

ORIGINAL RESEARCH

# Opioid-use disorder among patients on long-term opioid therapy: impact of final DSM-5 diagnostic criteria on prevalence and correlates

Joseph A Boscarino<sup>1</sup> Stuart N Hoffman<sup>1</sup> John J Han<sup>2</sup>

<sup>1</sup>Center for Health Research, <sup>2</sup>Department of Pain Medicine, Geisinger Clinic, Danville, PA, USA **Aims:** Previously, we estimated the prevalence and risk factors for prescription opioid-use disorder among outpatients on opioid therapy using the *Diagnostic and Statistical Manual of Mental Disorders* (DSM)-5 and DSM-4 criteria. However, at the time, the DSM-5 criteria were not finalized. In the current study, we analyzed these data using the final DSM-5 criteria and compared these results.

**Methods:** Using electronic records from a large US health care system, we identified outpatients receiving five or more prescription orders for opioid therapy in the past 12 months for noncancer pain (mean prescription orders =10.72; standard deviation =4.96). In 2008, we completed diagnostic interviews with 705 of these patients using the DSM-4 criteria. In the current study, we reassessed these results using the final DSM-5 criteria.

**Results:** The lifetime prevalence of DSM-5 opioid-use disorders using the final DSM-5 criteria was 58.7% for no or few symptoms (<2), 28.1% for mild symptoms (2–3), 9.7% for moderate symptoms (4–5), and 3.5% for severe symptoms (six or more). Thus, the lifetime prevalence of "any" prescription opioid-use disorder in this cohort was 41.3% (95% confidence interval [CI] =37.6–45.0). A comparison to the DSM-4 criteria indicated that the majority of patients with lifetime DSM-4 opioid dependence were now classified as having mild opioid-use disorder, based on the DSM-5 criteria (53.6%; 95% CI =44.1–62.8). In ordinal logistic regression predicting no/few, mild, moderate, and severe opioid-use disorder, the best predictors were age <65 years, current pain impairment, trouble sleeping, suicidal thoughts, anxiety disorders, illicit drug use, and history of substance abuse treatment.

**Conclusion:** Given the final DSM-5 criteria, including the elimination of tolerance and withdrawal, inclusion of craving and abuse symptoms, and introduction of a new graded severity classification, the prevalence of opioid-use disorders has changed, while many of the DSM-4 risk factors for opioid dependence were similar. To our knowledge, this is one of the first studies to compare the final results for DSM-5 versus DSM-4 prescription opioid-use disorders among a high-risk patient population.

**Keywords:** opioids, drug-use disorders, DSM-5, prescription drugs, pain, outpatients

#### Introduction

The prevalence of prescription opioid drug abuse and drug dependence in the USA has increased in the past decade, representing a serious public health threat.<sup>1-4</sup> The revised diagnostic criteria for the *Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition* (DSM-5) included significant modifications for prescription opioid-use disorder.<sup>5</sup> These changes included the exclusion of the tolerance and withdrawal symptoms for those taking opioids under appropriate medical supervision.<sup>6,7</sup> The rationale for this was that these symptoms would be "iatrogenic", rather than psychopathogenic,

Correspondence: Joseph A Boscarino Geisinger Clinic, 100 N Academy Avenue, MC 44-00, Danville, PA 17822-4400, USA Tel+1 570 214 9622 Fax+1 570 214 9451 Email jaboscarino@geisinger.edu in a therapeutic context.<sup>6,7</sup> In addition, given the conceptual problems noted in DSM-4 between "abuse" and "dependence", it was recommended to combine these symptom clusters into a single disorder of graded severity. <sup>6,8</sup> Previously, we compared the lifetime diagnostic results for the DSM-5 versus the DSM-4 for prescription opioid-use disorder, but this was based on the preliminary criteria proposed by the American Psychiatric Association (APA), not the final criteria.<sup>5,7</sup> Given that the final APA classification was modified, the objectives of the current study are to assess the prevalence of DSM-5 prescription opioid-use disorder among a highrisk patient population based on the final DSM-5 criteria, to identify the correlates associated with this final classification, and to compare the final DSM-5 criteria results to those using the DSM-4 criteria. Given the final changes proposed by the APA for prescription opioid-use disorder,<sup>5</sup> differences were expected.

#### **Methods**

## Sample

As described elsewhere, 4,7 the study subjects were randomly selected from primary and specialty care outpatients seen in the Geisinger Clinic, which is part of the Geisinger Health System (Danville, PA, USA), an integrated multihospital system that serves residents in 44 central and northeastern Pennsylvania counties. Patients were eligible for this study if they were 18 years of age or older, received care from one of nine community practice clinics or from three specialty clinics (including a pain, arthritis, and orthopedic clinic), and had five or more prescription opioid medication orders for nonmalignant pain in the electronic health record (EHR) in the past 12 months (mean prescription orders =10.72; standard deviation [SD] =4.96). Preliminary data analyses suggested that this cut-off point was associated with problematic drug use. It is noted that this count was based on the number of drug orders recorded in the EHR, not the actual number of prescriptions filled or refilled, which we did not have access to at the time. In the outpatient setting, opioids tend to be ordered by providers in small doses (ie, <30 pills) with no automatic refills prescribed. Consequently, numerous drug orders are seen in the EHR, but this does not mean that the average patient had ~11 months of exposure to opioid medications in the past 12 months.

#### Data collection

Telephone interviews for this study were completed from August 2007 through November 2008. Following patient notification letters, telephone recruitment was initiated. A total of 2,373 patients were initially identified as eligible for the survey and they were contacted by telephone. Of these patients, 234 were determined to be ineligible due to death, institutionalization, language barriers, illness, denial of opioid use, or being in the last group (number [n] =79) not contacted due to survey completion. Our survey completion rate was 705/2,139=33%, and our survey cooperation rate (ie, the percent interviewed after patient contact) was 51% (705/1,390).<sup>4,7</sup>

Following informed consent, structured diagnostic interviews were administered by telephone to assess depression, post-traumatic stress disorder (PTSD), general anxiety, and psychological trauma, 9-11 as well as opioid dependence, 12-14 tobacco dependence, 15 and childhood adversity. 16 We examined these domains because we wanted to determine the key factors that predict opioid disorders in clinical practice, as was done recently for PTSD. 17 The survey was administered by trained interviewers using a computer-assisted telephone interviewing system located at Geisinger's Center for Health Research in Danville, PA, USA. This study was approved by the Geisinger Clinic's Institutional Review Board.

# Lifetime DSM-5 and DSM-4 opioid disorders

For the present study, aberrant opioid use was defined based on both the DSM-4 criteria for dependence and the final DSM-5 criteria for opioid-use disorder published in 2013.<sup>5,18</sup> Because we assessed prescription opioid use at the beginning of the interview, we used a scale adapted for phone administration that had been previously validated. 14 The criteria for opioid dependence on this scale were concordant with DSM-4 nomenclature. 18 Lifetime DSM-4 prescription opioid dependence was defined as the presence of three or more prescription opioid symptoms (eg, health, family, functional, or legal problems) in the patient's history. Consistent with the recently released DSM-5 criteria for prescription opioid-use disorder, 5 however, we also counted the occurrence of opioid dependence symptoms, but excluded those for tolerance and withdrawal. In addition, we included one opioid-craving symptom and three prescription opioid abuse symptoms in this symptom count, but eliminated legal problems, as stipulated in the DSM-5.5 It is noted that the craving measure used was available for lifetime occurrence only. Thus, unlike our previous study,4 the current study is restricted to only lifetime opioid-use disorders. Typically, diagnostic interviews assess symptom counts for lifetime and for the past 12 months. 9,12 Based on these symptom counts, the DSM-5 nomenclature also included a severity classification, whereby patients

having no or few symptoms (<2), mild symptoms (2–3), moderate symptoms (4–5), and severe symptoms (six or more) were classified accordingly. In our previous DSM-5 comparison study,<sup>7</sup> this classification system was not finalized at the time and only included three categories: no or few symptoms (<2); moderate symptoms (2–3); and severe symptoms (four or more), which was a study limitation.

### Other study measures

Since our other study measures were discussed elsewhere, 4,7 we will briefly review them here. Our study also included alcohol dependence and tobacco (nicotine) dependence. Alcohol dependence was assessed based on DSM-4 criteria using the Composite International Diagnostic Interview scale. 12 The Fagerstrom Tolerance Scale (FTS) was used to assess nicotine dependence. 15,19 The FTS has good concurrent and predictive validity for nicotine dependence and it has been widely used.20 Major depression was assessed using a depression measure developed from the Structured Clinical *Interview for DSM*, <sup>11</sup> and it was validated in other surveys. <sup>21–23</sup> PTSD was based on the DSM-4, developed for telephone administration, and used in previous trauma studies. 9,24,25 We also used the Brief Pain Inventory to assess pain status, a widely used measure for chronic pain evaluation. <sup>26,27</sup> Trauma exposure was assessed using a childhood adversity and a lifetime trauma scale, respectively. Childhood adversity was assessed using a validated scale. 16,28,29 For trauma exposure, we used a measure that focused on the major lifetime traumatic events experienced (eg, forced sexual contact, being physically attacked, etc). This scale has been used and validated in previous studies. 9,24,25 Other study measures included a history of substance abuse treatments, psychotropic medication use, illicit drug use (eg, methamphetamine, heroin, cocaine use, etc), and self-reported health status.<sup>4,7</sup>

# Statistical analyses

With institutional review board approval, we analyzed selection bias using the EHR data to compare study respondents (total number [N] =705) to nonrespondents (N=1,434) by age, sex, and clinic type. We then used these results to develop case weights to adjust for possible selection bias. These weights corrected for the fact that women and specialty clinic patients tended to have higher survey response rates. Next, we examined the descriptive statistics for DSM-5 prescription opioid-use disorder by demographic and clinical risk factor characteristics (Tables 1 and 2). In these bivariate analyses, opioid-use disorder was dichotomized as moderate/severe (n=93) versus not moderate/severe (n=612), based on

the final APA criteria for the DSM-5.5 We also calculated point estimates for DSM-5 opioid-use disorder (with 95% confidence intervals [CIs]) and compared these to DSM-4 opioid dependence (Table 3). Next, we completed an ordinal logistic regression to identify the key risk factors for DSM-5 opioid-use disorder (Table 4). To identify the best adjusted model, variables with bivariate P-values < 0.05 were selected as covariates in these analyses. Variables that remained significant were retained in the final adjusted model. This adjusted regression model was used to assess risk factors for the severity of DSM-5 opioid-use disorder, classified as none/ few symptoms, mild symptoms, moderate symptoms, and severe symptoms, with "none/few" used as the reference category.<sup>30</sup> Since these patients were clustered within clinics, we also used the survey module in Stata (version 13.1) to adjust for patient clustering in all analyses.<sup>31</sup> Finally, it is noted that phone surveys produce similar results as in-person interviews. 32-34 The advantages of phone surveys include cost, patient privacy, convenience, and coverage. The limitation of phone surveys is that researchers have to simplify some survey response options because visual aids are typically unavailable during these interviews.<sup>32</sup> However, today, most diagnostic interviews and psychometric scales are available in both survey formats.<sup>21–24,32</sup>

#### **Results**

Nonresponse analyses suggested that nonparticipants tended to be male, not married, current smokers, seen in primary care clinics, and less physically ill than study participants (P<0.05). However, no differences were found in participation rates by race, employment status, obesity level, or the number of prescription opioid orders received in the past 3 years. Based on these results, case weights were developed to adjust for differences in participation by sex and clinic setting, and these weights were used in all analyses.

As shown in Table 1, using the final DSM-5 criteria, 13.2% of patients (95% CI =9.8–17.6) were classified as having moderate to severe lifetime opioid-use disorder. However, most demographic and clinical measures assessed were not associated with this disorder. Two noteworthy exceptions were for age less than 65 years (P<0.001) and for reporting fair/poor health status (P<0.01). The other opioid-use disorder associations found for medical demographic measures were for nonwhite race, being seen in a specialty clinic, and reporting greater pain impairment (P-values <0.05) (Table 1). By comparison, opioid-use disorder was significantly associated with a number of mental health comorbidities, including alcohol dependence,

Boscarino et al Dovepress

**Table I** Demographic and health characteristics of patients meeting the DSM-5 criteria for lifetime prescription opioid-use disorder (moderate or severe#)

Study variables	Total s	ample##		DSM-5 opioid-use disorder Moderate or severe		DSM-5 opioid-use disorder Not moderate or severe	
	N	Percent	(95% CI)	N	Percent	N	Percent
DSM-5 opioid-use disorder							
Moderate/severe							
% no	612	86.8	(82.4-90.2)				
% yes	93	13.2	(9.8–17.6)				
Age			, ,				
% 18–64 years old	558	79.3	(66.7–88.0)	90	16.1	468	83.9***
% 65+ years old	147	20.7	(12.0–33.3)	3	1.8	144	98.2
Sex			,				
% male	232	39.1	(34.3-44.1)	36	14.9	196	85.I
% female	473	60.9	(55.9–65.7)	57	12.1	416	87.9
Race			(55.7 55.1)	•			0
% white	694	98.4	(95.4–99.5)	90	12.8	604	87.2*
% nonwhite	11	1.6	(0.5–4.7)	3	34.2	8	65.8
Marital status			(0.0)	•	·	•	00.0
% married	458	65.6	(61.8–69.2)	58	12.8	400	87.2
% not married	247	34.4	(30.8–38.2)	35	14.0	212	86.0
Employment status	2.,,	5	(50.5 50.2)	33	1 1.0		00.0
% employed	179	25.7	(22.1–29.6)	19	11.9	160	89.1
% not employed	526	74.3	(70.4–77.9)	74	14.0	452	86.0
Household income	320	, 1.5	(70.1 77.7)	, ,	1 1.0	.52	00.0
% less than US\$30,000 per year	289	41.0	(33.0–49.5)	43	14.6	246	85.4
% greater than US\$30,000 per year	326	46.3	(40.8–52.1)	41	12.8	285	87.2
% refused/not reported	90	12.7	(8.6–18.3)	9	10.2	81	89.9
Education	70	12.7	(0.0–10.5)	,	10.2	01	07.7
% high school or less	351	50.2	(46.1–54.3)	41	11.7	310	88.3
% some college or more	352	49.8	(45.7–53.9)	52	14.7	300	85.3
Clinic setting	332	47.0	(43.7–33.7)	32	17.7	300	05.5
% primary care	563	83.I	(47.1–96.5)	68	12.2	495	87.8*
% specialty care	142	16.9	(3.5–52.9)	25	18.1	117	81.9
Reported health status	172	10.7	(3.3–32.7)	23	10.1	117	01.7
% good	570	80.8	(76.2–84.7)	61	10.7	509	89.3**
% fair/poor	135	19.2	(15.4–23.8)	32	23.6	103	76.4
•	133	17.2	(13.4–23.6)	32	23.6	103	70.4
Medically obese (BMI >30)	252	FO 3	(45.2 55.2)	40	127	204	07.4
% yes	353	50.3	(45.3–55.2)	49	13.6	304	86.4
% no	352	49.7	(44.8–54.7)	44	12.8	308	87.2
Reported average pain in past week	172	22.0	(10.4.30.3)	2.1	17.0	1.40	02.2
% high	173	23.8	(18.4–30.2)	31	17.8	142	82.2
% not high	532	76.2	(69.8–81.6)	62	11.7	470	88.3
Reported pain interference with life or							
work – greatly or extremely							
% yes	426	60.4	(55.4–65.1)	70	16.5	356	83.5*
% no	279	39.6	(34.9–44.6)	23	8.2	256	91.8
(N)	(705)			(93)		(612)	

**Notes:** "All percent results were adjusted/weighted for response bias and data clustering; Ns are unweighted. ""To aid in data interpretation, the total column is percentized by column and the disorder severity columns are percentized by row. \*P<0.05; \*\*P<0.05; and \*\*\*P<0.001.

Abbreviations: DSM-5, Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition; N, number; CI, confidence interval; BMI, body mass index.

major depression, generalized anxiety disorder, PTSD, childhood adversity, psychological trauma, illicit drug use, substance abuse treatment, psychotropic medication use, history of antisocial personality disorder, and other measures of psychological disturbance (*P*-values <0.01) (Table 2). We then cross-tabulated the results for patients based on the final DSM-5 versus the DSM-4

nomenclature, and these results are shown in Table 3. As seen, the majority of patients are now classified as having no or few symptoms (58.7%) and approximately one-third (28.1%) are classified as having mild symptoms. By comparison, only 13.2% (9.7% + 3.5% = 13.2%) are classified as having moderate to severe symptoms (Table 3). This DSM-5 classification appears to represent a departure from

**Table 2** Mental health characteristics of patients meeting the DSM-5 criteria for lifetime prescription opioid-use disorder (moderate or severe#)

Study variables	Total :	sample##		DSM-5 opioid-use disorder Moderate or severe		DSM-5 opioid-use disorder Not moderate or severe	
	N	Percent	(95% CI)	N	Percent	N	Percent
Lifetime alcohol dependence							
% yes	68	9.8	(7.1-13.3)	23	32.6	45	67.4**
% no	637	90.2	(86.7-92.9)	70	11.1	567	88.9
Lifetime tobacco dependence							
% yes	251	36.8	(32.1-41.9)	42	16.8	209	83.2*
% no	454	63.2	(58.2-68.0)	51	11.1	403	88.9
Lifetime major depressive disorder							
% yes	249	34.6	(29.8-39.7)	63	25.8	186	74.2***
% no	456	65.4	(60.3-70.2)	30	6.5	426	93.5
Lifetime generalized anxiety disorder							
% yes	89	12.6	(10.0–15.8)	26	30.1	63	69.9***
% no	616	87.4	(84.2–90.1)	67	10.8	549	89.2
Lifetime post-traumatic stress disorder							
% yes	97	13.3	(10.1–17.3)	29	30.6	68	69.4***
% no	608	86.7	(82.7–90.0)	64	10.5	544	89.5
History of high childhood adversity							
% yes	178	24.9	(21.3–28.8)	40	22.7	138	77.3***
% no	527	75. I	(71.2–78.7)	53	10.0	474	90.0
History of high exposure to psychological trauma							
% yes	161	23.0	(19.0–27.5)	41	24.6	120	75.4***
% no	544	77.0	(72.5–81.0)	52	9.8	492	90.2
History of illicit drug use							
% yes	167	24.1	(21.5-26.8)	39	22.7	128	77.3***
% no	432	75.9	(73.2-78.5)	54	10.2	484	89.8
History of any substance abuse treatment							
% yes	153	22.5	(20.3-25.0)	49	31.3	104	68.7***
, % no	552	77.5	(75.0–79.1)	44	7.9	508	92.1
Current psychotropic medication use			,				
% yes	434	61.1	(58.2-64.0)	81	18.7	353	81.3***
% no	271	38.9	(36.0–41.8)	12	4.5	259	95.5
Antisocial personality disorder	2/1	30.7	(30.0 11.0)	12	1.5	237	75.5
% yes	167	23.8	(21.6–26.1)	40	23.3	127	76.7**
			,				
% no	538	76.2	(73.9–78.4)	53	10.0	485	90.0
Ever sleeping problems							
% yes	461	64.4	(58.6–69.9)	83	18.1	378	81.9***
% no	244	35.6	(30.1–41.4)	10	4.3	234	95.7
Ever suicidal thoughts							
% yes	182	25.6	(22.2-29.4)	52	29.0	130	71.0***
% no	523	74.4	(70.6–77.8)	41	7.7	482	92.3
(N)	(705)			(93)		(612)	

**Notes:** \*\*All percent results were adjusted/weighted for response bias and data clustering; Ns are unweighted. \*\*\*To aid in data interpretation, the total column is percentized by column and the disorder severity columns are percentized by row. \*P < 0.05; \*\*P < 0.01; and \*\*\*P < 0.01.

Abbreviations: DSM-5, Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition; N, number; CI, confidence interval.

both the DSM-4 and the tentative DSM-5 criteria proposed by the APA in 2011.

Next, based on these bivariate results, a multivariate model was developed for the DSM-5 opioid-use disorder. Using the selection criteria discussed, eight predictor variables were identified for inclusion in the final model. Since the DSM-5 opioid-use disorder severity classifications include the presence of no/few symptoms, mild symptoms,

moderate symptoms, and severe symptoms, we used ordinal logistic regression to examine these outcomes (Table 4). As can be seen, the best predictors of higher opioid-use disorder severity include age less than 65 years, pain interference, trouble sleeping, suicidal thoughts, generalized anxiety, illicit drug use, and a history of substance abuse treatment (P-values <0.05). Among these variables, pain, trouble sleeping, suicidal thoughts, and substance abuse treatment are the

Table 3 Prevalence of lifetime DSM-5 prescription opioid-use disorder by lifetime DSM-4 prescription opioid dependence#

Diagnostic criteria*	DSM-5 opioid-use disorder: none/low (<2 symptoms)		DSM-5 opioid-use disorder: mild (2–3 symptoms)			DSM-5 opioid-use disorder: moderate (4–5 symptoms)			DSM-5 opioid-use disorder: severe (6+ symptoms)			
	%	95% CI	(n)	%	95% CI	(n)	%	95% CI	(n)	%	95% CI	(n)
DSM-4 no opioid dependence	86.0	82.2–89.1	(391)	14.0	11.0–17.8	(63)	0.0	_	(0)	0.0	_	(0)
DSM-4 opioid dependence	9.3	5.8-14.6	(23)	53.6	44.1-62.8	(135)	27.3	22.1-33.1	(69)	9.8	6.7-14.3	(24)
Total DSM-5 opioid-use disorder	58.7	54.2–63.1	(414)	28.1	25.4–30.9	(198)	9.7	7.3–12.8	(69)	3.5	2.2–5.5	(24)

Notes: "All percent results are adjusted/weighted for response bias and data clustering; ns are unweighted. \*P<0.0001.

**Abbreviations:** DSM-5, Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition; DSM-4, Diagnostic and Statistical Manual of Mental Disorders, Fourth Edition; CI, confidence interval; n, number.

most significant predictors of lifetime DSM-5 opioid-use disorder severity (*P*-values < 0.01).

#### **Discussion**

Based on the preliminary DSM-5 criteria available at the time, our earlier study suggests that 35% (95% CI =30.5–39.5) of the patients on long-term opioid therapy met the criteria for lifetime prescription opioid-use disorder. This was approximately the same percent that met the DSM-4 criteria for opioid dependence. The kappa coefficient for those meeting these preliminary DSM-5 and final DSM-4 criteria in our earlier study was high (kappa =0.873; P<0.0001), suggesting considerable overlap. This finding was surprising at the time, given the changes proposed in the DSM-5. As was seen in the current study, this does not appear to be the case using the final severity classifications issued by the APA in 2013, which included having no/few symptoms (<2), mild symptoms (<3), moderate symptoms (<5), and severe symptoms (<6). Thus, under the final APA nomenclature,

**Table 4** Ordinal logistic regression model predicting lifetime DSM-5 opioid-use disorders (N=705)#

OR	95% CI	P-value
1.86	1.14-3.04	0.013
2.01	1.38-2.93	< 0.001
1.81	1.28-2.55	0.001
1.65	1.29-2.12	< 0.001
1.63	1.11-2.38	0.013
1.40	0.99-2.00	0.058
1.75	1.10-2.78	0.018
2.28	1.59-3.26	< 0.001
2.23	1.66-2.81	-
4.05	3.43-4.68	-
5.66	4.89-6.42	_
	1.86 2.01 1.81 1.65 1.63 1.40 1.75 2.28 2.23 4.05	1.86 1.14–3.04 2.01 1.38–2.93 1.81 1.28–2.55 1.65 1.29–2.12 1.63 1.11–2.38 1.40 0.99–2.00 1.75 1.10–2.78 2.28 1.59–3.26 2.23 1.66–2.81 4.05 3.43–4.68

Notes: "All of the results are adjusted/weighted for survey response bias and data clustering. Ordinal contrasts include no opioid-use disorder (n=414), mild opioid-use disorder (n=198), moderate opioid-use disorder (n=69), and severe opioid-use disorder (n=24). Cuts 1-3 represent the coefficients used to estimate an ordinal model with 4 categories, representing no disorder, mild, moderate, and severe disorder.

**Abbreviations:** DSM-5, Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition; N, total number; OR, odds ratio; Cl, confidence interval; n, sample number.

only 13.2% of patients meet the criteria for moderate to severe opioid-use disorder, which is the subgroup that would likely be of highest risk for addiction. However, 41.3% (95% CI =37.6–45.0) of patients now met the criteria for having any lifetime prescription opioid-use disorder.

In the bivariate analyses, being younger and reporting a poorer health status were significantly associated with DSM-5 opioid-use disorder severity (Table 1), as was alcohol dependence, tobacco dependence, major depression, generalized anxiety disorder, a history of childhood adversity, exposure to psychological trauma, illicit drug use, substance abuse treatment, psychotropic medication use, a history of antisocial personality, sleep disturbance, and suicidal thoughts (Table 2). Noteworthy is that these are mostly the same risk factors that were previously reported for DSM-4 opioid dependence.<sup>4,35</sup>

In the multivariate analysis using ordinal logistic regression to predict opioid-use disorder severity (ie, no/ few, mild, moderate, or severe symptoms), significant risk factors included younger age, pain impairment, sleeping problems, suicidal thoughts, generalized anxiety, illicit drug use, and a history of substance abuse treatment (Table 4). It was noted that these multivariate risk factors for prescription opioid-use disorder are similar to those reported for DSM-4 opioid dependence, with the exception of a history of drug abuse and severity of addiction, both of which are now explicitly included in the DSM-5 and therefore were not assessed in the current study. Given the scope of the changes proposed for the DSM-5, these outcomes were not expected.

This study has its strengths and limitations. The study strengths include that this research was based on a random sample of outpatients seen in a large multispecialty group practice; that drug-use disorder was assessed based on the final DSM-5 criteria; and that subjects were identified through drug orders in the EHR, not based on patient self-report or treatment records. Study limitations include the following:

that our survey data were based on patient self-report; our survey completion rate was less than optimal; our current study only included lifetime opioid-use disorder; our drug-craving measure was limited; and our patients were drawn from a predominately Caucasian population in one US region. In addition to our low response rate, males were also underrepresented, as were nonwhite patients, all of which could have further skewed our results and may have affected the representativeness and validity of our study. As is common is survey research, however, we did use survey weights to adjust the data for potential response bias. 9-11

To our knowledge, this is one of the few studies to compare the final DSM-5 criteria to DSM-4 for prescription opioids and to report the results for DSM-5 moderate and severe opioid-use disorders.<sup>36</sup> We did not expect the prevalence of the DSM-5 disorders to vary significantly because our previous study used the criteria initially proposed by the APA in 2011 and found little variation. As shown in Table 3, however, there is more disparity than previously reported due to the changes in the final APA severity classifications. Nevertheless, the current findings appear to be consistent with recently reported observations.<sup>36</sup> The DSM-4 prevalence estimates we reported in 2010 appeared to be higher than those typically reported in the literature at the time.<sup>4,37</sup> As was shown, the lifetime prevalence of DSM-5 opioid-use disorders using the final DSM criteria was 28.1% for mild symptoms, 9.7% for moderate symptoms, and 3.5% for severe symptoms, suggesting that 41.3% of the patients had some symptoms (among those with more than one symptom), and 13.2% had moderate to severe symptoms. As we have noted, the prevalence of prescription opioid abuse in the USA has increased in the past decade, representing a major public health problem.<sup>1-3</sup> Our objective is to replicate these findings and to investigate key correlates of the DSM-5 criteria in the future. 4,35 Given that the DSM-5 addiction criteria were developed with a greater focus on the existing empirical evidence rather than on clinical consensus, 6,38 perhaps we will be more likely to discover robust correlates of these disorders,<sup>39</sup> which was a rationale for the DSM diagnostic nomenclature. 40 We now know that most psychiatric disorders are inherently complex and represent a multitude of genetic and environmental risk factors. 41 Given the refinement of the diagnostic criteria under the DSM-5, this discovery process might be more fruitful. As discussed, these changes included the elimination of the tolerance and withdrawal symptoms for those taking opioids under medical supervision, the elimination of legal problems, and the inclusion of abuse, dependence, and craving symptoms, all combined now into a single disorder of graded severity.

#### **Conclusion**

From a public health policy view, misuse and abuse of prescription opioids in the USA has increased significantly.<sup>42</sup> However, it has been noted recently that opioid dispensing and prescription opioid overdoses have actually decreased in the USA following major changes in the pharmaceutical market in late 2010.43 This included the introduction of abuse-deterrent extended-release oxycodone hydrochloride and the withdrawal of propoxyphene. 43 These two pharmaceutical market interventions appear to have reduced the prescription opioid overdoses, but heroin use rates have continued to increase.<sup>3,43</sup> The latter points to the complexity of contemporary addictionrelated disorders. 44,45 Opioid therapy for noncancer pain is still commonplace today, despite the lack of clinical evidence as to its benefit.<sup>2,46</sup> Previously, we developed a "risk-score" method to identify patients at risk for PTSD that included genetic risk information.<sup>47</sup> In the future, it might be possible to use data routinely collected in the EHR, together with the other pertinent patient information, to identify patients at highest risk for opioid abuse. 48 We think that the recently developed DSM-5 nomenclature for prescription opioid-use disorders could aid in this clinical endeavor.

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#### **Disclosure**

The authors report no conflicts of interest in this work.

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