### Ketamine Side Effect Tool (KSET)

The Ketamine Side Effect Tool (KSET) comprises a set of forms designed to monitor side effects across a course of treatment with ketamine or ketamine derivatives. It is based on evidence from a systematic review (Short et al. 2018)<sup>1</sup> of side effects associated with ketamine use in depression, combined with expert opinion. Although based on studies related to ketamine as a treatment for depression, the KSET has been developed to also be useful for other treatment indications, e.g., chronic pain. Most of the forms can be rated by either the clinician and/or patient and cover three distinct clinical stages:

**KSET – Screening** assesses medical issues which should be considered before a patient is prescribed ketamine treatment.

KSET – Baseline / KSET – Follow Up assesses for medium- and longer-term side effects and potential withdrawal symptoms related to ketamine treatment.

**KSET – Acute Treatment** assesses acute side effects associated with ketamine administration.

The forms do not replace the need for thorough psychiatric assessment and physical examination of patients considered for ketamine treatment, or for further investigations (including additional blood tests or urine toxicology) if adverse effects emerge during treatment. The KSET is an assessment and monitoring tool only and is not intended to provide clinicians or researchers with a comprehensive education on how ketamine or ketamine derivatives should be used. It can complement other policies and procedures already established at a clinical site and/or research protocol.

#### **KSET – Screening**

This form is designed to supplement the psychiatric and physical examinations conducted when assessing the safety of ketamine<sup>^</sup> treatment for a patient.

- **Page 1** of the form can be completed by either the patient or clinician. It enquires about medical conditions that are known to increase the risk of adverse effects with ketamine treatment. It also elicits information about any prior experience with ketamine to assist in determining the patient's risk of developing adverse effects, including dependency. There is a space to list allergies and medications used, as a prompt for clinicians to consider potential drug interactions with ketamine.
- **Page 2** of the form is to be completed by the clinician and includes baseline heart rate (HR) and blood pressure (BP) measurements, as well as a panel for recording whether important investigations have been completed. There is also a space for the clinician to document other relevant notes (e.g. length of current depression episode, number of previously failed antidepressant trials, past augmentation strategies).

<sup>&</sup>lt;sup>1</sup> Short B, Fong J, Galvez V, Shelker W, Loo C.K, 2018. Side-effects associated with ketamine use in depression: a systematic review, The Lancet Psychiatry 5(1) 65-78. <sup>^</sup> In these instructions, the word 'ketamine' refers to ketamine and ketamine derivatives.

## KSET— Screening (Page 1)

Complete when assessing suitability for ketamine treatment.

If pre-filled by patient, clinician should review

**Patient Name** 

Date of Birth

Have you experienced or currently experience any of the following?	Have you ever received or used ketar	nine for any reason? Tick	all that apply:					
Hallucinations (e.g., seeing, hearing, smelling or tasting things that are not present in reality)	Yes	No	<ul> <li>No previous use (please continue on to Allergies section)</li> <li>Depression treatment → If yes, was it helpful? □ Yes □ No</li> <li>Pain management → If yes, was it helpful? □ Yes □ No</li> <li>Other medical reasons (e.g., anaesthesia, sedation) → Please specify:</li></ul>					
Elevated/irritable mood (e.g., euphoria, agitation, recklessness, increased energy, increased confidence)	Yes	No						_
Dissociation – induced by a drug/substance (e.g., feeling disconnected from self, body, thoughts, surroundings)	Yes	No						
Dissociation – not related to drug/substance use	Yes	No	Please fill in the table to the best of yo	our ability regarding past ke	etamine use (as relevant):	1		1
High blood pressure (hypertension)	Yes	No		Depression treatment	Pain management	Recreationa	al use	
Heart/cardiovascular condition	Yes	No	When did you last receive	Last 3 months	Last 3 months	🗆 Last 3 mo	onths	ſ
Seizures/stroke/head injury/neurological disorder	Yes	No	ketamine?	□ 3—12 months ago	□ 3—12 months ago	□ 3—12 m	onths a	go
Glaucoma	Yes	No		> 12 months ago	> 12 months ago	□ > 12 mor	ths ago	)
Liver problems	Yes	No	How many times in total (lifetime)?					
Kidney problems	Yes	No	Route (e.g., injection, infusion					ſ
Bladder problems or problems passing urine (e.g., pain, burning, irrita- tion, increased frequency, difficulties passing and/or changes in colour/ smell of urine)	Yes	No	'drip', nasal, oral) Highest dose received Did you have any adverse		□ No			
Alcohol or drug misuse	Yes	No	reactions?	□ Yes—specify: □ Yes—specify:		□ Yes—specify:		
Chronic pain	Yes	No						
Problems with memory and/or concentration	Yes	No						
For females: Current pregnancy	Yes	No	Have you ever craved ketamine?				Yes	No
For females: Currently breastfeeding	Yes	No	If "Yes", please specify how recently:					
Current use of medications or supplements? If "Yes" please list all medications and supplements below	Ŷ	'es	No <b>Allergies</b>	N	inician ame			
					gnature			

# KSET— Screening (Page 2)

To be completed by clinician

Patient Name	
Date of Birth	
Medical Record No.	

Vital Signs		Wedical Record No.
Heart rate (per minute)		Clinician Notes
Blood pressure (mmHg)		
Investigations		
Liver Function Tests (LFTs) (Please attach results)	Test Date:	
Normal result		
Abnormal result(s)		
Urinalysis (Dipstick)	Test Date:	
(Please attach results if completed)		
Not applicable		
Normal result		
Abnormal result(s)		
500	Test Date:	
ECG	Test Date:	
(Please attach results if completed)		
Not applicable		
Normal result		
Abnormal result(s)		

Clinician Name	
Signature	
Date	

#### KSET – Baseline / KSET – Follow Up

The KSET – Baseline and KSET – Follow-Up are two complementary forms that can be used together to elicit details about any medium- and longer-term side effects which emerge during ketamine treatment (i.e., compared to pre-treatment baseline). Potential side effects are assessed in a systematic method via a table of symptoms, which can be completed by either the clinician or patient. Any symptoms present should be circled and rated according to the severity key. The severity of these symptoms can then be summed to give a total score to indicate the total side effect burden. This total score is to be viewed as a guide only.

**KSET – Baseline** should be completed before commencing ketamine treatment to allow for comparison of symptoms after the patient has received ketamine treatment. It needs to be completed once only at the beginning of the patient's ketamine treatment course. The recommended timeframe for enquiry is 'over the past month', but this may be modified if clinically appropriate.

**KSET – Follow Up** form(s) <u>should be completed at intervals determined by the treating clinician</u>, taking into consideration the patient's dose, route of administration and frequency of treatment. The recommended timeframe for enquiry is 'over the past month', but this may be modified if clinically appropriate. A section of the KSET – Follow Up form allows for the recording of <u>liver function tests</u> (LFTs) and <u>urinalysis</u>. These should be completed as clinically indicated (e.g., if the patient begins to subjectively report urological problems) or, in the case of LFTs, at clinician-specified follow-up time points.

Both the KSET – Baseline and KSET – Follow-Up forms also have sections to record <u>cognitive test results</u> to facilitate cognitive monitoring. If subjectively reported cognitive side effects emerge during treatment, or there is concern for cognitive side effects (e.g., due to cognitive impairment at pre-treatment, or if high doses, frequency or number of treatments are prescribed), it is recommended the patient be assessed using measures such as those detailed below and in the associated paper, Short et al. (in press)<sup>2</sup>. The cognitive domains in the table below are those which can be affected by ketamine. Several measures are listed for assessing each domain – any one of these tests can be used to assess that domain. The measures listed are standardized and validated tests suitable for repeated assessment (i.e., with alternative forms).

The KSET – Follow Up form(s) can then be <u>compared with KSET – Screening and KSET – Baseline</u> forms to assess whether there is a temporal relationship between ketamine treatment and the onset of new symptoms or changes. The follow-up form can also be used to assess for potential withdrawal symptoms once ketamine administration has been ceased.

Cognitive Domain	Examples of suitable measures for assessment of cognitive side effects with ketamine
Verbal episodic memory	Rey Auditory Verbal Learning Test <sup>3</sup> , Hopkins Verbal Learning Test-Revised <sup>4</sup> , California Verbal Learning Test 2 <sup>nd</sup> Ed <sup>5</sup> ,
	CogState International Shopping List Task <sup>6</sup>
Working memory/Attention	Symbol Digit Substitution Test <sup>7</sup> , CogState One Back Test <sup>6</sup> , Conners Continuous Performance Test 3 <sup>rd</sup> Ed <sup>8</sup>
Verbal fluency	Controlled Oral Word Association Test <sup>9</sup> , Delis-Kaplan Executive Function System Verbal Fluency subtest <sup>10</sup>

<sup>&</sup>lt;sup>2</sup>Short et al. KSET (in Press). <sup>3</sup>Rey, A., 1964. L'Examen Clinique en Psychologie. Presses Universitaires de France, Paris. <sup>4</sup>Benedict RHB, et al., 1998. Hopkins Verbal Learning Test revised: normative data and analysis of interform and test-retest reliability. Clin Neuropsychol 12: 43–55. <sup>5</sup>Delis, D.C., et al. 2000. California Verbal Learning Test, 2nd edn. The Psychological Corporation, San Antonio, TX.<sup>6</sup>Westerman, R., et al. 2001. Computer-assisted cognitive function assessment in pilots: How and why? ADF Health, 2, 29-36.<sup>7</sup>Smith, A., 1991. Symbol Digit Modalities Test. Western Psychological Services, Los Angeles, CA. <sup>8</sup>Conners, C.K., et al. 2000. Conners' Continuous Performance Test II: Computer Program for Windows Technical Guide and Software Manual. North Tonawanda, NY: Multi-Health Systems. <sup>9</sup>Benton, A.L., et al., 1989. Multilingual Aphasia Examination. AJA Associates, Iowa City, IA. <sup>10</sup>Delis, D.C., et al. 2001. The Delis-Kaplan Executive Function System: Examiner's Manual. The Psychological Corporation, San Antonio.

KSET— Baseline	If pr	e-fille	d by p	atient	Patient Name Date of Birth
<u>Questions for patient</u> "Have you experienced the following symptoms over the past month? If yes, how severe were they?"	Severity (circle) 0 = Never				Medical Record No. Weight
<u>Symptoms</u> Circle relevant items. Further details can be documented by the clinician in <b>Clinician Notes</b> (e.g., pre-existing conditions, treatments etc.)		1 = Mi 2 = Mi 3 = Se	oderat	e	Severity Key
Dissociation (e.g., felt disconnected from self, body, thoughts, surroundings; feeling strange and/or "spaced out")	0	1	2	3	<ul> <li>0 Never</li> <li>1 Mild - transient and easily tolerated</li> <li>2 Moderate - caused discomfort and/or interference with usual activities</li> <li>2 Several event discriminant discomfort and (on considerable interference with usual activities)</li> </ul>
Hallucinations (e.g., seeing, hearing, smelling or tasting things that were not present in reality)	0	1	2	3	3 Severe - caused significant discomfort and/or considerable interference with usual activities
Problems with memory and/or concentration	0	1	2	3	Clinician Notes
Anxiety	0	1	2	3	
Restlessness and/or agitation	0	1	2	3	
Elevated/irritable mood (e.g., euphoria, recklessness, increased energy, increased confidence)	0	1	2	3	
Insomnia, nightmares and/or unusual dreams	0	1	2	3	
Drowsiness, fatigue, and/or weakness	0	1	2	3	
Headache	0	1	2	3	
Abnormal movements (e.g., tremor, incoordination and/or spasms)	0	1	2	3	
Vision or hearing changes (e.g., blurred vision, double vision and/or tinnitus)	0	1	2	3	
Cardiovascular (e.g., shortness of breath, chest pain and/or palpitations)	0	1	2	3	
Diarrhoea and/or constipation	0	1	2	3	
Abdominal pain and/or cramps	0	1	2	3	
Nausea and/or vomiting	0	1	2	3	11
Skin changes (e.g., rash, itch, yellow discolouration)	0	1	2	3	
Problems passing urine (e.g., pain, burning, irritation, increased frequency, difficulties passing and/or changes in colour/smell of urine)	0	1	2	3	Cognitive Testing Clinician
A craving for ketamine (e.g., often thinking about wanting more ketamine)	0	1	2	3	Name of Test Test Date Score Name
Seeking and/or using non-prescribed ketamine	0	1	2	3	Signature
Other—specify:	0	1	2	3	Signature
	Tota	=			Date

#### **KSET – Acute Treatment**

This form is to be completed during each ketamine administration session. It consists of three pages, with sections colour-coded to indicate at what stage of the treatment session they should be completed:

- Yellow items should be completed prior to ketamine administration (i.e., pre-dose).
- Blue items should be completed after ketamine administration (i.e., post-dose).
- Green items (discharge checklist) should be completed at the time of patient discharge, e.g. at 120 minutes after treatment.

<u>Page 1</u> can be filled out by either the patient or clinician and is used to assess whether side effects have emerged <u>since the patient's last treatment session</u>. The questions are bundled in the table entitled **Pre-treatment Check**.

Page 2 allows the clinician or patient to:

- Assess and record <u>symptoms present</u> <u>Pre-Dose</u>\*
- Assess and record <u>symptoms present</u> <u>Post-Dose</u> after ketamine administration\*\*
- If a symptom emerges, the patient should be asked if it has Resolved at time of discharge (i.e., 120 mins), circling either yes or no.

Each symptom is rated according to the severity key. If a symptom has emerged, it should be circled. A 'Clinician Comments' box is available on pages 1 and 2 for the clinician to note any other relevant information elicited (e.g., medication changes since the last ketamine treatment, notes regarding emergent side effects).

\* Some of these symptom questions are repeated from the Pre-Treatment Checklist. This is with the aim of assessing whether they are present at the moment of pre-dose. The Pre-Treatment Checklist asks whether they have been present at any time since the last ketamine treatment (i.e., which may have been a week ago, not necessarily on the day the dose is due to be administered).

\*\* Although the form stipulates that the post-dose column can be completed at 60 minutes, the treating clinician should determine what time interval is most appropriate taking into account the route of ketamine administration, dosage and peak severity of symptoms. Please refer to your clinical site procedures and/or research protocols for guidance. Page 3 allows the clinician to complete:

- **Cardiovascular Monitoring**. Pre-treatment measurements should be taken before ketamine is administered (highlighted in yellow). The clinician should ensure that the pre-treatment BP reading is within safe limits before dosing; e.g., a reasonable upper threshold for BP may be 160/90mmHg. Subsequent observations could be done at 15-minutes and 30-minutes post-dose, then at 30-minute intervals thereafter. Note that what is considered a safe pre-dose BP and the exact timing of cardiovascular monitoring will depend on the patient, dose and route of administration. We recommend you check your clinical site procedures and/or research protocols.
- **Dosing Information** and an **Administration Site Check**. Prescription information can be added here as well as comments about administration site effects (e.g., IV/IM/SC site swelling, nasal congestion and/or bleeding associated with inhaled ketamine).
- Orientation & Discharge Assessments allow the clinician to assess whether the patient is safe for discharge after ketamine administration. This requires that a pre-treatment orientation score is obtained prior to ketamine administration (highlighted in yellow), for purpose of comparison. The <u>First Assessment</u> column can be completed at a time when the clinician considers a patient may be suitable for discharge. It involves an assessment of orientation along with other criteria which are listed in the table. If the answers on the checklist are all "Yes" the patient is likely to be suitable for discharge. If at least one answer is "No", a doctor should be consulted; if deemed necessary, the patient may be observed for a further 30 minutes, after which a <u>Second Assessment</u> can be completed. If the patient does not meet all the criteria at second assessment, a further period of observation may be required and recorded elsewhere.
- An **Overall Tolerability** assessment of ketamine relating to the treatment session can also be recorded. In completing this, the clinician should consider the side effect burden as well as cardiovascular and discharge checklist findings. The tolerability rating can then be used to inform future ketamine dosing.

Ketamine Side Effect Tool (KSET) Version 1.0

KSET— Acute Treatment (Page 1) If pre-filled by patient, clini	cian should review Patient Name
	Date of Birth
KSET—Acute Treatment Flowchart	Medical Record No.
Complete	Treatment Number
pre-treatment Administer post dose Comple	te discharge Clinician Notes
sections on Pages	ist on Pages and 3
1, 2 and 3 2 and 3	
Dro Trootmont Chack (consider before proceeding with treatment)	
Pre-Treatment Check (consider before proceeding with treatment)	
<u>Questions for patient</u> <i>"Have you experienced any of the following symptoms since your last treatment session?</i>	Not required at
If yes, how severe were they?"	first treatment
Severity       0 Never         1 Mild - transient and easily tolerated         2 Moderate - caused discomfort and/or interrupted usual activities         3 Severe - caused significant discomfort and/or considerable interference with usual activities	Severity (circle) 0 = Never 1 = Mild 2 = Moderate
Symptoms	3 = Severe
Problems with memory and/or concentration	0 1 2 3
Diarrhoea, constipation and/or abdominal pain/cramps Skin changes (e.g., rash, itch, yellow discolouration)	0 1 2 3 0 1 2 3
Problems passing urine (e.g., pain, burning, irritation, increased frequency, difficulties passing and/or changes in colour/smell of urine)	0 1 2 3
A craving for ketamine (e.g., often thinking about wanting more ketamine)	0 1 2 3
Seeking and/or using non-prescribed ketamine	0 1 2 3
Insomnia, nightmares and/or unusual dreams	0 1 2 3
Hallucinations (e.g., seeing, hearing, smelling or tasting things that were not present in reality)	0 1 2 3 Clinician Name
Elevated/irritable mood (e.g., euphoria, recklessness, increased energy, increased confidence)	0 1 2 3 Signature
Other—specify:	0 1 2 3 Date

# KSET— Acute Treatment (Page 2)

**Questions for patient** 

If pre-filled by patient, clinician should review

Severity Key

Patient Name	
Date of Birth	
Medical Record No.	
Treatment No.	

<i>"Have you experienced any of the following symp- toms today? If yes, how severe were they? "</i>	1 = N 2 = N	Noder	ate - d	ent and caused sed sigr	discor	nfort	and/o	or interf t and/c	Medical Record N Treatment No.	lo.		
Symptoms Circle relevant items and severity. Further details can be documented by the clinician in Clinician Notes (e.g., any other observations during treatment)	Pre Dose Time completed : 				(Ask patient to rate peak							Clinician Notes
Dissociation (e.g., felt disconnected from self, body, thoughts, surroundings; feeling strange and/or "spaced out")	0	1	2	3	0	1	2	3	Yes	No		
Hallucinations (e.g., seeing, hearing, smelling or tasting things that were not present in reality)	0	1	2	3	0	1	2	3	Yes	No		
Problems with memory and/or concentration	0	1	2	3	0	1	2	3	Yes	No		
Anxiety	0	1	2	3	0	1	2	3	Yes	No		
Restlessness and/or agitation	0	1	2	3	0	1	2	3	Yes	No		
Elevated/irritable mood (e.g., euphoria, recklessness, increased energy, increased confidence)	0	1	2	3	0	1	2	3	Yes	No		
Tearfulness	0	1	2	3	0	1	2	3	Yes	No		
Drowsiness, fatigue, and/or weakness	0	1	2	3	0	1	2	3	Yes	No		
Dizziness, lightheadedness, feeling faint and/or vertigo (e.g., felt like swaying or spinning)	0	1	2	3	0	1	2	3	Yes	No		
Headache	0	1	2	3	0	1	2	3	Yes	No		
Numbness and/or tingling of body parts	0	1	2	3	0	1	2	3	Yes	No		
Abnormal movements (e.g., tremor, incoordination and/or spasms)	0	1	2	3	0	1	2	3	Yes	No		
Vision changes (e.g., blurred vision)	0	1	2	3	0	1	2	3	Yes	No		
Hearing changes (e.g., hearing impairment or tinnitus)	0	1	2	3	0	1	2	3	Yes	No		
Dry mouth, increased salivation or metallic/unusual taste	0	1	2	3	0	1	2	3	Yes	No		
Cardiovascular (e.g., shortness of breath, chest pain and/or palpitations)	0	1	2	3	0	1	2	3	Yes	No		
Nausea and/or vomiting	0	1	2	3	0	1	2	3	Yes	No		
Skin changes (e.g., rash, itch, yellow discolouration)	0	1	2	3	0	1	2	3	Yes	No	Clinician Name	
Felt unusually hot, sweaty or cold	0	1	2	3	0	1	2	3	Yes	No		
Other – specify:	0	1	2	3	0	1	2	3	Yes	No	Signature	
Total =											Date	

### KSET— Acute Treatment (Page 3)

To be completed by clinician

Patient Name

								Date of Birth		
Cardiovascular Monitoring (Re	fer to your site's clir	nical/research	protocol regarding	time intervals)						
Time Intervals	Pre Treatment	15 min	30 min	60 min	90	min	120 min	Medical Record No.		
Time								Treatment No.		
Blood Pressure	/	/	/	/		/	/			
Heart Rate									t BP is within safe limits	
Other—Specify								before proceedin	g (details in manual)	
Orientation & Discharge As	cassmants							Dosing Information		
Onentation & Discharge As	sessments	-	Due Treestreest	Deat To		Dent	<b>-</b>	Time of Administration		
			Pre Treatment	Post-Tre Assessi			Treatment essment 2	Dosage		
				(120 mins			needed)	Route		
Time						,	,	Administration Site Check		
Orientation (tick if correct and	l record answer if inc	correct)								
Name								Comments		
Date of Birth										
Age										
Year								Overall	Tolerability	
Month								Select one answ	ver prior to discharge.	
Day										
Place								Consider both subjective patient reports and objective observations		
Total Score										
Discharge Checklist								Good, could tolerate <u>hig</u>	<u>her</u> dose.	
Orientation Score ≥ Pre Treatm	ent Orientation Sco	re		Yes	No	Yes	No	Moderate, could tolerate	e <u>same</u> dose.	
Blood pressure ≤ 120% Pre Trea	atment			Yes	No	Yes	No	Poor, could <u>not tolerate</u>	same dose.	
Heart Rate ≤ 120% Pre Treatme	ent			Yes	No	Yes	No			
Can walk unassisted				Yes	No	Yes	No			
Feeling physically well				Yes	No	Yes	No	Time of Discharge		
Alert, calm, and comfortable (i.	.e., not distressed)			Yes	No	Yes	No	Clinician Name		
Patient meets all criteria for dis	scharge			Yes	No	Yes	No			
If discharge criteria not met	at Post-Treatmen	t Assessmer	nt 1, complete Pos	st-Treatment A	ssessment	2 at a late	er time	Signature Date		

#### KSET – Baseline / KSET – Follow Up

The KSET – Baseline and KSET – Follow-Up are two complementary forms that can be used together to elicit details about any medium- and longer-term side effects which emerge during ketamine treatment (i.e., compared to pre-treatment baseline). Potential side effects are assessed in a systematic method via a table of symptoms, which can be completed by either the clinician or patient. Any symptoms present should be circled and rated according to the severity key. The severity of these symptoms can then be summed to give a total score to indicate the total side effect burden. This total score is to be viewed as a guide only.

**KSET – Baseline** should be completed before commencing ketamine treatment to allow for comparison of symptoms after the patient has received ketamine treatment. It needs to be completed once only at the beginning of the patient's ketamine treatment course. The recommended timeframe for enquiry is 'over the past month', but this may be modified if clinically appropriate.

**KSET – Follow Up** form(s) <u>should be completed at intervals determined by the treating clinician</u>, taking into consideration the patient's dose, route of administration and frequency of treatment. The recommended timeframe for enquiry is 'over the past month', but this may be modified if clinically appropriate. A section of the KSET – Follow Up form allows for the recording of <u>liver function tests</u> (LFTs) and <u>urinalysis</u>. These should be completed as clinically indicated (e.g., if the patient begins to subjectively report urological problems) or, in the case of LFTs, at clinician-specified follow-up time points.

Both the KSET – Baseline and KSET – Follow-Up forms also have sections to record <u>cognitive test results</u> to facilitate cognitive monitoring. If subjectively reported cognitive side effects emerge during treatment, or there is concern for cognitive side effects (e.g., due to cognitive impairment at pre-treatment, or if high doses, frequency or number of treatments are prescribed), it is recommended the patient be assessed using measures such as those detailed below and in the associated paper, Short et al. (in press)<sup>2</sup>. The cognitive domains in the table below are those which can be affected by ketamine. Several measures are listed for assessing each domain – any one of these tests can be used to assess that domain. The measures listed are standardized and validated tests suitable for repeated assessment (i.e., with alternative forms).

The KSET – Follow Up form(s) can then be <u>compared with KSET – Screening and KSET – Baseline</u> forms to assess whether there is a temporal relationship between ketamine treatment and the onset of new symptoms or changes. The follow-up form can also be used to assess for potential withdrawal symptoms once ketamine administration has been ceased.

Cognitive Domain	Examples of suitable measures for assessment of cognitive side effects with ketamine
Verbal episodic memory	Rey Auditory Verbal Learning Test <sup>3</sup> , Hopkins Verbal Learning Test-Revised <sup>4</sup> , California Verbal Learning Test 2 <sup>nd</sup> Ed <sup>5</sup> ,
	CogState International Shopping List Task <sup>6</sup>
Working memory/Attention	Symbol Digit Substitution Test <sup>7</sup> , CogState One Back Test <sup>6</sup> , Conners Continuous Performance Test 3 <sup>rd</sup> Ed <sup>8</sup>
Verbal fluency	Controlled Oral Word Association Test <sup>9</sup> , Delis-Kaplan Executive Function System Verbal Fluency subtest <sup>10</sup>

<sup>&</sup>lt;sup>2</sup>Short et al. KSET (in Press). <sup>3</sup>Rey, A., 1964. L'Examen Clinique en Psychologie. Presses Universitaires de France, Paris. <sup>4</sup>Benedict RHB, et al., 1998. Hopkins Verbal Learning Test revised: normative data and analysis of interform and test-retest reliability. Clin Neuropsychol 12: 43–55. <sup>5</sup>Delis, D.C., et al. 2000. California Verbal Learning Test, 2nd edn. The Psychological Corporation, San Antonio, TX.<sup>6</sup>Westerman, R., et al. 2001. Computer-assisted cognitive function assessment in pilots: How and why? ADF Health, 2, 29-36.<sup>7</sup>Smith, A., 1991. Symbol Digit Modalities Test. Western Psychological Services, Los Angeles, CA. <sup>8</sup>Conners, C.K., et al. 2000. Conners' Continuous Performance Test II: Computer Program for Windows Technical Guide and Software Manual. North Tonawanda, NY: Multi-Health Systems. <sup>9</sup>Benton, A.L., et al., 1989. Multilingual Aphasia Examination. AJA Associates, Iowa City, IA. <sup>10</sup>Delis, D.C., et al. 2001. The Delis-Kaplan Executive Function System: Examiner's Manual. The Psychological Corporation, San Antonio.

KSET— Follow-Up		lf pr	e-fille	ed by p	atient, clinician should review	Patient Name		
Questions for patient         "Excluding treatment sessions, have you experienced the following symptoms over the past month (or enter time range)?         If yes, how severe were they?"         Symptoms         Circle relevant items and provide further details in Clinician Notes (e.g., pre-existing conditions, treatments, etc.)	Severity (circle) 0 = Never 1 = Mild 2 = Moderate 3 = Severe				Severity Key O Never Mild- transient and easily tolerated Moderate- caused discomfort and/or interference with usual activities	Date of Birth Medical Record No. Clinician Notes		
Dissociation (e.g., felt disconnected from self, body, thoughts, surround- ings; feeling strange and/or "spaced out")	0	1	2	3	<b>3</b> Severe- caused significant discomfort and/or considerable interference with usual activities			
Hallucinations (e.g., seeing, hearing, smelling or tasting things that were not present in reality)	0	1	2	3	Date of last ketamine dose:			
Problems with memory and/or concentration	0	1	2	3	Investigations			
Anxiety	0	1	2	3	Liver Function Tests (LFTs) Test Date:			
Restlessness and/or agitation	0	1	2	-	(Attach results if completed)			
Elevated/irritable mood (e.g., euphoria, recklessness, increased energy, increased confidence)	0	1	2	3	<ul> <li>Not applicable</li> <li>Normal result</li> </ul>			
Insomnia, nightmares and/or unusual dreams	0	0       1       2       3         0       1       2       3         0       1       2       3         0       1       2       3         O       1       2       3         O       1       2       3         O       1       2       3						
Drowsiness, fatigue, and/or weakness	0							
Headache	0							
Abnormal movements (e.g., tremor, incoordination and/or spasms)	0	1	2	3	□ No			
Vision or hearing changes (e.g., blurred vision, double vision and/or tinnitus)	0	1	2	3	Urinalysis (Dipstick)Test Date:(Attach results if completed)	:		
Cardiovascular (e.g., shortness of breath, chest pain and/or palpitations)	0	1	2	3	Not applicable			
Diarrhoea and/or constipation	0	1	2	3	Normal result			
Abdominal pain and/or cramps	0	1	2	3	□ Abnormal result(s)			
Nausea and/or vomiting	0	1	2	3	Further tests required?			
Skin changes (e.g., rash, itch, yellow discolouration)	0	1	2	3	<ul> <li>Yes - specify due date:</li> <li>No</li> </ul>			
Problems passing urine (e.g., pain, burning, irritation, increased frequency, difficulties passing and/or changes in colour/smell of urine)	0	1	2	3	Cognitive Assessment			
A craving for ketamine (e.g., often thinking about wanting more ketamine)	0	1	2	3	Name of Test Test Date Score	Clinician		
Seeking and/or using non-prescribed ketamine	0	1	2	3		Name		
Other—specify:	0	1	2	3		Signature		
	Tot	al =				Date		