Schizophrenia and Autism as Contrasting Minds: Neural Evidence for the Hypo-Hyper-Intentionality Hypothesis

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Both schizophrenia (SCZ) and autism spectrum disorder (ASD) are characterized by mentalizing problems and associated neural dysfunction of the social brain. However, the deficits in mental state attribution are somehow opposed: Whereas patients with SCZ tend to over-attribute intentions to agents and physical events (“hyper-intentionality”), patients with ASD treat people as devoid of intentions (“hypo-intentionality”). Here we aimed to investigate whether this hypo-hyper-intentionality hypothesis can be supported by neural evidence during a mentalizing task. Using functional magnetic resonance imaging (fMRI), we investigated the neural responses and functional connectivity during reading others’ intention. Scanning was performed in 23 individuals with ASD, 18 with paranoid SCZ, and 23 gender and IQ matched control subjects. Both clinical groups showed reduced brain activation compared to controls for the contrast intentional vs physical information processing in left posterior superior temporal sulcus (pSTS) and ventral medial prefrontal cortex (vMPFC) for SCZ, and right pSTS in ASD. As predicted, these effects were caused in a group specific way: Relative increased activation for physical information processing in SCZ that was also correlated with positive PANNS score and relative decreased activation for intentional information processing in ASD. Additionally, we could demonstrate opposed connectivity patterns between the right pSTS and vMPFC in the clinical groups, i.e., increased for SCZ, decreased for ASD. These findings represent opposed neural signatures in key regions of the social brain as predicted by the hyper-hypo-intentionality hypothesis.

Key words: autism/schizophrenia/mentalising/intention/MPFC/pSTS

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Introduction

Autism spectrum disorder (ASD) and schizophrenia (SCZ) are 2 distinct neuropsychiatric disorders: ASD is characterized by deficits in social communication alongside stereotyped, repetitive behaviors, whereas SCZ is characterized by a combination of positive (hallucinations, delusions, and thought disorder) and negative symptoms (apathy, speech impairment, and flat affect). Despite these evident clinical discrepancies, ASD and SCZ both show altered development and function of the social brain, a specialized neural network dedicated to social cognition. Social cognition refers to automatic and controlled psychological processes that support social interaction. In particular, a specific cognitive ability, referred to as “theory of mind” (ToM), mentalizing or mind-reading, allows humans to represent and attribute different types of mental states to others, i.e., inferring beliefs and reading intentions. Two specific areas, the medial prefrontal cortex (MPFC) and the right posterior superior temporal sulcus (pSTS) along with the adjacent temporo-parietal junction (TPJ) are prominently involved in intention reading: The MPFC is activated in social situations, implying communicative intent and triadic social interaction, whereas the right pSTS/TPJ is engaged in processing more basic and simple intentions like recognizing a person’s goal.

ASD and SCZ share cognitive behavioral and neural dysfunction in mind-reading. The over-attribution of intentions to other people is a key symptom in SCZ, and psychotic symptoms, like delusions, may reflect a failure of affected individuals to monitor and distinguish their own from represented other persons’ mental states and
behavior. A behavioral study in which moving geometric figures enacted various social situations showed poor performance for SCZ compared to typically developing individuals (TD) implying that mind-reading deficits might lead paranoid patients to misunderstand others’ intentions. Few fMRI studies in SCZ have tested intention reading and reported fronto-temporo-parietal altered neural activation, although these findings were not consistently supported by a specific behavioral impairment.

Individuals with ASD appear to be “mind blind,” ie, have difficulties to understand and read others’ mind usually performing poor in tests of mind-reading. The geometric stimuli described above were also used for testing social attribution in ASD and reported a lack of spontaneous attribution of mental states. In ASD, reduced activation during processing of social information has been described in the right pSTS, as well as decreased fronto-temporal connectivity during mind-reading.

Consistent with these studies, it has been hypothesized that ASD and SCZ thus may be located at the extreme ends of a cognitive architecture ranging from a mechanistic hypo-intentional (to treat person as objects) to a mentalistic hyper-intentional (to treat objects as persons) mode of cognition, respectively. These opposed modes of cognition should be reflected by opposed patterns of brain activation and connectivity patterns between regions of the social brain. It has already been shown that overall brain connectivity is decreased in ASD but increased in SCZ.

To date, no fMRI-study has directly compared ASD and SCZ during a mentalizing task. To address the hyper-hypo-intentionality hypothesis, we conducted an fMRI experiment in which participants had to read different kinds of intentions, ie, private (1 person acting) and communicative (2 persons interacting) with a physical (non-intentional) condition serving as a control condition. It was demonstrated that patients with SCZ show a decreased differential activation (intentional minus non-intentional) of the MPFC and bilateral pSTS which, and this is relevant here, was driven mainly by increased activation during the (non-intentional) physical control condition. This is consistent with a hyper-intentional mode of cognition. For ASD patients, known for a more mechanistic cognition mode, we predicted also reduced activation, which however should be rather driven by reduced activation during the mentalizing condition, consistent with a hypo-intentional mode of cognition (figure 1).

In addition, 2 other predictions were tested. Based on previous results, we hypothesized that differences in neural function would be evident primarily in more complex social (communicative) intention reading compared to more simple (private) intention reading. This hypothesis is based primarily on our previous study in patients with SCZ who showed no aberrant activation patterns in simple, but only in complex intention reading. Furthermore, we aimed to explore whether the assumed differences between groups would also be reflected in opposed connectivity patterns of the right pSTS, a key region of intention attribution with other hubs of the social brain.

Methods and Materials

Participants

We included 23 right-handed patients (2 females) with ASD aged 14–33 years, mean IQ (SD) 105.9 (13.93). We also included 18 right-handed individuals (4 females) with paranoid SCZ, aged 14–32 years, mean IQ (SD) 101.4 (12.37). Twenty-three right-handed TD participants (4 females), aged 15–27 years, mean IQ (SD) 106.8 (10.9), were recruited from the community. The study was approved by the local ethics committee. Written informed consent was obtained from the subjects and their parents (for subjects <18 years). For more details about the 3 groups see supplementary material.

Experimental Design

We used a picture sequencing task similar to with a total of 30 comic strips shown via goggles, (each of 15 s duration) with a jittered inter-trial interval of 7–11 s showing a fixation-cross. Each comic strip included 2 phases. In Phase 1 (the story-phase) 3 pictures (3 s each) depicting an unfolding story plot were displayed sequentially. During Phase 2 (the choice-phase) 3 answer pictures were presented simultaneously (6 s). The participants’ task was to choose via button press the picture showing the logical story ending. Comic strips belonged to 1 of the
following 2 experimental conditions: (1) Private intention (PInt) representing a person’s intention with regard to her/his own action and plans, eg, observing a single person changing a broken bulb in order to read a book. (2) Communicative intention (CInt) representing the intention to communicate conveyed by communicative gestures, eg, observing a person requiring another person for a glass of water. In the control condition (third condition: physical causality or PhC), the comic strip showed non-intentional causal links among objects, eg, a ball blown by a gust of wind knocking over and breaking a glass of water.

In order to perform the task in the PInt and CInt conditions, it was necessary to read the intention of the characters. In the Ph-C condition (our control condition) the participants viewed scenes showing non-intentional causal links among objects, eg, a rock is rolling down a hill hitting a fence.

The experimental protocol was administered in 1 session (10 trials per condition) in pseudo-randomized order. Before scanning, each participant was trained outside the scanner with different comic strips for each category.

**Behavioral Statistics**

Descriptive statistics were calculated by \( \chi^2 \) and t-tests as appropriate. Participants’ reaction times and response accuracy were measured during scanning (for technical reasons, we collected behavioral data for 21 ASD subjects only). Data were analyzed with SPSS Statistics 17.0 by 1-way ANCOVA with age as a covariate (the SCZ group was older than the ASD and TD groups) and subsequent comparisons between means, using Bonferroni’s post hoc test (\( P < .05 \)).

**Imaging Parameters**

FMRI data were collected using a 3-Tesla Siemens Allegra scanner. Functional images were acquired using an echoplanar imaging (EPI) sequence. We obtained 384 whole brain scans. One volume consisted of 30 slices (slice thickness 3 mm + 0.75 mm gap, FOV = 192 mm, repetition time = 2 s, echo time = 30 ms, 64 \( \times \) 64 matrix, flip angle 80°).

**Data Analysis**

**Data Preprocessing.** The first 4 volumes were discarded in order to allow for T2-equilibration. Data preprocessing and statistical analyses were carried out with SPM8 (http://www.fil.ion.ucl.ac.uk/spm/software/spm8) following a standard sequence:

- Data were realigned, normalized with a voxel size of 2 mm \( \times \) 2 mm \( \times \) 2 mm and smoothed with an 8-mm Gaussian filter. For each trial, the variance of each voxel was estimated according to the general linear model. Intrinsic autocorrelations were accounted for by the first-order autoregressive AR(1) process, and low-frequency drifts were removed via a high-pass filtering (128 s).

**ROI Analysis.** This study based on a region of interest (ROI) analysis. Our ROIs were defined using an independent localizer “ToM” task\(^{15} \) in order to select 4 independent ROIs (ventral and dorsal MPFC, right and left pSTS). For more details for the localizer task and ROIs definition, see supplemental material. The rationale for the selection of these ROIs is based on the relevance of these regions considered to be the key brain areas during mind-reading\(^6\). Furthermore, based on our previous results, the MPFC and the bilateral pSTS were the unique brain regions showing aberrant activation in SCZ during an intention’s reading\(^ {18} \).

The first-level regression model consisted of 3 predictors defined for each condition (PhC, PInt, and CInt) modeled as an event (15 s) and convolved with the hemodynamic response function and 6 regressors describing residual motion.

In a second-level random effects group analysis, individual regionally specific effects of conditions for each subject were compared using a 3 \( \times \) 3 full factorial design (ANOVA) with condition and group as factors resulting in a \( t \)-statistic for every voxel. We also included an additional covariate “Age.” The significance threshold was set to \( P < .05 \) (k > 15) familywise error corrected for multiple comparisons within the a priori-defined regions of interest (right and left pSTS, ventral and dorsal MPFC) provided by the main effect activation of the localizer experiment. In order to compare the mean activations pairwise per condition, individual peak voxel data within the ROIs were extracted from the respective regressor and region and were subsequently analyzed externally using SPSS Statistics 17.0. Anatomical regions are reported according to standard atlases\(^ {36,37} \).

**Functional Connectivity**

We performed a functional connectivity analysis (fcMRI analysis) in order to explore the extent to which the right pSTS is connected to the rest of the brain during intention reading. A seed-based correlational analysis was performed using the CONN-fMRI functional connectivity toolbox (http://web.mit.edu/swg/software.htm\(^ {38} \)) and SPM8. After preprocessing, data were band-pass filtered (0.01 Hz < f < 0.15 Hz). Functional connectivity for each subject was examined during each trial. Regressors of no interest included the first-order derivatives of the 6 motion parameters, the 3 main condition effects (PhC, PInt, and CInt) as well as the eigenvectors from a principal component analysis on the white matter and cerebrospinal fluid voxels. Correlation maps were produced by extracting the residual BOLD time course from the seed region and computing Pearson’s correlation coefficients between the time course of the right pSTS and the time courses of all other voxels of the brain. Correlation coefficients were converted to normally distributed scores using Fisher’s
transformation to allow for second-level GLM analyses. The resulting contrast images were entered into a random-effects ANOVA included an additional covariate “Age.”

GLMs were calculated using the contrast images reflecting the connectivity change of the CInt and PhC conditions for the 3 groups. In fact, due to the functional results with no differences for the PInt>PhC, we performed the fcMRI group analysis for the CInt>PhC contrast only. Results were considered significant using an uncorrected $P < .001$ ($k > 15$) corrected for multiple comparisons.

Results

Behavioral Performance

For response accuracy we found a group effect $F(2,58) = 8.13; P = .001$ (ASD made more errors), but no effect of condition $F(2,58) = 0.79; P = .46$ or any interaction $F(4,88) = 0.92, P = .46$. For reaction times we found a group effect $F(2,58) = 3.42; P = .04$ (SCZ were slower), but again no effect of condition $F(2,58) = 1.27; P = .29$ or any interaction $F(2,58) = 0.86; P = .49$. See also supplementary table 2.

Functional Imaging Results

When comparing each intentional condition (PInt and CInt) with the physical control condition (PhC), differences in activation patterns between groups were found driven by specific aberrant activation within the 4 ROIs in pair-wise comparisons as predicted (table 1).

Table 1. Coordinates and Anatomic Localization for the ROI Analysis (Within and Between Analysis)

<table>
<thead>
<tr>
<th></th>
<th>PInt vs PhC</th>
<th>CInt vs PhC</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>x  y  z  Z</td>
<td>x  y  z  Z</td>
</tr>
<tr>
<td>Control</td>
<td>60  -54  18  5.5</td>
<td>56  -56  20  Inf</td>
</tr>
<tr>
<td></td>
<td>2    48  -18  6.26</td>
<td>0    56  32  3.97</td>
</tr>
<tr>
<td>Ventral MPFC</td>
<td>0    56  32  3.97</td>
<td>0    56  32  3.97</td>
</tr>
<tr>
<td>Dorsal MPFC</td>
<td>0    56  32  3.97</td>
<td>0    56  32  3.97</td>
</tr>
<tr>
<td>SCZ</td>
<td>54  -60  10  5.11</td>
<td>54  -60  10  5.11</td>
</tr>
<tr>
<td>ASD</td>
<td>56  -60  10  5.11</td>
<td>56  -60  10  5.11</td>
</tr>
<tr>
<td>Ventral MPFC</td>
<td>0    46  -20  6.31</td>
<td>0    54  34  4.92</td>
</tr>
<tr>
<td>Dorsal MPFC</td>
<td>-4    54  34  4.92</td>
<td>-4    54  34  4.92</td>
</tr>
<tr>
<td>CONTROL vs SCZ</td>
<td>-47    70  16  3.98</td>
<td>-46    70  16  3.98</td>
</tr>
<tr>
<td>Posterior STS</td>
<td>6     48  -20  3.91</td>
<td>6     48  -20  3.91</td>
</tr>
<tr>
<td>CONTROL vs ASD</td>
<td>60   -48  -2  3.95</td>
<td>60   -48  -2  3.95</td>
</tr>
<tr>
<td>Posterior STS</td>
<td>-4    54  34  4.22</td>
<td>-4    54  34  4.22</td>
</tr>
<tr>
<td>ASD vs SCZ</td>
<td>-4    54  34  4.22</td>
<td>-4    54  34  4.22</td>
</tr>
</tbody>
</table>

Note: STS, superior temporal sulcus; MPFC, medial prefrontal cortex; SCZ, schizophrenia; ASD, autism spectrum disorder.

TD Vs Patients With Paranoid SCZ. For PInt>PhC, no difference was observed between groups. For CInt>PhC, the TD group compared to the SCZ group showed higher activation in the left pSTS and ventral MPFC (table 1 and figure 2A).

In order to test the hypothesis of “hyper-intentionality” in the SCZ group and to replicate our results, the mean activation per condition in both groups in vMPFC was extracted and analyzed externally in order to test by which differences the group by condition effects were driven. Note that we do not claim $P$-values for the interaction contrast itself to be a significant finding, as this would be double dipping—the interaction was found to be significant already in the ROI analyses on a FWE-corrected level in SPM. The TD group showed increasing activation from PhC < PInt < CInt. In contrast, SCZ showed similar activation for PhC and for both experimental conditions (PInt and CInt) (figure 2A). Comparing directly the control condition (PhC) in controls and SCZ in a post hoc $t$-test, a strong difference between groups was observed ($P = .002$). Similar results were found for left pSTS activation ($P = .001$). Moreover, the SCZ group correlation analysis revealed that PhC activation in vMPFC correlated positively with the PANSS total score ($r = .73, P = .01$) (figure 2A). To summarize, the SCZ group revealed dysfunctional activation in vMPFC and left pSTS and these dysfunctional activations were primarily done by an increased activation in the control condition (Ph-C). For the vMPFC this abnormal activation in Ph-C was positively associated with the degree of illness.

TD Vs Patients With ASD. No difference was observed between TD and ASD for the contrast PInt>PhC. Instead, decreasing neural activation for CInt>PhC in the right pSTS was observed for ASD (table 1 and figure 2B). In order to test the hypothesis of “hyper-intentionality” in the ASD group mean activation per condition in both groups in the right pSTS was extracted and analyzed externally. Comparing directly the CInt condition by a post-hoc $t$-test in controls and ASD, CInt showed less activation in ASD ($P = .0056$).

ASC Vs SCZ. No differential activation for the contrast PInt>PhC was observed. In the CInt>PhC contrast ASD showed more activation in dMPFC compared to SCZ. The inverse contrast (SCZ>ASD) did not reveal any difference (table 1 and figure 2B).

In order to test the hypothesis of “hypo-intentionality” and “hyper-intentionality,” mean activation per condition in the dMPFC ROIs of ASD and SCZ were extracted and analyzed externally. Similar as reported above, the respective plots indicate increasing significant activation in the PhC condition for the SCZ group ($P = .0021$).

fcMRI SEED-Analysis. Functional connectivity analyses with the right pSTS (the seed region) was performed
Fig. 2. Interaction of mentalizing (CInt>PhC) for TD>SCZ (A), TD>ASD (B), and ASD>SCZ (C). (A) Bar plots indicate effect sizes at the peak voxel in MPFC and left pSTS for all conditions in TD and SCZ. The amount of MPFC activation for PhC in the SCZ correlated positively with the total positive PANSS score ($r = 0.73$, $P = .001$). (B) Bar plots indicate effect sizes at the peak voxel in right pSTS for all conditions in TD and ASD. (C) Bar plots indicate effect sizes at the peak voxel in dMPFC for all conditions in ASD and SCZ. ROI analysis $P < .05$ FWE corrected ($k > 15$). Dashed rectangles indicate the beta parameters for PhC and CInt. Blue: TD; Red: ASD; and Green: SCZ.
to identify connected voxels whose activation showed a stronger response during CInt>PhC. Whole brain analysis revealed different regions of the social brain connected to the right pSTS that responded more strongly to CInt than PhC (table 2).

The control group showed increasing connectivity during CInt compared to PhC with vMPFC and dMPFC, bilateral TPJ, bilateral pSTS, right temporal pole, bilateral fusiform gyrus (FG), left supplemental motor area (SMA), and bilateral dorsolateral prefrontal cortex (DLPFC). The SCZ group exhibited functional coupling to fewer areas: left pSTS and temporal pole, right FG, right DLPFC, and left inferior frontal gyrus. In ASD the right pSTS showed increased connectivity with the bilateral TPJ, right FG and left pSTS. In the between-group comparison, group differences were found for control group vs SCZ in connectivity strength of the pSTS with vMPFC. During PhC the right pSTS was more strongly coupled with the vMPFC in patients with SCZ (figure 3A). In contrast, the ASD group showed less connectivity between the right pSTS and vMPFC, not in the physical but during the intentional condition (CInt) (figure 3B).

Table 2. Seed-Based (Right pSTS) Connection Table for CInt>PhC (Within and Between Analysis)

<table>
<thead>
<tr>
<th></th>
<th>x</th>
<th>y</th>
<th>z</th>
<th>Z</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Control</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>TPJ</td>
<td>66</td>
<td>−52</td>
<td>24</td>
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<tr>
<td></td>
<td>−54</td>
<td>−68</td>
<td>20</td>
<td>4.28</td>
</tr>
<tr>
<td>Temporal pole</td>
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<td>−14</td>
<td>−16</td>
<td>4.15</td>
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<tr>
<td>Posterior STS</td>
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<td>−36</td>
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<td>3.72</td>
</tr>
<tr>
<td></td>
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<tr>
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<tr>
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<td>−24</td>
<td>3.54</td>
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<td></td>
<td>−36</td>
<td>−46</td>
<td>−30</td>
<td>4.09</td>
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<tr>
<td>Dorso lateral prefrontal cortex</td>
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<td>30</td>
<td>3.87</td>
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<td>3.72</td>
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<td>Ventral MPFC</td>
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<td>3.37</td>
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<tr>
<td><strong>Control vs ASD</strong></td>
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<td>54</td>
<td>−10</td>
<td>3.81</td>
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Note: STS, superior temporal sulcus; MPFC, medial prefrontal cortex; SCZ, schizophrenia; ASD, autism spectrum disorder.

To summarize, the right pSTS was dysfunctional connected with the vMPFC in both patients group. In particular for the SCZ group the abnormal connection was primarily apparent by increased connectivity strength in the control condition (Ph-C), whereas for the ASD group the anomaly was primarily done by reduced connectivity strength in the communicative intention.

**Discussion**

This is the first study comparing SCZ and autism with regard to the neural signatures of their mindreading deficits using fMRI. In a task specifically designed for reading different kind of intentions, we observed group specific activation and connectivity patterns in and between key regions of the social brain in line with the hyper-hypo-intentionality hypothesis. These were apparent only during complex social interactive but not during simple mentalizing conditions.

At the behavioral level, the ASD group showed overall reduced accuracy, whereas the SCZ group showed overall increased reaction times. Other studies reported similar findings, namely no specific behavioral impairments but specific neural deviations. Since we only observed a significant main effect, but no interactions, we cannot state that the observed behavioral effects are specific to mind-reading. Rather, they may reflect additional difficulties in both patient groups beyond the ability to read intention. An alternative explanation could be that these results are the consequence of the different way of intention processing: hyper-intentionality leads SCZ to attribute intentions to objects, hypo-intentionality leads ASD to treat persons as objects. In both cases, the stance toward persons and objects could be altered in a specific way but the result may be the same: to handle persons and objects in a similar way.

At the neural level, the SCZ group showed aberrant activation in vMPFC and left pSTS (figure 2A), core regions of the mentalizing network. Moreover, this was due mainly to increased activation in the control condition. In line with the hyper-intentionality hypothesis, patients with paranoid SCZ seem to process information about physical processes on the neural level in a similar way as information about mental states. Such an “intention detector” may become hyper-active in paranoid thinking as it is characteristic for SCZ. These neural findings replicate our results reported in an independent patient group. and they are also consistent with behavioral evidence that demonstrated that SCZ attributed intentions to objects. Likewise confirming our results, in a recent study on action observation hyper-activation in the ToM-network (in pSTS and MPFC) was reported in patients with SCZ during a non-social condition. Furthermore, the fact that activation in the MPFC was correlated with the degree of positive symptoms lends some additional support to our interpretation and
Reading Intentions in Autism and Schizophrenia

confirms an association between paranoid symptoms and abnormal brain activation during reading of intent and agency.40,41 This neurocognitive bias could be related to an exaggerated attentional focus on single visual elements in an observed social interaction, resulting in an incorrect reading of the social interaction itself.42 An alternative explanation is related to the strong decoupling difficulties of paranoid patients, which preclude them from differentiating between mental states belonging to different agents.18 Further evidence comes from connectivity analyses of our data demonstrating that a core region of the mind-reading network, the right pSTS, showed stronger coupling to a second key region of the mind-reading network (vMPFC) compared to TD (figure 3A) again during the control condition. The ASD group also showed group differences in activation within the right pSTS when inferring communicative intentions (figure 2B). However, in contrast to the SCZ group, in ASD this difference was driven mainly by less activation during the experimental, ie, CInt condition. This finding is exactly what we predicted for ASD based on the hypo-intentionality hypothesis. Behavioural studies in ASD showed that mental states are often processed similar to physical states of objects.30 Also eye-tracking data during video-clips viewing showed that individuals with ASD did not focus on the interactive actors but on irrelevant physical information present on the scene.43 Deviant activation of pSTS in ASD subjects during social cognition is well documented (for review see)44 and abnormal activation in ASD subjects was reported when they observed geometric forms that elicit mind-reading21 (whereas SCZ reported opposite results). Its abnormal involvement is described also for a variety of indicators of social information processing such as eye gaze, language perception, and joint attention.22,45,46 An fMRI study revealed that ASD responded to faces similarly as to objects.47 Additional evidence for the hypo-intentional hypotheses comes from our connectivity findings with decreased connectivity between right pSTS and vMPFC in patients with ASD (figure 3B) in line with the literature supporting hypo-connectivity in ASD23,25: Hypo-connectivity might result from a lack of early social experiences due to disturbed early developmental mechanisms involved in social cognition.48

Fig. 3. Group × connectivity interaction. Connectivity was calculated using the right pSTS as a seed region (ANOVA second-level for CInt>PhC, \( P = .001 \) uncorrected, \( k > 15 \)). (A) Group differences in connectivity strength for TD (blue) versus SCZ (green) for PhC. (B) Group differences in connectivity strength between TD (blue) vs ASD (red) for CInt. Few behavioural studies on social cognition directly compared SCZ and ASD revealing inconsistent results.49–51 One behavioural study using a huge number of participants, found that ASD showed reduced perception of agency, whereas SCZ showed increasing perception of agency in human but also in other entities (animals and objects).52 Inconsistent results come also from brain volumetric studies: A meta-analysis study reported similar brain volumes between ASD and SCZ, but also clear distinctions.53 In 2 different EEG-studies using the same paradigm, ASD and SCZ were separately tested regarding their social-emotional abilities54,55 revealing overlapping but opposite results (reduced Mu suppression in ASD and larger Mu suppression in SCZ). To our knowledge, only one imaging study on social cognition, namely during processing of trustworthy faces, compared both groups directly: ASD and SCZ reported similar hypo-activation in right amygdala, FG and left ventro-lateral prefrontal cortex.56 The authors argued for “deficit specific” rather than disorder specific abnormality. In our study, the SCZ group seems to be more affected on the neural level compared to ASD (figure 2C). The available data from the literature make it hard to delineate which psychopathological syndrome should be more affected.
We speculate that the presence of positive symptoms in the SCZ group might elicit more disruption on reading intentions, and although a general mind-reading deficit is described in ASD, a specific intention reading deficit in ASD is not well documented.

Clearly, there are several limitations to this study. A subgroup of the patients with SCZ and 1 patient with ASD were treated with atypical neuroleptics. Therefore, we cannot exclude that medication effects interfere with the interpretation of our findings. Additionally, the age range of our sample was rather large: Participants with SCZ were older than TD and ASD participants. Although age effects were partialled-out in all analyses, confounding influences of age on group comparability cannot definitely be excluded. Taken these limitations into account, we conclude that group differences in neural activation during intention reading in ASD and SCZ depend on the type of intention (communicative or private). Moreover, the group differences in brain activation and the results of our connectivity analyses are consistent with the idea that mind-reading problems in SCZ are due to an overactive intention-detection module, whereas mind-reading problems in ASD are rather due to an underactive intention-detector module.

Supplementary Material

Supplementary material is available at http://schizophreniabulletin.oxfordjournals.org.

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