Benzodiazepine Use and Misuse Among Adults in the **United States**

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Objective: Goals were to determine the prevalence of benzodiazepine use (as prescribed and misuse), characterize misuse, and examine variation by age.

Methods: A cross-sectional analysis was conducted of 2015 and 2016 National Survey on Drug Use and Health data limited to adults ≥18 (N=86,186) and data from those respondents reporting benzodiazepine use (N=10,290). Measurements included past-year prescription benzodiazepine use and misuse ("any way a doctor did not direct"), substance use disorders, mental illness, and demographic characteristics. Misuse was compared between younger (18-49) and older (≥50) adults.

Results: A total of 30.6 million adults (12.6%) reported pastyear benzodiazepine use-25.3 million (10.4%) as prescribed and 5.3 million (2.2%) misuse. Misuse accounted for 17.2% of overall use. Adults ages 50-64 had the highest prescribed use (12.9%). Those ages 18-25 had the highest

misuse (5.2%), and those ages \geq 65 had the lowest (.6%). Misuse and abuse of or dependence on prescription opioids or stimulants were strongly associated with benzodiazepine misuse. Benzodiazepine misuse without a prescription was the most common type of misuse, and a friend or relative was the most common source. Adults ages ≥50 were more likely than younger adults to use a benzodiazepine more often than prescribed and to use a benzodiazepine to help with sleep.

Conclusions: Benzodiazepine use among U.S. adults was higher than previously reported, and misuse accounted for nearly 20% of use overall. Use by adults ages 50-64 now exceeds use by those ages ≥65. Patients also prescribed stimulants or opioids should be monitored for benzodiazepine misuse. Improved access to behavioral interventions for sleep or anxiety may reduce some misuse.

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Benzodiazepines are prescribed to over 5% of the U.S. adult population, and use is growing (1-3), concentrated among middle-aged adults, for whom use increased nearly 50% from 1996 to 2013 (1). However, the prevalence of benzodiazepine use among adults ages 65 and older has been shown to be highest, at 8.6% (1). Prescribing to older adults has been considered potentially inappropriate for decades, given associated harms, including falls and fractures (4-7); however, the growth in benzodiazepine prescribing has been accompanied by increases in related adverse events for adults of all ages. One area of concern has been their combined use with opioids, given the increased risk of overdose and overdose death among opioid users coprescribed benzodiazepines (8-10). But benzodiazepines pose risks of their own. In an analysis of emergency department visits from 1999 to 2006 for poisoning from opioids, sedatives (sleep promoting), or tranquilizers (anxiolytics or muscle relaxants), the largest absolute growth was in benzodiazepine-related poisonings (11), and benzodiazepine-related overdose mortality grew nearly fivefold from 1996 to 2013 (1). Concerns related to benzodiazepine prescribing have spread beyond older adults and beyond coprescription with opioids (12).

The growth in adverse outcomes suggests that benzodiazepine prescribing and misuse in the United States have increased in tandem; however, less is known about benzodiazepine misuse. After marijuana use, prescription drug misuse-defined here as use in a manner other than prescribed or by a person to whom it was not prescribed—is the most common type of illicit drug use (13). Most recent benzodiazepine-related work has focused either on misuse in the context of opioid use (14-16) or on tranquilizer or sedative medication misuse—but not on benzodiazepines specifically (17, 18). The lack of information about misuse among older adults is particularly striking because they are prescribed benzodiazepines at the highest rates, are most at

HIGHLIGHTS

- Benzodiazepine use among 50- to 64-year-old adults now matches use among older adults (≥65).
- Among those 18 to 25, benzodiazepine misuse is as prevalent as use as prescribed.

risk of related adverse events, and have higher rates of use of alcohol and other substances than prior aging cohorts (19, 20). A recent systematic review of benzodiazepine and opioid misuse among older adults (21) found just one study that estimated potential benzodiazepine misuse among older adults in the United States (22). Given their widespread use, abuse potential (23), and related risks, surprisingly little is known about benzodiazepine misuse.

Addressing the growing problem of benzodiazepine use and misuse first requires information about the current scope and nature of misuse. As a result of a 2015 redesign, the National Survey on Drug Use and Health (NSDUH) now provides detailed information regarding prescription drug use and misuse in the United States, including the type of and reasons for misuse and the source of the misused medication. We used data from the redesigned NSDUH to develop national estimates of benzodiazepine use and misuse among U.S. adults and to determine whether characteristics associated with misuse varied by age, given the unique risks of these medications to older adults and the need for targeted interventions to reduce misuse.

METHODS

This analysis used data from the NSDUH, which measures the prevalence and correlates of drug use among the U.S. civilian, noninstitutionalized population (24). The survey uses a 50-state design with an independent, multistage area probability sample; is administered by RTI International; and is sponsored by the Substance Abuse and Mental Health Services Administration. Respondents complete two computer-assisted segments, one conducted by an interviewer and a second audio-assisted portion without interviewer help, intended to provide a private environment to increase the likelihood of honest reporting of illicit drug use.

NSDUH was redesigned in 2015 to collect more detailed information on the use and misuse of prescription medications, including benzodiazepines-in prior years, questions were limited to misuse exclusively (25). The pre-2015 NSDUH definition of misuse was limited to "nonmedical use," but the 2015 definition was revised to include "in any way a doctor did not direct." This analysis was limited to data from respondents ≥18 in the 2015 and 2016 survey years (N=86,186), the years available since the redesign.

Benzodiazepine Use or Misuse

Respondents could report benzodiazepine use through survey queries about tranquilizer or sedative use. NSDUH classifies tranquilizers as medications specifically for relief of anxiety or muscle spasms and sedatives as medications for insomnia. This analysis was limited to the 10,290 respondents who specifically reported benzodiazepine use in response to the tranquilizer and sedative items. [Additional details are provided in an online supplement to this article.]

For each medication class, NSDUH collected information on past-year use (that is, taken as prescribed) and misuse. Respondents were asked about the specific manner of misuse: without a prescription, in greater amounts or more often than prescribed, longer than prescribed, or any other use other than as prescribed. Next, they were asked about reasons for misuse: "to relax," "to experiment," "to get high," "for sleep," "for emotions," "to counter the effect of another drug," because they were "hooked," or another reason. Finally, respondents were asked about the source of medication for misuse (for example, their clinician or a friend or relative). For this analysis, the misuse category identifies a respondent who reported any past-year misuse, even though that respondent may have also used the benzodiazepine as prescribed. Presence of abuse or dependence was determined by DSM-IV criteria.

Other Respondent Characteristics

We included respondent self-rating of health and past-year presence of a major depressive episode, suicidal thinking, and mental illness by using a variable developed by NSDUH based on responses to items about psychological distress, functional impairment, symptoms of a major depressive episode, and suicidal ideation (26).

We included past-year alcohol, marijuana, and heroin use or abuse or dependence; past-year use of tobacco products; and past-year use, misuse, or abuse of or dependence on prescription opioids and stimulants.

Finally, respondents provided sociodemographic information, including age, gender, race-ethnicity, and household income.

Analysis

Analyses incorporated weights, clustering, and stratification, using NSDUH design elements to account for the complex survey design and generate nationally representative estimates. After determining population characteristics among adult respondents, we estimated the prevalence of benzodiazepine use-as prescribed, misuse, and any use-among adults overall and by age group. We compared demographic and clinical characteristics of adults with and without any benzodiazepine use by using chi-square tests. We determined the magnitude of the association between respondent characteristics and benzodiazepine use by using multivariable logistic regression (0, no benzodiazepine use; 1, any benzodiazepine use), adjusting for other sociodemographic and clinical characteristics.

We then limited the analysis to respondents who reported any past-year benzodiazepine use, split into those who reported use as prescribed or misuse. We compared characteristics of each group by using chi-square tests. Next, we completed bivariate logistic regression for each characteristic (0, use as prescribed; 1, misuse), including an interaction by age (younger [18-49] versus older [≥50]), to determine whether age moderated the association of each characteristic with misuse. We used ≥50 years as the age cutoff because prescription benzodiazepine use among those in later middle age approaches that of adults ages ≥65 (1),

and the youngest respondents in the Baby Boom cohort (those born in 1964) would have turned 50 just before the 2015 NSDUH. We used multivariable logistic regression to determine the characteristics associated with misuse among benzodiazepine users, adjusting for other demographic and clinical characteristics.

The final stage of analysis examined the characteristics of past-year benzodiazepine misuse overall and by age. We determined the type of, reason for, and source of misused benzodiazepine, along with the specific benzodiazepine misused.

Analyses were conducted with Stata, version 13.1, with two-sided tests and α =.05. The University of Michigan Medical School Institutional Review Board (IRB) considers

analyses of deidentified, publicly available data exempt from IRB approval.

RESULTS

Prevalence and Predictors of Any Benzodiazepine Use

An estimated 30.6 million adults per year in the United States (95% confidence interval [CI]=29.7-31.5 million) used benzodiazepines in the past year, an overall prevalence of 12.6% (CI=12.2%-12.9%), including misuse among 2.2% (CI=2.0%-2.3%) and use as prescribed among 10.4% (CI=10.1%-10.7%). Use among adults ages 50-64 was highest (Figure 1 and Table 1).

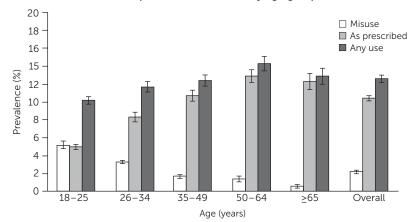
Women and non-Hispanic white respondents reported the highest rates of any past-year use. In the adjusted logistic regression model (Table 1), female gender, older age, and more education were all associated with increased odds of use, and respondents' race-ethnicity other than non-Hispanic white was associated with lower odds of any benzodiazepine use.

The presence of past-year mental illness was associated with increased odds of any use, as was worse self-rated health. In almost every instance, past-year use, misuse, or abuse of or dependence on tobacco, alcohol, marijuana, heroin, prescription opioids, or prescription stimulants was associated with any benzodiazepine use. Among all substances, prescription opioids were most strongly associated with benzodiazepine use at every level, from use as prescribed through abuse or dependence.

Prevalence and Predictors of Benzodiazepine Misuse

Among respondents who reported any benzodiazepine use, 25.3 million (CI=24.5-26.1 million) reported use as prescribed by their clinician, and 5.3 million (CI=5.0-5.6 million) reported misuse. Use as prescribed was highest among adults ages 50-64 (Figure 1 and Table 2). Misuse was highest among the youngest adults and decreased with age. Most benzodiazepine use among respondents ages 18-25

FIGURE 1. Prevalence of prescription benzodiazepine use and misuse among 2015 and 2016 NSDUH respondents (N=86,186), by age group^a



^a NSDUH, National Survey on Drug Use and Health.

was misuse (5.2%, CI=4.8%-5.6%). In contrast, misuse was reported by just .6% (CI=.4%-.8%) of adults ages \geq 65.

Bivariate logistic regressions testing for a moderating effect of age on the associations between respondent characteristics and misuse found a statistically significant interaction in just three instances (Table 2). Even though age itself was strongly associated with lower odds of misuse, because of the minimal evidence for a moderating effect, characteristic × age interactions were not included in the multivariable model. Females had lower odds than males of misuse; however, apart from age, no other demographic characteristic was associated with misuse. Fair or poor health self-rating was associated with lower odds of misuse, whereas any level of marijuana or alcohol use was associated with increased odds of benzodiazepine misuse. Prescription opioid use as prescribed was associated with lower odds of benzodiazepine misuse, whereas opioid misuse and abuse or dependence were the characteristics most strongly associated with benzodiazepine misuse.

Characteristics of Misuse Among Younger and Older Adults

The most common type of benzodiazepine misuse overall was use without a prescription, although this type of misuse use was less likely to be reported by respondents ages ≥ 50 , compared with younger adults (ages 18-49) (Table 3). Compared with younger adults, older respondents were more likely to report using their benzodiazepine more often than prescribed.

The most common reason for misuse overall was to relax or relieve tension, followed by to help with sleep. Older adults were significantly more likely than younger adults to endorse misuse to help with sleep, and they were much less likely to report misuse to get high.

The most common source of the misused medication for both younger and older groups was from a friend or relative. When all sources of benzodiazepines were combined—free, bought, or stolen-a friend or relative was the source for

TABLE 1. Overall characteristics and predictors of any past-year benzodiazepine use among all adult 2015 and 2016 NSDUH respondents $^{\rm a}$

	***	Benzodiazepine use ^b								
Characteristic	All respondents (%) (N=86,186)	None (%) (N=75,896)	Any (%) (N=10,290)	F ^c	df	р	AOR ^d	95% CI		
Overall	100.0	87.5	12.6							
Demographic	200.0	07.0	12.0							
Age				18.2	3, 138	<.001				
18-25 (reference)	14.3	89.8	10.2							
26-34	15.8	88.3	11.7				1.27***	1.18-1.36		
35-49	24.9	87.6	12.4				1.65***	1.52-1.78		
50-64	25.6	85.7	14.3				2.04***	1.86-2.22		
≥65	19.4	87.1	12.9				2.28***	2.04-2.54		
Gender				366.6	1, 50	<.001				
Male (reference)	48.2	90.5	9.5							
Female	51.8	84.6	15.4				1.84***	1.72-1.96		
Race-ethnicity				221.3	3, 124	<.001				
Non-Hispanic white (reference)	64.6	84.6	15.4							
Non-Hispanic black	11.8	92.8	7.2				.45***	.4050		
Hispanic	15.7	92.3	7.7				.59***	.5466		
Other	8.0	93.5	6.5				.45***	.3853		
Education				23.4	2, 84	<.001				
Less than high school (reference)	13.5	90.1	9.9							
High school graduate, some	56.0	86.6	13.4				1.28***	1.12-1.46		
college										
College graduate or higher	30.4	87.8	12.2				1.41***	1.18-1.69		
Household income				5.2	3, 143	.002				
<\$20,000 (reference)	17.4	86.2	13.8							
\$20,000-\$49,999	30.0	87.3	12.7				1.05	.96-1.15		
\$50,000-\$74,999	16.3	87.9	12.1				1.05	.94-1.17		
≥\$75,000	36.3	88.0	12.0				1.12	.99-1.27		
Medicaid				63.6	1, 50	<.001				
No (reference)	85.8	87.9	12.1							
Yes	14.2	84.9	15.1				1.01	.93-1.11		
Metropolitan area				12.0	2, 94	<.001				
Large (reference)	55.2	88.2	11.8							
Small	30.0	86.5	13.5				1.02	.96–1.09		
Nonmetropolitan area	14.8	86.9	13.1				.96	.87-1.05		
Past year general health										
Presence of a major depressive				912.5	1, 50	<.001				
episode ^e										
No	93.3	89.1	10.9							
Yes	6.7	66.4	33.6							
Suicidal ideation ^e				857.7	1, 50	<.001				
No	96.0	88.3	11.7							
Yes	4.0	68.6	31.4							
Presence of mental illness				215.9	2, 98	<.001				
None (reference)	81.9	90.9	9.1							
Mild	9.2	78.5	21.5				1.93***	1.74-2.15		
Moderate	4.8	70.7	29.3				2.47***	2.20-2.76		
Severe	4.2	58.5	41.5				3.80***	3.40-4.26		
Health self-rating				961.7	3, 142	<.001				
Excellent or very good (reference)	56.8	90.2	9.8							
Good	29.2	86.4	13.6				1.26***	1.16-1.38		
Fair or poor	14.0	78.4	21.6				1.78***	1.59-2.00		
Past-year substance use										
Tobacco				302.1	1 50	<.001				
None (reference)	69.2	89.5	10.5	JUL.1	1, 50	~.001				
Any	30.8	83.0	17.0				1.27***	1.16-1.38		
Alcohol	50.0	00.0	17.0	151.9	2 93	<.001	±. <i>L</i> /	1.10 1.00		
None (reference)	30.3	89.5	10.5	101.9	L, JJ	~.001				
Any	63.5	87.6	12.4				1.10*	1.00-1.21		
Abuse or dependence	6.2	75.9	24.1				1.53***	1.35-1.73		

continued

TABLE 1, continued

	All	Benzodiazepine use ^b								
Characteristic	respondents (%) (N=86,186)	None (%) (N=75,896)	Any (%) (N=10,290)	F ^c	df	р	AOR ^d	95% CI		
Marijuana				521.5	2, 96	<.001				
None (reference)	86.1	89.2	10.8							
Any	12.5	77.3	22.7				1.81***	1.66-1.98		
Abuse or dependence	1.4	69.7	30.3				1.97***	1.65-2.34		
Heroin				292.1	2, 96	<.001				
None (reference)	99.6	87.6	12.4							
Any	.1	44.3	55.7				2.92**	1.49-5.75		
Abuse or dependence	.2	35.8	64.2				2.65***	1.88 - 3.74		
Prescription opioid				1094.0	3, 137	<.001				
None (reference)	63.2	93.2	6.8							
Prescription use	32.3	80.0	20.0				2.54***	2.39 - 2.71		
Misuse	3.8	64.5	35.5				4.21***	3.66-4.85		
Abuse or dependence	.7	40.1	59.9				7.34***	5.51-9.76		
Prescription stimulant				565.5	3, 133	<.001				
None (reference)	93.3	88.9	11.1							
Prescription use	4.6	70.2	29.8				2.11***	1.89 - 2.36		
Misuse	1.9	62.6	37.4				2.71***	2.31-3.18		
Abuse or dependence	.2	45.7	54.3				2.21***	1.48-3.29		

^a NSDUH, National Survey on Drug Use and Health. All percentages are weighted.

nearly 70% of respondents reporting misuse. The next most common source was a single clinician.

Alprazolam was the most common benzodiazepine misused. Compared with younger adults, older adults were more likely to misuse lorazepam or diazepam. Respondents who reported misuse did so on 5.4 days (SE=.3) in the past month. Among those with benzodiazepine misuse, 4.6% (CI=3.7%-5.6%) and 6.8% (CI=5.6%-8.2%) met criteria for past-year abuse and past-year dependence, respectively.

DISCUSSION

In this cross-sectional analysis of survey data from U.S. adults, the annual prevalence of benzodiazepine use, when both prescription use and misuse were included, was 12.6% and exceeded 15% among women and non-Hispanic white respondents. By age, the highest rate of overall benzodiazepine use was among adults ages 50-64. More than 2% of respondents overall endorsed misuse, which was highest among the youngest adults (ages 18-25), for whom misuse exceeded as-prescribed use. In contrast, adults ages ≥65 had the lowest prevalence of misuse (.6%).

To our knowledge, this is the first analysis of benzodiazepine use in the United States to find that the highest prevalence of use is no longer among adults ages ≥65.

Although use among those ages ≥65 has not been declining (1, 3), the decades of evidence regarding safety concerns (4, 7, 27) and professional guidelines (6, 28) recommending limited use among older adults may have helped slow growth. In contrast, the aging Baby Boomers-who constituted nearly the entire 50-64 age group in this analysishave higher rates of alcohol and other substance use, compared with previous cohorts of aging adults (19, 20, 29). The high level of use among these late-middle-age adults means that potentially inappropriate prescribing of benzodiazepines to older adults may continue as this cohort ages.

The NSDUH redesign affords new insights into the nature of benzodiazepine misuse, which accounted for nearly 20% of all use among adults but which was much more common among younger adults. Although age largely did not moderate the patient characteristics associated with misuse, the nature of misuse varied between younger (ages 18–49) and older (ages ≥50) adults. Misuse without a prescription was the most common type of misuse, although this was more common among younger adults; older adults were more likely to use their benzodiazepine more often than prescribed. Misuse to help relax and to help sleep were the main reasons for misuse among both age groups, although sleep was a relatively larger driver of misuse among older adults. Relatively little misuse was for experimentation or to

b Percentages reflect past-year benzodiazepine use among respondents with each characteristic (for example, 10.2% of adults ages 18–25 reported past-year benzodiazepine use, and 89.8% did not). All estimates used NSDUH design elements to generate nationally representative estimates.

^c To account for complex survey design, Stata converts the usual Pearson chi-square into an F statistic with noninteger degrees of freedom, which have been

d AOR, adjusted odds ratio. The model was adjusted for all respondent demographic, general health, and substance use characteristics presented as rows in the table, except for major depressive episode and suicidal ideation. 0, no benzodiazepine use; 1, any benzodiazepine use.

e Presence of a major depressive episode and suicidal ideation were not included in the model per NSDUH instructions (26), because those variables are used to generate the presence of a mental illness variable.

^{*}p≤.05, **p≤.01, ***p≤.001.

TABLE 2. Past-year prevalence of use and misuse of prescription benzodiazepines among adult NSDUH respondents and predictors of benzodiazepine misuse, by characteristic^a

	All respond (N=86,1	ents (%) 86) ^b	Benzodiazepine users (%) (N=10,290) ^c							
Characteristic	As prescribed (N=7,757)	Misuse (N=2,533)	As prescribed (N=7,757)	Misuse (N=2,533)	F ^d	df	р	AOR ^e	95% CI	
Overall	10.4	2.2	82.8	17.2						
Demographic										
Age					188.2	3, 139	<.001			
18-25 (reference)	5.0	5.2	49.0	51.0						
26-34	8.3	3.3	71.4	28.6				.55**	.4666	
35-49	10.7	1.7	86.2	13.8				.33**	.2741	
50-64	12.9	1.4	90.3	9.7				.33**	.2643	
≥65	12.3	.6	95.6	4.4				.23**	.1534	
Gender					102.3	1, 50	<.001			
Male (reference)	7.3	2.3	76.3	23.7						
Female	13.3	2.1	86.5	13.5				.74**	.6190	
Race-ethnicity					4.9	3, 146	.003			
Non-Hispanic white	12.9	2.5	83.6	16.4		-,				
(reference)	12.5	2.0	00.0	10						
Non-Hispanic black	5.7	1.5	79.6	20.4				1.25	.94-1.65	
Hispanic	6.1	1.6	78.8	21.2				1.05	.79-1.38	
Other	5.4	1.1	82.4	17.6				.95	.67-1.35	
Education	0.1	1.1	02.1	17.0	9.4	2 100	<.001	.50	.07 1.00	
Less than high school	8.1	1.9	81.3	18.7	J. ¬	2, 100	<.001			
(reference)	0.1	1.5	01.5	10.7						
High school graduate, some	10.9	2.5	81.3	18.7				.99	.77-1.26	
	10.9	2.5	61.3	10.7				.99	.//-1.20	
college	10 E	1 7	86.2	17.0				00	67 1 22	
College graduate or higher	10.5	1.7	80.2	13.8	0.4	7 1 1 0	< 0.01	.90	.67–1.22	
Household income	40.0	7.0	70.6	24.4	9.4	5, 140	<.001			
<\$20,000 (reference)	10.9	3.0	78.6	21.4				0.0	70 4 27	
\$20,000-\$49,999	10.4	2.3	81.9	18.1				.98	.79-1.23	
\$50,000-\$74,999	10.3	1.8	85.0	15.0				.77	.58-1.03	
≥\$75,000	10.2	1.8	84.9	15.1				.84	.67–1.06	
Medicaid					7.0	1, 50	.01			
No (reference)	10.1	2.0	83.3	16.7						
Yes	12.1	3.0	80.2	19.8				.99	.79–1.23	
Metropolitan area					4.8	2, 95	.01			
Large (reference)	9.7	2.1	82.0	18.0						
Small	11.1	2.4	82.5	17.5				1.02	.83-1.25	
Nonmetropolitan	11.3	1.8	86.0	14.0				.84	.67–1.06	
Past year clinical Presence of major depressive					20.9	1 50	<.001			
episode ^f					20.9	1, 30	<.001			
	0.2	1.0	07.0	16.2						
No	9.2	1.8	83.8	16.2						
Yes Suicidal ideation ^f	26.4	7.2	78.5	21.5	02.0	1 50	< 0.01			
	0.0	1.0	047	15.7	92.9	1, 50	<.001			
No	9.9	1.8	84.3	15.7						
Yes	21.7	9.7	69.1	30.9	45.4	7 470	. 004			
Presence of mental illness				45.0	15.1	3, 139	<.001			
None (reference)	7.7	1.4	84.8	15.2						
Mild	18.0	3.6	83.5	16.5				.90	.72-1.12	
Moderate	22.8	6.5	77.9	22.1				1.24	.93-1.63	
Severe	31.7	9.8	77.3	22.7				1.19	.93–1.53	
Health self-rating					30.1	2, 96	<.001			
Excellent or very good (reference)	7.8	2.0	79.4	20.6						
Good	11.2	2.4	82.7	17.3				.85	.70-1.04	
Fair or poor	19.2	2.3	89.2	10.8				.61**	.47–.80	
•	17.5	2.5	03.L	10.0				.01	,	
Other past-year substance use					F.C.F. 4	4 50	- 001			
Tobacco	0.5	4.0	007	0.7	565.4	1, 50	<.001			
None (reference)	9.5	1.0	90.7	9.3				4 44-1-1	4 47 4 70	
Any	12.3	4.8	71.9	28.1				1.41**	1.17–1.70 continued	

TABLE 2, continued

	All respond (N=86,1	ents (%) 86) ^b	Benzodiazepine users (%) (N=10,290) ^c							
Characteristic	As prescribed (N=7,757)	Misuse (N=2,533)	As prescribed (N=7,757)	Misuse (N=2,533)	F ^d	df	р	AOR ^e	95% CI	
Alcohol					199.5	2, 90	<.001			
None (reference)	9.8	.8	92.8	7.2						
Any	10.4	2.0	83.5	16.5				1.42*	1.08 - 1.87	
Abuse or dependence	13.8	10.3	57.2	42.8				2.31**	1.73-3.09	
Marijuana					558.3	2, 99	<.001			
None (reference)	9.8	1.0	90.7	9.3						
Any	14.4	8.3	63.3	36.7				1.88**	1.58-2.24	
Abuse or dependence	11.7	18.6	38.7	61.3				2.39**	1.62 - 3.52	
Heroin					90.3	2, 86	<.001			
None (reference)	10.3	2.0	83.6	16.4						
Any	26.6	29.2	47.7	52.3				.91	.41-2.04	
Abuse or dependence	23.5	40.8	36.5	63.5				1.20	.76-1.91	
Prescription opioid					462.0	3, 144	<.001			
None (reference)	5.9	.9	86.2	13.8						
Prescription use	18.3	1.7	91.5	8.5				.69**	.5784	
Misuse	15.6	19.9	44.1	55.9				4.11**	3.26-5.18	
Abuse or dependence	24.4	35.6	40.7	59.3				4.81**	3.45-6.72	
Prescription stimulant					410.6	3, 127	<.001			
None (reference)	9.6	1.5	86.8	13.2						
Prescription use	24.1	5.7	80.9	19.1				.90	.74-1.09	
Misuse	13.3	24.1	35.6	64.4				2.46**	1.85-3.27	
Abuse or dependence	14.7	39.6	27.1	72.9				3.40**	1.72-6.70	

^a NSDUH, National Survey on Drug Use and Health. Percentages are weighted.

get high, and few respondents who reported benzodiazepine misuse met criteria for past-year abuse or dependence.

Taken together, these misuse findings raise questions about the underlying contributors to misuse as defined and identified in the NSDUH. Younger adults are more likely than other age groups to lack health insurance (30), whereas the most common reasons for misuse (for example, to relax or relieve tension) were reasons for which a clinician might prescribe a benzodiazepine or refer for behavioral treatment. Therefore, a significant proportion of NSDUHdefined "misuse" could reflect use for untreated symptoms among those with poor access to care—specifically, for behavioral treatments for insomnia (31, 32) or anxiety disorders (33, 34). The development of interventions, such as Web-based cognitive-behavioral therapy for insomnia, may help increase access to benzodiazepine alternatives for those who lack access to providers or insurance or both (35).

Clinicians should recognize their role as a source of misused benzodiazepines, either through medication that they prescribe but that is used other than as instructed or as

the source of prescribed medication given for misuse to a friend or relative. In addition to being mindful of their role as a potential source of misused medications, clinicians have an important role in understanding the reason for their patients' misuse to determine the appropriate intervention. If patients are consuming prescribed medication faster than expected, clinicians should determine the reason. Is it for inadequate symptom control? For additional indicationsfor example, prescribed for anxiety but also used for sleep? Is a patient allowing another family member to use some? Some misuse may be for symptoms appropriately treated by a benzodiazepine, but clinicians should be mindful of other potential reasons for misuse. An uninsured young person may use an older relative's prescribed benzodiazepine for insomnia rather than to get high, but this certainly was not the intention of the prescribing clinician.

Other substance use disorders were strongly associated with benzodiazepine use and misuse. Benzodiazepine misuse was most strongly associated with misuse of and abuse of or dependence on prescription opioids and stimulants.

b Values represent the percentage of all respondents with a given characteristic who reported use as prescribed or misuse (for example, 5.2% of adults ages 18-25 reported misuse).

^c Values represent the percentage of respondents with a given characteristic who reported use as prescribed or misuse (for example, among adults ages 18–25 who reported any benzodiazepine use, 51.0% reported misuse).

d To account for complex survey design, Stata converts the usual Pearson chi-square into a F statistic with noninteger degrees of freedom, which have been

e AOR, adjusted odds ratio. The model was adjusted for all respondent demographic, clinical, and substance use characteristics presented as rows in the table, except for major depressive episode and suicidal ideation. Because just three characteristic \times age interactions in bivariate tests were statistically significant (nonmetropolitan area [p=.02], any past-year use of tobacco [p=.002], and any past-year use of marijuana [p=.008]), characteristic × age interactions were not included in the adjusted model. 0, no benzodiazepine use; 1, any benzodiazepine use.

Presence of MDE and suicidal ideation were not included in the model per NSDUH instructions (26), because those variables are used to generate the presence of mental illness variable

^{*}p≤.01, **p≤.001.

TABLE 3. Characteristics of benzodiazepine misuse among 2015 and 2016 NSDUH respondents (N=2,533), by age group^a

		Age group (S				
		18-49	≥50			
Characteristic	Overall	(N=2,357)	(N=176)	ORb	95% CI	р
Type of misuse						
Without a prescription	73.9	78.4	56.4	.36	.25 to .51	<.001
Used in greater amounts than prescribed	16.0	15.0	20.2	1.44	.86 to 2.40	.16
Used more often than prescribed	10.1	8.1	17.9	2.47	1.57 to 3.90	<.001
Used longer than prescribed	4.9	4.7	5.7	1.24	.62 to 2.48	.53
Used in other ways other than as directed	18.7	17.6	23.2	1.41	.90 to 2.21	.13
Main reason for misuse						
Relax, relieve tension	47.1	48.6	41.7	.76	.53 to 1.08	.12
Experiment	6.0	6.5	4.0	.59	.21 to 1.68	.28
Get high	11.6	13.9	3.0	.19	.07 to 56	.003
Help with sleep	26.6	22.4	41.7	2.48	1.83 to 3.35	<.001
Help with feelings, emotions	10.1	10.4	9.1	.86	.43 to 1.76	.68
Increase or decrease effects of another drug	1.6	1.7	.9	.51	.11 to 2.35	.38
Hooked or have to have it	.6	.5	.9	1.89	.17 to 20.72	.60
Source of medication for misuse						
1 clinician	19.5	14.3	38.4	3.73	2.65 to 5.26	<.001
≥2 clinicians	.9	.6	1.9	3.27	.87 to 12.29	.08
Stolen from health care setting	.1	.2	.0			
Free from friend or relative	53.2	55.4	45.1	.66	.44 to .99	.04
Bought from friend or relative	12.3	14.2	5.6	.35	.15 to .84	.02
Stolen from friend or relative	3.3	3.2	3.5	1.08	.31 to 3.81	.90
Bought from dealer or stranger	7.7	8.9	3.1	.33	.10 to 1.07	.06
Medication misused						
Alprazolam	75.1	80.3	56.4	.32	.21 to.49	<.001
Lorazepam	15.4	12.6	26.0	2.44	1.54 to 3.87	<.001
Clonazepam	20.0	21.7	13.6	.57	.33 to 1.00	.05
Diazepam	20.5	17.8	30.5	2.04	1.21 to 3.43	.009
Other characteristics of benzodiazepine						
misuse						
Mean days of misuse in past month	5.4	5.0	7.1	2.10	30 to 4.51	.09
Meets past-year criteria for abuse (reference: no)	4.6	5.0	3.1	.60	.20 to 1.78	.35
Meets past-year criteria for dependence (reference: no)	6.8	6.8	6.6	.97	.50 to 1.90	.94

^a NSDUH, National Survey on Drug Use and Health. Percentages are weighted.

Prescription drug monitoring programs are an important tool for clinicians to understand which of their patients may be misusing other medications and would thus be at high risk of benzodiazepine misuse. The association of alcohol abuse or dependence with increased odds of benzodiazepine misuse is particularly concerning in light of the increased potential for fatal poisoning when these substances are combined (36, 37). This has received much less attention than the opioid-benzodiazepine combination, even though alcohol use disorders are more prevalent than prescription opioid use disorders.

Our findings that women, older respondents, and non-Hispanic white respondents reported higher rates of benzodiazepine use are consistent with previous work (1, 2, 38-40). In contrast, women and older respondents were less

likely than men and younger respondents to report misuse. This may further support the hypothesis that higher misuse is partly a function of limited access to a prescribed option-limited for younger individuals because of lack of insurance and for men because of minimal disclosure of mental health concerns to providers (41). However, higher misuse was not found among racial and ethnic minority groups, even though they have limited access to specialty mental health care and lower rates of insurance, compared with non-Hispanic whites (30).

Our estimate of the annual prevalence of any benzodiazepine use-12.6% overall and 10.4% as prescribed-is higher than other recent results. A study using data from the Medical Expenditure Panel Survey (MEPS) found that 5.6% of adults filled a benzodiazepine prescription in 2013 (1), and an analysis of data from the 2013-2014 National Health and Nutrition Examination Survey (NHANES) found benzodiazepine use among 4.2% of adults (42). Survey differences may partly account for the different estimates: NHANES assesses medica-

tion use in the past 30 days (43), whereas MEPS samples households, with information often provided by a single knowledgeable informant (44). MEPS respondents may underestimate the number of unique medications used in a year, specifically underreporting those used for a shorter duration or with fewer fills (45), which is how benzodiazepines may be prescribed. Finally, NSDUH is the only survey to include misuse. Our estimate of misuse is similar to an NSDUH estimate from 2002-2004 (17), although that analysis examined tranquilizers and sedatives overall and was not limited to benzodiazepines.

Our analysis had a number of limitations. NSDUH is crosssectional, and because of the 2015 redesign, we could not track trends in misuse over time. There was a potential for nonresponse bias. NSDUH response rates have been declining,

^b Younger adults are the reference group.

although this is unfortunately true for several federally administered national surveys (46). NSDUH is nationally representative but of the civilian population and thus does not include active-duty members of the military or institutionalized adults.

CONCLUSIONS

The study found a prevalence of benzodiazepine use among U.S. adults that is higher than previously reported. Misuse was highest among the youngest adults. Overall use among adults ages 50–64 was found to exceed that among adults ages ≥65. At the policy level, more widespread insurance coverage and access to behavioral treatments may reduce benzodiazepine use and misuse—some misuse may reflect limited access to health care generally and behavioral treatments specifically. Although clinicians should be mindful of the potential for benzodiazepine misuse among patients with any level of substance use, findings indicated that prescription opioid and stimulant use disorders were most strongly associated with benzodiazepine misuse.

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