Substance use and depressive and anxiety symptoms among out-of-school adolescent girls and young women in Cape Town, South Africa

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Background. There is a high prevalence of substance use among youth in South Africa (SA), and adolescent girls and young women (AGYW) experience high rates of depression and anxiety. Substance use behaviours and mental health are associated with other public health problems among AGYW such as HIV and unintended pregnancy. Therefore, understanding the relationship between substance use and mental health is imperative to improve AGYW's health.

Objectives. To examine the association between heavy drinking, marijuana, methamphetamine and methaqualone (Mandrax) use and depressive and anxiety symptoms among AGYW aged 16 - 19 years who have dropped out of school in Cape Town, SA.

Methods. Data for this report come from the baseline data of 500 participants of an ongoing cluster-randomised trial assessing the efficacy of a young woman-focused intervention to reduce substance use and HIV risk. After AGYW consented/assented to participate, they completed a urine drug screen and a baseline questionnaire.

Results. Logistic and negative binomial regressions, controlling for clustering at the neighbourhood level, revealed that frequency of depressive symptoms was significantly and positively related to a positive drug screen for Mandrax (β =0.07; p=0.03). All other associations between the frequency of depressive symptoms and substance use were not statistically significant (ps>0.05). The associations between frequency of anxiety symptoms and substance use were not statistically significant (ps>0.05).

Conclusions. Our findings highlight the need to address substance use, especially Mandrax use and its associated risk, and depression in an integrated, youth-friendly setting.

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South Africa (SA) is currently experiencing a high prevalence of substance use among young people.[1-3] Almost 50% of secondary learners or high school students report lifetime alcohol use, with 13% reporting their first drink before age 13. Approximately 13% of learners report lifetime use of marijuana (dagga), and 10% report using illicit drugs such as methamphetamine (tik) or methaqualone (Mandrax).[1] This high prevalence is concerning because substance use is associated with myriad adverse mental health outcomes among SA young people.[4-6]

The association between substance use and depression and anxiety disorders among adolescents and young adults in SA and other countries has been well documented.^[5,7] However, little of this research has focused specifically on adolescent girls and young women (AGYW) in SA. Almost 62% of AGYW in SA experience depression,[8] and 11% of youth experience generalised anxiety disorder. [9] Furthermore, SA women are more likely to have a lifetime prevalence of depression or anxiety than men.[10] AGYW are an important population to consider because their substance use behaviours and mental health are intricately tied to other public health problems facing this population, such as HIV[11] and unintended pregnancy.[12] For example, methamphetamine use[13,14] and mental health problems[15] are associated with condomless sex among AGYW. Therefore, to fully curtail public health issues such as HIV and unintended pregnancies, substance use and mental health among AGYW must be addressed.

Objectives

Given the prevalence of depression and anxiety among AGYW in SA,[8,16] the high prevalence of alcohol and other substance use among youth, [16] and the impact of substance use and mental health on HIV,[17] understanding the relationship between substance use and mental health is imperative to improve the physical and psychological wellbeing among this population. The objective of the current study is to examine the association between heavy drinking, marijuana, methamphetamine and Mandrax (which is often mixed with cannabis) use, and depressive and anxiety symptoms among AGYW aged 16 - 19 years old who have dropped out of school in Cape Town, SA. These are commonly used substances among young people in SA.[4,16] Based on similar research with other populations of people who use substances, we hypothesise that AGYW who report engaging in heavy drinking, marijuana use and methamphetamine use will report more frequent anxiety and depressive symptoms. [4,16,18] Furthermore, given the relative dearth of research regarding the association between Mandrax use and mental health symptoms, we will explore this association to better understand how this specific substance may be associated with depressive and anxiety symptoms.

Methods

Design

Data for this report come from the baseline data provided by participants of a cluster-randomised trial assessing the efficacy of a young woman-focused intervention to reduce substance use and HIV risk.

Setting

Participants were recruited from 24 community clusters across Cape Town, SA. Twelve communities were comprised predominately of those who self-identify as black African, and 12 were comprised predominately of those who self-identify as coloured (racial categories created during the Apartheid era in SA).

Participants and recruitment

Female adolescent peer role models were identified through the Western Cape Province's Department of Social Development's Expanded Public Works Programme (EPWP). The EPWP is a government programme that employs individuals from economically disadvantaged communities in a range of public and community services. EPWP beneficiaries, in combination with trained outreach workers, recruited participants using established community-based outreach methods that have been successful in other studies engaging young women.[19] The EPWP beneficiaries marketed the study by describing the study and the non-monetary benefits (e.g. HIV testing) of study participation, and helped to identify potential participants, but did not screen. Trained study outreach workers conducted the screening with AGYW individually, ensuring privacy and confidentiality. Study staff were multilingual in English, Afrikaans and/or isiXhosa. Eligibility criteria were the following: (i) aged between 16 and 19 years; (ii) self-identified as a woman; (iii) reported consuming 2 - 3 alcoholic drinks at least once in the past 30 days, or using illicit drugs at least once a week in the past 30 days; (iv) currently dropped out of school; (v) had been out of school for at least 6 months; (vi) had not completed matriculation (high school) or did not have an N3 (National Technical Certificate) certificate; (vii) reported condomless sex with a male partner in the past 90 days; (viii) reported living in one of the 24 target disadvantaged communities; (ix) had lived in the target community for at least 6 months; (x) planned to reside in the target community for the next year and could provide verifiable locator information; and (xi) provided verbal and written consent/ assent. A short cognitive test was given at the end of the eligibility screen, [20] and individuals were excluded if they had a score that indicated cognitive impairment. A total of 500 young women were enrolled in the study.

Procedures

If individuals were eligible and interested in participating in the study, they were scheduled for an intake appointment. During the intake appointment, young women were rescreened to ensure eligibility and given detailed information about the study through the consent/ assent process. Those who were interested in participating in the study and were ≥18 years of age provided consent. Those who were <18 years of age provided assent and were required to obtain written permission to participate by a trusted woman who was ≥25 years of age. Young women could identify this trusted woman (e.g. a sister) or

could agree to be introduced to a trusted female community member who had been identified by the study's staff. These trusted women consented for those <18 years of age in loco parentis (in place of the parent). This approach was used because if parents were to learn about a participant's substance use and sexual activity, it could result in conflict or violence in the family. The in loco parentis process has been used successfully in previous SA research with adolescents. [19]

Once they consented/assented to participate, participants underwent a breathalyser test for recent alcohol use and provided locator information (including having a photograph taken to identify the participant) to be used to contact them for follow-up. Participants then completed the baseline questionnaire via computer-assisted personal-interviewing technology, and audio computer-assisted selfinterviewing in the language of their choice (English, Afrikaans or isiXhosa) for sensitive topics, such as experiences of abuse. Next, participants provided urine samples and staff conducted pregnancy testing, screening for recent drug use (e.g. marijuana, opioids, Mandrax, cocaine, methamphetamine) using a rapid urine drug screen, conducted pre- and post-HIV counselling and rapid testing via finger prick, and provided participants with the test results. This appointment lasted approximately 2 hours, and participants were remunerated ZAR150 (~USD10.00) for their time.

Participants residing in communities that were randomised to the intervention arm were scheduled to participate in two group workshops focused on substance use and sexual risk reduction over the next ~2 weeks post enrollment. All participants were scheduled to return 6 and 12 months post intake for follow-up assessments. This article only reports data from the baseline appointment.

The SA co-investigator, SA co-project director, and the US principal investigator of this study are all psychologists who have extensive experience assisting with distressed participants (e.g. experiencing suicidal ideation). In addition, all staff were trained on a standard protocol to assist distressed participants, which included consulting with senior staff. All procedures were approved by ethics review boards at RTI International (ref. no. 0214954) and the SA Medical Research Council (ref. no. EC006-3/2016).

Main outcome measures

The Revised Risk Behavior Assessment^[21] tool was used to collect behavioural data. This instrument includes measures of sexual risk, mental health symptoms and substance use behaviours, and was modified for use in SA and combined with instrumentation from the US National Institute on Drug Abuse (NIDA) data harmonisation effort.[22]

Substance use

Participants' heavy alcohol use was defined by the extent to which participants reported the consumption of four or more alcoholic drinks a day. Heavy alcohol use was measured by assessing the number of days that participants had consumed four or more drinks in the past 30 days. Recent drug use was measured using a urine drug screen to detect metabolites of recent use of cocaine, methamphetamines, amphetamines, opioids, ecstasy and marijuana. Mandrax was measured through a separate urinalysis test. Participants also selfreported their lifetime and past-month use (use/not used; frequency of use in the past month) of marijuana, methamphetamines, heroin, cocaine, inhalants, ecstasy, other amphetamines, non-medical use of prescription medications and other drugs. Based on the results of the urine drug screen, participants were coded as 1 = positive or 0 = positivenegative for each drug tested. This variable was used as the outcome of the main analyses.

Depressive and anxiety symptoms

The frequency of participants' depressive and anxiety symptoms was assessed using the Drug Abuse Treatment for AIDS Risks Reduction depression scale (a 7-item scale; $\alpha = 0.70$) and anxiety scale (a 7-item scale; $\alpha = 0.79$). [23] Responses indicated the extent to which participants had experienced each symptom in the past 90 days and were on a Likert scale (0 = never to 4 = almost always). Higher scores indicated more frequent depressive or anxiety symptoms. This scale has demonstrated satisfactory reliability in other samples of women in SA.[24,25]

Covariates

Participants' engagement in sex work was assessed using the question, 'Have you ever traded sex to get the things you need (like drugs, money, food, clothing, shelter, transport, or any other goods)? This would include sex work.' Participants responded 1 = yes or 0 = no. Ethnicity was assessed using the question 'Do you consider yourself to be black African, coloured (i.e. mixed ancestry), Asian/Indian, white South African or another race or ethnic group?' from the screener. These racial categories represent classifications that were developed during the Apartheid era in SA.[26] Given the ethnic make-up of the targeted communities, all participants identified as either black African (coded as 0) or coloured (coded as 1). HIV status was assessed using a minimum of two rapid HIV tests (via finger prick). If the two results were concordant positive or negative, then no additional testing was done. If they were discordant, a third confirmatory test was done. Participants were classified as HIVpositive if two tests were positive. Results were coded as 0 = HIVnegative or 1 = HIV-positive.

Analysis

We used Stata version 15 (StataCorp, USA) for all the analyses. Descriptive analyses were performed and the associations among the outcomes of interest (heavy drinking, marijuana use, methamphetamine use and Mandrax use) as well as conceptually meaningful covariates (e.g. engagement in sex work, HIV status and ethnicity) were explored first. Next, separate logistic regression models were conducted to explore the association between biologically confirmed marijuana, methamphetamine and Mandrax use and the frequency of depressive and anxiety symptoms. Then, separate negative binomial regression models were conducted to examine the association between frequency of depressive and anxiety symptoms and days of heavy drinking. All analyses controlled for clustering at the neighbourhood level using a robust cluster estimator. P-values <0.05 and confidence intervals that did not include 1 indicated significance.

Results

Demographics

Characteristics of the sample are shown in Table 1. The mean (standard deviation (SD)) age of the participants was 17.8 (1.1). Approximately half of the participants identified as coloured and the other half identified as black African. Of note, almost half of the AGYW in the study reported that their household ran out of necessities at least monthly. Most participants tested positive for marijuana (38.6%) during the drug screen. Participants' mean (SD) scores regarding the frequency of depressive and anxiety symptoms were 9.6 (4.6) and 7.8 (5.3), respectively. These scores indicate that on average, participants experienced depressive or anxiety symptoms

Variable	n (%)*
Age, mean (SD; range 16 - 19)	17.76 (1.05)
Ethnicity	
Black African	246 (49.20)
Coloured	254 (50.80)
Unemployed	458 (91.60)
Highest level of education completed	
Standard 6 (Grade 8) or less	151 (30.20)
Standard 7 to 9 (Grade 9 to 11)	349 (69.80)
Age when first left school, mean (SD)	16.00 (1.31)
Would like to study further	420 (84.34)
Current relationship status	
Single, never married and not currently involved with a boyfriend	103 (20.68)
Have a boyfriend (or someone regularly have sex with)	384 (77.11)
Married	6 (1.20)
Separated	5 (1.00)
Live in own apartment/flat	81 (16.20)
Running out of money for necessities at least monthly	245 (49.00)
Ever engaged in sex work	27 (5.42)
Biological confirmed HIV-positive status	31 (6.20)
Ever drank alcohol	498 (99.60)
Days of heavy drinking in the past month, mean (SD)	3.93 (4.57)
Frequency of depressive symptoms (possible range 0 - 28), mean (SD)	9.59 (4.61)
Frequency of anxiety symptoms (possible range 0 - 28), mean (SD)	7.78 (5.31)
Lifetime physical abuse	78 (15.63)
Lifetime sexual abuse	47 (9.42)

slightly more than rarely. Alcohol (99.6%) and marijuana (81.4%) were the most commonly self-reported substances used during participants' lifetime. Table 2 displays information about the selfreported drug use of the sample.

Primary analysis

As shown in Table 3, negative binomial regression, controlling for neighbourhood cluster, revealed that the frequency of depressive symptoms was related to an increase in the number of days of self-reported heavy drinking in the past 30 days (β =0.02, p=0.07); however, this trend was not significant. Logistic regression revealed that self-reported frequency of depressive symptoms in the past 90 days was not significantly related to a positive drug screen for marijuana (p=0.69) or methamphetamine (p=0.13). However, frequency of depressive symptoms was significantly related to a positive drug screen for Mandrax, such that an increase in the frequency of depressive symptoms was related to a greater likelihood of testing positive for Mandrax (β=0.07; p=0.03). Frequency of anxiety symptoms was not significantly related to the number of heavy drinking days in the past month, use of marijuana, use of methamphetamine or use of Mandrax (ps>0.05).

Discussion

Globally, substance use and mental health disorders are public health epidemics.^[27] SA, in particular, has a considerable disease burden attributable to substance use and mental health disorders, [28] and a substantial number of those affected are youth.^[4,9,16] Moreover, AGYW are more likely to suffer from mental health disorders such

as depression and anxiety. However, little research has specifically examined the association between substance use and depression and anxiety among AGYW in SA. This is problematic because research suggests that the age of onset for many substance use and mental health disorders occurs before the age of 24, and those who are most vulnerable often experience these problems simultaneously.^[29,30] This study fills this gap by examining the correlates of depressive and anxiety symptoms among vulnerable AGYW in SA.

Statement of principal findings

Findings from the present study suggest that the frequency of depressive symptoms may be positively related to the use of Mandrax. Depressive symptoms and heavy alcohol use were also positively related, although this association was not significant. This study did not find significant associations between the frequency of depressive symptoms and marijuana and methamphetamine use. Similarly, there were no statistically significant associations between frequency of anxiety symptoms and the use of any substance examined.

Mandrax use as an illicit drug is unique to SA.[31,32] Some research has linked Mandrax to sexual risk.[33] Still, the association with mental health and Mandrax use has not been thoroughly examined. Therefore, our finding highlighting the association between depressive symptoms and Mandrax use is novel. This study illustrates the need for and potential benefit of further examination of the link between depressive symptoms and Mandrax use - and related outcomes such as sexual risk - among AGYW in SA.

This study did not demonstrate a link between the frequency of depressive symptoms and heavy alcohol use, marijuana use or

	Lifetime drug use,	Age of first use,	Drug use in the past month,	Times used drugs in the past month,
Substance	n (%)	mean (SD)	n (%)	mean (SD)
Marijuana	407 (81.40)	15.04 (1.79)	284 (57.37)	19.32 (38.85)
Methamphetamine (tik)	72 (14.40)	16.44 (1.51)	51 (10.20)	3.22 (19.66)
Cocaine	16 (3.20)	17.38 (1.31)	3 (0.60)	0.02 (0.45)
Heroin	3 (0.60)	14.67 (0.58)	1 (0.20)	0.08 (1.79)
Mandrax	58 (11.60)	16.41 (1.72)	39 (7.80)	1.54 (10.00)
Inhalant	12 (2.40)	14.50 (1.83)	2 (0.40)	0.01 (0.18)
Ecstasy	57 (11.40)	16.60 (1.49)	16 (3.20)	0.10 (0.91)
Other amphetamines	1 (0.20)	18.00 (-)	0 (-)	0 (-)
Prescription drugs	9 (1.80)	16.11 (1.69)	3 (0.60)	0.16 (2.45)

Table 3. Frequency of depression and anxiety symptoms predicting heavy alcohol use and biologically confirmed drug use, controlling for neighbourhood cluster and other covariates (N=500)

	Heavy drinking in				
Value	the past 30 days	Marijuana (dagga)	Methamphetamine (tik)	Mandrax	
Depressive symptoms in the past 90 days					
β (SE)	0.02 (0.01)	0.01 (0.03)	0.05 (0.03)	0.07 (0.03)	
95% CI	-0.0015 - 0.0412	-0.0397 - 0.0600	-0.0133 - 0.1069	0.0064 - 0.1245	
<i>p</i> -value	0.07	0.69	0.13	0.03*	
Anxiety symptoms in the past 90 days					
β (SE)	0.02 (0.01)	-0.004 (0.02)	0.03 (0.03)	0.003 (0.04)	
95% CI	-0.0038 - 0.0354	-0.0442 - 0.0369	-0.0284 - 0.0853	-0.0704 - 0.0761	
<i>p</i> -value	0.12	0.86	0.33	0.94	

SE = standard error; CI = confidence interval. *p<0.05. Based on bivariate correlations, models predicting heavy drinking in the past 30 days included race as a covariate. Models predicting a positive marijuana drug screen included race as a covariate. Models predicting a positive Mandrax drug screen included marijuana and methamphetamine use as covariates.

methamphetamine use, nor between anxiety symptoms and the substance use outcomes of interest. While these findings did not support our hypotheses, there are plausible explanations for these findings. The assessments of depressive and anxiety symptoms focused on frequency and not necessarily duration, severity, onset or impairment related to the symptoms. While frequency and these other dimensions of symptoms are related, they are not synonymous. Therefore, the association between symptoms of depression and anxiety and substance use may differ based on the dimension (e.g. frequency, severity, duration, impairment) that is assessed. For example, one may frequently experience trouble sleeping (a common symptom of depression and anxiety), but the association between sleep and substance misuse may be accounted for by impairments in functioning such as reductions in emotion regulation - which may be a stronger predictor of substance use than the frequency of the symptoms.^[34] As it relates to anxiety, the association between substance use and symptoms may depend on the particular type of anxiety disorder that one experiences. For example, the link between post-traumatic stress disorder and substance use has been consistently demonstrated among other populations such as adult women, [35] survivors of natural disasters^[36] and survivors of victimisation.^[37] Therefore, the associations may be more nuanced than the current article can examine due to the limitations of secondary data analysis.

Strengths

The present study may be among the first to explore the associations between depressive and anxiety symptoms and heavy drinking, marijuana, methamphetamine and Mandrax use among vulnerable AGYW in Cape Town. The examination of Mandrax use as a correlate of depressive and anxiety symptoms is a particular strength of this paper, given its prevalence among South Africans. [32,33,38] Additionally, our focus on out-of-school AGYW - a hard-to-reach population that has been understudied regarding their mental health - is a strength. Understanding substance use and mental health among this population is important given their role in HIV risk and unintended pregnancy among AGYW in SA.[39] Our sample was also recruited from communities in SA, which strengthens the external validity of the findings. As it relates to measurement, we used assessments that have been utilised and validated in similar samples in SA. Relatedly, recent substance use was assessed using a urine drug screen, which typically detects drug metabolites up to 2 - 30 days after use, depending on the substance of use $^{[40]}$ – which is more accurate than self-report. Collectively, these strengths contribute to the validity of our findings.

The strengths and implications should be interpreted considering study limitations. The data for this study were taken from baseline data of a larger trial examining the efficacy of a young womanfocused HIV prevention intervention. The assessment period for depressive and anxiety symptoms was the past 90 days, and the assessment period for heavy alcohol use was the past 30 days. The urine drug screen has an assessment period of up to 30 days. Given the assessment periods of each outcome, temporality may be suggested, but causality cannot be inferred. For example, it may be reasonable to believe that depressive symptoms in the past 90 days may predict Mandrax use in the past 30 days. However, drug use may also precipitate or exacerbate mental health symptoms. For example, Mandrax use or withdrawal could contribute to anxiety symptoms. Therefore, longitudinal assessments of substance use and mental health symptom assessments of substance use and mental health are necessary for future research to understand the causality of these associations. The assessment of depressive and anxiety symptoms was not diagnostic. Therefore, we are unable to discern if clinically diagnosed depression or anxiety are related to substance use. Future research should examine these associations using more diagnostic tools. Relatedly, future research may benefit from examining how different types of mood and anxiety disorders may be related to the use of different substances. The biological assessment of recent substance use also has some limitations. While rapid urine drug screens are more reliable than self-report, there is the risk of falsepositives due to secondary exposure to marijuana and the use of other medications (e.g. antidepressants, such as may result in false-positive tests for methamphetamine).[41] We did not conduct a comprehensive assessment of the medications that participants were taking, therefore we could not address this risk. Lastly, we did not assess how the composition of Mandrax that AGYW reported using was related to their mental health symptoms. Mandrax is often mixed with cannabis and smoked, and may be used with other substances.[2] Therefore, it will be important for future research to understand how the composition of Mandrax may differentially affect AGYW's mental health symptoms. These limitations notwithstanding, this study adds to the existing literature and serves as a basis to explore the nuanced associations between substance use and mental health among AGYW in Cape Town, SA.

Contributions

This study contributes to the nascent literature on the prevalence and correlates of depressive and anxiety symptoms among vulnerable young people in SA.[4,5,16] Overall, our results demonstrate the link between Mandrax and the frequency of depressive symptoms among AGYW in SA. Recently, this population has received much attention related to the need for HIV prevention because of the high incidence among this group. Also, access to reproductive health services to reduce unintended pregnancy among AGYW is of great concern in SA. Substance use has often been examined in the context of sexual risk behaviour. However, our findings suggest that depressive symptoms may also be important in understanding substance use.

Clinical and policy implications and recommendations

These findings have practical and policy implications. Recently, there has been an increase in the demand for substance use treatment among young people, with 37% of patients being <25 years of age. [42] There have also been efforts by the SA government to increase the provision of youth-friendly services in public health clinics. [39] Our findings highlight the need to address substance use and mental health challenges among AGYW. Given that there is a need for substance use treatment centres to see young people and that public health clinics are providing more youth-friendly services, these may be ideal settings for integrated care to treat substance use and mental health disorders. However, currently, in Cape Town, there are few substance use treatment centres that can treat those under 18. The results of the current study suggest that government stakeholders should consider providing substance use treatment centres with the resources to increase their capacity to treat youth and incorporate mental health treatment into their efforts to meet the demands of an increasingly younger client base. Furthermore, key decision-makers should extend efforts to increase the provision of youth-friendly services to include substance use treatment facilities. Lastly, it may also be important to build the capacity of public health clinics to provide youth-friendly mental health and substance use treatment.

Conclusions

Our findings highlight how depressive symptoms may contribute to Mandrax use among SA AGYW. Efforts seeking to reduce substance use among AGYW may consider addressing mental health challenges, especially given the high rate of depression among this population. Key decision-makers may consider incorporating mental health treatment within substance use treatment centres and equipping these centres with the resources to provide youth-friendly services. Addressing substance use and mental health among AGYW may also help to curtail public health challenges such as HIV and unintended pregnancy.

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Author contributions. All authors participated in the research and article preparation. WMW conceived of the study, oversaw drafts and finalised the manuscript; CPB conducted the analyses and led the development of the manuscript; TC, FAB, JWN and BNH contributed to writing and reviewing the manuscript.

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Conflicts of interest. None.

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