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Sexual orientation and gender identity disparities in substance use disorders during young adulthood in a united states longitudinal cohort\*

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#### Abstract

**Background:** This study examined associations of sexual orientation and gender identity with prevalence of substance use disorders (SUDs) and co-occurring multiple SUDs in the past 12-month during young adulthood in a United States longitudinal cohort.

**Methods:** Questionnaires self-administered in 2010 and 2015 assessed probable past 12-month nicotine dependence, alcohol abuse and dependence, and drug abuse and dependence among 12,428 participants of an ongoing cohort study when they were ages 20-35 years. Binary or multinomial logistic regressions using generalized estimating equations were used to estimate differences by sexual orientation and gender identity in the odds of SUDs and multiple SUDs, stratified by sex assigned at birth.

**Results:** Compared with completely heterosexuals (CH), sexual minority (SM; i.e., mostly heterosexual, bisexual, lesbian/gay) participants were generally more likely to have a SUD, including multiple SUDs. Among participants assigned female at birth, adjusted odds ratios (AORs) for SUDs comparing SMs to CH ranged from 1.61-6.97 (ps<.05); among participants assigned male at birth, AORs ranged from 1.30-3.08, and were statistically significant for 62% of the estimates. Apart from elevated alcohol dependence among gender minority participants assigned male at birth compared with cisgender males (AOR: 2.30; p<.05), gender identity was not associated with prevalence of SUDs.

**Conclusions:** Sexual and gender minority (SGM) young adults disproportionately evidence SUD, as well as co-occurring multiple SUDs. Findings related to gender identity and bisexuals assigned male at birth should be interpreted with caution due to small sample sizes. SUD prevention and treatment efforts should focus on SGM young adults. Keywords: Sexual Orientation; Gender Identity; LGBT; Substance Use Disorders (SUDs);

Young Adults; Longitudinal Cohort

#### **1. Introduction**

Substance use disorders (SUDs) affect more than 20 million individuals in the United States (U.S.) annually, increasing risk for psychiatric disorders, chronic diseases, and disruptions to social, family, and work lives (Patel et al., 2016; Whiteford et al., 2013). SUD prevalence peaks during young adulthood (Center for Behavioral Health Statistics and Quality, 2016), with co-occurrence of multiple SUDs also common during this time period, which increases clinical severity and complicates treatment (Falk et al., 2008; Moss et al., 2015). Previous research has established that, compared to completely heterosexual (CH) and cisgender individuals (i.e., gender identity corresponds with sex assigned at birth), sexual and gender minorities (SGMs; i.e., those identifying as lesbian, gay, bisexual, or transgender [LGBT], with same-sex sexual attractions or behaviors, or with a gender identity different than their birth sex) engage in greater substance use beginning in adolescence and extending throughout life (Buchting et al., 2017; Corliss et al., 2014; Day et al., 2017; De Pedro et al., 2017; De Pedro and Shim-Pelayo, 2018; Gerend et al., 2017; Gonzales et al., 2016; Newcomb et al., 2014; Talley et al., 2014). Despite evidence of SGMs' disproportionate substance use, only a small proportion of studies on substance use have assessed sexual orientation (2.3-6.5%) or gender identity (1.9-2.3%) (Flentje et al., 2015). Even fewer have examined more serious SUD outcomes by sexual orientation or gender identity or have focused on SUDs during young adulthood (Coulter et al., 2018; Goldberg et al., 2013; Kerridge et al., 2017; Medley, 2016). The present study addresses these gaps by examining associations between SGM statuses and past 12-month prevalence of SUDs in a community cohort of U.S. young adults.

SGM disparities in SUDs persist because SGMs use substance to cope with SGM-related minority stressors, including self-stigma and interpersonal and structural-level discrimination

(Felner et al., in press; McCabe et al., 2009a). Disparities may also be driven by differences in substance use norms within SGM communities (Felner et al., in press). For example, research indicates that sexual minorities (SMs) perceive greater availability of substances and have more tolerant use norms than do heterosexuals (Cochran et al., 2012; Mereish et al., 2017). Additionally, gender minority (GM) youth may perceive less risk associated with substance use than cisgender youth (Day et al., 2017).

#### 1.1 Sex Assigned at Birth, Sexual Orientation, Age, and SUD Risk

Research has found persistent variation in SUD risk by sex. In the general population, men experience single and co-occurring SUDs at higher levels than women (Falk et al., 2008). Among SMs, however, sex differences are typically reduced or even reversed, with greater sexual orientation disparities among adult women compared to men, and especially elevated rates among bisexual women (Coulter et al., 2018; Goldberg et al., 2013; Kerridge et al., 2017; McCabe et al., 2018; Medley, 2016). Nonetheless, studies have rarely tested whether sex modifies relationships between sexual orientation and SUDs by including interaction terms in statistical models.

Prevalence of SUDs tends to peak around age 25 and declines with age (Merikangas and McClair, 2012). Research examining SUDs among SMs, however, suggests a slower agenormative decline (Evans-Polce et al., 2019; Hughes et al., 2006; McKirnan and Peterson, 1989). Rarely have researchers compared sexual orientation or gender identity disparities in SUDs among individuals older than 25 years with those in younger age groups. Knowledge of how the magnitude of sexual orientation and gender identity differences in SUDs vary by birth sex and age can help identify subgroups in need of interventions.

#### 1.2 Gender Identity and SUD Risk

Research on how gender identity is associated with SUD risk is severely lacking, with available studies using small, subgroup samples (Flentje et al., 2014; Keuroghlian et al., 2015; Reisner et al., 2016). Studies also frequently lack cisgender comparison groups, preventing quantification of gender identity differences. Although most available research has examined substance use, rather than SUDs, among GMs, there is reason to suspect elevated SUD risk given GMs' 2-4 times higher likelihood of using substances compared to their cisgender peers (Coulter et al., 2015; Day et al., 2017; De Pedro et al., 2017; Gerend et al., 2017; Tupler et al., 2017).

#### 1.3 SGM Differences in Types of Drugs Used Among Those with SUDs

Research indicates that addiction potential, mortality risk, effective treatment, clinical outcomes, and public health impacts vary across drugs used (Magill and Ray, 2009; Nutt, 1996; Volkow et al., 2015). Given these implications, information on how sexual orientation and gender identity is associated with specific drug types can inform efforts to address SUD disparities.

#### 1.4 Study Aims

This study analyzed data from the longitudinal Growing Up Today Study (GUTS) when participants were aged 20-35 to estimate sexual orientation and gender identity differences in probable SUDs. Diagnostic and Statistical Manual of Mental Disorders, Fourth Edition (DSM-IV) criteria were used to assess past 12-month nicotine dependence, alcohol abuse/dependence, drug abuse/dependence, any SUD, and co-occurring multiple SUDs (2 or more). Because research demonstrates sex differences in associations between sexual orientation and substance outcomes (Hughes et al., 2016; Schuler et al., 2018), we estimated statistical interactions between 1) sexual orientation and birth sex, and 2) gender identity and birth sex, and present birth-sex-stratified estimates. We hypothesized that SGMs would be more likely than non-SGMs of their same birth sex to meet criteria for SUDs, and that sexual orientation differences would be larger among participants assigned female at birth compared to those assigned male. Additionally, we estimated statistical interactions between 1) sexual orientation and age, and 2) gender identity and age. We hypothesized that sexual orientation and gender identity differences in SUD risk would be larger in older (26-35 years) versus younger (20-25 years) periods. Among participants meeting criteria for a past 12-month drug use disorder, we examined associations of sexual orientation and gender identity with past 12-month specific drug use.

#### 2. Methods

#### 2.1 Participants and Procedures

Data are from two ongoing GUTS cohorts: GUTS1 and GUTS2. In 1996, GUTS1 participants (N=16,882) were aged 9-14 at baseline. In 2004, GUTS2 participants (N=10,923) were aged 10-15 at baseline. GUTS participants are children of Nurses' Health Study II (NHSII) participants. For GUTS1, 34,174 NHSII participants with eligible children were contacted to request permission to invite their children to join the cohort. For GUTS2, 20,700 NHSII participants with eligible children were similarly contacted. NHSII participants provided information on 26,765 (GUTS1) and 17,280 (GUTS2) eligible children, who were mailed an invitation to participate and a sex-specific questionnaire. More information about GUTS is available elsewhere (Field et al., 1999; Field et al., 2014). Data collection procedures (self-administered paper or web-based questionnaires) were approved by Partners Healthcare IRB.

The current analysis includes 17,496 observations from 8,701 GUTS1 and 3,727 GUTS2 participants of: (1) wave 2010 (GUTS1 only; 51% response rate) and (2) a 2015-2017 GUTS Substance Substudy (GUTS1 and 2; 73% response rate among 13,340 recent responders invited to participate). The proportion of participants included in the current analysis was lower among

those assigned male (32.4%) compared to those assigned female (55.2%; p<.0001) at birth due to greater attrition among males. Participants from the Midwest (44.8%), South (43.0%), and Northeast (43.6%) regions of the U.S. were less likely to be included in the analysis than participants from the West (48.9%; p<.0001). Additionally, included participants were younger at baseline than those excluded (mean age 12.1 vs. 12.2 years; p<.0001). No difference by race/ethnicity was observed (p=.37).

#### 2.2 Measures

*Sexual orientation*. Since 1999, GUTS questionnaires have consistently measured sexual orientation with a question from the Minnesota Adolescent Health Survey (Remafedi et al., 1992) that concurrently taps two dimensions (attraction and identity): "Which one of the following best describes your feelings?", with response options: 1) Completely heterosexual (attracted to persons of the opposite sex), 2) mostly heterosexual, 3) bisexual (equally attracted to men and women), 4) mostly homosexual, 5) completely homosexual (gay/lesbian, attracted to persons of the same sex), or 6) not sure. We included the following categories in analysis: completely heterosexual (CH), mostly heterosexual (MH), bisexual (BI), and lesbian/gay (LG) (mostly and completely homosexual combined). "Not sure" or missing information on sexual orientation were excluded from analysis due to small numbers (n=5). We modeled participants' sexual orientation response as time-varying and corresponding with the timing of their SUD assessment to account for potential changes in reports between 2010 and 2015.

*Gender identity*. At baseline, GUTS gathered information on birth sex. In 2010, 2014, and 2016, GUTS assessed participants' gender identity: "How do you describe yourself?", and response options: (1) Male, (2) Female, (3) Transgender, or (4) "None of the above" (2010) or "Do not identify as male, female or transgender" (2014, 2016). We classified participants as

GMs if they selected a response of transgender or none of the above, or if there was discordance between their gender identity and birth sex.

*Substance use disorders*. Past 12-month nicotine dependence, alcohol abuse/dependence, and drug abuse/dependence were assessed in 2010 and 2015-2017 with questions adapted from the National Survey on Drug Use and Health (NSDUH) corresponding to DSM-IV criteria for SUDs (American Psychiatric Association, 1994). We coded responses as evidencing or not evidencing symptoms of dependence (e.g., tolerance, withdrawal) and abuse (e.g., failure to fulfill major role obligation, physically hazardous), classifying participants as having probable substance dependence if they endorsed 3 or more of 7 dependence symptoms and as having probable abuse if they endorsed at least 1 of 4 abuse symptoms. We then created 4 SUD variables: nicotine dependence (dichotomous, yes versus no), alcohol use disorder (3-categories; none, abuse only, and dependence), drug use disorder (3-categories; none, abuse only, and dependence), drug use disorder (3-categories; no SUD, one SUD, and two or more SUDs).

GUTS questionnaires cover multiple health-related topics. Thus, to reduce participant burden, questions assessing drug use disorder for marijuana and other drugs were combined into a single set of questions. Although less comprehensive than the NSDUH's assessment of each drug separately, our approach is supported by findings that marijuana and other drug use frequently co-occur, and they have similar impacts on well-being (e.g., on school enrollment and employment) (Arria et al., 2013a; Arria et al., 2013b; Fergusson et al., 2008; Mack et al., 2017; Moss et al., 2015).

*Drug use*. Past 12-month use of marijuana, cocaine, heroin, MDMA/ecstasy, LSD/mushrooms, methamphetamine, amphetamines, and nonmedical use of prescription

benzodiazepines, painkillers, sleeping pills, and stimulants were assessed in 2010 and 2015-2017. Inhalants was assessed in 2010.

*Covariates*. Race/ethnicity (non-Hispanic White versus other), region of residence (Northeast, West, South, Midwest), cohort (GUTS1, GUTS2), and age at the time of SUD assessment (dichotomized into 20-25 years versus 26-35 years) were included in analyses as potential confounders.

#### 2.3 Statistical Analyses

Analyses were stratified by birth sex (assigned female, assigned male at birth). Unadjusted prevalences of past 12-month SUD outcomes were examined for each sexual orientation and gender identity subgroup. We estimated multivariable associations of SGM statuses with SUDs using generalized estimating equations with exchangeable correlations structure to account for non-independence of sibling clusters and repeated measures among individuals (Liang and Zeger, 1986). When exchangeable correlation structure did not yield convergence for three models estimating drug type (see Table 5), we used independence correlation structure. For nicotine dependence, binary logistic regression estimated adjusted odd ratios (AOR). For alcohol use disorders, drug use disorders, and co-occurring SUDs, multinomial logistic regression estimated AOR. To test whether birth sex modified relationships between sexual orientation and SUDs and gender identity and SUDs, we included sexual-orientation-bysex and gender-identity-by-sex interaction terms. To test whether age modified relationships between sexual orientation and SUDs and gender identity and SUDs, we included sexualorientation-by-age and gender-identity-by-age interaction terms stratified by birth sex. To estimate sexual orientation and gender identity differences in past 12-month use of specific drugs, we used binary logistic regression. In these analyses, we combined LGB participants into

one category due to small sample sizes. In all models, CH and cisgender participants were referent groups. Corresponding 95% confidence intervals (CI) and p-values were estimated. Multivariable models adjusted for age, race/ethnicity, cohort, region of residence, and birth sex (for models testing for sex-at-birth interactions). All analyses were performed with SAS software, version 9.4, with a significance level of 0.05.

#### 3. Results

#### 3.1 Sociodemographic Characteristics of Participants

Sociodemographic distributions of participants' observations are presented in Table 1. Approximately 19% of observations from participants assigned female at birth were mostly heterosexual, 3% were bisexual, 2% were lesbian, and 0.7% were GMs. Of observations from participants assigned male at birth, approximately 10% were mostly heterosexual, 1% were bisexual, 5% were gay, and 1% were GMs. More than 60% of observations were from participants aged 26-35, while less than 40% from participants aged 20-25.

#### 3.2 Prevalence of Past 12-Month SUDs by Sexual Orientation and Gender Identity

Table 2 presents past 12-month prevalences of SUDs by sexual orientation and gender identity stratified by sex assigned at birth. In most instances, prevalences of all SUD outcomes were higher among SGMs compared with non-SGMs. Comparisons within sexual minorities found one statistically significant difference, with the prevalence of drug dependence higher among bisexual women compared MH women.

#### 3.3 Interactions of Birth Sex with Sexual Orientation and Gender Identity on SUDs

In numerous instances, birth sex modified relationships with sexual orientation, with sexual-orientation differences larger among participants assigned female compared to assigned male at birth. Differences between MHs and CHs were larger among individuals assigned female for nicotine dependence (p=.02), alcohol abuse (p=.03), alcohol dependence (p=.02), drug dependence (p=.01), one SUD (p=.001), and multiple co-occurring SUDs (p=.01). Although differences between bisexuals and CHs were generally larger among individuals assigned female than assigned male, statistical significance was observed only for drug dependence (p=.006) and marginally for multiple co-occurring SUDs (p=.068), likely due to low power resulting from the small number of bisexual males. Differences between lesbian and gay participants compared to CHs were also larger among individuals assigned female than assigned male for nicotine dependence (p=.002), drug abuse (p=.04), drug dependence (p=.02), and multiple co-occurring SUDs (p=.003). Gender-identity-by-birth-sex interactions were not significant (ps>.05). For more information, see Appendix Table 1.

#### 3.4 Multivariable Associations of Sexual Orientation and Gender Identity with SUDs

Table 3 presents the multivariable associations of sexual orientation, gender identity, and other covariates with SUDs among participants assigned female at birth. The odds of evidencing each SUD and co-occurring multiple SUDs were greater among all SM groups compared to CHs. Associations between gender identity and SUDs were not statistically significant.

Table 4 presents the multivariable associations of sexual orientation, gender identity, and other covariates with SUDs among participants assigned male at birth. All SM groups had elevated odds for nicotine dependence and one SUD compared to CHs. MHs also had elevated odds of alcohol dependence, drug abuse and dependence, and having 2 or more SUDs. Gay men also evidenced elevated odds for alcohol abuse and dependence, drug dependence, and having 2 or more SUDs compared to CHs. GMs had significantly higher odds of alcohol dependence than their cisgender peers. Like patterns observed among participants assigned female, associations

between gender identity and SUDs were frequently smaller than associations of sexual orientation with SUDs.

#### 3.5 Interactions of Age with Sexual Orientation and Gender Identity

Statistical interactions between sexual-orientation-by-age and gender-identity-by-age were non-significant (all ps>.05), except for drug abuse among bisexual females, where differences between bisexuals and CHs were larger during ages 26-35 compared to 20-25 (p=.04). In most instances, prevalences of SUD outcomes across sexual orientation and gender identity groups were lower in the older versus younger age group.

# 3.6 Multivariable Associations of Sexual Orientation and Gender Identity with Types of Drugs Used

Table 5 presents the prevalence and multivariable associations of sexual orientation and gender identity with past 12-month drug use among participants evidencing a drug use disorder, stratified by birth sex. Regardless of sexual orientation and gender identity, marijuana was the most prevalent drug reported. Among participants assigned female, LGBs were more likely than CHs to report using MDMA/ecstasy and LSD/mushrooms and GMs were more likely than their cisgender counterparts to report using LSD/mushrooms. Among those assigned male, LGBs were more likely than CHs to report using methamphetamine and inhalant and GMs were more likely than cisgender peers to report using heroin, amphetamines, inhalants, and non-medical use of prescription painkillers. No differences were found between MH and CH in the prevalence of types of drugs used.

#### 4. Discussion

#### 4.1 Summary of Main Findings

Our study quantified sexual orientation and gender identity differences in SUD risk during young adulthood, when SUD prevalence in the general U.S. population is high (Center for Behavioral Health Statistics and Quality, 2016). We examined SUDs based on DSM-IV criteria including nicotine dependence, alcohol abuse and dependence, drug abuse and dependence, and multiple co-occurring SUDs. Aligning with previous literature (Goldberg et al., 2013; Kerridge et al., 2017; McCabe et al., 2009b), we found that SM status was associated with greater odds of past 12-month SUDs among young adults assigned female, and to a lesser extent among those assigned male. Co-occurrence of 2 or more SUDs in the past 12-months was also more common among SMs compared CHs, aligning with previous studies of lifetime SUD co-occurrence (Lee et al., 2015; Medley, 2016; Mereish et al., 2015).

Contrary to our hypothesis, age-related declines in SUD prevalence were largely similar across sexual orientation and gender identity groups. This finding may be due, in part, to our sample age range (20-35 years) and age periods compared in analysis (20-25 versus 26-35). Previous studies have shown differential age-related declines in alcohol problems between SMs and heterosexuals and noted the largest sexual orientation differences in ages 40 or older (Fredriksen-Goldsen et al., 2013; Hughes et al., 2006; McKirnan and Peterson, 1989). An analysis of representative U.S. data showed declines in the prevalence of tobacco and alcohol disorders among SMs between ages 26-35 but increases in prevalence between the mid-30s to mid-40s (Evans-Polce et al., 2019).

We uniquely examined how GM status is related to risk for SUDs. This is an important contribution as studies assessing SUDs by gender identity are limited and typically focused on substance use instead of abuse (Buchting et al., 2017; Coulter et al., 2015; Gerend et al., 2017;

Keuroghlian et al., 2015; Reisner et al., 2016). In contrast to findings related to sexual orientation, we did not find consistent evidence of greater prevalence of SUDs among GMs after accounting for sexual orientation in statistical models. The only exception is that GMs assigned male evidenced elevated odds for alcohol dependence. This lack of evidence, however, should be interpreted with caution considering small numbers of GM participants in GUTS and previous evidence indicating their disproportionate substance use (Buchting et al., 2017; Day et al., 2017; De Pedro et al., 2017; De Pedro and Shim-Pelayo, 2018; Gerend et al., 2017). Additional studies quantifying associations between gender identity and SUDs are needed.

#### 4.2 SUDs among SM Assigned Female at Birth

Among the general population, more people assigned male at birth report probable SUDs than do people assigned female at birth (Agabio et al., 2017; Chou et al., 2016; Evans et al., 2018). In contrast, we found SMs assigned female generally had similar or higher levels of SUDs compared to SMs assigned male, and sexual-orientation differences were larger in assigned females than assigned males. One reason is that comparisons between SM and CH women will yield relatively large effect sizes because CH women have the lowest levels of SUDs of all groups defined by sexual orientation and birth sex. Beyond this explanation, there is little insight into why SM women are at especially elevated risk, though some have proposed that SM women are at greater risk for minority-specific stressors and mood disorders, resulting in greater risk for SUDs (Goldberg et al., 2013; McCabe et al., 2009b).

#### 4.3 Drug Use and SGM Status Among Those with Drug Use Disorder

Among participants with a drug use disorder, we found that some subgroups of SGMs had elevated odds of reporting use of certain drugs (e.g., ecstasy, LSD, methamphetamine, inhalants) compared with CHs and cisgender participants. Studies examining sexual orientation

or gender identity differences in drug use among individuals with drug use disorders are rare; however, cross-sectional studies with participants of the NSDUH found that SM adults were significantly more likely than heterosexuals to report past-year marijuana and other drug use (Medley, 2016; Schuler et al., 2018). This indicates that SGMs may be more likely to use different substances than non-SGMs, which has implications for screening, intervention, and treatment (Magill and Ray, 2009; Volkow et al., 2015).

#### 4.4 Strengths/Limitations

The DSM-IV defined separate criteria for substance abuse and dependence, whereas in the updated DSM-5, abuse and dependence are combined into a single SUD diagnosis (e.g., alcohol use disorder). Studies comparing DSM-IV and DSM-5 SUD diagnostic criteria have shown increases (Agrawal et al., 2011; Bartoli et al., 2015; Goldstein et al., 2015; Kelly et al., 2014; Mewton et al., 2011; Peer et al., 2013), no differences (Bartoli et al., 2017; Hasin et al., 2013; Proctor et al., 2012), and decreases (Goldstein et al., 2015; Tuithof et al., 2014) in prevalence. Increases in SUD prevalences under DSM-5 may relate to the inclusion of "diagnostic orphans" in diagnoses—those who meet one or two DSM-IV criteria for dependence, but none for abuse (Peer et al., 2013; Proctor et al., 2012). Nonetheless, concordance of DSM-IV and DSM-5 diagnoses are acceptable, with concordance increasing with severity (Compton et al., 2013; Dawson et al., 2013; Denis et al., 2015), suggesting that our findings are likely similar to those resulting had we used DSM-5 criteria. Further research is needed to clarify this issue.

GUTS participants are not representative of the U.S. population as they are children of registered nurses and predominantly non-Hispanic White. The prevalence of SUDs in GUTS, however, is comparable to same-aged participants of the NSDUH (Substance Abuse and Mental Health Services Administration, 2017), as is the distribution of SGMs enrolled in GUTS

compared to population-based studies (Corliss et al., 2014; Krueger et al., 2018). Additionally, GUTS participants were not enrolled based on their sexual orientation or gender identity.

GUTS assessed sexual orientation with a single item tapping both identity and attraction. This limits direct comparisons between our findings and other studies assessing dimensions of sexual orientation (i.e., identity, attraction, and behavior) separately because research indicates these dimensions have different associations with substance involvement (McCabe 2005; McCabe 2009; Drabble 2005; Boyd 2019). Further, despite the large sample size, we were limited in our ability to detect within group differences among SGMs.

Despite these limitations, our study is strengthened by including multiple SGM subgroups, enabling examination of heterogeneous outcomes that may otherwise be obscured when combining SGM categories. Future research should include more diverse, nationally representative samples to enable examination of interactions between sexual orientation, gender identity, and other sociodemographic factors to further identify higher-risk SGM subgroups.

#### 4.5 Clinical and Public Health Relevance

Among the general population, young adults with SUDs experience disproportionate economic and public health burdens and have low utilization of SUD treatment (Center for Behavioral Health Statistics and Quality, 2016). For SGM young adults, these issues may be even more persistent, with one study finding that less than 4% of the 14-20% of SMs needing treatment actually accessing treatment (Medley, 2016). Specific barriers to treatment among SGMs include a lack of targeted interventions, differences in coping strategies and psychiatric comorbidities, discrimination within healthcare settings, lack of provider knowledge about SGM health needs, and lack of insurance (Hughes et al., 2016; Lee et al., 2016; Lyons et al., 2015; McCabe et al., 2013). Consequently, increasing access to treatment alone may be insufficient to address SGM SUD disparities. Efforts should also focus on bolstering the provision of culturally tailored, SGM affirming treatment which promotes resilience, coping, and wellness. Further, given high co-morbidity with other mental disorders, interventions are needed which integrate psychological and SUD treatment (Lee et al., 2015; Mereish et al., 2015).

#### 5. Conclusions

There is increased risk for substance use and disorders among SGM young adults, yet few studies have estimated sexual orientation and gender identity differences in SUDs during this developmental period when risk peaks. Our findings highlight the importance of examining sexual orientation and gender identity differences in assessing risk for SUDs among young adults.

#### References

- Agabio, R., Pisanu, C., Gessa, G.L., Franconi, F., 2017. Sex differences in alcohol use disorder. Curr. Med. Chem. 24, 2661-2670.
- Agrawal, A., Heath, A.C., Lynskey, M.T., 2011. DSM-IV to DSM-5: The impact of proposed revisions on diagnosis of alcohol use disorders. Addiction 106, 1935-1943.
- American Psychiatric Association, 1994. Diagnostic and statistical manual of mental disorders, Fourth ed. American Psychiatric Association, Washington, DC.
- Arria, A.M., Garnier-Dykstra, L.M., Caldeira, K.M., Vincent, K.B., Winick, E.R., O'Grady,
  K.E., 2013a. Drug use patterns and continuous enrollment in college: Results from a longitudinal study. J. Stud. Alcohol Drugs 74, 71-83.
- Arria, A.M., Garnier-Dykstra, L.M., Cook, E.T., Caldeira, K.M., Vincent, K.B., Baron, R.A., O'Grady, K.E., 2013b. Drug use patterns in young adulthood and post-college employment. Drug Alcohol Depend. 127, 23-30.
- Bartoli, F., Carra, G., Biagi, E., Crocamo, C., Dakanalis, A., Di Carlo, F., Parma, F., Perin, A.P.,
  Di Giacomo, E., Zappa, L., Madeddu, F., Colmegna, F., Clerici, M., 2017. Agreement
  between DSM-IV and DSM-5 criteria for alcohol use disorder among outpatients
  suffering from depressive and anxiety disorders. Am. J. Addict. 26, 53-56.
- Bartoli, F., Carra, G., Crocamo, C., Clerici, M., 2015. From DSM-IV to DSM-5 alcohol use disorder: An overview of epidemiological data. Addict. Behav. 41, 46-50.
- Buchting, F.O., Emory, K.T., Scout, Kim, Y., Fagan, P., Vera, L.E., Emery, S., 2017.Transgender use of cigarettes, cigars, and e-cigarettes in a national study. Am. J. Prev.Med. 53, e1-e7.

- Center for Behavioral Health Statistics and Quality, 2016. Key substance use and mental health indicators in the United States: Results from the 2015 National Survey on Drug Use and Health, HHS Publication No. SMA 16-4984, NSDUH Series H-51 ed.
- Chou, S.P., Goldstein, R.B., Smith, S.M., Huang, B., Ruan, W.J., Zhang, H., Jung, J., Saha, T.D.,
  Pickering, R.P., Grant, B.F., 2016. The epidemiology of DSM-5 nicotine use disorder:
  Results from the National Epidemiologic Survey on Alcohol and Related Conditions-III.
  J. Clin. Psychiatry 77, 1404-1412.
- Cochran, S.D., Grella, C.E., Mays, V.M., 2012. Do substance use norms and perceived drug availability mediate sexual orientation differences in patterns of substance use? Results from the California Quality of Life Survey II. J. Stud. Alcohol Drugs 73, 675-685.
- Compton, W.M., Dawson, D.A., Goldstein, R.B., Grant, B.F., 2013. Crosswalk between DSM-IV dependence and DSM-5 substance use disorders for opioids, cannabis, cocaine and alcohol. Drug Alcohol Depend. 132, 387-390.
- Corliss, H.L., Rosario, M., Birkett, M.A., Newcomb, M.E., Buchting, F.O., Matthews, A.K.,
  2014. Sexual orientation disparities in adolescent cigarette smoking: Intersections with race/ethnicity, gender, and age. Am. J. Public Health 104, 1137-1147.
- Coulter, R.W., Blosnich, J.R., Bukowski, L.A., Herrick, A.L., Siconolfi, D.E., Stall, R.D., 2015.
   Differences in alcohol use and alcohol-related problems between transgender- and nontransgender-identified young adults. Drug Alcohol Depend. 154, 251-259.
- Coulter, R.W.S., Jun, H.J., Calzo, J.P., Truong, N.L., Mair, C., Markovic, N., Charlton, B.M., Silvestre, A.J., Stall, R., Corliss, H.L., 2018. Sexual-orientation differences in alcohol use trajectories and disorders in emerging adulthood: Results from a longitudinal cohort study in the United States. Addiction Epub ahead of print.

- Dawson, D.A., Goldstein, R.B., Grant, B.F., 2013. Differences in the profiles of DSM-IV and DSM-5 alcohol use disorders: Implications for clinicians. Alcohol. Clin. Exp. Res. 37, E305-E313.
- Day, J.K., Fish, J.N., Perez-Brumer, A., Hatzenbuehler, M.L., Russell, S.T., 2017. Transgender youth substance use disparities: Results from a population-based sample. J. Adolesc. Health 61, 729-735.
- De Pedro, K.T., Gilreath, T.D., Jackson, C., Esqueda, M.C., 2017. Substance use among transgender students in California public middle and high schools. J. Sch. Health 87, 303-309.
- De Pedro, K.T., Shim-Pelayo, H., 2018. Prevalence of substance use among lesbian, gay, bisexual, and transgender youth in military families: Findings from the California Healthy Kids Survey. Subst. Use Misuse. Epub ahead of print.
- Denis, C.M., Gelernter, J., Hart, A.B., Kranzler, H.R., 2015. Inter-observer reliability of DSM-5 substance use disorders. Drug Alcohol Depend. 153, 229-235.
- Evans-Polce, R.J., Veliz, P.T., Boyd, C.J., Hughes, T.L., McCabe, S.E., 2019. Associations between sexual orientation discrimination and substance use disorders: Differences by age in US adults. Soc. Psychiatry Psychiatr. Epidemiol. Epub ahead of print.
- Evans, E.A., Upchurch, D.M., Simpson, T., Hamilton, A.B., Hoggatt, K.J., 2018. Differences by veteran/civilian status and gender in associations between childhood adversity and alcohol and drug use disorders. Soc. Psychiatry Psychiatr. Epidemiol. 53, 421-435.
- Falk, D., Yi, H.Y., Hiller-Sturmhofel, S., 2008. An epidemiologic analysis of co-occurring alcohol and drug use and disorders: Findings from the National Epidemiologic Survey of Alcohol and Related Conditions (NESARC). Alcohol Res. Health 31, 100-110.

- Felner, J.K., Wisdom, J.P., Williams, T., Katuska, L., Haley, S., Jun, H.-J., Corliss, H.L., in press. Stress, coping, and context: Examining substance use among LGBTQ young adults with probable substance use disorders. Psychiatr. Serv.
- Fergusson, D.M., Boden, J.M., Horwood, L.J., 2008. The developmental antecedents of illicit drug use: Evidence from a 25-year longitudinal study. Drug Alcohol Depend. 96, 165-177.
- Field, A.E., Camargo, C.A., Jr., Taylor, C.B., Berkey, C.S., Frazier, A.L., Gillman, M.W., Colditz, G.A., 1999. Overweight, weight concerns, and bulimic behaviors among girls and boys. J. Am. Acad. Child Adolesc. Psychiatry 38, 754-760.
- Field, A.E., Sonneville, K.R., Falbe, J., Flint, A., Haines, J., Rosner, B., Camargo, C.A., Jr.,
  2014. Association of sports drinks with weight gain among adolescents and young adults.
  Obesity. 22, 2238-2243.
- Flentje, A., Bacca, C.L., Cochran, B.N., 2015. Missing data in substance abuse research? Researchers' reporting practices of sexual orientation and gender identity. Drug Alcohol Depend. 147, 280-284.
- Flentje, A., Heck, N.C., Sorensen, J.L., 2014. Characteristics of transgender individuals entering substance abuse treatment. Addict. Behav. 39.
- Fredriksen-Goldsen, K.I., Kim, H.J., Barkan, S.E., Muraco, A., Hoy-Ellis, C.P., 2013. Health disparities among lesbian, gay, and bisexual older adults: Results from a populationbased study. Am. J. Public Health 103, 1802-1809.
- Gerend, M.A., Newcomb, M.E., Mustanski, B., 2017. Prevalence and correlates of smoking and e-cigarette use among young men who have sex with men and transgender women. Drug Alcohol Depend. 179, 395-399.

- Goldberg, S., Strutz, K.L., Herring, A.A., Halpern, C.T., 2013. Risk of substance abuse and dependence among young adult sexual minority groups using a multidimensional measure of sexual orientation. Public Health Rep. 128, 144-152.
- Goldstein, R.B., Chou, S.P., Smith, S.M., Jung, J., Zhang, H., Saha, T.D., Pickering, R.P., Ruan,
  W.J., Huang, B., Grant, B.F., 2015. Nosologic comparisons of DSM-IV and DSM-5
  alcohol and drug use disorders: Results from the National Epidemiologic Survey on
  Alcohol and Related Conditions-III. J. Stud. Alcohol Drugs. 76, 378-388.
- Gonzales, G., Przedworski, J., Henning-Smith, C., 2016. Comparison of health and health risk factors between lesbian, gay, and bisexual adults and heterosexual adults in the United States: Results from the National Health Interview Survey. JAMA Intern. Med. 176, 1344-1351.
- Hasin, D.S., O'Brien, C.P., Auriacombe, M., Borges, G., Bucholz, K., Budney, A., Compton,
  W.M., Crowley, T., Ling, W., Petry, N.M., Schuckit, M., Grant, B.F., 2013. DSM-5
  criteria for substance use disorders: recommendations and rationale. Am. J. Psychiatry.
  170, 834-851.
- Hughes, T.L., Wilsnack, S.C., Kantor, L.W., 2016. The influence of gender and sexual orientation on alcohol use and slcohol-related problems: Toward a global perspective.Alcohol Res. Curr. Rev. 38, 121-132.
- Hughes, T.L., Wilsnack, S.C., Szalacha, L.A., Johnson, T., Bostwick, W.B., Seymour, R., Aranda, F., Benson, P., Kinnison, K.E., 2006. Age and racial/ehnic differences in drinking and drinking-related problems in a community sample of lesbians. J. Stud. Alcohol. 67, 579-590.

- Kelly, S.M., Gryczynski, J., Mitchell, S.G., Kirk, A., O'Grady, K.E., Schwartz, R.P., 2014.
   Concordance between DSM-5 and DSM-IV nicotine, alcohol, and cannabis use disorder diagnoses among pediatric patients. Drug Alcohol Depend. 140, 213-216.
- Kerridge, B.T., Pickering, R.P., Saha, T.D., Ruan, W.J., Chou, S.P., Zhang, H., Jung, J., Hasin, D.S., 2017. Prevalence, sociodemographic correlates and DSM-5 substance use disorders and other psychiatric disorders among sexual minorities in the United States. Drug Alcohol Depend. 170, 82-92.
- Keuroghlian, A.S., Reisner, S.L., White, J.M., Weiss, R.D., 2015. Substance use and treatment of substance use disorders in a community sample of transgender adults. Drug Alcohol Depend. 152, 139-146.
- Krueger, E.A., Meyer, I.H., Upchurch, D.M., 2018. Sexual orientation group differences in perceived stress and depressive symptoms among young adults in the United States. LGBT Health 5, 242-249.
- Lee, J.H., Gamarel, K.E., Bryant, K.J., Zaller, N.D., Operario, D., 2016. Discrimination, mental health, and substance use disorders among sexual minority populations. LGBT Health 3, 258-265.
- Lee, J.H., Gamarel, K.E., Kahler, C.W., Marshall, B.D.L., van den Berg, J.J., Bryant, K., Zaller, N.D., Operario, D., 2015. Co-occurring psychiatric and drug use disorders among sexual minority men with lifetime alcohol use disorders. Drug Alcohol Depend. 151, 167-172.
- Liang, K.-Y., Zeger, S.L., 1986. Longitudinal data analysis using generalized linear models. Biometrika. 73, 13-22.

- Lyons, T., Shannon, K., Pierre, L., Small, W., Krüsi, A., Kerr, T., 2015. A qualitative study of transgender individuals' experiences in residential addiction treatment settings: Stigma and inclusivity. Subst. Abuse Treat. Prev. Policy. 10, 17.
- Mack, K.A., Jones, C.M., Ballesteros, M.F., 2017. Illicit drug use, illicit drug use disorders, and drug overdose deaths in metropolitan and nonmetropolitan areas— United States.
   MMWR Surveill. Summ. 66, 1-12.
- Magill, M., Ray, L.A., 2009. Cognitive-behavioral treatment with adult alcohol and illicit drug users: a meta-analysis of randomized controlled trials. J. Stud. Alcohol Drugs. 70, 516-527.
- McCabe, S.E., Boyd, C.J., Cranford, J.A., Teter, C.J., 2009a. Motives for nonmedical use of prescription opioids among high school seniors in the United States: Self-treatment and beyond. Arch. Pediatr. Adolesc. Med. 163, 739-744.
- McCabe, S.E., Hughes, T.L., Bostwick, W.B., West, B.T., Boyd, C.J., 2009b. Sexual orientation, substance use behaviors and substance dependence in the United States. Addiction 104, 1333-1345.
- McCabe, S.E., Matthews, A.K., Lee, J.G.L., Veliz, P., Hughes, T.L., Boyd, C.J., 2018. Tobacco use and sexual orientation in a national cross-sectional study: Age, race/ethnicity, and sexual identity– Attraction differences. Am. J. Prev. Med. 54, 736-745.
- McCabe, S.E., West, B.T., Hughes, T.L., Boyd, C.J., 2013. Sexual orientation and substance abuse treatment utilization in the United States: Results from a national survey. J. Subst. Abuse Treat. 44, 4-12.
- McKirnan, D.J., Peterson, P.L., 1989. Alcohol and drug use among homosexual men and women: Epidemiology and population characteristics. Addict. Behav. 14, 545-553.

- Medley, G., Lipari, R. N., Bose, J., Cribb, D. S., Kroutil, L. A., McHenry, G., 2016. Sexual orientation and estimates of adult substance use and mental health: Results from the 2015 National Survey on Drug Use and Health, NSDUH Data Review. Substance Abuse and Mental Health Services Administration, Rockville, MD.
- Mereish, E.H., Goldbach, J.T., Burgess, C., DiBello, A.M., 2017. Sexual orientation, minority stress, social norms, and substance use among racially diverse adolescents. Drug Alcohol Depend. 178, 49-56.
- Mereish, E.H., Lee, J.H., Gamarel, K.E., Zaller, N.D., Operario, D., 2015. Sexual orientation disparities in psychiatric and drug use disorders among a nationally representative sample of women with alcohol use disorders. Addict. Behav. 47, 80-85.
- Merikangas, K.R., McClair, V.L., 2012. Epidemiology of substance use disorders. Hum. Genet. 131, 779-789.
- Mewton, L., Slade, T., McBride, O., Grove, R., Teesson, M., 2011. An evaluation of the proposed DSM-5 alcohol use disorder criteria using Australian national data. Addiction 106, 941-950.
- Moss, H.B., Goldstein, R.B., Chen, C.M., Yi, H.Y., 2015. Patterns of use of other drugs among those with alcohol dependence: Associations with drinking behavior and psychopathology. Addict. Behav. 50, 192-198.
- Newcomb, M.E., Heinz, A.J., Birkett, M., Mustanski, B., 2014. A longitudinal examination of risk and protective factors for cigarette smoking among lesbian, gay, bisexual, and transgender youth. J. Adolesc. Health 54, 558-564.
- Nutt, D.J., 1996. Addiction: brain mechanisms and their treatment implications. Lancet 347, 31-36.

- Patel, V., Chisholm, D., Parikh, R., Charlson, F.J., Degenhardt, L., Dua, T., Ferrari, A.J., Hyman, S., Laxminarayan, R., Levin, C., Lund, C., Medina Mora, M.E., Petersen, I., Scott, J., Shidhaye, R., Vijayakumar, L., Thornicroft, G., Whiteford, H., 2016. Addressing the burden of mental, neurological, and substance use disorders: Key messages from Disease Control Priorities, 3rd edition. Lancet 387, 1672-1685.
- Peer, K., Rennert, L., Lynch, K.G., Farrer, L., Gelernter, J., Kranzler, H.R., 2013. Prevalence of DSM-IV and DSM-5 alcohol, cocaine, opioid, and cannabis use disorders in a largely substance dependent sample. Drug Alcohol Depend. 127, 215-219.
- Proctor, S.L., Kopak, A.M., Hoffmann, N.G., 2012. Compatibility of current DSM-IV and proposed DSM-5 diagnostic criteria for cocaine use disorders. Addict. Behav. 37, 722-728.
- Reisner, S.L., Biello, K.B., White Hughto, J.M., Kuhns, L., Mayer, K.H., Garofalo, R., Mimiaga, M.J., 2016. Psychiatric diagnoses and comorbidities in a diverse, multicity cohort of young transgender women: Baseline findings from project lifeskills. JAMA Pediatr. 170, 481-486.
- Remafedi, G., Resnick, M., Blum, R., Harris, L., 1992. Demography of sexual orientation in adolescents. Pediatrics 89, 714-721.
- Schuler, M.S., Rice, C.E., Evans-Polce, R.J., Collins, R.L., 2018. Disparities in substance use behaviors and disorders among adult sexual minorities by age, gender, and sexual identity. Drug Alcohol Depend. 189, 139-146.
- Substance Abuse and Mental Health Services Administration, 2017. Key substance use and mental health indicators in the United States: Results from the 2016 National Survey on Drug Use and Health, HHS Publication No. SMA 17-5044, NSDUH Series H-52. Center

for Behavioral Health Statistics and Quality, Substance Abuse and Mental Health Services Administration, Rockville, MD.

- Talley, A.E., Hughes, T.L., Aranda, F., Birkett, M., Marshal, M.P., 2014. Exploring alcohol-use behaviors among heterosexual and sexual minority adolescents: Intersections with sex, age, and race/ethnicity. Am. J. Public Health 104, 295-303.
- Tuithof, M., Ten Have, M., van den Brink, W., Vollebergh, W., de Graaf, R., 2014. The relationship between excessive alcohol consumption and alcohol use disorders according to DSM-IV and DSM-5. Alcohol. Clin. Exp. Res. 38, 249-256.
- Tupler, L.A., Zapp, D., DeJong, W., Ali, M., O'Rourke, S., Looney, J., Swartzwelder, H.S.,
  2017. Alcohol-related blackouts, negative alcohol-related consequences, and motivations for drinking reported by newly matriculating transgender college students. Alcohol. Clin. Exp. Res. 41, 1012-1023.
- Volkow, N.D., Koob, G., Baler, R., 2015. Biomarkers in substance use disorders. ACS Chem. Neurosci. 6, 522-525.
- Whiteford, H.A., Degenhardt, L., Rehm, J., Baxter, A.J., Ferrari, A.J., Erskine, H.E., Charlson,
  F.J., Norman, R.E., Flaxman, A.D., Johns, N., Burstein, R., Murray, C.J., Vos, T., 2013.
  Global burden of disease attributable to mental and substance use disorders: Findings
  from the Global Burden of Disease Study 2010. Lancet 382, 1575-1586.

	Assigned Fen	nale at Birth	Assigned Male at Birt				
	(N=11,	832)	(N=5,	664)			
Characteristic	N	%	N	%			
Sexual Orientation							
Completely heterosexual	8,907	75.3	4,777	84.3			
Mostly heterosexual	2,296	19.4	562	9.9			
Bisexual	387	3.3	52	0.9			
Gay/lesbian	242	2.1	273	4.8			
Gender Identity							
Cisgender	11,747	99.3	5,603	98.9			
Gender minority	85	0.7	61	1.1			
Age, years							
20-25	4,405	37.2	2,177	38.4			
26-35	7,427	62.8	3,487	61.6			
Race/Ethnicity							
White	10,986	92.9	5,275	93.1			
Other	846	7.2	389	6.9			
Cohort							
GUTS1	9,276	78.4	4,493	79.3			
GUTS2	2,556	21.6	1,171	20.7			
Region of Residence							
West	2,080	17.6	1,112	19.6			
Midwest	3,793	32.1	1,820	32.1			
South	2,136	18.1	950	16.8			
Northeast	3,823	32.3	1,782	31.5			

Table 1. Characteristics of repeated measures observations of Growing Up Today Study participants aged 20-35 years, by sex assigned at birth

(2010, 2015).

Note: N equals the number of observations over repeated measures. Percentages within variables sum to 100% except for rounding error.

**Table 2.** Past 12-month prevalences of substance use disorders among Growing Up Today Study participants aged 20-35 years, by sex assigned at

 birth (2010, 2015).

		Nicotine Alcohol Use Disorder					Drug Use Disorder						Number of SUDs								
	D	epend	lence		Abu	se	De	epend	ence		Abu	se	D	epend	lence	(	One S	UD	]	ſwo S	UDs
Characteristic	Ν	%	p-value	N	%	p-value	N	%	p-value	N	%	p-value	N	%	p-value	N	%	p-value	N	%	p-value
Assigned Female at	<u>Birth</u>																				
Sexual Orientation																					
Comp. hetero.	530	6.0		425	4.8		316	3.6		45	0.5		160	1.8		972	10.9		238	2.7	
Mostly hetero.	311	13.6	< 0.001	193	8.4	< 0.001	184	8.0	< 0.001	33	1.4	< 0.001	148	6.5	< 0.001	468	20.4	< 0.001	179	7.8	< 0.001
Bisexual	76	19.6	< 0.001	28	7.3	0.016	28	7.3	< 0.001	10	2.6	< 0.001	44	11.5	< 0.001	101	26.1	< 0.001	40	10.3	< 0.001
Lesbian	44	18.2	< 0.001	19	7.9	0.018	23	9.6	< 0.001	6	2.5	< 0.001	18	7.5	< 0.001	49	20.3	< 0.001	28	11.6	< 0.001
Gender Identity																					
Cisgender	950	8.1		661	5.6		540	4.6		91	0.8		360	3.1		1569	13.4		476	4.1	
Gender minority	11	12.9	0.159	4	4.7	0.850	11	12.9	0.003	3	3.5	0.033	10	11.8	0.0001	21	24.7	0.002	9	10.6	0.006
Assigned Male at Bi	rth_																				
Sexual Orientation																					
Comp. hetero.	449	9.4		365	7.7		367	7.7		75	1.6		267	5.6		839	17.6		305	6.4	
Mostly hetero.	81	14.4	0.002	52	9.3	0.091	66	11.8	0.001	17	3.1	0.006	67	12.0	< 0.001	125	22.2	0.001	68	12.1	< 0.001
Bisexual	12	23.1	0.004	5	9.8	0.488	6	11.8	0.266	2	3.9	0.184	5	9.8	0.186	18	34.6	0.002	5	9.6	0.143
Gay	37	13.6	0.049	28	10.3	0.044	42	15.4	< 0.001	6	2.2	0.362	27	10.0	0.007	77	28.2	< 0.001	29	10.6	0.002
Gender Identity																					
Cisgender	569	10.2		444	8.0		469	8.4		99	1.8		358	6.4		1042	18.6		399	7.1	

Gender minority	10 16.4	0.406	6	9.8	0.372	12 19.7	0.003	1	1.6	0.993	8 13.1	0.072	17 27.9	0.041	8 13.1	0.079
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Note: N equals the number of observations over repeated measures. Prevalence shown are unadjusted. Bivariate P-values comparing completely heterosexuals to sexual minorities and cisgenders to gender minorities estimated by generalized estimating equations with exchangeable correlation structure. Associations in bold are p < .05.

	Model 1: Nicotine			e Model 2: Alcohol Use Disorder				lodel 3: Drug	Disorder	Model 4: No. of Substance Disord				
	De	pendence		<u>Abuse</u>	De	pendence	-	Abuse	De	pendence	<u>Or</u>	ne SUD	<u>Tw</u>	vo+ SUDs
Characteristic	AOR	95% CI	AOR	95% CI	AOR	95% CI	AOR	95% CI	AOR	95% CI	AOR	95% CI	AOR	95% CI
Sexual Orientation														
Completely heterosexual	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
Mostly heterosexual	2.42	(2.07-2.84)	1.96	(1.63-2.36)	2.53	(2.08-3.08)	3.05	(1.93-4.82)	3.89	(3.06-4.95)	2.32	(2.03-2.64)	3.71	(3.00-4.60)
Bisexual	4.00	(2.98-5.38)	1.61	(1.08-2.41)	2.10	(1.37-3.22)	5.32	(2.46-11.5)	6.97	(4.72-10.3)	3.28	(2.55-4.22)	5.62	(3.77-8.38)
Lesbian	3.82	(2.63-5.54)	1.88	(1.13-3.13)	2.86	(1.73-4.72)	4.94	(1.94-12.6)	4.34	(2.58-7.29)	2.37	(1.66-3.38)	5.66	(3.56-9.00)
Gender Identity														
Cisgender	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
Gender minority	0.80	(0.39-1.67)	0.65	(0.24-1.77)	1.84	(0.86-3.94)	2.13	(0.38-11.9)	1.69	(0.77-3.72)	1.32	(0.76-2.30)	1.37	(0.54-3.45)
Age, Years														
20-25	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
26-35	0.75	(0.66-0.86)	0.71	(0.61-0.84)	0.66	(0.55-0.80)	0.56	(0.37-0.85)	0.55	(0.44-0.69)	0.72	(0.64-0.80)	0.61	(0.50-0.74)
Race/Ethnicity														
White	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
Other	1.09	(0.82-1.45)	0.78	(0.56-1.09)	0.76	(0.52-1.12)	1.01	(0.47-2.16)	1.12	(0.75-1.68)	0.74	(0.58-0.94)	1.14	(0.79-1.64)
Cohort														
GUTS1	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
GUTS2	0.43	(0.35-0.52)	1.22	(1.01-1.47)	0.72	(0.57-0.92)	0.87	(0.53-1.42)	0.76	(0.57-1.00)	0.82	(0.71-0.94)	0.54	(0.41-0.71)

**Table 3.** Results of multivariable generalized estimating equations regression models estimating past 12-month substance usedisorders among Growing Up Today Study participants aged 20-35 years assigned female at birth (2010, 2015).

Region of Residence									
West	1.00 -	1.00 -	1.00	- 1.00	- 1	- 00.	1.00	-	1.00 -
Midwest	<b>1.51</b> (1.20-1.89	) 1.05 (0.82-1.34	) 0.83 (0.0	63-1.10) 0.70	(0.38-1.30) 0	).91 (0.65-1.26)	1.01	(0.85-1.21)	1.25 (0.93-1.69)
South	<b>1.35</b> (1.07-1.69	) 0.93 (0.70-1.22	) 0.91 (0.0	67-1.24) 0.97	(0.45-1.83) 0	0.67 (0.75-1.00)	0.89	(0.73-1.08)	1.19 (0.86-1.65)
Northeast	<b>1.29</b> (1.00-1.67	) 1.08 (0.85-1.37	) 1.14 (0.8	88-1.48) 0.80	(0.51-1.43) 1	.03 (0.45-1.41)	1.08	(0.91-1.28)	1.32 (0.99-1.77)

Associations in bold are p < .05. Binary and multinomial logistic regression with exchangeable correlation structure estimated adjusted odds ratios (AOR) and 95% confidence intervals (CI). Participants without the substance use disorder were the referent group.

	Model 1: Nicotine		e Model 2: Alcohol Use Disorder				Model 3: Drug Use Disorder					Model 4: Number of SU		
	De	pendence		Abuse	De	ependence		Abuse	Dependence		One SUD		Tw	o+ SUDs
Characteristic	AOR	95% CI	AOR	95% CI	AOR	95% CI	AOR	95% CI	AOR	95% CI	AOR	95% CI	AOR	95% CI
Sexual Orientation														
Completely heterosexual	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
Mostly heterosexual	1.65	(1.27-2.14)	1.31	(0.96-1.78)	1.71	(1.27-2.30)	2.13	(1.26-3.62)	2.29	(1.69-3.12)	1.51	(1.21-1.89)	2.31	(1.71-3.11)
Bisexual	3.08	(1.51-6.24)	1.30	(0.53-3.20)	1.48	(0.61-3.61)	2.73	(0.59-12.63)	1.69	(0.66-4.34)	2.62	(1.43-4.82)	2.02	(0.73-5.58)
Gay	1.58	(1.08-2.33)	1.52	(1.00-2.30)	2.35	(1.58-3.50)	1.52	(0.65-3.54)	1.85	(1.17-2.91)	2.03	(1.48-2.77)	2.16	(1.37-3.41)
Gender Identity														
Cisgender	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
Gender minority	1.11	(0.48-2.55)	1.37	(0.60-3.15)	2.30	(1.12-4.69)	0.74	(0.09-6.42)	1.68	(0.68-4.19)	1.47	(0.82-2.63)	1.78	(0.63-5.01)
Age, Years														
20-25	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
26-35	0.62	(0.53-0.72)	0.75	(0.61-1.91)	0.62	(0.51-0.76)	0.71	(0.47-1.08)	0.69	(0.56-0.85)	0.73	(0.64-0.84)	0.54	(0.44-0.67)
Race/Ethnicity														
White	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
Other	0.86	(0.57-1.30)	1.48	(1.02-2.16)	0.99	(0.65-1.50)	0.38	(0.12-1.19)	0.97	(0.60-1.56)	1.02	(0.75-1.40)	0.99	(0.62-1.56)
Cohort														
GUTS1	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
GUTS2	0.44	(0.34-0.56)	0.88	(0.69-1.12)	0.50	(0.37-0.67)	1.08	(0.68-1.72)	0.73	(0.55-0.97)	0.69	(0.57-0.83)	0.50	(0.37-0.67)

Table 4. Results of multivariable generalized estimating equations regression models estimating past 12-month substance use disorders among

Growing Up Today Study participants aged 20-35 years assigned male at birth (2010, 2015).

Region of Residence														
West	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-	1.00	-
Midwest	0.95	(0.73-1.24)	0.99	(0.74-1.32)	0.81	(0.61-1.09)	0.79	(0.44-1.42)	0.50	(0.36-0.70)	0.85	(0.69-1.04)	0.72	(0.53-0.99)
South	0.98	(0.76-1.28)	1.20	(0.87-1.66)	0.97	(0.70-1.34)	1.13	(0.60-2.13)	0.61	(0.42-0.88)	1.03	(0.81-1.30)	0.90	(0.60-1.28)
Northeast	1.12	(0.84-1.50)	1.07	(0.80-1.44)	0.86	(0.72-1.27)	0.90	(0.51-1.60)	0.72	(0.53-0.97)	1.02	(0.83-1.26)	0.81	(0.64-1.09)

Associations in bold are p < .05. Binary and multinomial logistic regression with exchangeable correlation structure estimated adjusted odds ratios (AOR) and 95% confidence intervals (CI). Participants without the substance use disorder were the referent group.

Table

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**Table 5.** Prevalence and results of multivariable generalized estimating equations logistic regressionmodels estimating past 12-month drug use by sexual orientation and gender identity among Growing UpToday Study participants aged 20-35 years who evidenced past 12-month drug use disorder (2010, 2015).

			S	exual Orientat	Gender Identity						
	<u>CH</u>	<u>Mostly</u>	Heterose	<u>xual</u>	<u>Gay/Le</u>	sbian/Bis	sexual	<u>Cisgender</u>	Gende	r Minorit	У
Type of Drug	%	%	OR	95% CI	%	OR	95% CI	%	%	OR	95% CI
Assigned Female at Birth											
Marijuana	86.3	90.1	1.47	(0.78-2.80)	94.9	2.78	(0.91-8.43)	89.1	92.3	0.91	(0.16-5.07)
Cocaine	27.8	33.7	1.35	(0.87-2.10)	25.6	0.90	(0.50-1.65)	29.9	23.1	0.69	(0.15-3.09)
Heroin	2.0	3.9	1.91	(0.50-7.22)	6.4	3.09	(0.80-11.9)	3.1	15.4	4.33	(0.60-31.1)
MDMA/ecstasy	15.6	21.0	1.55	(0.91-2.65)	30.8	2.35	(1.24-4.44)	20.2	23.1	0.72	(0.19-2.77)
LSD/mushrooms	18.1	26.5	1.64	(0.99-2.71)	33.3	1.92	(1.03-3.59)	22.8	61.5	4.17	(1.15-15.1)
Methamphetamine	4.4	3.9	0.84	(0.27-2.62)	0.0			3.3	7.7	9.82	(0.31-309)
Amphetamines*	6.8	9.9	1.59	(0.77-3.28)	7.7	1.15	(0.43-3.11)	8.2	7.7	0.87	(0.11-6.94)
Inhalants	3.6	4.3			4.8			4.3	0.0		
Non-Medical Use of Pres	scription	Drugs									
Benzodiazepines	29.8	28.7	0.99	(0.63-1.55)	34.6	1.36	(0.77-2.40)	30.4	23.1	0.67	(0.18-2.45)
Painkillers	31.2	24.9	0.73	(0.45-1.17)	33.3	1.14	(0.65-2.02)	29.1	30.8	1.19	(0.37-3.82)
Sleeping pills	7.8	7.7	1.07	(0.51-2.24)	6.4	0.81	(0.27-2.45)	7.5	7.7	1.21	(0.12-12.2)
Stimulants	25.9	23.2	0.87	(0.55-1.39)	24.4	0.87	(0.45-1.67)	24.2	38.5	2.17	(0.69-6.84)
Assigned Male at Birth											
Marijuana	91.8	94.1	1.51	(0.56-4.13)	90.0	0.99	(0.31-3.18)	92.3	77.8	0.28	(0.05-1.57)
Cocaine	33.3	41.7	1.19	(0.71-1.98)	47.5	1.51	(0.72-3.17)	35.5	66.7	3.09	(0.64-15.0)
Heroin	4.4	7.1	1.45	(0.49-4.34)	2.5	0.41	(0.08-2.17)	4.4	22.2	13.9	(3.05-63.0)
MDMA/ecstasy	21.1	27.4	1.17	(0.66-2.07)	35.0	1.85	(0.91-3.78)	23.0	44.4	2.06	(0.47-8.96)
LSD/mushrooms	29.0	36.9	1.33	(0.79-2.24)	22.5	0.65	(0.29-1.46)	29.5	44.4	2.01	(0.43-9.43)
Methamphetamine	2.3	4.8	2.36	(0.69-8.13)	12.5	6.63	(1.52-29.0)	3.7	0.0		
Amphetamines	8.8	7.1	0.79	(0.31-2.02)	17.5	1.52	(0.55-4.24)	8.3	55.6	8.49	(2.38-30.3)
Inhalants*	4.8	5.9	1.33	(0.35-5.01)	40.0	10.6	(3.40-33.1)	7.4	57.1	6.95	(1.16-41.7)
Non-Medical Use of Pres	scription	Drugs									

Benzodiazepines*	25.7	22.6	0.87	(0.48-1.60)	37.5	1.63	(0.77-3.42)	25.8	44.4	1.89	(0.56-6.45)
Painkillers	37.1	32.1	0.82	(0.49-1.37)	30.0	0.64	(0.28-1.47)	35.0	66.7	6.19	(1.56-24.6)
Sleeping pills	5.6	2.4	0.35	(0.08-1.47)	15.0	2.14	(0.79-5.82)	5.5	22.2	4.67	(0.98-22.4)
Stimulants	26.9	35.7	1.33	(0.75-2.36)	35.0	1.21	(0.57-2.59)	28.5	66.7	3.61	(0.68-19.2)

Completely heterosexuals and cisgenders are the referent groups. Models adjusted for age, race/ethnicity, cohort, and region of residence and used exchangeable correlation structure except models noted with an asterisk (\*), which used independence correlation structure. Associations in bold are p < .05.

Supplementary Material Click here to download Supplementary Material: EGST-19-0459\_sm.doc

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#### Contributors

Hee-Jin Jun conceptualized the paper, performed the analyses, and wrote the manuscript.

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Sean Haley reviewed the manuscript and gave critical feedback.

S. Bryn Austin helped with study design, reviewed the manuscript and gave critical feedback.

Laura Katuska helped with data collection and reviewed the manuscript.

Heather L. Corliss obtained the funding, conceptualized the paper, and reviewed and revised the manuscript.

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#### **Conflict of Interest**

We have no potential conflicts of interest to disclose.

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# **Conflict of Interest**

We have no potential conflicts of interest to disclose.

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All authors have contributed to and approved the final article.

# Highlights

- Sexual minority young adults had elevated prevalence of past 12-month SUD.
- Sexual minorities were more likely to evidence multiple co-occurring SUDs.
- Orientation disparities in SUD were larger in people assigned female at birth.
- Gender-identity differences in SUD were smaller than sexual-orientation differences.
- Future research on SUDs should include sexual orientation and gender identity.