It is illegal to post this copyrighted PDF on any website. Prevalence of Kratom Use and Co-Occurring Substance Use Disorders in the United States

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K ratom (*Mitragyna speciosa*) is a novel psychoactive drug that is often used to self-manage pain and opioid withdrawal.¹⁻³ While kratom, in itself, has been found to lack the addictive potential of classic opioids and may have promising therapeutic properties,⁴ there are concerns about the safety of unregulated kratom products in the United States.⁵ As kratom has increasingly been found in association with overdoses, usually in the presence of other substances,¹ it is important to understand the co-occurring substance use disorders (SUDs) associated with kratom use. The aim of the present study was to describe the prevalence of kratom use and co-occurring substance use using the National Survey on Drug Use and Health (NSDUH).

Methods

We used data from the 2019 NSDUH (n = 56,136, weighted interview response rate 72.1% for adolescents, 64.2% for adults), the first year for which kratom data were available. NSDUH is conducted annually among the civilian, noninstitutionalized US population, aged 12 years and older. We calculated prevalence estimates of self-reported history of kratom use, as well as co-occurring past-year opioid, stimulant, alcohol, marijuana, sedative, and other SUDs. Prevalence ratios were calculated via log-binomial regression. We tabulated the number of SUDs per individual and examined if whether kratom use was associated with increased risk of multiple SUDs. Analyses were done with STATA 16; all estimates accounted for

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*Corresponding author: Kevin Y. Xu, MD, MPH, Department of Psychiatry, Washington University School of Medicine, 420 South Euclid Ave, Campus Box 8134, St Louis, MO 63110 (xukeviny@wustl.edu). Prim Care Companion CNS Disord 2021;23(4):21br02930 complex survey design. This study was exempted from review by the Washington University Institutional Review Board.

Results

As shown in Table 1, prevalence of lifetime kratom use in the United States was 1.5% (95% CI, 1.4%–1.6%). Among those who used kratom, 50.9% (46.8%–54.9%) used more than 1 year ago, 28.4% (24.4%–32.8%) used within the past year, and 20.7% (17.2%–24.6%) used within the past month. Most lifetime kratom users were male (61.2%, 56.6%–65.6%), white (81.9%, 78.1%–85.1%), and between the ages of 18 and 34 (55.2%, 49.9%–60.4%). Few users were under the age of 18.

Almost one-third (31%, 26.6%–35.7%) of kratom users had at least 1 SUD. Whereas the prevalence of opioid use disorder was 0.5% (0.4%–0.6%) among never users, this increased 18-fold to 8.9% (6.3%–12.5%) among lifetime kratom users. Kratom use was also associated with increased prevalence of prescription stimulant (16.5-fold, 9.2–29.5), methamphetamine (12.5-fold, 8.3–18.8), cocaine (14-fold, 8.5–23.1), and tranquilizer or sedative use disorders (16.8fold, 9.3–31.1).

Lifetime kratom use was also associated with a higher burden of multiple SUDs. Among never users, the pastyear prevalence of experiencing 3 or more co-occurring SUDs was 0.16%. However, this prevalence increased almost 20-fold to 2.81% among kratom users. In addition, whereas only 7.6% (7.2%–8.0%) of never users reported a major depressive episode in the last year, this increased to 26.7% (22.7%–31.0%) for kratom users.

Discussion

Nearly one-third of lifetime kratom users in the United States have at least 1 SUD; over one-fourth experienced past-year major depression. Lifetime kratom use is associated with a higher risk of other SUDs, especially opioids, stimulants, and sedatives. Importantly, the high burden of co-occurring SUDs among kratom users should not be interpreted to suggest that kratom causes SUDs; existing studies^{6–8} show that kratom itself does not typically produce a "high" in comparison to classic opioids, with the vast majority of users not meeting *DSM* criteria for kratom-related SUDs. Multiple studies have found that kratom alkaloids have unique binding properties that show potential for management of pain,⁹ as well as ameliorate opioid withdrawal symptoms with minimal



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	Table 1. Prevalence of Kratom Use and Co-Occurring Substance Use Disorders in the United States, 2019											

	Never	Used Kratom (n=	55,188)	Lifetir	ne Kratom Use (n	Never Used Kratom		
	Weighted				Weighted	Prevalence		
Variable	Frequency	Prevalence (%)	95% CI	Frequency	Prevalence (%)	95% Cl	Ratio	95% CI
Sex (female)	26,231	51.7	51.0-52.3	557	38.8	34.4-43.4	1.3	1.5-1.2
Age, y								
12 to 17	13,339	10.9	10.6–11.2	58	4.0	2.9-5.4	0.4	0.3–0.5
18 to 25	13,844	7.4	7.2-7.6	382	13.6	11.3–16.2	1.8	1.5-2.2
26 to 34	8,342	17.4	16.9–18.0	259	37.7	33.2-42.3	2.2	1.9–2.5
≥35	19,663	64.3	63.7-65.0	249	44.8	39.6-50.1	0.7	0.6–0.8
Race								
White	31,368	61.7	60.7-62.7	721	81.9	78.1-85.1	1.3	1.3–1.4
Black	7,210	12.2	11.4–13.0	46	4.4	2.8-6.6	0.4	0.2-0.6
Hispanic	10,749	17.4	16.6–18.2	99	7.6	5.4-10.6	0.4	0.3-0.6
Asian	2,686	5.9	5.4-6.4	11	0.9	0.4-2.0	0.2	0.1-0.3
Native American	1,022	1.0	0.8-1.1	22	1.2	0.6-2.4	1.3	0.7-2.4
Multirace	2,153	1.9	1.8-2.1	49	4.0	2.3-7.0	2.1	1.2-3.6
Past year major depressive episode	4,228	7.6	7.2-8.0	239	26.7	22.7-31.0	3.5	3.0-4.1
Past year substance abuse or depend	dence							
Opioids (heroin or pain reliever)	292	0.5	0.4-0.6	77	8.9	6.3-12.5	18.0	12.1-26.9
Heroin only	57	0.1	0.1-0.1	12	0.8	0.3-2.1		
Pain reliever only	214	0.4	0.3-0.5	48	6.2	4.0-9.4		
Heroin and pain reliever	21	0.04	0.02-0.1	17	2.0	1.2-3.3		
Prescription stimulants	131	0.2	0.1-0.2	26	2.8	1.6-4.6	16.5	9.2–29.5
Methamphetamine	197	0.3	0.3-0.5	44	4.3	2.9-6.5	12.5	8.3–18.8
Cocaine	180	0.3	0.2-0.4	42	4.0	2.6-6.0	14.0	8.5-23.1
Alcohol	3,170	5.1	4.8-5.4	193	17.8	14.8-21.3	3.5	2.9-4.2
Marijuana	1,474	1.6	1.5-1.8	130	10.6	7.8–14.2	6.6	4.8-9.0
Tranquilizer or sedative	142	0.2	0.2-0.3	32	3.6	2.2-6.1	16.8	9.3–30.6
Hallucinogens	79	0.1	0.1-0.1	11	1.4	0.7-2.8	16.8	9.0-31.1
Any substance use disorder ^a	3,698	5.9	5.6-6.3	301	31.0	26.6-35.7		
1	3,327	5.4	5.1-5.7	224	23.7	20.1-27.8		
2	278	0.4	0.4-0.5	48	4.5	2.9-6.9		
≥3	93	0.2	0.1-0.2	29	2.8	1.7-4.6		
^a Including opioids, prescription stim	ulants, methar	nphetamine, alco	hol, marijuan	a, prescriptior	tranguilizers or se	edatives, hall	ucinogens, co	caine.

central nervous system depression.¹⁰ More research using nationally representative data is needed to elucidate the interplay between kratom's therapeutic promise and safety concerns, especially given a current lack of quality control surrounding kratom products.⁵

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Potential conflicts of interest: Dr Bierut is listed as an inventor on US Patent 8080371 "Markers for Addiction," covering use of SNPs in determining the diagnosis, prognosis, and treatment of addiction. Dr Borodovsky serves on the board of directors and is treasurer of the nonprofit MySafeRx Inc but does not receive any financial compensation for this work. Drs Xu, Mintz, Glaser, and Grucza declare no financial interests. None of the authors have financial relationships with organizations that may have an interest in this work.

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REFERENCES

- Anwar M, Law R, Schier J. Notes from the field: kratom (*Mitragyna speciosa*) exposures reported to poison centers—United States, 2010–2015. *MMWR Morb Mortal Wkly Rep.* 2016;65(29):748–749.
- Schimmel J, Amioka E, Rockhill K, et al. Prevalence and description of kratom (*Mitragyna speciosa*) use in the United States: a cross-sectional study. *Addiction*. 2021;116(1):176–181.
- 3. Grundmann O. Patterns of Kratom use and health impact in the US results from an online survey. *Drug Alcohol Depend*. 2017;176:63–70.
- Veltri C, Grundmann O. Current perspectives on the impact of kratom use. Subst Abuse Rehabil. 2019;10:23–31.
- Prozialeck WC, Avery BA, Boyer EW, et al. Kratom policy: the challenge of balancing therapeutic potential with public safety. *Int J Drug Policy*. 2019;70:70–77.
- Grundmann O, Babin JK, Henningfield JE, et al. Kratom use in the United States: a diverse and complex profile. *Addiction*. 2021;116(1):202–203.
- Garcia-Romeu A, Cox DJ, Smith KE, et al. Kratom (*Mitragyna speciosa*): user demographics, use patterns, and implications for the opioid epidemic. *Drug Alcohol Depend*. 2020;208:107849.
- 8. Smith KE, Lawson T. Prevalence and motivations for kratom use in a sample of substance users enrolled in a residential treatment program. *Drug Alcohol Depend*. 2017;180:340–348.
- Todd DA, Kellogg JJ, Wallace ED, et al. Chemical composition and biological effects of kratom (*Mitragyna speciosa*): in vitro studies with implications for efficacy and drug interactions. *Sci Rep.* 2020;10(1):19158.
- Wilson LL, Harris HM, Eans SO, et al. Lyophilized kratom tea as a therapeutic option for opioid dependence. *Drug Alcohol Depend*. 2020;216:108310.