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### Altered alpha brain oscillations during multistable perception in schizophrenia

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#### ABSTRACT

Schizophrenia is a complex mental disorder with impairments in integrating sensory and cognitive functions, 16 leading to severe problems in coherent perception. This impairment might be accelerated during multistable 17 perception. Multistable perception is a phenomenon, where a visual pattern gives rise to at least two different 18 perceptual representations. 19

We addressed this issue by assessing event-related alpha oscillations during continuous viewing of an ambiguous 20 and unambiguous control stimulus. Perceptual reversals were indicated by a manual response, allowing differen-21 tiation between phases of reversion and non-reversion (that is perceptual stability) in both tasks. 22 During the ambiguous task, patients and controls showed a comparable number of perceptual reversals. Alpha 23

amplitudes in patients were larger in non-reversion phases, accompanied by a stronger decrease of alpha activity 24 preceding the perceptual reversal. This group difference was pronounced for lower alpha activity and not appar- 25 ent during the unambiguous task. 26

This indicates that ambiguous perception taps into the specific deficits that patients experience in maintaining 27 coherent perception. Given that top-down influences in generating a meaningful percept seems to be low in 28 patients, they appear more dependent on sensory information. Similar, bottom-up mechanisms might be more 29 important in triggering perceptual reversals in patients than in controls. 30

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1. Introduction Q7

1.1. Event-related brain oscillations in schizophrenia 37

Brain oscillatory activity indicates abnormal temporal integration 38 39 and interregional connectivity of brain networks as a core disturbance in schizophrenia. Such abnormal oscillatory activity has been shown 40 to be linked with a range of impairments in behavior and perception 41 (Basar, 2013; Basar and Güntekin, 2013; Basar-Eroglu et al., 2013; 4243Ford et al., 2007; Haenschel and Linden, 2011; Lee et al., 2003; Uhlhaas and Singer, 2010). 44

In the healthy brain, integrative brain functions are obtained 45 46 through the combined actions of multiple oscillations (Basar, 2008, 2011). Oscillatory activity is reported to be essential for the timing of 47 neuronal discharges (Buzsaki and Draguhn, 2004), and hence it is sug-48 49gested to provide a potential mechanism for precise temporal resolution 50in the encoding of information (e.g. Uhlhaas and Singer, 2010).

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Although schizophrenia is characterized by multiple impairments, a 51 single frequency band, the gamma-band, has currently captured most 52 attention in schizophrenia. Gamma band oscillations are modified in 53 amplitude, topography or coherence by a broad range of processes, in- 54 cluding attention, conscious awareness, multisensory and sensorimotor 55 integration, movement preparation, memory formation, feature inte- 56 gration, and stimulus selection (Basar, 2013; Sun et al., 2011). Several 57 studies report reduced gamma synchrony or amplitude in schizophre- 58 nia (e.g. Basar-Eroglu et al., 2009, 2011; Haenschel and Linden, 2011; 59 Haig et al., 2000; Herrmann and Demiralp, 2005; Lee et al., 2003; Spen- 60 cer et al., 2008). 61

Since, however, brain functions are realized in the interplay of 62 many frequency bands (Basar, 2008, 2011), it is not surprising that 63 many frequency bands seem to be deficient in schizophrenia (Basar 64 and Güntekin, 2013) under different levels of cognitive load 65 (e.g., Basar-Eroglu et al., 2008; Bates et al., 2009; Fehr et al., 2003). 66 Like gamma activity, alpha oscillations show distinct temporal-spa- 67 tial brain patterns and are modified in frequency, amplitude, topog- 68 raphy or coherence by a broad range of processes, including 69 attention, perceptual requirements, semantic content, memory for-70 mation, sensorimotor integration, movement preparation and exe-71 cution (Basar, 2012; Basar and Güntekin, 2012). Increases in 72 amplitude within the first 200 ms following an attended event have 73

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2

## **ARTICLE IN PRESS**

#### C. Basar-Eroglu et al. / International Journal of Psychophysiology xxx (2015) xxx-xxx

74 been suggested to indicate sensory processes, because the increase is 75mainly observable over primary sensory areas (Basar, 2012). Systematic suppression (i.e., decrease in power) of alpha activity is found in brain 76 77 regions actively engaged in the attentive processing of sensory information (e.g., Bastiaansen and Brunia, 2001; Pfurtscheller and Lopes da 78 79Silva, 1999; Schmiedt-Fehr et al., in press), often in parallel with an in-80 creases of alpha activity in task-irrelevant regions (e.g., Foxe and 81 Snyder, 2011; Jensen and Mazaheri, 2010). These specific modulations 82 are thought to enhance the neural responsiveness to attended informa-83 tion and disengage irrelevant neuronal generators, respectively.

Accordingly, recent studies of our research group indicate a more 84 general disturbance affecting the spatial distribution of oscillatory 85 brain activity in gamma as well as in slower frequency ranges, such as 86 delta, theta and alpha oscillations (Basar-Eroglu et al., 2007, 2008, 87 2009, 2013; Ergen et al., 2008; Schmiedt et al., 2005). For instance, pas-88 89 sive listening was found to elicit an altered distribution of event-related alpha activity in patients with schizophrenia compared to controls 90 91 (Basar-Eroglu et al., 2013). While healthy controls showed the common 92fronto-central auditory alpha response, patients had lower fronto-93 central and larger parieto-occipital alpha activity. Ideally, external 94input amplifies activity of specific neuronal populations, whereas the 95activities of the other populations are partially suppressed, which pro-96 vides an effective filtering mechanism (Basar, 2011, 2012; Basar and Güntekin, 2012). Hence, the larger parieto-occipital alpha activity in pa-97 tients with schizophrenia potentially indicates a disorder of the basic 98 neuronal filtering mechanism. 99

## 1.2. The potential functional significance of event-related alpha oscillations in multistable perception

102 Multistable visual perception is a phenomenon in which a visual pat-103tern gives rise to at least two different perceptual representations 104(e.g., the Necker cube). In order to maintain perceptual coherence only one perceptual alternative at a time is perceived. During continu-105ous observation, perception switches between all possible alternatives 106 although the physical properties of the stimulus remain unchanged 107(Long and Toppino, 2004). It is generally acknowledged that this phe-108 nomenon is driven by sensory information (bottom-up) as well expec-109 tations and prior experiences (top-down, Gregory, 1997; Long and 110 Toppino, 2004). Neurophysiological processes underlying perceptual 111 reversions have been investigated to disentangle bottom-up from top-112 113 down influences in multistable perception (Mathes et al., in press; Strüber et al., 2014). For instance, several studies reported a continuous 114 alpha decrease preceding perceptual reversals during multistable visual 115 116 perception (Isoglu-Alkac et al., 2000; Isoglu-Alkac and Strüber, 2006; Strüber and Herrmann, 2002). Isoglu-Alkac and Strüber (2006) suggest 117

that lower alpha suppression (i.e., amplitude decrease) reflects an auto- 118 matic arousal reaction, which triggers attentional processing during 119 perceptual switching in a bottom-up manner. Desynchronization of 120 higher alpha activity might be more closely related to top-down 121 processes during multistable perception (see also Klimesch, 1999). 122

#### 1.3. Present study

Pathological changes of alpha activity during visual multistable perception in schizophrenia may provide an appropriate model to investigate integrative processes of the brain and the potential impact of failures in object perception due to psychiatric illness. Several behavioral studies indicate that patients with schizophrenia show altered processing of visual ambiguous stimuli (e.g., McBain et al., 2011; 129 Tschacher et al., 2008). 130

The aim of this study was to assess event-related alpha oscillations 131 during perception of an ambiguous visual stimulus, termed stroboscop-132 ic alternative motion (SAM, see Fig. 1). EEG data were recorded in 133 patients with schizophrenia and healthy controls while they viewed 134 the SAM in two tasks that differed in terms of stimulus ambiguity 135 (*ambiguous* stimulus and an *unambiguous* control stimulus). We 136 compared alpha activity in both groups and both tasks during phases 137 of perceptual stability (*non-reversion*), on the one hand, and during per-138 ceptual reversals (*reversion*) on the other hand. 139

We assumed that the ambiguous (i.e., multistable) version of the vi- 140 sual pattern (i.e., SAM) would require considerably more cognitive con- 141 trol mediation, in contrast to an unambiguous visual pattern (see also 142 Mathes et al., in press). Since top-down control of ambiguous percep- 143 tion seems to be specifically impaired in schizophrenia (e.g., McBain 144 et al., 2011), we expected that disturbances in object perception 145 would be more apparent during ambiguous than unambiguous percep- 146 tion. The continuous alpha decrease should ideally reflect upon that, be- 147 cause it occurs specifically during ambiguous perception (Strüber and 148 Herrmann, 2002) and seems to reflect the interplay of bottom-up and 149 top-down mechanisms (Isoglu-Alkac and Strüber, 2006). Based on pre- 150 vious studies, we expected that the alpha-band would reflect abnormal 151 visual multistable perception in schizophrenia with a brain activation 152 pattern indicating failures in spatial-temporal integration (see e.g. 153 Basar-Eroglu et al., 2008; Basar-Eroglu et al., 2009). 154

### **2. Methods** 155

### 2.1. Participants

All patients were recruited as in-patients after clinical stabilization of 157 an acute psychotic episode. Fourteen patients were diagnosed with 158

### Stroboscopic Alternative Motion (SAM)



**Fig. 1.** Schematic representation of the stroboscopic alternative motion (SAM). During viewing of the SAM vertical apparent motion alternates with horizontal apparent motion (*ambiguous* stimulus, left) even though the physical properties of the stimulus remain unchanged. The *unambiguous* stimulus pattern (right) elicits apparent vertical and horizontal motion by externally controlled alternation of the presented stimulus displays. (+) = Fixation point.

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123

156

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