Substance Use Disorders (SUD) and Suicidal Behaviors in Adolescents: Insights From Cross-Sectional Inpatient Study

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Abstract

Objectives
To determine the demographic predictors of suicidal behaviors and measure the association between the spectrum of substance use disorders (SUD) and hospitalization for suicidal behaviors in the adolescent population.

Methods
We conducted a cross-sectional study using the nationwide inpatient sample and included 466,244 adolescent inpatients with psychiatric illnesses. The study sample was sub-grouped into suicidal (N = 182,454) and non-suicidal (N = 283,790) cohorts. The odds ratio (OR) of association for demographic characteristics and comorbid SUD in the suicidal group was evaluated using a logistic regression model with a P-value < 0.01.

Results
Our study population included 466,244 adolescent inpatients. Females had higher odds of suicidal behaviors (OR 1.45; 95% CI 1.431-1.470) compared to males. The most prevalent comorbid SUD among suicidal inpatients was cannabis (15.3%) but had a statistically non-significant association with suicidal behaviors (OR 0.98; 95% CI 0.95-0.99). Inpatients with alcohol use disorders had significantly increased odds of association with suicidal behaviors (OR 1.18; CI: 1.142-1.209) compared to other SUD. Among other substances (cannabis and stimulants), there existed a statistically non-significant association with hospitalization for suicidal behaviors.

Conclusion
Adolescent inpatients with comorbid alcohol use disorders were at 18% higher odds of hospitalization for suicidal behaviors. Our study provides a basis for more research while also suggesting potential avenues for early identification and intervention efforts for comorbid SUD in adolescents requiring psychiatric care to improve their prognosis and quality of life.

Introduction
Substance use disorders (SUD) have serious consequences for children and adolescents’ physical and mental health. The lifetime prevalence of any illicit substance use among adolescents is 42.9% [1]. The most commonly abused substance among teens is alcohol as they consume about 1/10th of all alcohol consumed in the United States (US), followed by tobacco and cannabis [2]. According to a 2019 youth risk behavior survey (YRBS) in the US, 14.8% of high school students used illicit substances, 14.3% misused prescription opioids, and 1.6% have injected illegal substances. The overall percentage of substance use has decreased among Caucasians and Hispanics, but it has remained unchanged in African Americans during this 10-year trend descriptive study. Although high-risk substance use behaviors have declined since 2009, nearly one in every seven adolescents still report drug use and misuse [3].

Several factors, including genetic predisposition, adverse familial and social environment, and substance abuse at a young age, could all contribute to addiction [4]. The “reward system” is a major
pathophysiological contributor to addiction in adolescents and the adult population. The brain receives far more dopamine after a "drug high" than the natural reward system would produce [4]. This dopaminergic reward pathway acts on the nucleus accumbens, providing positive reinforcement for substance abuse. The majority of stimulants exert their effect by increasing dopamine, norepinephrine, and serotonin levels in synapses [1]. Alcohol and sedatives/hypnotics, on the other hand, depress the central nervous system arousal by enhancing the action of gamma-aminobutyric acid (GABA) [1]. Research has demonstrated that certain developmental stages of adolescence result in higher vulnerability to SUD [5]. Furthermore, early substance exposure during a period when the brain is undergoing significant developmental changes is associated with a more chronic and severe course of SUD than is seen if started in older age groups [6].

Suicide accounts for approximately six percent of all adolescent deaths worldwide, and it is the second leading cause of death among children and youth aged 10 to 24 years old [7]. According to the YRBS 2019 report, approximately nine percent of youth in grades nine to 12 attempted suicide at least once during the year [3]. Major risk factors are being female, being of African American descent, exposure to bullying and violence, sexual orientation, alcohol and substance abuse, psychiatric comorbidities, poor familial, and social support [8]. SUD often coexist with psychiatric illnesses and further increases the suicide risk [9-11]. Adolescents with SUD and suicidal behaviors share similar risk factors that may interact with each other. Past studies have found adolescents with psychiatric comorbidities like depressive disorders and attention-deficit/hyperactivity disorder were at a higher risk of SUD, which can lead to impairment of overall functioning [12,13]. Our study focuses on identifying the demographic predictors of suicidal behaviors and assessing the relationship between the spectrum of SUD and suicidal behaviors in the adolescent population.

Materials And Methods

Study sample
We conducted a cross-sectional study using the national (nationwide) inpatient sample (NIS), which is the largest inpatient database in the US. The NIS constitutes data from 4,411 non-federal community hospitals across 44 states in the US [14]. We included 466,244 adolescent inpatients (aged 12 to 18 years) with a psychiatric illness as their primary discharge diagnosis. The sample was divided into two groups: those who did not engage in suicidal behavior/non-suicidal group (N = 283,790) and those who did engage in suicidal behavior/suicidal group (N = 182,454).

Variables
The demographic variables included in this study were age at admission, sex, race, and geographic region in the US. Comorbidities are co-diagnosed conditions found in inpatient records, and we included SUD such as alcohol, cannabis, stimulants (cocaine and amphetamine), and opioids [15].

Statistical analysis
We compared the distributions of demographic characteristics and comorbid SUD in adolescent inpatients without versus with suicidal behaviors by performing descriptive statistics and Pearson’s chi-square test for categorical variable and independent sample T-test for continuous variable (age). The odds ratio (OR) of association for demographic characteristics and comorbid SUD in the suicidal group was evaluated using a binomial logistic regression model with the non-suicidal group as the reference category. All analyses were conducted with the help of a statistical package for the social sciences (SPSS) version 26.0 (IBM Corp., Armonk, NY, USA) and with statistical significance was set at a two-sided P-value < 0.01.

Ethical approval
The NIS is publicly available de-identified data with the protection of patients, physicians, and hospital-related information; henceforward, we were not required to obtain institutional review board approval for our study [14].

Results
Our study population included 466,244 adolescent inpatients. 59.9% of our study population were females, while 40% were males. Caucasians made 62.3% of our population, and 39.1% of inpatients endorsed suicidal behaviors. A higher proportion of these inpatients with suicidal behaviors were females (65.3%) compared to 56.5% of females in the non-suicidal group. Suicidal behaviors were also more prevalent in Caucasians (65.3%), followed by blacks (14.9%), and Hispanics (12.7%). A higher proportion of adolescent inpatients with suicidal behaviors were prevalent in the midwest (37.7%) and the south (35.1%) but lower in the northeast (14.9%) and the west (12.3%) regions of the US.

The most prevalent comorbid SUD among suicidal inpatients was cannabis (15.3%), but it was significantly lower than that seen in the 16.4% non-suicidal group (P < 0.001). Approximately 6.6% of inpatients with suicidal behaviors had comorbid alcohol use, which was significantly higher than the non-suicidal cohort (six percent). Opioid use (2.6%) and stimulant use (2.4%) were observed in a higher proportion of the non-
suicidal cohort as shown in Table 1.

| Demographic                          | Suicidal (no) | Suicidal (yes) | P-value |
|--------------------------------------|---------------|----------------|---------|
| Number of inpatients                 | 283,790       | 182,454        | -       |
| Mean age (SD), in years              | 15.4 (1.9)    | 15.3 (1.8)     | 0.545   |
| Sex, in %                            |               |                |         |
| Male                                 | 43.5          | 34.7           | <0.001  |
| Female                               | 56.5          | 65.3           |         |
| Race, in %                           |               |                |         |
| Caucasian                            | 60.4          | 65.3           | <0.001  |
| Black                                | 18.6          | 14.9           |         |
| Hispanic                             | 13.3          | 12.7           | <0.001  |
| Others                               | 7.8           | 7.1            |         |
| Region, in %                         |               |                |         |
| Northeast                            | 23.1          | 14.9           | <0.001  |
| Midwest                              | 28.5          | 37.7           |         |
| South                                | 36.2          | 35.1           |         |
| West                                 | 12.2          | 12.3           |         |
| Comorbid substance use disorders     |               |                |         |
| Alcohol use disorder                 | 6.0           | 6.6            | <0.001  |
| Cannabis use disorder                | 16.4          | 15.3           | <0.001  |
| Stimulant use disorder               | 2.4           | 1.8            | <0.001  |
| Opioid use disorder                  | 2.6           | 1.6            | <0.001  |

TABLE 1: Descriptive characteristics of the study sample
SD: standard deviation

Females were more likely than males to be hospitalized for suicidal behaviors (OR 1.45; 95% CI 1.431-1.470). When compared with Caucasians, other races had a negative association with hospitalization for suicidal behaviors (P < 0.001). Likewise, compared to the northeast, the adolescent inpatients in the midwest (OR 1.98; 95% CI: 1.95-2.02), the west (OR 1.52; CI: 1.48-1.53), and the south (OR 1.51; CI: 1.48-1.53) had a significantly higher odds of hospitalization for suicidal behaviors.

Among SUD, only alcohol had significantly increased odds of association with hospitalization for suicidal behaviors (OR 1.18; CI: 1.142-1.209). Among other substances (cannabis and stimulants) there existed a statistically non-significant association with hospitalization for suicidal behaviors as shown in Table 2.
TABLE 2: Logistic regression analysis of odds ratio for suicidal behaviors in adolescent inpatients

| Variable                     | Odds ratio | 95% Confidence interval | P-value |
|------------------------------|------------|--------------------------|---------|
| Mean age                     | 0.99       | 0.990–0.997              | <0.001  |
| Sex                          |            |                          |         |
| Male                         | Reference  |                          |         |
| Female                       | 1.45       | 1.431–1.470              | <0.001  |
| Race, in %                   |            |                          |         |
| Caucasian                    | Reference  |                          |         |
| Black                        | 0.75       | 0.730–0.760              | <0.001  |
| Hispanic                     | 0.94       | 0.917–0.954              |         |
| Others                       | 0.87       | 0.852–0.896              |         |
| Region                       |            |                          |         |
| Northeast                    | Reference  |                          |         |
| Midwest                      | 1.98       | 1.95–2.02                | <0.001  |
| South                        | 1.51       | 1.48–1.53                |         |
| West                         | 1.52       | 1.48–1.53                |         |
| Comorbid substance use disorders |          |                          |         |
| Alcohol use disorder         | 1.18       | 1.142–1.209              | <0.001  |
| Cannabis use disorder        | 0.98       | 0.958–0.996              | 0.017   |
| Stimulant use disorder       | 0.92       | 0.855–0.962              | 0.013   |
| Opioid use disorder          | 0.66       | 0.628–0.692              | <0.001  |

Discussion

Suicidal behavior is a significant public health issue in the adolescent populations in the US. Suicidal behaviors encompass a continuum, ranging from suicidal ideation or thoughts of suicide to planning for suicide, suicide attempts, and death by suicide [16]. The mechanisms underlying suicidal behaviors are complex and heterogeneous, with multiple etiologies [17].

Our study provided an epidemiological view of distinct at-risk subgroups within this population, as well as identified at-risk subgroups with suicide risk among inpatient adolescents. We found that suicidal behaviors are more common in white females. Evidence suggests an increased rate of depression among females leading to increased suicidality [18]. The discrepancy could be due to various biological and psychosocial disparities in females. While female counterparts experience variable adverse stressors, their temperament and response to these stressors play an integral part in the development of depression [19].

Our findings are consistent with trends that show the geographic disparity in adolescent suicidality with higher prevalence in the midwest, the south, and the western regions of the US. There exists a correlation between suicide rates, and availability and utilization of psychiatric care and barriers to access psychiatric care, and regional differences in the state mental health parity laws. Also, there exist differences in psychiatric care resources and economic differences between urban and rural populations [20].

Our study found that alcohol use disorders significantly increased the odds of hospitalization for suicidal behaviors in adolescents. These results support a previous study by Swahn and Robert that has found that preteen alcohol use initiation was associated with suicidal ideation (increased by 1.89 times) and suicide attempts (increased by 2.71 times) relative to nondrinkers for both boys and girls [21]. An increased rate of suicides is seen in the contiguous post-consumption period and long-term use of alcohol. The theory is that alcohol could induce acute aggression and impulsivity during the post-consumption period, leading to
increased suicidal behaviors. And in the long term, alcohol consumption could cause biological changes and negative psychosocial events that could potentiate pre-existing issues influencing suicidal behaviors [22]. These deficits are more pronounced among youth who have used alcohol earlier in their life [23].

Although the prevalence of cannabis use disorder was highest among the suicidal inpatients, our study found no significant association between cannabis use and suicidal behaviors. A recent review and meta-analysis discovered a lack of consistent evidence linking acute cannabis use to increased risk for suicidality [24]. Evidence supports that chronic cannabis use can lead to suicidal behaviors, but the lack of systemic control for known risk factors tempers this finding. Our study demonstrated lower odds of opioid use in association with suicidal behaviors. However, as per the study by Wilkins et al., any prescription opioid misuse (current and past misuse) is associated with increased students’ risk of suicide-related behaviors but our study did not determine any causal association [25].

Our study had limitations. To begin with, we used administrative data that lacked patient-level clinical information, and inpatients’ diagnoses were based on the diagnostic codes, which may cause underreporting of comorbidities. Furthermore, because we conducted a cross-sectional study, we were unable to identify a causal relationship between suicidal behaviors and SUD. Despite this, the use of NIS offered an incomparable population-based perception of disease associations with systematic and temporal factors. Given the large sample size with the national representation of the population, results were appropriate for generalizability for the inpatient population.

Conclusions
It is essential to understand the impact of SUD on suicidal behaviors among adolescents. Adolescent inpatients with comorbid alcohol use disorders were at 18% higher odds of being hospitalized for suicidal behaviors. Our study provides a basis for more research while also suggesting potential avenues for early identification and intervention efforts for comorbid SUD in adolescents requiring psychiatric care to improve their prognosis and quality of life.

Additional Information

Disclosures

Human subjects: Consent was obtained or waived by all participants in this study. NA issued approval NA. The NIS is publicly available de-identified data with the protection of patients, physicians, and hospital-related information; henceforward, we were not required to obtain institutional review board approval for our study. Animal subjects: All authors have confirmed that this study did not involve animal subjects or tissue. Conflicts of interest: In compliance with the ICMJE uniform disclosure form, all authors declare the following: Payment/services info: All authors declare that no financial support was received from any organization for the submitted work. Financial relationships: All authors have declared that they have no financial relationships at present or within the previous three years with any organizations that might have an interest in the submitted work. Other relationships: All authors have declared that there are no other relationships or activities that could appear to have influenced the submitted work.

References

1. Diagnosis and treatment of drug abuse in family practice - american family physician monograph . (2003). Accessed: April 20, 2021: https://archives.drugabuse.gov/publications/diagnosis-treatment-drug-abuse-in-family-practice-american-family-physici...
2. Teen substance use & risks . (2020). Accessed: June 5, 2021: https://www.cdc.gov/ncbddd/fasd/features/teen-substance-use.html.
3. Youth risk behavior survey data summary & trends report: 2009-2019 . (2019). Accessed: April 20, 2021: https://www.cdc.gov/healthyyouth/data/yrbs/pdf/YRBSDataSummaryTrendsReport2019-508.pdf.
4. Principles of adolescent substance use disorder treatment: a research-based guide . (2021). Accessed: May 19, 2021: https://www.drugabuse.gov/publications/principles-adolescent-substance-use-disorder-treatment-research-based-guide/in...
5. Fowler JS, Volkow ND, Kasted CA, Chang I: Imaging the addicted human brain. Sci Pract Perspect. 2007, 3:4-16. 10.1151/spp07324
6. Wagner FA, Anthony JC: From first drug use to drug dependence; developmental periods of risk for dependence upon marijuana, cocaine, and alcohol. Neuropsychopharmacology. 2002, 26:479-88. 10.1016/S0895-1541(01)00367-0
7. Woltzky-Taylor KB, Ruggiero KJ, McCart MR, et al.: Has adolescent suicidality decreased in the United States? Data from two national samples of adolescents interviewed in 1995 and 2003. J Clin Child Adolesc Psychol. 2010, 39:64-76. 10.1080/15374410903401146
8. McKinnon B, Gariépy G, Sentenac M, Elgar FJ: Adolescent suicidal behaviours in 32 low- and middle-income countries. Bull World Health Organ. 2016, 94:540-50F. 10.2471/BLT.15.163295
9. Patel RS, Onyeaka H, Youssef NA: Suicidal ideation and attempts in unipolar versus bipolar depression: analysis of 131,740 adolescent inpatients nationwide. Psychiatry Res. 2020, 291:113231. 10.1016/j.psychres.2020.113231
10. Mathialagan K, Ceren Amuk O, Fikander N, Patel RS: Comorbid anxiety and suicidal behaviors in American adolescents with major depression. Cureus. 2020, 12:e8598. 10.7759/cureus.8598
11. Eskander N, Vadukapuram R, Zahid S, Ashraf S, Patel RS: Post-traumatic stress disorder and suicidal behaviors in American adolescents: analysis of 159,500 psychiatric hospitalizations. Cureus. 2020, 12:e8017. 10.7759/cureus.8017
12. Zahid S, Bodicherla KP, Eskander N, Patel RS: Attention-deficit/hyperactivity disorder and suicidal risk in major depression: analysis of 141,530 adolescent hospitalizations. Cureus. 2020, 12:e7949. 10.7759/cureus.7949
13. Amuk OC, Patel RS: Comorbid anxiety increases suicidal risk in bipolar depression: analysis of 9720 adolescent inpatients. Behav Sci (Basel). 2020, 10:108. 10.3390/bs10070108
14. Overview of the national (nationwide) inpatient sample. (2021). Accessed: April 20, 2021: https://www.hcup-us.ahrq.gov/nisoverview.jsp.
15. NIS description of data elements. (2021). Accessed: April 20, 2021: https://www.hcup-us.ahrq.gov/db/nation/nis/nisdde.jsp.
16. Hirsch JK, Ellis JR: Differences in life stress and reasons for living among college student suicide ideators and non-ideators. Coll Stud J. 1996, 30:577-86.
17. Bernanke J, Galfalvy HC, Mortali MG, et al.: Suicidal ideation and behavior in institutions of higher learning: a latent class analysis. J Psychiatr Res. 2017, 95:253-9. 10.1016/j.jpsychires.2017.09.003
18. Eisenlohr-Moul TA, Miller AB, Giletta M, Hastings PD, Rudolph KD, Nock MK, Prinstein MJ: HPA axis response and psychosocial stress as interactive predictors of suicidal ideation and behavior in adolescent females: a multilevel diathesis-stress framework. Neuropsychopharmacology. 2018, 43:2564-71. 10.1038/s41386-018-0206-6
19. Lee EJ, Jeong S: Analyzing gender differences in factors affecting depression among multicultural adolescents in South Korea: a cross-sectional study. Int J Environ Res Public Health. 2021, 18:3683. 10.3390/ijerph18073683
20. The relationship between mental health care access and suicide. (2018). Accessed: June 6, 2021: https://www.rand.org/research/gun-policy/analysis/essays/mental-health-access-and-suicide.html.
21. Swahn MH, Bossarte RM: Gender, early alcohol use, and suicide ideation and attempts: findings from the 2005 youth risk behavior survey. J Adolesc Health. 2007, 41:175-81. 10.1016/j.jadohealth.2007.05.005
22. Bagge CL, Sher KJ: Adolescent alcohol involvement and suicide attempts: toward the development of a conceptual framework. Clin Psychol Rev. 2008, 28:1283-96. 10.1016/j.cpr.2008.06.002
23. Cherpitel CJ, Borges GL, Wilcox HC: Acute alcohol use and suicidal behavior: a review of the literature. Alcohol Clin Exp Res. 2004, 28:185-28S. 10.1097/01.acl.0000127411.61654.14
24. Borges G, Bagge CL, Orozco R: A literature review and meta-analyses of cannabis use and suicidality. J Affect Disord. 2016, 195:63-74. 10.1016/j.jad.2016.02.007
25. Wilkins NJ, Clayton H, Jones CM, Brown M: Current prescription opioid misuse and suicide risk behaviors among high school students [PREPRINT]. Pediatrics. 2021, e2020030601. 10.1542/peds.2020-030601