## Adolescent Health, Medicine and Therapeutics

#### Open Access Full Text Article

REVIEW

# Screening for HIV and linkage to care in adolescents: insights from a systematic review of recent interventions in high-versus low- and middle-income settings

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Introduction: Compared to adults, adolescents and young adults have a higher incidence of HIV infection, yet lower rates of HIV testing. Few evidence-based interventions effectively diagnose new HIV infections among adolescents while successfully providing linkage to care. Methods: We conducted a systematic review of recent interventions to increase HIV testing among adolescents and young adults using data retrieved from PubMed and Google Scholar, and using abstracts presented at the International AIDS Society conferences and Conference on Retroviruses and Opportunistic Infections published between January 1, 2015, and April 28, 2018. Results: We identified 36 interventions (N=14 in high- income countries and N=22 in low- and middle-income countries) that were published in the literature (N=28) or presented at conferences (N=8). Interventions were categorized as behavioral/educational, alternate venue/self-testing, youth-friendly services, technology/mobile health, incentives, or peer-based/community-based interventions. The studies consisted of randomized controlled trials (RCTs), prospective and retrospective observational studies, and quasi-experimental/pre-post evaluations with variable sample sizes. Study designs, populations, and settings varied. All categories showed some degree of acceptability, yet not all interventions were effective in increasing HIV testing. Effectiveness was seen in more than one RCT involving technology/mobile health (2/3 RCTs) and alternative venue/self-testing (3/3 RCTs) interventions, and only in one RCT each for behavioral interventions, community interventions, and incentives. There were no effective RCTs for adolescent-friendly services. Data were limited on the number of new infections identified and on the methods to increase linkage to care after diagnosis.

**Conclusion:** Future studies should include combinations of proven methods for engaging adolescents in HIV testing, while ensuring effective methods of linkage to care. Keywords: adolescent, HIV, testing, interventions, barriers

#### Introduction

Worldwide in 2016, an estimated 2.1 million adolescents aged 10–19 years were living with HIV.1 Globally, one-third of all new HIV infections occurs among adolescents.2 Eighty percent of all adolescent HIV infections worldwide occur in sub-Saharan Africa where females are disproportionately affected compared to males.<sup>1,3</sup> In sub-Saharan Africa, less than a third of all adolescents have ever tested for HIV and only 20% of adolescent girls who are living with HIV know their HIV status.<sup>1,3</sup> There are limited evidence-based interventions targeting this population that effectively diagnose and link adolescents and young adults to care.

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In the US, more than 61,000 adolescents are living with HIV.<sup>4</sup> In 2016, of all the new HIV diagnoses among adolescents in the US, 81% were attributed to male-to-male sexual contact.<sup>4</sup> Despite the Centers for Disease Control and Prevention (CDC) recommendations for one-time HIV testing for all individuals aged 13–64 and annual testing in high-risk groups, testing rates among adolescents and young adults remain low.<sup>4,5</sup> Among high school aged men who have sex with men (MSM), only 21% ever had an HIV test.<sup>4</sup> Despite numerous interventions to increase HIV testing among high-risk adolescents and young adults, 44% of those living with HIV have not been tested and are unaware of their positive status.<sup>4</sup>

The Joint United Nations Programme on HIV and AIDS has set the target of 90-90-90 by the year 2020, describing the percentage goals for HIV testing, antiretroviral therapy (ART) initiation, and viral suppression, respectively.<sup>1</sup> In addition, the WHO has recommended moving to a treatment for all strategy to increase the number of individuals living with HIV who receive ART regardless of CD4 cell count or clinical stage.6 Worldwide, adolescents are falling well short of the targeted testing, ART initiation, and viral suppression goals. In the care continuum, estimates of viral suppression among all adolescents living with HIV are <10%.<sup>7,8</sup> The largest drop off in the continuum of care for adolescents is in HIV testing and linkage to care where only 41% know their diagnosis.<sup>4,8</sup> The ultimate goal of HIV testing is diagnosing new infections, linking individuals to care, and achieving viral suppression; yet there are significant gaps in evidence-based interventions to improve HIV testing and linkage to care for adolescents.

Adolescents face numerous barriers to HIV testing as indicated in Table 1. One of the most common psychological barriers to HIV testing among adolescents is lack of perceived risk.9,10 Other psychological barriers include fear of consequences of a positive test, worries about discrimination and rejection, stigma about HIV, sexual orientation, or gender identity.9,11-17 In addition, there are structural barriers to HIV testing among adolescents including never being offered an HIV test, inconvenient hours, lack of insurance, and parental consent.<sup>10,16,18-21</sup> Mistrust of the health care system and perception of poor attitudes of health care providers also hinder HIV testing for adolescents.<sup>11</sup> Social factors such as socioeconomic status, gender, and race can also impede HIV testing in adolescents. Interventions to improve HIV testing among adolescents should target these barriers to increase HIV testing and linkage to care.

Improving HIV testing and linkage to care is now recognized as a global health priority, and as a result, several interventions have been developed specifically targeting Table I Barriers to HIV testing for adolescents

Concerns about correct self-test operation <sup>82</sup>
Confidentiality using parental insurance/parental consent <sup>16,19</sup>
Conflicts with school hours <sup>20</sup>
Cost <sup>21,82</sup>
Drug use <sup>24</sup>
Fear of disclosure <sup>15</sup>
Fear of heterosexist bias <sup>13,83</sup>
Fear of job consequences <sup>84</sup>
Fear of rejection from partners/parents/peers <sup>9,14–17,84</sup>
Fear of results <sup>9,15–17</sup>
Inaccurate information from testing locations <sup>20</sup>
Inconvenience <sup>18</sup>
Lack of knowledge/low health literacy <sup>12,15,17</sup>
Lack of youth-friendly/Lesbian, Gay Bi, Trans (LGBT)-friendly spaces <sup>15</sup>
Lack of social support <sup>82</sup>
Never offered HIV test <sup>10</sup>
Patient-provider trust/communication <sup>11,19,83</sup>
Perceived healthy status/susceptibility/risk9-12,14
Privacy concerns <sup>18,19,83</sup>
Racial/ethnic differences <sup>23,24</sup>
Sex differences (male/female) <sup>24</sup>
Socioeconomic status <sup>25</sup>
Stigma, HIV-related <sup>9,12–14,16,1</sup>
Stigma, LGBT-related <sup>13,23</sup>

HIV testing uptake among adolescents.<sup>22</sup> We conducted a systematic review of interventions published between 2015 and 2018 targeting HIV testing among adolescents to highlight the lack of evidence-based, successful interventions that find new HIV infections among adolescents and successfully link them to care.<sup>23–25</sup>

## **Methods**

We performed a systematic review of HIV testing interventions targeting adolescents that were published between January 2015 and April 28, 2018. We adhered to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines.<sup>26</sup> We initially searched peer-reviewed journals written in English that were located in PubMed and Google Scholar and published after 2010. We then narrowed our results to those published on or after January 1, 2015. Our target population was adolescents and young adults aged 10–24 years. However, we included studies that included individuals outside of this age range provided that the intervention was targeted toward adolescents and young adults. Keywords searched included HIV, testing, and at least one of the following age terms: adolescent, adolescence, teen, youth, or young adults.

In addition, we searched for abstracts presented at the International AIDS Society Conference (IAS) and at the Conference on Retroviruses and Opportunistic Infections (CROI). We were able to search 2015–2018 CROI abstracts, but only 2015–2016 IAS abstracts were available online. Keywords searched included HIV and testing. Abstracts were then screened for at least one of the following age terms: adolescent, adolescence, teen, youth, or young adults.

Potential journal articles were uploaded into Covidence, a non-profit website working with the Cochrane database to improve systematic reviews. (www.covidence.org, Melbourne, Australia) Duplicates were removed. After initial screening of the title and abstract, two authors (BCZ and RJE) independently reviewed potential studies. Conflicts were resolved by reviewing the full text article and discussing inclusion/exclusion criteria. We excluded review articles, studies that did not include an intervention, that did not report primary data for HIV testing, or that were targeting children or adults outside our specified age range of 10–24 years. We then extracted data from the full text articles included in this review. We used the PRISMA guidelines in assessing the strength of evidence and bias for clinical trials and evaluated random sequence generation, allocation concealment, blinding, and outcome reporting.<sup>27</sup> Randomized controlled trials (RCTs) were considered less biased than pre–post evaluations, and prospective and retrospective observational studies. Interventions that were evaluated and published in peer reviewed journals were considered less biased than abstracts from conference proceedings that could only be judged by study design. Observational studies were evaluated for bias using the GRADE guidelines and included an assessment of eligibility, controls, loss to follow-up, and outcome consistency.<sup>28</sup>

#### Results

## Description of studies identified

Search results included 4,700 potential articles as indicated in the PRISMA flow-diagram in Figure 1. We excluded 85 duplicates,

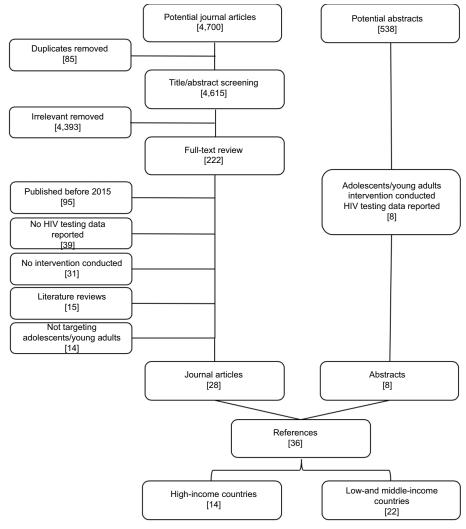


Figure I PRISMA figure of inclusion/exclusion criteria.

as well as 4,393 studies that were not relevant to adolescent HIV testing interventions. We reviewed 222 studies for eligibility based on the above criteria and excluded 194 of them: 95 were published between 2010 and 2014; 39 did not report primary data for HIV testing; 31 did not include an intervention involving adolescent HIV testing; 15 were literature reviews; and 14 were not targeting adolescents. We included 28 articles in our final review. In addition, among the 546 IAS and 292 CROI abstracts, 7 and 1 met our inclusion criteria, respectively. None of these were subsequently published in the literature.

We identified a total of 36 studies for this analysis. We separated the studies into those conducted in high-income countries (a total of 14 studies) and those conducted in low- and middle-income countries (a total of 22 studies) and arranged them by the type of intervention as indicated in Table 2. All of the studies from high-income countries took place in the US  $(N=14)^{16,29-41}$  and will be referred to as US from here forward. The low- and middle-income countries included South Africa (N=6),<sup>42-47</sup> Kenya (N=4),<sup>48-51</sup> Bangladesh (N=2),<sup>52,53</sup> Zambia (N=2),<sup>45,54</sup> Liberia (N=1),<sup>55</sup> Ethiopia (N=1),<sup>56</sup> Malawi (N=1),<sup>57</sup> Mozambique (N=1),58 Myanmar (N=1),59 Ghana (N=1),60 Indonesia (N=1),<sup>61</sup> Zimbabwe (N=1),<sup>62</sup> Uganda (N=1),<sup>48</sup> and Haiti (N=1)<sup>63</sup> with two studies taking place in more than one country.45,48 The interventions were organized into six categories as defined in Table 3. The interventions to increase HIV testing among adolescents and young adults consisted of behavioral/educational interventions (N=4)<sup>29,30,55,56</sup>, alternate venue/self-testing (N=11),<sup>31-34,42-45,50,57,58</sup> vouth-friendly services (N=2),<sup>59,60</sup> technology/mobile health (N=9),<sup>16,35-39,49,51,61</sup> incentives (N=3),53,54,62 and peer/community-based interventions (N=7).<sup>40,41,46-48,52,63</sup> The median sample size was 613 individuals (inter-quartile range=261-2,169). The types of studies included RCTs (N=13), 29,31,36,38,39,42,45,47,49,51,55,60,62 observational studies (N=15),16,32-35,41,44,58,61 and quasi-experimental/pre-post evaluations (N=8).30,37,40,46,52,54,56,59

Among the studies in the US, three studies contained information on new HIV diagnoses (ranging from 0.6% to 11.3% with a median of 3.2%).<sup>32,34,41</sup> All contained information on linkage to care (ranging from 85% to 100%). In low- and middle-income countries, nine studies included information on new HIV infections (ranging from 0.6% to 9.4% with a median of 3.4%).<sup>3,43-45,48,50,53,56,57</sup> Of these, three included information on the number linked to care (ranging from 50% to 100% with an absolute of 97% [94/97]).<sup>53,57,63</sup>

# **Bias assessment**

Of the 36 studies included in our review, 13 were RCTs and 23 were observational studies.<sup>16,30,32–35,37,40,41,44,46,52,54,56,58,59,61</sup>

Three of the RCTs<sup>39,42,62</sup> and five of the observational studies<sup>43,44,52,54,61</sup> were presented in abstracts; these studies were excluded from the risk of bias assessment due to insufficient information. Five of the remaining observational studies were pre–post evaluations of an intervention.

Of the remaining ten RCTs, both random sequence generation and allocation concealment were discernible for four studies from their study methods.<sup>36,51,55,60</sup> Two studies reported random sequence generation only.<sup>31,38</sup> Four RCTs did not report either random sequence generation or allocation concealment.<sup>29,45,47,49</sup> Given the nature of the interventions, blinding of participants and personnel was rarely possible and was reported in only one RCT.<sup>38</sup> In that study, only participants were blinded to arm allocation and blinding of outcome assessors was not reported. All RCTs report HIV testing outcomes as predefined primary or secondary outcomes.

All of the remaining 17 observational studies reported eligibility criteria that were applied consistently for all participants. Only one observational study included a control group.<sup>59</sup> Four observational studies reported loss to follow-up data.<sup>16,30,40,56</sup> Loss to follow-up rates ranged from 16% to 43%. Implementation challenges were noted for higher loss to follow-up rates.<sup>40,56</sup>

## Interventions

Below we report summaries of the individual interventions designed to increase HIV testing among adolescents separated by intervention category (behavioral/educational, alternative venue/self-testing, technology/mobile health, incentives, youth-friendly services, or peer/community) and country category (high-income versus low- and middle-income countries). Within each intervention category, interventions are listed beginning with the least biased (ie, RCTs followed by pre–post evaluations and observational studies). Interventions published in only abstract form are reported last.

# Behavioral/educational interventions (N=4)

There were two interventions from the US that provided a combination of educational material and behavioral interventions targeting adolescents interacting with the criminal justice system<sup>29,30</sup> with one RCT.<sup>29</sup> Letourneau et al randomized 105 adolescents attending juvenile drug court to receive standard care compared to risk reduction therapy.<sup>29</sup> The intervention involved adolescent-parent dyads in 24 weekly, 60–90 minute sessions involving cognitive behavior therapy and behavior management training with contingency-contracting with a point earning system. At the end of the study, there

Interventions to i	Interventions to increase adolescent HIV testing in high-income countries	HIV testing in hig	h-income count	riac	Interventions to increase adolescent HIV testing in high-income countries			
Authors	Publication	l ocation	Number	Ages	Study type	Intervention	Besults of intervention on	l inkage
	and date		of	Ages (target	auny type	description	testing	to care
			subjects	population)				
Behavioral/educa	<b>Behavioral/educational interventions</b>	2						
Letourneau,	Journal of	South	105	11–17	Randomized,	Risk Reduction	HIV testing prevalence	None
et al <sup>29</sup>	Substance	Carolina,		(juvenile	controlled	Therapy for	increased from 16% to 25%	
	Abuse	USA		drug court		Adolescents (RRTA,	for RRTA and decreased from	
	Treatment,			attendees)		n=45) versus UC	17% to 14% for UC baseline	
	2017					(n=60); RRTA: Family	versus I2-month follow-up	
						focused contingency	(not significant); non-significant	
						management with	between-group difference	
						adolescent-parent	RRTA versus UC (OR 2.15,	
						dyads (24 weekly	95% CI 0.49–9.36)	
						sessions, 60–90	_	
						minutes each)	_	
Donenberg,	Journal	USA	54	13-17	Pre-post	Preventing HIV/	HIV testing changes varied	None
et al <sup>30</sup>	of Child			(youth on	evaluation	AIDS among Teens	by sex (OR 2.99, P=0.11);	
	and Family			probation)		(PHAT Life), an	19% (n=6) of males reported	
	Studies, 2015					HIV-prevention	being tested for HIV in the	
						program for teens	previous 6 months at baseline,	
						on probation using	compared to 41% (n=13)	
						group role-plays,	who reported an HIV test	
						videos, games, and	in the previous 3 months at	
						skill development	follow-up; 36% (n=8) of females	
						activities	reported being tested for HIV	
							at both time points; HIV testing	
							increased for males (OR 2.99,	
							95% CI 1.42–6.31, P=0.004) but	
							not for females (OR 1.00, 95%	
							CI 0.33–3.06, P=1.00)	
Alternative venue/self-testing	e/self-testing							
Merchant,	AIDS and	USA	425	18–24	Randomized	Randomized 1:1:1	54% completed assigned test	None
et al <sup>31</sup>	Behavior,			(MSM)		for oral rapid test,	overall (62% oral test, 40%	
	2018					mail-in blood test,	mail-in blood,	
						or medical facility of	56% facility testing); oral	
						choice	and facility tests had greater	
							completion rates than mail-in	
							(P<0.01 each)	

Table 2 Interventions to increase HIV testing among adolescents, published between lanuary 1, 2015, and April 28, 2018

(Continued)

Interventions to	Interventions to increase adolescent HIV testing in high-income countries	t HIV testing in hig	ch-income count	ries				
Authors	Publication	Location	Number	Ages	Study type	Intervention	Results of intervention on	Linkage
	and date		of subjects	(target population)		description	testing	to care
Holliday et al <sup>32</sup>	Journal of Health Care for the Poor and Underserved, 2017	USA	2,385	18–24 (African Americans attending historically black colleges and universities)	Observational, prospective	Campus and Community HIV and Addiction Prevention (CCHAP): HIV (CCHAP): HIV (CCHAP): HIV (CCHAP): HIV testing plus 1-hour interactive peer-led HIV and substance- use workshop, and environmental strategies	2,383 tested (99.9%); I5 HIV infections (0.6%)	15/15 (100%)
Miller et al <sup>34</sup>	JAMA Pediatrics, 2017	ASU	3,301	13–24 (sexual minority males of color)	Observational, prospective	Multisite HIV testing program designed to encourage localized HIV testing programs focused on self-identified on self-identified sexual minority males evaluating universal testing, targeted testing, and combination testing	Universal testing: 35 sexual minority males, 1 (0.1%) new HIV infection; targeted testing: 236 sexual minority males, 16 (3.2%) new HIV infections; combination testing: 693 sexual minority males, 39 (2.1%) new HIV infections	None
Buzi et al <sup>33</sup>	Public Health Reports, 2016	Texas, USA	34,299	l 3–23 (family planning clinic attendees)	Observational, retrospective	Implementation of routine opt-out testing versus opt-in testing	50% increase in HIV testing in opt-out testing; during opt-out, 0.3% were HIV infected	None
								(Continued)

Interventions to	Interventions to increase adolescent HIV testing in high-income countries	It HIV testing in I	high-income cou	ntries				
Authors	Publication and date	Location	Number of	Ages (target	Study type	Intervention description	Results of intervention on testing	Linkage to care
			subjects	population)				
Technology/mobile health	ile health							
Bauermeister	AIDS and	NSA	130	15-24	Randomized,	Full Get Connected!	32 (25%) received HIV testing,	None
et al <sup>36</sup>	Behavior,			(MSM)	controlled	program versus	32% in full intervention,	
	2015					testing site	and 29% in locator only (no	
						locator control;	significant difference)	
						Get Connected!		
						a tailored online		
						HIV/STI testing		
						intervention designed		
						with input from a		
						youth advisory board		
Ybarra et al <sup>38</sup>	Pediatrics,	NSA	302	14–18	Randomized,	Randomized	Intervention more likely to	None
	2017			(MSM)	controlled	I:I Guy2Guy	receive HIV testing compared	
						intervention versus	to control: 55% versus 28%	
						self-esteem control;	(OR 3.42, 95% CI 1.65–7.09,	
						Guy2Guy: 5–10	P=0.001)	
						daily text messages		
						for 5 weeks,		
						content included		
						HIV information,		
						motivation,		
						behavioral skills,		
						HIV testing, healthy		
						relationships		

Interventions to	Interventions to increase adolescent HIV testing in high-income countries	t HIV testing in hi	gh-income coun	Itries				
Authors	Publication	Location	Number	Ages	Study type	Intervention	Results of intervention on	Linkage
	and date		of subjects	(target population)		description	testing	to care
Washington et al <sup>39</sup>	IAS, 2016	California, USA	142	l 8–30 (black MSM)	Randomized, controlled	Intervention versus control; intervention group watched five 60-second videos per week featuring vignettes from BMSM characters, and posted reflections using chat feature; control group viewed standard HIV text information	Retention rates of 71% for intervention group and 78% for control group; BMSM in intervention group was 7 times more likely to have tested for HIV (OR =7.00, 95% CI 1.72–28.33, P=0.006)	en Z
Dowshen et al <sup>37</sup>	AIDS and Behavior, 2015	Pennsylvania, USA	At least 1,500 interacted with campaign	13–17 (primary target); 18–24 (secondary target)	Pre-post evaluation	lknowUshould2 campaign to improve STIs/HIV knowledge and traditional media (print ads, t-shirts, radio, hotline) and new media (website, Facebook, Twitter, Instagram, YouTube)	Significant increase in proportion of CHOP Family Planning clinic visits at which HIV test was conducted (5.4% versus 19.0%, P<0.01)	е О Z
Aronson et al <sup>35</sup>	Journal of Mobile Technology in Medicine, 2016	New York, USA	00	18–24 (youth in ED who declined HIV testing)	Observational, prospective	Tablet-based sexual risk and substance abuse questionnaire and video on HIV testing; high- risk participants enrolled in weekly text messages for 12 weeks	30 (30%) individuals agreed to HIV testing after using tablet; 21 (70%) agreed to receive text messaging	None

Interventions to	Interventions to increase adolescent HIV testing in high-income countries	t HIV testing in his	zh-income count	riec				
Authors	Publication and date	Location	Number of	Ages (target	Study type	Intervention description	Results of intervention on	Linkage to care
			o. subjects	population)			9	
Solorio et al <sup>16</sup>	AIDS and Behavior, 2016	Washington, USA	50	18–30 (Latino MSM)	Observational, prospective	Tu Amigo Pepe pilot intervention: 16-week campaign included Spanish- language radio public service announcements, a website, social media outreach, a reminder system using mobile technology, print materials and a toll- free hotline	56% tested by the end of the campaign; B2% at baseline; 90% after campaign (OR 2.0; 95% Cl 0.8–5.4; P=0.16)	and Z
Community/peer engagement	engagement							
Shelley et al <sup>40</sup>	AIDS	USA	298	18–29	Pre-post	Mpowerment	Increase in HIV testing from	None
	Education			(MSM)	evaluation	(MP): community-	baseline (53.6%) to 3-month	
	puc							
							10110W-up (03.0%, FN =1.20,	
	Prevention,					mobilization	P<0.01) and 6-month follow-up:	
	2017					intervention to	(70.2%, PR =1.28, P<.001)	
						reduce sexual		
						risk behavior		
						by addressing		
						psychosocial		
						factors at individual,		
						interpersonal,		
						social, and		
						structural levels		
						(I) core group, (2)		
						formal outreach,		
						(3) M-groups, (4)		
						informal outreach,		
						(5) publicity, and (6)		
						the project space		
								(Continued)

Arthors         Indication         Lection         Murther         Ages         Study type         Intervention         Results of intervention on description           Concision- Controlio- sult         ADS, 2017         Gaorgia         435         18-34         Observations         Identified 49 (11.3%) HV           Concision- sult         ADS, 2017         Gaorgia         435         18-34         Observations         Identified 49 (11.3%) HV           Concision- sult         ADS, 2017         Gaorgia         435         18-34         Opservations         Identified 49 (11.3%) HV           Concision- sult         ADS, 2017         Gaorgia         Human         Addiatescin: Addiatescin: Apped         Identified 49 (11.3%) HV           alt         ADS, 2017         Gaorgia         Human         Addiatescin: Apped         Identified 40 (11.3%) HV           alt         Addiatescin:	AuthorsPublicationLocatioAuthorsand dateLocatioCamacho-AIDS, 2017Georgia,Camacho-AIDS, 2017Georgia,Gonzalez etAIDS, 2017Georgia,Gonzalez etAIDS, 2017Locatioal <sup>41</sup> Interventions to increase adolescent HIV tesLocatioAuthorsPublicationLocatioAuthorsPublicationLocatioInterventions to increase adolescent Altorsand dateBehavioral/educational interventionsLiberiaIssScience andPractice,201620162016	ation rgia,	Number of	Ages	Study type	Intervention	Results of intervention on	l inkada
and date         of subjects         (target subjects)         (target subjects)         description           at DS. 2017         Georgia, USA         435         BP-24         Observational, Prospective         Metropolitan Atlana Community Addiserent Rapid (MACART)         Metropolitan Addiana Community Addiserent Rapid (MACART)           in the section of the section of the the section of the section of	and date     and date       Camacho-     AIDS, 2017     Georgia, USA       Gonzalez et     USA       Gonzalez et     Locatio       al <sup>41</sup> Interventions to increase adolescent       Authors     Publication       Authors     Interventions       Behavioral/educational interventions       Firestone et     Global       Interventions     Liberia       15     Science and       Practice,     2016			)				LIINAGO
AIDS, 2017     Georgia, USA, USA     413     IB-24     Observational, Retropolitan Attanta Community Addressent Rapid Testing Infative (MCCRTI) intervention: intervention: intervention: intervention intervention intervention     Herropolitan Attanta Community (MCCRTI) intervention: intervention: intervention: intervention intervention       OID     ID-3     IB-24     Observational, Attanta Community (MCCRTI)     Merropolitan Attanta Community (MCCRTI)       ID-3     ID-3     ID-3     ID-4     ID-4       ID-4     ID-4	Camacho- Gonzalez et     AIDS, 2017     Georgia, USA       al <sup>41</sup> USA       al <sup>41</sup> Locatio       al <sup>41</sup> Interventions to increase adolescent       Authors     Publication       Authors     and date       Behavioral/educational interventions       If restone et     Global       If restone et     Global       If restone et     Science and       Practice,     2016			(target population)		description	testing	to care
toors to increase adolescent HIV testing in low- and middle-income countries       Publication     Location     Number     Ages     Study type     Intervention       and date     Location     Number     Ages     Study type     Intervention       alleducational intervention     subjects     Population)     description       alleducational intervention     subjects     Randomized, intervention     HealthyActions       tet     Global     Liberia     1,052     15-35     Randomized, intervention       tet     Global     Liberia     1,052     15-35     Randomized, intervention       science and     Science and     school     versus control;       Practice,     2016     pournal of the     Ethiopia       pound     pound     15-18     Pre-post     Three-month       Intervational     AIDS Society,     evaluation     conselor-delivered       2016     15-18     Pre-post     Three-month     conselor-delivered       2016     Ethiopia     730     15-18     post conselor-delivered       2016     Society,     AIDS Society,     evaluation     conselor-delivered       2016     Society,     Pre-post     Three-month     conselor-delivered       2016     Society,     Pre-post </td <td>Interventions to increase adolescent HIV tes       Authors     Publication       Authors     Publication       Authors     and date       Behavioral/educational interventions     Locatio       Firestone et     Global       Iss     Science and       Practice,     2016</td> <td></td> <td>435</td> <td>-24</td> <td>Observational, prospective</td> <td>Metropolitan Atlanta Community Adolescent Rapid Testing Initiative (MACARTI) intervention: combined nontraditional venue HIV testing, motivational interviewing, and case management</td> <td>Identified 49 (11.3%) HIV infections</td> <td>Higher for MACARTI compared to SOC (96% versus 57%, P&lt;0.001)</td>	Interventions to increase adolescent HIV tes       Authors     Publication       Authors     Publication       Authors     and date       Behavioral/educational interventions     Locatio       Firestone et     Global       Iss     Science and       Practice,     2016		435	-24	Observational, prospective	Metropolitan Atlanta Community Adolescent Rapid Testing Initiative (MACARTI) intervention: combined nontraditional venue HIV testing, motivational interviewing, and case management	Identified 49 (11.3%) HIV infections	Higher for MACARTI compared to SOC (96% versus 57%, P<0.001)
Publication         Location         Number         Ages         Study type         Intervention           and date         of         (Target         Population)         description         description           alleducational intervention         subjects         Population)         Health:         description           and date         Global         Liberia         1,052         I5–35         Randomized, intervention           etr         Global         Liberia         1,052         I5–35         Randomized, intervention           etr         Global         Liberia         1,052         I5–35         Randomized, intervention           etr         Global         Liberia         1,052         I5–35         Randomized, intervention           Practice,         Partice,         Science and         Youth)         Health/Actions:           Practice,         Partice,         Intervention         Science and         Nouth)           Journal of the         Ethiopia         730         I5–18         Pre-post           Journal of the         Ethiopia         730         I5–18         Pre-post           AIDS Society,         AIDS Society,         Ald Crastife evention         Science-costerfed, evented evention           20	thors Publication and date navioral/educational interventions stone et Global Health: Science and Practice, 2016	testing in low	- and middle-in	come countries				
and date     of     (Target     description       alleducational interventions     subjects     Population)     description       alleducational interventions     subjects     Population)     HealthyActions       alleducational intervention     Undenticed, intervention     HealthyActions       science and     Liberia     1,052     15-35     Randomized, intervention       Practice,     Cout-of-     controlled     intervention       Practice,     Nouth)     school     versus control:       Practice,     2016     versus control:     methyActions:       Dormal of the     Ethiopia     730     15-18     Pre-post       International     AIDS Society,     Pre-post     Three-month       International     2016     evaluation     conselor-delivered       2016     2016     isory and creative     intervention	and date aavioral/educational interventions stone et Global Health: Science and Practice, 2016		mber	Ages	Study type	Intervention	Results of intervention on	Linkage
alfeducational intervention alfeducational intervention alfeducational intervention alfeducational intervention begin be	navioral/educational interventions stone et Global Health: Science and Practice, 2016		viacte	(Target Population)		description	testing	to care
et Global Liberia 1,052 15–35 Randomized, HealthyActions Health: Science and Science and Science and Science and Science and Science and Science, Practice, School Practice, Sch	stone et Global Health: Science and Practice, 2016			(				
Health:       Health:         Science and       School         School       Practice,         Practice,       school         Practice,       school <td></td> <td></td> <td>1,052</td> <td>15–35</td> <td>Randomized,</td> <td>HealthyActions</td> <td>Control less likely to receive</td> <td>None</td>			1,052	15–35	Randomized,	HealthyActions	Control less likely to receive	None
Science and Practice,     school     school     versus control;       Practice,     youth)     HealthyActions:     HealthyActions:       2016     Coldentiation     Science and group learning on sexual and reproductive health     HealthyActions:       Journal of the     Ethiopia     730     15–18     Pre-post     Three-month       International     AIDS Society,     2016     evaluation     counselor-delivered       2016     2016     intervention     involving individual, group, and creative arts therapy	Science and Practice, 2016			(out-of-	controlled	intervention	HIV test (OR 0.45, 95% CI	
Practice,       Practice,       HealthyActions:         2016       2016       HealthyActions:         2016       Constant       6-day intensive         2016       Durnal of the       Ethiopia       730         15–18       Pre-post       Three-month         International       15–18       Pre-post       Three-month         AIDS Society,       2016       evaluation       counselor-delivered         2016       2016       psychosocial       involving individual,         2016       Ethiopia       730       15–18       psychosocial         2016       Ethiopia       730       15–18       psychosocial         AIDS Society,       Pre-post       Three-month       counselor-delivered         AIDS society,       Pre-post       Three-month       psychosocial         AIDS society,       Pre-post       Counselor-delivered       psychosocial         Pre-post       Pre-post       Pre-post       psychosocial       psychosocial	Practice, 2016			school		versus control;	0.38-0.53, P<0.001)	
2016     6-day intensive       2016     6-day intensive       group learning     most and       porrnal of the     Ethiopia     730       Journal     15-18     Pre-post       International     730     15-18       AIDS Society,     evaluation     client-centered,       2016     2016     psychosocial       intervention     intervention       2016     evaluation       group, and creative       arts therapy       counseling sessions	2016			youth)		HealthyActions:		
Journal of the     Ethiopia     730     15–18     Pre-post     Three-month       International     730     15–18     Pre-post     Three-month       AIDS Society,     evaluation     client-centered,     psychosocial       2016     2016     intervention     intervention       2016     evaluation     counselor-delivered     psychosocial       Three-month     evaluation     counselor-delivered       2016     evaluation     counselor-delivered       2016     evaluation     counselor-delivered       2016     evaluation     counselor-delivered       2016     evaluation     intervention						6-day intensive		
Journal of the     Ethiopia     730     15–18     Pre-post     Three-month       International     AIDS Society,     evaluation     client-centered,       2016     2016     intervention     intervention       2016     evaluation     intervention						group learning		
Journal of the     Ethiopia     730     15–18     Pre-post     Three-month       International     International     evaluation     client-centered,       AIDS Society,     E     evaluation     client-centered,       2016     2016     involving individual,     group, and creative       arts therapy     counseling sessions						on sexual and reproductive health		
evaluation client-centered, counselor-delivered psychosocial intervention involving individual, group, and creative arts therapy counseling sessions	Journal of the			15–18	Pre-post	Three-month	HIV testing increased by 80%	None
counselor-delivered psychosocial intervention involving individual, group, and creative arts therapy counseling sessions	International				evaluation	client-centered,	for females (AOR 1.8, 95%	
psychosocial intervention involving individual, group, and creative arts therapy counseling sessions	AIDS Society,					counselor-delivered	CI 1.13–2.97) and by 630%	
ividual, reative essions	2016					psychosocial	for males (AOR 7.3, 95% CI	
involving individual, group, and creative arts therapy counseling sessions						intervention	2.6–20.7)	
group, and creative arts therapy counseling sessions counseling sessions						involving individual,		
arts therapy     counseling sessions						group, and creative		
Counseling sessions						arts therapy		
-						counseling sessions		

Interventions to	Interventions to increase adolescent HIV testing in low- and	nt HIV testing in Ic		middle-income countries				
Authors	Publication	Location	Number	Ages	Study type	Intervention	Results of intervention on	Linkage
	and date		of subjects	(target population)		description	testing	to care
Alternate venue/self-testing	e/self-testing		-					-
Shanaube et	AIDS, 2017	Zambia and	15,456	15–19	Community-	PopART for	72.3% accepted intervention;	None
al <sup>45</sup>		South Africa			randomized,	Youth (P-ART-Y):	HCT uptake was 80.6%; 135	
					controlled	door-to-door	(1.6%) new HIV infections	
						combination	diagnosed; control arm data	
						prevention	was not yet available	
						delivered by trained		
						community health		
						workers		
Pettifor et	CROI, 2018	South Africa	284	18–24	Randomized,	Randomized into	96% randomized to choice arm	None
al <sup>42</sup>				(females	controlled	standard of care	chose self-testing; 97% who	
				only)		arm with invitation	returned for 3-month follow-	
						to local clinic for	up from choice arm reported	
						free HCT (n=144)	testing compared to 48% from	
						or choice arm	HCT arm (RR 2.00, 95% CI	
						offering free HCT	1.66–2.40); more peer referrals	
						or self-testing	from choice arm (66% of total	
						(n=140); five self-	peer referrals)	
						testing kits or five		
						testing invitations		
						given to each		
						woman, one for		
						herself and four for		
Hector et	PloS One	Mozambione	496	06-91	Ohservational	Assisted Oral self-	299 (60%) oral self-rests: 1 7%	None
al <sup>sa</sup>	2018		2	2	prospective	testing	HIV-infected	
Ahmed et	Tropical	Malawi	165	1–24	Observational,	Home- or facility-	Home-based: 2/156 new HIV	Home-
al <sup>57</sup>	Medicine and			(only	prospective	based HIV testing	diagnoses (1.3%); facility-based:	based: 1/2
	International			>I 5–24 data		and counseling	5/9 new HIV diagnoses (55.6%)	(20%);
	Health, 2017			reported in		offered for untested		Facility-
				this row)		children of known		based: 3/5
						adults living with		(%09)

Interventions to	Interventions to increase adolescent HIV testing in low- and	t HIV testing in low		middle-income countries				
Authors	Publication	Location	Number	Ages	Study type	Intervention	Results of intervention on	Linkage
	and date		of subjects	(target population)		description	testing	to care
Mugo et al <sup>so</sup>	Sexually Transmitted Infections, 2015	Kenya	1,490	18-29	Observational, prospective	Pharmacy workers referred clients purchasing medicine for fever, sexually transmitted infection symptoms, diarrhea, or body pains	353 (24%) were tested for HIV; 14 (4.0%) were newly diagnosed with HIV	Pone Z
Fatti et al <sup>43</sup>	1AS, 2016	South Africa	6,800	10-19	Observational, prospective	Three testing strategies employed as part of combination program in two districts: index client trailing, door- to-door testing, and campaign testing at events; routine HIV testing program data used	<ul> <li>4.756 (99.1%) consented to HIV testing; first HIV test for 90% of males and 85.7% of females (P&lt;0.0001); 7.5% of females tested HIV+ compared to 3.9% of males (P&lt;0.0001); HIV positivity was higher at campaigns (9.4%) than through index client trailing (6.0%) or door-to-door testing (5.9%) (P=0.019)</li> </ul>	e Z
Rousseau- Jermwa et al <sup>44</sup>	IAS, 2016	South Africa	1,285	12-24	Observational, prospective	Nurse-led, counselor- supported Tutu Teen Truck (TTT) mobile clinic offering HTC offering HTC services at targeted locations (taxi ranks, shopping/ community centers, sports fields, opposite schools)	45.6% had no prior HIV test before TTT; 2.7% were newly diagnosed	Zone
			-					(Continued)

ntervenuons to	increase adolescer	Interventions to increase adolescent HIV testing in low- and		middle-Income countries				
Authors	Publication and date	Location	Number of	Ages (target	Study type	Intervention description	Results of intervention on testing	Linkage to care
			subjects	population)			0	
Technology/mobile health	ile health							
Njuguna et	Sexually	Kenya	600	18–24	Community-	Randomized by	HIV testing within 6 months:	None
49	Transmitted			(females	randomized,	college to SMS	67% from the intervention arm	
	Diseases,			only)	controlled	intervention	and 51% from the control arm	
	2016					or control;	(HR 1.57, 95% CI 1.28–1.92)	
						intervention: weekly		
						SMS on HIV and		
						reproductive health		
Mugo et al <sup>s1</sup>	PloS One,	Kenya	410	18–29	Randomized,	Randomized 1:1	Repeat test attendance was	None
)	2016			evaluated for	controlled	to enhanced	41% (85/207) for standard	
				acute HIV		versus standard	group and 59% (117/199) for	
				infection		appointment	the enhanced group (RR 1.4,	
						reminders to return	95% CI 1.2–1.7)	
						for repeat HIV		
						testing; standard		
						reminders: a dated		
						appointment		
						card; enhanced		
						reminders: a dated		
						appointment card		
						plus SMS and phone		
						call reminders, or		
						in-person reminders		
						for participants without a phone		
Nevendorff,	IAS, 2016	Indonesia	Not	Unspecified	Retrospective,	YKP-friendly	66% increase in HIV testing of	67%
et al <sup>61</sup>			specified	(young key	observational	training for service	YKPs compared to baseline	increase
				population)		providers; YKP	data	in YKP
						sensitization		receiving
						workshop		ART
						for outreach		compared
						workers; online		to baseline
						communication		data
						l platform developed		

Interventions	Interventions to increase adolescent HIV testing in low- and middle-income countries	ant HIV testing in	low- and middle	-income countries				
Authors	Publication and date	Location	Number of subjects	Ages (target population)	Study type	Intervention description	Results of intervention on testing	Linkage to care
Incentives			-				-	
al <sup>62</sup>	IAS, 2016	Zimbabwe	2,796	8-17	Randomized, controlled	Households with eligible children were randomized to one of three arms: 26% standard of care (no monetary incentive; n=735), 41% monetary prize draw (10, 5, or 0 USD with probability of picking as 0.06, 0.07, and 0.90, respectively: n=1,155), and 32% monetary incentive (2 USD; n=906)	Overall 35.7% of those eligible were tested; 15% in SOC arm, 37% in prize draw arm, and 48% in monetary incentive arm attended HIV testing; significantly more adolescents attended HIV testing with monetary incentive	Zone
Chapman et al <sup>st</sup>	IAS, 2015	Zambia	1,813	I I – I 7 (orphans and vulnerable children)	Pre-post evaluation	Survey before and after participation in the STEPS program (Sustainability through Economic Strengthening, Prevention, and Support); 86% response rate at endline (1.813/2.099)	More likely to report having had an HIV test after STEPS (21% pre versus 28% post)	None
Oyewale et al <sup>53</sup>	Current Opinions in HIV and AIDS, 2016	Bangladesh	239	15-24 (MSM and transgender individuals)	Observational, prospective	Effectiveness of voucher scheme to access HIV testing	160 (76%) tested; I HIV infection	1/1 (100%)

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nterventions to	increase adolesce	Interventions to increase adolescent HIV testing in low- and middle-income countries	ow- and middle-	income countries				
Authors	Publication	Location	Number of	Ages	Study type	Intervention	Results of intervention on	Linkage
	and date		or subjects	(target population)		description	testing	to care
Youth-friendly services	rvices							-
Aninanya et	PloS One,	Ghana	2,664	15-17	Community-	26 communities	9.7% increase in testing with	None
al <sup>60</sup>	2015				randomized,	randomized to	intervention (OR 1.16, 95% CI	
					controlled	intervention or	0.85-1.58, P>0.05)	
						comparison;		
						Intervention:		
						school- based		
						curriculum out-of-		
						school outreach.		
						community		
						mobilization.		
						and health-		
						worker training		
						in vourth-friandly		
						houdi-menuly hooks comisers:		
						comparison:		
						only community		
						mobilization and		
						health-worker		
						training		
Aung et al <sup>59</sup>	Journal of	Myanmar	613	15-24	Non-	Link Up	HIV testing increased from	None
	Adolescent			(MSM)	randomized,	intervention	45% to 57% for Link Up and	
	Health, 2017				community	townships	stayed the same for control at	
					controlled	versus control	29%; no significant difference	
						townships; Link	between Link Up and control	
						Up intervention:	(AOR 1.45, 95% CI 0.66–3.17,	
						community- and	P=0.35)	
						clinic-based		
						services that were		
						youth friendly and		
						tailored to meet		
						the specific needs		
						of YMSM including		
						peer education and		
						outreach and youth		
						MSM-friendly clinic		

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Interventions to i	Interventions to increase adolescent HIV testing in low- and	: HIV testing in low		middle-income countries				
Authors	Publication	Location		Ages	Study type	Intervention	Results of intervention on	Linkage
	and date		of	(target		description	testing	to care
			subjects	population)				
Peer/community interventions	interventions							
Rotheram-	AIDS and	South Africa	142	18–25	Community-	Randomized	29% testing in immediate	None
Borus et al <sup>47</sup>	Behavior,			(unemployed	randomized,	neighborhoods	versus 24% in delayed (no	
	2016			males)	controlled	to immediate	significant difference)	
						intervention or		
						delayed control;		
						Intervention:		
						Grassroot Soccer		
						program with		
						trained coaches,		
						random rapid		
						diagnostic tests for		
						alcohol/drugs, and		
						vocational training		
Hershow et	Sport in	South Africa	1,953	12–16	Pre-post	Survey before and	69% tested for HIV	None
al <sup>46</sup>	Society, 2015			(females	evaluation	after participation		
				only)		in SKILLZ Street		
						intervention		
						developed by		
						Grassroot Soccer		
						(GRS); SKILLZ		
						Street: Female		
						coaches deliver		
						an afterschool		
						education program		
						consisting of 10,		
						2 hour biweekly		
						sessions		
								(Continued)

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Authors	Publication	Authors Publication Location Num	Number	ber Ages	Study type	Intervention	<b>Results of intervention on</b>	Linkage
	and date		of subjects	(target population)		description	testing	to care
Hossain et al <sup>s2</sup>	IAS, 2016	Bangladesh	1,005	18–24 (female sex workers)	Pre-post evaluation	Link Up peer outreach intervention at select brothels compared to comparison brothels without intervention	68% in intervention brothels reported contact with peer educator; significantly higher odds of HIV testing uptake (AOR 1.76, 95% CI 1.04–2.96) and receipt of HIV test results (AOR 6.56, 95% CI 1.79–24.12) with peer educator contact; multivariate analyses showed no significant difference between improvement at intervention sites versus comparison sites	Zone
Reif et al <sup>63</sup>	AIDS Patient Care and STDs, 2016	Haiti	3,425	10-24	Observational, prospective	Community-based adolescent HIV testing campaign with community sensitization and active recruitment by CHWs	3,348 (98%) accepted an HIV test; HIV prevalence was 2.65% (n=89)	89/89
al <sup>48</sup>	AIDS, 2016	Uganda and Kenya	116,326	10-24	Observational, prospective	SEARCH hybrid strategy: 2-week community health campaign that included HIV testing, followed by home-based testing of community health campaign nonparticipants	86,421 (89%) adolescents tested for HIV; 1,843 (2.1%) diagnosed positive	Zone

standard of care.

Category	Definition	Examples
Behavioral/	Interventions that provided information on	Cognitive behavior therapy
educational	HIV and/or focused on behavioral change	Behavior management training
Alternative	Interventions that provided HIV counseling	Mobile testing vans
venue/self-testing	and testing outside of traditional health care	In-home testing
	facilities	Oral, home self-testing
Technology/	Interventions using social media, internet, or	Text messaging
mobile health	mobile phones	Online chat groups
Incentives	Interventions that provided monetary or	Cash
	coupon reimbursement for HIV testing	Prize draw
Youth-friendly	Interventions that focused explicitly on	Health-worker training
services	targeting services for an adolescent population	Youth-friendly facilities
Peer/community	Interventions that used interactions with	Youth soccer programs
	trained community members, peers, or groups	Community testing campaigns or events

Table 3 Categories of HIV testing interventions

was no difference in HIV testing in the intervention group (25%) compared to those in standard care (14%) (95% CI 0.49–9.36). This RCT did not report random sequence generation, allocation concealment, or blinding.

In an uncontrolled observational study, Donenberg et al enrolled 54 adolescents aged 13–17 years who had been arrested into the program Preventing HIV/AIDS among Teens (PHAT life), which used group format role plays, videos, and games as an HIV prevention program.<sup>30</sup> There was no change in HIV testing among females, while there was a significant increase in testing from 19% at baseline to 41% (P=0.004) at the end of the intervention for males. Both programs were labor intensive, required multiple visits over time and did not significantly increase HIV testing for the defined populations at the end of the program nor did they report the number of new HIV infections diagnosed.

In low- and middle-income countries, two studies provided educational materials and behavioral interventions<sup>55,56</sup> with one RCT.<sup>55</sup> In Liberia, 1,052 out-of-school adolescents and young adults aged 15–35 years were randomized to receive standard care versus HealthyActions, a 6-day intensive group learning on sexual and reproductive health.<sup>55</sup> At the completion of the study, participants in the control group (42%) were less likely to undergo HIV testing (OR 0.45; 95% CI 0.38–0.53; *P*<0.001) than the intervention group (88%). This RCT included both random sequence generation and allocation concealment but was not blinded.

A separate non-controlled observational study in Ethiopia enrolled 730 adolescents aged 15–18 years in a 3-month client-centered, counselor-delivered psychosocial intervention that involved individual, group, and creative arts therapy counseling sessions.<sup>56</sup> At the end of the study, both females (AOR 1.8; 95% CI 1.13–2.97) and males (AOR 7.3; 95% CI 2.6–20.7) were more likely to have received an HIV test compared to before enrollment. The authors did not report on the number who tested positive or linkage to care. Both of these interventions were labor and training intensive and required multiple visits over time.

# Alternate venues/self-testing (N=11)

In the US, there were four studies involving alternative venue testing strategies for adolescents<sup>31–34</sup> with only one RCT.<sup>31</sup> Merchant et al randomized 425 MSM aged 18–24 to perform an oral rapid self-test, self-mail in blood test, or testing at a local medical facility.<sup>31</sup> Of those enrolled, only 54% completed their assigned test. Oral self-testing (62%) and facility-based testing (56%) were superior (*P*<0.01 each) to mail in blood testing (40%). This RCT reported random sequence generation but did not report allocation concealment or blinding nor did they report the number of HIV diagnoses or linkage to care.

The alternative strategies investigated by the non-RCT studies included testing on campus at historically black colleges and universities,<sup>32</sup> opt-out testing at family planning clinics,<sup>33</sup> and targeted testing tailored to sexual minority men.<sup>34</sup> All of these strategies reported increased HIV testing among adolescents. An observational study evaluating the provision of HIV testing on campus among 2,385 students attending historically black colleges and universities over a 2-year period detected new HIV infections in 0.6% of those tested.<sup>32</sup> The investigators were able to link 100% (N=15) of individuals newly diagnosed with HIV to care. A separate observational study of 3,301 sexual minority men of color aged 13–24 found that targeted testing detected the highest number of new HIV infections (6.3%) compared to universal testing (0.1%) and combination of universal and targeted test-

ing (5.6%).<sup>34</sup> This study was not controlled or blinded, and did not report on loss to follow-up. A retrospective analysis evaluated the historical effect of opt-out testing compared to opt-in testing among 34,299 individuals aged 13–23 years attending family planning clinics.<sup>33</sup> They found a 50% increase in HIV testing during the opt-out period with 0.3% new HIV diagnoses.

In low- and middle-income countries, there were seven studies evaluating alternative HIV testing strategies<sup>42–45,50,57,58</sup> with two RCTs.<sup>42,45</sup> A community RCT in Zambia and South Africa involving 15,456 adolescents aged 15–19 years found that door-to-door testing had high uptake (81%) with 1.6% of those tested newly diagnosed with HIV.<sup>45</sup> This RCT did not include random sequence generation, allocation concealment, or blinding. In South Africa, an RCT involving 284 females aged 18–24 years showed that home-based self-testing had higher uptake (97%) than referrals to local clinic (48%) (RR 2.00, 95% CI 1.66–2.40).<sup>42</sup> This study was reported in abstract form.

There were five non-RCT studies that evaluated alternative venue HIV testing among adolescents.43,44,50,57,58 In an observational study of 496 individuals aged 16-20 years from Mozambique, uptake of oral self-testing was 60% with 1.7% newly diagnosed with HIV.58 A prospective, observational study of 165 adolescents aged 15-24 years in Malawi evaluated untested adolescents with known HIV-infected adult family members by use of household testing and found 9.7% new HIV infections with 77% successfully linking to care.57 A prospective, observational study offering HIV testing to 1,490 symptomatic youth aged 18-29 years presenting to pharmacies in Kenya found low uptake for testing (24%); however, of those tested, 4% were newly diagnosed with HIV.<sup>50</sup> A prospective, observational analysis of 4,800 adolescents in South Africa aged 10-19 years evaluated three testing strategies: index client trailing, door-to-door testing, and campaign testing at events.<sup>43</sup> In this study, 4,756 (99.1%) agreed to HIV testing. Diagnosing new HIV infections was highest with testing campaigns (9.4%), followed by index client trailing (6.0%) and door-to-door testing (5.9%; P=0.019). Another South African observational study that evaluated 1,285 youth aged 12-24 years utilizing mobile HIV testing trucks specifically targeting adolescents found a high uptake of first-time HIV testers (45.6%) and found 2.7% of individuals testing to be newly diagnosed with HIV.44

# Technology/mobile health (N=9)

In the US, there were six technology or mobile health interventions<sup>16,35–39</sup> with three RCTs. <sup>36,38,39</sup> Ybarra et al randomized

302 MSM aged 14-18 years to receive a self-esteem control versus Guy2Guy program that involved daily text messaging for 5 weeks providing HIV information, motivation and behavioral skills, the importance of HIV testing, and healthy relationships. Individuals randomized to the intervention were more likely to undergo HIV testing (OR 3.42; 95% CI 1.65–7.09; P=0.001).<sup>38</sup> This RCT included random sequence generation but did not report allocation concealment or blinding. Washington et al randomized 142 black MSM aged 18-30 years old to watch five 60-second videos per week that included vignettes from black MSM characters and to participate in reflections using a group chat feature compared to a control group that received standard information via text messaging. Individuals in the intervention group were more likely to undergo HIV testing compared to the control group (OR 7.0; 95% CI 1.72-28.33; P=0.0006).<sup>39</sup> This RCT was reported in abstract form. Bauermeister et al randomized 130 MSM aged 15-24 years to use of an online HIV site testing locator or Get Connected!, a tailored online HIV/sexually transmitted infection intervention with a website, where the logo and online materials were designed with input from a youth advisory board.<sup>36</sup> When randomized to the intervention or use of an online testing site locator, there was no statistical difference in HIV testing rates in those receiving the full intervention (32%) compared to those receiving the testing locator only (29%, P>0.05 exact not reported). This RCT reported both random sequence generation and allocation concealment. None of these studies reported on the number of new HIV diagnoses or linkage to care.

Among the non-randomized studies in the US evaluating technology, Dowshen et al reported an observational pre/post evaluation of the IknowUshould2 campaign which used traditional media (print/radio) and technology-based media such as websites, Facebook, Twitter, Instagram, and YouTube to promote HIV testing among 1,500 adolescents who interacted with the campaign.<sup>37</sup> Over the 9 months of the program, there was a significant increase in visits to a family planning clinic for HIV testing (5.4%-19%; P<0.01). This study did not control for temporal differences or other possible confounding factors. Solorio et al reported an observational evaluation of 50 Latino MSM using the Tu Amigo Pepe intervention, a 16-week campaign in the US that included Spanish-language radio announcements, a website, social media outreach, text message reminders, and a toll-free hotline.16 The intervention did not significantly increase HIV testing (90%) from baseline (82%) (OR 2.0; 95% CI 0.8-5.4; P=0.16). Neither study reported the number of new HIV infections or linkage to care. Aronson et al conducted a tablet-based intervention for 100 young adults aged 18–24 presenting an emergency department who had declined HIV testing.<sup>35</sup> After the intervention, 30% of youth tested and 70% agreed to participate in a 12-week program of weekly text messages.

In low- and middle-income countries, there were three interventions using technology or mobile health49,51,61 including two RCTs.<sup>49,51</sup> In Kenya, a randomized text messaging trial involving 600 women aged 18-24 found that women in the intervention group were more likely (67%) to receive HIV testing within 6 months compared to control (51%) (HR 1.57; 95% CI 1.28–1.92).49 This RCT did not include random sequence generation, allocation concealment, or blinding nor did they report the number of new HIV diagnoses. Another RCT in Kenya randomized 410 individuals aged 18-19 years who were evaluated for acute HIV infection to receive appointment reminders via text message compared to enhanced reminders including dated appointment cards, text message reminders, and phone call reminders to increase repeat HIV testing.<sup>51</sup> Repeat HIV testing was 41% in the standard group and 59% in the enhanced group (RR 1.4; 95% CI 1.2–1.7). This RCT reported both random sequence generation and allocation concealment but did not report the number of new HIV diagnoses.

In a retrospective observational study from Indonesia, a combination intervention that included key-populationfriendly services and outreach with an online communication forum found a 66% increase in HIV testing compared to baseline and a 67% increase in those receiving ART compared to baseline.<sup>61</sup>

# Incentives (N=3)

Three studies involved interventions that provided incentives to adolescents for undergoing HIV testing, all of which were conducted in low- and middle-income countries: Bangladesh,<sup>53</sup> Zambia,<sup>54</sup> and Zimbabwe<sup>62</sup> with only one RCT.<sup>62</sup> Dakshina et al randomized 2,796 individuals aged 8–17 years in Zimbabwe into three arms: standard of care (no incentive) versus a prize draw (6% chance of getting \$10, 7% chance of getting \$5, and 90% of getting \$0) versus a guaranteed monetary incentive of \$2.<sup>62</sup> Overall 35.7% of individuals were tested: 15% in the standard of care; 37% in prize draw; 48% in monetary incentive. The investigators identified 11 new HIV infections: 4 (0.3%) in the monetary incentive arm, 7 (0.8%) in the prize-draw arm, and 0 in the standard of care arm. This RCT was reported in abstract form; therefore, a full bias assessment could not be conducted.

An observational pre/post evaluation in Zambia evaluated 1,813 orphans and vulnerable children aged 11–17 years who

participated in the STEPS program (Sustainability through Economic Strengthening, Prevention and Support).<sup>54</sup> Individuals enrolled in the program were more likely to have had an HIV test after the program (28%) compared to prior to the intervention (21%). The authors did not report the number of infections identified or linkage to care. An observational study in Bangladesh evaluated 239 MSM and transgender individuals aged 15–24 years in using a voucher system to access HIV testing.<sup>53</sup> Of the vouchers distributed, 160 (76%) were returned for testing and 1 (0.6%) individual was found to be HIV infected and subsequently linked to care.

# Youth-friendly services (N=2)

There were two interventions involving the use of youthfriendly services to improve HIV testing among adolescents, both in low- and middle-income countries: Myanmar<sup>59</sup> and Ghana<sup>60</sup> with one RCT.<sup>60</sup> A community randomized trial in Ghana evaluated 2,664 adolescents aged 15–17 years who participated in youth-friendly health services, school-based curriculum, outreach, and community mobilization with health-worker training in youth-friendly service compared to control with community mobilization and youth-friendly services only.<sup>60</sup> Compared to the control, adolescents receiving the full curriculum had a 9.7% increase in HIV testing, which was not statistically significant (OR 1.16; 95% CI 0.85–1.58). This RCT reported both random sequence generation and allocation concealment.

A non-randomized, controlled study in Myanmar evaluated 613 MSM aged 15–24 years who participated in the Link Up intervention compared to control townships.<sup>59</sup> The intervention consisted of community and clinic-based youthfriendly services and included peer education, outreach, and a youth-/MSM-friendly clinic. HIV testing increased from 45% to 57% in the intervention group and was unchanged at 29% in the control group (OR 1.45; 95% CI 0.66–3.17). Neither study reported on the number of new HIV diagnoses during the intervention nor did they discuss linkage to care.

# Peer/community interventions (N=7)

In the US, there were two non-randomized, non-controlled studies that evaluated community interventions to increase HIV testing among adolescents. Mpowerment, a community-level mobilization intervention targeting MSM of color aged 18–29 years, found increased testing from 54% at baseline to 70% at 6 months (P<0.001), but it did not report the number of newly diagnosed or linked to care.<sup>40</sup> Metropolitan Atlanta Community Adolescent Rapid Testing Initiative (MACARTI) combined non-traditional venue HIV testing, motivational

interviewing, and case management targeting youth and identified 11.3% of testers with new HIV infections, 96% linked to care compared to 57% under standard of care (P<0.001).<sup>41</sup>

In low- and middle-income countries, five studies assessed community- or peer-focused interventions<sup>46–48,52,63</sup> with one RCT.<sup>47</sup> A community randomized trial in South Africa evaluated the Grassroot Soccer program that included trained coaches, educational/vocational training, and use of rapid HIV diagnostics. Among the 142 males, the program did not increase HIV testing (29%) compared with delayed enrollment into the program (24%).<sup>47</sup> This RCT did not report random sequence generation, allocation concealment, or blinding.

In a separate observational study of the Grassroot Soccer intervention among 1,953 females, 69% of participants tested for HIV; however, there was no comparison or control group.<sup>46</sup> Neither study reported on the number of new HIV diagnoses or linkage to care. In Bangladesh, an observational study of the Link Up peer outreach program targeted HIV testing among 1,005 young adult female sex workers working in brothels, but it did not find a significant difference in HIV testing in brothels that had the intervention compared to those that did not participate in the intervention.<sup>52</sup> In Haiti, an observational evaluation of a community-based adolescent HIV testing campaign tested 3,348 individuals, of whom 98% offered testing.<sup>63</sup> They diagnosed 89 (2.7%) new HIV infections, all of which were linked to the clinic the same day. In Uganda and Kenya, an observational study of a combination approach of community-based testing followed by home-based HIV testing for community members not participating in the campaign tested 86,421 (88%) of adolescents.48 The authors reported that 1,843 (2.1%) individuals were newly diagnosed with HIV; however, they did not report on linkage to care.

## Discussion

In this systematic review of HIV testing interventions among adolescents in high- versus low-/middle-income settings, we found 36 studies including 13 RCTs; yet only six studies discussed linkage to care. The primary purpose of screening for any disease is early diagnosis, so that diagnosed individuals can be promptly treated; this is true for HIV testing. Therefore, it is important that interventions to increase HIV testing not only address barriers to HIV testing but also include methods to effectively link newly diagnosed individuals to care.

Interventions that use technology and mobile health can address psychological barriers such as perceived risk,

stigma, disclosure, and fear of rejection to increase HIV testing in adolescents. In the US, there were two RCTs and one pre-post evaluation that evaluated interventions using mobile health and technology to target key populations that significantly increased HIV testing among adolescents.37-39,64 In low- and middle-income countries, two RCTs and one retrospective observational study found that text messaging interventions targeting high-risk key populations significantly increased HIV testing among adolescents.49,51,61 In most settings, adolescents and young adults found text messaging and mobile health technology interventions acceptable and feasible.35-39,65-68 However, these interventions did not address structural barriers to HIV testing and none of the nine studies reported the numbers of new HIV diagnoses or linkage to care. Linking newly HIV diagnosed individuals is critical to improving the continuum of care among adolescents living with HIV.7,8

Non-traditional HIV testing venues and oral self-testing can overcome structural barriers such as inconvenience, insurance, and parental consent to improve HIV testing among adolescents.<sup>69</sup> In the US, one RCT and three observational studies found high levels of acceptability, but they offered limited data on linkage to care, which is particularly important for alternative venue testing.<sup>31–34,69</sup> Oral self-testing was an acceptable HIV testing method among adolescents and could be a method employed to expand HIV testing programs; however, there are limited data on linkage to care after self-testing among adolescents.<sup>31,58</sup> Within traditional health care venues, opt-out testing appears to improve HIV testing among adolescents compared to opt-in testing and may increase HIV testing over provider-initiated counseling and testing in high-prevalence areas.<sup>33,70,71</sup> In low- and middle-income countries, door-to-door and mobile HIV testing was found to be feasible, acceptable, and led to large numbers of adolescents obtaining HIV testing; however, data on linkage to care is absent in those studies.<sup>43-45</sup> In Malawi, door-to-door contact tracing of children born to HIV-infected mothers was a high yield method for detecting perinatal HIV in children and younger adolescents and could be expanded to other high HIV prevalence countries.<sup>57</sup> Providing flexible, alternative strategies outside of traditional health care settings appears to be an acceptable, feasible, and effective method of increasing HIV testing among adolescents in the US and low- and middle-income countries. In lowand middle-income countries, alternative testing sites such as home self-testing, in-home door-to-door testing, testing campaigns, and pharmacy-based testing appear

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to be acceptable and feasible alternatives for HIV testing among adolescents; however, little is known about linkage to care after new HIV diagnosis in alternative venues. Of the eleven interventions involving alternative venue HIV testing, only two evaluated linkage to care, limiting the generalizability of these approaches.<sup>32,57</sup>

Community testing events and mobilization have the potential to overcome structural and psychosocial barriers by easing access, making testing normative, and providing social support; therefore, they have the ability to test large numbers of individuals and can be used to target high-risk, key populations.<sup>40,41,46,48,52,57,63</sup> The only RCT in this category did not find efficacy in increasing HIV testing.<sup>47</sup> However, six observational studies reported high levels of HIV testing in community interventions. In addition, two interventions reported successful linkage to care after HIV testing, making this an important component for future HIV testing modalities for adolescents.<sup>41,57,63</sup>

Other HIV testing interventions had mixed results and lower quality evidence for increasing HIV testing among adolescents. Interventions that involved behavioral change or education in the US were labor and resource intensive and showed variable results.<sup>29,30</sup> These interventions appeared to be more effective in low- and middle-income countries compared to the US; however, none of these studies discussed the number of newly diagnosed HIV infections nor the number linked to care. Offering incentives for HIV testing among adolescents can address motivation for HIV testing but does not overcome many structural or psychosocial barriers to HIV testing, and sustainability may be challenging.<sup>53,54,62</sup> In addition, there is limited data on the effectiveness of incentives on linkage to care after HIV testing. The specific use of youth-friendly testing facilities can address some structural and psychosocial barriers to HIV testing; however, these interventions did not have statistically superior effects compared to traditional testing sites.59,60

Though critical to informing the implementation and scale-up of effective interventions, none of the studies reported resource utilization or costs related to HIV testing or the intervention. With resource utilization and cost data, health policy models can project the long-term impact of interventions beyond the time-horizon of traditional studies. This type of analysis is particularly important when considering testing interventions among youth, in whom the effects of HIV infection and treatment may not manifest for years or decades.<sup>5</sup> Few studies have reported on the cost-effectiveness of adolescent-specific prevention or testing interventions.<sup>72–76</sup> Such information is invaluable for policymakers to under-

stand optimally deploying combinations of universal and targeted testing in specific settings and warrants more study moving forward.<sup>77</sup>

Current studies are exploring alternative methods and venues to improve HIV testing and linkage to care among high-risk adolescents.<sup>78</sup> Mpower is an ongoing communitylevel intervention in the targeting young MSM using peer educators to engage high-risk youth to increase HIV testing.<sup>79</sup> Other investigators are exploring the use of oral, self-testing with video counseling for transgender youth in the US.<sup>80</sup> In Kenya, a large-scale study is evaluating alternative testing venues (community versus home) and testing modalities such as oral self-testing, home testing, mobile testing, or facility-based testing.<sup>81</sup> Results of these studies may add valuable information for the development of multicomponent interventions to increase HIV testing among adolescents.

Future research should also focus on expanding the geographic reach of interventions for HIV testing among adolescents and young adults. In particular, this review did not identify any recently published interventions to increase HIV testing in Latin America or in high-income countries other than the US. Although the majority of HIV infections among adolescents occur in sub-Saharan Africa, HIV affects adolescents globally. It is important that effective interventions are identified that address culture-specific barriers and target local at-risk populations.<sup>3</sup>

#### Conclusion

To diagnose more HIV infections among adolescents, it is important to target high-risk populations, minimize barriers to HIV testing, and make testing easier and more widely available. One intervention is unlikely to address all of the barriers to HIV testing among adolescents and would be unlikely to succeed across all settings. Therefore, future interventions should utilize multiple components and expand on the successful use of mobile health technology, alternate venue testing, and community mobilization while stressing the importance of linkage to care. High-quality RCTs are needed to identify optimal combinations of interventions that increase HIV testing among adolescents while focusing on diagnosing new HIV infections and providing linkage to care.

# **Author contributions**

All authors contributed to data analysis, drafting or revising the article, gave final approval of the version to be published, and agree to be accountable for all aspects of the work.

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