### Research methods in psychiatry

# Development of a checklist of short-term and long-term psychological symptoms associated with ketamine use

Ni FAN<sup>1</sup>, Ke XU<sup>2</sup>, Yuping NING<sup>1</sup>, Daping WANG<sup>1</sup>, Xiaoyin KE<sup>1</sup>, Yi DING<sup>1</sup>, Bin SUN<sup>1</sup>, Chao ZHOU<sup>1</sup>, Xuefeng DENG<sup>3</sup>, Robert ROSENHECK<sup>2</sup>, Hongbo HE<sup>1,\*</sup>

**Background:** Ketamine is an increasingly popular drug of abuse in China but there is currently no method for classifying the psychological effects of ketamine in individuals with ketamine dependence.

**Aims:** Develop a scale that characterizes the acute and long-term psychological effects of ketamine use among persons with ketamine dependence.

**Methods:** We developed a preliminary symptom checklist with 35 dichotomous ('yes' or 'no') items about subjective feelings immediately after ketamine use and about perceived long-term effects of ketamine use that was administered to 187 inpatients with ketamine dependence recruited from two large hospitals in Guangzhou, China. Exploratory factor analysis (EFA) was conducted on a randomly selected half of the sample to reduce the items and to identify underlying constructs. Confirmatory factor analysis (CFA) was conducted on the second half of the sample to assess the robustness of the identified factor structure.

**Results:** Among the 35 symptoms, the most-reported acute effects were 'floating or circling' (94%), 'euphoric when listening to rousing music' (86%), and 'feeling excited, talkative, and full of energy' (67%). The most-reported long-term symptoms were 'memory impairment' (93%), 'personality changes' (86%), and 'slowed reactions' (81%). EFA resulted in a final 22-item scale best modelled by a four-factor model: two factors representing chronic symptoms (social withdrawal and sleep disturbances), one about acute psychotic-like symptoms, and one that combined acute drug-related euphoria and longer-term decreased libido. CFA showed that these 4 factors accounted for 50% of the total variance of the final 22-item scale and that the model fit was fair (Goodness of Fit Index, GIF=83.3%; Root Mean Square Error of Approximation, RMSEA=0.072).

**Conclusions:** A four-factor model including social withdrawal, sleep disturbance, psychotic-like symptoms, and euphoria at the time of drug use provides a fair description of the short-term and long-term psychological symptoms associated with ketamine use. Future work on the 22-item version of the scale with larger samples is needed to confirm the validity of this 4-factor structure, to assess the scale's test-retest reliability, and to determine whether or not it can be useful in the differential diagnosis and monitoring of treatment of individuals with ketamine dependence.

Keywords: ketamine; substance-related disorders; signs and symptoms; questionnaire; factor analysis; China

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<sup>&</sup>lt;sup>1</sup> Guangzhou Brain Hospital (Guangzhou Hui Ai Hospital), Guangzhou Medical University, Guangzhou, Guangdong Province, China

<sup>&</sup>lt;sup>2</sup> Department of Psychiatry, Yale University School of Medicine, New Haven, CT, United States

<sup>&</sup>lt;sup>3</sup> Guangzhou Baiyun Voluntary Drug Rehabilitation Hospital, Guangzhou, Guangdong, China

<sup>\*</sup>correspondence: vglut2@126.com

#### 1. Background

Ketamine is an N-methyl-D-aspartate (NMDA) receptor antagonist that can induce various effects including sedation, relaxation, euphoria, and hallucinations. At low doses, ketamine induces sedation and pain relief; at high doses it can induce an anesthetic dissociative state. Since ketamine was first synthesized in the 1960s, it has been widely used in anesthesia and pain management and is currently in the WHO Directory of Essential Drugs. [1] However, during recent decades ketamine has been increasingly used as a recreational drug. In China, where ketamine is referred to as 'K-powder', it is becoming a popular drug of abuse among illegal drug users, accounting for 21.5% of all substance abuse disorders in 2004 and 40% of all substance abuse disorders in 2009. [2-5] Studies have found that a single intravenous injection of a sub-anesthetic dose of ketamine (0.5mg/kg) in individuals with refractory depression can induce a pronounced anti-depressant effect within 1 to 2 hours that is sustained for up to one week. [6,7] Among healthy volunteers, a single intravenous injection of ketamine can induce schizophrenia-like symptoms. [8-10] Compared with the predominantly positive psychotic symptoms produced by amphetamines, ketamineinduced psychotic symptoms include positive symptoms (i.e., hallucinations and delusions), negative symptoms, and cognitive impairment. [11] Ketamine use has also been associated with cystitis and other physical problems. [12] Based on these findings, some researchers have proposed using a ketamine model of schizophrenia to study the glutamate neural pathway believed to be important in schizophrenia.[13]

Despite its increasing popularity in China, there have been few reports about the effects of ketamine use among chronic ketamine users. To provide a profile of ketamine-related symptoms in this population, we developed a preliminary 35-item symptom checklist of ketamine-related symptoms based on our clinical experience with such patients, administered it to a sample of ketamine users recruited from two hospitals in Guangzhou (a major metropolitan city in south China), and used exploratory and confirmatory factor analyses to identify the subset of items in the checklist that best represent the underlying clinical domains related to the short-term and long-term symptoms of ketamine use.

#### 2. Methods

#### 2.1 Recruitment

The recruitment of participants in the study is shown in Figure 1. The Department of Substance Dependence at Guangzhou Brain Hospital has 40 beds, and the Voluntary Rehabilitation Ward of the Guangzhou Baiyun Mental Hospital has 100 beds. Two psychiatrists and a graduate student from the Guangzhou Brain Hospital recruited all participants, visiting the Baiyun Mental Hospital twice a week over the study enrollment period. They screened a convenience sample of 630 individuals with a history of ketamine use at the time of inpatient admission to the two participating hospitals from January 2012 to December 2013. Participants meeting the following inclusion and exclusion criteria were considered for the study: (a) receiving inpatient ketamine detoxification treatment or treatment for symptoms related to ketamine use; (b) met the DSM-

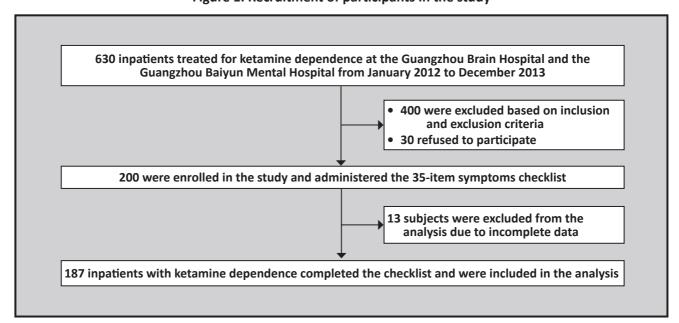


Figure 1. Recruitment of participants in the study

IV-TR<sup>[14]</sup> diagnostic criteria for substance dependence; (c) no history of dependence on psychoactive drugs other than ketamine and tobacco; (d) other than tobacco and alcohol, did not use other drugs besides ketamine during the prior 6 months; (e) no serious physical illness; (f) had no history of a head injury that resulted in loss of consciousness; (g) no significant visual or hearing impairments; and (h) no history of other mental disorders. All eligible individuals were asked to participate in the study. Persons who signed the informed consent subsequently participated in a 2-hour interview about their current drug use patterns that included completion of a 35-item symptom checklist (described below).

This study was approved by the research ethics committee of the Guangzhou Brain Hospital.

#### 2.2 Assessment

To our knowledge, there is no previous report of a scale used to assess clinical symptoms related to ketamine abuse. We constructed a preliminary 35-item symptom checklist that assess 19 acute effects of ketamine use (i.e., those that occur immediately after using ketamine) and 16 chronic symptoms that emerge after long-term use of ketamine (shown in Table 2). The items included in this preliminary checklist were based on a previous report that included individuals' descriptions of their experiences related to ketamine use[15] and on our own knowledge and clinical experience with ketamine dependence. Each of the 35 items was rated on a binary scale ('yes' or 'no') based on patients' responses to a structured interview conducted by a psychiatrist. All interviews were conducted within one week of admission. (The Chinese version of the checklist used in the study is available with the Chinese translation of this article on the website of the Shanghai Archives of Psychiatry).

We also collected basic demographic information on each subject, and information about the age of onset, duration, and frequency of use of ketamine and other substances including tobacco and alcohol.

#### 2.3 Data analysis

A total of 13 patients had incomplete data, so the analysis was based on data from 187 individuals. The results for these 187 individuals were randomly assigned to two groups using a random number table. The first group (n=94) was used for the exploratory factor analysis (EFA) and the second group (n=93) was used for the confirmatory factor analysis (n=93).

The scree plot from the initial EFA (on the first half of the sample) of all 35 items and consideration of the item loadings in different models (i.e., the 3-factor, 4-factor, and 5-factor models) were used to decide on the best model to represent the results. The item-reduction process was conducted as follows. For each of the three possible models we removed items that had loadings of less than 0.30 on all factors or that had

loadings of greater or equal to 0.30 on two or more factors (i.e., cross-loading). After removing these items the EFA was re-run and the loadings re-examined; if there were still items with no loadings of 0.30 or greater or items with cross-loadings they were removed and the EFA was re-run on the remaining items. This process was repeated until all remaining items had loadings of 0.30 or greater on only one factor. The three potential final models were then examined and the model in which the factors contained the most logical combination of items (symptoms) was selected as the final model.

This final model with factor-specific items was then tested using CFA based on the results of the second group of respondents. The validity of the factor structure identified in the EFA was assessed using various fit indices including the Goodness of Fit Index (GFI) and the Root Mean Square Error of Approximation (RMSEA). The internal consistency of the subscales of the final questionnaire was assessed using alpha values. SPSS 18.0 and Mplus (version 7) software was used for data analyses.

#### 3. Results

#### 3.1 Sample characteristics

Table 1 shows the characteristics of the sample. A total of 187 individuals completed the interviews including 173 (92.5%) males and 14 (7.5%) females. Their mean (sd) age was 26.2 (5.0) years (range=15 to 44). The mean duration of ketamine use was 6.3 (3.1) years and the mean duration of ketamine dependence was 3.1 (2.0) years. Most participants were using ketamine daily prior to admission. The mean amount of ketamine used during the prior month was 3.4 (2.7) g/day. Our enrollment criteria excluded individuals who had previously been dependent on other illegal substances; nevertheless, the majority of these ketamine-dependent individuals had previously used (but not been dependent on) other types of illegal psychoactive substances.

#### 3.2 Acute and chronic symptoms of ketamine use

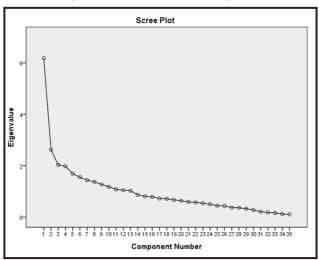
The reported prevalence of short-term and long-term effects of ketamine use are listed in Table 2. The most common short-term effects experienced immediately after using ketamine were 'weightless floating or circling' (94%), 'euphoric when listening to rousing music' (86%), and 'excited, talkative, and full of energy' (67%). The most common long-term symptoms the individuals reported after using ketamine for more than a year were 'memory loss' (93%), 'personality changes' (86%), and 'slow reaction time' (81%).

## 3.3 Item reduction and characteristics of the final model

As shown in Figure 2, the EFA of the initial 35 items in the first half of the sample identified 12 factors with eigenvalues >1. Based on the scree plot, we considered

Table 1. Characteristics of the 187 participants							
Male, n (%)	173 (92.5%)						
Age in years, mean (sd)	26.2 (4.95)						
Education in years, mean (sd)	10.5 (2.73)						
Marital status (single, n [%])	101 (54.0%)						
Employed, n (%)	135 (72.2%)						
Family history of mental illness, n (%)	12 (6.4%)						
Family history of drug abuse, n (%)	59 (31.6%)						
Past admission due to ketamine use, n (%)	77 (41.2%)						
Age of the first ketamine use, mean (sd)	20.0 (5.0)						
Age of onset of dependence, mean (sd)	23.1 (5.0)						
Years of ketamine use, mean (sd)	6.3 (3.1)						
Years of ketamine dependence, mean (sd)	3.1 (2.0)						
Average dosage before admission (g/day)	3.4 (2.7)						
Maximum dosage before admission (g/day)	6.9 (6.0)						
Ketamine use before admission (n, %)							
every day ≥4 times/week <4 times/week	141 (75.4%) 14 (7.5%) 32 (17.1%)						
History of other psychoactive drug use, n (%)	158 (84.5%)						
Current smoker <sup>a</sup> , n (%)	180 (96.3%)						
Current alcohol drinker <sup>b</sup> , n (%)	124 (66.3%)						
<sup>a</sup> smokes at least 3 days every week <sup>b</sup> drinks alcohol at least one day per week over prior six months							

Figure 2. Scree plot of exploratory factor analysis of a 35-item checklist of short-term and longterm symptoms of ketamine use among 94 inpatients with ketamine dependence



three possible models, a 3-factor model, a 4-factor model, and a 5-factor model. After completing the item-reduction steps (described in the methods) for each of these models, the 22-item version of the 4-factor model provided the most coherent grouping of items (data provided on request), so this was selected as the final model and as the final version of the scale.

The loadings of the 22 items on these 4 factors in the EFA are shown in the Table 3. In the EFA, the first factor included 5 long-term symptoms associated with social withdrawal; the second factor included 4 longterm symptoms associated with sleep disturbance; the third factor included 8 acute symptoms associated with psychotic-like experiences and irritability; and the fourth 'other symptoms' factor included 5 symptoms that combined long-term reports of decreased sex drive and reports of excitement and euphoria at the time of taking ketamine. Based on the results using the first random half of the sample, these 4-factors accounted for 49% of the variance in the 22-item scale and the internal consistency (based on Cronbach alpha) of the four factors – the subscales of the overall scale – range from 0.63 to 0.86.

#### 3.4 Assessment of the final model for the scale

The validity and robustness of this 22-item, 4-factor model of the acute and chronic symptoms of ketamine abuse identified using EFA based on the first half of the sample was assessed using CFA based on the second half of the sample. As shown in Table 3, the distribution of items across factors and the loading of the items of the 4 factors in the CFA is very similar to that in the EFA. All the items loaded on the appropriate factors, and, with the exception of the 'weightless floating or circling' item, all items had loadings of 0.30 or greater on a single factor. In the CFA these 4 factors accounted for 50% of the variance. The internal consistency of the items included in the social withdrawal and sleep disturbance subscales was excellent (alpha=0.86 and 0.82, respectively), that for the psychotic-like symptoms subscale was fair (0.69), and that for the 'other symptoms' factor was poor (0.41). The correlation between the factors were all quite low, ranging from -0.03 to 0.28. The fit parameters for this 4-factor model were fair (GFI=83.3%: RMSEA= 0.072, 90% confidence interval=0.062, 0.083).

#### 4. Discussion

#### 4.1 Main Finding

In this study, we found that among the 19 short-term symptoms of ketamine use, the most common symptoms reported by individuals who are dependent on ketamine are a floating or circling sensation, euphoria, and excitement. Among the 16 long-term symptoms considered, the most commonly reported symptoms were memory loss, personality changes, slow reaction time, lethargy, and decreased libido.

Table 2. Prevalence of ketamine-related symptoms in 187 inpatients with ketamine dependence and loadings on 4-factor model in exploratory factor analysis (with varimax rotation) in a random sample of 94 of these individuals

Reported symptoms		alence	loading on 4 factors				
		%	1	2	3	4	
SHORT-TERM SYMPTOMS							
Weightless floating or circling	176	94.1%	0.093	0.166	<u>0.381</u>	0.129	
2. Euphoric when listening to rousing music	161	86.1%	0.024	0.036	<u>0.509</u>	0.038	
3. Difficulty falling asleep when used before bedtime	143	76.5%	0.069	0.023	<u>0.395</u>	0.220	
4. Excited, talkative, and full of energy	126	67.4%	<u>0.307</u>	0.130	0.055	0.153	
5. Hallucinations	100	53.5%	0.085	<u>0.713</u>	0.187	0.028	
6. Out of body experience	97	51.9%	0.187	0.148	0.241	-0.007	
7. Distorted or twisted body parts	92	49.2%	0.018	<u>0.364</u>	<u>0.317</u>	0.163	
8. Restlessness, irritation, and short temper	90	48.1%	0.278	<u>0.398</u>	0.132	-0.040	
9. Visual hallucinations	63	33.7%	0.064	<u>0.516</u>	0.220	0.068	
10. Sexual arousal	63	33.7%	-0.073	0.093	0.104	-0.113	
11. Auditory hallucinations	59	31.6%	0.075	<u>0.713</u>	0.051	-0.047	
12. Coma or loss of consciousness	55	29.4%	0.133	0.063	0.162	0.025	
13. Stiffness as if being made of wood, rubber or plastic	55	29.4%	0.050	0.284	0.072	0.299	
14. Electric currents running through the body	33	17.6%	-0.065	<u>0.372</u>	0.263	-0.002	
15. Delusions	20	10.7%	0.151	<u>0.543</u>	-0.034	0.096	
16. Violence	17	9.1%	0.148	<u>0.346</u>	0.001	-0.065	
17. Tactile hallucinations	12	6.4%	0.043	<u>0.527</u>	-0.147	0.130	
18. Parallel existence with the universe	7	3.7%	0.004	0.165	0.193	<u>0.322</u>	
19. Sudden realization of the mystery of existence and self	4	2.1%	-0.003	-0.046	0.241	0.161	
LONG-TERM SYMPTOMS							
20. Memory loss	173	92.5%	0.276	-0.036	<u>0.435</u>	0.190	
21. Personality changes	160	85.6%	<u>0.684</u>	0.183	0.206	0.074	
22. Slow reaction time	151	80.7%	<u>0.515</u>	-0.093	<u>0.488</u>	0.057	
23. Lethargy	131	70.1%	<u>0.715</u>	0.079	0.267	-0.002	
24. Low libido	131	70.1%	0.064	0.163	<u>0.734</u>	-0.122	
25. Decreased work efficiency	131	70.1%	<u>0.570</u>	0.035	<u>0.401</u>	0.124	
26. Reduced quality of sleep	126	67.4%	0.197	-0.032	0.107	<u>0.848</u>	
27. Decreased sex drive	118	63.1%	0.037	0.127	<u>0.689</u>	-0.142	
28. Unsociable and eccentric	118	63.1%	<u>0.816</u>	0.103	-0.021	0.160	
29. Reduced social activities	114	61.0%	<u>0.837</u>	0.103	-0.070	0.165	
30. Difficulty falling asleep	106	56.7%	0.120	-0.030	0.098	<u>0.793</u>	
31. Easily waken during sleep	90	48.1%	0.145	0.019	0.075	<u>0.749</u>	
32. Short temper or bad temper	83	44.4%	<u>0.330</u>	<u>0.330</u>	0.191	0.157	
33. Reduced number of friends	79	42.2%	<u>0.711</u>	0.093	-0.088	0.194	
34. Paranoia	60	32.1%	<u>0.369</u>	<u>0.330</u>	0.029	0.005	
35. Early awakening	39	20.9%	0.145	0.123	-0.045	<u>0.579</u>	
Eigenvalue of each factor			6.18	2.62	2.03	1.98	
Proportion of total variance of each factor			17.70%	7.50%	5.80%	5.66%	

Table 3. Factor loadings (with varimax rotation) of the final 22-item, 4-factor model in the exploratory factor analysis (n-94) and the confirmatory factor analysis (n=93)<sup>a</sup>

	Factor loadings of 22 retained items based on 4-factor exploratory factor analysis				Factor loadings of 22 retained items based on 4-factor confirmatory factor analysis			
	1	2	3	4	1	3 <sup>b</sup>	<b>2</b> <sup>b</sup>	4
Reduced social activities	0.88	0.16	0.10	-0.01	0.88	0.14	0.12	-0.03
Unsociable and eccentric	<u>0.86</u>	0.16	0.09	0.03	<u>0.85</u>	0.15	0.12	0.02
Reduced number of friends	<u>0.73</u>	0.18	0.10	-0.07	<u>0.74</u>	0.18	0.08	-0.06
Personality changes	<u>0.70</u>	0.11	0.16	0.27	<u>0.72</u>	0.09	0.14	<u>0.31</u>
Lethargy	<u>0.70</u>	0.05	0.12	0.24	<u>0.71</u>	0.05	0.09	0.28
Reduced quality of sleep	0.16	<u>0.90</u>	0.00	0.11	0.17	<u>0.90</u>	-0.01	0.12
Difficulty falling asleep	0.07	<u>0.83</u>	-0.02	0.11	0.07	<u>0.83</u>	-0.06	0.15
Easily waken during sleep	0.12	<u>0.77</u>	0.05	0.05	0.12	<u>0.78</u>	0.07	0.02
Early awakening	0.16	<u>0.55</u>	0.09	-0.05	0.16	<u>0.58</u>	0.17	-0.11
Hallucinations	0.05	0.07	<u>0.79</u>	0.22	0.00	0.11	<u>0.72</u>	0.29
Auditory hallucinations	0.07	0.01	<u>0.72</u>	0.15	0.02	0.03	<u>0.69</u>	0.16
Visual hallucinations	0.00	0.09	<u>0.62</u>	0.17	0.04	0.11	<u>0.57</u>	0.21
Delusions	0.13	0.10	<u>0.52</u>	0.02	0.14	0.06	<u>0.51</u>	0.00
Tactile hallucinations	0.04	0.09	<u>0.51</u>	-0.19	0.03	0.12	<u>0.53</u>	-0.08
Restlessness, irritation, and short temper	0.25	-0.03	<u>0.42</u>	0.08	0.19	-0.03	<u>0.53</u>	0.01
Violence	0.11	-0.05	<u>0.39</u>	-0.09	0.11	-0.17	<u>0.36</u>	-0.13
Electric currents running through the body	-0.02	-0.06	<u>0.35</u>	0.19	0.00	-0.03	<u>0.49</u>	0.11
Low libido	0.05	-0.11	0.13	<u>0.83</u>	-0.02	-0.07	0.20	<u>0.82</u>
Decreased sex drive	0.05	-0.13	0.07	<u>0.82</u>	-0.01	-0.06	0.15	<u>0.79</u>
Euphoric when listening to rousing music	0.04	0.10	0.02	<u>0.51</u>	0.09	0.06	0.03	<u>0.55</u>
Difficulty falling asleep when used before bedtime	0.10	0.23	0.00	<u>0.41</u>	0.19	0.18	-0.05	<u>0.43</u>
Weightless floating or circling	0.10	0.16	0.18	<u>0.33</u>	0.19	0.21	0.12	<u>0.25</u>
Eigenvalue of each factor	4.62	2.52	1.86	1.76	4.69	1.97	2.58	1.68
Proportion of total variance of each factor	21.0%	11.5%	8.5%	8.0%	21.3%	9.0%	11.8%	7.7%
Alpha value of items included in each factor	0.86	0.80	0.69	0.63	0.86	0.81	0.69	0.41

<sup>&</sup>lt;sup>a</sup> 4 Factors included: (1) Social withdrawal; (2) Sleep disturbance; (3) Psychotic-like symptoms; and (4) Other symptoms. Cumulative variance explained by the 4 factors was 48.9% in the exploratory factor analysis and 49.7% in the confirmatory analysis. For the confirmatory factor analysis, GFI=83.3% and RMSEA= 0.072 (90% confidence interval, 0.062, 0.083)

A series of studies by Muetzelfeldt and colleagues, [15] Curran and colleagues, [16] and Morgan and colleagues, [17] used a 12-item, self-report questionnaire to ask 20 recreational ketamine users about their subjective experiences with ketamine on the night of use, 3 days later, and 3 years later. They observed a 80% drop in the amount of ketamine use (in both frequency and dosage) among these recreational users three years after the baseline assessment. They reported that

commonly occurring short-term effects included bodily numbness, impaired concentration and memory, lack of coordination, unsteadiness, out of body experiences, altered perception, and altered reality; the impaired concentration and memory persisted at the 3-day and 3-year follow-up assessments. Our findings parallel these results, but in our sample of inpatients with chronic ketamine dependence (who used higher doses of ketamine for more prolonged periods than recreation

<sup>&</sup>lt;sup>b</sup> To simplify the presentation in the table, factors 2 and 3 in the confirmatory factor analysis is are reversed so that the order of factors matches that in the exploratory factor analysis

users) social withdrawal and sleep disturbance were also prominent long-term symptoms.

Like ketamine use, amphetamine use has been associated with euphoria and a high prevalence of hallucinations but, somewhat unlike the symptoms of ketamine use, amphetamine use is also commonly associated with the occurrence of different types of delusions (reported in 30% to 52% of users) including erotomania, persecutory delusions, delusions of jealousy, grandiose delusions, and delusions of control. [18] Our study found delusions a relatively uncommon shortterm symptom following ketamine use, occurring in only 11% of the sample. A previous study that compared the occurrence of various psychiatric symptoms between amphetamine and ketamine users using the Brief Psychiatric Rating Scale [19] found higher levels of cognitive dysfunction, motor hyperactivity, and hostility among amphetamine users and higher levels of anxiety, depression, and motor retardation among ketamine

Exploratory factor analysis identified a subset of 22 items from the original pool of 35 items that segregated into four factors, representing the underlying groupings of symptoms that occur with ketamine use. The confirmatory factor analysis conducted on the second half of the sample confirmed the validity of this 22item, 4-factor structure, but the model fit parameters (GFI=83%, RMSEA=0.072) were only fair, so further work will be needed to refine the scale. The first three factors considered by the final version of the scale - longterm social impairment, long-term sleep disturbances, and short-term psychotic-like symptoms - appeared reasonably consistent and theoretically feasible; however, the final factor – which combines short-term euphoria and long-term deceases in sexual functioning - had poor internal consistency so it may not be a useful construct for understanding ketamine-related symptoms.

#### 4.2 Limitations

All participants were inpatients seeking voluntary treatment for addiction, so it is uncertain whether or not the pattern of symptoms they report are similar to those of ketamine users who do not seek treatment. We tried to eliminate the effect of other psychoactive drugs by excluding patients who had used other psychoactive drugs (other than tobacco and alcohol) in the previous 6 months, but over 80% of participants reported prior use of other psychoactive drugs more than 6 months previously so we cannot be sure that all the long-term symptoms they report are actually due to ketamine use. We consulted previous reports and used our clinical experiences in the construction of the preliminary 35item symptom checklist, but the list may have missed some relevant symptoms. Moreover, the checklist focuses on subjective feelings and mental symptoms so it does not include potentially important somatic symptoms related to ketamine use (e.g. urinary tract symptoms). [20] Conducting exploratory and confirmatory factor analysis usually requires large samples, so our sample of 187 individuals may not have been large enough to generate stable results from the factor analysis, particularly because 4 of the 35 symptoms in the preliminary scale were reported by less than 10% of respondents. Moreover, the inter-rater reliability and test-retest reliability of the symptom checklist was not assessed. Finally, it is uncertain how sensitive the symptom checklist will be to changes in the prevalence and severity of long-term symptoms of ketamine use during periods of abstinence.

#### 4.3 Implications

Despite the increasing popularity of ketamine abuse in contemporary China, the symptoms related to ketamine use are poorly understood. To the best of our knowledge, this is the first report on the profile of short-term and long-term symptoms that are associated with ketamine use in China. We administered an initial symptom checklist with 35 items selected from reports in other countries and our own clinical experience to a large sample of individuals with ketamine dependence and used exploratory and confirmatory factor analysis to identify the 22 symptoms that best capture the shortterm and long-term clinical symptoms of ketamine abuse. These 22 key symptoms segregate into four dimensions, two sets of long-term symptoms (i.e., social withdrawal and sleep disturbance), one set of psychoticlike short-term symptoms, and one dimension that included both short-term items related to euphoria and excitement at the time of ketamine use and items about decreased sex drive that occurs after long-term use of ketamine. The first three dimensions appear reasonably robust but the last dimension is less internally consistent. Future work with this scale in larger samples is needed to confirm this 4-factor structure of ketamineuse-related symptoms, to assess the test-retest reliability of the scale, and to determine whether or not the scale can be useful in the differential diagnosis and monitoring of treatment of individuals with ketamine dependence.

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#### **Conflict of Interest**

The authors declare no conflict of interest related to this manuscript.

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#### **Ethical approval**

The study was approved by the research ethics committee of the Guangzhou Brain Hospital.

#### Informed consent

All participants provided written informed consent to participate in this study.

#### 与氯胺酮滥用相关的短期和长期心理症状清单的编制

范妮,许柯,宁玉萍,王达平,柯晓殷,丁毅,孙彬,周超,邓雪峰,Rosenheck R,何红波

**背景:** 氯胺酮滥用在中国日渐增多,但目前还没有一个方法可以将氯胺酮对依赖个体的心理影响作用进行分类。

**目的**:编制能反映氯胺酮依赖人群中因该物质滥用所致的急性心理影响和长期心理影响的量表。

方法: 我们编制了一个初步的症状清单,共有 35 个条目("是"或"否"的二分法问题),内容是关于使用氯胺酮后即刻产生的主观感受以及氯胺酮滥用后感受到的长期作用。从中国广州市两个大医院招募氯胺酮依赖住院患者,共入组 187 例。随机选择一半的样本进行探索性因子分析 (exploratory factor analysis, EFA),以精简条目并确定量表的基本结构。对另一半样本进行验证性因子分析 (confirmatory factor analysis, CFA),以评估所提取因子结构的稳健性。

结果: 35 个症状中常见的急性期症状是"身体的漂浮或盘旋感"(94%)、"在动感音乐的环境下心情更加舒畅、欣快感"(86%)、和"兴奋话多,精力充沛"(67%)。常见的慢性症状是"记忆力下降"(93%)、"性格变化"(86%)和"反应变慢"(81%)。探索性因子分析最终得到的最佳模型量表,共22 个条

目,构成四个因子:两个因子代表慢性症状(社交退缩和睡眠障碍),一个因子是关于急性精神病样症状的,而第四个因子整合了急性的药物所致的欣快感和长期的性欲下降。验证性因素分析显示这 4 个因子能解释最终 22 个条目量表总方差的 50%,并且模型拟合良好(拟合指数,GFI=83.3%;近似误差均方根,RMSEA=0.072)。

结论: 四因子模型(社交退缩、睡眠障碍、精神病样症状和使用毒品时的欣快感)能较为客观地反映氯胺酮滥用相关的短期和长期心理症状。对这一 22 个条目的量表,今后的研究工作需要扩大样本量,以验证实该四因子结构的效度,评估量表的重测信度,并确定是否能将该量表用于氯胺酮依赖的鉴别诊断和治疗监测。

**关键词:** 氯胺酮; 物质相关障碍; 体征和症状; 因子分析; 中国

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Dr. Fan obtained her bachelor's degree from Chongqing Medical College in 1997, her master's degree in Neuropsychology from Jinan University (Guangdong) in 2003, her Ph.D. from Louisiana State University (USA) in 2009, and post-doctoral training at Yale University from 2009 until 2012. She then joined Guangzhou Brain Hospital where she is currently an associate professor and the deputy director of the Institute of Psychiatry and Neuroscience. Her main research interest is the mechanisms of ketamine dependence.