



Cognition-emotion interactions are modulated by working memory capacity in individuals with schizophrenia

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ABSTRACT

Prior research provides evidence for aberrant cognition-emotion interactions in schizophrenia. In the current study, we aimed to extend these findings by administering the “distractor devaluation” task to 40 individuals with schizophrenia and 32 demographically matched healthy controls. The task consisted of a simple visual search task for neutral faces, followed by an evaluative response made for one of the search items (or a novel item) to determine whether prior attentional selection results in a devaluation of a previously unattended stimulus. We also manipulated working memory demands by preceding the search array with a memory array that required subjects to hold 0, 1, or 2 items in working memory while performing the search array and devaluation task, to determine whether the normative process by which attentional states influence evaluative response is limited by working memory capacity. Results indicated that individuals with schizophrenia demonstrated the typical distractor devaluation effect at working memory load 0, suggesting intact evaluative response. However, the devaluation effect was absent at working memory loads of 1 and 2, suggesting that normal evaluative responses can be abolished in people with schizophrenia when working memory capacity is exceeded. Thus, findings provide further evidence for normal evaluative response in schizophrenia, but clarify that these normal experiences may not hold when working memory demands are too high.

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1. Introduction

There has been a resurgence of interest in the area of emotional functioning in schizophrenia (Kring and Moran, 2008). Much of this work has focused on the nature of emotional experience in response to standard stimuli such as pictures, movies, smells, and film clips (Berenbaum and Oltmanns, 1992; Kring et al., 1993; Earnst and Kring, 1999; Horan et al., 2006; Strauss and Herbener, 2011). While there are exceptions (e.g., Strauss et al., 2010a), most studies indicate that patients and healthy controls report experiencing a similar magnitude of positive emotion when exposed to positive stimuli (see Cohen and Minor, 2010 for a meta-analysis), thereby challenging traditional concepts of anhedonia as a diminished capacity for pleasure (see Strauss and Gold, 2012 for a new perspective). However, it is clear that not all aspects of emotional experience are normal in schizophrenia. For example, patients also report increased levels of negative emotion in response to neutral and pleasant stimuli (see Trémeau et al., 2009; Cohen and Minor, 2010) and display reductions in pleasure-seeking and goal-directed behavior (Gard et al., 2007; Fossias and Remington, 2010; Oorschot et al., 2011). Thus, an important question remains to

be answered: why is it that patients' intact ability to experience emotions does not result in motivated behavior?

One possibility is that dysfunctional cognition-emotion interactions prevent these intact emotional experiences from being translated into motivated, goal-directed actions. Individuals with schizophrenia display a range of abnormalities in cognition-emotion interactions, and these have often been linked to greater severity of negative symptoms, particularly anhedonia and avolition. For example, patients display impairments in long-term emotional memory (Herbener et al., 2007; Herbener, 2008), emotional working memory capacity and maintenance (Anticevic et al., 2011; Gard et al., 2011; Kring et al., 2011; Ursu et al., 2011), reward learning (Waltz and Gold, 2007; Waltz et al., 2007; Gold et al., 2008; Strauss et al., 2011a,c; Gold et al., 2012), and dysfunctional emotion-attention interactions (Kinderman et al., 2003; Strauss et al., 2008; Besnier et al., 2011; Strauss et al., 2011b). The aforementioned studies have primarily investigated how emotional stimuli interact with cognition. However, in healthy individuals, it is known that the reverse is also possible, i.e., basic cognitive processes can influence subjective emotional experience and evaluative response. Given the nature and severity of cognitive impairment in schizophrenia (Heinrichs and Zakzanis, 1998), there is reason to suspect that cognitive processes may not influence evaluative response normally when cognitive demands are high.

The “distractor devaluation” paradigm may offer a novel means of testing this possibility and extending the literature on cognition-emotion interactions in schizophrenia (Fenske et al., 2004; Raymond

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et al., 2005; Goolsby et al., 2009). In this task, participants perform two procedures. First, they are asked to complete a simple attentional visual search task (e.g., two faces are presented on the screen, one male and one female, which are tinted in red or blue hue. Participants are given the task of identifying the color in which the male faces, the search target, are tinted). Then, on a subsequent screen, they make an evaluative response in relation to one of the search items (or a novel item) to determine whether prior attentional selection results in differences in evaluative response to the previously attended or unattended stimulus (e.g., subjects see a single neutral face that was either a target in the search array or a novel face and rate it on trustworthiness using a 1–9 scale). Studies using this paradigm consistently indicate that this attentional selection manipulation influences subsequent evaluative responses, whereby healthy people devalue (i.e., give lower ratings) stimuli that they have been led to ignore (i.e., distractors) on the basis of task instructions relative to stimuli that have been subjected to selective attention (i.e., targets).

An inhibition-based theory has been applied to explain this effect. When multiple stimuli are in competition for selective attention, inhibitory processes are enacted and then associated with the mental representation of the stimuli that were unattended (Raymond et al., 2003; Tipper et al., 2003; Kessler and Tipper, 2004). When an unattended stimulus is subsequently presented, the inhibitory processes are re-evoked, causing the unattended stimulus to be emotionally devalued relative to an attended target. Interestingly, the distractor devaluation effect is modulated by working memory, such that healthy subjects show devaluation in the absence of working memory demands and when demands are low to moderate, but fail to show devaluation at higher loads when capacity is exceeded (Goolsby et al., 2009). This has been explained by the fact that at higher loads, processing capacity is fully engaged by the central task, which prevents resources from being devoted to the inhibitory processing that produces the devaluation effect. Thus, attentional selection implicitly influences evaluative response, but this influence may be diminished when working memory demands are too high.

In the current study, we employed the distractor devaluation paradigm to examine cognition-emotion interactions in schizophrenia, and manipulated working memory load to determine whether working memory capacity differentially influences evaluative response in patients relative to controls. We hypothesized that patients would show normal distractor devaluation in the absence of working memory demands (i.e., lower ratings for prior distractors relative to targets), but expected patients to fail to show devaluation at higher loads. Such a pattern of findings would suggest that evaluative processes are normal in people with schizophrenia, but interact with other cognitive deficits and break down under cognitively demanding situations.

2. Methods and materials

2.1. Participants

Participants included 40 patients meeting DSM-IV criteria for Schizophrenia (SZ) and 32 Healthy Controls (CN). Persons with Schizophrenia were recruited through the Outpatient Research Program at the Maryland Psychiatric Research Center, and evaluated during a period of clinical stability evinced by no changes in medication type or dosage for a period greater than or equal to four weeks. Consensus diagnosis was established via a best-estimate approach based upon multiple interviews and a detailed psychiatric history. This diagnosis was subsequently confirmed using the Structured Clinical Interview for DSM-IV (SCID). All SZ participants were taking antipsychotic medication at the time of treatment (see Table 1 note).

Control subjects were recruited by means of random digit dialing, word-of-mouth among recruited participants, and through the use of newspaper advertisements. Controls had no current Axis I or II diagnoses as established by the SCID (First et al., 1997) and SID-P (Pfohl et al., 1997), no family history of psychosis, and were not taking psychotropic

Table 1
Demographic and clinical characteristics of controls (CN) and persons with Schizophrenia (SZ).

| | CN (n = 32) | SZ (n = 40) |
|--------------------------------|----------------|----------------|
| Age | 40.41 (10.14) | 40.17 (10.41) |
| Parental education | 13.31 (1.91) | 13.6 (2.53) |
| WASI estimated IQ | 114.25 (11.50) | 92 (13.29) |
| WTAR SS | 108.94 (12.04) | 94.85 (14.48) |
| % male | 62.5 | 57.5 |
| Ethnicity | | |
| American Indian/Alaskan native | 0.0% | 2.50% |
| Black/African American | 37.50% | 37.50% |
| Mixed race | 3.10% | 0.0% |
| White | 59.40% | 60.00% |
| MATRICES battery | | |
| Processing speed | 54.16 (8.47) | 34.28 (12.94) |
| Working memory | 50.03 (9.72) | 35.69 (10.71) |
| Verbal learning | 53.47 (12.19) | 36.08 (10.41) |
| Visual learning | 47.88 (11.32) | 30.82 (14.04) |
| Social cognition | 56.41 (7.80) | 38.85 (11.51) |
| Attention/vigilance | 53.53 (8.20) | 38.58 (11.87) |
| Reasoning/problem solving | 53.34 (9.89) | 40.73 (10.50) |
| Overall | 54.09 (9.32) | 28.5 (13.14) |

Note. WASI Estimated IQ = Wechsler Abbreviated Scale of Intelligence full-scale estimated IQ; WTAR SS = Wechsler Test of Adult Reading (WTAR) scaled score. Patients were prescribed various antipsychotic medications, either alone (clozapine: n = 15; risperidone: n = 6; olanzapine: n = 4; fluphenazine: n = 3; aripiprazole: n = 1; haloperidol: n = 1; quetiapine: n = 1; thiothixene: n = 1; ziprasidone: n = 1) or in combination with another antipsychotic (risperidone and clozapine: n = 2; aripiprazole and haloperidol: n = 1; olanzapine and clozapine: n = 1; paliperidone and quetiapine: n = 1; clozapine and aripiprazole: n = 1; olanzapine and risperidone: n = 1).

medications. All participants denied a history of significant neurological injury or disease, and significant medical or substance use disorders within the last six months. Participants were routinely screened for substance use by means of urine toxicology upon admission to the subject pool, and in any instance where substance use was suspected. All participants provided informed consent for a protocol approved by the University of Maryland Institutional Review Board.

Controls and SZ participants did not significantly differ in age: $F(1,72) = 0.01$, $p = 0.93$, parental education: $F(1,72) = 0.29$, $p = 0.60$, gender: $X^2(1,72) = 0.19$, $p = 0.67$, or ethnicity: $X^2(3,72) = 2.05$, $p = 0.56$. Patients had lower Wechsler Abbreviated Scale of Intelligence (WASI) estimated full-scale intelligence quotients, $F(1,72) = 56.10$, $p < 0.001$, and lower scores on all MATRICES battery composite scores (all $ps > 0.001$) (see Table 1).

2.2. Distractor devaluation task

In the Distractor Devaluation task, participants were asked to complete three primary procedures per trial: working memory encoding and recall, visual search, and evaluative response (see Fig. 1 for sample trial sequence). Each trial began with a 2000 ms working memory array, which was to be remembered by the participant. Images included in the working memory array were a neutral grayscale male face, a neutral grayscale female face, or a placeholder. Placeholders were created by scrambling grayscale facial images into a 20×20 grid (see Fig. 1 for an example). In conditions where working memory load was high (WM-2 conditions) both images were faces. In moderate memory load conditions (WM-1) one image was a face and the other a placeholder. In conditions without working memory load (WM-0) both images were placeholders (see Fig. 1). The working memory array was followed by a 1000 ms retention interval during which a blank screen was displayed. Then a search display appeared, consisting of two faces, one male and one female, which were presented in different hue for 300 ms. Participants were asked to make a dichotomous judgment and identify whether a male or the female face (gender pre-specified) was presented in blue or red tint, as quickly and accurately as possible via button press. The to-be-identified gender remained constant within the experimental

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