

---

Faculty of Human & Social Development

Faculty Publications

---

This is a post-print version of the following article:

The Additive Effects of Depressive Symptoms and Polysubstance Use on HIV Risk Among Gay, Bisexual, and Other Men Who Have Sex With Men

Card, K. G., Lachowsky, N. J., Armstrong, H. L., Cui, Z., Wang, L., Sereda, P., Jollimore, J., Patterson, T. L., Corneil, T., Hogg, R. S., Roth, E. A., & Moore, D. M.

2018

[This article is distributed under the terms of the Creative Commons Attribution License. <https://creativecommons.org/licenses/by-nc-nd/4.0> ]

The final publication is available at:

<https://doi.org/10.1016/j.addbeh.2018.03.005>

---

Citation for this paper:

Card, K. G., Lachowsky, N. J., Armstrong, H. L., Cui, Z., Wang, L., Sereda, P., Jollimore, J., Patterson, T. L., Corneil, T., Hogg, R. S., Roth, E. A., & Moore, D. M. (2018). "The Additive Effects of Depressive Symptoms and Polysubstance Use on HIV Risk Among Gay, Bisexual, and Other Men Who Have Sex With Men." *Addictive Behaviors*, 82, 158–165. <https://doi.org/10.1016/j.addbeh.2018.03.005>



# HHS Public Access

Author manuscript

*Addict Behav.* Author manuscript; available in PMC 2019 July 01.

Published in final edited form as:

*Addict Behav.* 2018 July ; 82: 158–165. doi:10.1016/j.addbeh.2018.03.005.

## The Additive Effects of Depressive Symptoms and Polysubstance Use on HIV Risk Among Gay, Bisexual, and Other Men Who Have Sex with Men

Kiffer G. Card<sup>1,2</sup>, Nathan J. Lachowsky<sup>4,8</sup>, Heather L. Armstrong<sup>1,3</sup>, Zishan Cui<sup>1</sup>, Lu Wang<sup>1</sup>, Paul Sereda<sup>1</sup>, Jody Jollimore<sup>5</sup>, Thomas L. Patterson<sup>6</sup>, Trevor Corneil<sup>3</sup>, Robert S. Hogg<sup>1,2</sup>, Eric A. Roth<sup>7</sup>, and David M. Moore<sup>1,3</sup>

<sup>1</sup>British Columbia Centre for Excellence in HIV/AIDS, Vancouver, British Columbia, Canada

<sup>2</sup>Faculty of Health Sciences, Simon Fraser University, Burnaby, British Columbia, Canada

<sup>3</sup>Faculty of Medicine, University of British Columbia, Vancouver, British Columbia, Canada

<sup>4</sup>School of Public Health and Social Policy, University of Victoria, Victoria, British Columbia, Canada

<sup>5</sup>Community Based Research Centre for Gay Men's Health, Vancouver, British Columbia, Canada

<sup>6</sup>University of California – San Diego, San Diego, California, United States

<sup>7</sup>Department of Anthropology, University of Victoria, Victoria, British Columbia, Canada

<sup>8</sup>Centre for Addictions Research of British Columbia, Victoria, British Columbia, Canada

### Abstract

**Introduction**—Among gay, bisexual, and other men who have sex with men (GBM), collinearity between polysubstance use and mental health concerns has obscured their combined effects on HIV risk with multivariable results often highlighting only one or the other.

**Methods**—We used mediation and moderation analyses to examine the effects of polysubstance use and depressive symptoms on high-risk sex (i.e., condomless anal sex with serodiscordant/unknown status partner) in a sample of sexually-active GBM, aged >16 years, recruited in Metro Vancouver using respondent driven sampling. Hospital Anxiety and Depression Scale scores assessed mental health. Alcohol Use Disorder Identification Test scores assessed alcohol disorders. Poly-use of multiple drug types (e.g., stimulants, sedatives, opiates, hallucinogens) was assessed over the previous six months.

---

<sup>§</sup>Corresponding Author: Kiffer G. Card, c/o Faculty of Health Sciences, 8888 University Drive, Burnaby, BC V5A 1S6, 778-988-4269, kcard@sfu.ca.

#### Data Availability

Data are available through the B.C. Centre for Excellence in HIV/AIDS. Please contact Dr. Robert S. Hogg (rhogg@sfu.ca) for additional information.

**Publisher's Disclaimer:** This is a PDF file of an unedited manuscript that has been accepted for publication. As a service to our customers we are providing this early version of the manuscript. The manuscript will undergo copyediting, typesetting, and review of the resulting proof before it is published in its final citable form. Please note that during the production process errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.

**Results**—Among 719 predominantly white (68.0%), gay-identified (80.7%) GBM, alcohol use was not associated with increased prevalence of high-risk sex. Controlling for demographic factors and partner number, an interaction between polysubstance use and depressive symptoms revealed that the combined effects were additively associated with increased odds for high-risk sex. Mediation models showed that polysubstance use partially mediated the relationship between depressive symptoms and high-risk sex.

**Conclusion**—An interaction effect between polysubstance use (defined by using 3 or more substances in the past six months) and depressive symptoms (defined by HADS scores) revealed that the combination of these factors was associated with increased risk for high-risk sex – supporting a syndemic understanding of the production of HIV risk.

### Keywords

Gay and bisexual men; Polysubstance use; Alcohol use; HIV risk; Depressive Symptoms

---

## 1. Introduction

Gay, bisexual, and other men who have sex with men (GBM) represent a disproportionate percentage of new HIV infections in North America (Hogg et al., 2012). In Canada, some estimates suggest that GBM are as much as 71 times as likely as other men to contract HIV (Public Health Agency of Canada, 2014); and despite expanded prevention efforts and advancements in HIV care, the number of new diagnoses among GBM has remained stable over the past decade (Public Health Agency of Canada, 2014, 2015).

Among several salient factors contributing to HIV risk, adverse mental health conditions among GBM are regularly identified as covariates of increased sexual risk (Drabkin et al., 2013; Sander et al., 2013). For instance, depression has been associated with a nearly 10-fold increase in serodiscordant condomless anal sex (Reisner et al., 2009). Furthermore, depression is highly prevalent in the general population, among sexual minorities, and among people living with HIV (Kessler, Chiu, Demler, & Walters, 2005; Klein, 2014; Lai, Cleary, Sitharthan, & Hunt, 2015). However, the mechanisms that underlie the association between depression and HIV risk remain poorly understood.

One possible explanation for this association is the high prevalence of substance use among people living with depression. A recent meta-analysis of 22 epidemiological studies reported that individuals who used illicit drugs had 3.8 times greater odds of reporting depression (Lai et al., 2015). Similarly, recent examinations of the relationship between substance use and poor mental health among GBM found that depression was associated with 3.5 times greater odds of reporting a doctor-diagnosed substance use disorder (Lachowsky et al., 2017). Together these findings highlight significant intercorrelations between substance use and mental health – a phenomena which has come to be identified with syndemics theory (Stall et al., 2003). Summarizing the relevance of syndemics theory to the present subject, syndemics describe the additive effects of two or more health conditions that contribute to excess burden of disease among vulnerable persons. Being two regularly identified syndemic factors that impact the health of GBM (Halkitis et al., 2015; Wilson et al., 2014), the combined effect of co-occurrent substance use and mental health challenges is likely to

increase risk for HIV acquisition and transmission more than the effect of either concern individually.

Indeed, two potential mechanisms have already been identified that link substance use to condomless anal sex (Moss & Albery, 2009) – a common risk factor for HIV. The first proposed mechanism demonstrates a proximal suppressant effect on cognitive capacity whereby intoxicated persons become more easily aroused and less capable of implementing risk management strategies, such as condom use (Rehm, Shield, Joharchi, & Shuper, 2012; Scott-Sheldon, Carey, Cunningham, Johnson, & Carey, 2016). The second proposed mechanism posits that escape and pleasure motivated sexual and substance use expectancies – perhaps undergirded by mental health conditions – confound the relationship between substance use and sexual risk, and that, in fact, individuals use psychoactive substances as a means to escape emotional and psychological stressors and enhance sexual pleasure – abstaining from condoms for this same reason (George, Stoner, Norris, Lopez, & Lehman, 2000; Sternberg, 2011).

However, because substance use and mental health are highly collinear, one or the other is often found to be non-significant when examining the factors that contribute to HIV risk (Deuba et al., 2013; Kelly et al., 2013; Parsons, Lelutiu-Weinberger, Botsko, & Golub, 2013). Furthermore, recent rights-based models of substance use, mental health, and sexual behavior have taken issue with the pathologizing of these phenomenon (De Block & Adriaens, 2013; Kardefelt-Winther et al., 2017; Wasserman & Wasserman, 2016); and indeed a number of studies report that only at the extremes are substance use and mental health conditions linked to sexual risk behavior (Fendrich, Avci, Johnson, & Mackesy-Amiti, 2013) – suggesting that significant socio-structural traumas, rather than particular typologies of substance use or mental health conditions, contribute to HIV risk. These considerations highlight the need to more carefully evaluate the relationships between mental health, substance use, and sexual risk – particularly among vulnerable populations such as GBM.

Consistent with a syndemic conception of the above described relationships, the present analysis used mediation and moderation analyses to examine the association between high-risk sexual behavior, depression scores, and polysubstance use among a sample of GBM recruited using respondent-driven sampling (RDS). This allowed us to achieve more reliable population-based estimates of mental health, substance use, and sexual risk – adding to the array of sampling methods which have sought to improve on traditional convenience samples, including random-digit-dialing (Stall et al., 2003), venue-based time-location sampling (Matthews et al., 2016), and national surveillance sampling (Cochran & Mays, 2000).

Using these approaches, we hypothesized that: (i) polysubstance use would partially mediate the association between depressive symptoms and high-risk sex, and (ii) an interaction term between polysubstance use and depressive symptoms would better characterize these associations in multivariate models. In doing so, the present analysis leverages previously established conventions to offer the following strengths: (i) focuses only on sexual behavior which poses a risk for HIV transmission (Jin et al., 2015); (ii) recruits GBM in a way that

allows for adjustments to account for sampling bias (Heckathorn, 2011); (iii) uses validated scales of alcohol dependence and depressive symptoms as opposed to participant reported past history of doctor diagnosed depression to reduce misclassification bias (Ferguson, 2000; McGinnis et al., 2013); and (iv) focuses on polysubstance use rather than any substance use – the latter of which has poor specificity among GBM (McCarty-Caplan, Jantz, & Swartz, 2014).

## 2. Methods

### 2.1. Study Protocol

We used respondent driven sampling (RDS) to recruit a cohort of GBM in Metro Vancouver, British Columbia. Inclusion criteria restricted participation in the study to individuals who were (i) 16 years or older, (ii) gender identified as male, (iii) lived in the Metro Vancouver area, (iv) had sex with another man in the previous 6 months, and (v) could complete a questionnaire in English. Initial seeds were recruited from community contacts and geosocial networking applications (Lachowsky, et al., 2016), following formative research which mapped the relevant in-person and online networks in Metro Vancouver (Forrest et al., 2014). Respondents completed a computer-administered self-interview (CASI) at our study site in Vancouver's downtown West End (an area traditionally thought of as Vancouver's "gay neighbourhood") and then a nurse-administered questionnaire and sexual health check-up with serology for HIV, HCV, and syphilis. Ethical approval for this study was granted by the research ethic boards of Simon Fraser University, the University of British Columbia, the Providence Healthcare Research Institute, and the University of Victoria.

### 2.2. Dependent Variable

High-risk sexual behavior was assessed by asking participants to report the HIV serostatus of sexual partners with whom they had had condomless anal sex (CAS) in the past six months (P6M). Individuals who reported CAS with a partner who was serodiscordant or whose HIV status they did not know were classified as having engaged in "high-risk sex," and individuals who reported not engaging in CAS or who engaged in CAS with only seroconcordant partners were classified as not having engaged in "high-risk sex." This cut-off provides a more accurate representation of HIV risk as there is reduced risk for HIV transmission between known-concordant partners (Jin et al., 2015).

### 2.3. Independent Variables

Substance use behaviour was measured by self-report and participants were asked to report use of drugs in the following categories over the P6M: stimulants (i.e., crack/cocaine), prescription stimulants (i.e., Ritalin, Concerta, Adderall), methamphetamine (i.e., crystal methamphetamine, speed), inhalants (i.e., nitrous oxide), sedatives (i.e., gamma-hydroxybutyric acid), benzodiazepine, barbiturates), hallucinogens (e.g., lysergic acid diethylamide, psilocybin, ketamine, methylenedioxymethamphetamine), street opioids (i.e., heroin), prescription opioids (i.e., morphine, codeine, Oxycontin, Percocet), erectile dysfunction drugs, poppers (e.g., alkyl nitrate), cannabis, and other drugs. Polysubstance use was classified by the number of categories from which a participant reported using a drug.

Continuous counts were trichotomized as (i) no substance use, (ii) one or two categories of substance use, and (iii) three or more categories of substance use. For the purposes of the present study, alcohol use was assessed separately from polysubstance use using the Alcohol Use Disorder Identification Test (AUDIT; Saunders, Aasland, Babor, de la Fuente, & Grant, 1993). This was done because the majority of participants reported alcohol consumption over the P6M. The AUDIT is a brief screening measure that assesses harmful alcohol use and dependence and has been previously shown to have high sensitivity and specificity among both HIV-negative and HIV-positive men, even compared to other alcohol screening measures (McGinnis et al., 2013). The scale consists of three items measuring alcohol consumption, three items measuring drinking behavior, two items assessing adverse reactions, and two items assessing alcohol-related problems. Items are scored on zero-weighted five-point Likert-scales and final scores range from 0 (no harmful or dependent alcohol use) to 40 (harmful or dependent alcohol use). Clinical guidelines developed for the AUDIT suggest that scores >20 warrant treatment of alcohol dependence, scores 16–19 warrant counseling and continued monitoring, and lower scores require limited to no intervention (Miller, 2014). Balancing the sensitivity and specificity of these categorizations, the AUDIT scale was dichotomized in the present study with values between 0 and 15 indicating “low or medium risk” and scores between 16 and 40 indicating “harmful use and possible dependence” (Lachowsky et al., 2017).

To assess depressive symptoms, the depression subscale score from the Hospital Anxiety and Depression Scale (HADS; Snaith, 2003) was used. This subscale consists of seven items (e.g., “I feel as if I am slowed down”) scored on a zero-weighted four-point Likert scale. Final scores range from 0 (no depressive symptoms) to 21 (severe depressive symptoms). Based on original score cut-offs and literature showing that more severe, rather than less severe, depression is associated with abnormal sexual risk (Fendrich et al., 2013; Zigmond & Snaith, 1983), scale scores were dichotomized with scores less than or equal to 10 indicating “normal/borderline depressive symptoms” and scores greater than 10 indicating “abnormal depressive symptoms” (Whelan-Goodinson, Ponsford, & Schönberger, 2009). Further, diagnostic evaluations of the HADS depression subscale have shown that cut offs around 10 have acceptable specificity and sensitivity (Honarmand & Feinstein, 2009; Hung, Liu, Wang, Yao, & Yang, 2012; Whelan-Goodinson et al., 2009) and may better represent depression than a history of doctor diagnosed depression (Golden, Conroy, & O’Dwyer, 2007).

#### 2.4. Other Covariates

To control for extraneous variation, baseline cross-sectional data used in this analysis included the following sociodemographic variables: age, sexual identity (gay, bisexual, other), race/ethnicity (White, Asian, Indigenous, Latin American, other), annual income (< \$30,000 vs. \$30,000), HIV status (negative, positive), and participant’s reported number of male anal sex partners in the P6M.

#### 2.5. Statistical Analysis

All analyses used RDS-generated weights to adjust for network size and homophily using RDSAT (version 7.1.46, Voltz et al., 2012) and SAS (version 9.4, SAS, Cary, North

Carolina, USA). Bivariate statistics for demographic, mental health, and substance use factors associated with “any high-risk sex” (vs. none) were generated using simple logistic regression to identify candidate variables for multivariate models. To adjust for the confounding effects of sociodemographic variables and partner frequency, multivariable logistic regression models were constructed using a backwards selection approach wherein at each step the variable with the largest Type-III p-value was omitted until the Akaike information criterion was optimized (minimized). Only variables with a univariable p-value <0.20 were initially included. Moderation effects of polysubstance use and depressive symptoms were examined by including an interaction term between the two variables. A mediation analysis also examined the effect of polysubstance use (i.e., reporting three or more substances vs. none) on the relationship between depressive symptoms and high-risk sex. A partial posterior method (i.e., an inferential approach that assesses the weighting of the null hypothesis by assessing the distribution of the mediating effect under the null hypothesis of no mediation) was used to test the significance of indirect effects of depression on high-risk sex via polysubstance use (Biesanz, Falk, & Savalei, 2010; Falk & Biesanz, 2016). This method provides a p-value interpretable in the same way as the p-value from Sobel’s test, but offers greater power that is comparable to other commonly used alternatives (e.g., bootstrap and Monte Carlo). Finally, a ratio of the unadjusted and adjusted effects on the association between depression and high-risk sex was used to calculate the proportion of variance explained by polysubstance use.

### 3. Results

A total of 719 respondents were recruited, of which 119 were seeds. Of these, 703 provided valid responses for our primary outcome variable. Table 1 provides RDS-adjusted study characteristics for the study sample, stratified by the primary outcome variable of interest (i.e., high-risk sex). Univariate p-values are also provided testing the association between each factor and the outcome of interest. Overall, 23.4% of respondents were HIV-positive, 80.7% were gay identified, 68.0% were White, the median age was 33 years old (Q1-Q3: 26–47), 65.6% had at least some post-secondary education, 74.3% made less than \$30,000 per year, 19.0% had moderate or severe HADS depression sub-scale scores, 57.1% had moderate or severe HADS anxiety sub-scale scores, 36.1% reported using >3 substances (excluding alcohol), 13.2% had AUDIT scale scores indicating harmful use and possible dependence, and 35.9% reported serodiscordant or unknown CAS (i.e., high-risk sex) in the P6M with an additional 26.0% reporting CAS only with a seroconcordant partner. Prevalence estimates for each substance used in the past six months are also provided in Supplemental Table 1.

Univariate interactions with HADS Depression were examined for (i) AUDIT scores ( $p = 0.686$ ) and (ii) polysubstance use ( $p = 0.032$ ), and an interaction term was selected for inclusion in final multivariable modeling. Neither the effects of period prevalent alcohol use (P6M, OR = 1.05, 95% CI: 0.68, 1.61), nor elevated (i.e., harmful/possible dependence) AUDIT scores (P6M, OR = 1.29, 95% CI: 0.84, 1.99) were significantly associated with high-risk sex. Setting alcohol use aside, Table 2 shows that the effects of polysubstance use on high-risk sex varied across depression levels, as did the effects of depression across polysubstance use levels. Among those with normal/borderline HADS scores, those who

used 3 or more substances were more likely to report high-risk sex (AOR = 1.65, 95%CI: 1.01, 2.68) than those without any substance use; and among those with abnormal HADS Depression Scores, the association between the highest level of polysubstance use and high-risk sex increased thirteen-fold (aOR = 20.76, 95% CI: 1.66, 260.40). Presenting an alternative way of interpreting these results: among those with the highest level of polysubstance use (>3 reported substances), depression was positively associated with high-risk sex (aOR: 5.44, 95% CI: 1.56, 19.00); whereas at lower levels of polysubstance use, this relationship was not significant (1–2 substances: aOR: 1.18, 95% CI: 0.35, 3.99; 0 substances: aOR: 0.43, 95% CI: 0.05, 4.09).

As shown in Figure 1, univariate results showed that abnormal HADS depression scores were associated with high-risk sex (OR: 2.77, 95% CI: 1.20, 6.36) and with the highest level of polysubstance use (OR: 2.62, 95% CI: 1.06, 6.45). When adjusting for polysubstance use, the odds between depression and engaging in high-risk sex diminished to 2.41 (95% CI: 1.03, 5.62) and those with the highest level of polysubstance use had 2.11 times greater odds of engaging in high-risk sex (95% CI: 1.39, 3.21). Partial posterior p-value for indirect effect indicated that the mediation effect was significant ( $p = 0.026$ ). These results suggest that, 18.2% of the effect of depression on high-risk sex was mediated by polysubstance use.

## 4. Discussion

### 4.1. Primary Findings

In the present study we found that among 719 GBM recruited using RDS in Vancouver, Canada, neither period-prevalence alcohol use (any vs. none) nor AUDIT scores were associated with increased prevalence of high-risk sex (defined by condomless anal sex with a serodiscordant or unknown status partner). However, these findings should be interpreted within the broader literature on this topic which has produced mixed findings (Colfax et al., 2004; Sander et al., 2013; Vosburgh, Mansergh, Sullivan, & Purcell, 2012). Setting alcohol use aside and controlling for age, ethnicity, income, and number of sexual partners, an interaction effect between polysubstance use (defined by using 3 or more substances in the P6M) and depressive symptoms (defined by abnormal HADS scores) revealed that the combination of these factors was associated with greater likelihood of high-risk sex – supporting a syndemic understanding of the production of GBM's sexual behaviour and HIV risk (Halkitis, Moeller, et al., 2013; Kurtz, Buttram, Surratt, & Stall, 2012; Singer, Bulled, Ostrach, & Mendenhall, 2017).

With respect to previous research, our findings offer several important insights into the relationship between mental health, substance use, and HIV risk. First, they suggest that while alcohol use may be a driving factor for condomless sex, it is not necessarily a driving factor for HIV risk (Sandfort et al., 2017). In previous analyses we showed that alcohol use was associated with higher odds of condomless anal sex (Card et al., 2017; Lachowsky, et al., 2016), but in the present analyses, alcohol use was not associated with condomless anal sex with a serodiscordant or unknown serostatus partner. This suggests that alcohol use may have a complicated relationship with participant's patterns of seroadaptation, strategies used by GBM to facilitate sexual intimacy and sensation seeking during sexual intercourse (McKirman, Ostrow, & Hope, 1996). This does not rule out the possibility that alcohol use

has an inhibitory cognitive effect on condom use, but does suggest that the inhibition may have a less pronounced effect on partner selection.

With that said, the interaction between polysubstance use and depression underscores the syndemic effects of comorbid health conditions on shaping patterns of HIV risk. As noted by Stall et al., there has long been irrefutable evidence of the disparities in disease concentration among marginalized groups such as MSM (Stall, Coulter, Friedman, & Plankey, 2015). The present study expands this body of literature by providing evidence for an interaction effect between syndemic factors. Specifically, we have shown that the effects of depression and polysubstance use on high-risk sex were only statistically significant at the highest cut-offs for these measures (See also, Fendrich et al., 2013), suggesting that in addition to possible dose-response relationships, there is also a synergistic relationship wherein the combined effects of comorbidities, in addition to the singular morbidities themselves, have a profound effect on shaping high-risk sexual behavior (Singer et al., 2017). In summary, our findings are consistent with empirical research showing that syndemic production of HIV through depression, polydrug use, and sexual trauma is strongly predictive of HIV seroconversion among GBM (Guadamuz et al., 2014; Mimiaga et al., 2015).

Examining a rationale for the association between depression and substance use, our mediation analysis underscores the need to better describe the mechanisms that underlie the association between depression and condomless anal sex. While we, and others (Hutton, Lyketsos, Zenilman, Thompson, & Erbedding, 2004), have hypothesized that substance use would have a large mediating effect in this relationship, in our study it explained less than 20% of the direct effect of depression on high-risk sex. This underscores the need to examine syndemics not only with respect to their behavioral endpoints, but also the antecedent biological factors that prime these behaviors (Singer et al., 2017). Regarding sexual risk, these include the endocrinological effects of glucocorticoid regulation that aggravates the stress-response, which in turn impacts the neural-cognitive processes underlying risk perception and sensation seeking behavior (Arnsten, 2009; Goldey & van Anders, 2012; Harrison, Ratcliffe, Mitchell, & Smith, 2014; Mehta, Welker, Zilioli, & Carré, 2015; Shabani, Dehghani, Hedayati, & Rezaei, 2011).

## 4.2. Implications

The implications of our research highlight the importance of not only considering the comorbid effects of depression and substance use when screening for individuals who are at greatest risk for HIV infection, but also providing competent holistic care that can address the combination of these comorbidities as well as the underlying factors that give rise to them (Halkitis, Moeller, et al., 2013; Safren, Blashill, & O'Cleirigh, 2011; Wim, Christiana, & Marie, 2014). With an increasing number of GBM undergoing more formal evaluations of HIV risk behavior as part of the expansion of PrEP, these screenings may provide opportunities for identifying and addressing mental health concerns as well. Integrated care services may thus provide a new standard of care compared to services only addressing traditional prevention and care strategies (Halkitis, Wolitski, & Millett, 2013). As part of these interventions, identifying and enhancing patterns of resilience among GBM is believed

to be a productive approach for alleviating health disparities (Herrick et al., 2011). Serosorting, for example, has been advanced as a resiliency strategy, but its effectiveness has been called into question by high partner frequency among those engaging in CAS (Kurtz et al., 2012) – again emphasizing the need for combined prevention focusing on traditional risk-related indicators (e.g., partner number).

### 4.3. Limitations

This study has several important limitations. First, while we have relied on previously established and validated cut-offs for assignment of depression and alcohol use disorders, it is possible that different schema would have produced different results. This is of particular concern given the small sample of individuals meeting the threshold for abnormal depression – which potentially reduced our power to detect significant effects. Second, our decision to not include alcohol use in our classification of polysubstance use was important for practical reasons related to the high prevalence of alcohol use in our sample. This might make it difficult to compare our polysubstance use measure to those used by others. However, with changes to the DSM-5, which has sought to disaggregate alcohol use and other substance use disorders (APA, 2013), we hope that studies will increasingly distinguish between these factors. Third, as the data used in the present analyses assessed period-prevalence at a population-level, future analyses should examine event- and network-level covariates, as these measures have previously been shown to produce divergent results (Vosburgh et al., 2012). Fourth, as polysubstance use is difficult to define (i.e., which drugs should be included, at what frequencies, and how many drugs qualify as “poly”), our results may not capture the full gradient of substance use patterns. More nuanced models of substance use behavior are needed, such as cluster or latent class analyses. Fifth, while CAS with serodiscordant or unknown status partners is a better proxy for HIV risk than CAS with serconcordant partners, the increasing availability of pre-exposure prophylaxis and antiretroviral therapies to prevent HIV transmission/acquisition, may make this measure of HIV risk an inadequate proxy for true epidemiological HIV risk in the future. Conversely, as recent seroconversion is often associated with being unaware of one’s HIV status, individuals may believe they or their partner is HIV-negative, when in fact they are in a stage of acute infection or long-term non-diagnosis. It is therefore possible that strategic deployment of risk management strategies has the possibility to prevent HIV transmission for even those who experience multiple syndemic comorbidities. This may introduce some misclassification error, especially considering that substance use (alcohol use in particular) has been previously correlated with unawareness of infection among HIV-positive men (Vagenas et al., 2014). Fifth, it should be noted that due to the lack of relevant empirical comparisons of RDS and other sampling techniques (e.g., online samples, time-location samples), it is unclear exactly how these methods compare in terms of recruiting samples of MSM in which there is minimal bias. As such, readers should be attentive in comparing the results of the present study to those previously generated by alternative recruitment methods.

## 5. Conclusion

In conclusion, our findings underscore the need for integrated care in the management of comorbid mental health and substance use conditions, which together promote HIV risk.

Furthermore, incomplete mediation between depression and HIV risk by substance use underscores the need to further evaluate the mechanisms by which depression contributes to HIV risk. Given the present study's support for the syndemic production of HIV-risk, such models might begin by examining other previously identified syndemic factors.

## Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

## Acknowledgments

The authors would like to thank the Momentum Study participants, office staff and community advisory board, as well as our community partner agencies, Health Initiative for Men, YouthCO HIV and Hep C Society, and Positive Living Society of BC.

**Funding** This study is funded by National Institute on Drug Abuse (R01DA031055-01A1) and the Canadian Institutes for Health Research, including through a Foundation Grant awarded to RSH (MOP-107544, 143342, PJT-153139). NJL was supported by a CANFAR/CTN Postdoctoral Fellowship Award. DMM and NJL are supported by Scholar Awards from the Michael Smith Foundation for Health Research (#5209, #16863). HLA is supported by a Postdoctoral Fellowship Award from the Canadian Institutes of Health Research (Grant # MFE-152443). KGC is funded by a UWW-Engage Fellowship.

## Abbreviations

**GBM Gay** bisexual, and other men who have sex with men

## References

- APA. The Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition.: DSM 5. American Psychiatric Association; 2013.
- Arnsten AFT. Stress signalling pathways that impair prefrontal cortex structure and function. *Nature Reviews Neuroscience*. 2009; 10(6):410–422. <https://doi.org/10.1038/nrn2648>. [PubMed: 19455173]
- Biesanz JC, Falk CF, Savalei V. Assessing Mediation Models: Testing and Interval Estimation for Indirect Effects. *Multivariate Behavioral Research*. 2010; 45(4):661–701. <https://doi.org/10.1080/00273171.2010.498292>. [PubMed: 26735714]
- Card KG, Lachowsky NJ, Cui Z, Shurgold S, Armstrong HL, Rich AJ, ... Hogg RS. An Event-Level Analysis of the Interpersonal Factors Associated With Condomless Anal Sex Among Gay, Bisexual, and Other Men Who Have Sex With Men (MSM) With Online-Met Partners. *AIDS Education and Prevention: Official Publication of the International Society for AIDS Education*. 2017; 29(2):154–174. <https://doi.org/10.1521/aeap.2017.29.2.154>. [PubMed: 28467159]
- Cochran SD, Mays VM. Lifetime prevalence of suicide symptoms and affective disorders among men reporting same-sex sexual partners: results from NHANES III. *American Journal of Public Health*. 2000; 90(4):573–578. [PubMed: 10754972]
- Colfax G, Vittinghoff E, Husnik MJ, McKirnan D, Buchbinder S, Koblin B. ... EXPLORE Study Team. Substance use and sexual risk: a participant- and episode-level analysis among a cohort of men who have sex with men. *American Journal of Epidemiology*. 2004; 159(10):1002–1012. [PubMed: 15128613]
- De Block A, Adriaens PR. Pathologizing sexual deviance: a history. *Journal of Sex Research*. 2013; 50(3–4):276–298. <https://doi.org/10.1080/00224499.2012.738259>. [PubMed: 23480073]
- Deuba K, Ekström AM, Shrestha R, Ionita G, Bhatta L, Karki DK. Psychosocial Health Problems Associated with Increased HIV Risk Behavior among Men Who Have Sex with Men in Nepal: A Cross-Sectional Survey. *PLOS ONE*. 2013; 8(3):e58099. <https://doi.org/10.1371/journal.pone.0058099>. [PubMed: 23516434]

- Drabkin AS, Sikkema KJ, Wilson PA, Meade CS, Hansen NB, DeLorenzo A, ... Mayer G. Risk patterns preceding diagnosis among newly HIV-diagnosed men who have sex with men in New York City. *AIDS Patient Care and STDs*. 2013; 27(6):333–341. <https://doi.org/10.1089/apc.2012.0313>. [PubMed: 23730703]
- Falk CF, Biesanz JC. Two Cross-Platform Programs for Inferences and Interval Estimation About Indirect Effects in Mediational Models. *SAGE Open*. 2016; 6(1):2158244015625445. <https://doi.org/10.1177/2158244015625445>.
- Fendrich M, Avci O, Johnson TP, Mackesy-Amiti ME. Depression, substance use and HIV risk in a probability sample of men who have sex with men. *Addictive Behaviors*. 2013; 38(3):1715–1718. <https://doi.org/10.1016/j.addbeh.2012.09.005>. [PubMed: 23254224]
- Ferguson JM. Depression: Diagnosis and Management for the Primary Care Physician. *Primary Care Companion to The Journal of Clinical Psychiatry*. 2000; 2(5):173–178.
- Forrest, JL, Stevenson, B., Rich, A., Michelow, W., Pai, J., Jollimore, J., ... Roth, EA. Community mapping and respondent-driven sampling of gay and bisexual men's communities in Vancouver, Canada. *Culture, Health & Sexuality*. 2014. <https://doi.org/10.1080/13691058.2014.881551>
- George WH, Stoner SA, Norris J, Lopez PA, Lehman GL. Alcohol expectancies and sexuality: a self-fulfilling prophecy analysis of dyadic perceptions and behavior. *Journal of Studies on Alcohol*. 2000; 61(1):168–176. [PubMed: 10627112]
- Golden J, Conroy RM, O'Dwyer AM. Reliability and validity of the Hospital Anxiety and Depression Scale and the Beck Depression Inventory (Full and FastScreen scales) in detecting depression in persons with hepatitis C. *Journal of Affective Disorders*. 2007; 100(1–3):265–269. <https://doi.org/10.1016/j.jad.2006.10.020>. [PubMed: 17156850]
- Goldey KL, van Anders SM. Sexual thoughts: links to testosterone and cortisol in men. *Archives of Sexual Behavior*. 2012; 41(6):1461–1470. <https://doi.org/10.1007/s10508-011-9858-6>. [PubMed: 21993767]
- Guadamuz TE, McCarthy K, Wimonasate W, Thienkrua W, Varangrat A, Chaikummao S, ... van Griensven F. Psychosocial Health Conditions and HIV Prevalence and Incidence in a Cohort of Men Who have Sex with Men in Bangkok, Thailand: Evidence of a Syndemic Effect. *AIDS and Behavior*. 2014; 18(11):2089–2096. <https://doi.org/10.1007/s10461-014-0826-8>. [PubMed: 24989128]
- Halkitis PN, Kapadia F, Bub KL, Barton S, Moreira AD, Stults CB. A Longitudinal Investigation of Syndemic Conditions Among Young Gay, Bisexual, and Other MSM: The P18 Cohort Study. *AIDS and Behavior*. 2015; 19(6):970–980. <https://doi.org/10.1007/s10461-014-0892-y>. [PubMed: 25192900]
- Halkitis PN, Moeller RW, Siconolfi DE, Storholm ED, Solomon TM, Bub KL. Measurement model exploring a syndemic in emerging adult gay and bisexual men. *AIDS and Behavior*. 2013; 17(2): 662–673. <https://doi.org/10.1007/s10461-012-0273-3>. [PubMed: 22843250]
- Halkitis PN, Wolitski RJ, Millett GA. A holistic approach to addressing HIV infection disparities in gay, bisexual, and other men who have sex with men. *The American Psychologist*. 2013; 68(4): 261–273. <https://doi.org/10.1037/a0032746>. [PubMed: 23688093]
- Harrison C, Ratcliffe JM, Mitchell M, Smith MA. Cortisol reactivity to psychosocial stress is greater in sexual risk takers. *Health Psychology and Behavioral Medicine*. 2014; 2(1):221–230. <https://doi.org/10.1080/21642850.2014.889571>. [PubMed: 25750779]
- Heckathorn DD. Snowball versus Respondent Driven Sampling. *Sociological Methodology*. 2011; 41(1):355–366. <https://doi.org/10.1111/j.1467-9531.2011.01244.x>. [PubMed: 22228916]
- Herrick AL, Lim SH, Wei C, Smith H, Guadamuz T, Friedman MS, Stall R. Resilience as an Untapped Resource in Behavioral Intervention Design for Gay Men. *AIDS and Behavior*. 2011; 15(1):25–29. <https://doi.org/10.1007/s10461-011-9895-0>.
- Hogg RS, Heath K, Lima VD, Nosyk B, Kanters S, Wood E, ... Montaner JSG. Disparities in the Burden of HIV/AIDS in Canada. *PLoS ONE*. 2012; 7(11):e47260. <https://doi.org/10.1371/journal.pone.0047260>. [PubMed: 23209549]
- Honarmand K, Feinstein A. Validation of the Hospital Anxiety and Depression Scale for use with multiple sclerosis patients. *Multiple Sclerosis (Houndmills, Basingstoke, England)*. 2009; 15(12): 1518–1524. <https://doi.org/10.1177/1352458509347150>.

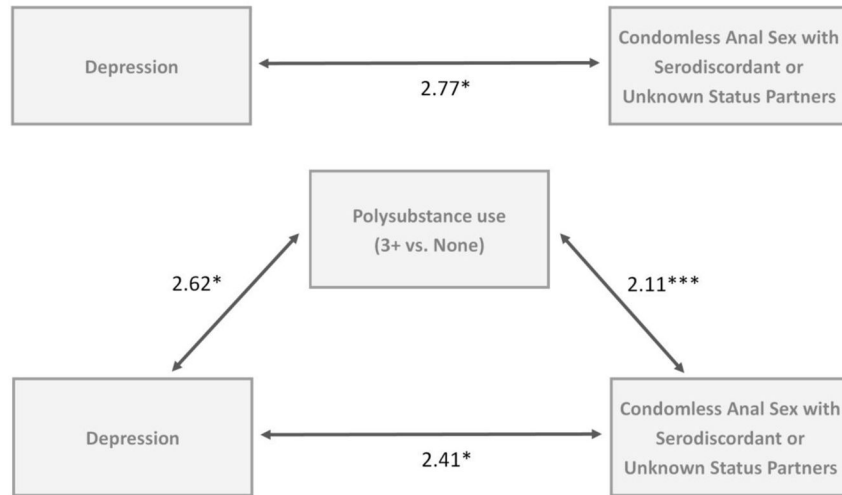
- Hung CI, Liu CY, Wang SJ, Yao YC, Yang CH. The cut-off points of the Depression and Somatic Symptoms Scale and the Hospital Anxiety and Depression Scale in detecting non-full remission and a current major depressive episode. *International Journal of Psychiatry in Clinical Practice*. 2012; 16(1):33–40. <https://doi.org/10.3109/13651501.2011.617456>. [PubMed: 22122659]
- Hutton HE, Lyketsos CG, Zenilman JM, Thompson RE, Erbeling EJ. Depression and HIV risk behaviors among patients in a sexually transmitted disease clinic. *The American Journal of Psychiatry*. 2004; 161(5):912–914. <https://doi.org/10.1176/appi.ajp.161.5.912>. [PubMed: 15121659]
- Jin, F., Prestage, GP., Mao, L., Poynten, IM., Templeton, DJ., Grulich, AE., Zablotska, I. “Any Condomless Anal Intercourse” is No Longer an Accurate Measure of HIV Sexual risk Behavior in Gay and Other Men Who have Sex with Men; *Frontiers in Immunology*. 2015. p. 6 <https://doi.org/10.3389/fimmu.2015.00086>
- Kardefelt-Winther D, Heeren A, Schimmenti A, van Rooij A, Maurage P, Carras M, ... Billieux J. How can we conceptualize behavioural addiction without pathologizing common behaviours? *Addiction (Abingdon, England)*. 2017; 112(10):1709–1715. <https://doi.org/10.1111/add.13763>.
- Kelly JA, St Lawrence JS, Amirhanian YA, DiFranceisco WJ, Anderson-Lamb M, Garcia LI, Nguyen MT. Levels and predictors of HIV risk behavior among Black men who have sex with men. *AIDS Education and Prevention: Official Publication of the International Society for AIDS Education*. 2013; 25(1):49–61. <https://doi.org/10.1521/aeap.2013.25.1.49>. [PubMed: 23387951]
- Kessler RC, Chiu WT, Demler O, Walters EE. Prevalence, Severity, and Comorbidity of Twelve-month DSM-IV Disorders in the National Comorbidity Survey Replication (NCS-R). *Archives of General Psychiatry*. 2005; 62(6):617–627. <https://doi.org/10.1001/archpsyc.62.6.617>. [PubMed: 15939839]
- Klein H. Depression and HIV Risk Taking Among Men Who Have Sex with Other Men and Who Use the Internet to Find Partners for Unprotected Sex. *Journal of Gay & Lesbian Mental Health*. 2014; 18(2):164–189. <https://doi.org/10.1080/19359705.2013.834858>. [PubMed: 26877831]
- Kurtz SP, Buttram ME, Surratt HL, Stall RD. Resilience, syndemic factors, and serosorting behaviors among HIV-positive and HIV-negative substance-using MSM. *AIDS Education and Prevention: Official Publication of the International Society for AIDS Education*. 2012; 24(3):193–205. <https://doi.org/10.1521/aeap.2012.24.3.193>. [PubMed: 22676460]
- Lachowsky NJ, Dulai JJS, Cui Z, Sereda P, Rich A, Patterson TL, ... Moore DM. Lifetime Doctor-Diagnosed Mental Health Conditions and Current Substance Use Among Gay and Bisexual Men Living in Vancouver, Canada. *Substance Use & Misuse*. 2017; 52(6):785–797. <https://doi.org/10.1080/10826084.2016.1264965>. [PubMed: 28379111]
- Lachowsky NJ, Lal A, Forrest JI, Card KG, Cui Z, Sereda P, ... Hogg RS. Including Online-Recruited Seeds: A Respondent-Driven Sample of Men Who Have Sex With Men. *Journal of Medical Internet Research*. 2016; 18(3):e51. <https://doi.org/10.2196/jmir.5258>. [PubMed: 26980147]
- Lachowsky NJ, Tanner Z, Cui Z, Sereda P, Rich A, Jollimore J, ... Roth EA. An Event-Level Analysis of Condom Use During Anal Intercourse Among Self-Reported Human Immunodeficiency Virus-Negative Gay and Bisexual Men in a Treatment as Prevention Environment. *Sexually Transmitted Diseases*. 2016; 43(12):765–770. <https://doi.org/10.1097/OLQ.0000000000000530>. [PubMed: 27832026]
- Lai HMX, Cleary M, Sitharthan T, Hunt GE. Prevalence of comorbid substance use, anxiety and mood disorders in epidemiological surveys, 1990–2014: A systematic review and meta-analysis. *Drug and Alcohol Dependence*. 2015; 154(Supplement C):1–13. <https://doi.org/10.1016/j.drugalcdep.2015.05.031>. [PubMed: 26072219]
- Matthews DD, Herrick AL, Coulter RWS, Friedman MR, Mills TC, Eaton LA, ... Stall RD. Running Backwards: Consequences of Current HIV Incidence Rates for the Next Generation of Black MSM in the United States. *AIDS and Behavior*. 2016; 20(1):7–16. <https://doi.org/10.1007/s10461-015-1158-z>. [PubMed: 26267251]
- McCarty-Caplan D, Jantz I, Swartz J. MSM and drug use: A latent class analysis of drug use and related sexual risk behaviors. *AIDS and Behavior*. 2014; 18(7):1339–1351. <https://doi.org/10.1007/s10461-013-0622-x>. [PubMed: 24065437]
- McGinnis KA, Justice AC, Kraemer KL, Saitz R, Bryant KJ, Fiellin DA. Comparing Alcohol Screening Measures Among HIV Infected and Uninfected Men. *Alcoholism, Clinical and Experimental Research*. 2013; 37(3):435–442. <https://doi.org/10.1111/j.1530-0277.2012.01937.x>.

- McKirman DJ, Ostrow DG, Hope B. Sex, drugs and escape: A psychological model of HIV-risk sexual behaviours. *AIDS Care*. 1996; 8(6):655–670. <https://doi.org/10.1080/09540129650125371>. [PubMed: 8993716]
- Mehta PH, Welker KM, Zilioli S, Carré JM. Testosterone and cortisol jointly modulate risk-taking. *Psychoneuroendocrinology*. 2015; 56(Supplement C):88–99. <https://doi.org/10.1016/j.psyneuen.2015.02.023>. [PubMed: 25813123]
- Miller, WR. *Motivational Enhancement Therapy Manual: A Clinical Research Guide for Therapists Treating Individuals with Alcohol Abuse and Dependence*. Echo Point Books & Media; 2014. (Reprint ed. edition)
- Mimiaga MJ, O’Cleirigh C, Biello KB, Robertson AM, Safren SA, Coates TJ, ... Mayer KH. The effect of psychosocial syndemic production on 4-year HIV incidence and risk behavior in a large cohort of sexually active men who have sex with men. *Journal of Acquired Immune Deficiency Syndromes (1999)*. 2015; 68(3):329–336. <https://doi.org/10.1097/QAI.0000000000000475>. [PubMed: 25501609]
- Moss AC, Albery IP. A dual-process model of the alcohol-behavior link for social drinking. *Psychological Bulletin*. 2009; 135(4):516–530. <https://doi.org/10.1037/a0015991>. [PubMed: 19586160]
- Parsons JT, Lelutiu-Weinberger C, Botsko M, Golub SA. Predictors of day-level sexual risk for young gay and bisexual men. *AIDS and Behavior*. 2013; 17(4):1465–1477. <https://doi.org/10.1007/s10461-012-0206-1>. [PubMed: 22614745]
- Public Health Agency of Canada. *Population-Specific HIV/AIDS Status Report: Gay, Bisexual, Two-Spirit and Other Men Who Have Sex With Men - Public Health Agency of Canada*. 2014 Oct 23. Retrieved August 4, 2015, from <http://www.phac-aspc.gc.ca/aids-sida/publication/ps-pd/men-hommes/index-eng.php>
- Public Health Agency of Canada. *HIV in Canada: Surveillance summary tables, 2014–2015*. 2015. Retrieved from [https://www.canada.ca/en/publichealth/services/publications/diseases-conditions/hiv-in-canada-surveillance-summary-tables-2014-2015.html?\\_ga=2.98622750.1479042079.1512497634-1113029221.1506106857](https://www.canada.ca/en/publichealth/services/publications/diseases-conditions/hiv-in-canada-surveillance-summary-tables-2014-2015.html?_ga=2.98622750.1479042079.1512497634-1113029221.1506106857)
- Rehm J, Shield KD, Joharchi N, Shuper PA. Alcohol consumption and the intention to engage in unprotected sex: systematic review and meta-analysis of experimental studies. *Addiction (Abingdon, England)*. 2012; 107(1):51–59. <https://doi.org/10.1111/j.1360-0443.2011.03621.x>.
- Reisner SL, Mimiaga MJ, Skeer M, Bright D, Cranston K, Isenberg D, ... Mayer KH. Clinically Significant Depressive Symptoms as a Risk Factor for HIV Infection Among Black MSM in Massachusetts. *AIDS and Behavior*. 2009; 13(4):798–810. <https://doi.org/10.1007/s10461-009-9571-9>. [PubMed: 19462228]
- Safren SA, Blashill AJ, O’Cleirigh CM. Promoting the sexual health of MSM in the context of comorbid mental health problems. *AIDS and Behavior*. 2011; 15(Suppl 1):S30–34. <https://doi.org/10.1007/s10461-011-9898-x>. [PubMed: 21331799]
- Sander PM, Cole SR, Stall RD, Jacobson LP, Eron JJ, Napravnik S, ... Ostrow DG. Joint effects of alcohol consumption and high-risk sexual behavior on HIV seroconversion among men who have sex with men. *AIDS (London, England)*. 2013; 27(5):815–823. <https://doi.org/10.1097/QAD.0b013e32835cff4b>.
- Sandfort TGM, Knox JR, Alcalá C, El-Bassel N, Kuo I, Smith LR. Substance Use and HIV Risk Among Men Who Have Sex With Men in Africa: A Systematic Review. *Journal of Acquired Immune Deficiency Syndromes (1999)*. 2017; 76(2):e34–e46. <https://doi.org/10.1097/QAI.0000000000001462>. [PubMed: 28903126]
- SAS. Version 9.4. Cary, NC, USA: SAS Institute Inc; n.d.
- Saunders JB, Aasland OG, Babor TF, de la Fuente JR, Grant M. Development of the Alcohol Use Disorders Identification Test (AUDIT): WHO Collaborative Project on Early Detection of Persons with Harmful Alcohol Consumption--II. *Addiction (Abingdon, England)*. 1993; 88(6):791–804.
- Scott-Sheldon LAJ, Carey KB, Cunningham K, Johnson BT, Carey MP. Alcohol Use Predicts Sexual Decision-Making: A Systematic Review and Meta-Analysis of the Experimental Literature. *AIDS and Behavior*. 2016; 20(0 1):19–39. <https://doi.org/10.1007/s10461-015-1108-9>.

- Shabani S, Dehghani M, Hedayati M, Rezaei O. Relationship of serum serotonin and salivary cortisol with sensation seeking. *International Journal of Psychophysiology: Official Journal of the International Organization of Psychophysiology*. 2011; 81(3):225–229. <https://doi.org/10.1016/j.ijpsycho.2011.06.015>. [PubMed: 21854814]
- Singer M, Bulled N, Ostrach B, Mendenhall E. Syndemics and the biosocial conception of health. *The Lancet*. 2017; 389(10072):941–950. [https://doi.org/10.1016/S0140-6736\(17\)30003-X](https://doi.org/10.1016/S0140-6736(17)30003-X).
- Snaith RP. The Hospital Anxiety And Depression Scale. *Health and Quality of Life Outcomes*. 2003; 1:29. <https://doi.org/10.1186/1477-7525-1-29>. [PubMed: 12914662]
- Stall R, Coulter RWS, Friedman MR, Plankey MW. Commentary on “Syndemics of psychosocial problems and HIV risk: A systematic review of empirical tests of the disease interaction concept” by A. Tsai and B. Burns. *Social Science & Medicine*. 2015; 145:129–131. <https://doi.org/10.1016/j.socscimed.2015.07.016>. [PubMed: 26254086]
- Stall R, Mills TC, Williamson J, Hart T, Greenwood G, Paul J, ... Catania JA. Association of Co-Occurring Psychosocial Health Problems and Increased Vulnerability to HIV/AIDS Among Urban Men Who Have Sex With Men. *American Journal of Public Health*. 2003; 93(6):939–942. [PubMed: 12773359]
- Sternberg E. A self-fulfilling prophecy: linking belief to behavior. *Annals of the New York Academy of Sciences*. 2011; 1234:98–99. <https://doi.org/10.1111/j.1749-6632.2011.06190.x>. [PubMed: 21988254]
- Vagenas P, Ludford KT, Gonzales P, Peinado J, Cabezas C, Gonzales F, ... Altice FL. Being Unaware of Being HIV-Infected is Associated with Alcohol Use Disorders and High Risk Sexual Behaviors Among Men Who Have Sex with Men in Peru. *AIDS and Behavior*. 2014; 18(1) <https://doi.org/10.1007/s10461-013-0504-2>.
- Voltz, E., Wejnert, C., Cameron, C., Spiller, M., Barash, V., Degani, I., Heckathorn, D. Respondent-Driven Sampling Analysis Tool (RDSAT) (Version 7.1). Ithaca, New York: Cornell University; 2012.
- Vosburgh HW, Mansergh G, Sullivan PS, Purcell DW. A review of the literature on event-level substance use and sexual risk behavior among men who have sex with men. *AIDS and Behavior*. 2012; 16(6):1394–1410. <https://doi.org/10.1007/s10461-011-0131-8>. [PubMed: 22323004]
- Wasserman, T., Wasserman, LD. *Depathologizing Psychopathology*. Springer; Cham: 2016. *Pathologizing Everyday Life*; p. 7-12. [https://doi.org/10.1007/978-3-319-30910-1\\_2](https://doi.org/10.1007/978-3-319-30910-1_2)
- Whelan-Goodinson R, Ponsford J, Schönberger M. Validity of the Hospital Anxiety and Depression Scale to assess depression and anxiety following traumatic brain injury as compared with the Structured Clinical Interview for DSM-IV. *Journal of Affective Disorders*. 2009; 114(1–3):94–102. <https://doi.org/10.1016/j.jad.2008.06.007>. [PubMed: 18656266]
- Wilson PA, Nanin J, Amesty S, Wallace S, Cherenack EM, Fullilove R. Using Syndemic Theory to Understand Vulnerability to HIV Infection among Black and Latino Men in New York City. *Journal of Urban Health : Bulletin of the New York Academy of Medicine*. 2014; 91(5):983–998. <https://doi.org/10.1007/s11524-014-9895-2>. [PubMed: 25155096]
- Wim VB, Christiana N, Marie L. Syndemic and other risk factors for unprotected anal intercourse among an online sample of Belgian HIV negative men who have sex with men. *AIDS and Behavior*. 2014; 18(1):50–58. <https://doi.org/10.1007/s10461-013-0516-y>. [PubMed: 23681697]
- Zigmond AS, Snaith RP. The hospital anxiety and depression scale. *Acta Psychiatrica Scandinavica*. 1983; 67(6):361–370. [PubMed: 6880820]

### Highlights

- Increased odds for high-risk sex was seen only at the highest levels of concurrent polysubstance use and depressive symptoms.
- About one-fifth (18.2%) of the effect of depressive symptoms on condomless anal sex with serodiscordant or unknown status partners was explained by polysubstance use – highlighting a significant need for integrated healthcare services for those experiencing concurrent substance use and mental health conditions.
- Neither Alcohol Use, nor Alcohol Use Disorder Identification Test Scores were associated with condomless anal sex with serodiscordant or unknown status partners, despite being previously associated with condomless anal sex.



**Figure 1.** Crude and Mediated Effects (Odds Ratios) Examining the Effects of Depression and Polysubstance use on Condomless Anal Sex with Serodiscordant or Unknown Status Partners.  
\*  $p < 0.05$ , \*\*  $p < 0.01$  \*\*\*  $p < 0.001$

**Table 1**

Sample Characteristics, Stratified by Engagement in High-Risk Sex

Variable	Full Sample (n = 703)		High-risk sex <sup>A</sup>				Bivariate Association	
	N	RDS%	No (n = 441)		Yes (n = 262)			p-value
Age <sup>B</sup>	33	25.47	31	25.44	37	27.49		0.0039
Sexual Identity								
Gay	597	80.1	368	62.5	229	37.5		0.0990
Bisexual	65	15.8	43	71.2	22	28.8		
Other	41	4.1	30	75.9	11	24.1		
Race/Ethnicity								
White	532	68.7	330	67.1	202	32.9		< 0.0001
Asian	67	9.1	49	71.8	18	28.2		
Indigenous	48	10.0	31	48.8	17	51.2		
Latin American	30	7.4	13	35.4	17	64.6		
Other	26	4.8	18	82.6	8	17.4		
Annual Income (CAD)								
<30,000	445	74.0	300	66.9	145	33.1		0.0236
30,000	258	26.0	141	57.8	117	42.2		
HIV Test Result								
Negative	508	77.8	334	68.1	174	31.9		0.0004
Positive	195	22.2	107	53.7	88	46.3		
HADS Depression <sup>C</sup>								
Normal/Borderline	657	93.6	418	64.5	239	35.5		0.0716
Abnormal	40	6.4	18	50.4	22	49.6		
HADS Anxiety <sup>C</sup>								
Normal/Borderline	519	70.9	343	65.1	176	34.9		0.2074
Abnormal	178	29.1	93	60.0	85	40.0		
Alcohol Use <sup>A</sup>								

Author Manuscript

Author Manuscript

Author Manuscript

Author Manuscript

Variable	Full Sample (n = 703)		High-risk sex <sup>A</sup>		Bivariate Association p-value		
	N	RDS%	No (n = 441)	Yes (n = 262)			
No	104	17.3	66	65.2	38	34.8	0.8258
Yes	599	82.7	375	64.1	224	35.9	
AUDIT Zone <sup>D</sup>							
Low/Medium Risk	602	86.0	387	65.1	215	34.9	0.2418
Harmful/Possible Dependence	97	14.0	51	59.0	46	41.0	
Polysubstance Use <sup>A</sup>							
0	184	24.5	130	70.9	54	29.1	0.0002
1-2	303	43.6	200	68.0	103	32.0	
3+	216	31.9	111	53.1	105	46.9	

<sup>A</sup>In the Past Six Months;

<sup>B</sup>(Median, Q1, Q3);

<sup>C</sup>Hospital Anxiety and Depression Scale

<sup>D</sup>Alcohol Use Disorder Identification Test

**Table 2**

Multivariable Adjusted Results for Associations with High-risk Sex

<b>Variable</b>	<b>AOR (95% CI)</b>
<b>Age</b>	<b>1.02 (1.01, 1.04)</b>
<b>Sexual Identity</b>	
Gay	Not selected
Bisexual	
Other	
<b>Race/Ethnicity</b>	
White	Ref
Asian	1.25 (0.67, 2.32)
Indigenous	<b>2.15 (1.18, 3.94)</b>
Latin American	<b>4.03 (2.02, 8.01)</b>
Other	0.48 (0.18, 1.28)
<b>Annual Income (CAD)</b>	
<\$30,000	Ref
\$30,000	<b>1.63 (1.10, 2.40)</b>
<b>HIV Test Result</b>	
Negative	Not selected
Positive	
<b>Number of Male Anal Sex Partners <sup>A</sup></b>	<b>1.05 (1.03, 1.07)</b>
<b>AUDIT</b>	
Low/Medium Risk	Not Selected
Harmful/Possible Dependence	
<b>Poly Substance Use</b>	
HADS Depression: Normal/Borderline	
Polysubstance Use: 0	Ref
Polysubstance Use: 1–2	1.15 (0.74, 1.80)
Polysubstance Use: 3+	<b>1.65 (1.01, 2.68)</b>
HADS Depression: Abnormal	
Polysubstance Use: 0	Ref
Polysubstance Use: 1–2	3.16 (0.26, 39.06)
Polysubstance Use: 3+	<b>20.76 (1.66, 260.40)</b>

<sup>A</sup>Past Six Months; HADS = Hospital Anxiety and Depression Scale; AUDIT = Alcohol Use Disorder Identification Test.