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Psychosocial interventions for reducing alcohol consumption in sub-Saharan African settings: a systematic review and metaanalysis

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Abstract

Background: Harmful alcohol use is a leading cause of morbidity and mortality in sub-Saharan Africa (sSA); however, the effects of non-pharmacological alcohol interventions in this region are unknown.

Design: A systematic review and meta-analysis of the available literature through March 14, 2019 was undertaken. Two authors extracted and reconciled relevant data and assessed risk of bias. Meta-analyses were conducted. The review protocol is registered on PROSPERO (CRD42019094509).

Setting: Studies conducted in sSA were eligible for inclusion.

Participants: Individuals participating in interventions aimed at reducing alcohol use.

Interventions: Randomized and non-randomized controlled trials testing non-pharmacological interventions (psychosocial and structural) on alcohol consumption in sSA.

Measurements: Eligible outcomes included the Alcohol Use Disorders Identification Test (AUDIT) scores; alcohol abstinence; measures of drinking quantity and frequency; and biomarkers of alcohol consumption.

Corresponding author: Katelyn M. Sileo, PhD, One UTSA Circle, San Antonio, TX, 78212, katelyn.sileo@utsa.edu; 860-977-8447. **Declaration of competing interests:** The authors have no conflicts of interests to disclose.

Findings: Nineteen intervention trials (18 scientific manuscripts) testing psychosocial interventions (no structural intervention included), judged of moderate quality, were included in meta-analyses. A beneficial effect was identified for psychosocial interventions on alcohol abstinence at 3–6 months (OR=2.05, 95% CI=1.20–3.48, k=5, n=2,312, I² = 79%) and 12–60 months (OR=1.91, 95% CI=1.40–2.61, k=6, n=2,737, I² = 63%) follow-up. There were no statistically significant effects found for AUDIT score (2–3 month: MD= –1.13, 95% CI: –2.60–0.34, k=6, n=992, I²=85%; 6 month: MD= –0.83, 95% CI= –1.92–0.26, k=6, n=1081, I²=69%; 12 month: MD= –0.15, 95% CI = –1.66–1.36, k=4; n=677; I² = 75%), drinks per drinking day (3 months: MD: –0.22, 95% CI = –2.51–2.07, k=2, n=359, I²=82%; 6–36 months: MD= –0.09, 95% CI= –0.49–0.30, k=3, n=1450, I²=60%), or percent drinking days (3 months: MD= –4.60, 95%= –21.14–11.94; k=2; n=361; I² = 90%; 6–9 months: MD=1.96, 95% CI= –6.54–10.46; k=2; n=818; I² = 88%).

Conclusion: Psychosocial interventions show promise at increasing self-reported alcohol abstinence in sSA, but clinical, methodological, and statistical heterogeneity across meta-analytic outcomes suggests results should be interpreted with caution.

Introduction

Harmful alcohol use is the seventh leading risk factor for morbidity and mortality globally and has been causally linked to more than 230 diseases and injuries.^{1,2} Although the World Health Organization (WHO) Africa Region has relatively low alcohol per capita consumed (APC) (6.3 liters per person),² APC is high among those who drink (18.4 liters) and is among the highest in the world in some sub-Saharan African (sSA) countries.² Consequently, the region experiences a disproportionately high level of alcohol-related harms.³ Alcohol use is of special concern in sSA given the high prevalence of HIV and tuberculosis,^{4,5} for which alcohol is a risk factor for infection, a catalyst to disease progression, and interferes with treatment adherence and efficacy.^{6,7}

With limited availability of pharmacologic alcohol treatments in low-income settings,⁸ feasible and effective non-pharmacological approaches are needed. Psychosocial interventions, or psychologically based approaches to alcohol reduction, are the most commonly studied non-pharmacological approaches to alcohol reduction (e.g., cognitive-behavioral therapy, brief interventions [BI], family therapy, 12-step programs).⁹ Systematic reviews and meta-analyses of psychosocial interventions have demonstrated efficacy for alcohol reduction in specific settings and subpopulations in resource-rich settings.^{9–12} To-date, one narrative review of BI for alcohol in sSA showed positive results for BIs in health care settings¹³ and a more recent scoping review assessed the amount and types of alcohol interventions in sSA.¹⁴ However, there have been no meta-analyses to quantitatively synthesize the effect of alcohol interventions on consumption.

Structural interventions are another important non-pharmacological approach to alcohol reduction, especially given the alcohol industry's rapid expansion and limited regulation in Africa.^{15,16} Structural interventions aim to change the environments in which risk behavior occurs, such as limiting alcohol availability. Although no systematic reviews exist to-date to assess their effect on alcohol consumption, an increasing number of intervention trials have

demonstrated success in structural approaches at reducing alcohol consumption and related problems in diverse populations and contexts.¹⁷

With an increasing number of studies focused on the evaluation of alcohol interventions in sSA,^{13,14} along with distinct patterns of drinking, comorbidities, and cultural and environmental contexts in this setting, a review focusing exclusively on these types of interventions in sSA is warranted. The consolidation of existing evidence can inform decisions on which interventions should be scaled up to reduce harmful drinking in these settings. Therefore, we conducted a systematic review and meta-analysis to assess the effect of non-pharmacological interventions on alcohol reduction in sSA settings.

Methods

Search strategy and selection criteria

In this systematic review and meta-analysis, Embase, Medline, PsycINFO, EBSCO, CINAHL, and Cochrane CENTRAL were searched on December 21, 2017 for published reports in English from the earliest available date per database. This search was rerun on March 14, 2019. The search protocol is provided in Table S1. Reports were also hand-searched and supplementary data sent by study authors was included.

Inclusion criteria

To be included, studies had to be a randomized or nonrandomized controlled trial, conducted in sSA, assessing a non-pharmacological intervention aimed at alcohol reduction, and measuring at least one alcohol consumption outcome at follow-up greater than one month post-intervention. Eligible comparator groups included interventions unrelated to alcohol, usual care for alcohol or other services, brief feedback on an alcohol screening tool, alcohol or other informational materials, wait-list, and nothing.

Eligible outcomes included alcohol biomarkers (i.e., urine/blood analysis, breathalyzer tests) and self-reported measures: total score for the Alcohol Use Disorders Identification Test (AUDIT),¹⁸ alcohol abstinence (i.e., no drinking vs. any drinking), measures of drinking quantity (e.g., average number of drinks in a specific time period, such as in the prior week or per drinking day) and frequency (e.g., percent drinking days in a specified time period), and drinking intensity (e.g., binge/heavy episodic drinking, such as 4–5 drinks in a 2 hour time period). However, only 3 of the 12 studies reporting binge drinking outcomes had similar definitions, for which the timeframes and standard drink definitions used were unclear. The variability in these outcomes warranted a narrative synthesis. Therefore, the review results were divided into two companion manuscripts to ensure adequate space for reporting. Specifically, the present meta-analysis and a systematic review for the non-pooled outcomes¹⁹ were submitted simultaneously for peer review.

Exclusion criteria

Reasons for exclusion included alcohol reduction not being a primary goal of the intervention; alcohol reduction only being addressed in the context of sexual behavior; no comparator condition; comparator was another evidence-based or 'bona-fide' alcohol

intervention (i.e., non-inferiority trial) and studies without data to be included in the metaanalysis (i.e., not reported and not provided after requested from study authors).

Screening procedures

One author (KS) screened all titles and abstracts, which underwent a targeted review by a second author (AM). If there was disagreement, studies were included in the full-text review. Four authors (AM, JW, KS, SK) and two research assistants reviewed full-text reports and assessed their eligibility in pairs. Disagreements were resolved by discussion and consensus reached between the reviewers or by a third author.

Data extraction and quality assessment

Two reviewers (AM, KS) independently extracted all outcome data into standardized, piloted data collection forms. Characteristics of each study (e.g., design, intervention) were extracted by one reviewer and checked for accuracy by a second reviewer. Per GRADE handbook recommendations, the Population, Intervention, Comparator, Outcome (PICO) framework was used to inform the structure of the data extraction form.²⁰ In this framework, every row of extracted data represents the components of a study essential to answering the review's research questions. The data extraction form was stratified one step further by utilizing the intervention, outcome, population trio (IOPT) structure.²¹ In this structure, each row of extracted data represents a unique data point for analysis, reflecting one intervention and comparator combination (e.g., BI versus standard of care), one outcome (e.g., AUDIT score), in a specific population (e.g., male only, female only, both genders) at a given follow-up interval (e.g., 3 months). In most cases, multiple IOPTs were extracted from each study, which were then grouped together in the meta-analysis by outcome and follow-up interval. All outcome data were independently extracted by both reviewers, compared, and reconciled through discussion. Corresponding authors of studies were contacted to collect relevant data not reported in the paper. Of 15 data requests made, 13 authors responded (response rate: 86%).

Study quality was assessed using the Cochrane Collaboration Tool for Assessing Risk of Bias,²⁰ and three additional bias categories from the GRADE handbook (see Table S2).²² Assessment of risk of bias occurred at the time of data extraction and was assessed at the IOPT level as well as the study level. Each reviewer (AM, KS) independently rated each of the items as low risk, high risk, or unclear. Discrepancies were resolved by discussion. If consensus could not be reached, a third author was asked to break the tie.

Data analysis

Meta-analysis was done to synthesize the effect estimate for alcohol reduction interventions. All analyses were conducted in RevMan version 5. Results of trials with comparable outcomes were pooled using the random effects model and 95% confidence intervals (CIs). For continuous outcomes, mean differences (MD) were calculated between the intervention and comparison groups with 95% CIs. For dichotomous outcomes, odds ratios (ORs) with 95% CIs are presented. Outcomes were compared at different follow-up points, categorizing 2–3 months as short-term, 6–9 months as medium-term, and 12 months or longer as longterm follow-up. To make the maximum number of comparisons between studies, categories

were sometimes merged for meta-analysis (i.e., short-medium [3–6 months], medium-long [6–36 months]). In cases where multiple data points were available from one study within one follow-up interval, we used the longest follow-up period.

In cases where data (i.e., mean/standard deviations; n per outcome) were not available nor provided by the corresponding authors, effect estimates and standard errors (SE) were extracted, if available, or calculated using alternative statistics. Analysis was then conducted under the generic inverse variance outcome in RevMan. In some cases, data transformations were made to synthesize data (e.g., transforming a categorical drinking frequency variable that could not be synthesized with continuous frequency outcomes into no drinking vs. any drinking; transforming number of drinking days in prior 30 days into percent drinking days).

Most cluster randomized controlled trials (CRCTs) accounted for clustering in their analysis. However, too few reported comparable effect and variance measures for our outcomes to use the generic inverse variance method. Therefore, we calculated the design effect by extracting the average cluster size and the intra-cluster coefficient (ICC). We then divided the original sample size by the design effect to reduce the size of the trial to its "effective sample size."²⁰ The ICC was obtained from the published report or from the study authors.

Heterogeneity between comparable trials was tested using a standard chi-squared test and I² statistics, using a p-value of 0.10 or less to determine heterogeneity.^{23,24} I² values are interpreted as low (0%–40%), moderate (30%–60%), substantial (50%–90%), and considerable (75%–100%) heterogeneity.²⁵ We did not conduct quantitative investigations of heterogeneity or subgroup analyses due to the small number of studies per outcome/ follow-up interval with comparable results, as these analyses are not recommended with less than 10 studies.²⁰ We qualitatively explore select factors that might affect heterogeneity through the visual assessment of funnel plots when possible. The symmetry of funnel plots were similarly only visually assessed for publication bias due to an insufficient number of studies (<10) to quantitatively test for symmetry.²⁰ If sufficient information had been available, planned formal subgroup analyses (e.g., gender, intervention dose) were outlined in the study protocol, which was registered on January 5, 2019 with PROSPERO (CRD42019094509) after the initial search and review of studies commenced.

Role of the funding source

This study had no direct funding. Sponsors of the study authors had no role in study design, data collection, data analysis, data interpretation, or writing of the report. The authors had full access to all of the data in the study and had final responsibility for the decision to submit for publication.

Results

The database search identified a total of 1,282 unique citations after the exclusion of duplicates. Six additional studies were identified through hand-searching and correspondence with study authors. Of these citations, 77 reports underwent full-text screening, 53 were excluded for reasons outlined in Figure 1 (a full list of ineligible studies reviewed as full-text is available in Table S3). Of the 24 reports judged eligible

for meta-analysis, 5 were excluded due to missing data on eligible outcomes. In total, 18 studies^{26–43} reported in 19 scientific manuscripts⁴⁴ met criteria for inclusion and contributed data for meta-analysis for the following outcomes: AUDIT score (k=11),^{26,30–34,38,39,41–43} drinks per drinking day (DDD) (k=3),^{29,35,39} percent drinking days (PDD) (k=3),^{28,29,39} and alcohol abstinence (k=7).^{26–29,35–37} No eligible studies included alcohol biomarkers.

Summary of study characteristics

The 18 studies included spanned seven sSA countries: South Africa (k=9), $^{30-36,40}$ Kenya (k=5), $^{26-29,38}$ Namibia (k=1), 37 Rwanda (k=1), Uganda (k=2), 39 Tanzania (k=1), 26 and Zambia (k=1) 26 (including one multi-country study). 26 Study designs were primarily RCTs (k=11). $^{27-29,31-33,37,39-41,43}$ Eight studies tested interventions focused solely or primarily on alcohol reduction (k=5) $^{30-32,38,39}$ or abstinence (k=3), $^{27-29}$ as opposed to dual or multi-outcome focused interventions (k=10). $^{26,33-37,40-43}$ The majority of studies included special subpopulations: people living with HIV (k=7), $^{26,28,29,39-41,43}$ adolescents (n=3, including one with adolescents living with HIV), 36,37,41 female sex workers (k=1), 27 pregnant women (k=2), 35,42 and TB patients (k=1). 30

Nineteen interventions were tested across the 18 studies (one 3-armed study tested an intervention and an enhanced version of that intervention).³⁸ All studies evaluated psychosocial interventions; one structural-level intervention was identified but was excluded because of a lack of appropriate data for meta-analysis. Seven interventions^{27,30–32,38,42} were based on or expanded upon WHO's alcohol BI manual,18 five explicitly stated using motivational interviewing or motivational therapy (inclusive of two of the WHO BI studies),^{27,33,38,39,42} three were grounded in cognitive behavioral therapy,^{28,29,41} and four were informed by a behavior change model.^{30–32,37} The most common intervention setting was a health facility (k=8), 26-30,39,41,42 followed by community venues (k=5), 31-34,40schools (k=3),³⁶⁻³⁸ and participants' homes (k=2).^{35,43} Three interventions were delivered in a single-session, ^{31,32,39} two did not reported the number of sessions, ^{26,34} while all others were multi-session interventions. Comparator groups were: feedback on AUDIT results and/or general information or an educational leaflet on alcohol (k=6), 30-33,38,42standard-of-care for a range of services (k=7),^{26,29,35,39–41,43} nutrition/lifestyle intervention (k=4),^{27,28,36,40} and delayed intervention (k=2).^{34,37} Study design and intervention details are provided in Table 1.

Meta-analysis results

The meta-analyses that focused on AUDIT scores found no statistically significant differences between intervention and comparator at 2–3 months, 6-months, or 12-months post-intervention (Figure 2). No statistically significant differences were found for DDD at 3 or 6–36 months or PDD at 3 or 6–9 months (Figures 3 & 4). The meta-analysis of trials on alcohol abstinence showed a beneficial effect of psychosocial interventions versus comparator at 3–6 months post-intervention. The effect on alcohol abstinence was also statistically significant for trials assessing long-term follow-up (12–60 months) (Figure 5).

A moderate to considerable level of heterogeneity was identified across all analyses (I² between 60% and 90%). Qualitative comparison identified one factor that appeared to drive

differences in effects on abstinence. Exploratory funnel plots in Figure 6 demonstrates larger effect sizes for studies that included drinking (any drinking or specified risk-level) at baseline in their inclusion criteria compared to studies that did not.

Publication bias results

Figure S1 presents funnel plots for AUDIT scores and abstinence outcomes to assess publication bias. Too few studies (k=2-3) were reported to make comparisons for PDD and DDD.²⁰ The plots were overall symmetrical; therefore, no publication bias was detected. Studies tended to cluster at the top of the plot, indicating more publication of studies with larger sample sizes. Clustering at one end can indicate small study bias. However, this concern is mitigated; small studies with positive effects were not more likely to be published; rather, larger studies were more likely to be published regardless of effect.

Risk of bias assessment results

In general, studies evaluated with the Cochrane risk of bias tool were of moderate quality (see Figure 7). Randomization procedures were properly described in 75% of studies, and half of the studies reported details on allocation concealment. No studies blinded both participants and study personnel, and less than 25% blinded outcome assessment – potential sources of performance and detection bias. Other weaknesses included a lack of published study protocols resulting in high risk for selective reporting bias, and flawed measurement of exposure (i.e., a lack of information on intervention dose and fidelity). See Figure S2 and Table S4 for the full risk of bias assessment per study and outcome.

Discussion

Despite high rates of alcohol-related morbidity and mortality in sSA,² this is the only metaanalysis to compare the effect of psychosocial interventions versus a comparator on alcohol consumption among individuals in sSA to-date. Our results are specific to psychosocial interventions; only one structural intervention met our inclusion criteria,⁴⁵ but did not have data available for meta-analysis. Two main findings emerged from this systematic review and meta-analysis. First, psychosocial interventions appear to have a benefit on alcohol abstinence at both short to medium and long-term follow up. Second, psychosocial interventions showed no significant effect on AUDIT score, DDD, and PDD.

Our review reinforces the need for research aimed to develop and test alcohol interventions in sSA. In line with Francis et al.'s scoping review,¹⁴ this review demonstrates that the number of studies with this aim is disproportionately low compared to the burden of alcohol use problems in the region and are heavily concentrated in South Africa and Kenya. Still, the beneficial effect identified for alcohol abstinence outcomes shows promise for the use of alcohol psychosocial interventions in sSA with this aim.

Potential sources of heterogeneity and varied effects

Drinking at baseline and alcohol-only vs. multi-component interventions—The heterogeneity identified across outcomes was moderate to high. While the total number of studies per outcome restricted quantitative comparisons, qualitative exploratory analysis

identified larger effect sizes among studies that included any type of drinking at baseline as part of the study's inclusion criteria compared to those that did not for alcohol abstinence (Figure 6). These differences may be attributed to regression to the mean, or more room for change for those already drinking at baseline. However, the same two studies in the short-medium term follow-up assessment that included those drinking at baseline were also the only two studies to focus solely on alcohol reduction. Therefore, we cannot tease apart the effects of these subcomponents within the short-medium analyses. It is possible that alcohol-focused interventions had a stronger impact than those with multiple outcomes, as we are unable to determine the intervention dose specific to alcohol in these studies. Given multiple alcohol-involved "syndemics" in African settings (i.e., two or more epidemics interacting synergistically to contribute to excess burden of disease in a population), about 45% of the interventions tested included alcohol reduction as a subcomponent of a multicomponent intervention aimed at more than one health behavior.

Population—It is possible that the observed abstinence effects are influenced by overreporting of self-reported alcohol abstinence due to social desirability, especially for certain populations (e.g., pregnant women).⁴⁷ No clear patterns emerged in intervention effect by population, but the wide variability of populations included limited even qualitative comparison. However, studies measuring alcohol abstinence exclusively included subpopulations for which alcohol abstinence was an appropriate goal (i.e., HIV populations, female sex workers, pregnant women, adolescents). It is possible that these studies achieved greater effects than other alcohol outcomes given unique motivations to not drink among these subpopulations.

Measurement bias—The effect sizes for abstinence may be exaggerated by the binary nature of the measure.⁴⁶ The null findings for AUDIT scores could also be an effect of measurement bias. The AUDIT is designed to identify high-risk drinking patterns, with half of the questions assessing occurrence of alcohol-related problems or negative consequences of alcohol use in the past year. Of the 8 studies reporting AUDIT change at less than 12 months follow-up, 2 did not explicitly state changing the timeframe of the questions to match their shorter follow-up period. These studies were included with the assumption that this change was made, but it is possible that the scale's timeframe was not modified for all studies, reducing the likelihood that change would be observed in less than 12 months. Further, two of the AUDIT questions assess current or past lifetime harmful drinking. Thus, studies with less than 12 months follow-up may not show significant change in AUDIT scores even when the questions are modified to assess change in a 2 to 6 month timeframe. No studies included alcohol biomarkers, such as blood alcohol concentration (BAC) or Phosphatidylethanol (PEth), which has been shown to be more reliable than self-report in African cohort studies.⁴⁷ Taken together, these findings highlight self-reported and inconsistent alcohol outcome measurement as a weakness of the alcohol-focused intervention literature in sSA.

Comparators—Comparators can drive effect size magnitude. The wide variability of comparators within each outcome assessment limited the ability to make any meaningful conclusions about their influence on effect size across outcomes. However, a number

of BI studies and one CBT study found no significant differences between intervention participants compared to minimal intervention, but reductions in drinking were observed in both treatment arms. ^{31–33,41,42} This nuance is not apparent in our meta-analysis results but may be a driver of the AUDIT meta-analyses' null effects.

Intervention—At this stage, the picture remains unclear on which intervention approaches show the most promise. Alcohol interventions for groups with special health concerns or other reasons not to drink (i.e., adolescents, female sex workers) that showed promise were conducted across a wide set of settings, using a range of psychosocial approaches, including CBT, MI, and other broad psychosocial group and individual-focused approaches. More research is needed to provide pointed policy and practice recommendations on which interventions work in different settings. Future research will also be needed to inform the cost-effectiveness and feasibility of scaling up these approaches in resource-limited settings. A cost-benefit analysis associated with the Kenya CBT study²⁸ included in this review reported CBT can be effectively and economically task-shifted to paraprofessionals in Kenya.⁴⁸ Additional costing studies, along with hybrid implementation studies that simultaneously assess implementation and effectiveness, can inform the feasible scale up of alcohol interventions in settings with resource-constraints.

Individual-level focus—This review demonstrates alcohol interventions in sSA to-date are overwhelmingly focused on individual, rather than structural-level, change. Despite a large number of BI studies based on MI and the WHO SBI guidelines, evidence for change in AUDIT scores using this approach remains limited in sSA, contrary to a body of literature supporting moderate effects using this approach in well-resourced settings.¹⁰ Beyond the measurement and methodological limitations already noted, a possible explanation for the underwhelming effects of these interventions may be their lack of focus on the social and physical environment.⁴⁹ Alcohol outlet density,⁵⁰ aggressive alcohol marketing,⁵¹ and lax alcohol regulation and policy enforcement⁵² are prevalent contributors to alcohol consumption in African settings. More rigorous research that tests interventions altering the social and physical environment, or other structural approaches, are needed.

Limitations

Limitations of this systematic review include challenges in the ability to synthesize all eligible studies due to the disparate measurement of alcohol outcomes at varying time points, resulting in low statistical power for some outcomes assessed. Low statistical power may have contributed to the null findings for DDD and PDD outcomes in particular. While variability of outcome measures is a known issue in the alcohol intervention field,⁵³ our broad inclusion criteria likely also contributed to the broad set of outcomes identified. The especially variable measurement of heavy episodic/binge drinking limited our ability to present these findings alongside the outcomes in this review, which are included in a forthcoming narrative synthesis.¹⁹ In addition, we identified significant heterogeneity across studies with limitations in our ability to conduct comprehensive, quantitative assessments of differences by study design, intervention, and population, as discussed above. Our qualitative comparisons are exploratory in nature, and should be reviewed with caution

as they include less than ten studies per outcome/timepoint.²⁰ As the alcohol intervention literature in sSA continues to grow, this should be a focus of future reviews.

Our inclusion criteria allowed for both non-randomized and randomized controlled trials. Though only two non-randomized trials were included, they bring inherent risk of selection bias. Moreover, the risk of bias assessment identified risk in randomization and allocation concealment in a number of randomized studies. These and other risks of bias identified should be considered in the interpretation of our findings. Moreover, several interventions were "pilot" studies which may be less robust in design and intervention content – a reflection of the developmental stage of the alcohol intervention literature specific to $sSA.^{29,34,40}$ The studies were judged as moderate quality, demonstrating a need for added rigor in the assessment of future alcohol interventions through randomized controlled trials.

Conclusion

This review highlights the need for more research testing alcohol interventions in sSA. Null findings were identified for interventions assessing change in AUDIT, DDD, and PDD across a range of sSA contexts. However, the review showed some promise for psychosocial interventions to promote alcohol abstinence. Given the wide scope of this review, significant heterogeneity was identified across studies. As the pool of research grows in this area, more direct investigations of differences across population, setting, design, and intervention type would provide more pointed guidance on the context-specific application of research to alcohol policy and programming in sSA.

With detrimental health and societal effects of harmful alcohol use affecting sSA and limited access to pharmacological alcohol interventions, research to develop acceptable and feasible non-pharmacological interventions for sSA should be prioritized. The literature on alcohol-focused interventions in sSA would benefit from more rigorous designs, consistency across alcohol outcomes, the inclusion of alcohol biomarker outcomes, and the systematic assessment of structural approaches to alcohol reduction in addition to the current literature focused on individual-level psychosocial interventions.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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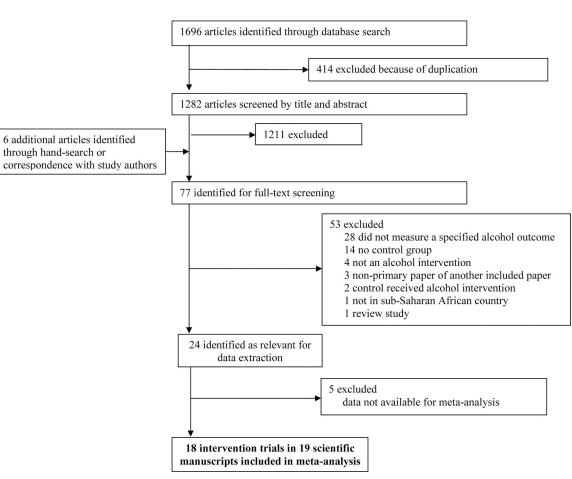


Figure 1. Studies included in systematic review and meta-analysis

Adapted from the 2009 PRISMA Flow Diagram. Moher D, Liberati A, Tetzlaff J, Altman DG, The PRISMA Group. Preferred Reporting Items for Systematic Reviews and MetaAnalyses: The PRISMA Statement. PLoS Med 2009;**6**(7): e1000097. doi:10.1371/journal.pmed1000097

Short-term follow-up (2-3 months)

	Exp	erimen	tal	0	Control			Mean Difference	Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI	
Takahashi, 2018	8.07	3.34	49	11.5	3.32	44	16.1%	-3.43 [-4.79, -2.07]		
Takahashi, 2018	8.33	3.34	49	11.5	3.32	44	16.1%	-3.17 [-4.53, -1.81]		
Marais, 2011	0.46	3.84	80	2.43	3.91	68	16.5%	-1.97 [-3.22, -0.72]		
Chaudhury, 2016	1.96	14.63	65	2.25	19.53	66	4.7%	-0.29 [-6.19, 5.61]		
Senyonyi, 2012	0.26	1.55	81	0.47	2.57	34	17.4%	-0.21 [-1.14, 0.72]	+	
Peltzer, 2013	5	6.1	84	4	5.9	37	12.8%	1.00 [-1.31, 3.31]		
Wandera, 2017	7.6 5.4 144 6.5 5.6 1			147	16.4%	1.10 [-0.16, 2.36]				
Total (95% CI)	552 440						100.0%	-1.13 [-2.60, 0.34]	•	
Heterogeneity: Tau ² =	= 3.02; C	hi ² = 40	.59, df :	= 6 (P <	0.0000	1); = 1	85%			-
Test for overall effect						- 25			-20 -10 0 10 Favours [experimental] Favours [control]	20

Medium follow-up (6 month follow-up)

	Expe	erimen	tal	C	ontrol			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Takahashi, 2018	6.64	3.32	49	9.4	3.38	48	16.8%	-2.76 [-4.09, -1.43]	-
Takahashi, 2018	7.72	3.32	47	9.4	3.38	48	16.7%	-1.68 [-3.03, -0.33]	-8-
Peltzer, 2013*	2.4	4.8	84	3.6	6.2	57	13.3%	-1.20 [-3.11, 0.71]	
Rotheram-Borus, 2016*	16* 6.7 5.6 41 7				7.4	39	8.7%	-0.70 [-3.59, 2.19]	
Pengpid, 2013a	11.3	7.6	59	11.6	7.7		8.7%	-0.30 [-3.20, 2.60]	
Wandera, 2017	7.7	5.2	163	7.5	5.6		17.8%	0.20 [-0.98, 1.38]	+
Pengpid, 2013b	7 4.5 129			6.3 4.6	4.6	109	17.9%	0.70 [-0.46, 1.86]	
Total (95% CI)			572			509	100.0%	-0.83 [-1.92, 0.26]	•
Heterogeneity: Tau ² = 1.3	9; Chi ² =	19.33	, df = 6	(P = 0.0))04); lª	= 69%		-	
Test for overall effect: Z =					1				-20 -10 0 10 20 Favours [experimental] Favours [control]

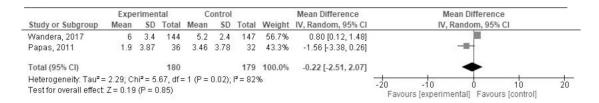
Long-term follow-up (12 month follow-up)

		erimen			ontrol			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Rendall-Mkosi, 2013	12.91	8.94	61	17.16	8.86	64	14.4%	-4.25 [-7.37, -1.13]	
Pengpid, 2013b	7.2	5.8	143	7.3	6.8	139	27.6%	-0.10 [-1.58, 1.38]	
Bachanas, 2016 [*]	4.89	0.68	60	4.96	0.62	63	37.2%	-0.07 [-0.30, 0.16]	
Pengpid, 2013a	13.5	7.3	79	11	6.4	68	20.8%	2.50 [0.29, 4.71]	
Total (95% CI)			343			334	100.0%	-0.15 [-1.66, 1.36]	+
Heterogeneity: Tau ² =	1.59; Chi	² = 12.	06, df=	3 (P =	0.007)	; I ² = 75	5%		
Test for overall effect: 2									-20 -10 0 10 Favours [experimental] Favours [control]

Figure 2. Results of meta-analyses with Alcohol Use Disorders Identification Test (AUDIT) score by follow-up period

Note: *indicates cluster randomized controlled trials (CRCT) for which the sample size was adjusted by design effect

Short-term follow-up (3 months)



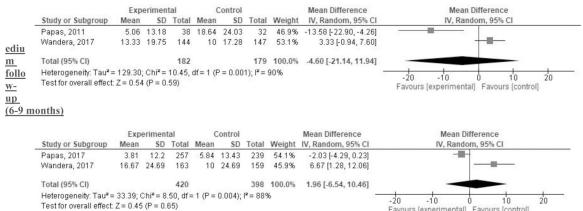
Medium to long-term follow-up (6-36 months)

	Expe	erimen	tal	С	ontrol			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Papas, 2017	1.29	2.96	257	1.86	3.12	239	27.3%	-0.57 [-1.11, -0.03]	8
Rotheram-Borus, 2015 *	2	0.96	330	2	0.93	302	50.9%	0.00 [-0.15, 0.15]	
Wandera, 2017	6	2.8	163	5.7	3.2	159	21.8%	0.30 [-0.36, 0.96]	*
Total (95% CI)			750			700	100.0%	-0.09 [-0.49, 0.30]	•
Heterogeneity: Tau ² = 0.0	7; Chi ² =	5.02,	df = 2 (P = 0.08	3); I ^z =	60%			-20 -10 0 10 20
Test for overall effect: Z =	0.45 (P	= 0.65))						Favours [experimental] Favours [control]

Figure 3. Results of meta-analyses with drinks per drinking day (DDD) by follow-up period Note: *indicates cluster randomized controlled trials (CRCT) for which the sample size was

adjusted by design effect

Short-term follow-up (3-months)



Favours [experimental] Favours [control]

Figure 4.

Results of meta-analyses with percentage of drinking days (PDD) by follow-up period

Short to medium-term follow-up (3-6 months)

	Experim	ental	Contr	rol		Odds Ratio	Odds Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
Papas, 2011	25	36	12	32	14.0%	3.79 [1.38, 10.37]	· · · · · · · · · · · · · · · · · · ·
L'Engle, 2014	186 34		93	355	25.7%	3.27 [2.38, 4.50]	
Stanton, 1998	167	209	118	170	23.1%	1.75 [1.10, 2.80]	
Bachanas, 2016 *	45 51 43			53	12.9%	1.74 [0.58, 5.21]	
Rotheram-Borus, unpublished	518	573	436	487	24.3%	1.10 [0.74, 1.65]	
Total (95% CI)		1215		1097	100.0%	2.05 [1.20, 3.48]	+
Total events	941		702				
Heterogeneity: Tau ² = 0.26; Chi ²	= 19.44, dt	= 4 (P =	= 0.0006)	; I ² = 79	1%		
Test for overall effect: Z = 2.64 (F	= 0.008)	- 11					0.05 0.2 1 5 20 Favours [control] Favours [experimental]

Long-term follow-up (12-60 months)

			xperimental			Odds Ratio		Odds Ratio
Study or Subgroup	log[Odds Ratio]	SE	Total	Total	Weight	IV, Random, 95% CI		IV, Random, 95% CI
Zule, 2014	1.2837	0.5493	42	42	6.6%	3.61 [1.23, 10.59]		
L'Engle, 2014	1.1047	0.1563	350	360	23.3%	3.02 [2.22, 4.10]		
Smith, 2008 *	0.5311	0.1746	225	333	22.1%	1.70 [1.21, 2.39]		
Stanton, 1998	0.4484	0.2353	201	158	18.2%	1.57 [0.99, 2.48]		
Rotheram-Borus, unpublished *	0.4353	0.1629	477	443	22.9%	1.55 [1.12, 2.13]		
Bachanas, 2016	0.1178	0.5297	53	53	6.9%	1.13 [0.40, 3.18]		
Total (95% CI)			1348	1389	100.0%	1.91 [1.40, 2.61]		•
Heterogeneity: Tau ² = 0.08; Chi ² :	= 13.66, df = 5 (P =	0.02); 2=	63%					<u> </u>
Test for overall effect: Z = 4.08 (P							0.05 0.2 Favour	1 5 s [control] Favours [experiment

Figure 5. Results of meta-analyses with alcohol abstinence by follow-up period

Note: *indicates cluster randomized controlled trials (CRCT) for which the sample size was adjusted by design effect; note the direction of the intervention effect differs from the previous continuous outcomes to reflect the desired outcome of greater abstinence.

Short to medium-term follow-up (3-6 months) Long-term follow-up (12-60 months) "T "T "T "T

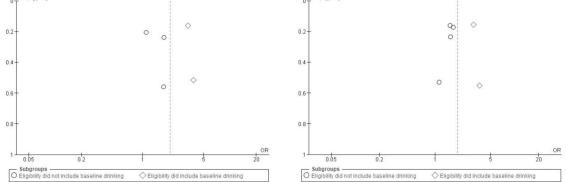


Figure 6.

Funnel plot of comparison: Alcohol abstinence outcome by subgroups, eligibility did include baseline drinking vs. eligibility did not include baseline drinking

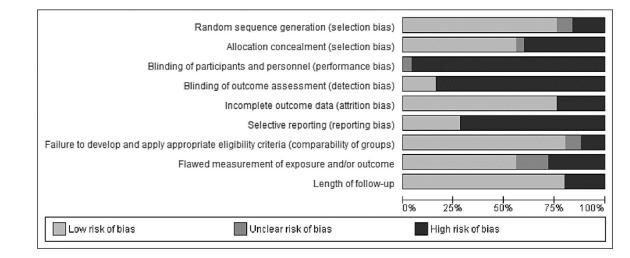


Figure 7.

Risk of bias graph: review authors' judgements about each risk of bias items presented as percentages across all included studies

_											
	Alcohol outcome(s) for meta-analysis	AUDIT score Abstinence	AUDIT score	Abstinence	AUDIT score	Drinks per drinking day Percent drinking days Abstinence	Percent drinking days	AUDIT score	AUDIT score	AUDIT score	AUDIT score
	Comparator /time & attention matched?	HIV standard-of-care / No	HIV standard-of-care for social work support services / No	Nutrition control intervention / Yes	AUDIT screening and alcohol booklet (2 sessions) / No	Routine medical care / No	Group healthy lifestyles education control intervention / Yes	Health education leaflet on responsible drinking / No	Feedback on AUDIT + health education leaflet on responsible drinking / No	Health education leaflet on responsible drinking / No	Information pamphlet on fetal alcohol syndrome
	Format and setting	Individual and group format, delivered by health workers and lay workers, in a health facility	Individual and group (family) format delivered by a trained counselor in participants' homes	Individual format delivered by nurse counselors in a health facility setting	Individual format delivered by trained research staff in a health facility setting	Group format delivered by a trained counselor in a health facility setting	Group format delivered by a trained counselor in a health facility setting	Individual format delivered by a lay counselor in a health facility	Individual format delivered by a nurse research assistant in a public venue	Individual format delivered by a nurse research assistant in a public venue	Individual format delivered by a lay worker in a venue of participants' choice
	Dose	NR	90 minutes (single session carried out over everal, plus additional follow-ups to check progress)	120 minutes (six 20- minute sessions over 6 months)	120 minutes (4 sessions: first session 1 hour, follow-ups 20 minutes)	540 minutes (6 weekly 90-minute sessions)	540 minutes (6 weekly 90-minute sessions)	30-40 minutes(Two 15- 20-minute sessions within 1 month)	20 minutes (single session)	20 minutes (single session)	5 sessions (duration NR)
	Counseling/theoretical approach	NR	Psychosocial family approaches teated to restlience building, family communication, parenting skills, HTV psycho-education; engagement of formal and informal supports	WHO Brief Intervention, Motivational Interviewing	WHO Brief Intervention, Motivational Interviewing	Cognitive Behavioral Therapy	Cognitive-Behavioral Therapy	WHO Brief Intervention + content informed by Information- Motivation-Behavioral Skills Model	WHO Brief Intervention + content informed by Information- Motivation-Behavioral Skills Model	WHO Brief Intervention + content informed by Information- Motivation-Behavioral Skills Model	Motivational interviewing
	Intervention focus	HIV prevention, with alcohol components	Family strengthening with content on HIV, violence reduction, and alcohol reduction	Alcohol abstinence, some focus on alcohol and sexual risk	Alcohol abstinence, pregnancy-focus	Alcohol abstinence	Alcohol abstinence	Alcohol reduction	Alcohol reduction	Alcohol reduction	Fetal alcohol syndrome prevention
	%female*	58.10%	68.29%	100.00%	100.00%	NR (females and males included)	NR (females and males included)	25.70%	12.70%	27.60%	100.00%
	Age [*] (SD)	36 (SD NR)	41.03 (8.76)	27.5 (6.6)	25 (SD NR)	37.07 (8.40)	NR	36.7 (10.9)	21.9 (3.5)	35.6 (11.45)	29.8 (SD NR)
	Total N*	3538	62	818	194	75	614	1196	152	392	165
	Population group	PLHIV attending clinical care	HIV-infected caregivers with at least one school aged child; women and men: Alcohol analyses only run for those who drank at baseline	Female Sex Workers, moderate risk drinkers (AUDIT = $7-19$)	Pregnant women reporting any drinking	PLHIV reporting binge or hazardous drinking (AUDIT-C = 3 or 6 or more drinks per occasion at least monthly)	PLHIV reporting binge or hazardous drinking (AUDIT-C = 3 or 6 or more drinks per occasion at least monthly)	Tuberculosis outpatients misusing alcohol (AUDIT = 8 or more for men; AUDIT = 7 or more for women)	University students screened as at-risk drinkers (AUDIT > 8)	Outpatients screened as hazardous or harmful drinkers (AUDIT = $8-19$ for men and $7-19$ for women	Women at high-risk of alcohol-affected pregnancy (not using contraceptives
	Study design	CRCT	RCT	RCT	CRCT	RCT	RCT	CRCT	RCT	RCT	RCT
	Data years	2009– 2011	NR	2011– 2012	2007– 2008	2009	2012– 2016	2011– 2012	2011– 2012	2011– 2012	2007– 2008
	Country	Kenya, Tanzania, Zambia	Rwanda	Kenya	South Africa	Kenya	Kenya	South Africa	South Africa	South Africa	South Africa
	Author, year / peer-reviewed	Bachanas, 2016 / peer-reviewed	Chaudhury, 2016 / peer-reviewed	L'Engle, 2014 / peer-reviewed	Marais, 2011 / peer-reviewed	Papas, 2011/peer- reviewed	Papas, 2017/not peer-reviewed	Peltzer, 2013/peer- reviewed	Pengpid, 2013a / peer-reviewed	Pengpid, 2013b / peer-reviewed	Rendall-Mkosi, 2013/peer-reviewed

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Table 1.

Summary of study characteristics of alcohol interventions in sub-Saharan Africa included in meta-analysis

Author, year / peer-reviewed	Country	Data years	Study design	Population group	Total N*	Age [*] (SD)	%female*	Intervention focus	Counseling/theoretical approach	Dose	Format and setting	Comparator /time & attention matched?	Alcohol outcome(s) for meta-analysis
				and engaging in risk drinking)								prevention and a handbook onwoman's health / No	
Rotheram-Borus, 2015/peer-reviewed & Rotheram- Borus, unpublished manuscript/not peer reviewed	South Africa	2009– 2014	CRCT	Pregnant women	1238	26.53(5.63)	100.00%	Maternal health package (HIV, TB, actopid, mental health, breastfeeding, malnutrition healthy behaviorchange)	NR	240 minutes(Eight 30- minute sessions, 4 prenatal and 4 postnatal) averaging about 30 minutes each	Individual format delivered by a Community Health Worker in participant's homes	Standard-of-care services offered at health clinic within 5 km radius / No	Drinks per drinking day Abstinence (unpublished data)
Rotheram-Borus, 2016/peer-reviewed	South Africa	NR	CRCT	Unemployed young men (age 18–25)	142	21.9 (1.9)	0.00% (men only)	HIV prevention, alcohol and drug reduction	Psychosocial approaches (e.g., goal setting, problem solving, praise, social rewards). Subset of men received vocational training.	NR	Individual and group format delivered by uraned soccer coaches in the context of a community soccer program	Delayed intervention / Yes	AUDIT score
Senyonyi, 2016 / peer-reviewed	Uganda	2011	RCT	HIV-infected adolescents	171	15.24 (1.96)	54.80%	Sexual risk and substance use reduction	Cognitive Behavioral Therapy	640 minutes (Eight 80- minute sessions over 8 weeks)	Group format delivered by a trained counselor in a health facility	Standard group counseling / Yes	AUDIT score
Smith, 2008/peer- reviewed	South Africa	2003	NRCT	Adolescent students	2176	14 (0.86)	51.50%	Leisure, life skills, substance use, and sexuality education intervention	NR	15 hours (Twelve 50- minute lessons in grade 8, followed by 6 booster lessons in grade 9)	Group format delivered by educators in a school setting	Standard-of-care (Life Orientation curriculum from the South AfricanDepartment of Education) / No	Abstinence
Stanton, 1998 / peer-reviewed	Namibia	1996– 1997	RCT	Adolescents (aged 15-18)	515	17 (medi an, SD NR)	54.00%	HIV, sexual health, alcohol reduction, violence reduction	Social Cognitive Theory	28 hours (14 sessions, 2 hours each)	Group format delivered by volunteer teachers and an out- of-school youth in an after- school program	Delayed intervention control / Yes	Abstinence
Takahashi, 2018 / peer-reviewed	Kenya	2015	NRCT	Hazardous or hamful drinkers (AUDIT = 8–19)	161	43.6 (SD NR)	18.25%	Alcohol reduction	Two intervention arms: 1) WHO Brief Intervention 2) WHO Brief Intervention + Motivational Therapy	 Brief Intervention Brief Intervention minutes (three 5-20- minute sessions) Brief Intervention Brief Intervention 4 additional session, duration NR 	 Brief Intervention Only: Individual format delivered by a format delivered by a community Health Worker in a local school Brief Intervention Brief Inte	General information about alcohol / No	AUDIT' score
Wandera, 2017 / peer-reviewed	Uganda	2013– 2014	RCT	PLHIV identified as hazardous drinkers (AUDIT-C > 2)	337	39 (medi an, IQR: 32–46)	34.40%	Alcohol reduction	Motivational interviewing	20–30 minutes (single session)	Individual format delivered by a trained counseling in a health facility	Standardized positive prevention counseling / Yes	AUDIT score Drinks per drinking day Percent drinking days
Zule, 2014/peer- reviewed	South Africa	2008– 2011	RCT	WLHIV (age 18–33) who drink	84	23.35 (3.95)	100.00%	HIV-focused intervention, focused on: alcohol and drug use, sexual risk, violence, gender inequality	NR	4 hours (Four 1-hour sessions delivered over 2 contact points)	Group format delivered by a peer educator in a community setting	Two control groups combined for analysis: Nurrition intervention and HIV Counseling and Testing group	Abstinence
* indicates as reporte Alcohol Use Disorde	ed at baseline; ers Identificati	RCT = ran ion Test; DI	domized cc DD = Mear	* indicates as reported at baseline; RCT = randomized controlled trial; CRCT = cluster randomized controlled trial; NRCT = non-randomized quasi-experimental Alcohol Use Disorders Identification Test; DDD = Mean drinks per drinking day; PDD = percentage of days drank; IQR = interquartile range; NR = Not reported	er randomi)D = perce	ized controlled to	rial; NRCT = 1 ank; IQR = int	on-randomized quasi-ex erquartile range; NR = P	sperimental controlled tric	al; PLHIV=people living v	<pre># indicates as reported at baseline; RCT = randomized controlled trial; NRCT = non-randomized quasi-experimental controlled trial; PLHIV=people living with HIV; women living with HIV=WLHIV; TB = Tuberculosis; AUDIT = Auohol Use Disorders Identification Test; DDD = Mean drinking day; PDD = percentage of days drank; IQR = interquartile range; NR = Not reported</pre>	IV=WLHIV; TB = Tubercule	sis; AUDIT =

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