

The role and clinical correlates of complex PTSD in people with psychosis

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Conflict of interest statement

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest

Author contribution statement

PP, FV, KB and WS contributed to the conception and design of the study. FV, WS, CC and RPB were involved in collecting the data. PP analysed the data, and wrote the first draft of the manuscript. FV, KB, WS and RPB contributed to the final draft of the manuscript. All authors contributed to the article and approved its submission.

Keywords

Trauma, psychosis, PTSD, CPTSD, Mediation, disturbance of self-organisation

Abstract

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Traumatic experiences and post-traumatic stress are highly prevalent in people with psychosis, increasing symptom burden, decreasing quality of life and moderating treatment response. A range of post-traumatic sequelae have been found to mediate the relationship between trauma and psychotic experiences including the 'traditional' symptoms of post-traumatic stress disorder (PTSD). The International Classification of Diseases-11th Edition recognises a more complex post-traumatic presentation, complex PTSD (cPTSD), which captures both the characteristic symptoms of PTSD alongside more pervasive post-traumatic sequelae known as 'disturbances in self-organisation' (DSOs). The prevalence and impact of cPTSD and DSOs in psychosis remains to be explored. In the first study of this kind, 144 participants with psychosis recruited from North West United Kingdom mental health services completed measures assessing trauma, PTSD and cPTSD symptoms and symptoms of psychosis. Forty-per-cent of the sample met criteria for cPTSD, compared to 10% who met diagnostic criteria for PTSD. PTSD mediated the relationship between trauma and positive symptoms; DSOs did not contribute to explaining this relationship. Both PTSD and DSOs mediated the relationship between trauma and affective symptoms but did not explain a significant proportion of variance in negative symptoms. Cognitive and excitative symptoms of psychosis did not correlate with trauma, PTSD or DSO scores. This findings indicate the possible value of adjunct therapies to manage cPTSD symptoms in people with psychosis, pending replication in larger epidemiological samples and longitudinal studies.

Contribution to the field

Research has shown that people with psychosis often have a history of traumatic life experiences. As such, post-traumatic stress disorder (PTSD) is highly common in people with psychosis. Such complex trauma histories may lead to complex PTSD - a more severe and enduring presentation than PTSD that requires distinct trauma-focused interventions. This paper is the first (to the authors' knowledge) to investigate the rates of complex PTSD in a sample of people with psychosis, and explore its clinical correlates. This provides a basis for future research into the prevalence and impact of complex PTSD in people with psychosis.

Ethics statements

Studies involving animal subjects

Generated Statement: No animal studies are presented in this manuscript.

Studies involving human subjects

Generated Statement: The studies involving human participants were reviewed and approved by National Institute for Health Research, National Health Service. The patients/participants provided their written informed consent to participate in this study.

Inclusion of identifiable human data

Generated Statement: No potentially identifiable human images or data is presented in this study.

Data availability statement

Generated Statement: The data analyzed in this study is subject to the following licenses/restrictions: The dataset analysed in this study is not publicly available to protect the sensitivity and confidentiality of the participants.. Requests to access these datasets should be directed to Peter Panayi, peter.panayi@manchester.ac.uk.

nreview



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17 Abstract

- 18 Traumatic experiences and post-traumatic stress are highly prevalent in people with psychosis,
- 19 increasing symptom burden, decreasing quality of life and moderating treatment response. A range of
- 20 post-traumatic sequelae have been found to mediate the relationship between trauma and psychotic
- 21 experiences, including the 'traditional' symptoms of post-traumatic stress disorder (PTSD). The
- 22 International Classification of Diseases-11th Edition recognises a more complex post-traumatic
- 23 presentation, complex PTSD (cPTSD), which captures both the characteristic symptoms of PTSD
- 24 alongside more pervasive post-traumatic sequelae known as 'disturbances in self-organisation'
- 25 (DSOs). The prevalence and impact of cPTSD and DSOs in psychosis remains to be explored. In the
- 26 first study of this kind, 144 participants with psychosis recruited from North West United Kingdom
- 27 mental health services completed measures assessing trauma, PTSD and cPTSD symptoms and
- symptoms of psychosis. Forty-per-cent of the sample met criteria for cPTSD, compared to 10% who

29 met diagnostic criteria for PTSD. PTSD and DSOs mediated the relationship between trauma and 30 positive symptoms, controlling for dataset membership. Both PTSD and DSOs mediated the 31 relationship between trauma and affective symptoms but did not explain a significant proportion of 32 variance in negative symptoms. Cognitive and excitative symptoms of psychosis did not correlate 33 with trauma, PTSD or DSO scores. These findings indicate the possible value of adjunct therapies to 34 manage cPTSD symptoms in people with psychosis, pending replication in larger epidemiological

35 samples and longitudinal studies.

36

37 1 Introduction

38 Traumatic life events and adverse childhood experiences may lead to various psychosocial 39 difficulties. Perhaps the most notable consequence of such experiences is Post-Traumatic Stress 40 Disorder (PTSD), characterized by re-experiencing (e.g., intrusive trauma memories), hyperarousal 41 (e.g., irritability and hypervigilance) and avoidance of trauma reminders (1). Cohort studies have 42 recorded a prevalence of PTSD as high as 7.8% in England (2). PTSD has been shown to predict 43 adverse physical and mental health outcomes (3,4), as well as reduced quality of life and social 44 functioning (5,6).

45

46 Following the classification of PTSD in the third edition of the Diagnostic and Statistical Manual of 47 Mental Disorders (7), neither clinicians nor trauma survivors felt it captured the complex difficulties 48 endorsed by victims of multiple or prolonged traumas. Hence, Herman (8) introduced the concept of 49 complex PTSD (cPTSD). This includes the above core symptoms of PTSD, as well as broader and 50 more severe symptomatology (e.g., cognitive, affective, and relational disturbance). Following a 51 body of empirical research demonstrating a quantitative (9) and qualitative (10) distinction between PTSD and cPTSD, the most recent International Classification of Diseases (11th Edition; ICD-11, 1) 52 53 has recognised cPTSD as a separate, sibling diagnosis to PTSD (11). This includes the symptoms of 54 PTSD as above, alongside other symptoms collectively referred to as 'disturbances of self-55 organisation' (DSOs), including negative self-concept, emotional dysregulation and interpersonal 56 difficulties. Prior studies indicate that core PTSD symptoms may be more severe in people with

57 cPTSD (12,13), that cPTSD is associated with increased burden (14), and requires disparate

58 treatment approaches to PTSD (15).

59

60 Complex trauma histories are highly common among people with schizophrenia spectrum conditions 61 (16), with meta-analyses consistently implicating trauma as an aetiological factor in psychosis (17– 62 19). Trauma exposure may lead to symptoms of PTSD – namely, re-experiencing and memory 63 intrusions – that may be appraised as anomalous experiences (e.g., hearing insults or phrases of past 64 abusers appraised as an external voice in the present moment) (20). Systematic reviews have 65 indicated that PTSD symptoms mediate the relationship between trauma and psychosis (21–23). 66 Despite consistent evidence of the involvement of PTSD symptoms in the pathway between trauma 67 and psychosis, there are no studies to our knowledge investigating the potential role of cPTSD.

68

The involvement of these additional post-traumatic sequelae in the trauma-psychosis relationship is 69 70 plausible, considering previous studies which considered similar mediators. Systematic reviews have 71 shown that emotion dysregulation, negative thoughts about the self and interpersonal difficulties 72 mediate the relationship between trauma and psychosis (21–23). Meta-analytic evidence is 73 concordant with these findings, showing that emotion dysregulation, negative self-concept and 74 attachment difficulties predict specific psychotic symptoms following trauma exposure (24). Further 75 plausibility for the role of cPTSD in the pathway from trauma to psychosis stems from repeated 76 childhood trauma, an risk factor for cPTSD (25) that is highly common among those with psychosis 77 (16). Similarly, PTSD is highly comorbid with psychosis (26,27), moderating treatment outcomes 78 and reducing quality of life (28,29). This requires replication in a psychosis sample to inform 79 assessment and intervention.

80

81 Negative symptoms of psychosis remain largely unexplained by PTSD. Strauss et al. (2011) found 82 those meeting criteria for deficit schizophrenia (i.e., negative symptoms related to the illness itself 83 lasting longer than 12 months; 29) were at lower risk of PTSD than those displaying secondary 84 negative symptoms. Consistently, subsequent meta-analytic evidence (32) has found small, non-

85 significant effects of trauma-focused cognitive-behavioural therapies on negative symptoms.

86 Together, these findings suggest PTSD symptoms may not mediate the relationship between trauma

87 and negative psychotic symptoms. DSOs may, however, play a role in this relationship as opposed to

88 PTSD, owing to their apparent clinical similarity with certain negative symptoms. For instance,

B9 DSOs may present as emotional numbing and anhedonia (25,33) as well as social withdrawal (34),

90 consistent with negative symptom presentations in psychosis. The limited evidence-based

91 interventions for negative symptoms and their associated burden (35) makes this mediation

92 hypothesis worth exploring.

93

94 Psychotic symptomatology is not restricted to positive and negative domains. Affective difficulties 95 are also common in people with psychosis, with anxiety and major depressive disorders affecting up 96 to 1 in 3 people at their first episode (36). Systematic reviews suggest that these difficulties correlate 97 with psychotic symptom severity, distress and content (37), and decrease quality of life (38), making 98 affective problems key targets in psychological interventions for psychosis. PTSD is associated with 99 a greater risk of anxiety and depression (39), and cPTSD even more so (40). Hence, post-traumatic 91 sequelae may play a maintaining role in affective difficulties among people with psychosis.

101

102 Other symptom domains of psychosis have also been identified; namely, cognitive and excitative 103 difficulties. PTSD symptoms do not correlate with cognitive difficulties in people with psychosis 104 (41,42). However, those with an alleged neurodevelopmental predisposition to psychosis may be at 105 greater risk of childhood victimization, especially bullying from peers (43). Thus, this pathway may 106 interact with a trauma pathway to psychosis. The positive/negative symptom solutions typically used 107 in the scoring of the PANSS may not capture such cognitive difficulties; factor analytic studies have 108 identified a more complex underlying structure to the PANSS comprising positive, negative, 109 cognitive, affective and excitative symptoms (44–46). Therefore, it is possible that post-traumatic 110 sequelae more complex than PTSD - i.e., cPTSD - may lead to nuanced psychotic symptoms, such111 as cognitive/excitative symptoms. Given that adults with PTSD following childhood maltreatment 112 scored significantly higher on cognitive dysfunction than those without such experiences (47), and 113 also given that cPTSD is often associated with prolonged, repeated traumatic experiences during

- 114 childhood, it is plausible that cPTSD may incur greater cognitive consequences than PTSD.
- 115 Preliminary findings support this hypothesis, demonstrating that childhood trauma is linked to
- 116 subjective and objective cognitive difficulties among people with psychosis, including working
- 117 memory and attention (48,49). Therefore, it is possible that DSOs rather than core PTSD symptoms
- 118 may contribute to explaining cognitive and/or excitative symptoms of psychosis.

119

This study aimed to describe rates of cPTSD and PTSD in a trauma-exposed sample of people with psychosis, assess clinical differences in symptom severity between trauma groups, and explore the relative contribution of cPTSD via PTSD and DSOs in explaining the relationship between trauma and psychotic symptoms. Using a parallel mediation framework (illustrated in Figure 1), we tested separate mediational models to test whether PTSD and DSO symptoms mediated between trauma and positive, negative, affective, cognitive and excitative symptoms.

126

127 **2 METHOD**

128 2.3 Study design

129 This study employed a correlational design, combining data collected from a feasibility randomised

130 controlled trial testing the feasibility and acceptability of a Eye-Movement Desensitisation and

131 Reprocessing for psychosis intervention (the EASE trial) (57), and a previous research project within

132 the University of Manchester Complex Trauma & Resilience Research Unit (58). Both samples were

133 recruited from North West of England mental health services with ethical approval received from an

134 National Health Service (NHS) research ethics committee.

135

136 2.4 Participants

137 This study employed the baseline sample of the EASE trial (n = 66) and the full sample recruited by

138 Capodonico and colleagues (n = 85), thus N = 151.

139

140 The inclusion criteria for the parent studies are reported in full elsewhere (57,58). Generally, these

- 141 included adults with a schizophrenia-spectrum diagnosis (or who met diagnostic criteria), that were
- 142 registered with local NHS mental health services, and had capacity to provide informed consent at
- 143 the time of recruitment. Exclusion criteria for both studies were requirement of an interpreter, a
- 144 primary diagnosis of substance misuse, intellectual disability or gross cognitive dysfunction.
- 145 In addition to those set out by the parent studies, inclusion criteria for this study included
- 146 endorsement of at least one traumatic life event on the Trauma and Life Events checklist (TALE; 33),
- 147 with International Trauma Questionnaire (60) scores anchored to traumas identified by the TALE.
- 148 Participants must also have completed the Positive and Negative Syndrome Scale (61).
- 149

150 **2.5 Measures**

151 The Positive and Negative Syndrome Scale (PANSS; 47) is a 30-item semi-structured clinical

152 interview used to measure psychotic symptoms and general psychopathology. Items are scored on a

153 Likert scale from 1 ('Absent') to 7 ('Extreme'), with higher scores indicating more severe symptoms.

154 This study employed a pentagonal model of the PANSS, in accordance with factor analytic evidence

155 (44,45). This is comprised of 5 factors: positive, negative, cognitive (measuring cognitive

156 disorganization), affective (measuring anxiety and depression) and excitative (measuring activity and

157 hostility); see Supplementary Table 1. Cronbach's α in this sample for these subscales were good,

158 ranging from .73 to .83, aside from the negative subscale, where $\alpha = .66$. This is comparable to prior

159 research (62). All PANSS interviews were administered by trained and supervised research

160 assistants/workers who completed thorough reliability assessments against 'gold standard' scores

161 produced by expert PANSS raters. Raters across datasets demonstrated excellent inter-rater reliability

162 with 'gold standard' scores, with baseline intra-class correlation coefficients (ICCs) ranging from .86

163 to .94 in the EASE dataset, and .90 in the Campodonico dataset.

164

165 The Trauma and Life Events Questionnaire (TALE; 33) is a 22-item self-report checklist assessing

166 traumatic and difficult life experiences. Each event is scored for its occurrence, whether this was

167 more than once, and whether this occurred when the participant was under/over 16 or both. The

- 168 number of traumas endorsed is summed to derive a traumatic experiences score. The TALE
- 169 demonstrates good test-retest reliability and convergent validity with related trauma measures (33).
- 170

171 The International Trauma Ouestionnaire (ITO; 46) is an 18-item self-report scale assessing the 172 presence and severity of PTSD and DSOs within the past month. Items are administered in relation to 173 an index trauma identified on the TALE as affecting the individual most in the past month. PTSD and 174 DSO subscales are each comprised of 3 symptom clusters, themselves comprised of 2 items each. 175 Both subscales include 3 additional items measuring functional impairment associated with the 176 symptoms captured by each subscale. All items are scored on a 5-point Likert scale from 0 ('Not at 177 all') to 4 ('Extremely'). As per the ITQ diagnostic algorithm, a probable diagnosis of PTSD is 178 identified when a participant presents with a score of 2 or more on at least one item in each PTSD 179 cluster, plus a score of 2 or more on at least one functional impairment item associated with these 180 symptoms. The cPTSD threshold includes that of PTSD, as well as a score of 2 or more on at least 181 one item in each DSO cluster, plus a score of 2 or more on at least one functional impairment item 182 associated with these symptoms. PTSD and DSO items were totaled to derive continuous severity 183 scores, with higher scores indicating higher severity. Scores within each DSO cluster were summed 184 to derive continuous DSO scores for use in planned secondary analyses. Both PTSD and DSO subscales demonstrated high internal consistency in this sample ($\alpha = .83$ and .87, respectively). 185

186

187 **2.6 Procedures**

Participants were first introduced to either parent study by members of their clinical teams, and signposted to the research team, if interested in participating. Participants then met with a researcher for more information about the relevant study at a mutually convenient location or a digital meeting (during the COVID-19 pandemic in affected periods of the EASE trial). Information sheets were then provided, and informed consent taken. Following informed consent, participants completed a battery of measures that included the PANSS, TALE and ITQ, in addition to other measures (see (57,58).

194

195 **2.7 Data analysis**

196 Diagnostic algorithms outlined above were applied to ITQ scores to group participants into those

197 who met criteria for PTSD, cPTSD or neither. Independent samples t-tests were then used to explore

198 mean differences in psychotic symptoms scores between these trauma groups. Correlation, regression

and finally mediation analyses were used to examine the role of PTSD and DSO symptoms in the

200 relationship between trauma and symptom dimensions. The Baron and Kenny (1986) requirement for

201 the presence of a significant direct effect was used here as a conservative option to indicate

202 subsequent mediation analyses, to minimize Type I error.

203

Out of 151 participants available from the parent studies, 144 people met inclusion criteria for this

study. No cases were identified as multivariate outliers per Mahalanobis (64), Cook's (65) or

206 Leverage analyses of distance. The data were then checked for normality and homoscedasticity; both

207 assumptions were met per Curran and colleagues' criteria (66).

208

209 Within this dataset, less than 20% of the data were missing. Little's MCAR test was not significant, $\chi^2 = 257.59(251), p = .37$, suggesting the data were missing at random. Thus, the missing scores 210 211 were imputed where possible. Eight participants were missing PANSS scores on specific items likely 212 obfuscated by remote assessments in the EASE trial; these participants were excluded from analyses 213 involving these items. Where one of the two scores on an ITO symptom was missing, this was 214 imputed with the score on the other item. Where an ITQ functional impact score was missing, this 215 was imputed with the mean of the other two available scores. One participant chose not to respond to 216 one item on the TALE; this was not imputed, as trauma checklists are likely not missing at random. 217 Then, specific items were summed to derive continuous TALE, PTSD, DSO, and PANSS subscale 218 scores.

219

220 Twenty-four participants in the EASE trial completed their assessments remotely due to the COVID-

221 19 lockdown restrictions on face-to-face working. This impacted the confidence of negative PANSS

ratings reliant on factors more easily observed in person (e.g., gesticulation). Analyses were rerun

with these participants excluded (n = 120), to check the validity of the full dataset. This did not affect

the outcomes of the study. As such, the following findings are in relation to the full combined

sample.

226

227	To capture potential confounding differences between parent datasets, dataset membership was
228	entered as a covariate in our analyses. This did not affect the outcomes of PANSS-negative or -
229	affective analyses. As such, the analyses of these outcomes reported below do not include covariates.
230	Dataset did, however, change the outcomes of analyses of PANSS-positive scores. Thus, the analyses
231	thereof include dataset as a covariate. Bonferroni-corrected independent samples <i>t</i> -tests and bivariate
232	Pearson correlations were used to assess the role of gender and age, respectively, as potential
233	covariates. The only significant finding was a small, positive correlation between PANSS-cognitive
234	scores and age ($r = .19$, $p = .027$). Age would therefore be included in analyses of PANSS-cognitive
235	scores, but these were not indicated by bivariate correlations (see below).
236	
237	3 RESULTS
238	3.3 Descriptive statistics
239	Demographic and clinical characteristics were aggregated across datasets, reported in Table 1.
240	Participants reported a mean of 9.7 traumatic life experiences on the TALE. The most common
241	experiences endorsed were loss or permanent separation from a close friend or relative (81%),
242	bullying (70%), emotional (66%) and physical (62%) abuse.
243	
244	3.4 The frequency of cPTSD
245	The ITQ diagnostic algorithm was applied to ITQ scores of the sample to delineate groups of
246	participants meeting ICD-11 criteria for PTSD and cPTSD. Among those who met criteria for a post-
247	traumatic stress diagnosis (50.7%), cPTSD was far more common (40.3%) than PTSD (10.4%).
248	
249	3.5 Symptom severity between trauma group

250 A between-subjects MANOVA found a significant overall trauma diagnosis group differences across 251 PANSS subscales ($F_{10, 252} = 4.702$, p < .001, Wilk's $\Lambda = .710$). Follow-up univariate ANOVAs 252 detected significant differences on positive ($F_2 = 6.02$, p = .003, $\eta_p^2 = .09$), negative ($F_2 = 5.94$, p = .09) .003, $\eta_p^2 = .08$) and affective ($F_2 = 17.16$, p < .001, $\eta_p^2 = .21$) PANSS subscales. No significant 253 differences among cognitive ($F_2 = 1.30$, p = .276, $\eta_p^2 = .02$) or excitative ($F_2 = 6.14$, p = .345, $\eta_p^2 = .02$) 254 255 .02) subscales were observed. Post-hoc Tukey's tests were used to investigate significant differences 256 between groups. These suggested that positive symptoms were significantly higher among those 257 meeting criteria for cPTSD (M = 16.86, SD = 3.93) than those who met criteria for neither cPTSD 258 nor PTSD (M = 13.93, SD = 5.13), as were negative symptoms (cPTSD: M = 15.41, SD = 5.40; none: 259 M = 12.66, SD = 4.21). Affective symptoms were significantly higher among those meeting criteria 260 for cPTSD (M = 17.51, SD = 4.22) compared to those meeting criteria for PTSD (M = 13.86, SD =261 3.59) or neither (M = 12.40, SD = 5.26).

262

263 Another between-subjects MANOVA was used to assess differences among PTSD, DSO and TALE scores. This was also significant ($F_{6, 264} = 27.16$, p < .001, Wilk's $\Lambda = .38$). Follow-up univariate 264 ANOVAs showed significant differences between PTSD ($F_2 = 72.80$, p < .001, $\eta_p^2 = .52$), DSO ($F_2 =$ 265 26.89, p < .001, $\eta_p^2 = .29$) and TALE scores ($F_2 = 14.95$, p < .001, $\eta_p^2 = .18$). Post-hoc Tukey's tests 266 were used in an attempt to replicate prior findings of PTSD symptom severity in people with cPTSD. 267 268 PTSD scores were significantly higher among those meeting criteria for PTSD (M = 15.93, SD =269 4.42) and cPTSD (M = 18.66, SD = 3.98) than those who did not meet criteria (M = 8.09, SD = 5.32). 270 DSO scores were significantly higher among those meeting criteria for cPTSD (M = 18.98, SD =271 3.60) than those meeting criteria for PTSD (M = 9.80, SD = 4.90) or neither (M = 11.90, SD = 7.06). 272 Lastly, TALE scores were significantly higher among those meeting criteria for cPTSD (M = 11.61, 273 SD = 2.93) than those meeting PTSD criteria (M = 8.93, SD = 3.73) or neither (M = 8.39, SD = 3.57).

274

275 **3.6 Hierarchical Regressions**

276 Bivariate Pearson correlations are presented in Table 1. These did not indicate a need for further

277 regressions on excitatory subscales, as no significant relationships were observed with predictor

278 variables. Hierarchical regressions were then used to assess whether DSO scores predicted positive,

- 279 negative, cognitive and affective PANSS scores, and whether these associations survived the addition
- 280 of PTSD scores as a covariate (see Table 2 for coefficients). As neither PTSD nor DSOs were
- 281 significant predictors of negative or cognitive subscale scores, exploratory mediation analyses were
- 282 only assessed for positive and affective subscales.
- 283

284 **3.7 Mediation analyses**

285 **3.7.1 Positive symptoms**

- 286 Exploratory parallel mediation analyses were conducted via the SPSS PROCESS macro (model 4;
- 40) to assess whether PTSD and/or DSOs mediate the relationship between TALE and positive
- 288 PANSS scores, including dataset as a covariate. TALE scores significantly predicted PTSD (b = .597,
- 289 $t_{137} = 3.989, p < .001, 95\%$ CI [.349, .845]) and DSOs ($b = .530, t_{137} = 3.648, p < .001, 95\%$ CI [.290,
- 290 .771]). Dataset also significantly predicted both PTSD (b = -3.463, $t_{137} = -3.097$, p = .002, 95% CI [-
- 291 5.315, -1.611]) and DSOs (b = -4.008, $t_{137} = -3.689$, p < .001, 95% CI [-5.807, -2.209]). When
- 292 controlling for PTSD, DSOs and dataset, TALE scores no longer predicted positive PANSS scores (b
- 293 = .144, t_{135} = 1.338, p = .183, 95% CI [-.034, .321]). Both PTSD (b = .240, t_{135} = 3.650, p < .001,
- 294 95% CI [.131, .349]) and DSO scores ($b = .143, t_{135} = 2.118, p = .036, 95\%$ CI [.031, .255])
- significantly mediated the relationship between TALE scores and positive PANSS scores. Dataset
- 296 significantly predicted positive PANSS scores ($b = 1.770, t_{135} = 2.235, p = .027, 95\%$ CI [.458,
- 3.082]). Regression statistics are presented in Table 3, and results displayed graphically in Figure 2.

298

299 **3.7.2 Affective symptoms**

- 300 A similar mediation analysis was conducted to assess whether PTSD and DSOs mediate the
- 301 relationship between TALE and affective PANSS scores. TALE scores significantly predicted PTSD

302 $(b = .711, t_{132} = 3.998, p < .001, 95\%$ CI [.464, .958]) and DSOs $(b = .649, t_{132} = 4.375, p < .001, t_{132} = 4.375$

- 303 95% CI [.403, .895]). When controlling for PTSD and DSOs, TALE scores no longer predicted
- 304 positive PANSS scores (b = .086, $t_{130} = .840$, p = .403, 95% CI [-.084, .257). PTSD scores
- 305 significantly mediated the relationship between TALE scores and positive PANSS scores (b = .212,
- 306 $t_{130} = 3.342, p = .001, 95\%$ CI [.107, .317]), as did DSO scores ($b = .328, t_{130} = 5.139, p < .001, 95\%$

307 CI [.222, .434]). Regression statistics are presented in Table 3, and results displayed graphically in
 308 Figure 3.

309 4 **DISCUSSION**

To the authors' knowledge, this is the first study to investigate the frequency and correlates of cPTSD in a trauma-exposed sample of people with psychosis. A higher proportion of the sample met criteria for cPTSD than PTSD. The results suggest that participants meeting criteria for cPTSD presented with significantly higher positive, negative and affective symptoms than those who did not meet criteria for a trauma diagnosis. These did not differ from those meeting criteria for PTSD, aside from affective symptoms, which were significantly higher in the cPTSD group.

316

317 The frequency of cPTSD in our sample may suggest that PTSD symptoms in psychosis occur in the 318 context of cPTSD. In this sense, our mediation models may hint at the role of cPTSD in the trauma-319 psychosis pathway. Consistent with hypotheses, PTSD and DSO symptoms mediated the relationship 320 between trauma and positive symptoms. Of note, the DSO path became significant following the 321 inclusion of dataset as a covariate. The mediation analyses were rerun in each dataset to tentatively 322 investigate the validity of this effect; the coefficients of the DSO path closely resembled that of the 323 adjusted, pooled analysis reported above, though these did not reach statistical significance in either 324 dataset. This pattern of results suggests the samples of each parent dataset were inadequately 325 powered to test these multivariate models individually. This preserves the validity of our results, but 326 clearly requires further replication in larger samples. Also as hypothesized, both PTSD and DSOs 327 were found to mediate the relationship between trauma and affective symptoms. However, significant 328 relationships between excitative symptoms, PTSD and DSOs were not observed, and neither PTSD 329 nor DSOs significantly predicted negative or cognitive symptoms, contraindicating further regression 330 and mediation analyses.

331

332 Our main aim of assessing how common cPTSD is relative to PTSD in a trauma-exposed sample of

333 people with psychosis was met. The considerable proportion of participants meeting criteria for

334 cPTSD is consistent with prior research demonstrating the prevalence of difficulties consistent with

335 DSOs among those with psychosis (22,23,68). The finding that cPTSD may be more common than 336 PTSD is consistent with other clinical samples (e.g., 11,34.42). This may be unsurprising, 337 considering the prevalence of complex interpersonal trauma in this population, thought to underpin 338 cPTSD (see (25). We did not replicate prior findings of increased PTSD symptom severity in people 339 meeting cPTSD (12,13) criteria, though our descriptive statistics between groups hint at these 340 findings in larger samples of people with psychosis. These results demonstrate the importance of 341 trauma-informed approaches to assessment and treatment of those with psychosis, owing to the 342 potentially high prevalence of cPTSD that larger epidemiological studies could confirm. Again, this 343 could contextualize the reliability of PTSD mediating the trauma-psychosis relationship in terms of 344 cPTSD, with specific symptoms thereof predicting specific psychotic phenomena. This is consistent 345 with prior findings (24), but requires further empirical investigation.

346

347 The second aim of this study exploring the relative contribution of cPTSD via PTSD and DSOs in 348 explaining the relationship between trauma and psychotic symptoms was also met. The finding that 349 PTSD symptoms mediate the relationship between trauma and positive symptoms is consistent with 350 prior systematic reviews (21–23) and meta-analyses (24). Indeed, it may be that re-experiencing 351 symptoms of PTSD underpin positive symptoms in certain psychosis subgroups (70). The finding 352 that DSOs also mediate this is not surprising, owing to the theoretical consistency between these and 353 related constructs previously shown to predict hallucinations and delusions (21,23). Notably, 354 however, the effect size of the PTSD path is almost double that of the DSO path, suggesting the way 355 in which cPTSD may precipitate or maintain positive symptoms may primarily occur via PTSD 356 symptoms, rather than DSOs. This finding could be explained by the sensory phenomenology of re-357 experiencing symptoms, compared to the psychological phenomenology of DSOs in cognitive and 358 emotional patterns (71), leaving core PTSD symptoms open to interpretation as anomalous 359 experiences (20). That said, re-experiencing symptoms have been shown not to correlate with 360 positive PANSS scores (72). Indeed, recent network analyses have demonstrated that trauma-related 361 beliefs and hypervigilance may be more closely related to positive symptoms of psychosis than reexperiencing symptoms (73). Therefore, it may be that the complexity of trauma histories among 362 363 those with cPTSD lead to trauma beliefs dissimilar to those with PTSD that predict positive 364 symptoms. However, qualitative (10) and quantitative (74) research suggests that trauma-related 365 beliefs among those with cPTSD are similar to those with PTSD. The present study therefore requires

366 replication to assess the reliability of these findings, to further delineate the mechanism by which

- 367 DSOs may impact psychotic symptoms.
- 368

369 Our findings suggest that cPTSD may underpin the affective difficulties among people with 370 psychosis. This would be unsurprising, considering cPTSD is more strongly correlated with anxiety 371 and depressive disorders than PTSD (75,76). Affective difficulties (i.e., anxiety and depressive 372 disorders) are highly comorbid with psychosis (27). Prospective studies suggest anxiety and 373 depression mediate the relationship between childhood victimisation and adolescent psychotic-like 374 experiences (77), consistent with the affective pathway to psychosis (78). Affective difficulties may 375 also maintain psychoses, posing a higher risk of maladaptive appraisals and behavioural responses to 376 psychotic experiences that perpetuate the experience (79). Together with our findings, this literature 377 could imply an aetiological or maintaining role of cPTSD in the affective pathway to psychosis. This 378 requires exploration in longitudinal studies, which could indicate the adjustment of trauma-focused cognitive behaviour therapy for psychosis to account for cPTSD symptoms. Considering our model 379 380 clustered DSOs together, such research may investigate symptom-specific relationships between 381 cPTSD and psychosis to uncover finer mechanisms for therapeutic targets. These were not explored 382 here following considerations of statistical power.

383

384 The small correlation between negative symptoms of psychosis and all trauma-related variables is 385 consistent with the literature. Meta-analytic evidence has found no relationship between childhood 386 trauma and negative symptoms (80), and that TF-CBT may not lead to significant improvements in 387 this domain (32). Whilst reliable, these findings seem at odds with the consistency between negative 388 and post-traumatic sequelae (e.g., interpersonal difficulties and social withdrawal, (34)), as well as 389 recent network analyses demonstrating paths between specific adverse childhood experiences and 390 negative symptoms (81). Negative symptoms may be divided conceptually into experiential and 391 expressive subgroups (82); it may be that the PANSS score does not reflect this complexity that is 392 captured by, for instance, the Scale for the Assessment of Negative Symptoms (83). Similar 393 arguments may be applied to the measurement of positive symptoms in this study, considering 394 experiences such as hallucinations and delusions were collapsed into a single score, despite research

395 demonstrating specific trauma pathways to each (24,70). Future studies may therefore adopt a 396 symptom-specific approach to assess whether and how trauma and post-traumatic sequelae may 397 predict specific psychotic symptoms. Prior studies have also shown no relationship between 398 cognitive or excitative symptoms and PTSD (41,42); our findings are consistent with these. It is 399 outlined above that high cognitive/excitative symptoms may reflect a subgroup of individuals 400 experiencing psychosis via a neurodevelopmental pathway. Whilst atypical neurodevelopment may 401 pose a risk for childhood victimization, this may not necessarily incur post-traumatic 402 symptomatology (43). This further demonstrates the importance of assessing for trauma-related 403 difficulties in psychosis, to inform whether said difficulties are incorporated into formulations and 404 treatment plans.

405

406 A number of methodological limitations should be noted. For instance, the relatively small number of 407 participants meeting criteria for PTSD may have reduced statistical power – G*Power analyses 408 indicate an achieved power of 0.4 in the comparison of positive symptoms between trauma groups, 409 far below the acceptable 0.8 (84). Our finding that those meeting criteria for neither PTSD nor 410 cPTSD significantly differed in psychotic symptoms to those meeting cPTSD criteria may indicate a 411 stepwise increase in psychotic symptom severity along the spectrum of trauma-related diagnoses. 412 Future research in larger samples may better assess this, to delineate the clinical utility of diagnosing 413 cPTSD separately from PTSD among those with psychosis.

414

415 The unidimensional measurement of emotional dysregulation on the ITQ may not represent the 416 complexity of affective difficulties among those with cPTSD. The ITQ currently includes two items 417 - one assessing emotional hyperactivation and the other hypoactivation - that are summed to derive a 418 single emotional dysregulation score, as per ICD-11 guidance (85). Prior studies suggest that a 419 bifactor structure of this DSO reflecting these dimensions - and measured by more items - may 420 provide a better fit (69,86). This would align with research (e.g., 42) demonstrating the specificity of 421 certain traumas with specific affective regulatory difficulties. Further, as a self-report measure, 422 diagnostic categories assigned by the ITQ may not be as valid as, for instance, a structured clinical 423 interview. The ITQ is the only validated psychometric measure of cPTSD, though Lechner-

- 424 Meichsner and Steil (2021) propose updates to the Clinician-Administered PTSD Scale to diagnose
- 425 cPTSD. Perhaps these limitations may be considered in ongoing development of the ITQ.
- 426

427 Our models could not be adjusted for potential clinical confounds. Substance misuse is common 428 among trauma survivors (89) and has been shown to predict psychosis (90), but was unfortunately 429 not measured by the parent datasets. This could arguably act as a covariate in our model, though this 430 is unclear based on prior systematic reviews (21). Another potential confound is dissociation – an 431 associated, but not core, feature of both PTSD and cPTSD that has been shown to mediate the 432 relationship between trauma and psychosis (21,23). Similarly, further exploration of contextual 433 factors (e.g., lack of emotional resources) that may moderate the expression of complex trauma 434 symptoms and their relationships with psychotic outcomes is warranted. Future research controlling 435 for these confounds could elucidate further the relative contribution of cPTSD in explaining the 436 trauma-psychosis relationship.

437

438 Owing to the cross-sectional nature of this study, causal interpretations of our model should be very 439 tentative. The possibility of reverse causation from positive and/or affective symptoms to PTSD and 440 DSOs cannot be completely discounted, though previous prospective studies make this unlikely (91). 441 Paradigms focused on finer-grained measurement of cPTSD and psychosis may prove fruitful in 442 future research refining these models. One paradigm - ecological sampling methods (ESM) – may 443 address both questions, enabling a fine-grained assessment of whether DSOs and psychosis interact 444 in the flow of daily life. One study (92) found emotion regulation predicted psychosis symptoms in 445 daily life, despite no association between retrospective measures of the same variables, 446 demonstrating the utility of ESM in overcoming common methodological limitations (e.g., recall 447 bias).

448

449 The sample may constitute a limitation of this study. For one, it is predominantly white. The

450 prevalence of psychosis in minority ethnic groups (93) coupled with the trauma of systemic racial

451 discrimination could mean psychotic symptoms and DSOs interact differently in minoritised groups

452 than in their white counterparts. Indeed, systemic racism may decrease self-concept clarity (94),

453 known to be common in people with psychosis (95) building the plausibility of this argument. Future

454 research into this area may explore the generalisability of these findings to black and minority ethnic

455 populations; such research may have implications for sociodevelopmental pathways to psychosis (see

456 (96). Including those with affective psychosis may pose a further limitation of this sample,

457 considering biological sequelae robustly mediate the relationship between trauma and bipolar

458 disorder, as opposed to symptoms of post-traumatic stress (97). This heterogeneity may have diluted

459 the effects of cPTSD in this sample. However, diagnoses were not collected using gold-standard

460 tools. They were validated with clinical services and medical notes, though exact confounding effects

461 are therefore difficult to determine. Future studies into the impact of cPTSD in psychosis may choose

462 to focus specifically on non-affective psychosis samples to avoid such effects.

463

464 Our findings suggest a potentially nuanced impact of cPTSD in people with psychosis. Rates of 465 cPTSD in our sample suggest that the 'core' symptoms of PTSD (i.e., re-experiencing, hyperarousal 466 and avoidance) mostly occur in the context of cPTSD, impacting or maintaining positive symptoms. 467 Both symptom domains of cPTSD (i.e., PTSD and DSOs) may play a role in the maintenance of 468 positive symptoms and affective difficulties (i.e., anxiety and depression) among people with 469 psychosis. Therefore, whilst trauma-focused interventions may be effective at addressing symptoms 470 of trauma and psychosis in people with psychosis (98), a broader range of treatment options may 471 need to be developed to address DSO-related difficulties. For example, dialectic behavioural skills 472 training, demonstrated as most effective when used as an adjunct therapy (99) and more effective for 473 cPTSD than cognitive processing therapy (100). In any case, diagnostic criteria on the ITQ require a 474 functional impact of DSOs in at least one domain, demonstrating an increased treatment need among 475 those with psychosis and comorbid cPTSD. Future research testing the reliability of our findings in 476 larger, more representative samples may therefore have important assessment and treatment 477 implications as trauma-informed care becomes the norm in psychosis services. Such research could 478 employ intense longitudinal designs and adjust for potential confounds to further refine our models.

479

480 **4.3 Conclusions**

- 481 This is the first study to the authors' knowledge investigating the frequency of cPTSD in a trauma-
- 482 exposed psychosis sample. In accordance with prior research, cPTSD was more common than PTSD
- 483 in this sample, comprised of both early and chronic psychosis presentations. The functional
- 484 impairment required to meet cPTSD criteria, as well as the potentially maintaining role cPTSD in
- 485 positive and affective psychotic symptoms, demands further research into the impact of this
- 486 comorbidity. This impact may be more nuanced than first thought, with symptom-specific
- 487 relationships affecting individuals in different ways. Future research may investigate the relationship
- 488 between symptoms of cPTSD and psychosis at a momentary level, to assess this potentially dynamic
- 489 interplay between cPTSD and psychosis, as well as the reliability of these results.
- 490

491 **5** Conflict of Interest

The authors declare that the research was conducted in the absence of any commercial or financial
 relationships that could be construed as a potential conflict of interest.

494 6 Author Contributions

495 PP, FV, KB and WS contributed to the conception and design of the study. FV, WS, CC and RPB
496 were involved in collecting the data. PP analysed the data, and wrote the first draft of the manuscript.
497 FV, KB, WS and RPB contributed to the final draft of the manuscript. All authors contributed to the
498 article and approved its submission.

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- and not necessarily those of the NIHR or the Department of Health and Social Care.

505 8 Data Availability Statement

- 506 The datasets analysed for this study are not available due to the sensitivity of the dataset and
- 507 participant confidentiality.

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874



875 Tables

876 **Table 1.** Bivariate Pearson correlations between PANSS subscales and trauma variables

	Positive	Negative	Cognitive	Affective	Excitative	TALE score	PTSD	DSO	М	SD
Positive	-	.13	.21*	.46**	.12	.26**	.45**	.38**	15.4	4.8
Negative		-	.17	.39**	03	.06	.28**	.28**	13.8	4.9
Cognitive			-	.12	.52**	09	17*	15	13.1	4.3
Affective				-	05	.32**	.55**	.61**	14.5	5.2
Excitative					-	13	13	10	4.9	2.3
TALE score						-	.38**	.37**	9.7	3.6
PTSD							-	.59**	13.2	6.9
DSO								-	14.5	6.8

Note: PANSS = Positive and Negative Syndrome Scale, TALE = Trauma and Life Events Checklist, PTSD = Post-traumatic Stress Disorder, DSO = Disturbances of Self-Organisation, M = mean, SD = standard
 deviation.

879 *Significant at the p < .05 level

880 **Significant at the p < .01 level

Predictor	Positive	Negative	Cognitive	Affective
Step 1				
DSOs	.383**	.284*	154	.606**
F	23.737	11.501	3.138	76.773
R^2	.212	.086	.016	.363
Step 2				
DSOs	.178	.184	079	.436**
PTSD	.344**	.171	125	.295**
ΔF	13.533	2.761	1.349	13.267
ΔR^2	.077	.019	.010	.058

881	Table 2. Standardised coefficients of hierarc	chical regressions predicting PANSS scores

882 Note: DSOs = Disturbance of Self Organisation; PTSD = Post-Traumatic Stress Disorder

883 **p* < .01

884 ***p* < .001

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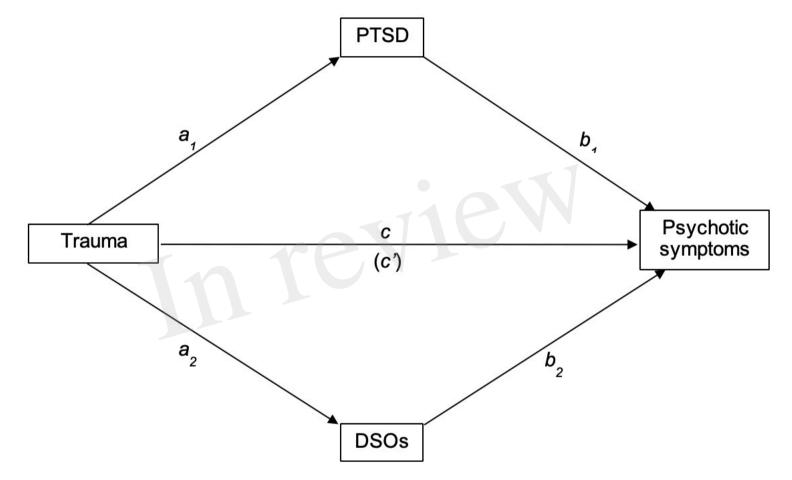
	Р	ositive symptom	Affective symptoms			
Path	R^2	F	р	R^2	F	р
<i>a</i> ₁	.201	17.274	< .001	.177	27.456	< .001
b_1	.211	18.351	< .001	.139	20.596	<.001
c'	.063	10.496	.001	.097	15.505	< .001
С	.257	11.687	< .001	.224	12.125	<.001

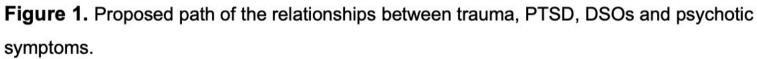
890 **Table 3.** Regression statistics for the mediation pathways predicting psychotic symptoms

891 Note: $a1 = \text{TALE} \rightarrow \text{PTSD}, a2 = \text{TALE} \rightarrow \text{DSO}, c' = \text{TALE} \rightarrow \text{PANSS-Positive}, c = \text{TALE} \rightarrow \text{PTSD} + \text{DSO} \rightarrow \text{PANSS-Positive}.$ Positive symptom

892 coefficients include dataset as a covariate.

893





Note: PTSD = Post-Traumatic Stress Disorder, DSOs = Disturbances of Self-Organisation

Figure 2.JPEG

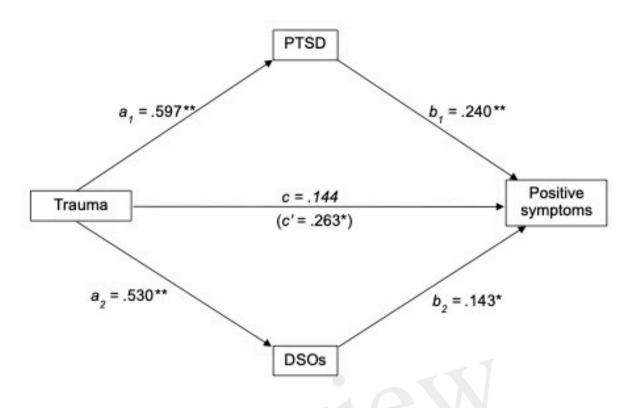
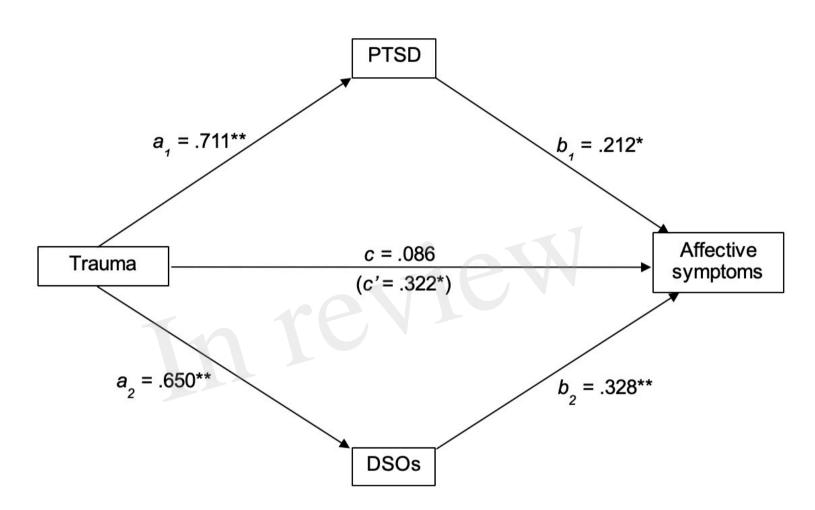


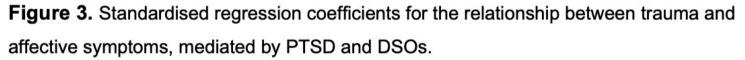
Figure 2. Standardised regression coefficients for the relationship between trauma and positive symptoms, mediated by PTSD and DSOs, including dataset as a covariate.

**,p < .005

*p < .05

Note: PTSD = Post-Traumatic Stress Disorder, DSOs = Disturbances of Self-Organisation





**p < .005

*p < .05

Note: PTSD = Post-Traumatic Stress Disorder, DSOs = Disturbances of Self-Organisation