	South Africa, rural	2009-2011	Prospective population-based cohort of participants offered HBHCT as a standard-of-care service & in the context of a trial of HBHCT versus facility-based HCT.	$\geq 14$	Public clinic	$\leq 200$	Obtaining a CD4 count	3	Referral & at least 3 follow-up counselling visits <sup>g</sup>	410 (492)	248 (60.5, 55.6-65.3)	Not reported	41 (16.5, 12.1-21.8)	33 (80.5, 65.1-91.2)



Author, year	Country, setting	Study period	Study design	Age eligibility (years)	HIV care provider	CD4 ART eligibility threshold (cells/ $\mu$ L)	Linkage to care definition	Linkage assessment time (months)	Linkage strategies	Number in analysis (Number of HIV-positive persons in study) <sup>a</sup>	Linked n (%; 95% CI)	Initiated CTX <sup>b</sup> n/N (%; 95% CI)	Eligible for ART <sup>c</sup> n (%; 95% CI)	Initiated ART n (%; 95% CI)
Labhardt, 2014 [22]	Lesotho, rural	2011	Community-based cluster randomised trial of HBHCT versus mobile HCT	All ages	Public clinic	$\leq 350$	Registration at clinic	1	Referral & POC CD4 count testing	37 (39)	9 (24.3, 11.8-41.2)	Not reported	7 (77.8, 40.0-97.2)	1 (14.3, 0.36-57.9)
Becker, 2014 [46]	Malawi, peri-urban	2009	Population-based uncontrolled intervention study of couple HBHCT & couple family planning services	15-49 (female); $\geq 15$ (male)	Public clinic	$\leq 250$	Registration at clinic	1 week	Referral & provision of funds for transport to clinic to participants who disclosed their HIV status to their partners <sup>h</sup>	15 (46)	8 (53.3, 26.6-78.7)	Not reported	Not reported	Not reported
Nakigozi, 2011 [43]	Uganda, rural	2005-2008	Retrospective population-based cohort of participants receiving HBHCT or other community-based HCT in an HIV surveillance program	15-49	Research clinic	$\leq 250$	Registration at clinic	6	Referral & follow-up counselling <sup>i</sup>	1137 (1451)	781 (68.7, 65.9-71.4)	781/781 (100.0)	237 (30.3, 27.1-33.7)	225 (94.9, 91.3-97.4)
Tumwebaze, 2012 [44]	Uganda, rural & peri-urban	2010-2011	Population-based uncontrolled intervention study of HBHCT & a combination of linkage strategies	$\geq 18$	Public clinic	$\leq 250$	Registration at clinic	3	Referral, CD4 count laboratory testing (results returned to participant a week later), & follow-up counselling (1, 2 & 3 months)	77 (152)	63 (81.8, 71.4-89.7)	63/63 (100.0)	13 (20.6, 11.5-32.7)	8 (61.5, 31.6-86.1)

Author, year	Country, setting	Study period	Study design	Age eligibility (years)	HIV care provider	CD4 ART eligibility threshold (cells/ $\mu$ L)	Linkage to care definition	Linkage assessment time (months)	Linkage strategies	Number in analysis (Number of HIV-positive persons in study) <sup>a</sup>	Linked n (%; 95% CI)	Initiated CTX <sup>b</sup> n/N (%; 95% CI)	Eligible for ART <sup>c</sup> n (%; 95% CI)	Initiated ART n (%; 95% CI)
Barnabas, 2014 [13]	South Africa & Uganda, rural	2011-2013	Population-based uncontrolled intervention study of HBHCT & a combination of linkage strategies	$\geq 18$	Public clinic	$\leq 350$	Registration at clinic	12	Referral, POC CD4 count testing, follow-up counselling (1, 3, 6, 9, & 12 months), & viral load testing (0 & 6 months)	229 (635)	227 (99.1, 96.9-99.9)	2/12 (16.7, 2.1-48.4) SA; 86/110 (78.2, 69.3-85.5) UG	74 (32.6, 26.5-39.1)	59 (79.7, 68.8-88.2)
van Rooyen, 2013 [18]	South Africa, rural	2011-2012	Population-based uncontrolled intervention study of HBHCT & a combination of linkage strategies	$\geq 18$	Public clinic	$\leq 200$ ; $\leq 350$ (from August 2011)	Registration at clinic	6	Referral, POC CD4 count testing, & follow-up counselling (1, 3, & 6 months)	73 (201)	70 (95.9, 88.5-99.1)	Not reported	35 (50.0, 37.8-62.2)	19 (54.3, 36.6-71.2)
Barnabas, 2016 [14]	South Africa & Uganda, rural	2013-2015	Household randomised controlled trial of referral only versus referral plus other linkage strategies after HIV diagnosis through HBHCT or mobile HCT (6 linkage strategies) <sup>j</sup>	$\geq 16$	Public clinic	$\leq 350$ ; $\leq 500$ (from January 2015)	Registration at clinic	9	Referral only	82 (226)	70 (85.4, 75.8-92.2)	2/6 (33.0, 4.3-77.7) SA; 20/25 (80.0, 59.3-93.2) UG	37 (52.9, 40.6-64.9)	28 (75.7, 58.8-88.2)
									Referral & POC CD4 count testing	81 (213)	73 (90.1, 81.5-95.6)	1/4 (25.0, 0.63-80.6) SA; 37/38 (97.4, 86.2-99.9) UG	43 (58.9, 46.8-70.3)	25 (58.1, 42.1-73.0)
									Referral & counsellor clinic linkage facilitation	104 (231)	102 (98.1, 93.2-99.8)	1/3 (33.3, 0.84-90.6) SA; 18/21 (85.7, 63.7-97.0) UG	51 (50.0, 39.9-60.1)	39 (76.5, 62.5-87.2)

Author, year	Country, setting	Study period	Study design	Age eligibility (years)	HIV care provider	CD4 ART eligibility threshold (cells/ $\mu$ L)	Linkage to care definition	Linkage assessment time (months)	Linkage strategies	Number in analysis (Number of HIV-positive persons in study) <sup>a</sup>	Linked n (%; 95% CI)	Initiated CTX <sup>b</sup> n/N (%; 95% CI)	Eligible for ART <sup>c</sup> n (%; 95% CI)	Initiated ART n (%; 95% CI)
									Referral, POC CD4 count testing, & counsellor clinic linkage facilitation	72 (206)	69 (95.8, 98.3-99.1)	0/3 (0.0) SA; 23/27 (85.2, 66.3-95.8) UG	43 (62.3, 49.8-73.7)	23 (53.5, 37.7-68.8)
									Referral & follow-up counselling (1, 3, & 6 months)	87 (229)	80 (92.0, 84.1-96.7)	0/12 (0.0) SA; 30/30 (100.0) UG	41 (51.3, 39.8-62.6)	31 (75.6, 59.7-87.6)
									Referral, POC CD4 count testing & follow-up counselling (1, 3, & 6 months)	85 (220)	81 (95.3, 88.4-98.7)	1/5 (20.0, 0.51-71.6) SA; 28/28 (100.0) UG	46 (56.8, 45.3-67.8)	36 (78.3, 63.6-89.1)

<sup>a</sup>Only adults ( $\geq 18$  years) newly diagnosed with HIV through HBHCT were included in the analysis.

<sup>b</sup>Uptake of CTX prophylaxis is shown separately for South Africa (SA) and Uganda (UG) because eligibility criteria are different in each country.

<sup>c</sup>Based on locally recommended CD4 count eligibility threshold during the study period.

<sup>d</sup>Children aged <13 years were offered HBHCT if their biological mothers were HIV-positive or deceased

<sup>e</sup>Follow-up counselling was offered to individuals who did not link to care within 3 months of referral; number and timing of follow-up visits were not specified.

<sup>f</sup>Information on ART eligibility and initiation was not available for persons who linked to the public health facilities. Hence, the denominator used for ART eligibility (n=141) is the number of persons who linked to the research clinics.

<sup>g</sup>Timing of follow-up visits was not specified.

<sup>h</sup>The number of participants who disclosed their HIV status to their partners was not reported.

<sup>i</sup>Number and timing of follow-up visits were not specified.

<sup>j</sup>Results for linkage to care and other outcomes are presented separately for each of the six linkage strategies.

**Table 2: Risk of bias within studies**

Author, year	Selection of participants <sup>a</sup>	Outcome ascertainment <sup>b</sup>	Loss to follow-up <sup>c</sup>
Genberg, 2015 [12]	HBHCT coverage could not be estimated because the size of the target population was not reported. <i>Unclear risk</i>	Clinic-verified data. No information on persons who may have linked to clinics outside the study area. <i>Unclear risk</i>	No participant follow-up
Dalal, 2013 [19]	HBHCT coverage was 82%. <i>Low risk</i>	Self-report for all participants. <i>High risk</i>	48% (881/1839) loss to follow-up. <i>High risk</i> <sup>d</sup>
Wringe, 2012 [47]	HBHCT coverage could not be estimated because the total number of persons in the target population was not reported. <i>Unclear risk</i>	Clinic-verified data. No information on persons who may have linked to clinics outside the study area. <i>Unclear risk</i>	No participant follow-up
Parker, 2015 [21]	HBHCT was conducted in only 26% of households in the target area due to time constraints. It is not clear how these households were selected. <i>High risk</i>	Clinic-verified data. Participants who were referred health facilities outside the study area were excluded from analysis. <i>Unclear risk</i>	No participant follow-up
Velen, 2016 [23]	HBHCT was offered to household contacts of TB patients selected through convenience sampling. <i>High risk</i>	Self-report for all participants. <i>High risk</i>	12% (3/26) loss to follow-up. <i>Low risk</i>
Iwujii, 2016 [24]	HBHCT coverage was 64%. <i>High risk</i>	Clinic verified data. No information on persons who may have linked to clinics outside the study area. <i>Unclear risk</i>	16% (58/358) loss to follow-up. <i>Low risk</i> <sup>e</sup>
Naik, 2015 [45]	HBHCT coverage could not be estimated because the size of the population targeted for testing in the participating communities was not reported. <i>Unclear risk</i>	Self-report & clinic-verified data or self-report only (29% of participants). <i>High risk</i>	18% (79/438) loss to follow-up. <i>Low risk</i> <sup>f</sup>
Labhardt, 2014 [22]	HBHCT coverage could not be estimated because the total number of persons in the target population was not reported. <i>Unclear risk</i>	Clinic-verified data. No information on persons who may have linked to clinics outside the study area. <i>Unclear risk</i>	No participant follow-up
Becker, 2014 [46]	HBHCT was offered to all eligible participants in one village and approximately one-third in two other villages (no information on how participants in these two villages were selected). <i>High risk</i>	Self-report for all participants. <i>High risk</i>	2% (1/46) loss to follow-up. <i>Low risk</i>
Nakigozi, 2011 [43]	Annual HIV testing coverage in the cohort is >90%. However, 21% of the persons who tested HIV-positive were excluded from the analysis because they had either refused to learn their HIV results (3%) or received their HIV result less than six months before data-set closure (18%). <i>High risk</i>	Self-report & clinic-verified data. No information on whether self-reported linkage was confirmed with clinic records for all participants. <i>Unclear risk</i>	Loss to follow-up not reported. <i>Unclear risk</i>
Tumwebaze, 2012 [44]	HBHCT coverage was 80%. <i>Low risk</i>	Self-report for all participants. <i>High risk</i>	2% (3/152) loss to follow-up. <i>Low risk</i>

Author, year	Selection of participants <sup>a</sup>	Outcome ascertainment <sup>b</sup>	Loss to follow-up <sup>c</sup>
Barnabas, 2014 [13]	HBHCT coverage was 96%. <i>Low risk</i>	Self-report & review of documentation issued to patient by the HIV clinic. <i>Low risk</i>	10% (62/635) loss to follow-up. <i>Low risk</i>
van Rooyen, 2013 [18]	HBHCT coverage was 91%. <i>Low risk</i>	Self-report & review of documentation issued to patient by the HIV clinic. <i>Low risk</i>	2% (5/201) loss to follow-up. <i>Low risk</i>
Barnabas, 2016 [14]	HBHCT coverage could not be estimated because the size of the population targeted for testing in the participating communities was not reported. <i>Unclear risk</i>	Self-report & review of documentation issued to patient by the HIV clinic. <i>Low risk</i>	3% (40/1325) loss to follow-up. <i>Low risk</i>

<sup>a</sup>In population-based studies, there was low risk of bias if HBHCT coverage (defined as the number of persons accessing HBHCT out of the total resident population) was  $\geq 80\%$ , high risk if HBHCT coverage was  $< 80\%$  and unclear risk if there was no information on coverage. In non-population based cohort studies, there was low risk if participants were randomly selected, high risk if the selection was non-random and unclear risk if there was insufficient information on participant selection.

<sup>b</sup>There was low risk of bias if ascertainment of linkage to care was by both self-report & examination of HIV clinic records or documentation issued to patients by the HIV clinic for  $\geq 80\%$  of the participants, high risk if ascertainment was by self-report only and unclear risk if there was insufficient information on ascertainment of linkage outcomes for some study participants.

<sup>c</sup>There was low risk if retention of HIV-positive persons identified through HBHCT was  $\geq 80\%$ , high risk if retention was  $< 80\%$  and unclear risk if information was not available.

<sup>d</sup>Only applicable to HIV-positive individuals who were newly identified through HBHCT.

<sup>e</sup>Only applicable to newly identified HIV-positive persons who were referred to care at least 12 months before the end of the first phase of the trial i.e. May 2014.

<sup>f</sup>Only applicable to HIV-positive individuals who were not engaged in care at baseline.

## References

1. UNAIDS. Global AIDS update 2016. Geneva: UNAIDS; 2016 [Accessed 15 October 2016]; Available from: [http://www.unaids.org/sites/default/files/media\\_asset/global-AIDS-update-2016\\_en.pdf](http://www.unaids.org/sites/default/files/media_asset/global-AIDS-update-2016_en.pdf).
2. Fox MP, Rosen S. Patient retention in antiretroviral therapy programs up to three years on treatment in sub-Saharan Africa, 2007-2009: systematic review. *Trop Med Int Health*. 2010;15 Suppl 1:1-15.
3. Kranzer K, Zeinecker J, Ginsberg P, Orrell C, Kalawe NN, Lawn DS, et al. Linkage to HIV Care and Antiretroviral Therapy in Cape Town, South Africa. *PLoS ONE*. 2010;5(11):e13801.
4. Matovu JK, Makumbi FE. Expanding access to voluntary HIV counselling and testing in sub-Saharan Africa: alternative approaches for improving uptake, 2001-2007. *Trop Med Int Health*. 2007;12(11):1315-22.
5. Suthar AB, Ford N, Bachanas PJ, Wong VJ, Rajan JS, Saltzman AK, et al. Towards universal voluntary HIV testing and counselling: a systematic review and meta-analysis of community-based approaches. *PLoS Med*. 2013;10(8):e1001496.
6. UNAIDS. How AIDS changed everything — MDG6: 15 years, 15 lessons of hope from the AIDS response. Geneva2015 [Accessed 14 June 2016]; Available from: [http://www.unaids.org/sites/default/files/media\\_asset/MDG6Report\\_en.pdf](http://www.unaids.org/sites/default/files/media_asset/MDG6Report_en.pdf).
7. WHO. Consolidated guidelines on HIV testing services 2015. 5Cs: Consent, Confidentiality, Counselling, Correct results, and Connection Geneva WHO; 2015 [Accessed 12 October 2016]; Available from: [http://apps.who.int/iris/bitstream/10665/179870/1/9789241508926\\_eng.pdf](http://apps.who.int/iris/bitstream/10665/179870/1/9789241508926_eng.pdf).
8. Mabuto T, Latka MH, Kuwane B, Churchyard GJ, Charalambous S, Hoffmann CJ. Four models of HIV counseling and testing: utilization and test results in South Africa. *PLoS ONE*. 2014;9(7).
9. Sharma M, Ying R, Tarr G, Barnabas R. Systematic review and meta-analysis of community and facility-based HIV testing to address linkage to care gaps in sub-Saharan Africa. *Nature*. 2015;528:S77-S85.
10. WHO. Planning, implementing and monitoring home-based HIV testing and counselling: a practical handbook for Sub-Saharan Africa. Geneva 2012 [Accessed 21 March 2017]; Available from: [http://apps.who.int/iris/bitstream/10665/75366/1/9789241504317\\_eng.pdf?ua=1](http://apps.who.int/iris/bitstream/10665/75366/1/9789241504317_eng.pdf?ua=1).
11. Sabapathy K, Van den Bergh R, Fidler S, Hayes R, Ford N. Uptake of home-based voluntary HIV testing in sub-Saharan Africa: a systematic review and meta-analysis. *PLoS Med*. 2012;9(12):e1001351.
12. Genberg BL, Naanyu V, Wachira J, Hogan JW, Sang E, Nyambura M, et al. Linkage to and engagement in HIV care in western Kenya: An observational study using population-based estimates from home-based counselling and testing. *Lancet HIV*. 2015;2(1):e20-e6.
13. Barnabas RV, van Rooyen H, Tumwesigye E, Murnane PM, Baeten JM, Humphries H, et al. Initiation of antiretroviral therapy and viral suppression after home HIV testing and counselling in KwaZulu-Natal, South Africa, and Mbarara district, Uganda: A prospective, observational intervention study. *Lancet HIV*. 2014;1(2):e68-e76.
14. Barnabas RV, van Rooyen H, Tumwesigye E, Brantley J, Baeten JM, van Heerden A, et al. Uptake of antiretroviral therapy and male circumcision after community-based HIV testing and strategies for linkage to care versus standard clinic referral: a multisite, open-label, randomised controlled trial in South Africa and Uganda. *Lancet HIV*. 2016 3(5):e212-20.

15. Fylkesnes K, Sandoy IF, Jürgensen M, Chipimo PJ, Mwangala S, Michelo C. Strong effects of home-based voluntary HIV counselling and testing on acceptance and equity: a cluster randomised trial in Zambia. *Soc Sci Med.* 2013;86:9-16.
16. Bigogo G, Amolloh M, Laserson KF, Audi A, Aura B, Dalal W, et al. The impact of home-based HIV counseling and testing on care-seeking and incidence of common infectious disease syndromes in rural western Kenya. *BMC Infect Dis.* 2014;14:376.
17. Medley A, Ackers M, Amolloh M, Owuor P, Muttai H, Audi B, et al. Early uptake of HIV clinical care after testing HIV-positive during home-based testing and counseling in western Kenya. *AIDS Behav.* 2013;17(1):224-34.
18. van Rooyen H, Barnabas RV, Baeten JM, Phakathi Z, Joseph P, Krows M, et al. High HIV testing uptake and linkage to care in a novel program of home-based HIV counseling and testing with facilitated referral in KwaZulu-Natal, South Africa. *J Acquir Immune Defic Syndr.* 2013;64(1):e1-8.
19. Dalal W, Feikin DR, Amolloh M, Ransom R, Burke H, Lugalia F, et al. Home-based HIV testing and counseling in rural and urban Kenyan communities. *J Acquir Immune Defic Syndr.* 2013;62(2):e47-54.
20. Doherty T, Tabana H, Jackson D, Naik R, Zembe W, Lombard C, et al. Effect of home based HIV counselling and testing intervention in rural South Africa: cluster randomised trial. *Bmj.* 2013;13(346).
21. Parker LA, Jobanputra K, Rusike L, Mazibuko S, Okello V, Kerschberger B, et al. Feasibility and effectiveness of two community-based HIV testing models in rural Swaziland. *Trop Med Int Health.* 2015;20(7):893-902.
22. Labhardt ND, Motlomelo M, Cerutti B, Pfeiffer K, Kamele M, Hobbins MA, et al. Home-Based Versus Mobile Clinic HIV Testing and Counseling in Rural Lesotho: A Cluster-Randomized Trial. *PLoS Med.* 2014;11(12):e1001768.
23. Velen K, Lewis JJ, Charalambous S, Page-Shipp L, Popane F, Churchyard GJ, et al. Household HIV Testing Uptake among Contacts of TB Patients in South Africa. *PLoS ONE.* 2016;11(5):e0155688.
24. Iwuji CC, Orne-Gliemann J, Larmarange J, Okesola N, Tanser F, Thiebaut R, et al. Uptake of Home-Based HIV Testing, Linkage to Care, and Community Attitudes about ART in Rural KwaZulu-Natal, South Africa: Descriptive Results from the First Phase of the ANRS 12249 TasP Cluster-Randomised Trial. *PLoS Med.* 2016;13(8).
25. Menzies N, Abang B, Wanyenze R, Nuwaha F, Mugisha B, Coutinho A, et al. The costs and effectiveness of four HIV counseling and testing strategies in Uganda. *AIDS.* 2009;23(3):395-401.
26. Tabana H, Nkonki L, Hongoro C, Doherty T, Ekstrom AM, Naik R, et al. A Cost-Effectiveness Analysis of a Home-Based HIV Counselling and Testing Intervention versus the Standard (Facility Based) HIV Testing Strategy in Rural South Africa. *PLoS ONE.* 2015;10(8).
27. Helleringer S, Kohler HP, Frimpong JA, Mkandawire J. Increasing uptake of HIV testing and counseling among the poorest in sub-Saharan countries through home-based service provision. *J Acquir Immune Defic Syndr.* 2009;51(2):185-93.
28. Mantell JE, Dicarolo AL, Remien RH, Zerbe A, Morris D, Pitt B, et al. 'There's no place like home': perceptions of home-based HIV testing in Lesotho. *Health Educ Res.* 2014;29(3):456-69.
29. Wachira J, Kimaiyo S, Ndege S, Mamlin J, Braitstein P. What is the impact of home-based HIV counseling and testing on the clinical status of newly enrolled adults in a large HIV care program in Western Kenya? *Clin Infect Dis.* 2012;54(2):275-81.

30. WHO. Service delivery approaches to HIV testing and counselling (HTC): a strategic HTC programme framework. Geneva, Switzerland 2012 [Accessed 24 June 2016]; Available from: [http://apps.who.int/iris/bitstream/10665/75206/1/9789241593877\\_eng.pdf](http://apps.who.int/iris/bitstream/10665/75206/1/9789241593877_eng.pdf).
31. WHO. Global HIV/AIDS response: epidemic update and health sector progress towards universal access: progress report 2011. Geneva, Switzerland: World Health Organization,; 2011 [Accessed 1 January 2017]; Available from: [http://apps.who.int/iris/bitstream/10665/44787/1/9789241502986\\_eng.pdf](http://apps.who.int/iris/bitstream/10665/44787/1/9789241502986_eng.pdf).
32. Ganguli I, Bassett IV, Dong KL, Walensky RP. Home testing for HIV infection in resource-limited settings. *Curr HIV/AIDS Rep.* 2009;6(4):217-23.
33. WHO. Consolidated guidelines on the use of antiretroviral drugs for treating and preventing HIV infection: recommendations for a public health approach. 2016 [Accessed 15 November 2016]; Available from: [http://apps.who.int/iris/bitstream/10665/208825/1/9789241549684\\_eng.pdf?ua=1](http://apps.who.int/iris/bitstream/10665/208825/1/9789241549684_eng.pdf?ua=1).
34. Siegfried N, Uthman OA, Rutherford GW. Optimal time for initiation of antiretroviral therapy in asymptomatic, HIV-infected, treatment-naïve adults. *Cochrane Database Syst Rev.* 2010;17(3):CD008272.
35. Severe P, Juste MA, Ambroise A, Eliacin L, Marchand C, Apollon S, et al. Early versus standard antiretroviral therapy for HIV-infected adults in Haiti. *N Engl J Med.* 2010;363(3):257-65.
36. Kitahata MM, Gange SJ, Abraham AG, Merriman B, Saag MS, Justice AC, et al. Effect of early versus deferred antiretroviral therapy for HIV on survival. *N Engl J Med.* 2009;360(18):1815-26.
37. Cohen MS, Chen YQ, McCauley M, Gamble T, Hosseinipour MC, Kumarasamy N, et al. Prevention of HIV-1 infection with early antiretroviral therapy. *N Engl J Med.* 2011;365(6):493-505.
38. Phelps BR, Ahmed S, Amzel A, Diallo MO, Jacobs T, Kellerman SE, et al. Linkage, initiation and retention of children in the antiretroviral therapy cascade: an overview. *AIDS.* 2013;27(2):S207-13.
39. Govindasamy D, Ferrand RA, Wilmore SM, Ford N, Ahmed S, Afnan-Holmes H, et al. Uptake and yield of HIV testing and counselling among children and adolescents in sub-Saharan Africa: a systematic review. *J Int AIDS Soc.* 2015;18(20182).
40. The Cochrane collaboration. *Cochrane Handbook for Systematic Reviews of Interventions*, version 5.1.0 [updated March 2011]. The Cochrane Collaboration; 2011 [Accessed 7 August 2016]; Available from: <http://handbook.cochrane.org>.
41. Moher D, Liberati A, Tetzlaff J, Altman DG, Prisma Group. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. *PLoS Med.* 2009;6(7):e1000097.
42. Plazy M, Farouki KE, Iwuji C, Okesola N, Orne-Gliemann J, Larmarange J, et al. Access to HIV care in the context of universal test and treat: challenges within the ANRS 12249 TasP cluster-randomized trial in rural South Africa. *J Int AIDS Soc.* 2016;19(1).
43. Nakigozi G, Makumbi F, Reynolds S, Galiwango R, Kagaayi J, Nalugoda F, et al. Non-enrollment for free community HIV care: findings from a population-based study in Rakai, Uganda. *AIDS Care.* 2011;23(6):764-70.
44. Tumwebaze H, Tumwesigye E, Baeten JM, Kurth AE, Revall J, Murnane PM, et al. Household-based HIV counseling and testing as a platform for referral to HIV care and medical male circumcision in Uganda: a pilot evaluation. *PLoS ONE.* 2012;7(12):e51620.
45. Naik R, Doherty T, Jackson D, Tabana H, Swanevelder S, Thea DM, et al. Linkage to care following a home-based HIV counselling and testing intervention in rural South Africa. *J Int AIDS Soc.* 2015;18(19843).



46. Becker S, Taulo FO, Hindin MJ, Chipeta EK, Loll D, Tsui A. Pilot study of home-based delivery of HIV testing and counseling and contraceptive services to couples in Malawi. *BMC Public Health*. 2014;14(1309).
47. Wringe A, Floyd S, Kazooba P, Mushati P, Baisley K, Urassa M, et al. Antiretroviral therapy uptake and coverage in four HIV community cohort studies in sub-Saharan Africa. *Trop Med Int Health*. 2012;17(8):e38-48.
48. Department of Health: Republic of South Africa. National consolidated guidelines for the prevention of mother-to-child transmission of HIV (PMTCT) and the management of HIV in children, adolescents and adults. Pretoria, South Africa: Department of Health; 2014 [Accessed 15 July 2016]; Available from: <http://www.kznhealth.gov.za/family/HIV-Guidelines-Jan2015.pdf>.
49. Uganda Ministry of Health. National Antiretroviral Treatment Guidelines for Adults, Adolescents, and Children. Kampala 2008 [Accessed 15 July 2016]; Available from: [http://www.who.int/hiv/amds/uganda\\_moh\\_treatment\\_guidelines.pdf](http://www.who.int/hiv/amds/uganda_moh_treatment_guidelines.pdf).
50. Kenya Ministry of Medical Services. Guidelines for antiretroviral therapy in Kenya Nairobi: Kenya Ministry of Medical Services,; 2011 [Accessed 15 July 2016]; 4:[Available from: <http://healthservices.uonbi.ac.ke/sites/default/files/centraladmin/healthservices/Kenya%20Treatment%20Guidelines%202011.pdf>].
51. Lubogo D, Ddamulira JB, Tweheyo R, Wamani H. Factors associated with access to HIV care services in eastern Uganda: the Kumi home based HIV counseling and testing program experience. *BMC Fam Pract*. 2015;16:162.
52. WHO. Guidelines on co-trimoxazole prophylaxis for HIV-related infections among children, adolescents and adults: recommendations for a public health approach. Geneva: World Health Organisation; 2006 [Accessed 15 July 2016]; Available from: <http://www.who.int/hiv/pub/guidelines/ctxguidelines.pdf?ua=1>.
53. Date AA, Vitoria M, Granich R, Banda M, Fox MY, Gilks C. Implementation of co-trimoxazole prophylaxis and isoniazid preventive therapy for people living with HIV. *Bull World Health Organ*. 2010;88(4):253-9.
54. MacPherson P, MacPherson EE, Mwale D, Bertel Squire S, Makombe SD, Corbett EL, et al. Barriers and facilitators to linkage to ART in primary care: a qualitative study of patients and providers in Blantyre, Malawi. *J Int AIDS Soc*. 2012;15(2):18020.
55. Muhamadi L, Nsabagasani X, Tumwesigye MN, Wabwire-Mangen F, Ekstrom AM, Peterson S, et al. Inadequate pre-antiretroviral care, stock-out of antiretroviral drugs and stigma: policy challenges/bottlenecks to the new WHO recommendations for earlier initiation of antiretroviral therapy (CD<350 cells/microL) in eastern Uganda. *Health Policy*. 2010;97(2-3):187-94.
56. Hayes JR, Moulton HL. Cluster randomised trials. Boca Raton: Taylor & Francis; 2009. 338 p.
57. ClinicalTrials.gov. The Effectiveness of a Counselling Intervention on the Uptake of HIV Care Services Among HIV Infected Patients in Uganda. 2016 [Accessed July 2016]; Available from: <https://clinicaltrials.gov/ct2/show/NCT02497456?term=NCT02497456&rank=1>.
58. ClinicalTrials.gov. Same-day ART Initiation Versus Standard of Care After Positive HIV-test Result in Rural Lesotho (CASCADE). ClinicalTrials.gov; 2016 [Accessed 14 October 2016]; Available from: <https://clinicaltrials.gov/ct2/show/NCT02692027>.
59. ClinicalTrials.gov. High-yield HIV Testing, Facilitated Linkage to Care, and Prevention for Female Youth in Kenya (GIRLS). 2016 [Accessed 15 July 2016]; Available from: <https://clinicaltrials.gov/ct2/show/NCT02735642>.

60. ClinicalTrials.gov. The Effectiveness of a Counselling Intervention on the Uptake of HIV Care Services Among HIV Infected Patients in Uganda. 2015 [Accessed 15 July 2015]; Available from:  
<https://clinicaltrials.gov/ct2/show/NCT02497456?term=NCT02497456&rank=1>.
61. ClinicalTrials.gov. Enhanced Linkage to HIV Care Following Home-Based HIV Testing in Rural Uganda. 2015 [Accessed 15 July 2016]; Available from:  
<https://clinicaltrials.gov/ct2/show/study/NCT02545673>.