

Taking it at ‘face value’: the use of face processing strategies in bipolar disorder and schizophrenia

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Abstract

Objective: Use of appropriate face processing strategies is important for facial emotion recognition, which is known to be impaired in schizophrenia (SZ) and bipolar disorder (BD). There is preliminary evidence of abnormalities in the use of face processing strategies in the former, but there has been no explicit attempt to assess face processing in patients with BD.

Method: 28 BD I, 28 SZ and 28 healthy control participants completed tasks assessing featural and configural face processing. The facial inversion effect was used as a proxy of second order configural face processing and compared to featural face processing performance (which is known to be relatively less affected by facial inversion).

Results: Controls demonstrated the usual second-order inversion pattern. In the BD group, the absence of a second-order configural inversion effect in the presence of a disproportionate bias toward a featural inversion effect was evident. Despite reduced accuracy performance in the SZ group compared to controls, this group unexpectedly showed a normal second-order configural accuracy inversion pattern. This was in the context of a reverse inversion effect for response latency, suggesting a speed-vs-accuracy trade-off.

Conclusions: To our knowledge, this is the first study to explicitly examine and contrast face processing in BD and SZ. Our findings indicate a generalised impairment on face processing tasks in SZ, and the presence of a second-order configural face processing impairment in BD. It is possible that these face processing impairments represent a catalyst for the facial emotion recognition deficits that are commonly reported in the literature.

Keywords: facial emotion recognition; emotion processing; psychosis; euthymia, bipolar I, cognition; neuropsychology; mental illness; mood disorder

Bipolar disorder (BD) and schizophrenia (SZ) are complex mental disorders characterized by poor psychosocial functioning (Van Rheenen & Rossell, 2014a) and impaired cognition extending across both social and non-social domains (Gogos, Joshua, & Rossell, 2010; Rossell, Van Rheenen, Joshua, O'Regan, & Gogos, 2014; Rossell & Van Rheenen, 2013; Van Rheenen & Rossell, 2013b, 2014b, 2014c; Van Rheenen & Rossell, 2014d). There is a growing literature suggesting that these latter deficits are strongly predictive of the former; with impairments in facial emotion recognition in particular, often cited as a potentially important contributing factor for impaired interpersonal functioning (Brekke, Kay, Lee, & Green, 2005; Kee, Green, Mintz, & Brekke, 2003). In the SZ literature, there have been some attempts to determine the underlying mechanisms associated with these emotion recognition aberrations, with findings pointing toward a potential role for general cognitive ability as well as perceptual face processing per se (Fakra, Jouve, Guillaume, Azorin, & Blin, 2015; Joshua & Rossell, 2009; Kohler, Bilker, Hagendoorn, Gur, & Gur, 2000; Sergi et al., 2007). In the BD literature however, there has been far less attention focused on these lines of enquiry (Van Rheenen, Meyer, & Rossell, 2014).

Sufficient processing of visual information and the use of appropriate face processing strategies are a necessary pre-requisite for intact facial emotion recognition. In SZ, impaired performance on face processing tasks suggests that a failure of these prerequisites may at least partially account for some of the facial emotion recognition impairments commonly observed in the disorder (Bortolon, Capdevielle, & Raffard, 2015; Joshua & Rossell, 2009; Rossell et al., 2014; Shin et al., 2008). In BD, no studies have comprehensively investigated the use of typical face processing strategies in and of themselves, although there have been some attempts to address the influence of basic face processing ability in emotion recognition studies in BD (Addington & Addington, 1998; Getz, Shear, & Strakowski, 2003; Van Rheenen & Rossell, 2013a). This generally occurs in the context of control tasks that require the discrimination of gender or identity. Such tasks, when used as a proxy for face processing ability, are limited

since the discrimination of faces can be accomplished on the basis of matching local featural information (such as the eyebrows or the nose) in a piecemeal fashion (Duchaine & Weidenfeld, 2003). Yet a large body of research shows that the use of top-down processing strategies that incorporate facial information more globally, may be an even more important means of face recognition than the processing of isolated featural information (Tanaka & Farah, 1993). Thus, normal performance on gender or identity discrimination tasks does not necessarily imply normal face processing ability. As a result intact performance on such tasks is not sufficient to conclude intact face processing in BD generally (Duchaine & Weidenfeld, 2003).

Typically, face processing requires the initial identification of a normal facial configuration represented by two eyes, above a nose, above a mouth. As all faces share this same *first-order* relationship, face perception also relies on the processing of spatial relationships between elements in the face, including the distances between local features (Maurer, 2002). This *second-order configural processing* is an important top-down perceptual skill for distinguishing identities, but also for the appropriate identification of facial expressions that reflect subtle muscular changes in the spatial positioning and relationships between local features (Bombari et al., 2013; Derntl, Seidel, Kainz, & Carbon, 2009; Fakra et al., 2015).

A known proxy for the use of configural strategies is the *facial inversion effect*, which describes performance impairments in the processing of faces when they are inverted by 180° (Renzi et al., 2013). Given that this effect is substantially less pronounced when faces are digitally manipulated to express changes in featural rather than spatial information, the extent of disruption for the processing of inverted compared to the upright faces is widely recognised as representing the extent to which second-order configural face processing strategies are relied on during face recognition (Maurer, 2002). Indeed, significant research shows a differential effect in which there is an absence or smaller inversion effect on featural compared to second-order configural processing tasks in the healthy population (Freire, Lee, & Symons, 2000; Leder & Bruce, 2000).

In patients with SZ, there is evidence that shows that there is a reduction in susceptibility to this facial inversion effect compared to controls (Shin et al., 2008). The implications that this has for emotion processing has been demonstrated recently, in a study indicating that a smaller inversion effect reduces the capacity for accurate identification of emotional expressions in this disorder (Fakra et al., 2015). Given increasing evidence that BD patients have similar, albeit less severe cognitive deficits in emotion recognition as their SZ counterparts (Van Rheenen & Rossell, 2014c), it is possible that the configural processing deficits that have been previously found in SZ extend to BD as well.

To explore this notion we aimed to investigate the use of configural face processing strategies in BD; the first study of its kind to our knowledge. Here, we examined performance differences in a sample of individuals with BD and SZ compared to controls. We expected that in controls, upright faces would be processed more accurately and efficiently than inverted faces, but that this effect would be more pronounced for manipulations of second-order relationships than local featural elements. Given evidence of impaired configural processing in SZ alongside preferential processing of local (compared to global) information (Schwartz Place & Gilmore, 1980; Silverstein et al., 2006; Wells & Leventhal, 1984), we expected that the SZ group would be less susceptible to the configural inversion effect than controls, with BD patients performance falling intermediate to the two other groups. The relative strength of the configural over featural inversion effect in the BD group and the effect of current symptomatology on face processing in the clinical groups generally, remained open questions.

Method

This study was approved by the relevant Hospital and University review boards and abided by the Declaration of Helsinki. Written informed consent was obtained from each participant before the study began.

Participants

The clinical sample comprised 28 individuals with BD-I and 28 individuals with SZ.

Patients were recruited via community support groups and community care units and were all out-patients. Diagnosis was ascertained using the Structured Clinical Interview for DSM-IV (SCID)(First, Spitzer, Gibbon, & Williams, 1996). Current symptomology was acquired using the Beck Depression Inventory (BDI) (Beck & Steer, 1987) and the Positive and Negative Syndrome Scale (PANSS) (Kay, Fiszbein, & Opler, 1987): global positive, negative, general and composite scores were calculated. All BD patients were tested during a period of clinical stability (i.e., not currently meeting criteria for a mood or psychotic episode) as determined by the investigators via their SCID interview, however 12 BD individuals were considered to be depressed on the basis of their BDI score (≤ 10). Sixteen SZ patients were considered to have current psychotic symptoms (PANSS P1 and/or P3 scores > 3). None of the patients included in the sample experienced any co-morbid Axis 1 diagnoses at the time of testing.

A sample of 28 healthy control participants was recruited via newspaper advertisements. Control participants were excluded if they had any history of psychiatric disorder or a first degree relative with SZ, BD or Schizoaffective Disorder (based on the SCID). In addition, participants from all three groups met the following criteria: a) No history of neurological disorder or head trauma, b) No diagnosable current drug or alcohol abuse disorder and no illicit drug use in the previous 24 hours, c) English spoken as first language, d) Aged between 18-65 years and e) Estimated premorbid IQ > 85 based on the National Adult Reading Test (NART) (Nelson & Willison, 1991), f) no ECT in past 12 months

Within the BD group, one patient was taking antipsychotic medication alone (atypical), four were taking mood stabilisers alone, six were taking antipsychotics (all atypical) and mood stabilisers, six were taking antidepressants and mood stabilisers, two were taking antipsychotics (all atypical), antidepressants and mood stabilisers, one was taking an antidepressant and an antipsychotic (atypical), one was taking an antidepressant and a benzodiazepine, one was taking

a mood stabiliser and a sedative, one was taking an antipsychotic (atypical), mood stabiliser and a benzodiazepine, one was taking an antipsychotic (atypical), mood stabiliser and an anti-cholinergic and three were medication free.

Within the SZ group, 15 patients were taking antipsychotic medication alone (one typical, 14 atypical, two combination), four were taking antipsychotics (four atypical, one combination) and an antidepressant, one was taking an antipsychotic (atypical) and a mood stabiliser, one was taking a combination of an antipsychotic (atypical), an antidepressant and a mood stabiliser, one was taking an antipsychotic (atypical), antidepressant and a benzodiazepine, three were taking antipsychotics (atypical) and benzodiazepines, two were taking antipsychotics (atypical), benzodiazepines and anti-cholinergic and one was medication free. Group averages of chlorpromazine equivalents (CPZ) are given in Table 1.

[INSERT TABLE 1 ABOUT HERE]

Materials

*Featural and second-order configural face processing*¹. Featural and spacing manipulation sets were used to assess the facial inversion effect after featural and configural change. Both tasks had the same design and face stimuli, however differed in the type of manipulation completed. The *featural set* manipulated the features of the face while maintaining consistent configural information. The *spacing set* manipulated the spatial distances between features, that is second-order configural face information. The two stimuli sets were created using gray-scaled emotionally neutral faces from the Pictures of Facial Affect series (Ekman & Friesen, 1976). One white adult female face was selected as the template face and digital manipulations were made in line with previous work (Freire et al., 2000). For the featural set, the eyes, nose and mouth were selected from four other female faces and digitally pasted over the features in the

¹ All participants also completed a first-order configural processing task. A description of this task and the results of its analysis are presented in the supplementary materials.

template face. Thus, four manipulations in addition to the template face were created, resulting in five distinct face images. The eye-nose-mouth replacement features did not differ in brightness or contrast to the template face. Further, they were selected to closely match the iris size and width of the nose and mouth to that of the template face. This clearly altered the featural information within the original template face, while maintaining the configural information as well as external features. While featural manipulation may result in slight alterations in the configural information, care was taken to minimise these changes.

For the spacing set, the eyes (and eyebrows) were either moved horizontally in or out by 2 pixels, or the eyes and mouth were moved vertically down or up by 3 pixels. Thus, like the featural task, four manipulations in addition to the template face were created, resulting in five distinct face images. This disrupted the second-order configural information within the template face, while maintaining the featural and first-order information. All face images were 320 x 480 pixels in size. During stimuli editing, the blurring tool was used to maintain continuity of skin shade. Therefore, all face manipulations were subtle, resulting in normal looking faces with careful attempts made to avoid distinctiveness or grotesqueness. Figure 1 presents an example of the task stimuli.

[INSERT FIGURE 1 ABOUT HERE]

Face pairs were presented on the screen for eight seconds, followed by a fixation cross (+) for 500ms in between each trial. Participants were required to determine if the two faces presented were the same or different. This discrimination design was selected to minimise the memory demand apparent in other similar tasks (LeGrand, Mondloch, Maurer, & Brent, 2001). Responses were made via a two-button press. Participants were instructed to respond as quickly yet as accurately as possible to indicate whether the faces were the ‘same’ or ‘different’. As soon as the participant responded, the task progressed to the next trial. The buttons were labelled so participants would not forget which button was which.

After reading the instructions, participants completed three practice trials for each task.

Thereafter, for both tasks, each of the five face images was paired with itself eight times and each other twice (once to the left, once to the right), creating a total of 80 face pairs, thus half the face pairs showed identical (same) faces and the other half showed different faces. Each face pair was presented in upright and inverted orientation, thus participants completed 160 randomised trials for each task. The order of task completion was counterbalanced.

Statistical analysis

Demographic and clinical group differences were assessed via Chi-Square tests or one-way between-groups analysis of variance (ANOVA) with post-hoc Fisher's LSD tests. Given group differences in premorbid IQ and gender, a validity check was performed to examine associations between these demographic variables and accuracy, response time and difference scores (i.e., upright-inverted performance scores on the featural and spacing tasks) on the face processing tasks using Pearson's/Spearman's correlations (alpha set at a conservative $p < .01$ to correct for multiple testing) in each of the three groups. As neither of the variables significantly correlated with these demographic variables in any of the groups, we did not covary for them in subsequent analyses.

As the featural and spacing tasks shared the same design, we incorporated both tasks into a two (task; featural or spacing)*two (orientation; upright, inverted) * three (group; controls, BD, SZ) repeated measures ANOVA with post-hoc Fisher's LSD tests to investigate differential effects of featural and configural manipulation on performance. Follow up paired samples t-tests separated by group were used to compare performance on upright and inverted conditions as well as the inversion difference scores in both spacing and featural tasks to each other. Independent samples t-tests comparing each group to each other were used to compare the inversion difference score (i.e., performance in the upright minus the inverted condition) on the spacing task. Effect sizes were calculated in Cohen's d and for clarity, are only reported in positive form in text. Pearson's correlations were used to examine associations between clinical symptomatology and task

performance in the clinical groups. All post-hoc/follow-up tests and correlations were corrected using a conservative α of $p<.01$.

Results

Demographics

As can be seen in Table 1, the SZ patients had a significantly lower premorbid IQ than controls. PANSS and BDI ratings in SZ patients were significantly higher than for BD patients. SZ patients were also on a higher average dose of antipsychotics in CPZ equivalents. There were no group differences between SZ and BD groups in terms of age, age of illness onset or illness duration.

Featural and spacing task analysis

Accuracy: Figures 2a and 2b show accuracy performance for the spacing and featural tasks across groups. There was a main effect of task ($F(1,81)=145.11$, $p<.001$) and orientation ($F(1,81) = 68.151$, $p<.001$) with all participants performing better in the featural ($M=84.72$, $SD=13.44$) compared to the spacing ($M=69.78$, $SD=13.83$) task ($d=1.24$), as well as in the upright ($M=80.57$, $SD=12.75$) compared to the inverted ($M=73.93$, $SD=13.07$) orientations ($d=0.51$). A main effect of group ($F(2,81)=21.61$, $p<.001$) indicated that SZ patients ($M=67.78$, $SD=11.29$) performed worse than controls ($M=85.37$, $SD=7.03$) overall ($p<.001$, $d=1.87$), with BD patients ($M=78.60$, $SD=11.36$) performing significantly better than the former group ($p<.001$, $d=0.96$) and worse than the latter ($p<.01$, $d=0.72$).

[INSERT FIGURE 2 ABOUT HERE]

There were trends for an orientation*group ($F(2,81)=2.61$, $p=.08$) interaction. Significant task*group ($F(2,81) = 3.86$, $p=.03$) and task*orientation*group ($F(2,81)=4.86$, $p<.01$) interactions were also present. Follow up analysis revealed that the SZ group showed an inversion effect on both the spacing ($t(27)=5.63$, $p<.001$) and the featural tasks ($t(27)=2.62$, $p=.01$). Similarly, the control

group also showed a significant inversion effect on the spacing task ($t(27)=5.03$, $p<.001$) and the featural task ($t(27)=4.36$, $p<.001$). In contrast, the BD group showed a significant inversion effect on the featural task ($t(27)=4.72$, $p<.001$) but not the spacing task ($t(27)=1.48$, $p=.15$). There was a trend for a significantly stronger inversion effect (ie, the difference score) for the spacing compared to the featural task for SZ patients ($t(27)=2.33$, $p=.03$) only (see Figure 3a).

Comparison of the spacing inversion effects across groups indicated that BD patients were significantly less susceptible to the effect relative to controls ($t(54)=2.92$, $p<.001$, $d=0.78$) and SZ patients ($t(54)=3.77$, $p<.001$, $d=0.86$). There were no differences in the strength of the spacing inversion effect between the SZ and control groups (all p 's $>.05$). Figure 4a presents a graphical representation of these difference scores across groups.

[INSERT FIGURE 3 ABOUT HERE]

Response time. Figures 2c and 2d shows response time performance for the two tasks across groups. There was a main effect of task ($F(1,80)=14.17$, $p<.001$) and orientation ($F(1,80)=5.34$, $p<.01$) with all participants responding faster in the featural ($M=2151.98$, $SD=466.16$) compared to the spacing ($M=2328.68$, $SD=550.19$) task ($d=0.35$), as well as in the upright ($M=2199.34$, $SD=438.47$) compared to the inverted ($M=2281.8152$, $SD=540.85$) orientations ($d=0.08$). A main effect of group ($F(2,80)=4.04$, $p=.02$) indicated that SZ patients ($M=2075.28$, $SD=498.52$) had shorter latencies compared to BD patients ($M=2415.95$, $SD=433.39$) overall ($p<.01$, $d=0.73$). Although SZ patients were faster than controls ($M=2225.14$, $SD=402.10$, $d=0.30$) and BD patients were slower overall, neither difference was significant (both p 's $>.05$; C/BD $d=0.41$).

Significant task*group ($F(2,80)=4.52$, $p<.01$), orientation*group ($F(2,80)=10.13$, $p<.001$), task*orientation ($F(1,80)=19.55$, $p<.001$) and task*orientation*group ($F(2,80)=4.10$, $p=.02$) interaction effects were also present. Follow up analyses indicated that on the featural task, both the BD ($t(27)=-4.61$, $p<.001$, $d=0.45$) and control ($t(27)=-6.90$, $p<.001$, $d=0.58$) groups showed an

inversion effect by performing significantly faster in the upright compared to the inverted condition. A featural inversion effect was not apparent in the SZ group however (i.e., performance did not significantly differ across conditions ($t(27) = -.37$, $p=.72$, $d=0.05$). On the spacing task the control group showed the normal inversion effect ($t(27)=-5.75$, $p<.001$). As performance speed did not significantly differ across upright and inverted conditions in the BD group ($t(27) = -.44$, $p=.66$), a spacing inversion effect was not apparent. The SZ group on the other hand showed a trend for a reverse inversion pattern, responding faster in the inverted compared to the upright spacing condition ($t(26)=2.36$, $p=.03$).

Analyses of the extent of the inversion effects across the featural and spacing tasks indicated a larger difference score for the former compared to the latter ($t(27)=-4.25$, $p<.001$) in the BD group, while the difference score was larger for the latter compared to the former in the SZ group ($t(26) = -2.78$, $p\leq .01$). There was no significant difference in the strength of the inversion effect on the featural and spacing tasks in the control group ($t(27)=.47$, $p=.64$. Figure 3b provides a graphical representation of effect sizes of the difference scores between these two tasks for each of the three groups.

Comparison of the spacing inversion effect across groups indicated that BD patients were significantly less susceptible to the inversion effect relative to controls ($t(54)=2.81$, $p<.01$, $d=0.75$). Significant differences in the strength of the effect were evident for the SZ group compared to controls ($t(38.57)=4.66$, $p<.001$, $d=1.26$), while a trend-level difference was also evident comparing the two clinical groups ($t(53)=-2.12$, $p=.04$, $d=0.57$; see Figure 4b).

[INSERT FIGURE 4 ABOUT HERE]

Post-hoc analysis

Given that the mean accuracy performance on the spacing task in the SZ group was just above chance level, we re-ran all within-group analysis in the SZ group removing patients whose spacing inversion error rate was above 50% ($n=9$). This was to further understand whether the

unexpected inversion effect in this group was related to the particularly poor performance on the inverted condition in some SZ individuals. However this analysis yielded no difference to the response time or accuracy results.

Correlations with symptom scores

There were no correlations between any of the variables of interest and symptomatology scores on the PANSS or BDI in any of the groups

Discussion

Despite the potential for impaired perceptual face processing to contribute to the facial emotion recognition deficits that are increasingly replicated in BD and SZ research, there has been limited attention focused on the extent to which typical face-processing strategies are used in patients with these disorders; particularly BD. Here we examined a group of individuals with BD and SZ compared to controls, in the first study to examine and contrast featural and configural face processing in these disorders.

In terms of control performance, we were able to replicate the normal inversion effect for the second-order configural processing task (Freire et al., 2000), which suggests that healthy individuals do process upright faces more efficiently than inverted faces. Further, mean accuracy inversion differences between the spacing and featural tasks indicated that faces that had been featurally manipulated were not as vulnerable to this effect compared to faces that had been configurally manipulated. Although this difference was not significant, the effect was in the medium size range. Taken together, this pattern of results speaks to claims that face inversion disrupts configural processing more so than featural processing, suggesting that there may be different mechanisms involved in these two face-processing strategies, at least in part (Leder & Bruce, 2000; Schwaninger, Lobmaier, Wallraven, & Collishaw, 2009).

The most prominent finding of the current study was the absence of an inversion effect on the measure of second-order configural processing in BD, where the strength of the inversion effect was substantially lesser than that found in the SZ and control groups. This, coupled with an abnormal preferential inversion effect for the featural task over the spacing task suggests that in the BD group, there is impairment in the processing of second-order configural face information in the context of a disproportionate reliance on featural face processing. This is further supported by an absence of group differences in the strength of the inversion effect on the featural processing measure in BD patients compared to controls, together with indications that the lack of configural inversion effect in this group was likely attributable to accuracy impairments (relative to controls) for the processing of upright faces - which are primarily reliant on second-order configural processing. This is as opposed to improved performance (relative to controls) for inverted faces - which are reliant on featural or ‘object’ information (see Figure 2a for a visual comparison of performance in both conditions between BD versus controls.).

Further, a reverse inversion pattern was evident for some individuals with BD (see negative error bar in Figure 4a for example), which suggests that these individuals processed inverted faces *more efficiently* than upright faces. This pattern certainly speaks to the assertion that inverted faces activate a more general visual processing mechanism i.e., one that processes isolated facial features or ‘objects’; in these BD individuals this mechanism appears to be intact and unable to compensate for aberrant processing of upright configurally manipulated faces (de Gelder, Bachoud-Lévi, & Degos, 1998; Farah, Wilson, Maxwell Drain, & Tanaka, 1995). In combination these findings are consistent with the theory that inversion increases dependence on featural processing at the expense of configural processing efficiency (Maurer, 2002), and suggest that featural *but not configural* face processing is intact in this group.

The prediction that the SZ group would be less susceptible to the facial inversion effect compared to controls was not supported by the current accuracy findings. This is in direct opposition to a number of recent studies that have demonstrated disturbed second-order configural

processing in this disorder (Fakra et al., 2015; Joshua & Rossell, 2009; Shin et al., 2008). The response time findings for the SZ group did however, support an argument for disturbed second-order configural processing, although this was unexpectedly revealed in the context of a reverse inversion pattern. Therefore it is suggested that the lack of a significant accuracy inversion effect seen here, likely reflects the product of a speed versus accuracy trade-off in patients with SZ.

Indeed, a main effect of orientation indicated that inverted faces were more difficult to process overall. On the second-order configural task this effect appeared to be even more pronounced in the SZ group, such that mean-level performance in SZ patients (versus controls) was lower in both the upright and inverted conditions, but response latencies were substantially increased for the latter condition only. This effect remained when participants performing at chance level on the inverted condition were removed. In light of known deficits in speed of processing and psychomotor behaviour in SZ, this unusually increased response time in the presence of a pronounced accuracy error rate in the inverted spacing condition, potentially indicates a lack of effortful responding in this group in this arguably more challenging condition; it is certainly plausible that had this impulsive responding been normalised or decreased compared to controls as usually occurs in patients with SZ, performance in the inverted condition would have improved. Consequently, this would likely have reduced the extent of the inversion related differences between the two orientation conditions, which would be in line with the existing literature.

While this suggestion is purely speculative, the growing literature indicating a performance preference for the processing of local over more global top-down facial information in SZ patients does speak to the contention that a speed versus-accuracy trade off masked an observation of reduced susceptibility to the facial inversion effect in SZ here (Chen, Nakayama, Levy, Matthysse, & Holzman, 2003; Johnson, Lowery, Kohler, & Turetsky, 2005). Regardless, it is unlikely that any alterations in response latency would have been enough to compensate entirely for aberrant accuracy performance in this group, since SZ patients consistently show impaired group-level performance on perceptual and emotion recognition tasks in general (Johnson et al., 2005; Rossell

et al., 2014; Rossell, Van Rheenen, Groot, Gogos, & Joshua, 2013). Consistent with this, worse performance in SZ across both the spacing and featural tasks in general suggests that abnormalities in face specific (upright - configural) *and* more general visual processing mechanisms (inversion-featural) contribute to abnormalities on face processing tasks in SZ.

In sum, our results indicate significant accuracy impairments on measures of face processing in SZ and highlight impairment in the use of second-order configural face processing strategies in BD compared to controls. However, despite the novelty of these latter findings, our results should still be interpreted within the confines of some limitations. Firstly, although based on existing measures, the facial processing tasks were newly developed in our lab and have not been validated in other clinical samples. This notwithstanding, ceiling performance was not evident on either the featural or spacing tasks, and both tasks did reliably reveal group differences and support the facial inversion pattern expected in controls. In future studies we aim to replicate these findings with the same methodology. Secondly, given that patients were on different medications, we could not adequately control for medication effects. Although we found no significant bivariate correlations between CPZ equivalent scores and task performance, these scores only account for a subclass of the medications in use in the sample. Thus, it remains possible that other medications may have still had an influence on performance. Thirdly, there was a gender imbalance across participants from each of the three groups. Although provisions were made to explore gender as an influential factor within initial correlation analyses, future investigations should attempt to include equal numbers of males and females across groups. Finally, it should be noted that the BD and SZ groups had significant differences in their positive, negative and general symptomatology scores, with many of the patients with SZ demonstrating moderate - severe psychotic symptoms. Although there were no correlations between PANSS scores and face processing performance in either the BD or SZ groups, it remains possible that differences in the clinical state of individuals in both groups partially affected the results.

Despite these limitations, our novel study does shed light on the use of typical face processing strategies in BD and SZ, suggesting impairment in second-order configural processing for BD and a more generalised impairment in the processing of faces for SZ. Importantly, configural face processing enables the distinction of individual facial identities and also likely aids in facial expression recognition (Bombari et al., 2013; Derntl et al., 2009; Fakra et al., 2015). It is therefore possible that emotion recognition impairments in some individuals with BD and SZ may be related to subtle deficits in the processing of the spacing and distances between local features. Future studies would do well to examine this further.

Financial Support and Acknowledgements

This work was supported by an NHMRC early career fellowship (TEVR), a Menzies Foundation Scholarship (NJ) and the Harold and Cora Brennan Trust Grant as well as the Marian & E.H. Flack Trust

Conflicts of interest

The authors declare that they have no conflicts of interest

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Table 1. *Demographics and clinical characteristics of the sample*

	Controls	BD		SZ		Group comparisons	Post-hoc comparisons
		M	SD	M	SD		
Age		42.43	11.29	41.96	11.35		
Gender (Male/Female)	13/15		9/19		21/7		
Premorbid IQ		113.96	7.57	109.14	9.88	106.46	11.09
BDI		2.57	3.10	9.29	9.60	14.32	11.53
CPZe		-	-	79.02	126.56	482.22	296.91
Age of illness onset			-	22.07	9.54	22.78	6.73
Illness duration		-	-	19.93	11.26	17.00	10.65
PANSS P		-	-	10.07	3.11	13.07	4.52
PANSS N		-	-	8.82	2.00	12.89	5.97
PANSS G		-	-	20.79	3.07	24.93	6.66
PANNS COMPOSITE (P+N+G)		-	-	39.68	5.30	50.54	14.03

CPZe = chlorpromazine equivalents; PANSS = Positive and Negative Syndrome Scale



Figure 1. Featural Manipulation (top) and Spacing Manipulation (bottom).

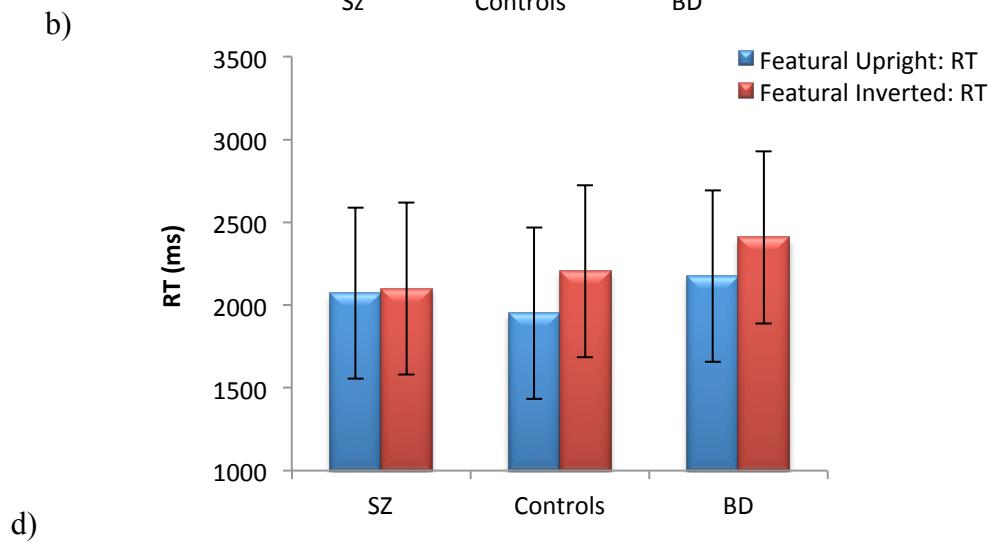
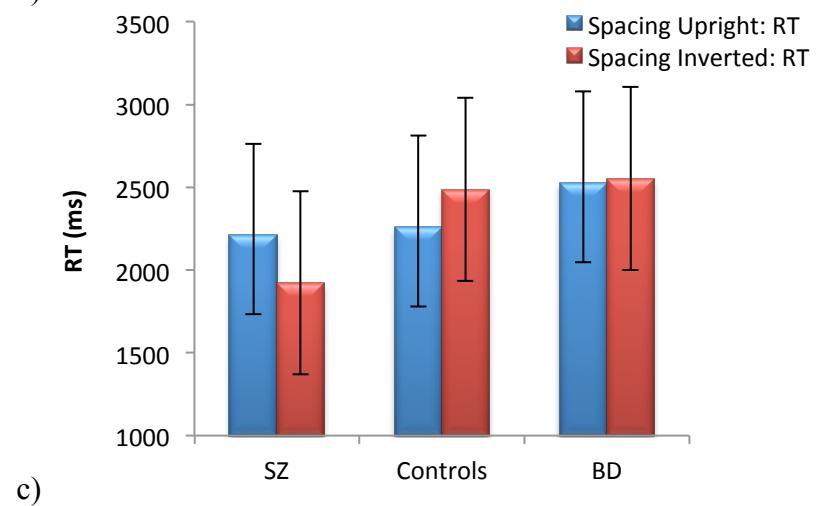
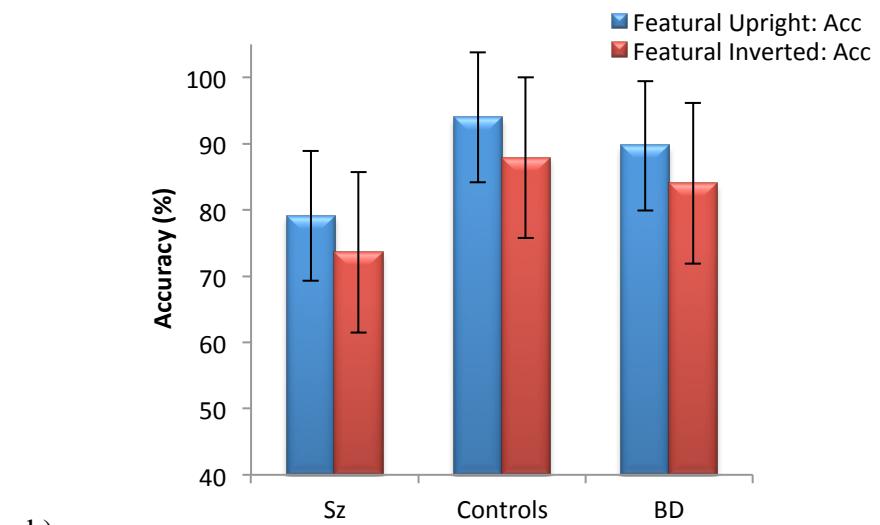
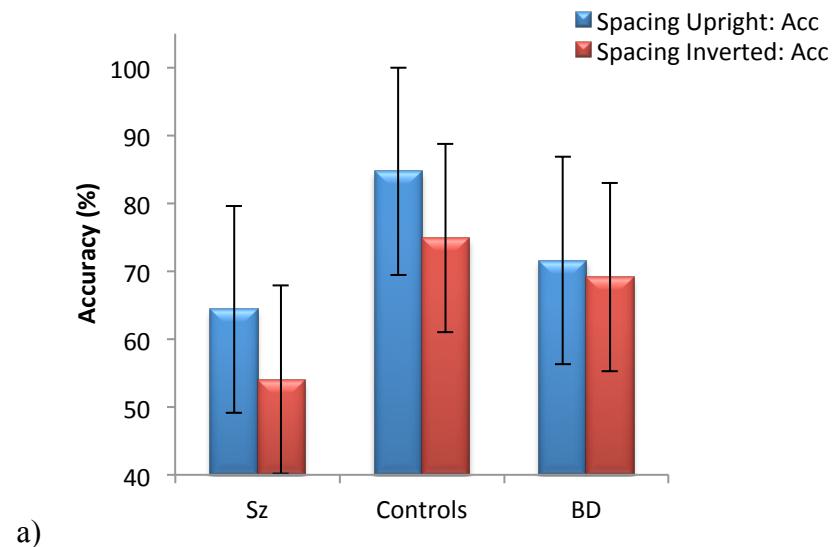


Figure 2. Group performance on the Spacing and Featural Tasks. a) Accuracy - Spacing Task. b) Accuracy - Featural Task. c) Response time - Spacing Task. d) Response time - Featural Task. Error bars represent standard deviations. Acc= accuracy; RT = response time

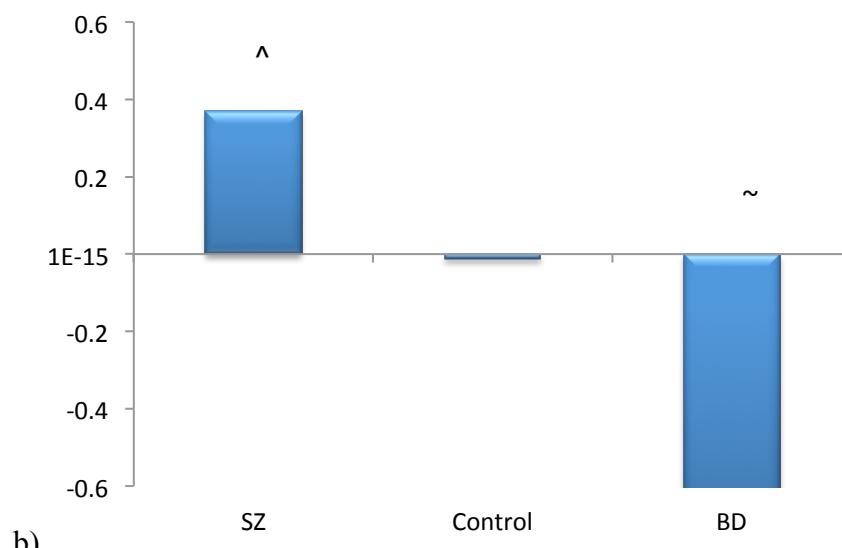
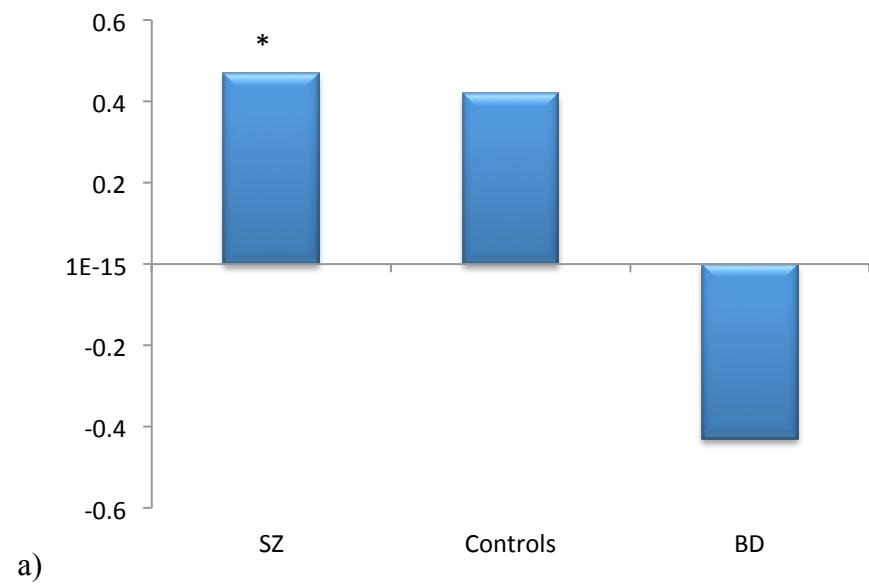


Figure 3. Effect size differences between the Featural and Spacing inversion effects (difference scores) across groups. a) Accuracy and b) response time; * $p < .01$ larger inversion effect for Spacing over Featural Task; ~ $p < .01$ reverse inversion bias favouring the Featural over Spacing Task; ^ $p < .01$ larger reverse Spacing inversion effect compared to normal Featural inversion effect in SZ; note that although it appears that the normal Spacing over Featural inversion effect is evident in the SZ group, the Spacing inversion effect, albeit larger compared to the Featural inversion effect, was in reverse. Note larger inversion effects in the Featural compared to the Spacing Task are represented as negative. Error bars represent standard deviations

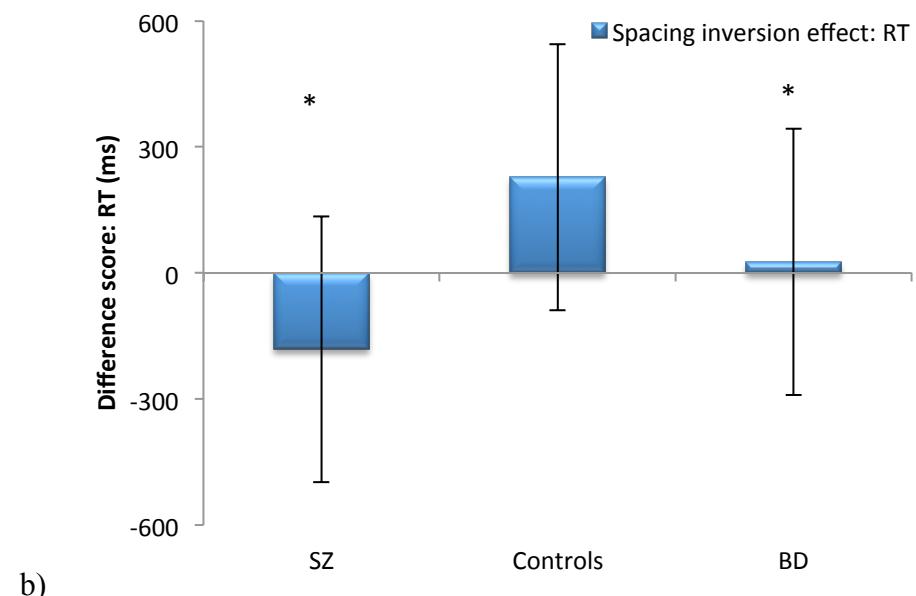
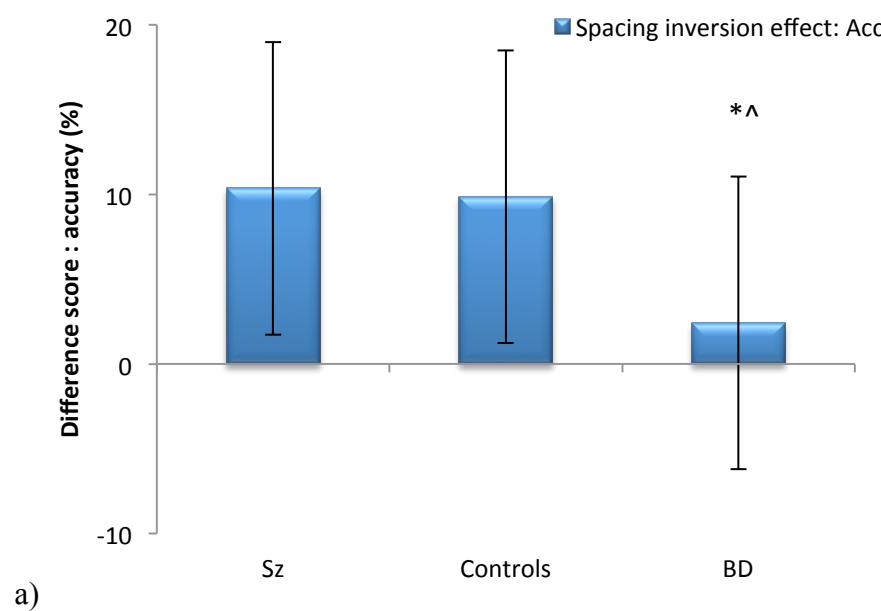


Figure 4. Difference scores (i.e., extent of inversion effect) on the Spacing Task per group; a) accuracy difference score and b) response time difference score. Note the lack of inversion effect on the Spacing Task for both accuracy and response time in the BD group. * $p<.01$ compared to controls; $^p<.01$ between clinical groups. Note the opposite response time effect in the SZ group. Error bars represent standard deviation. Acc= accuracy; RT = response time.

First order configural face processing task

A *Scrambled Faces Task* was used to assess first-order configural face processing. This task was designed to disrupt first-order information, while maintaining featural and second-order configural information. Twelve different gray-scaled emotionally neutral faces (six males, six females) from the Pictures of Facial Affect series (Ekman & Friesen, 1976) were used. Each face was presented as either normal, that is, with no alteration or disruption ('face' condition), or scrambled ('scrambled-face' condition). Scrambled faces involved a disruption of the first-order relations. Images were edited such that the normal first-order configuration of the eyes above the nose above the mouth was changed. This method was adapted from previous research on this type of face processing (1994). Photo editing software was used to capture the eyes region or the combined nose and mouth region of the face. The locations of the two regions were then switched to preserve the featural information and the specific second-order configural information, i.e. the features remained the same shape and distance apart, however, were reordered in terms of location. The blurring tool was used to maintain continuity of skin shade. New 'scrambled-faces' showed the nose above the mouth above the eyes. Images were 290 x 390 pixels in size. Supplementary Figure 1 presents an example of the task stimuli.

Participants were presented with a randomised sequence of normal or scrambled faces. Each face was presented in the centre of the screen for 2000ms seconds followed by a fixation cross (+) for 1500ms. Participants were told they were going to see a picture on the screen and they were to decide if the picture was a real face or not. Via a two-button press participants were instructed to respond as quickly and accurately as possible to indicate whether the picture was either a 'face' or a 'non-face'. After reading the instructions, participants completed five practice trials followed by 72 randomised

experimental trials in total. Each face and associated scrambled face was presented three times (36 ‘faces’ and 36 ‘non-faces’). The task took approximately four and a half minutes to complete.



Figure 1. Face (left) and Scrambled Face (right)

Statistical analysis

A two (condition; face, scrambled face) * three (group; controls, BD, SZ) repeated measures ANOVA with post-hoc LSD tests was conducted to investigate performance on the Scrambled Faces Task across groups.

Results

Supplementary Table 1 presents the accuracy and response time scores on the Scrambled Faces Task per group.

Table 2. Accuracy and response time performance on the Scrambled Faces Task per group

		Controls		BD		SZ	
		M	SD	M	SD	M	SD
Accuracy (%)	Overall	98.76	1.33	98.41	2.24	94.39	10.09
	Face	99.11	1.52	97.94	2.73	93.85	13.64
Response time (ms)	Scrambled face	98.41	2.06	98.87	2.21	94.94	8.35
	Overall	684.89	112.57	715.16	123.09	792.08	208.67
	Face	690.71	120.92	730.34	136.92	793.36	229.42
	Scrambled face	679.08	115.85	699.97	119.48	790.80	197.97

BD= bipolar disorder; SZ=schizophrenia

Accuracy. There was no significant main effect of condition ($F(1,80)=.42$, $p=.52$) and no condition by group interaction ($F(2,80) = .70$, $p=.50$) but there was a group effect ($F(2,80)=4.48$, $p=.01$), with SZ patients performing less accurately than controls ($p=.03$) and BD patients ($p=.05$) overall.

Response time. There was no main effect of condition ($F(1,80)=2.70$, $p=.10$) and no condition by group interaction ($F(2,80) = 8.1$, $p=.45$) but there was a group effect ($F(2,80)=3.57$, $p=.03$), with SZ patients performing slower than controls overall ($p=.04$).

Correlations. Only one noteworthy association was found in regards to the correlations between symptom scales and performance; the negative symptom score of the PANSS correlated negatively with accuracy in the face condition of the Scrambled Faces Task in SZ patients ($r=-.49$, $p=.01$), such that performance was worse in those with greater negative symptomatology.