

# Does Childhood Trauma Impact Outcomes in Adults Receiving Ketamine for Depression?

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## BACKGROUND

Childhood trauma is associated with inferior treatment response to conventional antidepressants. This relationship has not been extensively explored with infusion ketamine therapy.

## STUDY AIM

To compare antidepressant effects of repeated IV ketamine in adults with TRD with histories of self-reported childhood trauma who sought treatment at an outpatient ketamine clinic.

## METHODS

### Sample:

Seventy-three adults received at least 4 infusions of intravenous ketamine (0.5-1.0 mg/kg over 40 minutes) on a twice-weekly or once-weekly basis at a private outpatient clinic.

### Measures:

The **QIDS-SR** was used to assess depression severity at immediately prior to each infusion for a total of 4 observations.

The **Childhood Trauma Questionnaire (CTQ)** was administered at pre-treatment to assess severity of childhood trauma, yielding a total trauma score (TT) and subscale scores for childhood history of physical neglect (PN) physical abuse (PA) emotional neglect (EN) emotional abuse (EA) and sexual abuse (SA).

Cut-off scores between mild, moderate and severe physical, sexual, emotional abuse, physical neglect, and emotional neglect (8, 6, 9, 8, 10, respectively) have been previously established. These cut-off scores were used to devise dichotomous categories (i.e. none-to-minimal, mild-to-severe) for analyses.

### Data Analytic Plan

Repeated measures **General Linear Models** tested effects of childhood trauma on QIDS-SR scores.

**Fisher Exact Tests** compared response ( $\geq 50\%$  reduction from baseline QIDS-SR score) and remission (QIDS-SR  $\leq 5$ ) rates between patients reporting scores above and below established cut points for PN, PA, EN, EA and SA.

## KEY REFERENCES

- 1) Nanni V, Uher R, Danese A. Childhood maltreatment predicts unfavorable course of illness and treatment outcome in depression: a meta-analysis. *Am J Psychiatry* 2012; 169: 141–151.
- 2) Williams LM, DeBattista C, Duchemin A-M, Schatzberg AF, Nemeroff CB: Childhood trauma predicts antidepressant response in adults with major depression: data from the randomized international study to predict optimized treatment for depression. *Transl Psychiatry* 2016 May 3; 6(5):e799
- 3) Bernstein BP, Fink L: Childhood Trauma Questionnaire: A Retrospective Self-Report. San Antonio, Tex. The Psychological Corp; 1998

## RESULTS

Fig 1

Interaction of Emotional Neglect x Treatment Number on Depression Severity

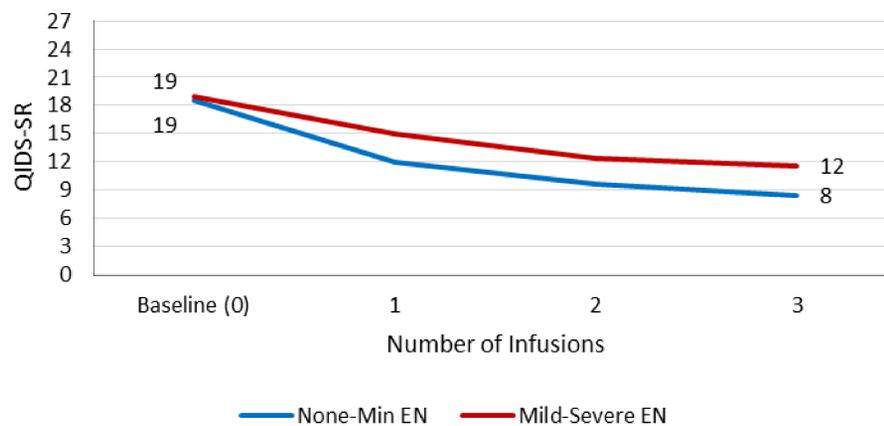


Fig 2

Rates of Response for Patients with None-Minimal vs Mild-Severe Neglect and Abuse

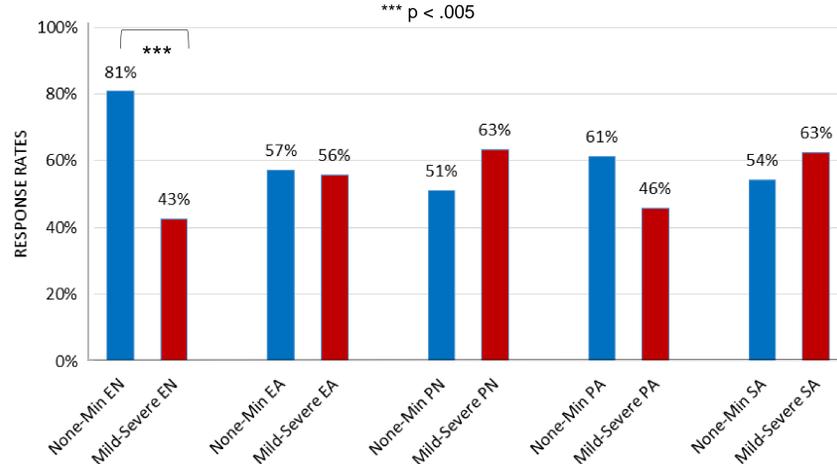
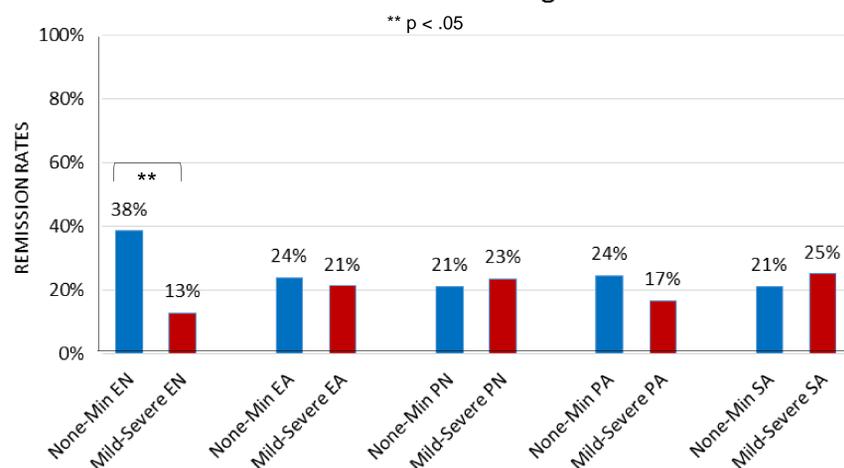


Fig 3

Rates of Remission for Patients with None-Minimal vs Mild-Severe Neglect and Abuse



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Table 1. Patient Demographics & Clinical Data

	N	%
Female	42	58
Mean Age (SD)	41.9	(14.9)
Comorbid Pain Disorder	8	11
Comorbid Anxiety Disorder	41	56
Comorbid PTSD	7	10
Previous ECT Treatment	11	15
Average Number of Acute Tx Infusions (SD)	5.6	(0.7)

Descriptive data on the study sample are provided in Table 1. Ketamine significantly reduced QIDS-SR scores (Mean QIDS score change=8.4, SD=5.9,  $F(2.41, 168.8)=110.41$   $p<.0001$ ); 56% of patients were ketamine responders, and 22% were ketamine remitters.

Total trauma scores did not associate with outcomes. There was a main effect of emotional abuse (EA;  $F(1,69)=4.14$ ,  $p=.046$ ) and emotional neglect (EN;  $F(1,69)=4.61$ ,  $p=.035$ ) due to higher QIDS-SR scores throughout treatment for the mild-severe EA and mild-severe EN groups.

Analyses also showed an emotional neglect (EN) by infusion interaction ( $F(2.45,168.99)=3.33$ ,  $p=.029$ , indicating a weaker antidepressant effect in the Mild-Severe Emotional Neglect group than the None-Minimum EN group (Fig 1).

Additionally, Mild-Severe EN was associated with lower response (Mild-Severe EN=43% vs None-Min EN=81%,  $p=0.003$ ) and remission rates (Mild-Severe EN=13% vs None-Min EN=38%,  $p=0.017$ ) (Fig 2 & Fig 3).

Gender and age did not impact outcomes.

## DISCUSSION

- ❖ Emotional abuse and neglect are associated with greater depression severity throughout treatment
- ❖ Ketamine improved depression irrespective of childhood trauma severity, although patients with a self-reported history of emotional neglect benefitted less than patients with minimal or no history of emotional neglect.
- ❖ Response and remission rates with high childhood trauma are comparable to conventional antidepressant response rates in patients without childhood trauma, suggesting increased efficacy of ketamine for patients with childhood trauma.

- ❖ Future research should address the effects of concurrent treatment and previous treatment history.