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# The Mediating Role of Partner Selection in the Association Between Transactional Sex and HIV Incidence Among Young Women

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**Objective:** In sub-Saharan Africa, transactional sex is associated with an increased risk of HIV infection in adolescent girls and young women, but the mechanisms for this relationship remain unclear. We hypothesize that young women who report transactional sex may have multiple partners and older partners, thereby increasing their HIV risk.

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**Setting:** We used longitudinal data from the HPTN 068 trial in rural South Africa where young women aged 13-20 who were HIV-negative at enrolment (n = 2362) were followed approximately annually for up to 6 years.

**Methods:** We used the parametric g-formula to estimate the total effect of time-varying, frequent transactional sex (receipt of gifts/ money at least weekly versus monthly or less) on HIV incidence and the controlled direct effect for mediation in a simulated cohort using 20,000 bootstrapped observations. We calculated rates and hazard ratios (HRs) over the entire study period.

**Results:** The HR for the total effect of frequent transactional sex on HIV incidence was 1.56 (95% confidence interval: 1.28 to 1.85). However, this effect was mediated by partner age (>5+) and number of partners (>1) and the HR was attenuated to 1.09 (95% confidence interval: 0.90 to 1.28) when setting both partner age and partner number constant.

**Conclusion:** Both partner age difference and partner number mediate the relationship between transactional sex and incident HIV infection. Through this mediation analysis, we provide important longitudinal evidence to suggest that young women who engage in frequent transactional sex select multiple partners, often older male partners that may be part of higher risk sexual networks.

**Key Words:** adolescent girls, young women, transactional sex, mediation analysis, causal pathways, HIV incidence, older partners, multiple partners

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### **INTRODUCTION**

In sub-Saharan Africa (SSA) adolescent girls and young women (hereafter young women) aged 15–24 bear a disproportionate burden of the HIV epidemic; over 25% of new HIV infections in this region occur in this population.<sup>1–3</sup> Transactional sex defined as "noncommercial, non-marital sexual relationships motivated by an implicit assumption that sex will be exchanged for material support or other benefits" is considered to be a contributing factor to the high HIV infection rates observed among young women in the region.<sup>4–7</sup> Both cross-sectional<sup>6,8</sup> and longitudinal evidence<sup>7,9</sup> has demonstrated that young women reporting transactional sex are at a higher risk of HIV acquisition. In addition,

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previous analysis of these data by Kilburn et al<sup>7</sup> found that the effect of transactional sex is particularly driven by relationships in which a sexual partner provides money and/or gifts frequently, meaning at least weekly, as opposed to monthly.

Structured by gender inequality, transactional sex takes place across a range of economic contexts; from those characterized by poverty and insecure livelihoods to those marked by income inequality and consumerist aspirations.<sup>10</sup> The discourse around women's motivations for engaging in transactional sex have included: fulfilment of basic needs in impoverished settings; the expectation that men should provide for their partners in relationships; and efforts to improve one's social status.<sup>4,12,13</sup> Not all sexual relationships characterized by or involving transactional sex are risky for HIV infection. Qualitative evidence suggests that transactional sex is an expectation embedded in adolescent romantic relationships; only certain aspects related to male provision or dependence on partners for money or material support, result in young women's weakened negotiating position within the relationship that make it risky for HIV infection.<sup>11</sup>

Transactional sex is also associated with other dimensions of HIV risk in women. These include different forms of partner violence and abuse,<sup>12,13</sup> alcohol consumption or patronizing venues that serve alcohol,<sup>15,16</sup> and nonuse or inadequate use of condoms, although there is no clear association with condom use, possibly because of reporting and measurement bias.<sup>17,18</sup> Furthermore, there is evidence that young women who report transactional sex are more likely to have multiple partners,<sup>12,14</sup> and to have older partners.<sup>13</sup> However, a study by Jewkes et al in South Africa showed an increase in incident HIV among young women who reported transactional sex with an on-going or once-off partner. This finding was independent of partner number or age.<sup>9</sup> Therefore, the question still remains as to why transactional sex is risky for HIV.

In particular, examining the causal pathways between transactional sex and increased HIV risk is important for improving the health and well-being of young women in SSA, and for improving our HIV prevention response. Longitudinal evidence examining the pathway from transactional sex to HIV acquisition is limited. A cross-sectional study from Swaziland suggests a measure of gender inequality—constrained agency of young women—and offers an explanation of the pathway.<sup>21</sup> This was further clarified by a cross-sectional analysis in rural South Africa that found that young women who engage in transactional sex are at risk for HIV due to their choice of partners and the sexual networks of those partners.<sup>6</sup>

Our aim is to investigate whether the association between frequent transactional sex and HIV acquisition is mediated by young women having multiple and older sexual partners given their role in influencing young women's HIV risk. We have conceptualized multiple sexual partners as being on the pathway between frequent transactional sex and HIV, because young women might engage in transactional acts with multiple partners, primarily motivated by the need to obtain items or status.<sup>4</sup> For older partners, young women who report frequent transactional sex are more likely to have age-disparate partnerships, because of the following reasons: young women may pick older partners because of men's ability to provide gifts and money to the young women, for their social and educational maturity, for the belief that they are better sexual partners, and they may be perceived as more marriageable.<sup>15–17</sup> There is usually an established power dynamic between these older men and young women, in turn making it more difficult for young women to negotiate safe sexual behavior, especially in transactional relationships.<sup>18</sup>

To our knowledge, there are no longitudinal studies that formally test these causal pathways.<sup>4</sup> Hence, our aim is to address this knowledge gap using longitudinal data collected from a randomized controlled trial with young rural South African women.

## METHODS

## **Study Population and Sample**

This paper is a secondary analysis of longitudinal data of participants enrolled in a phase 3, individually randomized conditional cash transfer trial in rural South Africa (HPTN 068). The primary objective of the trial was to determine whether providing cash transfers, conditional on school attendance, reduced the risk of HIV acquisition in young women aged 13–20 years. Data collection was conducted in rural Mpumalanga Province, South Africa.<sup>19–21</sup> Further details on the study design, questionnaires and laboratory procedures are available in the baseline and main trial publications.<sup>22,23</sup>

The trial included young women living in 28 villages within the Agincourt Health and Socio- Demographic Surveillance System (AHDSS) area run by the MRC/Wits Rural Public Health and Health Transitions Research Unit.24 At baseline, the trial enrolled 2533 young women in grades 8, 9, 10, or 11 at selected schools within the AHDSS study site. Participants were excluded if they were pregnant or married at baseline. Participants were seen annually from baseline at 12, 24, and 36 months until the study completion date or their planned high-school completion date, whichever came first.<sup>23</sup> One additional visit took place 1-2 years after the study ended (a postintervention visit) for all participants; thus participants could have up to 4 follow-up visits over 6 years.<sup>7</sup> Young women were in different grades at enrolment and could have had fewer than 4 visits if they were expected to graduate before the end of the study period. Each annual study visit included an audio computer-assisted self-interview (ACASI) with the young woman and HIV testing for those who were negative at the previous visit. An additional HIV test was conducted for some girls around the time of expected graduation from high school or when the study was completed to capture more person-time in the study, if eligibility was met (termed the "graduation visit"). This test was typically around 6 months after the previous annual visit.23

To measure HIV incidence and mediation, our analytical sample (n = 2362) included participants who were HIV negative at baseline enrolment and had at least one follow-up visit.<sup>7</sup> We did not exclude sexually inactive young women as a meaningful proportion of the incident HIV infections (20%) occurred in those that did not report any sexual activity and we wanted to extrapolate findings to all young women.<sup>7</sup> Kilburn et al's<sup>7</sup> paper that showed whether there is an association between transactional sex and HIV incidence used the same dataset and provides a sensitivity analysis that shows the association among only those who reported ever having sex (see Table A1, Supplemental Digital Content, http://links.lww.com/QAI/B395).

## Measures

The outcome variable, *HIV incidence* was determined using HIV tests conducted at baseline and at each follow-up visit. HIV testing procedures included using 2 HIV rapid tests performed in parallel followed by a confirmatory test, if one or both rapid results were HIV reactive. Detailed procedures for HIV testing and laboratory procedures are described in the trial paper.<sup>23</sup>

Our exposure variable is *frequent transactional sex* as Kilburn et al<sup>7</sup> showed not only an association between transactional sex and HIV incidence in this cohort, but that the effect was strongest among those who engaged in transactional sex with frequent exchanges. Frequent exchanges were defined as receiving money weekly or gifts "often" or "always," in contrast to infrequent exchanges (having received money once or monthly and gifts "a few times" or "once" or "none"). We constructed a binary exposure variable for this analysis to equal 1 for transactional sex with a partner that gave money or gifts frequently (frequent transactional sex) and 0 if either (1) no transactional sex; or (2) transactional sex with infrequent exchange (infrequent transactional sex). Furthermore, in modelling the exposures and outcomes for the simulations, we included a dummy variable for infrequent transactional sex so that "none" served as reference group.

We defined the mediator of *having an older partner* as having had at least one sexual or nonsexual partner >5 years older at each follow-up visit. Partners with whom there was no reported sexual relationship were included to account for potential misreporting about sexual behaviors. The mediator of the *number of sexual partners* was defined as having zero, 1, or >1 sex partners in the 12 months before each followup visit.

We selected confounders based on previous literature on transactional sex and HIV infection and our directed acyclic graph (shown as Appendix 2, Supplemental Digital Content, http://links.lww.com/QAI/B395). We included the exposure-outcome baseline confounders of age of young woman, intervention arm assignment to account for the original trial design, and quartiles of per-capita household consumption. Time-varying controls include schooling (high school attainment versus enrolled in high school), ever pregnant, physical intimate partner violence (IPV), herpes simplex virus-2. We also included the exposure-outcome and mediator-outcome time-varying confounders for depression from the Center for Epidemiologic Studies depression scale (CES-D), a 20-item scale with a cut-off score  $\geq 16^{25}$  and wealth quartiles, represented as the lag of time-varying log of per capita expenditure in all models. The construction of the specific variables—schooling, IPV, and per-capita household consumption—has been referenced in earlier papers.<sup>7,26</sup>

## **Statistical Analysis**

To explore the mediating effect of frequent transactional sex on HIV incidence, we used an adaptation to the parametric g-formula for mediation analysis that allows us to empirically model both time-varying confounding and mediators within longitudinal, survival data.<sup>27</sup> We examined total effects and controlled direct effects (CDE) of mediators of interest. Mediator effects were examined both separately and jointly in a simulated cohort of 400 estimates using approximately 20,000 bootstrapped samples (inflating the original baseline sample by 8).

Using the counterfactual approach to causal mediation, we define the total effect as the Hazard Ratio (HR) of the effect of frequent transactional sex on incident HIV, if it were possible to observe all participants under each possible exposure plan: Y(1) = frequent transactional sex and Y(0) = no frequent transactional sex. Mediators in this model are left at the natural value they would have taken under each exposure plan.<sup>28</sup> We estimated the CDEs of the 2 mediators (older partners and number of sexual partners) on the relationship between frequent transactional sex and incident HIV. CDEs are defined as the effect of exposure on an outcome while keeping the mediator "controlled" at level M for everyone, but switching exposure from control, Y(0), to treatment, Y(1). Mediation is the attenuation of the total effect closer to the null (ie, the HR of the CDE is closer to 1). We also attempted to estimate the CDE for the mediators, condom use, and low sexual power, but did not include them in the final model because of measurement concerns and issues with missing values (see "discussion").

In general, CDEs require fewer assumptions about no measured confounding of exposure and outcome relationships than the natural direct and indirect effects. In particular, the CDE does not require that mediator-outcome confounders are unaffected by previous exposure, a difficult assumption to demonstrate without randomization of mediators.<sup>29</sup> In our study, CDEs represent the hypothetical scenario if we were able to set mediators to a riskier level in the sample (eg, increasing number of sexual partners). We examined the CDE under several different "scenarios" including (1) setting all young women to have an older partner; (2) setting young women to have one sexual partner (3) setting young women to have more than one sexual partner; and (4) setting all young women to have both an older partner and more than one sexual partner. We checked for interactions between the exposure and mediators and are not including them because of sparse data.

To estimate the total effect and the CDEs using the parametric g-formula, we undertook the following steps (details in Appendix 1, http://links.lww.com/QAI/B395): First, we expanded the dataset to 8 times the sample size (around 20,000 observations) and pulled a random sample with replacement.<sup>29,30</sup> Next, we fitted pooled logistic regression models (ordered logit model for number of sexual partners) for every time-varying outcome, exposure, and

confounder used in the analysis. Third, using Monte Carlo simulation, we used baseline confounders and coefficients obtained from the logit models in the first step to simulate the predicted probabilities of every time-varying outcome across each of the 4 follow-up time points. We repeated this process under each exposure plan, Y(1) and Y(0), to estimate the risk of HIV incidence across both potential outcomes. We then used this predicted HIV incidence to estimate the HR of the total effect and CDEs using Cox Proportional Hazard models. We also report the rate of HIV incidence per person year and the difference between them taken as an average across the simulated sample. We repeated all steps for each hypothetical scenario to estimate the CDEs. We calculated 95% confidence intervals (CIs) of rates, HRs and rate differences using the SD of the point estimate from 400 simulated samples. We used STATA version 15.1 for all analyses.

## RESULTS

Table 1 provides baseline characteristics for the entire sample (n = 2362) of young women. At baseline, the median age was 15 years, 26.2% reported ever having sex and the

**TABLE 1.** Baseline Characteristics of HIV-Negative Young Women Aged 13–20 in Agincourt, South Africa With at Least One Follow-up Visit (N = 2362)\*

	Median (IQR) or n (%)
Young women's age at baseline (yrs)	15 (14–17)
Average monthly per capita household expenditure (South African Rand)	289.2 (184.5–477.6)
Ever had sex	618 (26.2)
Age in years at first sex (sexually active girls)	16 (14–16)
CCT beneficiary arm	1215 (51.4)
Older partner (5+ years)	129 (5.6)
Sex partner number	
0	1715 (73.6)
1	476 (20.4)
>2	138 (5.9)
Prevalent HSV-2 infection	90 (3.8)
Ever pregnant	192 (8.2)
Any alcohol use	51 (2.2)
Double or single orphan	468 (20.0)
Physical IPV	245 (10.6)
Condom use at last sex (sexually active girls)	426 (69.5)
High relationship power (sexually active girls)	227 (37.9)
Children's depression inventory score $\geq 16$ §	415 (18.5)
Any transactional sex past 12 months†	82 (3.6)
Frequent transactional sex past 12 months‡	38 (1.7)

\*Baseline data collection occurred between March 2011 and December 2012.

\*Ever transactional sex is operationalized as whether a young woman reported that she felt that she had to have sex with a male partner who gave her money or gifts. \*Frequent transactional sex is defined as transactional sex with a partner that frequently provided material items (gave money weekly and gifts "often" or "always"). These frequent exchanges are in comparison to infrequent exchanges, defined as having received money once or monthly and gifts "a few times" or "once" infrequently.

§We report on the Children's depression inventory score at baseline, but used the Center for Epidemiologic Studies Depression Scale (cutoff score ≥16) in the models. CCT, conditional cash transfer.

median age of first sex was 16 years. With a low proportion of sexually active participants in the entire sample, sexual behaviors such as past year transactional sex [n = 82 (3.6%)] and frequent transactional sex [n = 38 (1.7%)] were low. Furthermore, of all young women, 5.6% (n = 129) reported having a partner >5 years older, 20.4% (n = 476) had one partner, 73.6% (n = 1715) had zero partners, and 5.9% (n = 138) had more than one partner in the past year.

Table 2 provides mean characteristics of young women by frequency of transactional sex across time. For this table, we split the table into 2 study visit periods: during the main trial (3 years) and postintervention visit (1 year). A higher percentage of young women who reported frequent transactional sex during the main and postintervention trial had an older partner (>5 years older) (29% versus 0.7%), and higher mean numbers of sexual partners compared with those who reported infrequent or no transactional sex (mean: 1.2 versus 0.3). For sexually active young women, the proportion using condoms during last sex was almost the same between those engaging in frequent or infrequent transactional sex across study periods. Furthermore, those that did not report frequent transactional sex had higher sexual relationship power compared with those who reported frequent transactional sex (57.7% versus 36.7%) in the main trial period. In addition, Table 2 shows that a higher proportion of young women that engaged in frequent transactional sex had older partners (>5 years older) and number of partners in the postintervention visit compared with the main trial. We have included sample observations of our key covariates through 2  $\times$  2 tables (cross-tabulations) to demonstrate that the cells of our covariates have sufficient sample numbers. This table, included as Appendix 3, Supplemental Digital Content, http:// links.lww.com/QAI/B395 shows the exposure (frequent transactional sex) by each confounder and mediators (older partner and multiple partners) by each confounder, pooled over all intervals.

Table 3 displays the total effect and CDEs of frequent transactional sex on HIV incidence by different levels of mediators. The total effect in Table 3 indicates that if the mediators had taken on their natural values (represented by the coefficients that the simulation model shows before we set the mediators), the incidence rate of HIV per person year over 6 years of follow-up was  $\sim$ 5% if all young women had frequent transactional sex and  $\sim 3\%$  if all had infrequent or no transactional sex. The HR for the total effect was 1.56 (95% CI: 1.28 to 1.85). Table 3 also shows the CDE for the effect of frequent transactional sex on HIV incidence under different scenarios, such as having an older partner >5 years, having one sex partner and more than one sex partner, each individually. We observed attenuation from the total effect (as HR reaches 1) for CDEs after setting individual mediators to: all young women have an older partner (HR: 1.38; 95% CI: 1.17 to 1.59), sex partner number is set to one partner (HR 1.23; 95% CI: 1.00 to 1.46) and then more than one sex partner (HR: 1.22; 95% CI: 1.00 to 1.45), as also depicted in Figure 1.

Furthermore, when jointly setting the 2 mediators having an older partner and more than one sex partner— CDEs are strongly attenuated in comparison to the total

	During the Main Trial (3 Visits) Frequent Transactional Sex		Post-intervention (1 Visit) Frequent Transactional Sex	
	Yes	No	Yes	No
Age	17.9	16.9	20.3	20.1
Condom use during last sex (sexually active girls)	73.1%	73.3%	74.8%	79.2%
High sexual relationship power (sexually active girls)	36.7%	57.7%	53.0%	56.4%
Ever pregnant	55.0%	15.0%	55.3%	35.9%
Older partner (5+ years older)	29.0%	0.7%	40.6%	21.7%
No. of partners (mean)	1.2	0.3	1.3	0.7
Alcohol use	4.3%	1.7%	6.4%	4.2%
Physical IPV	39.1%	21.9%	17.6%	8.5%

**TABLE 2.** Mean Characteristics of Young Women By Whether She Engaged in Frequent Transactional Sex Across Study Visits

During the main trial there were 449 observations over the 3 follow-up visits defined as having frequent transactional sex and 2314 that did not have frequent transactional sex. At the postintervention visit 188 young women reported frequent transactional sex and 1571 that did not have frequent transactional sex.

effect. In this joint scenario, the HR is the closest to one out of all scenarios (HR 1.09, 95% CI: 0.90 to 1.28) (Table 3). This result is also demonstrated by the cumulative incidence curves in Figure 2B. When we set the mediators, the curves for frequent and no frequent transactional sex (either none or nonfrequent exchanges) with HIV incidence are uniform during years 1–3 especially (during the main trial) and diverge later during the postintervention study period.

#### DISCUSSION

This longitudinal mediation analysis examined the pathways between frequent transactional sex and HIV incidence among a sample of secondary school young women aged 13–20 in rural South Africa. Young women's partner

characteristics, such as having an older partner (>5 years) and the number of sexual partners in the past year mediated the relationship between frequent transactional sex and incident HIV, suggesting that a large proportion of the effect of frequent transactional sex on HIV acquisition is the result of partner selection. To our knowledge, this is the first study that uses formal mediation analysis methods to delineate the causal pathways between transactional sex and HIV incidence. Previous research has shown that a partner age difference of 5–10 years is associated with higher HIV risk<sup>31–33</sup> and that young women who engage in transactional sex, thus increasing their HIV risk.<sup>34</sup> Furthermore, evidence from South Africa shows that young women's negotiating power

TABLE 3. Total and CDE of Frequent Transactional Sex on HIV Incidence By Different Levels of Mediators (Partner Number and Partner Age Difference)

	HIV		
	Incidence Rate (%)	RD (95% CI)	HR (95% CI)
Total effect			
Frequent TS (weekly)	5.0	1.8 (1.2 to 2.2)	1.56 (1.28 to 1.85)
Nonfrequent TS or none	3.3		
CDE: Having an older partner			
Frequent TS (weekly)	7.3	2.0 (1.4 to 2.6)	1.38 (1.17 to 1.59)
Nonfrequent TS or none	5.3		
CDE: Having 1 sex partner			
Frequent TS (weekly)	4.9	0.8 (0.3 to 1.5)	1.23 (1.00 to 1.46)
Nonfrequent TS or none	4.0		
CDE: Having more than 1 sex partner			
Frequent TS (weekly)	6.1	1.0 (0.4 to 1.7)	1.22 (1.00 to 1.45)
Nonfrequent TS or none	5.1		
CDE: Having older partner and more than one sex			
partner			
Frequent TS (weekly)	8.9	0.7 (-0.1  to  1.5)	1.09 (0.90 to 1.28)
Nonfrequent TS or none	8.2		

Incidence Rates, Rate Differences (RD), and HRs calculated from 400 Monte Carlo simulations with 20,000 observations randomly pulled with replacement from all follow-up visits (main trial and postintervention).



**FIGURE 1.** CDEs showing the effect of transactional sex on HIV incidence under different scenarios using older partner and partner number as mediators.

for condom use is often compromised by partner age disparities and economic dependence that increases HIV risk.  $^{33,35}$ 

We also observed that young women who engaged in more frequent transactional sex reported lower sexual relationship power than those who engaged less or not at all. In this study, we saw this particularly among younger women as part of the main trial study visits. Not seen here, but shown in other studies, women in multiple and concurrent relationships report less consistent condom use and are more likely to report transactional sex, and difficulty in both negotiating condoms and not being able to influence timing and nature of sex.<sup>36</sup> In this analysis, young women engaging in frequent transactional sex have a higher risk of HIV. These young women may be more dependent on male partners, thus reducing their decision-making power when practicing safer sex. In addition, when male partners provide money/gifts frequently, the imbalance in power may be more acute, as shown by qualitative research in South Africa.<sup>37</sup> This finding aligns with research by Luke<sup>17</sup> in Kenya that demonstrates that resources obtained from within the relationship decrease young women's negotiating power. Furthermore, Luke's (2005) research on the value of transfers and condom use showed that the larger the value of the gift, the less likely safer sex would be practised.<sup>38</sup> Our results also suggest that young women with multiple partners may be part of a network

of higher-risk male partners that increase their exposure to HIV, as has been shown through an ecological analysis of epidemiological data in 14 West African countries.<sup>39,40</sup> These high-risk male partners are likely older men with their own networks of sexual partners who may have more power in relationships, thus compromising young women's ability to negotiate condom use.<sup>35,41,42</sup>

As far as we know, this paper is one of the only longitudinal studies that formally tests the mechanisms through which transactional sex increases HIV acquisition in young women. It is based on a biological outcome measure of HIV, not self-reported sexual behaviors. However, there are a few limitations to consider. We recognize that violations or near violations of positivity can be of concern in this causal analysis given the small proportion of young women who, at baseline, report any transactional sex, who report sex with older partners, and who report multiple sex partners. This can cause some participants to have high probability, as fewer individuals within a given covariate stratum have the exposure. Hence, the probability of those "rare" individuals who do have the exposure become more extreme. To show that there has not been a violation of positivity, we have included a table (see Appendix 3, Supplemental Digital Content, http://links.lww.com/QAI/B395) that indicates that the cells for our covariates have sufficient sample numbers. Furthermore, our models converge, and the standard errors



**FIGURE 2.** Cumulative HIV incidence by frequent transactional sex and by time since study enrolment in a Monte Carlo sample of 20,000 observations accounting for confounding. A, Illustrates the total effect of the frequent transactional sex on HIV. B, Illustrates the CDE under the condition that young women have an older partner and have more than one sex partner.

and coefficients are reasonable suggesting that data sparsity is not an issue. Relatedly, as noted in Kilburn et al, there might have been underreporting or misreporting of sexual behaviors, especially as we found incident HIV among young women that did not report ever having sex. This was despite the use of ACASI to minimize reporting bias. However, we conducted a sensitivity analysis among sexually active girls only and found similar results. There might have been some misreporting when it came to sexual partner number and partner age difference, in particular, because of issues of social-desirability and recall bias. If we assume that young women are not telling the truth, for example, they are reporting having a lower number of partners or underestimating their partner's age, it may result in underestimating the effect. However, there is no reason to suspect that misreporting of sexual activity would be associated with HIV status, as HIV testing occurred after young women had answered the ACASI questions. We had also planned to include condom use as a mediation variable, but the condom use variable showed collinearity with older partners and multiple sexual partners. Given that our results show that almost all the mediation was through older partners and partner number, we extrapolated that lack of condom use within such partnerships may increase HIV risk, but we could not explicitly test it in our model. Furthermore, we planned to test low sexual relationship power using the sexual relationship power scale, but had issues with a high percentage of missing values that could not be addressed with multiple imputation methods. However, again we can extrapolate how low power relates to older partners and condom use negotiation to explain the mechanism further. Finally, our analysis assumes no unmeasured confounding that is impossible to assess in the data. However, we explored measured confounding by examining the effect of adding and removing different variables to our models.

### CONCLUSION

To reduce the burden of HIV faced by young women in SSA, we need to examine transactional sex and the pathways to HIV infection. Our analysis demonstrates that for young women engaging in frequent transactional sex, having older and multiple sexual partners helps explain their increased HIV risk. This may be due to the underlying risk profile of these older men that young women have as sexual partners and the density of sexual networks. Interventions addressing transactional sex should target young women and men's gendered expectations of male provision<sup>4</sup> and promote notions of equitable relationships that include critical reflections on agency and power to influence young women's choice in partners.<sup>29,43,44</sup> Furthermore, programs that tackle relationship dynamics, individual beliefs and psycho-social aspects of adolescence, alongside economic opportunities for young women transitioning to adulthood are promising, especially if tailored to the socio-economic context to reduce reliance on risky partnerships.<sup>45</sup> From a research perspective, using improved measures for transactional sex to capture primary motivations is important to understand risk and having better measures for sexual relationship power may enable further confirmation of these pathways.<sup>46</sup> Research should also focus on influences that shape engagement in transactional sex and the underlying developmental trajectories of these young women within overarching systems of gendered social and economic inequalities in different contexts.

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#### REFERENCES

- 1. UNAIDS. Global AIDS Update. Geneva, Switzerland: UNAIDS; 2016.
- Wilson CM, Wright PF, Safrit JT, et al. Epidemiology of HIV infection and risk in adolescents and youth. *J Acquir Immune Defic Syndr.* 2010; 54(suppl 1):S5–S6.
- UNAIDS. Global Report: UNAIDS Report on the Global AIDS Epidemic 2012. Geneva, Switzerland: UNAIDS; 2012.
- Stoebenau K, Heise L, Wamoyi J, et al. Revisiting the understanding of transactional sex in sub-Saharan Africa: a review and synthesis of the literature. Soc Sci Med. 2016;168:186–197.
- Wamoyi J, Stobeanau K, Bobrova N, et al. Transactional sex and risk for HIV infection in sub-Saharan Africa: a systematic review and metaanalysis. J Int AIDS Soc. 2016;19:20992.
- Ranganathan M, Heise L, Pettifor A, et al. Transactional sex among young women in rural South Africa: prevalence, mediators and association with HIV infection. J Int AIDS Soc. 2016;19:20749.
- Kilburn K, Ranganathan M, Stoner MCD, et al. Transactional sex and incident HIV infection in a cohort of young women from rural South Africa. *AIDS*. 2018;32:1669–1677.
- Dunkle KL, Jewkes RK, Brown HC, et al. Transactional sex among women in Soweto, South Africa: prevalence, risk factors and association with HIV infection. *Soc Sci Med.* 2004;59:1581–1592.
- Jewkes R, Dunkle K, Nduna M, et al. Transactional sex and HIV incidence in a cohort of young women in the stepping stones trial. *J AIDS Clin Res.* 2012;3:158.
- The Joint United Nations Programme on HIV and AIDS (UNAIDS). *Transactional Sex and HIV Risk: From Analysis to Action*. Geneva, Switzerland: Joint United Nations Programme on HIV/AIDS and STRIVE; 2018.
- Ranganathan M, MacPhail C, Pettifor A, et al. Young women's perceptions of transactional sex and sexual agency: a qualitative study in the context of rural South Africa. *BMC Public Health.* 2017;17:666.
- Moore AM, Biddlecom AE, Zulu EM. Prevalence and meanings of exchange of money or gifts for sex in unmarried adolescent sexual relationships in Sub-Saharan Africa. *Afr J Reprod Heal*. 2007;11:44–61.
- 13. Luke N. Confronting the "sugar daddy" stereotype: age and economic asymmetries and risky sexual behavior in urban Kenya. *Int Fam Plan Perspect*. 2005;31:6–14.
- Okigbo CC, McCarraher DR, Chen M, et al. Risk factors for transactional sex among young females in post-conflict Liberia. *Afr J Reprod Health.* 2014;18:133–141.
- Harrison A, Newell ML, Imrie J, et al. HIV prevention for South African youth: which interventions work? A systematic review of current evidence. *BMC Public Health.* 2010;10:102.
- Leclerc-Madlala S. Transactional sex and the pursuit of modernity. Soc Dyn. 2003;29:213–233.
- Luke N. Age and economic asymmetries in the sexual relationships of adolescent girls in sub-Saharan Africa. *Stud Fam Plann.* 2003;34:67–86.
- Maganja RK, Maman S, Groves A, et al. Skinning the goat and pulling the load: transactional sex among youth in Dar es Salaam, Tanzania. *AIDS Care*. 2007;19:974–981.

- 19. Agincourt Health and Population Unit. *Changing Lives in Rural South Africa: Annual Research Brief*, 2011. MRC/Wits Rural Public Health and Health Transitions Research Unit (Agincourt), South Africa.
- Madhavan S, Townsend N. The social context of children's nutritional status in rural South Africa. Scand J Public Health Suppl. 2007;69: 107–117.
- Collinson M, Tollman S, Kahn K. Migration, settlement change and health in post-apartheid South Africa: triangulating health and demographic surveillance with national census data. *Scand J Public Health Suppl.* 2007;69:77–84.
- Pettifor A, MacPhail C, Selin A, et al. HPTN 068: a randomized control trial of a conditional cash transfer to reduce HIV infection in young women in South Africa—study design and baseline results. *AIDS Behav.* 2016;20:1863–1882.
- Pettifor A, MacPhail C, Hughes JP, et al. The effect of a conditional cash transfer on HIV incidence in young women in rural South Africa (HPTN 068): a phase 3, randomised controlled trial. *Lancet Glob Heal*. 2016;4: e978–e988.
- Kahn K, Collinson MA, Gómez-Olivé FX, et al. Profile: Agincourt health and socio-demographic surveillance system. *Int J Epidemiol.* 2012;41: 988–1001.
- Radloff LS. The CES-D scale: a self-report depression scale for research in the general population. *Appl Psychol Meas.* 1977;1:385–401.
- Stoner MCD, Pettifor A, Edwards JK, et al. The effect of school attendance and school dropout on incident HIV and HSV-2 among young women in rural South Africa enrolled in HPTN 068. *AIDS*. 2017;31: 2127–2134.
- Lin SH, Young J, Logan R, et al. Parametric mediational g-formula approach to mediation analysis with time-varying exposures, mediators, and confounders. *Epidemiology*. 2017;28:266–274.
- 28. Daniel MR, dE Stavola BL, Cousens SN. gformula: estimating causal effects in the presence of time-varying confounding or mediation using the g-computation formula. *STATA J.* 2011;11:479–517.
- 29. Stoner MCD, Edwards JK, Miller WC, et al. Does partner selection mediate the relationship between school attendance and HIV/Herpes Simplex virus-2 among adolescent girls and young women in South Africa. J Acquir Immune Defic Syndr. 2018;79:20–27.
- Keil AP, Edwards JK, Richardson DB, et al. The parametric g-formula for time-to-event data: intuition and a worked example. *Epidemiol.* 2014; 25:889–897.
- Schaefer R, Gregson S, Eaton JW, et al. Age-disparate relationships and HIV incidence in adolescent girls and young women: evidence from Zimbabwe. *AIDS*. 2017;31:1461–1470.

- Ott MQ, Barnighausen T, Tanser F et al. Age-gaps in sexual partnerships: seeing beyong "sugar daddies." *AIDS*. 25:861–863, 2011.
- Stoner MCD, Nguyen N, Kilburn K, et al. Age-disparate partnerships and incident HIV infection in adolescent girls and young women in rural South Africa. *AIDS*. 2019;33:83–91.
- Pettifor AE, Rees HV, Kleinschmidt I, et al. Young people's sexual health in South Africa: HIV prevalence and sexual behaviors from a nationally representative household survey. *AIDS*. 2005;19:1525–1534.
- Leclerc-Madlala S. Age-disparate and intergenerational sex in southern Africa: the dynamics of hypervulnerability. *AIDS*. 2008;22(suppl 4):S17–S25.
- 36. Steffenson AE, Pettifor AE, Seage GR, et al. Concurrent sexual partnerships and human immunodeficiency virus risk among South African youth. *Sex Transm Dis.* 2011;38:459–466.
- Stern E, Buikema R. The relational dynamics of hegemonic masculinity among South African men and women in the context of HIV. *Cult Health Sex.* 2013;15:1040–1054.
- 38. Luke N. Exchange and condom use in informal sexual relationships in urban Kenya. *Econ Dev Cult Change*. 2006;54:319–348.
- Prudden HJ, Beattie TS, Bobrova N, et al. Factors associated with variations in population HIV prevalence across West Africa: findings from an ecological analysis. *PLoS One.* 2015;10:1–15.
- Halperin DT, Epstein H. Concurrent sexual partnerships help to explain Africa's high HIV prevalence: implications for prevention. *Lancet.* 2004; 364:4–6.
- 41. Jewkes R, Morrell R, Sikweyiya Y, et al. Transactional relationships and sex with a woman in prostitution: prevalence and patterns in a representative sample of South African men. *BMC Public Health.* 2012;12:325.
- 42. Swidler A, Watkins SC. Ties of dependence: AIDS and transactional sex in rural Malawi. *Stud Fam Plann.* 2007;38:147–162.
- 43. Duflo E, Dupas P, Kremer M, et al. Education and HIV/AIDS Prevention: Evidence from a Randomized Evaluation in Western Kenya (English). Policy, Research working paper; no. WPS 4024. Washington, DC: Background paper to the 2007 World Development Report; 2007.
- 44. Dunbar MS, Maternowska MC, Kang M, et al. Findings from SHAZI: a feasibility study of a microcredit and life-skills HIV prevention intervention to reduce risk among adolescent female orphans in Zimbabwe. J Prev Interv Community. 2010;38:147–161.
- 45. Jewkes R, Morrell R. Sexuality and the limits of agency among South African teenage women: theorising femininities and their connections to HIV risk practices. *Soc Sci Med.* 2012;74:1729–1737.
- 46. Wamoyi J, Ranganathan M, Kyegombe N, et al. Improving the measurement of transactional sex in sub-Saharan Africa: a critical review. *J Acquir Immune Defic Syndr.* 2019;80:367–374.