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#### JUMPING TO CONCLUSIONS AT FIRST ONSET OF PSYCHOSIS PREDICTS LONGER ADMISSIONS, MORE COMPULSORY ADMISSIONS AND POLICE INVOLVEMENT OVER THE NEXT 4 YEARS; the GAP Study.

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#### ABSTRACT

**Background.** Jumping to Conclusions (JTC), which is the proneness to require less information before forming beliefs or making a decision, has been related to formation and maintenance of delusions. Using data from the GAP case-control study of first-episode psychosis (FEP), we set out to test whether the presence of JTC would predict poor clinical outcome at 4 years.

**Methods.** 123 FEP patients were assessed with the positive and negative syndrome scale (PANSS), Global Assessment of Functioning (GAF), and the probabilistic reasoning "Beads" task at the time of recruitment. The sample was split into two groups based on the presence of JTC bias. Follow-up data over an average of 4 years were obtained concerning clinical course and outcomes (remission, intervention of police, use of involuntary treatment – the Mental Health Act (MHA) -, and inpatient days).

**Results.** FEP who presented JTC at baseline were more likely during the follow up period to be detained under the MHA [adjusted OR=15.62, 95% confidence interval (CI) 2.92-83.54, p=0.001], require intervention by the police (adjusted OR=14.95, 95% CI 2.68-83.34, p=0.002) and have longer admissions (adjusted IRR=5.03, 95% CI 1.91-13.24, p=0.001). These associations were not accounted for by socio-demographic variables, IQ and symptom dimensions.

**Conclusions.** JTC in FEP is associated with poorer outcome as indicated by defined by more compulsion police intervention, and longer periods of admission.. Our findings raise the question of whether the implementation of specific interventions to reduce JTC, such as Metacognition Training, may be a useful addition in early psychosis intervention programs. *Abstract word count: 249* 

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# 1 1. INTRODUCTION

Psychosis especially schizophrenia may be a disabling condition and classically is associated
with poor clinical outcome. Nonetheless, recent findings confirm that prognosis is not
universally poor and that the course need not be one of inexorable decline (Hopper *et al.*,
2007; Morgan *et al.*, 2014; Revier *et al.*, 2015). Interest has therefore shifted towards
identifying predictors of outcome for treatment planning in order to ameliorate the adverse
impact of the illness (White *et al.*, 2009; Juola *et al.*, 2013; Friis *et al.*, 2016).

Neurocognitive deficits and negative symptoms are important drivers of disability in psychosis
(Breier *et al.*, 1991; Wieselgren, Lindström and Lindström, 1996; Ho *et al.*, 1998; Green *et al.*,
2000; Lipkovich *et al.*, 2009; Faber *et al.*, 2011), as is social cognition (Couture, Penn and
Roberts, 2006; Fett *et al.*, 2011; Pinkham, 2014). However, which aspects of cognition are
most predictive of prognosis remains elusive.

One area to attract attention in this regard is reasoning and cognitive biases (van Hooren et 13 14 al., 2008). Jumping to conclusions (JTC) defines a tendency to form beliefs and to make a 15 decision about an event without having enough information about it, sometimes referred to as 16 data-gathering bias (Freeman, Pugh and Garety, 2008). JTC plays a central role in psychological and neuropsychological theories of delusions as it is considered to lead to the 17 rapid acceptance of implausible ideas and to prevent consideration of more realistic alternative 18 19 explanations of events (Freeman and Garety, 2014). A recent meta-analysis (Dudley et al., 20 2016) summarized evidence that people with psychosis make decisions on the basis of little 21 evidence, have a more extreme reasoning style than people with other mental health conditions or healthy controls and that JTC is linked with a higher probability of having 22 delusions. Moreover, there is evidence that such hasty reasoning is predictive of less 23 24 improvement over time in delusions (So et al., 2014).

In her theoretical model of psychosis, Garety et al. (2001) proposed JTC as one of the biased 25 conscious appraisal processes which are crucial in contributing to the perception of anomalous 26 experiences as personally significant and externally caused when the subject is in search for 27 an explanation (Garety et al., 2001). Garety and Freeman (1999) found empirical support for 28 29 JTC, externalizing attributional biases and deficits in understanding social situations and the intentions of others, to be specific biases in these processes (Garety and Freeman, 1999). 30 Interestingly, these biases and deficits, which are considered to be part of social cognition, are 31 also included in similar constructs as metacognition (Lysaker et al., 2005, 2013), which has 32 led to the development of specific intervention programmes, as the Metacognition Training 33 designed by Moritz, which has specific modules dedicated to the bias (Moritz and Woodward, 34 2007). This latter intervention has been reported to improve positive symptoms (Aghotor et 35 al., 2010; Moritz et al., 2011, 2013; Favrod et al., 2014) in patients with schizophrenia, which 36 renders the JTC bias as especially interesting for research into the outcome of the disorder. 37

There is also evidence of a link between JTC and the more classic construct of neurocognition. JTC has been shown to be negatively associated with neuropsychological performance in FEP samples (Falcone *et al.*, 2015; González *et al.*, 2017), but also with lower IQ scores among healthy relatives of patients with schizophrenia (Van Dael *et al.*, 2006) and controls with high levels of psychotic experiences (Mortimer *et al.*, 1996; Van Dael *et al.*, 2006). Nonetheless, whether the relationship between neurocognition and JTC moderate the link with positive symptoms remain elusive (Andreou *et al.*, 2015). 45 To the best of our knowledge, only one study has examined whether JTC moderates outcome of the disorder by measuring its impact on functional outcome (Andreou et al., 2014). The 46 results did not support a predictor role of the bias, although this may have been because of 47 48 the length of follow-up was only 6 months and the sample was small. We decided to examine reasoning bias as a possible predictor of the course and clinical outcome in the medium to 49 long term, using data from a well-characterized sample of patients presenting to psychiatric 50 51 services for the first time with psychosis. The aim of our study was to examine independent associations between JTC and clinical outcome at 4 years after the first contact with mental 52 health services. For this purpose, we selected remission and several clinical outcomes of 53 54 service use such as days of admissions, use of the Mental Health Act (MHA) for involuntary treatment and instances of police involvement during an admission to a psychiatric unit. 55

As JTC has been related in several studies to the presence of delusions (Falcone *et al.*, 2015;

57 McLean *et al.*, 2016), and to a lesser extent to persistence of delusion severity (Falcone *et al.*,

58 2015), we hypothesized that those patients who presented more severe reasoning bias at 59 baseline would have worse clinical prognosis.

# 60 2. METHODOLOGY

#### 61 2.1. Participants

Participants for this study were recruited as part of the National Institute of Health Research 62 Biomedical Research Centre Genetics and Psychosis (GAP) study conducted in South 63 London, UK. Further details of the study are available in Di Forti et al (Di Forti et al., 2015). 64 65 Briefly, the GAP study comprised individuals aged 18-65 years who presented to the 66 psychiatric services of the South London and Maudsley (SLaM) National Health Service (NHS) Foundation Mental Health Trust between December 2005 and October 2010 with a first 67 episode of psychosis (FEP) (International Classification of Diseases (ICD)-10; F20-F29 and 68 F30-F33; and Diagnostic and Statistical Manual of Mental Disorders (DSM)-IV; 295.1-298.9) 69 (WHO, 1992; American Psychiatric Association, 2000). Diagnosis at the moment of 70 recruitment was determined by administration of the Schedules for Clinical Assessment in 71 Neuropsychiatry (SCAN) (WHO, 1994) and was later validated by using the computerized 72 Operational Criteria system (version 2004) (McGuffin et al., 1991). Cases were excluded if 73 there was evidence of: 1) psychotic symptoms precipitated by an organic cause; 2) transient 74 psychotic symptoms resulting from an acute intoxication as defined by ICD-10: 3) head injury 75 causing clinically significant loss of consciousness; and 4) learning disability (IQ<70) as 76 assessed by the Wechsler Adult Intelligence Scale-Third Edition (Wechsler, 1997). The 77 original GAP sample comprised N=431 FEP cases; of these, information on JTC at baseline 78 79 was available for 123 cases (28% of the original GAP sample(Falcone et al., 2015)). This subsample with information on JTC did not differ significantly from the full GAP sample in 80 terms of gender, age, education level, diagnosis or ethnicity (Supplementary Table 1). Further, 81 82 83% (N=102) of the subsample (N=123) was successfully traced four years after first contact with mental health services. Therefore, data presented here are based on these 102 cases. 83 Ethical permission was obtained from the SLaM and the Institute of Psychiatry Research 84 Ethics Committee. All patients gave informed written consent after reading a detailed 85 information sheet. 86

#### 87 2.2. Measures at baseline

#### 88 2.2.1. Socio-demographic characteristics

89 Demographic data were collected using the Medical Research Council (MRC) Sociodemographic Schedule modified version (Mallett et al., 2002) and supplemented by clinical 90 91 records. For educational level, we divided the sample into three categories: No qualification, 92 school education (GCSE, 'O' levels and 'A' levels) and tertiary education (vocational, college, university or professional qualification). We dichotomized the domains of employment 93 (employed vs. unemployed), marital (in a stable relationship vs. no relationship) and living 94 95 arrangements status (independent living vs. no independent living). For lifetime use of alcohol and illegal drugs, we collected data from the GAP baseline measures, and split both into ever 96 used (1) vs. never used (0). Lifetime use of cannabis was assessed with the Cannabis 97 98 Experience Questionnaire modified version (Di Forti et al., 2009), dividing patients into those who reported ever having used cannabis (1) and those who reported never having used it (0). 99 100 Ethnicity was self-ascribed using categories employed by the 2001 UK Census 101 (http://www.ons.gov.uk/ons/guide-method/census/census-2001/index.html). Due to small 102 numbers in some ethnic categories, we combined them into three broad ethnic groups: white 103 (all white groups); black (all black groups); and other (encompassing Asian, mixed ethnicity 104 and other ethnicities).

#### 105 2.2.2. Clinical assessments at baseline.

Duration of untreated psychosis (DUP) was defined as the difference between the date of the appearance of the first positive psychotic symptom and the date of initiation of treatment with antipsychotics, in weeks (Norman and Malla, 2001).

The baseline diagnoses were made from interviews and mental health records utilizing the 109 Operational Criteria Checklist (McGuffin et al., 1991) and were grouped using ICD-10 into: 110 affective psychosis group (patients diagnosed with codes F30-33) or schizophrenia-spectrum 111 112 disorders group (ICD-10 codes F20-29) (Trotta et al., 2016). For those who did not meet 113 criteria based on ICD-10 we extracted the diagnosis from the same OPCRIT assessment 114 based on DSM-IV, grouping them into: affective psychosis (patients diagnosed with bipolar disorder, manic episode with psychosis or major depression with psychotic features - codes 115 296-296.9-) or schizophrenia-spectrum disorder (for schizophrenia, schizoaffective disorder, 116

delusional disorder and psychotic disorder NOS - codes 295.1-295.9 and 297.1-298.9-).

- 118 Symptomatology at baseline was rated on the Positive and Negative Syndrome Scale
- 119 (PANSS) (Kay, Fiszbein and Opler, 1987), from face-to-face interviews in the week
- 120 preceding the assessment. The 30 items that comprise the scale are divided into positive,
- 121 negative and general psychopathology scales. A confirmatory factor analyses (CFAs) was
- 122 conducted of the Wallwork/Fortgang five-factor model (Wallwork et al., 2012). This
- 123 Wallwork/Fortgang five-factor model has been shown to be the most robust PANSS factorial
- solution for exploring symptom profiles in FEP patients (Langeveld *et al.*, 2013) and has
- previously used in this sample (Ajnakina *et al.*, 2016, 2017). Global Assessment of
- 126 Functioning (GAF) was used to measure both overall symptoms severity and disability
- associated with the illness at the study entry (Endicott *et al.*, 1976). We complimented the
- 128 GAF with the Clinical General Impression scale (CGI) (Guy, 1976).
- Baseline General intelligence was assessed using a brief version of the Wechsler Adult Intelligence Scale—Third Edition (WAIS-III) (Wechsler, 1997) which includes a standardized
- 131 set of 5 tasks (Information, Block Design, Matrix Reasoning, Digit Symbol Coding and Digit
- 132 Span) to give a prorated intelligence quotient (IQ).

#### 133 2.2.3. Jumping to conclusions

For the measurement of the Jumping to Conclusions (JTC) bias participants had to complete 134 both versions of the probabilistic reasoning 'Beads' Task (Garety et al., 2005), with beads in 135 85:15 and 60:40 ratios. Participants are shown two jars containing coloured beads in opposite 136 ratios (e.g., in one of them the proportion was 85 black versus 15 orange and in the other the 137 ratio was reversed). Beads are then drawn from one of the two jars (randomly chosen by the 138 computer), one at a time. After each draw the participants can either make a guess about 139 which jar the beads come from or request a new bead. Participants will only have a single trial 140 in which they can ask as many beads as they need to make their final decision. The two main 141 variables classically extracted from the "Beads Task" are: a continuous variable defined as the 142 numbers of draws before making a decision, so-called "draws to decision" (DTD); and a 143 dichotomous variable (JTC/no JTC), in which JTC bias has been operationally defined as 144 reaching a decision after fewer than three beads (Garety et al., 2005). 145

146

In this study, dichotomous rating was preferred since the distribution of the DTD variable is 147 not normal, the colour of each drawn and the sequence could influence the next decision; and 148 there is better fit with clinical validity of the dichotomous approach, as we are mainly interested 149 in identifying the extreme responders from the rest. Moreover, the use of dichotomous scoring 150 seems to be superior in predicting change in delusion conviction (So et al., 2012). 151 Consequently, and based on previously published papers using different types of Beads 152 153 Tasks, we considered a hasty decision on any of the two versions of the task, to be evidence of the tendency to JTC (Garety et al., 2005; Ross et al., 2011; Jolley et al., 2014; So et al., 154 2014; So and Kwok, 2015). 155

156

# 157 2.3 Tracing patients at follow-up

Approximately 4 years (mean=4.0, S. D=0.13) after first contact with psychiatric services for 158 159 psychosis, we sought to trace all FEP cases included in the original GAP study with JTC 160 scores available at baseline and who had given consent for their clinical records to be accessed at follow-up. A thorough database search was carried out using the electronic 161 psychiatric records that are the primary clinical record keeping system within the SLaM Trust 162 (Electronic Patient Journey System (ePJS)). To trace those patients who dropped out from the 163 services we contacted their last known General Practitioners via mail seeking further 164 information about the patient's whereabouts and health; then patients themselves were 165 contacted wherever possible. All deaths and emigrations up to and including those that 166 occurred during the final year of follow-up were identified by a case-tracing procedure with the 167 Office for National Statistics for England and Wales and the General Register Office for 168 Scotland Further details are available in Ajnakina et al (Ajnakina et al., 2017). 169

# 170 **2.4 Data at follow up**

171 At follow-up, extensive information was extracted across clinical and social domains, and 172 patterns of care, from electronic psychiatric clinical records using the WHO Life Chart 173 Schedule (LCS) extended version (Sartorius *et al.*, 1996).

#### 174 2.4.1 Clinical outcomes

175 Remission was defined as an absence of overt psychotic symptoms for  $\geq$ 6 months similar to 176 earlier work conducted in the same geographical region (Morgan *et al.*, 2014; Revier *et al.*, 2015) and in line with operational criteria (Andreasen *et al.*, 2005) using information extracted
 from clinical records. This measure of remission was neither dependent on absence of non psychotic symptoms (e.g. depressed mood, neurotic manifestations), nor on whether patients
 were receiving treatment with antipsychotic medications during remission.

# 181 2.4.2 Service use

Utilising the LCS extended version (Sartorius et al., 1996) and excluding hospital admission 182 on first contact with mental health services for psychosis, we extracted detailed information 183 on circumstances of each re-admission including all compulsory admissions (i.e., admissions 184 exercised under MHA legislation) and instances when police was involved at the time of, or 185 shortly before, hospital admissions throughout the 4-year follow up period. Compulsory 186 admission under the MHA has been previously used for analysing pathways of care in similar 187 samples of the area of South London (Davies et al., 1996; Morgan et al., 2005b); and more 188 recently instances of police involvement has been also applied to measure pattern of care in 189 this same sample (Ajnakina et al., 2017). Using the admission and discharge dates for each 190 re-admission, we calculated the total length of inpatient stays in psychiatric wards during the 191 entire follow up period. 192

#### 193 **2.5 Analyses**

All analyses were conducted in STATA release 14 (STATACorp, 2015).

195 Descriptive statistics. The basic characteristics of the sample including socio-demographics (gender, age, ethnicity and educational level) and clinical information (DUP, diagnosis and 196 length of follow-up) were described using frequencies, percentages, mean and standard 197 deviations (SD), median and interguartile ranges (IQRs). The comparisons between FEP 198 groups based on the presence or not of the bias were made using chi-square, Student's t test 199 or Wilcoxon-Mann-Whitney tests when appropriate. Normality of all variables was assessed 200 computing Shapiro-Wilk normality test. . For the investigation of cross-sectional relationships 201 at follow-up between JTC and clinical outcome, we ran Wilcoxon-Mann-Whitney tests for "days 202 of hospitalization" as it is not normally distributed, and Chi-square for the three dichotomous 203 categorical variables: MHA intervention, police intervention and remission (yes/no). Effect 204 sizes were calculated for all the statistical tests using Cohen's d for t-test and Cramer's V ( $\Phi$ c) 205 for chi-square. When Mann-Whitney test is used, effect sizes from z values were calculated. 206

*Confirmatory Factor Analysis (CFA).* The detailed description of methods employed to conduct
CFA using this sample is available in Ajnakina et al. (Ajnakina *et al.*, 2016). Briefly, CFA was
conducted to evaluate the statistical fit (Stefanovics *et al.*, 2014) of the Wallwork/Fortgang fivefactor model of psychosis (Wallwork *et al.*, 2012) in this sample. This model includes positive
(P1, P3, P5, G9), negative (N1, N2, N3, N4, N6 and G7), excited (P4, P7, G8 and G14),
disorganised/concrete (P2, N5, G11), and depressed (G2, G3, G6) factors; which were used
as confounders for regression analyses.

Association analysis. The main hypothesis was tested by four regressions with the dichotomous rating of JTC as the dependant variable (JTC=1; no JTC=0). a Negative Binomial regression for the not normally distributed count dependent variable "days of hospitalization" as it was over dispersed; and logistic regression analyses for the binary variables (clinical remission, MHA intervention and police intervention). These regressions were calculated both unadjusted and adjusted for age, gender, ethnicity, IQ, symptom dimensions and functional level measured by GAF. In the regression model for predicting days of admission, we performed further adjustment of the model including the dichotomy variables of MHA and

222 police intervention, as they both could be confounders for longer admission.

# 223 3. RESULTS

Follow-up was successfully completed for 82.9% (n=102) of baseline patients, with a total of 224 17.1% lost (n=21). Of those 21, 2 (1.6%) had died, 6 (4.9%) had emigrated, 4 (3.3%) were 225 excluded as we did not have available information on follow-up and in 9 (7.3%) the attempt 226 at contact was unsuccessful. When comparing baseline sociodemographic characteristics of 227 228 those subjects lost at follow-up with those included, we did not find any significant difference in age, gender, educational level, ethnicity nor diagnosis (please refer to the Supplementary 229 Table 2). Nonetheless we found significant differences in baseline symptomatology, with 230 more severe symptoms in the lost subjects compared with those who completed follow-up 231 (positive subscale of PANSS 16.6 vs 13.52 respectively; t(113)=.21, p=0.03; GAF symptom 232 subscale 42.22 vs 52.8 respectively, t(105)=2.11, p=0.04). Hence, it is not possible to reject 233 234 the possibility of attrition bias.

# 235 **3.1 Patient characteristics at baseline**

Sociodemographic and clinical information of the baseline sample is shown in Table 1. The
mean age at first contact was 29.4 years (S.D.=10.03); n=75 (61%) of the sample were men,
38% were of white ethnicity and n=58 (49%) had tertiary education. Of the total 123 patients,
n=63 (51.2%) had shown JTC and n=92 (74.8%) had received a diagnosis of a schizophrenia
spectrum disorder.

#### 241

# [insert Table 1 here]

Table 2a shows the comparison of baseline sociodemographic and clinical characteristics 242 243 between patients with and without JTC bias (JTC/no JTC groups). We found no statistically 244 significant differences in age, gender, DUP, follow-up time, ethnicity or educational level. In 245 terms of lifetime substance use, there was no statistically significant differences regarding consumption of cannabis by JTC. As far as social functioning is concerned, no difference was 246 found in employment or marital status according to JTC, but there was a significant difference 247 in living arrangements, with a higher proportion of non-independent living in those presenting 248 the bias (46.8% vs 27.1%;  $X^2$  (1)= 5, p=0.03). 249

250

# [insert Table 2a here]

The baseline clinical status characteristics are presented in Table 2b. There was no statistically significant difference in diagnosis between JTC groups (p=0.72). To our surprise,, we did not find significant differences in scores on the Positive Symptoms subscale of the PANSS(14.78  $\pm 6.11$  vs 13.27  $\pm 5.38$ ; p=0.22). We found significant differences in IQ (mean of  $85.95\pm14.31$ for those presenting JTC and  $94.96\pm13.94$  for those without the bias; d=0.64 p=0.001). Functionality measured by the GAF scale was worse in patients with JTC (mean of  $55.38 \pm$ 16.7) than the patients without it (mean of  $61.92 \pm 16.63$ ; t=2.02 d=0.39 p=0.05).

258

# [insert Table 2b here]

259 **3.2 Clinical presentation over the follow up period** 

260 The comparison of clinical outcome domains during follow-up is presented in Table 3. Fifty-

eight percent (n=29 of 50) of the individuals with the bias and 64% (n=31 of 48) of those

without JTC achieved remission criteria in the long-term; this difference was not statistically

263 significant ( $\chi^2$ =0.45, df=1, p=0.5).

# 264 **3.3 Service use over the follow up period**

265 The comparison of service use variables during follow-up are also presented in Table 3.

In the follow-up period, on average, patients presenting JTC had more inpatient days (median=56, IQR=20-158), than those without the bias (median=15.5, IQR=0-93.5; U=-2.08, p=0.04). A higher proportion of patients with the JTC bias (N=27, 65.8%) were detained under the MHA than those without JTC (N=13, 37.1%;  $X^2$  (1)= 6.24; p= 0.01) The percentage subject to police involvement in compulsory admission during follow-up show a tendency to be higher in the patients with JTC [N=23 (56.1%) out of 41] than in those without the bias [12 (34.3%) out of 35], but this did not reach significance [X<sup>2</sup> (1)=3.62; p= 0.06].

273 [insert Table 3 here]

# **3.4 Predicting effects of JTC on long-term clinical outcome.**

275 Predictors effects of JTC on long-term clinical outcome are presented in Table 4.

Regression analyses showed that the presence of JTC predicted more inpatient days 276 277 (IRR=2.18, 95% CI 1.01-4.72, p=0.05) and more proneness to intervention under the mental health act (OR=5.9, 95% CI 2.04-17.05, p=0.001) and by the police (OR=4.18, 95% CI 1.46-278 11.98 p=0.008) in the moment of admission. After adjusting for age, gender, ethnicity, IQ and 279 280 symptoms and disability measured by GAF, the effect remained significant for days of hospitalization (adjusted IRR=5.03, 95% CI 1.91-13.24, p=0.001), use of the MHA (adjusted 281 OR=15.62, 95% CI 2.92-83.54, , p=0.001) and police involvement (adjusted OR=14.95, 95% 282 283 CI 2.68-83.34, p=0.002). There was no predictive effect on remission.

284

# [insert Table 4 here]

Additionally, all the OR with 95% confidence intervals and p values for all covariates of four

full regression models are presented in the Supplementary material (Supplementary Table 3).

# 288 4. DISCUSSION

The presence of JTC at baseline was associated with subsequent greater risk of compulsory admissions under the MHA and higher risk of police intervention at follow up, confirming our preliminary hypothesis. Why that should be is as yet unclear. One possible explanation is that JTC is related to behaviours such as impulsivity, beyond general cognitive and executive impairments, although evidence for this association has not been demonstrated conclusively (Moritz and Woodward, 2005; Rubio *et al.*, 2011; Lunt *et al.*, 2012); however, the studies claiming to look at this association have not employed a specific measure of impulsivity.

Another possible explanation for our findings could be a relationship between data-gathering bias and risk of aggressive behaviour or violence. The link between schizophrenia and violence has been previously studied (Fazel *et al.*, 2009), but it has been challenging to identify the processes underlying this association. Despite discrepancies in the literature, neurocognition seems to be one of the core risk factors for violence, mediated by several proximal and more direct risk factors(O'Reilly *et al.*, 2015). In addition, it has been shown that positive psychotic symptoms, including persecutory ideation, increase the risk of minor and serious violence (Swanson *et al.*, 2006). Given the relationship between JTC and the proneness and maintenance of delusion ideation, this could also explain part of the association of JTC and the higher need for police and MHA interventions in the moment of admission.

One mediating factor may be insight. Preliminary work on this cohort showed that JTC bias 306 and cognition were each associated with poorer recognition of illness (Wiffen et al., 2010). 307 Furthermore, the relationship between JTC and insight has been explored in schizophrenia 308 patients in forensic settings and it was found a direct correlation between the information 309 310 patients consider in making decisions and their clinical insight (Kuokkanen et al., 2016). Given that poor clinical insight was found to be a risk factor for violence, and following the same 311 rationale (Alia-Klein et al., 2007), lack of insight could be a possible additional explanation for 312 the need of police intervention in those presenting hasty-decision style; and indeed poor 313 insight is strongly associated with involuntary treatment in hospital (David et al., 1992). Thus, 314 insight could have had an impact in the association between JTC and the intervention of police 315 and MHA, so further studies focusing on the relationship between JTC and insight should be 316 encouraged. Nonetheless, the interpretation of the need for police intervention as a proxy for 317 social or behavioural disruption should be cautious. Previous work from the same area found 318 319 that the higher police involvement at time of admission in the African Caribbean group was 320 explained by the greater likelihood of such families to seek help from police themselves in 321 these situations than White British families (Morgan et al., 2005b).

The other effect predicted by JTC was number of days of hospitalization. This outcome is one of the most employed markers of service use in order to study the clinical course of psychosis (Johnstone *et al.*, 1990; Geddes *et al.*, 1994; Takei *et al.*, 1998; Lehtinen *et al.*, 2000; Cahn *et al.*, 2002; Möller *et al.*, 2002; Thara, 2004; Nordentoft *et al.*, 2010; Heslin *et al.*, 2016). As JTC is thought to be critical in delusion formation by contributing to erroneous interferences (Garety, Hemsley and Wessely, 1991), it is reasonable to expect an effect on inpatient length of stay.

Contrary to our hypothesis, an association between JTC and remission rates at follow-up was 329 not significant. As noted above, JTC is linked to with the persistence of delusion ideation 330 331 (Falcone et al., 2015) and hence less improvement in delusions over time (So et al., 2014). As our remission outcome was based on Remission in Schizophrenia Working Group (RSWG) 332 (Andreasen et al., 2005) which includes delusions, we expected that the presence of 333 334 reasoning bias at baseline would have been inversely related to the achievement of remission at follow-up. One of the reasons for the lack of effect could have been a ceiling effect for our 335 sample, as reported rates of RSWG remission range between 48-61% for first-episode 336 337 patients (Lambert et al., 2010); and in our patients, the percentages of remission were above that range. 338

The relationship between the presence of JTC and cognitive impairments has been widely studied in both schizophrenia patients and healthy participants, especially with working memory (Garety *et al.*, 2013), executive functions (Woodward *et al.*, 2009; Rubio *et al.*, 2011), verbal memory (Keefe, Eesley and Poe, 2005; Lee and Park, 2005; Forbes *et al.*, 2009; Fatouros-Bergman *et al.*, 2014), and cognitive processing speed (Ochoa *et al.*, 2014). The potential effect of neurocognition on clinical outcome is well known (Lam, Raine and Lee, 345 2014). In our study, we found a negative association between IQ and JTC, but the associations 346 in the regressions models remain statistically significant when IQ was added as confounder in 347 the adjusted model. Nonetheless, the evidence supporting the association between some 348 neurocognitive domains and JTC talks in favour of encouraging new studies with bigger 349 samples to include not only IQ but also other cognitive measures as confounder factors in the 350 association of JTC and clinical outcome.

Our evidence that the presence of JTC appears to be linked to poor outcome variables makes 351 it a potential target for therapies aiming to improve the prognosis of the illness. There is a body 352 of literature showing that changes in JTC are directly linked with changes in symptomatology 353 (Dudley et al., 2013; Sanford et al., 2013; Garety et al., 2015). Furthermore, psychotherapeutic 354 355 interventions have been developed for overcoming biases in metacognition, including JTC, which have been proven to be effective in reducing the tendency to make hasty decisions and 356 in improving outcome for people with psychosis, such as the Maudsley Review Training 357 Programme (Waller et al., 2011), Metacognitive Training (MCT) (Moritz et al., 2014a, b) and 358 Social Cognition and Interaction Training (SCIT)(Roberts et al., 2014). Our study raises the 359 question of whether the implementation of these specific interventions to reduce Jumping to 360 conclusions may be a useful addition in early psychosis intervention programs. 361

# 362 Limitations and Strengths

Among the main limitations in our study are the lack of inclusion of more neurocognitive 363 measures and the small sample size, which limited the inclusion of more confounders in the 364 regression models. Nonetheless, a general IQ measure was used as confounder. Another 365 limitation is the lack of information regarding co-morbid diagnosis at baseline and follow-up 366 that may have had an impact in clinical outcome. A potential methodological limitation is the 367 368 information bias arising from loss and missing data at follow-up. Although we completed follow-369 up for 82.9% of patients, we found statistically significant differences in symptomatology 370 between those who completed the follow-up and those who were lost. Specifically, those lost participants had higher scores in the subscale of positive symptoms of PANSS. Our findings 371 may therefore not be generalizable to the more severely affected patients. Lastly, it should be 372 noted that by only performing one trial per version, as widely used in JTC beads tasks, may 373 allow results to be affected by miscomprehension (Balzan et al., 2012), with a tendency to 374 overestimate the presence of JTC. 375

On the other hand, one of the strengths of our study is the inclusion in the regression analyses of ethnicity and other important socio-demographic confounders. It has been shown in previous works with a sample from the same area that there exists a higher risk of compulsory admission and longer admission among African–Caribbean and Black African patients (Morgan *et al.*, 2005*a*; Ajnakina *et al.*, 2017), but a strong association between JTC and compulsory admission remained in our study after adjusting by ethnicity.

# 382 Conclusions

JTC is a data-gathering bias that has been consistently proved to be associated to psychotic patients. Our study found that its presence at FEP is associated with worse clinical outcome, reflected by more days of admissions, greater need for compulsory hospitalization and police intervention. This raises the question of whether more efforts should be devoted to developing and applying therapies focusing on JTC as possible target.

#### **BIBLIOGRAPHY**

Aghotor, J, Pfueller, U, Moritz, S, Weisbrod, M, Roesch-Ely, D (2010). Metacognitive training for patients with schizophrenia (MCT): Feasibility and preliminary evidence for its efficacy. *Journal of Behavior Therapy and Experimental Psychiatry* **41**, 207–211.

391 Ajnakina, O, Trotta, A, Oakley-Hannibal, E, Di Forti, M, Stilo, SA, Kolliakou, A, Gardner-

392 Sood, P, Gaughran, F, David, AS, Dazzan, P, Pariante, C, Mondelli, V, Morgan, C,

**Vassos, E, Murray, RM, Fisher, HL** (2016). Impact of childhood adversities on specific

394 symptom dimensions in first-episode psychosis. *Psychological Medicine* **46**, 317–326.

Ajnakina, O, Lally, J, Di Forti, M, Kolliakou, A, Gardner-Sood, P, Lopez-Morinigo, J,

- Dazzan, P, Pariante, CM, Mondelli, V, MacCabe, J, David, AS, Gaughran, F, Murray, RM,
   Vassos, E (2017). Patterns of illness and care over the 5 years following onset of psychosis
   in different ethnic groups; the GAP-5 study. *Social psychiatry and psychiatric epidemiology* 52, 1101–1111.
- Alia-Klein, N, O'Rourke, TM, Goldstein, RZ, Malaspina, D (2007). Insight into illness and
   adherence to psychotropic medications are separately associated with violence severity in a
   forensic sample. Aggressive Behavior 33, 86–96.
- 403 American Psychiatric Association (2000). *Diagnostic and statistical manual of mental* 404 *disorders : DSM-IV.* 4th ed. American Psychiatric Publishing: Washington, DC .
- Andreasen, NC, Carpenter, WT, Kane, JM, Lasser, RA, Marder, SR, Weinberger, DR
   (2005). Remission in Schizophrenia: Proposed Criteria and Rationale for Consensus.
   American Journal of Psychiatry 162, 441–449.
- Andreou, C, Treszl, A, Roesch-Ely, D, Köther, U, Veckenstedt, R, Moritz, S (2014).
   Investigation of the role of the jumping-to-conclusions bias for short-term functional outcome
   in schizophrenia. *Psychiatry research* 218, 341–7.
- Andreou, C, Schneider, BC, Balzan, R, Luedecke, D, Roesch-Ely, D, Moritz, S (2015).
  Neurocognitive deficits are relevant for the jumping-to-conclusions bias, but not for
  delusions: A longitudinal study. *Schizophrenia research. Cognition* 2, 8–11.
- Balzan, RP, Delfabbro, PH, Galletly, CA, Woodward, TS (2012). Over-adjustment or
  miscomprehension? A re-examination of the jumping to conclusions bias. *Australian & New*Zealand Journal of Psychiatry 46, 532–540.
- Breier, A, Schreiber, JL, Dyer, J, Pickar, D (1991). National Institute of Mental Health
  longitudinal study of chronic schizophrenia. Prognosis and predictors of outcome. Archives
  of general psychiatry 48, 239–46.
- 420 Cahn, W, Hulshoff Pol, HE, Lems, EBTE, van Haren, NEM, Schnack, HG, van der
  421 Linden, JA, Schothorst, PF, van Engeland, H, Kahn, RS (2002). Brain volume changes in
  422 first-episode schizophrenia: a 1-year follow-up study. *Archives of general psychiatry* 59,
  423 1002–10.
- 424 **Couture, SM**, **Penn, DL**, **Roberts, DL** (2006). The functional significance of social cognition 425 in schizophrenia: a review. *Schizophrenia bulletin* **32**, S44-63.
- 426 Van Dael, F, Versmissen, D, Janssen, I, Myin-Germeys, I, van Os, J, Krabbendam, L
- 427 (2006). Data Gathering: Biased in Psychosis?. *Schizophrenia Bulletin* **32**, 341–351.

- 428 **David, A, Buchanan, A, Reed, A, Almeida, O** (1992). The assessment of insight in 429 psychosis. *The British Journal of Psychiatry* **161**.
- 430 Davies, S, Thornicroft, G, Leese, M, Higgingbotham, A, Phelan, M (1996). Ethnic
   431 differences in risk of compulsory psychiatric admission among representative cases of
   432 psychosis in London. *British Medical Journal (Clinical research ed.)* 312, 533–7.
- 433 Dudley, R, Daley, K, Nicholson, M, Shaftoe, D, Spencer, H, Cavanagh, K, Freeston, M
  434 (2013). 'Jumping to conclusions' in first-episode psychosis: A longitudinal study. *British*435 *Journal of Clinical Psychology* 52, 380–393.
- 436 Dudley, R, Taylor, P, Wickham, S, Hutton, P (2016). Psychosis, Delusions and the
  437 'Jumping to Conclusions' Reasoning Bias: A Systematic Review and Meta-analysis.
  438 Schizophrenia Bulletin 42, 652–665.
- 439 Endicott, J, Spitzer, RL, Fleiss, JL, Cohen, J (1976). The global assessment scale. A
   440 procedure for measuring overall severity of psychiatric disturbance. *Archives of general* 441 *psychiatry* 33, 766–71.
- 442 Faber, G, Smid, HGOM, Van Gool, AR, Wunderink, L, Wiersma, D, van den Bosch, RJ
- (2011). Neurocognition and recovery in first episode psychosis. *Psychiatry research* 188, 1–
  6.
- Falcone, MA, Murray, RM, Wiffen, BDR, O'Connor, JA, Russo, M, Kolliakou, A, Stilo, S,
   Taylor, H, Gardner-Sood, P, Paparelli, A, Jichi, F, Di Forti, M, David, AS, Freeman, D,
- Jolley, S (2015). Jumping to conclusions, neuropsychological functioning, and delusional
   beliefs in first episode psychosis. *Schizophrenia bulletin* 41, 411–8.
- 449 Falcone, MA, Murray, RM, O'Connor, JA, Hockey, LN, Gardner-Sood, P, Di Forti, M,
- Freeman, D, Jolley, S (2015). Jumping to conclusions and the persistence of delusional beliefs in first episode psychosis. *Schizophrenia research* **165**, 243–6.
- Fatouros-Bergman, H, Cervenka, S, Flyckt, L, Edman, G, Farde, L (2014). Meta-analysis
  of cognitive performance in drug-naïve patients with schizophrenia. *Schizophrenia Research*158, 156–162.
- Favrod, J, Rexhaj, S, Bardy, S, Ferrari, P, Hayoz, C, Moritz, S, Conus, P, Bonsack, C
  (2014). Sustained antipsychotic effect of metacognitive training in psychosis: A randomizedcontrolled study. *European Psychiatry* 29, 275–281.
- 458 **Fazel, S**, **Gulati, G**, **Linsell, L**, **Geddes, JR**, **Grann, M** (2009). Schizophrenia and Violence: 459 Systematic Review and Meta-Analysis. *PLoS Medicine* **6**, e1000120.
- 460 Fett, A-KJ, Viechtbauer, W, Dominguez, M-G, Penn, DL, van Os, J, Krabbendam, L
- 461 (2011). The relationship between neurocognition and social cognition with functional
  462 outcomes in schizophrenia: a meta-analysis. *Neuroscience and biobehavioral reviews* 35,
  463 573–88.
- Forbes, NF, Carrick, LA, McIntosh, AM, Lawrie, SM (2009). Working memory in
   schizophrenia: a meta-analysis. *Psychological medicine* 39, 889–905.

466 Di Forti, M, Morgan, C, Dazzan, P, Pariante, C, Mondelli, V, Marques, TR, Handley, R,
467 Luzi, S, Russo, M, Paparelli, A, Butt, A, Stilo, SA, Wiffen, B, Powell, J, Murray, RM
468 (2009). High-potency cannabis and the risk of psychosis. *The British journal of psychiatry :*469 the journal of mental science 195, 488–91.

Di Forti, M, Marconi, A, Carra, E, Fraietta, S, Trotta, A, Bonomo, M, Bianconi, F,
 Gardner-Sood, P, O'Connor, J, Russo, M, Stilo, SA, Margues, TR, Mondelli, V, Dazzan,

- 472 P, Pariante, C, David, AS, Gaughran, F, Atakan, Z, Iyegbe, C, et al. (2015). Proportion of
- patients in south London with first-episode psychosis attributable to use of high potency
- 474 cannabis: a case-control study. *The lancet. Psychiatry* **2**, 233–8.
- Freeman, D, Garety, P (2014). Advances in understanding and treating persecutory
  delusions: a review. Social psychiatry and psychiatric epidemiology 49, 1179–89.
- Freeman, D, Pugh, K, Garety, P (2008). Jumping to conclusions and paranoid ideation in
  the general population. *Schizophrenia Research* 102, 254–260.
- 479 Friis, S, Melle, I, Johannessen, JO, Røssberg, JI, Barder, HE, Evensen, JH, Haahr, U,
- Ten Velden Hegelstad, W, Joa, I, Langeveld, J, Larsen, TK, Opjordsmoen, S, Rund, BR,
   Simonsen, E, Vaglum, PW, McGlashan, TH (2016). Early Predictors of Ten-Year Course in
   First-Episode Psychosis. *Psychiatric services (Washington, D.C.)* 67, 438–43.
- 483 Garety, P, Waller, H, Emsley, R, Jolley, S, Kuipers, E, Bebbington, P, Dunn, G, Fowler,
- 484 **D**, **Hardy**, **A**, **Freeman**, **D** (2015). Cognitive Mechanisms of Change in Delusions: An 485 Experimental Investigation Targeting Reasoning to Effect Change in Paranoia.
- 485 Experimental investigation rargeting Reasoning to Effect Cha 486 Schizophrenia Bulletin **41**, 400–410.
- 487 **Garety, PA**, **Kuipers, E**, **Fowler, D**, **Freeman, D**, **Bebbington, PE** (2001). A cognitive model 488 of the positive symptoms of psychosis. *Psychological Medicine* **31**, 189–195.
- 489 Garety, PA, Freeman, D, Jolley, S, Dunn, G, Bebbington, PE, Fowler, DG, Kuipers, E,
- 490 **Dudley, R** (2005). Reasoning, Emotions, and Delusional Conviction in Psychosis. *Journal of* 491 *Abnormal Psychology* **114**, 373–384.
- 492 Garety, PA, Gittins, M, Jolley, S, Bebbington, P, Dunn, G, Kuipers, E, Fowler, D,
- Freeman, D (2013). Differences in cognitive and emotional processes between persecutory and grandiose delusions. *Schizophrenia bulletin* **39**, 629–39.
- 495 **Garety, PA**, **Freeman, D** (1999). Cognitive approaches to delusions: a critical review of 496 theories and evidence. *The British journal of clinical psychology* **38**, 113–54.
- 497 **Garety, PA**, **Hemsley, DR**, **Wessely, S** (1991). Reasoning in deluded schizophrenic and 498 paranoid patients. Biases in performance on a probabilistic inference task. *The Journal of* 499 *nervous and mental disease* **179**, 194–201.
- Geddes, J, Mercer, G, Frith, CD, MacMillan, F, Owens, DG, Johnstone, EC (1994).
   Prediction of outcome following a first episode of schizophrenia. A follow-up study of
- Prediction of outcome following a first episode of schizophrenia. A follow-up study of
   Northwick Park first episode study subjects. *The British journal of psychiatry : the journal of mental science* 165, 664–8.
- González, LE, López-Carrilero, R, Barrigón, ML, Grasa, E, Barajas, A, Pousa, E,
   González-Higueras, F, Ruiz-Delgado, I, Cid, J, Lorente-Rovira, E, Pélaez, T, Ochoa, S,
   Ochoa, S (2017). Neuropsychological functioning and jumping to conclusions in recent
   onset psychosis patients. *Schizophrenia Research* 195, 366-371.
- Green, MF, Kern, RS, Braff, DL, Mintz, J (2000). Neurocognitive deficits and functional
  outcome in schizophrenia: are we measuring the "right stuff"?. *Schizophrenia bulletin* 26,
  119–36.
- 511 **Guy, W** (1976). *ECDEU assessment manual for psychopharmacology*. U. Department of 512 Health, and Welfare, 534-537
- 513 Heslin, M, Lappin, JM, Donoghue, K, Lomas, B, Reininghaus, U, Onyejiaka, A,
- 514 Croudace, T, Jones, PB, Murray, RM, Fearon, P, Doody, GA, Dazzan, P, Craig, TJ,
- 515 Morgan, C (2016). Ten-year outcomes in first episode psychotic major depression patients

- 516 compared with schizophrenia and bipolar patients. *Schizophrenia research* **176**, 417–22.
- Ho, B-C, Nopoulos, P, Flaum, M, Arndt, S, Andreasen, NC (1998). Two-Year Outcome in
  First-Episode Schizophrenia: Predictive Value of Symptoms for Quality of Life. *American Journal of Psychiatry* 155, 1196–1201.

van Hooren, S, Versmissen, D, Janssen, I, Myin-Germeys, I, à Campo, J, Mengelers, R,

- van Os, J, Krabbendam, L (2008). Social cognition and neurocognition as independent
   domains in psychosis. *Schizophrenia research* 103, 257–65.
- Hopper, K, Harrison, G, Janca, A, Sartorius, N (2007). *Recovery from Schizophrenia an International Perspective.* Oxford University Press: New York, NY..
- 525 Johnstone, EC, Macmillan, JF, Frith, CD, Benn, DK, Crow, TJ (1990). Further
- 526 investigation of the predictors of outcome following first schizophrenic episodes. *The British* 527 *journal of psychiatry : the journal of mental science* **157**, 182–9.
- Jolley, S, Thompson, C, Hurley, J, Medin, E, Butler, L, Bebbington, P, Dunn, G,

Freeman, D, Fowler, D, Kuipers, E, Garety, P (2014). Jumping to the wrong conclusions?
 An investigation of the mechanisms of reasoning errors in delusions. *Psychiatry research* 219, 275–82.

- Juola, P, Miettunen, J, Veijola, J, Isohanni, M, Jääskeläinen, E (2013). Predictors of
- short- and long-term clinical outcome in schizophrenic psychosis--the Northern Finland 1966
- 534 Birth Cohort study. *European psychiatry : the journal of the Association of European* 535 *Psychiatrists* **28**, 263–8.
- 536 **Kay, SR**, **Fiszbein, A**, **Opler, LA** (1987). The positive and negative syndrome scale 537 (PANSS) for schizophrenia., *Schizophrenia bulletin* **13**, 261–76.
- 538 **Keefe, RSE**, **Eesley, CE**, **Poe, MP** (2005). Defining a cognitive function decrement in 539 schizophrenia. *Biological Psychiatry* **57**, 688–691.
- Kuokkanen, R, Lappalainen, R, Repo-Tiihonen, E, Tiihonen, J, Aho-Mustonen, K
  (2016). Cognitive Insight, Clinical Insight, and Reasoning in Schizophrenia: A Pilot Study in a
  Forensic Setting. *Journal of Forensic Psychology Practice* 16, 253–267.
- 543 **Lam, BYH**, **Raine, A**, **Lee, TMC** (2014). The relationship between neurocognition and 544 symptomatology in people with schizophrenia: social cognition as the mediator. *BMC* 545 *psychiatry* **14**, 138.
- Lambert, M, Karow, A, Leucht, S, Schimmelmann, BG, Naber, D (2010). Remission in schizophrenia: validity, frequency, predictors, and patients' perspective 5 years later.
- 548 *Dialogues in clinical neuroscience* **12**, 393–407.
- Langeveld, J, Andreassen, OA, Auestad, B, Faerden, A, Hauge, LJ, Joa, I,
- Johannessen, JO, Melle, I, Rund, BR, Røssberg, JI, Simonsen, E, Vaglum, P, Larsen,
- **TK** (2013). Is there an optimal factor structure of the Positive and Negative Syndrome Scale
- in patients with first-episode psychosis?. *Scandinavian Journal of Psychology* **54**, 160–165.
- Lee, J, Park, S (2005). Working Memory Impairments in Schizophrenia: A Meta-Analysis.
   Journal of Abnormal Psychology 114, 599–611.
- 555 **Lehtinen, V, Aaltonen, J, Koffert, T, Räkköläinen, V, Syvälahti, E** (2000). Two-year 556 outcome in first-episode psychosis treated according to an integrated model. Is immediate
- neuroleptisation always needed?. European psychiatry : the journal of the Association of
- 558 *European Psychiatrists* **15**, 312–20.

- 559 Lipkovich, IA, Deberdt, W, Csernansky, JG, Sabbe, B, Keefe, RS, Kollack-Walker, S.
- (2009). Relationships among neurocognition, symptoms and functioning in patients with
   schizophrenia: a path-analytic approach for associations at baseline and following 24 weeks
- of antipsychotic drug therapy. *BMC Psychiatry* **9**, 44.
- 563 Lysaker, PH, Carcione, A, Dimaggio, G, Johannesen, JK, Nicolo, G, Procacci, M,
- Semerari, A (2005). Metacognition amidst narratives of self and illness in schizophrenia:
   associations with neurocognition, symptoms, insight and quality of life. *Acta Psychiatrica Scandinavica* 112, 64–71.
- 567 Lysaker, PH, Gumley, A, Luedtke, B, Buck, KD, Ringer, JM, Olesek, K, Kukla, M,
- Leonhardt, BL, Popolo, R, Dimaggio, G (2013). Social cognition and metacognition in
   schizophrenia: evidence of their independence and linkage with outcomes. *Acta Psychiatrica Scandinavica* 127, 239–247.
- 571 **Mallett, R, Leff, J, Bhugra, D, Pang, D, Zhao, JH** (2002). Social environment, ethnicity and schizophrenia. *Social Psychiatry and Psychiatric Epidemiology* **37**, 329–335.
- 573 **McGuffin, P, Farmer, A, Harvey, I.** (1991). A Polydiagnostic Application of Operational 574 Criteria in Studies of Psychotic Illness. *Archives of General Psychiatry* **48**, 764.
- McLean, BF, Mattiske, JK, Balzan, RP (2016). Association of the Jumping to Conclusions
   and Evidence Integration Biases With Delusions in Psychosis: A Detailed Meta-analysis.
   Schizophrenia bulletin 43, 344-354.
- Möller, H-J, Bottlender, R, Gross, A, Hoff, P, Wittmann, J, Wegner, U, Strauss, A (2002).
   The Kraepelinian dichotomy: preliminary results of a 15-year follow-up study on functional
   psychoses: focus on negative symptoms. *Schizophrenia research* 56, 87–94.
- Morgan, C, Mallett, R, Hutchinson, G, Bagalkote, H, Morgan, K, Fearon, P, Dazzan, P,
  Boydell, J, McKenzie, K, Harrison, G, Murray, R, Jones, P, Craig, T, Leff, J, AESOP
  Study Group (2005a). Pathways to care and ethnicity. 1: Sample characteristics and
  compulsory admission. *British Journal of Psychiatry* 186, 281–289.
- Morgan, C, Mallett, R, Hutchinson, G, Bagalkote, H, Morgan, K, Fearon, P, Dazzan, P,
  Boydell, J, McKenzie, K, Harrison, G, Murray, R, Jones, P, Craig, T, Leff, J,AESOP
  Study Group (2005b). Pathways to care and ethnicity. 2: Source of referral and helpseeking. British Journal of Psychiatry **186**, 200, 296
- seeking. *British Journal of Psychiatry* **186**, 290–296.
- 589 Morgan, C, Lappin, J, Heslin, M, Donoghue, K, Lomas, B, Reininghaus, U, Onyejiaka,
- A, Croudace, T, Jones, PB, Murray, RM, Fearon, P, Doody, GA, Dazzan, P (2014).
   Reappraising the long-term course and outcome of psychotic disorders: the AESOP-10
- 592 study. *Psychological medicine* **44**, 2713–26.
- Moritz, S, Kerstan, A, Veckenstedt, R, Randjbar, S, Vitzthum, F, Schmidt, C, Heise, M,
   Woodward, TS (2011). Further evidence for the efficacy of a metacognitive group training in
   schizophrenia. *Behaviour Research and Therapy* 49, 151–157.
- 596 Moritz, S, Veckenstedt, R, Bohn, F, Hottenrott, B, Scheu, F, Randjbar, S, Aghotor, J, 597 Köther, U, Woodward, TS, Treszl, A, Andreou, C, Pfueller, U, Roesch-Ely, D (2013).
- 598 Complementary group Metacognitive Training (MCT) reduces delusional ideation in 599 schizophrenia. *Schizophrenia Research* **151**, 61–69.
- Moritz, S, Andreou, C, Schneider, BC, Wittekind, CE, Menon, M, Balzan, RP,
- 601 **Woodward, TS** (2014). Sowing the seeds of doubt: a narrative review on metacognitive 602 training in schizophrenia. *Clinical psychology review* **34**, 358–66.
- 603 Moritz, S, Veckenstedt, R, Andreou, C, Bohn, F, Hottenrott, B, Leighton, L, Köther, U,

- 604 Woodward, TS, Treszl, A, Menon, M, Schneider, BC, Pfueller, U, Roesch-Ely, D (2014).
- 505 Sustained and 'Sleeper' Effects of Group Metacognitive Training for Schizophrenia. *JAMA* 506 *Psychiatry* **71**, 1103.
- 607 **Moritz, S**, **Woodward, TS** (2005). Jumping to conclusions in delusional and non-delusional schizophrenic patients. *The British journal of clinical psychology* **44**, 193–207.
- Moritz, S, Woodward, TS (2007). Metacognitive training in schizophrenia: from basic
   research to knowledge translation and intervention. *Current Opinion in Psychiatry* 20, 619–
   625.
- 612 Mortimer, A, McKay, A., Quemada, I, Clare, L, Eastwood, N, McKenna, P. (1996).
- 613 Delusions in Schizophrenia: A Phenomenological and Psychological Exploration. *Cognitive* 614 *Neuropsychiatry* **1**, 289–304.
- 615 Nordentoft, M, Øhlenschlæger, J, Thorup, A, Petersen, L, Jeppesen, P, Bertelsen, M
- 616 (2010). Deinstitutionalization revisited: a 5-year follow-up of a randomized clinical trial of
- 617 hospital-based rehabilitation versus specialized assertive intervention (OPUS) versus
- 618 standard treatment for patients with first-episode schizophrenia spectrum disorder.
- 619 *Psychological medicine* **40**, 1619–1626.
- Norman, RM, Malla, AK (2001). Duration of untreated psychosis: a critical examination of the concept and its importance. *Psychological medicine* **31**, 381–400.
- O'Reilly, K, Donohoe, G, Coyle, C, O'Sullivan, D, Rowe, A, Losty, M, McDonagh, T,
   McGuinness, L, Ennis, Y, Watts, E, Brennan, L, Owens, E, Davoren, M, Mullaney, R,
   Abidin, Z, Kennedy, HG (2015). Prospective cohort study of the relationship between
   neuro-cognition, social cognition and violence in forensic patients with schizophrenia and
   schizoaffective disorder. *BMC psychiatry* 15, 155.
- Ochoa, S, Haro, JM, Huerta-Ramos, E, Cuevas-Esteban, J, Stephan-Otto, C, Usall, J,
   Nieto, L, Brebion, G (2014). Relation between jumping to conclusions and cognitive
   functioning in people with schizophrenia in contrast with healthy participants. *Schizophrenia research* 159, 211–7.
- Pinkham, AE (2014). Social cognition in schizophrenia. *The Journal of clinical psychiatry* 75, 14–9.
- 633 Revier, CJ, Reininghaus, U, Dutta, R, Fearon, P, Murray, RM, Doody, GA, Croudace, T,
- Dazzan, P, Heslin, M, Onyejiaka, A, Kravariti, E, Lappin, J, Lomas, B, Kirkbride, JB,
- **Donoghue, K**, **Morgan, C**, **Jones, PB** (2015). Ten-Year Outcomes of First-Episode
- Psychoses in the MRC ÆSOP-10 Study. *The Journal of nervous and mental disease* 203,
  379–86.
- Roberts, DL, Combs, DR, Willoughby, M, Mintz, J, Gibson, C, Rupp, B, Penn, DL
  (2014). A randomized, controlled trial of Social Cognition and Interaction Training (SCIT) for
  outpatients with schizophrenia spectrum disorders. *The British journal of clinical psychology*53, 281–98.
- Ross, K, Freeman, D, Dunn, G, Garety, P (2011). A randomized experimental investigation
   of reasoning training for people with delusions. *Schizophrenia bulletin* 37, 324–33.
- Rubio, JL, Ruiz-Veguilla, M, Hernández, L, Barrigón, ML, Salcedo, MD, Moreno, JM,
- 645 **Gómez, E**, **Moritz, Š**, **Ferrín, M** (2011). Jumping to conclusions in psychosis: A faulty 646 appraisal. *Schizophrenia research* **133**, 199–204.
- 647 **Sanford, N**, **Lecomte, T**, **Leclerc, C**, **Wykes, T**, **Woodward, TS** (2013). Change in jumping 648 to conclusions linked to change in delusions in early psychosis. *Schizophrenia Research*

649 207–208.

Sartorius, N, Gulbinat, W, Harrison, G, Laska, E, Siegel, C (1996). Long-term follow-up of
 schizophrenia in 16 countries. A description of the International Study of Schizophrenia
 conducted by the World Health Organization. Social psychiatry and psychiatric epidemiology
 31, 249–58.

So, SH, Freeman, D, Dunn, G, Kapur, S, Kuipers, E, Bebbington, P, Fowler, D, Garety,
 PA (2012). Jumping to conclusions, a lack of belief flexibility and delusional conviction in
 psychosis: A longitudinal investigation of the structure, frequency, and relatedness of
 reasoning biases. *Journal of Abnormal Psychology* 121, 129–139.

- So, SH, Peters, ER, Swendsen, J, Garety, PA, Kapur, S (2014). Changes in delusions in
   the early phase of antipsychotic treatment An experience sampling study., *Psychiatry Research* 215, 568–573.
- So, SH, Kwok, NT (2015). Jumping to conclusions style along the continuum of delusions:
   delusion-prone individuals are not hastier in decision making than healthy individuals. *PloS* one Public Library of Science10, e0121347.
- 664 **STATACorp** (2015). Stata Statistical Software: Release 14. College Station, TX: StataCorp 665 LP.
- 666 Stefanovics, EA, Elkis, H, Zhening, L, Zhang, XY, Rosenheck, RA (2014). A cross-
- national factor analytic comparison of three models of PANSS symptoms in schizophrenia.
   *Psychiatry Research* 219, 283–289.
- 669 Swanson, JW, Swartz, MS, Van Dorn, RA, Elbogen, EB, Wagner, HR, Rosenheck, RA,
- 670 **Stroup, TS**, **McEvoy, JP**, **Lieberman, JA** (2006). A National Study of Violent Behavior in 671 Persons With Schizophrenia. *Archives of General Psychiatry* **63**, 490.
- Takei, N, Persaud, R, Woodruff, P, Brockington, I, Murray, RM (1998). First episodes of
   psychosis in Afro-Caribbean and White people. An 18-year follow-up population-based study.
   The British journal of psychiatry : the journal of mental science 172, 147–53.
- 675 **Thara, R** (2004). Twenty-year course of schizophrenia: the Madras Longitudinal Study., 676 *Canadian journal of psychiatry. Revue canadienne de psychiatrie* **49**, 564–9.
- 677 Trotta, A, lyegbe, C, Di Forti, M, Sham, PC, Campbell, DD, Cherny, SS, Mondelli, V,
- Aitchison, KJ, Murray, RM, Vassos, E, Fisher, HL. (2016). Interplay between
- Schizophrenia Polygenic Risk Score and Childhood Adversity in First-Presentation Psychotic
   Disorder: A Pilot Study. *PLOS ONE* **11**, e0163319.
- Waller, H, Freeman, D, Jolley, S, Dunn, G, Garety, P (2011). Targeting reasoning biases in
  delusions: a pilot study of the Maudsley Review Training Programme for individuals with
  persistent, high conviction delusions. *Journal of behavior therapy and experimental psychiatry* 42, 414–21.
- 685 Wallwork, RS, Fortgang, R, Hashimoto, R, Weinberger, DR, Dickinson, D (2012).
- 686 Searching for a consensus five-factor model of the Positive and Negative Syndrome Scale 687 for schizophrenia. *Schizophrenia Research* **137**, 246–250.
- 688 Wechsler, D (1997). Wechsler Adult Intelligence Scale® Third Edition.
- 689 White, C, Stirling, J, Hopkins, R, Morris, J, Montague, L, Tantam, D, Lewis, S (2009). 690 Predictors of 10-year outcome of first-episode psychosis. *Psychological medicine* **39**, 1447–
- 690 Predictors of 10-year outcome of first-episode psychosis. *Psychological medicine* **39**, 1447–
   691 56.

- 692 **WHO** (1992). *The ICD-10 Classification of Mental and Behavioural Disorders: Clinical* 693 *descriptions and diagnostic guidelines*Geneva.
- 694 **WHO** (1994). Schedules for Clinical Assessment in Neuropsychiatry: Version 2, Manual.
- Wieselgren, IM, Lindström, E, Lindström, LH (1996). Symptoms at index admission as
   predictor for 1-5 year outcome in schizophrenia. *Acta Psychiatrica Scandinavica* 94, 311–
   319.
- Wiffen, B, Falcone, MA, O'Connor, J, Russo, M, Murray, R, David, A (2010). Bias or
   deficit? Differences in IQ and insight in first episode psychosis patients who demonstrate a
   'jumping to conclusions' response to the beads task. *Schizophrenia Research* 117, 446.
- Woodward, TS, Mizrahi, R, Menon, M, Christensen, BK (2009). Correspondences
   between theory of mind, jumping to conclusions, neuropsychological measures and the
- symptoms of schizophrenia. *Psychiatry Research* **170**, 119–123.