# **ORIGINAL ARTICLE**

# Nonprescribed use of tranquilizers and use of other drugs among Brazilian students

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**Objectives:** To describe patterns of nonprescribed use of tranquilizers by students aged 10 to 18 years and assess the sociodemographic characteristics of these adolescents and their use of other substances.

**Methods:** A randomized and stratified sample of 47,979 students from state and private schools of the 27 Brazilian state capitals completed a self-report questionnaire. Poisson regression was used to estimate the associations between tranquilizer use and sociodemographic factors, as well as the use of other psychotropic substances.

**Results:** The lifetime prevalence of nonprescribed use of tranquilizers was 3.9%. Use was most common among girls, wealthier adolescents, and those from private schools. An association was found between use of tranquilizers and lifetime use of alcohol (prevalence ratio [PR] = 3.15; 95% confidence intervals [95%CI] 2.58-3.85), tobacco (PR = 2.61; 95%CI 2.31-2.95), illicit drugs (PR = 3.70; 95%CI 3.19-4.29), and other prescription drugs (PR = 7.03; 95%CI 6.18-7.99). As the number of substances adolescents reported having used increased, so did the nonprescribed use of tranquilizers.

**Conclusions:** Nonprescribed use of tranquilizers by adolescents might indicate the use of other substances, including high-risk combinations such as tranquilizers and alcohol. The risks of this association should be addressed during the early stages of drug prevention programs.

**Keywords:** Adolescents; antianxiety agents; epidemiology; gender differences; psychoactive substance use disorder

# Introduction

Nonprescribed use of prescription drugs – use without a physician's prescription, even when the intended use is self-medication – has been increasing among youths worldwide, promoted both by ease of access and by the false perception that medicines are completely safe.<sup>1-4</sup> The prevalence of this behavior varies among countries and age groups, and between genders. A recent systematic review reported the lifetime prevalence of such use by drug class among adolescents in the U.S. The most commonly used medicines were opioids (13-18%), followed by stimulants, including amphetamines (approximately 5%), and sedatives (approximately 4%). Most studies on the use of tranquilizers or anxiolytics have failed to examine lifetime use rates; however, past-year use rates range from 2 to 4.9% among individuals.<sup>5</sup>

The use of tranquilizers in Brazil gained attention during the late 1980s due to the suggestion that part of the population was abusing anxiolytics. Most such drugs are of the benzodiazepine class, and analyses of medical prescription records in pharmacies have shown that these drugs are the psychotropic agents most often prescribed

by medical professionals in Brazil.<sup>7</sup> Recent research has indicated that tranquilizers are used over much longer periods than recommended and might cause dependence.<sup>8</sup>

Previous studies of Brazilian school populations indicate that tranquilizers are the psychotropic agents most commonly implicated in nonprescribed use among adolescents, unlike in the U.S., where opioids are much more common. A nationwide survey of Brazilian state schools found that 4.1% of participants had ever used nonprescribed tranquilizers, whereas only 0.3% had used opioids. A study using a similar methodology found an even higher prevalence (7.7%) in private and state schools in the municipality of Passo Fundo, state of Rio Grande do Sul. Carantee in the province of the provin

Despite growing use of these drugs without a prescription, there is a shortage of information from developing countries. In a large pharmaceutical market such as Brazil, knowledge of the access of vulnerable populations – including adolescents – to these substances is important, given the failures associated with controlling psychotropic medications and the high rate of self-medication. <sup>11,12</sup>

Within this context, this study sought to: 1) estimate the prevalence of nonprescribed use of tranquilizers among 10-to-18-year-old students attending the primary and secondary schools of the state and private educational systems in all 27 state capitals of Brazil; 2) identify the

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most vulnerable adolescents by estimating the associations between sociodemographic characteristics and nonprescribed use of these substances; and 3) assess whether nonprescribed use of tranquilizers by this population is associated with use of other psychotropic substances.

#### **Methods**

#### Sample

This cross-sectional study was conducted across all 27 Brazilian state capitals by the Brazilian Center for Information on Psychotropic Drugs (CEBRID), with funding from the National Secretariat for Drugs Policies (SENAD), between March and October 2010. A selfreport questionnaire was administered to state and private-school students in classrooms. The sample was recruited from a national registry of all Brazilian state and private schools (Instituto Nacional de Estudos e Pesquisas Educacionais Anísio Teixeira, INEP). Subsamples were drawn for each city, which generated 54 subsamples: 27 from state schools and 27 from private schools. Given the small number of private schools in Boa Vista, state of Roraima, and the refusal of a great number of private schools in the city of Rio de Janeiro to participate in the survey, data from both capitals were pooled into a single subsample for each city. Three strata were considered for each subsample: schools providing primary education only, schools providing both primary and secondary education, and schools providing secondary education only. A proportional sample was based on the number of students (estimated using the number of classrooms in each school) within each stratum and a randomized process used to select an average of three clusters (classrooms) in each school. Finally, all students in the selected classrooms were then invited to participate in the survey.

A total of 923 schools participated in this study: 545 state schools and 378 private schools. The average acceptance rate was 86% (94% among state schools and 73% among private schools). The student participation rate, which represents the number of students who participated in the survey divided by the number of students who officially enrolled in the classrooms drawn, was 83% (79% in state and 92% in private schools). Only 0.3% of students who were invited to participate in the survey refused to take part (0.2% from private and 0.4% from state schools). This process generated a sample of 51,154 guestionnaires. A dummy guestion concerning drug use was included, and all questionnaires with positive responses to that question were removed (n=263). As all analyses were restricted to students between the ages of 10 and 18, the final sample consisted of 47,979 students: 17,370 from secondary schools and 30.609 from primary schools.

#### Procedure

An investigator with experience in survey research supervised each Brazilian state capital. This group of

investigators completed a 3-day training session at Universidade Federal de São Paulo (UNIFESP), administered by CEBRID, and subsequently replicated this training for the team of field researchers (i.e., interviewers) in their states.

The questionnaires were administered in classroom to students who agreed to participate in the study. On average, the questionnaires were completed in 40 minutes, in the presence of a previously trained interviewer and in the absence of school officials. The procedures were standardized and performed on the same day for all classes selected from the same school to avoid confounders.

Written informed consent was obtained from the principal of each school clarifying that the adolescents' participation in the study was voluntary, and they would be free to discontinue their participation at any time. They were also notified that the information collected was anonymous, confidential, and intended for academic purposes only. The UNIFESP Research Ethics Committee approved this study with protocol no. 0348/08.

#### Instrument

The study questionnaire was based on a specific research instrument developed by the World Health Organization to examine student substance use. <sup>13</sup> This measure has been used in prior CEBRID surveys. <sup>9</sup> Additional questions were included on the basis of the European School Survey Project on Alcohol and Other Drugs (ESPAD). <sup>14</sup>

The questionnaire comprised the following sections:

#### Sociodemographic variables

Data concerning gender, age (open question), type of school (state or private), and country region were collected. The Brazilian Association of Research Companies (ABEP) scale of socioeconomic status was applied. This assessment collects data including the level of education of the head of the household, ownership of household goods (including a TV, DVD player, cars, refrigerators, and so on), and the number of domestic workers in the household (maid, driver, gardener, and so on). The ABEP scale classifies participants into the following social statuses, in descending order: A1, A2, B1, B2, C1, C2, D, and E. Participants were subsequently categorized as lower (C, D, and E), middle (B), or upper (A) socioeconomic class for the purposes of the present analysis.

#### Nonprescribed use of tranquilizers

Questions concerning the nonprescribed use of tranquilizers were included. Questions regarding lifetime use, past-year use, and past-month use were answered with "yes" or "no". The medicines mentioned as examples were diazepam (and its trade names Dienpax® and Valium®), Lorax® (lorazepam), Rohypnol® (flunitrazepam), Psicosedin® (chlordiazepoxide), Rivotril® (clonazepam), alprazolam (and its trade names Apraz® and Frontal®),

Dalmadorm® (flurazepam), Dormonid® (midazolam), bromazepam (and its trade names Lexotan® and Somalium®), and Olcadil® (cloxazolam). The questionnaire noted that intake of teas and naturally occurring substances, including valerian, should not be considered part of this category. A question regarding frequency of use was designed for past-month users, with possible answers ranging from 1 to 5 days (sporadic use), 6 to 19 days (frequent use), and 20 days or more (heavy use). Age at first use was also examined in the form of an openended question, as was the name of the last used medicine. This variable considered generic and trade names, which were converted to the generic nomenclature after grouping.

#### Use of other substances

Several questions assessed the use of other psychoactive substances. In the present analysis, we collected data on lifetime use of alcohol, tobacco, illicit drugs (cannabis, cocaine/crack, or ecstasy), and nonprescribed use of other prescription drugs (benzydamine, anabolic steroids, amphetamines, or opiates). A variable, number of drugs, was constructed by adding the lifetime use of different drugs, except for alcohol and tobacco, by the respondent.

#### Data analyses

All analyses were performed using STATA version 11. The significance level was set at 5%. Prevalence was expressed with regard to the sample weight using confidence intervals. Student's *t*-test was used to examine the mean differences in participant age at the first use of tranquilizers by gender.

A Poisson regression model was constructed to estimate the unadjusted associations between tranquilizer use and age, gender, level of education, type of school (state vs. private), socioeconomic status, and country region. These associations were subsequently estimated using a fitted model. Level of education was excluded from this model because it was collinear with age, and socioeconomic status was excluded because it was collinear with type of school. Other Poisson regression models were constructed to estimate the associations between tranquilizer use and lifetime use of alcohol, tobacco, illicit drugs (cannabis, cocaine/crack, or ecstasy), other prescription drugs (benzydamine, anabolic steroids, amphetamines, or opiates), and the number of substances previously used. These models were fit using gender, age, and type of school. The results were expressed as prevalence ratios with 95% confidence intervals and p-values.

#### Results

### Sample characteristics

Table 1 describes the sample profile (n=47,979). Overall, 62.1% of participants were from primary schools, and 78.2% were from state schools. The largest age group

was 13-to-15-year-olds (45.2%), and females made up the majority of the sample (52.2%).

The distribution of the weighted proportion of students per region was based on representations per capital. Thus, although the absolute number of participants from the Southeast region of Brazil was not the largest, this region comprised the largest proportion of students (42.8%). The vast majority of students were from low-income and middle-class socioeconomic backgrounds (64.3%). Many respondents were grouped into a single "not reported" category (26.5%) because they failed to provide all the data required to assess their socioeconomic status.

# Lifetime use of tranquilizers and associated sociodemographic factors

Of the total students interviewed, 2,124 (3.9%) reported a lifetime nonprescribed use of tranquilizers (95%Cl 3.7-4.2). Students from the oldest age group were most likely to report this use (Table 1): the probability of a student reporting lifetime nonprescribed use of tranquilizers increased by 15% with each year of life. This use was also higher among secondary school students (PR = 1.55; 95%Cl 1.35-1.78) as compared with primary school students. However, age and level of education were collinear in the fitted model. Girls showed a higher prevalence of tranquilizer use, and were two times more likely to use than boys (95%CI 1.76-2.26). An association was found between lifetime nonprescribed use of tranquilizers and highest socioeconomic status. This association was noted in the bivariate analysis with regard to both a higher prevalence of users among students from private schools as compared with those from state schools (PR = 1.92; 95%CI 1.69-2.18) and an increasing prevalence ratio found among the middle and upper socioeconomic statuses as compared with the lower status (PR = 1.63; 95%CI 1.40-1.89 and PR = 2.52; 95%CI 2.12-3.00, respectively). Private-school students were more than twice as likely to report using tranquilizers as public school students in the fitted model. Lastly, living in the South and Center-West regions was associated with use as compared with the Northeast region after adjusting for age, gender, and type of school (PR = 1.50; 95%CI 1.28-1.77 and PR = 1.18; 95%CI 1.01-1.37, respectively).

Figure 1 shows the prevalence of lifetime nonprescribed tranquilizer use by age group and gender. As shown above, the proportion of users increases by age, although this increase was most significant among female adolescents.

Figure 2 shows the lifetime prevalence of nonprescribed tranquilizer use per Brazilian state capital by type of school (state or private). The highest prevalence was recorded at private schools in Belo Horizonte, state of Minas Gerais (9%), and the lowest rate was recorded at state schools in Belém, state of Pará (2%). The prevalence of use in private schools was higher than in state schools for most Brazilian state capitals, except Goiânia (state of Goiás) and Macapá (state of Amapá).

**Table 1** Sociodemographic characteristics of the total sample and according to lifetime use of nonprescribed tranquilizers and the prevalence rate by Poisson regression

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	n	Total (n=47,979) wgt% (95%CI)	n	Lifetime use of tranquilizers (n=2,124) wgt% (95%CI)	PR1 (95%CI)	p-value	PR2 (95%CI)	p-value
Age (years)* 10-12 13-15 16-18 Missing	13,785 21,807 12,387	27.5 (25.8-29.2) 45.2 (43.7-46.7) 27.4 (25.6-29.1)	324 1,037 763 150	2.2 (1.8-2.6) 4.1 (3.7-4.5) 5.3 (4.8-5.9)	1.15 (1.12-1.19)	< 0.001	1.17 (1.14-1.21)	< 0.001
Gender Male Female Missing	22,813 25,001 165	47.8 (47.2-48.5) 52.2 (51.5-52.8)	686 1,432 22	2.6 (2.3-2.9) 5.1 (4.7-5.5)	1.00 1.97 (1.74-2.23)	< 0.001	1.00 2.00 (1.77-2.27)	< 0.001
Level of education Primary Secondary	30,609 17,370	62.1 (59.8-64.3) 37.9 (35.7-40.2)	,	3.2 (2.9-3.6) 5.0 (4.6-5.5)	1.00 1.55 (1.35-1.78)	< 0.001	-	
Type of school State Private	28,759 19,220	78.2 (76.6-79.7) 21.8 (20.3-23.4)	990 1,134	3.3 (3.0-3.5) 6.2 (5.7-6.9)	1.00 1.92 (1.69-2.18)	< 0.001	2.09 (1.85-2.37)	< 0.001
Socioeconomic status Lower (C-E) Middle (B) Upper (A) Not informed	13,750 15,708 6,796 11,725	33.2 (32.2-34.2) 31.1 (30.0-32.2) 9.2 (8.2-10.3) 26.5 (25.6-27.4)	456 822 514 332	3.0 (2.7-3.4) 4.9 (4.5-5.4) 7.6 (6.7-8.7) 2.5 (2.1-2.9)	1.00 1.63 (1.40-1.89) 2.52 (2.12-3.00) 0.83 (0.68-1.00)	< 0.001 < 0.001 0.056	-	
Country region Northeast North Midwest South Southeast	16,503 10,619 7,486 5,055 8,316	26.2 (24.8-27.6) 11.8 (10.8-12.9) 11.7 (10.7-12.7) 7.6 (6.8-8.3) 42.8 (40.9-44.7)	725 444 353 274 328	4.0 (3.7-4.4) 3.7 (3.2-4.2) 4.5 (4.0-5.2) 5.5 (4.8-6.2) 3.4 (3.0-4.0)	1.00 0.90 (0.77-1.06) 1.12 (0.96-1.32) 1.35 (1.15-1.58) 0.85 (0.71-1.00)	0.224 0.156 < 0.001 0.064	1.00 1.03 (0.88-1.21) 1.18 (1.01-1.37) 1.50 (1.28-1.75) 0.94 (0.80-1.11)	0.684 0.028 < 0.001 0.473

95%CI = 95% confidence interval; PR1 = prevalence ratio on bivariate analysis; PR2 = prevalence ratio adjusted by age, gender, and type of school (level of education was excluded from this model because it was collinear with age, and socioeconomic status was excluded because it was collinear with type of school); wgt% = weighted proportions.

The overall lifetime prevalence of nonprescribed use of tranquilizers was 3.3% (95%Cl 3.0-3.6) in state schools and 6.3% (95%Cl 5.7-6.9) in private schools.

# Patterns of tranquilizer use

Of the 2,124 adolescents who reported lifetime use of tranquilizers, 1,403 (65.7%) reported past-year use (95%CI 63.0-68.0). However, only 658 adolescents (32.7%) indicated past-month use (95%Cl 29.8-35.7), and the frequency of this use suggested that most of these adolescents were not heavy users. Of the 658 students who reported past-month use, 534 (81.0%) reported sporadic use (95%Cl 76.0-85.0), whereas 51 adolescents (9.0%) reported frequent use (95%Cl 6.4-12.0), and 73 participants (10.1%) were heavy users (95%CI 7.3-13.7). The mean age at first use was 13.2 years (standard error [SE] = 0.06). Although boys were less likely to use tranquilizers than girls, their first use tended to be at a younger age (12.8 vs. 13.5 years; p < 0.001). The tranquilizer most frequently used by adolescents was diazepam, which accounted for approximately 60% of the reports. The other medicines cited and their respective proportions are shown in Figure 3.

Association between lifetime use of tranquilizers and use of other substances

Lifetime tranquilizer use was associated with lifetime use of other substances, both legal and illicit (Table 2). The

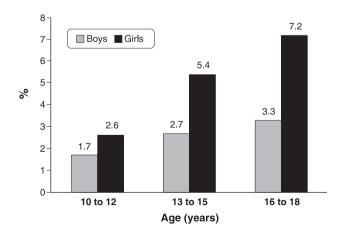
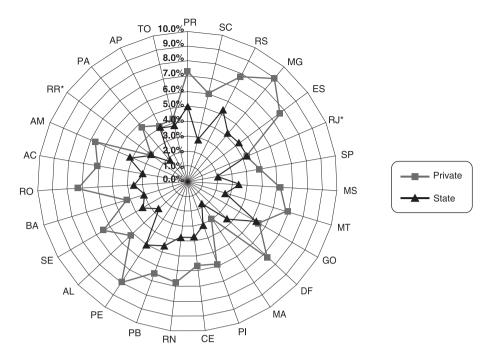


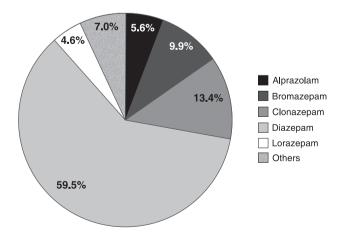
Figure 1 Lifetime nonprescribed use of tranquilizers by age range, according to gender (n=47,979)

<sup>\*</sup> Age was used as a continuous variable in the regression model.



**Figure 2** Lifetime nonprescribed use of tranquilizers by state capital according to type of school (n=47,979). \* RJ and RR had data from private and state schools analyzed together.

lifetime prevalence of alcohol use was three times higher among adolescents who reported having used tranquilizers compared with non-users, and the lifetime prevalence of tobacco use was 2.6 times higher, even after accounting for the major sociodemographic variables associated with tranquilizer use (i.e., age, gender, and type of school). Tranquilizer use was four times greater (PR = 3.70; 95%CI 3.19-4.29) among adolescents who reported using at least one illicit drug (cannabis, cocaine/crack, or ecstasy). Similarly, students who reported nonprescribed use of other prescription drugs were most likely to use tranquilizers with regard to the variables examined (PR = 7.03; 95%CI 6.18-7.99). The number of substances used by adolescents was also associated



**Figure 3** Benzodiazepines most commonly used by adolescents without a medical prescription as self-reported (n=2,124)

with tranquilizer use: as the number of substances reportedly used increased, so did the prevalence of nonprescribed tranquilizer use.

#### **Discussion**

Overall, lifetime nonprescribed use of tranquilizers was reported by 3.9% of adolescents in this sample (3.3% of students from state schools and 6.3% of those from private schools). This estimated prevalence is very similar to that found in a survey conducted 6 years earlier in state schools only 9 (3.8% among students aged 10 to 18 years).

However, the prevalence in private schools (6.3%) was found to be twice as high as that found in state schools. This estimate was similar to the overall prevalence found in 31 European countries (5.6%), ranging from 1.5% in the Ukraine to 13.6% in Lithuania. Likewise, the last edition of Monitoring the Future, a survey conducted in U.S. schools, reported a 6.3% prevalence, similar to that found among Brazilian private-school students. <sup>17</sup>

The findings of this study also suggest that the lifetime prevalence of nonprescribed tranquilizer use among students from private schools in Brazil is similar to the prevalence found among Brazilian adults. Nonprescribed use of tranquilizers was examined in a representative nationwide sample of the 107 largest Brazilian cities via household surveys of 12-to-65-year-olds in 2005. Nearly one in six respondents (5.6%) reported nonprescribed use of tranquilizers at some time during their lives. However, such comparisons should be interpreted with caution, as the methods employed to ascertain nonprescribed use differed between the two surveys.

**Table 2** Lifetime use of other substances, overall and according to lifetime nonprescribed use of tranquilizers, and prevalence ratios by Poisson regression

	n	Total (n=47,979) wgt% (95%Cl)	n	Lifetime use of tranquilizers (n=2,124) wgt% (95%CI)	PR (95%CI)	p-value
Alcohol						
No Yes	19,557 28,308	40.4 (39.1-41.7) 59.6 (58.3-60.9)	332 1,788	1.5 (1.2-1.7) 5.6 (5.2-5.9)	1.00 3.15 (2.58-3.85)	< 0.001
Missing	114	39.0 (38.3-00.9)	4	5.0 (5.2-5.9)	3.13 (2.36-3.63)	< 0.001
Tobacco						
No	40,746	84.1 (83.3-84.8)	1,412	3.0 (2.7-3.2)	1.00	
Yes Missing	7,206 27	15.9 (15.2-16.7)	711	8.7 (7.9-9.6)	2.61 (2.31-2.95)	< 0.001
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Illicit drugs*	45.040	04.0 (04.4.05.0)	4 700	0.0 (0.1.0.5)	4.00	
No Yes	45,018 2,743	94.8 (94.4-95.2) 5.2 (4.8-5.6)	1,733 388	3.3 (3.1-3.5) 13.1 (11.6-14.8)	1.00 3.70 (3.19-4.29)	< 0.001
Missing	218	5.2 (4.6-5.0)	3	13.1 (11.0-14.8)	3.70 (3.19-4.29)	< 0.001
Prescription drugs						
No	45,576	96.2 (95.9-96.5)	1,583	3.1 (2.9-3.3)	1.00	
Yes	2,083	3.8 (3.5-4.1)	525	24.6 (22.1-27.2)	7.03 (6.18-7.99)	< 0.001
Missing	320		3			
Number of substances <sup>‡</sup>						
None	43,342	91.3 (90.8-91.7)	1,378	2.8 (2.6-3.0)	1.00	
One	2,910	5.9 (5.5-6.2)	431	12.2 (10.8-13.8)	4.03 (3.46-4.71)	< 0.001
Two	868	1.9 (1.8-2.2)	160	16.8 (13.6-20.7)	6.10 (4.84-7.68)	< 0.001
Three or more Missing	408 451	0.9 (0.8-1.0)	130 25	31.7 (25.1-39.2)	10.92 (8.67-13.76)	< 0.001

95%CI = 95% confidence interval; PR = prevalence ratio adjusted by sociodemographic factors (age, gender, and type of school); wgt% = weighted proportions.

This use was higher among older respondents, which corroborates other studies of school-age adolescents. <sup>19-21</sup> However, the cumulative effect of the question concerning lifetime use might explain this finding.

The scientific literature has established that women are more likely to use psychotropic medications recreationally as compared with men, most likely because they visit medical services more often and have a higher prevalence of anxiety symptoms. <sup>22,23</sup> In the present study, the prevalence of use among adolescent girls was almost twice that observed in boys across all age groups. Another factor associated with higher use among girls is their more frequent sharing of prescriptions and medications. <sup>24</sup>

In contrast to the international literature, in which low income is generally a risk factor for drug use, <sup>25,26</sup> Brazilian studies have found a higher use of certain drugs — especially alcohol — among socioeconomic statuses with greater purchasing power. <sup>27,28</sup> A survey of psychotropic medication use conducted in Passo Fundo, state of Rio Grande do Sul, found that the prevalence of tranquilizer use was 1.3 times higher among private schools. <sup>10</sup> A possible hypothesis to explain this finding is that parents' higher purchasing power increases access to healthcare services and, consequently, increases adolescents' access to medications, which are often within reach in their own homes. A culture of tranquilizer use might also exist among adolescents who interact in

the same social circles (e.g., private schools). The greater use observed in private schools as compared with state schools was evident in approximately all Brazilian state capitals, despite the discrepancy between the overall prevalence of use among state capitals. The regression model indicated a higher rate of tranquilizer use in the South and Center-West regions compared with the Northeast region. Possible sample differences in purchasing power might explain this finding; however, additional research is required to better understand this phenomenon.

The present study also found an association between nonprescribed use of tranquilizers and use of other substances, both legal and illicit. American adolescents who reported using sedatives or tranquilizers recreationally were approximately eight times more likely to report a past-year use of illicit drugs than non-users. 19 In a study of 18-to-24-year-olds, past-year tranquilizer use was strongly associated with patterns of alcohol abuse.<sup>29</sup> Another study indicated that lifetime nonprescribed use of prescription medications was associated with a greater likelihood of using illicit drugs and binge drinking (five or more drinks on a single occasion) over the past year among university students. These findings were more significant when analyzing the use of tranquilizers for recreational purposes (illicit drugs odds ratio [OR] = 15.7, p < 0.001; binge drinking OR = 12.2, p < 0.01).30

<sup>\*</sup> Illicit drugs: cannabis, cocaine/crack, or ecstasy.

<sup>†</sup> Prescription drugs: benzydamine, anabolic steroids, amphetamines, or opioids.

<sup>\*</sup> Only illicit drugs and nonprescribed use of prescription drugs.

Adolescents who reported using prescription benzodiazepines recreationally might have more anxiety or other mental disorder symptoms, including those associated with social phobia. The self-medication hypothesis of addictive disorders<sup>31</sup> suggests that adolescent interest in self-medication might explain their use of other substances. Importantly, substance abuse is a multifactorial phenomenon that encompasses a range of biological, psychological, and social factors. Thus, the factors that promote an adolescent's first use of a substance might be the same factors that induce nonprescribed use of prescription anxiolytics.

Although the cross-sectional nature of our study does not allow us to establish causality, another possible hypothesis for this behavior is based on the social learning theory. This theory postulates that behaviors are learned and reproduced by observing others (i.e., social models).<sup>32</sup> Peralta & Steele<sup>33</sup> concluded that social learning theory at least partially explained the nonprescribed use of psychotropics in a sample of university students.

Nonprescribed use of prescription medicines by adolescents might be affected by their observation of adults who use these medications, including their reasons for use (e.g., before a job interview or after an argument with their partner). Observing adults who use tranquilizers for reasons related to daily life might induce adolescents to seek relief from their own discomfort via prescription drugs. This behavior is potentially harmful to the development of adolescents, who are still learning their limits and potentials.

In Brazil, several factors appear to create a context conducive to the inappropriate use of medications. One such strong effect is the influence of the pharmaceutical industry<sup>34</sup> on the prescriber/user/seller triad, which incorporates a culture of medication reinforced by the symbolic power of medicine.<sup>35</sup> This concept might be assimilated as a fanciful belief that medicines are unlikely to cause harm because they are produced with stringent quality controls by a pharmaceutical company and used by patients under the scrutiny of licensed physicians. This reasoning might explain why our findings revealed a stronger association between the use of tranquilizers and that of other medicines with potential for abuse.

Certain limitations should be considered. Crosssectional surveys are designed to collect data about outcome and exposure at the same time; therefore, the association of factors found in this study does not imply causality. The sample is representative of students from Brazilian state capitals: therefore, extrapolation of these findings to adolescents who do not attend school or who live in cities with different characteristics should be done with caution. Seventeen per cent of students were absent on the day of data collection. Previous studies have shown that absent students are more likely to use drugs<sup>36,37</sup>; if this is true for tranquilizers, we might have underestimated the nonprescribed use among Brazilian students. Despite being the most costeffective epidemiological study method for large samples, the self-report method is subject to under- and over-reporting.<sup>38</sup> The use of a dummy question regarding drug use minimized over-reporting, and questionnaires were excluded from the analysis for this reason. However, the strength of this survey was that it reported unpublished nationwide data concerning nonprescribed use of tranquilizers by adolescents. This survey had a good response rate and voluntary participation.

The assessment of nonprescribed use of tranquilizers by Brazilian adolescents provides key information from a public health standpoint. Moreover, these data help prevent trivialization of this behavior by highlighting the risks that result from the chronic and acute use of anxiolytics in combination with other substances (e.g., alcohol). The analyses presented in this paper raised specific concerns that should be studied to prevent this behavior. First, although access to tranquilizers by adolescents was not assessed in this specific analysis, evidence suggests that nonprescribed use of tranquilizers is associated with family influence and previous medical prescription of tranquilizers.<sup>39</sup> Therefore, physicians should advise their patients about personal medical use and the risks associated with using substances without therapeutic monitoring so as to reduce the availability of prescription drugs to adolescents. Pharmacists are responsible for exerting greater control over the sale of these medications; furthermore, guidance regarding the storage and proper disposal of psychotropic agents should be followed.

Another key point is the need to include specific information about medicines in drug prevention programs. These programs should expand their discourse, which currently focuses on drugs such as cannabis and cocaine/crack, to substances with less visibility but great relevance throughout life (i.e., tranquilizers). Prescription drugs immediately follow alcohol, tobacco, and solvents with regard to age at first use; therefore, addressing this content in the classroom is appropriate. Prevention programs should work with adolescents to search for strategies that might alleviate issues related to anxiety and insomnia without resorting to the use of medicines. In addition, they should provide information on the potential for dependence and the risks associated with nonprescribed use of tranquilizers.

Finally, periodic epidemiological surveys are essential to assess changes in user profiles over time and promote immediate health measures to reach this population.

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#### **Disclosure**

The authors report no conflicts of interest.

#### References

- 1 Manchikanti L. Prescription drug abuse: what is being done to address this new drug epidemic? Testimony before the Subcommittee on Criminal Justice, Drug Policy and Human Resources. Pain Physician. 2006;9:287-321.
- 2 Sung HE, Richter L, Vaughan R, Johnson PB, Thom B. Nonmedical use of prescription opioids among teenagers in the United States: trends and correlates. J Adolesc Health. 2005;37:44-51.
- 3 Johnston LD. Prescription drug use by adolescents: what we are learning and what we still need to know. J Adolesc Health. 2009;45:539-40.
- 4 Kuehn BM. Prescription drug abuse rises globally. JAMA. 2007;297:1306.
- 5 Young AM, Glover N, Havens JR. Nonmedical use of prescription medications among adolescents in the United States: a systematic review. J Adolesc Health. 2012;51:6-17.
- 6 Nappo S, Carlini EA. Preliminary finding: consumption of benzodiazepines in Brazil during the years 1988 and 1989. Drug Alcohol Depend. 1993;33:11-7.
- 7 Noto AR, Carlini EdA, Mastroianni PC, Alves VC, Galduróz JCF, Kuroiwa W, et al. Analysis of prescription and dispensation of psychotropic medications in two cities in the State of São Paulo, Brazil. Rev Bras Psiquiatr. 2002;24:68-73.
- 8 de Souza ARL, Opaleye ES, Noto AR. Contextos e padrões do uso indevido de benzodiazepínicos entre mulheres. Cien Saude Coletiva. 2013;18:1131-40.
- 9 Galduroz JC, Noto AR, Fonseca AM, Carlini EA. V Levantamento Nacional sobre o Consumo de Drogas Psicotrópicas entre Estudantes do Ensino Fundamental e Médio da Rede Pública de Ensino nas 27 Capitais Brasileiras. São Paulo: CEBRID; 2005.
- 10 Dal Pizzol Tda S, Branco MM, Carvalho RM, Pasqualotti A, Maciel EN, Migott AM. [Non-medical use of psychoactive medicines among elementary and high school students in Southern Brazil]. Cad Saude Publica. 2006;22:109-15.
- 11 Mastroianni Pde C, Lucchetta RC, Sarra Jdos R, Galduróz JC. Estoque doméstico e uso de medicamentos em uma população cadastrada na estratégia saúde da família no Brasil. Rev Panam Salud Publica. 2011:29:358-64.
- 12 Arrais PS, Coelho HL, Batista Mdo C, Carvalho ML, Righi RE, Arnau JM. [Profile of self-medication in Brazil]. Rev Saude Publica. 1997;31:71-7.
- 13 Smart RG, Hughes DPH, Johnston LD. Methodology for students drug-use surveys. Geneva: WHO; 1980.
- 14 Hibell S, Guttormsson U, Ahlström S, Balakireva O, Bjarnason T, Kokkevi A, et al. The 2007 ESPAD report substance use among students in 35 European Countries. Stockholm: The Swedish Council for Information on Alcohol and Other Drugs (CAN); 2009.
- 15 Associação Brasileira de Empresas de Pesquisa (ABEP). Critério de classificação econômica Brasil [Internet]. 2008 [cited 2013 Oct 30]. http:// www.abep.org/novo/Content.aspx?ContentID=302&SectionCode= CCER
- 16 Kokkevi A, Fotiou A, Arapaki A, Richardson C. Prevalence, patterns, and correlates of tranquilizer and sedative use among European adolescents. J Adolesc Health. 2008;43:584-92.
- 17 Johnston LD, O'Malley PM, Bachman JG, Schulenberg JE. Monitoring the future. National results on drug use. 2012 overview. Key findings on adolescent drug use. Ann Arbor: Institute for Social Research; 2013.
- 18 Fonseca AM, Galduroz JC, Noto AR, Carlini EL. Comparison between two household surveys on psychotropic drug use in Brazil: 2001 and 2004. Cien Saude Colet. 2010;15:663-70.

- 19 McCabe SE, Boyd CJ, Young A. Medical and nonmedical use of prescription drugs among secondary school students. J Adolesc Health. 2007;40:76-83.
- 20 Andersson B, Hibell B, Beck F, Choquet M, Kokkevi A, Fotiou A, et al. Alcohol and drug use among European 17-18 year old students -Data from the ESPAD Project. Sweden: Swedish Council for Information on Alcohol and Other Drugs (CAN). 2007.
- 21 Johnston LD, O'Malley PM, Bachman JG, Schulenberg JE. Monitoring the future. National results on adolescent drug use. Overview of key findings. 2009; Bethesda; : National Institute on Drug Abuse; 2010.
- 22 Simoni-Wastila L, Ritter G, Strickler G. Gender and other factors associated with the nonmedical use of abusable prescription drugs. Subst Use Misuse. 2004;39:1-23.
- 23 Simoni-Wastila L, Yang HW, Lawler J. Correlates of prescription drug nonmedical use and problem use by adolescents. J Addict Med. 2008;2:31-9.
- 24 Boyd CJ, McCabe SE, Cranford JA, Young A. Prescription drug abuse and diversion among adolescents in a southeast Michigan school district. Arch Pediatr Adolesc Med. 2007;161:276-81.
- 25 Goodman E, Huang B. Socioeconomic status, depressive symptoms, and adolescent substance use. Arch Pediatr Adolesc Med. 2002;156:448-53.
- 26 Baumann M, Spitz E, Guillemin F, Ravaud JF, Choquet M, Falissard B, et al. Associations of social and material deprivation with tobacco, alcohol, and psychotropic drug use, and gender: a population-based study. Int J Health Geogr. 2007;6:50.
- 27 Locatelli D, Sanchez Z, Opaleye E, Carlini C, Noto A. Socioeconomic influences on alcohol use patterns among private school students in Sao Paulo. Rev Bras Psiguiatr. 2012;34:193-200.
- 28 Pratta EM, dos Santos MA. Adolescence and the consumption of psychoactive substances: the impact of the socioeconomic status. Rev Lat Am Enfermagem. 2007;15:806-11.
- 29 McCabe SE, Cranford JA, Boyd CJ. The relationship between pastyear drinking behaviors and nonmedical use of prescription drugs: prevalence of co-occurrence in a national sample. Drug Alcohol Depend. 2006;84:281-8.
- 30 McCabe SE, Boyd CJ, Teter CJ. Subtypes of nonmedical prescription drug misuse. Drug Alcohol Depend. 2009;102:63-70.
- 31 Khantzian EJ. The self-medication hypothesis of addictive disorders: focus on heroin and cocaine dependence. Am J Psychiatry. 1985;142:1259-64.
- 32 Bandura A. Social learning theory. Englewood Cliffs: Prentice-Hall; 1977
- 33 Peralta RL, Steele JL. Nonmedical prescription drug use among US college students at a Midwest university: a partial test of social learning theory. Subst Use Misuse. 2010;45:865-87.
- 34 Mastroianni PC, Vaz AC, Noto AR, Galduroz JC. Psychoactive drug advertising: content analysis. Rev Saude Publica. 2008;42:968-71.
- 35 Barros JA. [New tendencies in medicalization]. Cien Saude Colet. 2008;13:579-87.
- 36 Tavares BF, Béria JU, Silva de Lima M. Prevalência do uso de drogas e desempenho escolar entre adolescentes. Rev Saude Publica. 2001;35:150-8.
- 37 Moura YG, Sanchez ZM, Opaleye ES, Neiva-Silva L, Koller SH, Noto AR. Drug use among street children and adolescents: what helps? Cad Saude Publica. 2012;28:1371-80.
- 38 Dunn J, Ferri CP. Epidemiological methods for research with drug misusers: review of methods for studying prevalence and morbidity. Rev Saude Publica. 1999;33:206-15.
- 39 Opaleye ES, Noto AR, Sanchez ZM, Amato TC, Locatelli DP, Gossop M, et al. Nonprescribed use of tranquilizers or sedatives by adolescents: a Brazilian national survey. BMC Public Health. 2013;13:499.