

**IMPROVED SOCIAL FUNCTIONING FOLLOWING SOCIAL
RECOVERY THERAPY IN FIRST EPISODE PSYCHOSIS: DO SOCIAL
COGNITION AND NEUROCOGNITION IMPROVE FOLLOWING
THERAPY, AND DO THEY PREDICT TREATMENT RESPONSE?**

Running Head: Cognition and functional improvement in psychosis

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Abstract

There is a need to develop and refine psychosocial interventions to improve functioning in First Episode Psychosis (FEP). Social cognition and neurocognition are closely linked to functioning in psychosis; examinations of cognition pre- and post- psychosocial intervention may provide insights into the mechanisms of these interventions, and identify which individuals are most likely to benefit.

Method: Cognition was assessed within a multi-site trial of Social Recovery Therapy (SRT) for individuals with FEP experiencing poor functioning (<30h weekly structured activity). Fifty-nine participants were randomly allocated to the therapy group (SRT + Early intervention), and 64 were allocated to treatment as usual group (TAU - early intervention care). Social cognition and neurocognition were assessed at baseline and 9 months; assessors were blind to group allocation. It was hypothesized that social cognition would improve following therapy, and those with better social cognition prior to therapy would benefit the most from SRT.

Results: There was no significant impact of SRT on individual neurocognitive or social cognitive variables, however, joint models addressing patterns of missingness demonstrate improvement across a number of cognitive outcomes following SRT. Further, regression analyses showed those who had better social cognition at baseline were most likely to benefit from the therapy ($\beta = .350$; 95% CI = .830 to 8.891; $p = .019$).

Conclusion: It is not clear if SRT impacts on social cognitive or neurocognitive function, however, SRT may be beneficial in those with better social cognition at baseline.

Keywords: Cognition; Functioning; Psychosocial intervention; Psychosis; Social disability.

1. Introduction

Many young people with first episode psychosis (FEP) have persisting difficulties with their social functioning, even when receiving high quality care under an Early Intervention Service (EIS; Hodgekins et al., 2015a). This highlights the need for new interventions to target social impairments in early psychosis.

Widespread impairments in social cognition, defined as the mental operations underlying social interactions (Adolphs, 2008), and neurocognition (e.g. memory), are evident in psychosis and are closely associated with functional impairments (Allott et al., 2011a; Couture et al., 2006). Whilst there are number of evidence-based psychosocial interventions targeting poor functioning in psychosis, the impact of these interventions on cognitive impairment is not well established (Kurtz, 2011). There is also considerable heterogeneity in therapy response (Fizdon et al., 2020). Thus, to ensure that psychosocial interventions are being delivered appropriately, it is important to understand which factors contribute to change, and identify individuals who are most likely to benefit (Allott et al., 2011b).

Neurocognitive variables have been identified as robust predictors of treatments response to a number of psychosocial interventions, demonstrating that poorer neurocognitive function prior to intervention has a rate-limiting impact on treatment progress (Kurtz, 2011; Kurtz and Mueser, 2008). Verbal memory, working memory, executive functions, problem-solving and attention have been shown to predict progress on proximal measures designed to assess acquisition of specific therapy-related skills following: social skills training (Kern et al., 1992; Kurtz and Mueser, 2008), cognitive remediation (Fizdon et al., 2020), and vocational rehabilitation (Bell and Bryson, 2001). The importance of baseline neurocognitive variables in predicting change in more distal outcomes has also been demonstrated. Verbal memory and

executive function were shown to predict long-term competitive employment outcomes following a supported employment programme (McGurk and Mueser, 2006), and composite neurocognition and social cognition predicted improvement in functioning following community rehabilitation (Brekke et al., 2007). Studies have failed to demonstrate a moderating influence of neurocognition on outcomes in response to Cognitive Behavioural Therapy (CBT) for psychosis, however, in these studies, the primary outcome was positive symptom reduction as opposed to improvement in functioning (Allott et al., 2011a; Garety et al., 1997).

Social cognition and neurocognition are also shown to improve following psychosocial interventions. In a meta-analysis evaluating a social cognition training programme, large effect sizes were found for facial affect recognition, moderate-sized effects for Theory of Mind (ToM), and small to medium effects for attribution style, in addition to moderate to large effects for community functioning (Kurtz et al., 2016). A meta-analysis has also reported on lasting improvements in social cognition, neurocognition, and functioning following an integrative psychosocial intervention with adjunctive cognitive remediation (Roder et al., 2006; Roder et al., 2011). These findings may demonstrate a reciprocal relationship between social cognition and neurocognition with functional improvement.

Despite these findings, interventions addressing poor functioning have largely focused on those with enduring illness. Interventions delivered earlier in a person's functional trajectory might be more effective at preventing disability from becoming entrenched (Griffiths et al., 2018). In early psychosis, the effectiveness of a specialised Social Recovery Therapy (SRT) has been demonstrated in a recent randomized controlled trial (Fowler et al., 2017). Individuals' receiving SRT alongside EI showed an 8.1-hour greater increase of weekly structured activity,

compared to those receiving EIS alone (95% CI 2.5 to 13.7; $p = 0.005$). SRT is designed to motivate individuals to re-engage with their social environment, whilst using CBT techniques, such as cognitive reappraisal strategies, to overcome barriers to successful social engagement (Fowler et al., 2009).

Although cognition was not addressed by the SRT, given the evidence that cognition is implicated in a range of evidence-based psychosocial interventions (Kurtz, 2011), we might expect to see a change in cognition post SRT, or that baseline cognition may limit the progress made via therapy. Further, with regards to the nature of the SRT and reliance on social and interpersonal skills to engage in a therapeutic relationship and re-engage in a social environment, it was hypothesized that social cognition is more likely implicated with SRT. Finally, it was apparent *a priori* that many individuals did not feel ready to return to work, and as such, the focus of the SRT was to encourage engagement in a range of social activities, rather than a return to main roles such as work or education. This, together with evidence showing that progress following vocational interventions is associated with neurocognition (e.g. attention and problem solving, which might be considered important for obtaining employment), we expect that SRT may be less reliant on neurocognitive functions.

Nevertheless, exploration of both social cognition and neurocognition pre- and post-SRT may be important to understand the dynamics of therapy, and explore which individuals are most likely to benefit. No studies to date have examined the impact of cognition on treatment response to a CBT intervention targeting functional impairment in psychosis. This study was designed to add value to the assessment of the effectiveness of SRT in a recent NIHR SUPEREDEN trial (reported elsewhere in Fowler et al. 2017), by addressing two main questions:

- (1) Does SRT lead to improvements in social cognition and neurocognition?
- (2) Do baseline social cognitive and/or neurocognitive variables predict which individuals are more likely to benefit from the SRT?

2. Method

2.1. Design

This was a sub-study within the multi-site NIHR SUPEREDEN trial, which was a single-blind, proof-of-principle trial, comparing SRT plus EIS care, against standard care from EIS alone (referred to as treatment as usual – TAU). SRT draws on psychological intervention and multi-systemic assertive outreach case management to promote social recovery. The trial is registered (ISRCTN61621571), and further details on the specific therapeutic approach can be found in Fowler *et al.* (2017). EIS care was provided under four recognised centres of excellence with high fidelity to the EIS model, indicating that the centres provided a comprehensive range of interventions including: CBT for psychosis, family interventions, supported employment, social support, assertive outreach case management, and medical and psychopharmacological management (Fowler *et al.*, 2017).

2.2. Sample

The inclusion criteria were as follows: (1) individuals aged 16-35 years with non-affective psychosis; (2) low level of structured activity after 12 months of EIS [defined as 30 hours or less per week on the Time Use Survey; (Short, 2006)]; (3) receiving EIS care between 12-30 months. Participants were excluded if they were: (1) not proficient in English language to engage in SRT; (2) deemed too unwell to partake; (3) diagnosed with a neurological disorder;

(4) had a learning disability; (5) had a history of severe head injury (more than 5 minutes loss of consciousness, or an overnight hospital stay).

One hundred and fifty-five individuals from four UK EIS centers (Birmingham, Lancashire, Norfolk and Sussex) consented to the trial and 122 completed cognitive assessments. The breakdown of group allocation post-randomization was as follows: 59 participants were allocated to the SRT plus EIS group, and 63 participants were allocated to the TAU group (EIS alone). An analysis comparing those who completed the cognitive assessments at baseline with those who did not (Table 1), showed no group differences on demographics, structured activity and clinical characteristics.

2.3. Measures

2.3.1. Functioning

The primary hypothesis of the trial was that SRT (plus EIS) would lead to improvements in social recovery (assessed as time spent in structured activity at 9 months). Structured activity was assessed using the Time Use Survey [TUS (Short, 2006)]. The TUS measures time spent over the last month in activities such as work, education, socializing, leisure, sports, chores and childcare. The average hours per week spent in ‘structured activity’ was calculated.

2.3.2. Cognitive study measures (Secondary outcome measure)

2.3.2.1. Neurocognition

To minimize participant burden and maximise engagement, a brief neurocognitive battery was selected. Verbal skills are shown to be most impaired in psychosis (Allott et al., 2011b), and as such, the battery included two verbal assessments: *verbal learning and memory* [Logical

Memory subtest - Wechsler Memory Scale Revised – IV; WMS-IV (Wechsler, 1987)]; and *verbal comprehension* [Vocabulary subtest Wechsler Adult Intelligence Scale – IV; WAIS-IV; (Wechsler, 1981)]. In addition, one non-verbal test was used: *problem-solving and visuospatial skills* [Block Design subtest WAIS-IV; (Wechsler, 1981)]. Raw scores on the subtests were converted into age standardized scores (range: 1-19), with a mean of 10 and a standard deviation of 3.

2.3.2.2. *Social Cognition*

Given the interpersonal nature of the intervention, we hypothesized that SC was more likely to explain response to SRT. We therefore included four social cognition sub-domains most commonly impaired in psychosis: *Theory of Mind* [Picture sequencing task; (Langdon et al., 2014)]; *emotion recognition* (Mayer-Salovey-Caruso Emotional Intelligence Test – Perceiving Emotions; MSCEIT; (Mayer, 2002)); *Attribution of blame bias* [Ambiguous Intentions Hostility Questionnaire; AIHQ; (Combs et al., 2007)]; and *social perception* [The Social Knowledge Questionnaire; SKQ ; (Cutting and Murphy, 1990)].

2.4. *Study Procedure*

Participants were randomly allocated using a computer program to either the treatment group (SRT + standard EIS care) or TAU (standard EIS care). Those allocated to the treatment group received SRT for 9-months by a CBT accredited therapist. Those allocated to TAU group continued to receive standard care under EIS. Participants completed a battery of cognitive assessments in addition to the main trial assessments at baseline (prior to randomization), and at 9-month follow-up. Although all trial participants were given the opportunity to complete the cognitive test battery at baseline, only 79% agreed to complete (see *section 2.2.*). Researchers were blind to group allocation. Where un-blinding occurred,

assessments were conducted by another researcher blind to group allocation. Consent was obtained on one occasion at the start of the trial. Trial researchers were trained on the use of cognitive assessments and completed regular inter-rater reliability checks in line with the NIHR SUPEREDEN protocol (Fowler *et al.*, 2017). The study was approved by the Black Country NHS research ethics committee (REC reference: 12/WM/009).

2.5. Statistical Analyses

2.5.1. Analysis 1: Impact of SRT on Social Cognition and Neurocognition

To explore the impact of SRT on each of the social cognitive and neurocognitive variables, a mixed 2x2 multivariable Analysis of Variance (MANOVA) was used. To determine a treatment effect, the ‘group x time’ interaction was inspected. A Bonferroni correction was applied to reduce the likelihood of a Family-Wise-Error (Haynes, 2013). Cohen’s *f* was used as a measure of effect size, with small, medium and large effects corresponding to values of .01, .025, and .04, respectively (Cohen, 1988).

The assumption of multivariable normality was met, and there were no problematic outliers. There was no indication from scatterplots of non-linearity, and there was no evidence of multicollinearity or singularity in the data (tolerance >.2, variance inflation factor <5; Pearson correlations <.7).

In the initial analysis, no imputation was conducted, but due to higher attrition in the control group by follow-up, data was assumed missing not at random. Missingness was addressed using joint multivariate modelling, where the continuous outcome score (using Gaussian error) was modelled with the observed loss to follow-up (with Bernoulli error) to describe the joint probability of the observed outcome. Joint modelling assumes a relationship between the

observed outcome and missing data points via an informative missingness parameter; although such a parameter cannot be estimated from the observed data, it can be specified with associated uncertainty (Chaimani et al., 2018). In this instance, we made the assumption that those who did not remain in the study, had poorer outcomes on the cognitive measures than those who remained. The justification of this assumption is given by the main trial analysis, whereby in this group who uniformly displayed extreme social withdrawal at baseline, by 15-months, the proportion of those lost to follow-up was more than two-times higher in the TAU group compared to the SRT group, implying a benefit on maintenance of engagement via therapy, which was reflected by an improvement on the primary outcome (Fowler et al., 2017). Joint modelling was undertaken here and in the main trial analysis, and was conducted using SAS Proc Glimmix (version 9.4, SAS Institute, CARY, NC).

2.5.2. Analysis 2: Social cognition and neurocognition as predictors of response to SRT

A Change score (difference in hours per week spent in structured activity between pre- and post SRT), was entered as a dependent variable in a linear regression, with social cognitive and neurocognitive scores entered as predictor variables. To handle the large number of variables entered into the regression, a backward method was employed to find an optimal model of the most parsimonious predictors of treatment response (Lindsey and Sheather, 2010). A post-hoc Bonferroni correction was applied in the case of significant regression models (Haynes, 2013).

3. Results

3.1. The Sample: cognitive study

Of the full trial sample, 79% (122) completed cognitive assessments at baseline. Demographic and clinical comparisons between the trial sample and the sub-group completing cognitive assessments (by group allocation) are shown in Table 1. There were no differences on demographic or clinical characteristics between the two groups at baseline.

At 9-month follow-up (post SRT), data were available for 109 of the 122 participants (89%) on the primary outcome (TUS). Not all cognitive assessments were completed by each participant who returned for follow-up. Table 2 provides a breakdown of missing data for each of the cognitive measures at 9-month follow-up. There was higher percentage of missing cognitive data in the control group, which supported an assumption that data were missing not at random (Table 2).

*****Please insert Table 1 here*****

3.2. Analysis 1: Impact of SRT on social cognition and neurocognition

Group by time MANOVAs were conducted for each of the social cognitive and neurocognitive variables. Means and standard deviations are presented in Table 2. The only significant group differences over time was observed for attribution of blame bias (95% CI = 0 to 0.437; $p = 0.49$), but this this did not retain its significance following a Bonferroni correction. There were no other significant differences observed on either of the social cognitive or neurocognitive variables. The group x time interactions are presented in Table 3. Due to the bias in the pattern of missingness (Table 2); the data was regarded as missing not at random. Based on the assumption that loss to follow-up was associated with poor scores on the cognitive measures, results of the joint models indicated significant differences on several of the cognitive variables (Table 3), including: verbal comprehension, non-verbal problem solving, ToM, social knowledge and attribution of blame bias.

*****Please insert Table 2 here*****

*****Please insert Table 3 here*****

3.3. Data Analysis 2: Social cognition and Neurocognition as predictors of response to SRT.

The change score on TUS was entered into a backward linear regression as a dependent variable. The final model included social knowledge and attribution of blame bias, accounting for 11% of variance (Table 4). Baseline social knowledge was the only significant independent predictor of change in structured activity ($\beta = .350$; 95% CI = .830 to 8.891; $p = .019$); individuals with better social knowledge at baseline appeared more likely to improve their structured activity post SRT. To illustrate the magnitude of this effect, individuals in the SRT group scoring in the top quartile at baseline on average, increased their structured activity by 11 hours more than those who scored below the 50th percentile.

*****Please insert Table 4 here*****

4. Discussion

4.1. Main Findings

First, we investigated whether there was a change on the social cognitive and neurocognitive variables following a specialized psychosocial intervention. In our main analysis, we did not identify robust statistically significant benefits of SRT on either neurocognition or social cognition over the follow-up interval, compared with TAU. Although attribution of blame bias was significant post SRT, this effect was lost once a Bonferroni correction was applied, suggesting this finding was perhaps due to chance.

Overall, findings may suggest that SRT had no impact on social cognition or neurocognition, but given the high level of missing data at 9 months, this finding might be biased if those who dropped out had poorer outcomes on the cognitive measures. As data were considered missing not at random, with the assumption that missingness was associated with a poorer outcome (an

assumption strongly supported by the primary and secondary analyses of main trial), joint models may be considered to provide the least biased assessment of the outcome (Hickey et al., 2016). Results of the joint modelling (multivariate) analyses demonstrated nominally significant treatment effects for several neurocognitive and social cognitive outcomes. Whilst this is encouraging, we are unable to make firm conclusions due to the lack of treatment effect in our main analysis.

Plausible reasons for the lack of treatment effect could be as follows: first, the SRT did not seek to address cognitive impairments. Second, in prior studies evaluating psychosocial interventions, these studies focused on homogenous groups with enduring illness, where the effects of illness chronicity, such as medication and repeated relapse, may contribute to impaired functioning (Roder et al., 2011). In contrast, the present sample represented a clinically heterogeneous subgroup with FEP who were deemed to have ‘severe social disability’, and likely to be a ‘chronic’ sample of the future. Comparing the severity of cognitive impairment reported in other early psychosis studies (typically 1 SD below the mean), impairments were also evident in this sample, and most pronounced for social knowledge, ToM, verbal memory and verbal comprehension (Cutting and Murphy, 1990; Langdon et al., 2014; Lin et al., 2011; Combs et al., 2007; Simons et al., 2016). Participants were mildly impaired on problem solving, and no impairment was observed for social perception, hence it is not surprising that we didn’t see a treatment effect for these variables. Interestingly, by follow-up, the neurocognitive scores were above the impairment threshold for both groups. Whilst this might represent a natural recovery process, improvement over time might simply reflect a practice effect.

A second aim of this study was to investigate whether social cognition and neurocognition predicted response to SRT. We found that individuals who had better social knowledge prior to therapy were more likely to improve their structured activity post therapy. Social knowledge may therefore have a moderating influence on response to SRT, expanding on prior research showing that baseline cognition has a rate-limiting impact on functional change following psychosocial rehabilitation (Brekke et al., 2007). We postulated that those who have a better understanding of the social world may be more able to use adaptive cognitive reappraisal strategies to engage in social activity and a psychosocial intervention, compared to those who struggle to understand social situations (Rowland et al., 2013). They may also form a better therapeutic alliance leading to a better therapy response (Allott et al., 2011b). But it must be acknowledged that no other social cognitive sub-domain predicted treatment response, and chance is a plausible explanation for the observation given the high number of variables in the model.

4.2. Implications and future directions

Firstly, the potential to identify individuals at an early stage who are more likely to respond to therapy could ensure that costly interventions (for both health service and service user), are delivered to those who are more likely to benefit. An *a priori* study comparing response to SRT between those with 'poor' and 'good' social knowledge would be a direct test of this assumption. A second implication is the potential to guide interventions for those who are less responsive; improving social cognition might improve response to SRT. Refining SRT by incorporating a social cognition intervention may enhance its efficacy in those with deficits in social cognition (Bartholomeusz et al., 2013). Indeed, integrative psychosocial interventions with adjunctive cognitive training are shown to have greater cognitive and real-world functional improvements in schizophrenia (Horan and Green, 2017; Roder et al., 2011). The

application of an integrated, multi-dimensional treatment approach combining SRT with a form of social cognitive intervention may produce more durable and sustained therapeutic effects for individuals with poor functioning in FEP.

4.3. Study Strengths and Limitations

The study benefits from being part of a large scale, multi-site, randomized controlled trial, where researchers were blind to group allocation. Participants were recruited from several centers across the UK, which included urban, rural and town settings, making the sample highly diverse and representative. Nevertheless, despite a substantial proportion of the overall sample completing cognitive assessments at baseline, 20% of the trial sample did not. This may have biased the findings, as those with poorer cognitive function may have chose not to engage in cognitive assessments. There was also a significant number of participants who did not complete the cognitive assessments (up to 40%; Table 4), resulting in inadequate power to detect a posited significant post-intervention effect. Further, there was a bias for missing data in the control arm at follow-up, and it is plausible that individuals with missing data, on average, had lower scores on the cognitive measures compared to those who remained in the study, making the findings difficult to interpret. Although we attempted to address this bias with joint modelling, we are unable to draw on conclusions without replication in a larger sample. Further, we cannot rule out that the higher percentage of missing cases simply reflected the challenge of retaining TAU participants in trials. This group of young people in particular are an incredibly challenging group to engage and retain in research. The pattern of missingness was evident across all measures in the main trial, and twice-higher in the TAU group 6-months post-intervention (Fowler et al., 2017). As the SRT aims to promote social engagement, and research participation requires a level of social engagement, this may demonstrate that missingness in this instance was related to the outcomes of interest. A final limitation is the

selection of cognitive subtests and their ecological validity and sensitivity to detect changes following improvements in structured activity. In particular, vocabulary and block design may be insensitive to detect fluctuations in cognitive performance (Nuechterlein et al., 2008). Further, a number of social cognitive measures, including the AIHQ used in this study, have been deemed unsuitable for assessing social cognition in early psychosis (Ludwig et al., 2017; Pinkham et al., 2014). Future research should aim to select appropriate measures identified by consensus groups (e.g. MATRICS), to ensure the reliability of findings across studies (Nuechterlein et al., 2008; Pinkham et al., 2014).

4.4. Conclusion

Results of the main analysis suggest that SRT had no impact on social cognition or neurocognition, however, joint models suggest an improvement on several social cognitive and neurocognitive variables following treatment. Those who had better social knowledge at baseline were also more likely to improve their structured activity post SRT. Although no firm conclusions can be made about the treatment effect on cognition, the findings have implications for targeted intervention for individuals in the early phase of psychosis with poor social cognition. Combining SRT with a social cognitive intervention may maximize its impact by providing more durable and sustained treatment effects, particularly for those who may not respond to SRT alone.

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6. References

- Adolphs, R., 2008. The Social Brain: Neural Basis of Social Knowledge. *Annual Review of Psychology* 60(1), 693-716.
- Allott, K., Liu, P., Proffitt, T.M., Killackey, E., 2011a. Cognition at illness onset as a predictor of later functional outcome in early psychosis: systematic review and methodological critique. *Schizophr Res* 125(2-3), 221-235

Allott, K., Alvarez-Jimenez, M., Killackey, E.J., Bendall, S., McGorry, P.D., Jackson, H.J., 2011b. Patient predictors of symptom and functional outcome following cognitive behaviour therapy or befriending in first-episode psychosis. *Schizophrenia Research* 132(2), 125-130.

Allott, K., Liu, P., Proffitt, T.M., Killackey, E., 2011b. Cognition at illness onset as a predictor of later functional outcome in early psychosis: systematic review and methodological critique. *Schizophr Res* 125(2-3), 221-235.

Bartholomeusz, C.F., Allott, K., Killackey, E., Liu, P., Wood, S.J., Thompson, A., 2013. Social cognition training as an intervention for improving functional outcome in first-episode psychosis: a feasibility study. 7(4), 421-426.

Bell, M.D., Bryson, G., 2001. Work Rehabilitation in Schizophrenia: Does Cognitive Impairment Limit Improvement? *Schizophrenia Bulletin* 27(2), 269-279.

Brekke, J.S., Hoe, M., Long, J., Green, M.F., 2007. How neurocognition and social cognition influence functional change during community-based psychosocial rehabilitation for individuals with schizophrenia. *Schizophrenia bulletin* 33(5), 1247-1256.

Chaimani, A., Mavridis, D., Salanti, G., Higgins, J.P.T., White, I.R., 2018. Allowing for Informative Missingness in Aggregate Data Meta-Analysis with Continuous or Binary Outcomes: Extensions to Metamiss. *The Stata Journal* 18(3), 716-740.

Cohen, J., 1988. *Statistical Power Analysis for the Behavioral Sciences* 2nd Edition ed. Lawrence Erlbaum Associates, Publishers, Hillsdale, NJ.

Combs, D.R., Adams, S.D., Penn, D.L., Roberts, D., Tiegreen, J., Stem, P., 2007. Social Cognition and Interaction Training (SCIT) for inpatients with schizophrenia spectrum disorders: Preliminary findings. *Schizophrenia Research* 91(1), 112-116.

Combs, D.R., Penn, D.L., Wicher, M., Waldheter, E., 2007. The Ambiguous Intentions Hostility Questionnaire (AIHQ): A new measure for evaluating hostile social-cognitive biases in paranoia. *Cognitive Neuropsychiatry*, 12(2), 128-143.

Couture, S.M., Penn, D.L., Roberts, D.L., 2006. The Functional Significance of Social Cognition in Schizophrenia: A Review. *Schizophrenia Bulletin* 32(suppl_1), S44-S63.

Cutting, J., Murphy, D., 1990. Impaired ability of schizophrenics, relative to manics or depressives, to appreciate social knowledge about their culture. *British Journal of Psychiatry*, 157, 355-258. <https://doi.org/10.1192/bjp.157.3.355>

Fiszdon, J.M., Kurtz, M.M., Parente, L., Choi, J., 2020. What variables predict cognitive remediation associated improvement in individuals with psychosis? *Schizophr Res Cogn* 19, 100148.

Fowler, D., Hodgekins, J., French, P., Marshall, M., Freemantle, N., McCrone, P., Everard, L., Lavis, A., Jones, P.B., Amos, T., Singh, S., Sharma, V., Birchwood, M., 2017. Social recovery therapy in combination with early intervention services for enhancement of social recovery in patients with first-episode psychosis (SUPEREDEN3): a single-blind, randomised controlled trial. *The Lancet Psychiatry*, 5, 1, 41-50. [https://doi.org/10.1016/S2215-0366\(17\)30476-5](https://doi.org/10.1016/S2215-0366(17)30476-5).

Fowler, D., Hodgekins, J., Painter, M., Reilly, T., Crane, C., Macmillan, I., Mugford, M., Croudace, T., Jones, P.B., 2009. Cognitive behaviour therapy for improving social recovery in psychosis: a report from the ISREP MRC Trial Platform study (Improving Social Recovery in Early Psychosis). *Psychological Medicine* 39(10), 1627-1636.

Garety, P., Fowler, D., Kuipers, E., Freeman, D., Dunn, G., Bebbington, P., Hadley, C., Jones, S., 1997. London-East Anglia randomised controlled trial of cognitive-behavioural therapy for psychosis. II: Predictors of outcome. *Br J Psychiatry* 171, 420-426.

Griffiths, S.L., Wood, S.J., Birchwood, M., 2018. Vulnerability to psychosocial disability in psychosis. *Epidemiology and psychiatric sciences* 28(2): 140-145. <https://doi.org/10.1017/S2045796018000495>.

- Haynes, W., 2013. Bonferroni Correction, in: Dubitzky, W., Wolkenhauer, O., Cho, K.-H., Yokota, H. (Eds.), *Encyclopedia of Systems Biology*. Springer New York, New York, NY, pp. 154-154.
- Hickey, G.L., Philipson, P., Jorgensen, A., Kolamunnage-Dona, R., 2016. Joint modelling of time-to-event and multivariate longitudinal outcomes: recent developments and issues. *BMC Medical Research Methodology* 16(1), 117.
- Hodgekins, J., Birchwood, M., Christopher, R., Marshall, M., Coker, S., Everard, L., Lester, H., Jones, P., Amos, T., Singh, S., Sharma, V., Freemantle, N., Fowler, D., 2015a. Investigating trajectories of social recovery in individuals with first-episode psychosis: A latent class growth analysis. *The British Journal of Psychiatry*, 207, 536-543.
- Hodgekins, J., French, P., Birchwood, M., Mugford, M., Christopher, R., Marshall, M., Everard, L., Lester, H., Jones, P., Amos, T., Singh, S., Sharma, V., Morrison, A.P., Fowler, D., 2015b. Comparing time use in individuals at different stages of psychosis and a non-clinical comparison group. *Schizophrenia Research* 161(2), 188-193.
- Horan, W.P., Green, M.F., 2017. Treatment of social cognition in schizophrenia: Current status and future directions. *Schizophrenia Research*. 203, 3-11.
<https://doi.org/10.1016/j.schres.2017.07.013>
- Kern, R.S., Green, M.F., Satz, P., 1992. Neuropsychological predictors of skills training for chronic psychiatric patients. *Psychiatry Res* 43(3), 223-230.
- Kurtz, M.M., 2011. Neurocognition as a predictor of response to evidence-based psychosocial interventions in schizophrenia: what is the state of the evidence? *Clinical psychology review* 31(4), 663-672.
- Kurtz, M.M., Gagen, E., Rocha, N.B.F., Machado, S., Penn, D.L., 2016. Comprehensive treatments for social cognitive deficits in schizophrenia: A critical review and effect-size analysis of controlled studies. *Clinical Psychology Review* 43(Supplement C), 80-89.
- Kurtz, M.M., Mueser, K.T., 2008. A meta-analysis of controlled research on social skills training for schizophrenia. *Journal of Consulting and Clinical Psychology* 76(3), 491-504.
- Langdon, R., Connors, M.H., Still, M., Ward, P.B., Catts, S., 2014. Theory of mind and neurocognition in early psychosis: a quasi-experimental study. *BMC Psychiatry* 14(1), 316.
- Lauronen, E., Miettunen, J., Veijola, J., Karhu, M., Jones, P.B., Isohanni, M., 2007. Outcome and its predictors in schizophrenia within the Northern Finland 1966 Birth Cohort. *European Psychiatry* 22(2), 129-136.
- Lin, A., Wood, S.J., Nelson, B., Brewer, W.J., Spiliotacopoulos, D., Bruxner, A., Broussard, C., Pantelis, C., Yung, A.R., 2011. Neurocognitive predictors of functional outcome two to 13 years after identification as ultra-high risk for psychosis. *Schizophrenia Research* 132(1), 1-7.
- Lindsey, C., Sheather, S., 2010. Variable Selection in Linear Regression. *The Stata Journal* 10(4), 650-669.
- Ludwig, K.A., Pinkham, A.E., Harvey, P.D., Kelsven, S., Penn, D.L., 2017. Social cognition psychometric evaluation (SCOPE) in people with early psychosis: A preliminary study. *Schizophrenia Research* 190(Supplement C), 136-143.
- Mayer, J.D., Salovey, P., & Caruso, D. R. , 2002. Mayer– Salovey–Caruso Emotional Intelligence Test (MSCEIT) item booklet. Multi-Health Systems In Publishers, Toronto, Ontario, Canada.
- McGurk, S.R., Mueser, K.T., 2006. Cognitive and clinical predictors of work outcomes in clients with schizophrenia receiving supported employment services: 4-year follow-up. *Adm Policy Ment Health* 33(5), 598-606.
- Nuechterlein, K.P.D., Green, M.F.P.D., Robert S. Kern , P.D., Lyle E. Baade , P.D., Deanna M. Barch , P.D., Jonathan D. Cohen , M.D., Ph.D. , Susan Essock , P.D., Wayne S. Fenton , M.D., Frederick J. Frese , I., Ph.D. , James M. Gold , P.D., Terry Goldberg , P.D., Robert K.

Heaton , P.D., Richard S.E. Keefe , P.D., Helena Kraemer , P.D., Raquelle Mesholam-Gately , P.D., Larry J. Seidman , P.D., Ellen Stover , P.D., Daniel R. Weinberger , M.D., Alexander S. Young , M.D., M.S.H.S. , Steven Zalcman , M.D., Stephen R. Marder , M.D., 2008. The MATRICS Consensus Cognitive Battery, Part 1: Test Selection, Reliability, and Validity. *American Journal of Psychiatry* 165(2), 203-213.

Pinkham, A.E., Penn, D.L., Green, M.F., Buck, B., Healey, K., Harvey, P.D., 2014. The Social Cognition Psychometric Evaluation Study: Results of the Expert Survey and RAND Panel. *Schizophrenia Bulletin* 40(4), 813-823.

Roberts, D.L., Combs, D.R., Willoughby, M., Mintz, J., Gibson, C., Rupp, B., Penn, D.L., 2014. A randomized, controlled trial of Social Cognition and Interaction Training (SCIT) for outpatients with schizophrenia spectrum disorders. *British Journal of Clinical Psychology* 53(3), 281–298.

Roder, V., Mueller, D.R., Mueser, K.T., Brenner, H.D., 2006. Integrated Psychological Therapy (IPT) for Schizophrenia: Is It Effective? *Schizophrenia Bulletin* 32(suppl_1), S81-S93.

Roder, V., Mueller, D.R., Schmidt, S.J., 2011. Effectiveness of Integrated Psychological Therapy (IPT) for Schizophrenia Patients: A Research Update. *Schizophrenia Bulletin* 37(suppl_2), S71-S79.

Rowland, J., Hamilton, M., Vella, N., Lino, B., Mitchell, P., Green, M., 2013. Adaptive Associations between Social Cognition and Emotion Regulation are Absent in Schizophrenia and Bipolar Disorder. *Frontiers in Psychology* 3, 607.

Short, S., 2006. Review of the UK 2000 Time Use Survey. . London: Office for National Statistics.

Simons, C.J.P., Bartels-Velthuis, A.A., Pijnenborg, G.H.M., Genetic, R., Outcome of Psychosis, I., 2016. Cognitive Performance and Long-Term Social Functioning in Psychotic Disorder: A Three-Year Follow-Up Study. *PLoS One* 11(4), e0151299-e0151299.

Wechsler, D., 1981. WAIS-R : manual : Wechsler adult intelligence scale--revised. Harcourt Brace Jovanovich [for] Psychological Corp., New York, NY.

Wechsler, D., 1987. Manual for the Wechsler Memory Scale-Revised. The Psychological Corporation, San Antonia.

Tables

Table 1. Comparison of demographic and clinical characteristics (by trial allocation) between the full trial sample compared with the sub-group completing cognitive assessments.

Item	SRT group allocation in the cognitive sub-sample (n=59)	SRT group allocation in the trial sample (n=75)	95% CI of the difference and p-value	TAU group allocation in the cognitive sub-sample (n=63)	TAU group allocation in the trial sample (n=79)	95% CI of the difference and p-value
Mean Age (years)	25.89	24.8	-1.88 to 1.56 (<i>p</i> =.851)	24.8	24.2	-1.35 to 1.87 (<i>p</i> =.750)
Gender:						
Male (n; %)	44; 72.9	56; 74.7	0.52 to 2.38 (<i>p</i> =.929) ^a	54; 84.4	60; 76.0	0.69 to 3.81 (<i>p</i> =.369) ^a
Years in Education (Mean)	12.1	12	-0.66 to 0.37 (<i>p</i> =.584)	12.2	12	-0.65 to 0.46 (<i>p</i> =.738)
DUP (days; mean)	341	240	-411.21 to 209.62 (<i>p</i> =.522)			
Time Use Survey	10	11	-1.85 to 3.31 (<i>p</i> =.577)	270	285	-118.94 to 148.33 (<i>p</i> =8.28)
Symptoms (PANSS; mean)				11.6	12	-2.49 to 3.16 (<i>p</i> =.814)
Positive	13.3	13.3	-1.64 to 1.66 (<i>p</i> =.989)	14.1	14.6	-1.39 to 2.39 (<i>p</i> =.603)
Negative	15.6	15.5	-2.27 to 2.11 (<i>p</i> =.940)	16.8	16.6	-2.24 to 1.81 (<i>p</i> =.834)
General	32.4	32.8	-2.60 to 3.42 (<i>p</i> =.790)	32.3	33.7	-1.59 to 4.27 (<i>p</i> =.367)

^a Yates' Correction for continuity. SRT = Social Recovery Therapy. TAU = Treatment as Usual. Cognitive sub-sample = participants who completed cognitive assessments and were included in the present study.

Table 2. Means and standard deviations for the individual cognitive variables by group allocation, along with missing data at 9-month follow-up.

Cognitive measures	Baseline		9-month Follow-up		Missing data at 9-month follow-up (n; %)	Missing data at 9-month follow-up (n; %)
	SRT M±SD	TAU M±SD	SRT M±SD	TAU M±SD	SRT Group (n=59)	TAU Group (n=63)
Verbal learning and memory ^a	8.65 (3.75)	6.88 (2.95)	8.94 (3.77)	7.83 (3.53)	12 (20%)	21 (33%)
Verbal comprehension ^a	8.49 (3.51)	7.38 (2.85)	9.14 (3.23)	8.31 (3.22)	12 (20%)	23 (36.5%)
Non-verbal problem solving ^a	8.71 (3.08)	7.74 (3.33)	9.39 (3.37)	8.76 (2.78)	11 (19%)	22 (35%)
Theory of Mind ^b	4.44 (0.98)	4.35 (1.07)	4.74 (0.94)	4.56 (1.05)	11 (19%)	22 (35%)
Social Knowledge ^b	7.00 (1.87)	6.48 (1.55)	7.16 (1.75)	6.95 (1.53)	7 (12%)	20 (32%)
Attribution of blame bias ^b	2.94 (0.91)	2.85 (0.88)	2.49 (0.98)	2.84 (0.93)	9 (15%)	25 (40%)
Social perception ^b	98.36 (18.60)	95.44 (22.59)	101.92 (18.69)	96.86 (18.22)	18 (31%)	21 (33%)

^aNeurocognitive sub-domain. ^bSocial Cognition sub-domain. Group = Social Recovery Therapy + Early Intervention vs. Treatment as usual (Early Intervention). ‡p value from joint modelling (multivariate) analyses. The p value is the effect of treatment allocation, derived from the joint model of loss to follow up and item.

Table 3. MANOVA results testing for treatment effects in each of the social cognitive and neurocognitive variables, along with results of the joint models.

	Effect size (95% CI)	<i>p</i> – value	‡ <i>p</i> value
<i>Group x Time Interaction Effect</i>			
Verbal learning and memory ^a	0.088 (0 to 0.346)	.197	0.138
Verbal comprehension ^a	0 (0 to 0.263)	.603	0.004
Non-verbal problem solving ^a	0 (0 to 0.288)	.452	0.025
Theory of Mind ^b	0 (0 to 0.272)	.565	0.011
Social Knowledge ^b	0.092 (0 to 0.346)	.187	0.012
Attribution of blame bias ^b	0.196 (0 to 0.437)	.049 [#]	0.008
Social perception ^b	0 (0 to 0.281)	.562	0.359

^aNeurocognitive sub-domain. ^bSocial Cognition sub-domain. [#]No longer significant after applying a Bonferroni correction. ‡*p* value from joint modelling (multivariate) analyses – the *p* value is the effect of treatment allocation, derived from the joint model of loss to follow up and item.

Table 4. Optimal regression model predicting change in Time Use post Social Recovery Therapy using baseline social cognitive and neurocognitive predictors.

Predictors	β	95% CI	Adjusted <i>R</i> ²	<i>p</i> – value
Social Knowledge	.350*†	.830 to 8.891	0.110	0.029
Attribution of Blame Bias	.278	-.358 to 15.866		

* ≤ 0.05 . † Retains Significance with a post-hoc Bonferroni Correction.

