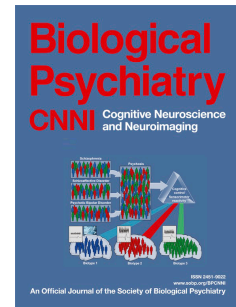


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Neural basis of impaired emotion recognition in adult attention deficit hyperactivity disorder

Agnieszka Zuberer^{1,2*}, Lena Schwarz¹, Benjamin Kreifelts¹, Dirk Wildgruber¹, Michael Erb³, Andreas Fallgatter¹, Klaus Scheffler^{3,4}, Thomas Ethofer^{1,3}

¹ Dept. for Psychiatry and Psychotherapy, University of Tübingen, Tübingen, Germany

² Department of Psychiatry and Psychotherapy, Jena University Hospital, Jena, Germany

³ Department for Biomedical Magnetic Resonance, University of Tübingen, Tübingen, Germany

³ Max-Planck-Institute for Biological Cybernetics, Tübingen, Germany

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Corresponding author:

Agnieszka Zuberer, PhD

Department of Biomedical Magnetic Resonance

Otfried-Müller-Str. 51

72076 Tübingen

Germany

Email: azuberer@gmail.com

Abstract**Background**

Deficits in emotion recognition have been repeatedly documented in patients diagnosed with attention deficit hyperactivity disorder (ADHD), but their neural basis is unknown so far.

Methods

In the current study, adult ADHD patients (n=44) and healthy controls (HC) (n=43) underwent functional magnetic resonance imaging (fMRI) during explicit emotion recognition of stimuli expressing affective information in face, voice, or face-voice combinations. The employed experimental paradigm allowed us to delineate areas for processing audiovisual information based on their functional activation profile including bilateral posterior superior temporal gyrus/middle temporal gyrus (STG/MTG), amygdala, medial prefrontal cortex, and precuneus as well as the right posterior thalamus.

Results

As expected, unbiased hit rates for correct classification of the expressed emotions were lower in ADHD patients than healthy controls irrespective of the sensory modality. This deficit at behavioral level was accompanied by lower activation in ADHD patients versus HC in the cortex adjacent to the right STG/MTG as well as the right posterior thalamus which represent key areas for processing socially relevant signals and their integration across modalities. The cortex adjacent to the right pSTS was the only brain region which showed a significant correlation between brain activation and behavioral data (unbiased hit rates).

Conclusions

Altogether these results provide first evidence for a potential neural substrate of the observed impairments in emotion recognition in an adult ADHD. (219 words)

1 **Introduction**

2 Attention-Deficit/Hyperactivity Disorder (ADHD) is characterized by inattention, motor hyperactivity
3 and hyperactivity (1). It interferes with impulse and emotional control leading to impaired social
4 interactions and subsequent mental health problems (2). ADHD was first described in children, but it
5 is meanwhile accepted that it persists into adulthood in roughly every second patient although with
6 partly altered symptomatology (1, 2). Social interaction deficits of ADHD patients are well known (3)
7 and behavioral research documented impaired emotion recognition for facial expressions (4) and
8 prosody in children (5-7) and adults (8-11) with ADHD. Such emotion recognition deficits correlate
9 with attention parameters (11, 12), but inattention cannot fully explain impairments in emotion
10 perception (11) arguing for a social cognition deficit of ADHD patients in its own right (13).

11 Two imaging studies investigating processing of facial emotions revealed hyperactivation of the
12 amygdalae during overt (14) as well as subliminal presentation (15) while a third study argued for
13 hypoactivation of the amygdala which is modulated by medication with methylphenidate (16).
14 Concerning emotional prosody, a near-infrared spectroscopy (NIRS) study revealed hypoactivation to
15 anger prosody within the right superior temporal gyrus (17) which has been implicated in procession
16 of emotional voices (18). So far, however, it is unknown whether the brain areas responsible for
17 binding emotional information expressed in voice and face (i.e. audiovisual integration areas) exhibit
18 an altered activation profile and to which extent such differences correlate with behavior.

19 In this study, we examined processing of emotional information presented in voice, face, or face-voice
20 combinations in adult ADHD patients versus healthy controls (HC) using functional magnetic
21 resonance imaging (fMRI). The employed experiment was evaluated in a behavioral study (11) in
22 ADHD patients as well as an fMRI study in HC (19). Thus, we had straightforward hypotheses
23 regarding behavioral effects (i.e. lower recognition rates in ADHD patients versus HC for all
24 emotions) as well as activations in bilateral posterior superior temporal gyrus/middle temporal gyrus
25 (STG/MTG) and right posterior thalamus (19). We specifically investigated whether lower recognition
26 rates of emotions in ADHD patients are accompanied by decreased functional activations in these
27 areas and to which extent activation in these areas correlates with behavior.

28

29 Material and methods**30 *Participants***

31 44 patients with ADHD of combined presentation (10 women, mean age \pm standard deviation (SD)
32 30.0 ± 7.0 years) and 43 HC (18 women, 28.2 ± 6.5 years) were included. All participants were
33 German native speakers, right-handed (Edinburgh Handedness Inventory, 20), and had normal or
34 corrected to normal vision. The education was assessed in years not counting repetition of classes.
35 Stimulant medication (methylphenidate: 13 patients, atomoxetine: 2 patients, dexamphetamine: 1
36 patient) was discontinued for at least five half-life periods. None of the participants were treated with
37 any other psychotropic medication. Patients were recruited via the outpatient clinic of the University
38 Hospital for Psychiatry and Psychotherapy Tübingen. ADHD diagnosis was established on the basis of
39 a clinical interview by a trained psychiatrist or psychologist assessing DSM-V diagnostic criteria for
40 the combined presentation in adults and the self-report questionnaires ADHS-SB (“ADHS-
41 Selbstbeurteilungsskala”, 21) and WURS-K (“Wender-Utah-Rating-Scale-Kurzform“, 22) as well as
42 third-party anamnesis on childhood symptoms. HC were included if they were never diagnosed with
43 or suspected to suffer from ADHD and were clearly below the diagnostic threshold of any present or
44 childhood ADHD subtype ($< 90\%$ of the clinical threshold obtained via questionnaires). Verbal
45 intelligence was assessed using the Mehrfachwortschatztest (MWT-B, Multiple-Choice Vocabulary
46 Intelligence Test, 23). None of the participants suffered from major depression or any other current
47 serious psychiatric condition including drug use, as determined by the German Version of the
48 Structured Clinical Interview (SKID-I, 24). To exclude autistic symptoms all participants completed
49 the Adult Asperger Assessment, 25). Depressive symptoms were captured with the BDI-II (“Beck
50 Depression Inventory”, 26). The study was performed according to the Code of Ethics of the World
51 Medical Association (Declaration of Helsinki) and the protocol was approved by the ethics committee
52 of the medical faculty of the Eberhard-Karls-University Tübingen. All participants gave written
53 informed consent prior to participating.

54

55 *Stimulus material and experimental design*

56 The stimuli were created with the help of professional actors who were videotaped while speaking
57 single German words in a neutral, angry, disgusted, happy, or seductive tone of voice with a congruent
58 facial expression (for more details, see 11). The stimuli were evaluated by an independent group of 31
59 healthy participants (mean age: 27.3 ± 6.0 years, 16 females). Based on this evaluation we selected the
60 stimuli to guarantee that they are well recognized (minimum recognition rate for each stimulus: 80%)
61 and that recognition rates during audiovisual presentation are balanced across emotions (mean
62 recognition rates: neutral = 93.4%, happy = 95.1%, seductive = 94.1%, angry = 95.1%, disgusted =
63 95.2%). During three fMRI sessions these stimuli were presented as audiovisual color videos (AV),
64 mute color videos (V), or auditory sound sequences (A) without visual presentation (60 stimuli per
65 modality). During each of the three fMRI sessions 20 stimuli per modality were presented. Participants
66 indicated the expressed emotion on a 5-point circular scale with the German words for “happiness”;
67 “eroticism”, “neutral”, “anger”, and “disgust”. The word “neutral” was placed on top of the circle
68 whereas the two positive connoted expressions “eroticism” and “happiness” were shown on one side
69 and the negative connoted emotions on the other side. To avoid possible laterality effects, the scale
70 was flipped horizontally for half of the participants of both groups. A white dot was randomly placed
71 at one of the five categories and participants had 6 s to choose the response which was fitting best to
72 the expressed affect by moving the white dot clockwise by a button of the response device (see Fig. 1).
73 This design was previously evaluated for application in fMRI studies targeting audiovisual integration
74 (19) and was chosen to remove potential biases from spatial arrangement of response alternatives.
75 While this design effectively removes any association between the number of motor responses and a
76 given category it does not allow to determine omission errors as the randomly chosen starting point of
77 the white dot might already correspond to the category which the participant judges as fitting best to
78 the presented stimulus (and thus this trial requires no response). **The number of trials without motor**
79 **responses was comparable across the two groups (ADHD: 22.8 ± 1.5 , HC: 24.4 ± 1.5 , $p = 0.22$)**
80 **indicating that there was no systematic bias due to inattention.**
81 The order of the stimuli within the three imaging sessions was randomized. Stimulus onset occurred
82 on average every 10.5 s (range: 9.0 - 12.0 s) and was jittered in TR/4 with a fixation cross presented in
83 between stimulus presentation. Stimulus presentation and response recording relied on Presentation

84 software (Neurobehavioral Systems, www.neurobs.com). A screen was placed at the head end of the
85 MRI scanner, visible for participants through a mirror attached in front of their eyes. Sound was
86 presented via MR-compatible headphones. Responses were conveyed using a fiber optic response
87 system (Celeritas Fiber Optic Button Response System, Psychology Software Tools).

88

89 *Analysis of behavioral data*

90 We evaluated emotion classification accuracy by determining unbiased hit rates (HU, 27) which were
91 obtained for each emotion by multiplying the simple hit rates with their respective positive predictive
92 value to incorporate both sensitivity and specificity of the responses. HU values were arcsine
93 transformed and then submitted to a repeated-measures analysis of variance (ANOVA) with emotion
94 (neutral, happy, seductive, angry, disgusted) and modality (AV, V, A) as within-subject factor and
95 group (ADHD, HC) as between-subjects factor. Gender and BDI scores were included as covariates of
96 no interest. P-values were corrected for effects of non-sphericity (28).

97

98 *MRI data acquisition*

99 Imaging was performed using a 3 Tesla scanner (Siemens PRISMA, Erlangen, Germany) and a 20
100 channel head coil. We acquired a high-resolution structural 3D magnetization prepared rapid
101 acquisition gradient echo (MPRAGE) T1-weighted scan (TR = 2.3 s, TE = 4.16 ms, TI = 0.9 s, flip
102 angle 9°, voxel size = 1x1x1 mm³), a fieldmap for image distortion correction (TR = 0.4 s, TE(1) =
103 5.19 ms, TE(2) = 7.65 ms, flip angle = 60°, voxel size = 3x3x3 mm³) and 441 functional images (72
104 transversal slices acquired interleaved, TR = 1.5 s, TE = 34 ms, flip angle = 70°, voxel size = 2x2x2
105 mm³, multiband acceleration factor 3).

106

107 *Analysis of fMRI data*

108 FMRI data were analyzed with the statistical parametric mapping software (SPM12, Wellcome
109 Department of Imaging Neuroscience, London, UK, www.fil.ion.ucl.ac.uk/spm/software/spm12).
110 Preprocessing included realignment and unwarping to correct for movement as well as static and

111 movement-dependent field distortions (29), coregistration to the anatomical image, normalization to
112 the standard space of the Montreal Neurological Institute (MNI, resampled voxel size 2x2x2 mm)
113 based on the unified segmentation algorithm (30), and smoothing with an isotropic Gaussian filter of 5
114 mm full width at half maximum.

115 Statistical analysis relied on a general linear model (GLM, 31) with three regressors (A, V, and AV
116 trials) modelling presentation of the stimuli with a boxcar function of two seconds duration convolved
117 with the hemodynamic response function. A high-pass filter with a cut-off frequency of 1/128 Hz
118 filtered out low-frequency components. The error term of the GLMs was modelled as a first-order
119 autoregressive process (AR coefficient = 0.2) plus white noise to account for serial autocorrelations.

120 Contrast images from the first-level GLMs were submitted to second-level random effects analyses.
121 Assignment of activation clusters relied on the automatic anatomical labelling tool (32). We used
122 conjunction analyses (33) to define areas for audiovisual integration ($AV > V \cap (AV > A)$) using a
123 voxel-wise family-wise error (FWE) corrected height threshold of $p < 0.05$ and a cluster threshold of k
124 > 25 voxels. Based on previous research, the regions-of-interest (ROIs) defined by these second-level
125 conjunction analyses were expected to include audiovisual integration areas within bilateral
126 STG/MTG and right posterior thalamus (19). Averaged responses of brain regions delineated by this
127 conjunction analysis were obtained for each subject and submitted to repeated-measures ANOVAs
128 with modality (AV, V, A) as within-subject factor and group (ADHD, HC) as between-subjects factor.
129 Please note that the ROIs were defined by a modality-dependent conjunction analysis and thus the
130 main effect of modality is inevitably significant and therefore only reported for the sake of
131 completeness. This is not the case for the main effect of group or the interaction between group and
132 modality as they are orthogonal to the definition of the ROIs on the basis of the conjunction analysis.

133 Gender and BDI scores were included as covariates of no interest. P-values were corrected for effects
134 of non-sphericity (28). We additionally investigated whether the ROIs exhibited a significant
135 correlation between activation and behavior (HU).

136

137 **Results**

138 *Demographic and psychometric data*

139 Demographic and psychometric data are presented in Table 1. There were no significant differences
140 between the two groups regarding age or educational level, but ADHD patients exhibited significantly
141 lower levels of verbal intelligence as measured by the MWT-B ($p < 0.01$). As expected, significantly
142 higher levels of self-reported problems regarding inattention, hyperactivity, and impulsivity were
143 found regarding childhood (WURS-K) and present symptoms (ADHS-SB) in ADHD patients than HC
144 (both $p < 0.001$). Although none of the participants fulfilled the clinical criteria for a major depressive
145 disorder, ADHD patients exhibited significantly higher scores ($p < 0.01$) for depressive symptoms as
146 assessed by the BDI-II.

147

148 ***Behavioral data***

149 Mean HUs depending on the sensory modality and emotion are shown in Fig. 2 for ADHD patients
150 and HC (grey and white bars, respectively). The repeated-measures 5x3x2 ANOVA with emotion
151 (neutral, happy, seductive, angry, disgusted) and modality (AV, V, A) as within-subjects factor and
152 group (ADHD, HC) as between-subjects factor as well as gender and BDI scores as covariates of no
153 interest revealed a significant main effect of emotion ($F(4, 328) = 6.56, p < 0.001$), modality ($F(2,$
154 $164) = 31.75, p < 0.001$), and group ($F(1, 82) = 4.65, p < 0.05$). The main effect of emotion was due
155 to the fact that HUs (averaged across modalities and groups) were significantly higher (all $T(86) >$
156 4.54 , all $p < 0.001$) for stimuli expressing happiness ($70 \pm 1\%$) and seduction ($70 \pm 2\%$) than anger
157 ($62 \pm 1\%$), disgust ($65 \pm 1\%$), or neutrality ($64 \pm 1\%$). The main effect of modality was due to
158 higher HUs (averaged across emotions and groups) for AV stimuli ($82 \pm 1\%$) than V stimuli ($69 \pm 1\%$)
159 and A stimuli ($47 \pm 1\%$). All statistical comparisons across the three modalities were significant (all
160 $T(86) > 11.57$, all $p < 0.001$). The main effect of group was driven by significantly ($T(85) = 2.49, p <$
161 0.05) lower HUs (averaged across emotions and modalities) in ADHD patients ($64 \pm 1\%$) than HC (69
162 $\pm 2\%$). There was a significant interaction between emotion and modality ($F(8, 656) = 4.54, p <$
163 0.001). This interaction reflected significantly higher HUs (averaged across groups) for AV and V
164 stimuli expressing happiness than the other four categories (all $T(85) > 2.55$, all $p < 0.05$) while
165 seduction was significantly better recognized than the other four categories for A stimuli (all $T(85) >$
166 10.4 , all $p < 0.001$). None of the interactions including the factor group were significant.

167

168 *fMRI data*

169 In agreement with our a priori hypothesis, the conjunction analysis revealed significant activations in
170 bilateral STG/MTG and right posterior thalamus. In addition, bilateral hippocampus/amygdala, medial
171 superior frontal gyrus (SFG), precuneus, and gyrus rectus as well as right temporal pole were
172 identified in this analysis (see Fig. 3 and Table 2).

173 The repeated-measures 2x3 ANOVA revealed a significant main effect of modality (all $F(2, 162) >$
174 4.68 , all $p < 0.05$) in all ROIs which is to be expected as the ROIs were defined by a modality-specific
175 effect. A significant main effect of group was only found in right STG/MTG ($F(1, 81) = 6.34$, $p <$
176 0.05) and right thalamus ($F(1, 81) = 4.79$, $p < 0.05$) which was due to lower activation in ADHD
177 patients than HC (see Fig. 2, all $T(84) > 3.30$, all $p < 0.05$). None of the other brain areas delineated by
178 the conjunction analysis exhibited a significant main effect of group (all $F(1, 81) < 1.32$, all $p > 0.38$).
179 There was no interaction between modality and group in any of the ROIs (all $F(2, 162) < 2.41$, all $p >$
180 0.13). The only brain area which showed a significant correlation between activation and Hus was the
181 right STG/MTG. In this brain area, the correlation between activation (beta estimates) and HUs was r
182 $= 0.18$ for AV and V trials as well as $r = 0.22$ for A trials (all $p < 0.05$). Separate analyses for ADHD
183 patients and HC revealed that these correlations are mostly driven by the control group for AV trials
184 (HC: $r = 0.38$, ADHD: $r = -0.07$) and A trials (HC: $r = 0.28$ and ADHD: $r = 0.09$). For V trials similar
185 correlation coefficients were observed in both groups (HC: $r = 0.16$, ADHD: $r = 0.14$).

186

187 **Discussion**

188 To our knowledge, this is the first neuroimaging study investigating the neural basis of emotion
189 recognition in adult ADHD patients. We utilized an explicit emotion categorization task (19) and
190 presented emotions in different sensory modalities (facial expressions, prosody, and their
191 combination) allowing us to delineate key areas for emotion processing including posterior
192 STG/MTG, thalamus, and amygdala. Adult ADHD patients exhibited diminished activation in the
193 right posterior STG/MTG and the thalamus suggesting that processing of emotions is disrupted both in

194 early and late processing. The only ROI showing a correlation between recognition rates and
195 activation was the right STG/MTG suggesting its activity has direct consequences for behavior.

196

197 *Behavior*

198 Participants identified audiovisually presented emotions better than auditory or visual stimuli alone.

199 This enhanced performance when combining auditory and visual emotional information is a well-
200 established finding (11, 19, 34, 35) as the combination across modalities yields richer information.

201 Thus, reductions of the uncertainty translate to better accuracy in task performance (36). There is
202 initial evidence that this cross-modal effect might not result from “post-perceptual decision under
203 attentional control” (34), but instead reflects an early-on integration process (36, 37) with a limited
204 role of awareness.

205 It should be noted, that the emotion recognition rates obtained in this study during fMRI were lower
206 than in a previous behavioral study (11) which is most probably due to distractions caused by the
207 scanner noise and the less comfortable situation. The main result with lower recognition rates in
208 ADHD patients than HC, however, was replicated. This emotion recognition deficit in ADHD was not
209 confined to a specific emotional category. Although this is in line with recent work (11, 38), previous
210 meta-analyses suggested that patients with ADHD show a robust deficiency in recognizing negative
211 emotions, particularly for anger and fear (13, 39). This observation is typically discussed as a deficit in
212 decoding cues that signal socially relevant negative feedback (e.g. 40). There are several aspects that
213 could explain the discrepancy to our results. First, we did not include fear stimuli as we decided to
214 choose emotions with strong relevance for social interactions rather than those signaling threat-related
215 information important for survival. However, no bias for anger was found in our study either.
216 Secondly, most of the studies on this topic employed several negative emotions (5, 41-43), but only
217 one positive emotion (i.e. happiness, for a discussion on this issue, see 44). This represents a possible
218 confound as it requires discrimination between distinct categories of negative, but not positive
219 emotions. We avoided such confounds of emotion-specific differences in task difficulty by including
220 the same number of positive and negative stimuli/categories and using stimuli with similar recognition
221 rates across emotions (11). In line with these previous findings, our behavioral results suggest that the

222 emotion independent performance decrement in ADHD might reflect a general sensory encoding
223 deficiency (11) which occurs similarly for all emotions used in our study.

224

225 *Activation*

226 The experimental design was adapted from a previous fMRI study in HC (19) which identified
227 bilateral posterior STG/MTG and right thalamus as central nodes for processing facial and vocal
228 emotions. We replicated these findings, but also delineated significant results for medial SFG,
229 precuneus, and gyrus rectus as well as right temporal pole which is most probably due to the higher
230 statistical power provided by a larger number of participants (87 versus 24 subjects) as well as higher
231 signal-to-noise ratio afforded by a higher field strength (3T versus 1.5T) and faster sampling (TR = 1.5
232 s versus 2.0 s) by using multiband EPI. The only ROIs that showed a significant difference with lower
233 activation in ADHD patients, however, were the right posterior STG/MTG and the right posterior
234 thalamus. The posterior STG/MTG has consistently been reported to be involved during audiovisual
235 integration of facial and vocal information (19, 35, 45, 46). Furthermore, audiovisual percepts, such as
236 the McGurk illusion, depend on the activation level in this area (47) and can be disrupted by means of
237 transcranial magnetic stimulation (TMS, 48). While this suggests that the posterior STG/MTG is
238 involved in audiovisual integration, there is also evidence that it participates in attentional processes
239 (49-51) and shifts of attention to social cues (52, 53). As hypoactivation within this area and emotion
240 recognition deficits of ADHD patients were not confined to the bimodal condition, but similarly
241 occurred for the two unimodal conditions, this finding might rather reflect deficient allocation of
242 attentional resources to relevant cues than disrupted audiovisual integration. The same pattern of brain
243 activation was observed in the right posterior thalamus indicating that altered brain activity is not
244 restricted to higher-order areas, but arises also in early processing stages. This is in excellent
245 agreement with electrophysiological studies indicating alterations of early and late event-related
246 potentials (ERP, 54, 55) during processing of facial emotions in ADHD patients.

247 Two previous fMRI studies described increased activation (14, 15) while a third reported diminished
248 activation within the amygdala (16) of ADHD patients. In our study, no significant difference in
249 activation was found within the amygdala. A likely explanation for these discrepant findings across

250 studies is the nature of the employed tasks as it is long established that the amygdala show activation
251 differences during implicit versus explicit processing (56, 57) or rating of self- versus other-related
252 emotions (14). However, other differences in study design including participants (adults versus
253 children or adolescents), stimuli (dynamic videos versus photographs), and employed emotions might
254 also contribute to variability of amygdalar responses across studies.

255

256 *Association between behavior and activation*

257 Better emotion recognition was related to higher activations in the posterior STG/MTG in HC
258 replicating previous results (19). This relationship between activation and behavior was not confined
259 to audiovisual stimuli, but also occurred for auditory and visual stimuli. A possible explanation for this
260 observation is that electrophysiological results obtained in multisensory regions of the macaque
261 revealed that only a minority of about 20% of cells within these areas are sensitive to stimuli from
262 more than one modality (58, 59) and that high-resolution fMRI revealed that the posterior STG/MTG
263 has a patchy organization of unimodal and multimodal units (45). Furthermore, neuroimaging data
264 revealed that sharpening of neural responses as measured by repetition suppression (60) within this
265 area occurs also for unimodal and bimodal stimuli (35). In contrast to HC, ADHD patients did not
266 show a significant association between behavior and activation levels of the right posterior STG/MTG.
267 This finding is in line with NIRS data indicating that ADHD patients exhibit a higher variability of
268 oxygenated hemoglobin within this area during perception of emotional faces (61). Unfortunately, this
269 NIRS study was based on a passive viewing paradigm making it impossible to draw conclusions with
270 respect to behavioral consequences of this phenomenon.

271 Our findings showing a relationship between activity in the right STG/MTG and emotion recognition
272 nicely dovetails with electrophysiological results indicating that late ERP components, such as the P3,
273 correlate with behavioral measures obtained during emotional face perception (55). We believe that a
274 relationship between brain activation and behavior is the strongest evidence which fMRI alone can
275 provide for the assumption that a certain brain area is implicated in a cognitive process. Further
276 evidence for a causal involvement requires combination with other methods, such as TMS, to show
277 that disruption of activity impacts behavior (48). It has, however, been pointed out that altered

278 activation without concurrent behavioral effect might still provide complementary information about
279 'hidden' processes not observable by traditional behavioral measures (62). Thus, we do not propose
280 that the lack of a correlation between activity in the right posterior thalamus and emotion recognition
281 signifies that diminished activity in this area is irrelevant. It is possible that disrupted activity in early
282 regions could affect downstream processing areas which might explain why the correlation between
283 activation and behavior in the posterior STG/MTG was observed for HC, but not ADHD patients.

284

285 *Limitations*

286 It should be noted that ADHD patients were recruited from an outpatient clinic dedicated to
287 diagnosing adults first-time. We took careful measures to ensure relevant symptoms were present in
288 childhood, including primary school certificates encompassing descriptions of behavior. Nevertheless
289 it is possible that the ADHD patients included in this study were milder affected than their peers
290 diagnosed in childhood right away as they did not require consultation until facing heightened
291 environmental demands in adulthood. Our screening for comorbid disorders relied on the SKID-I.

292 None of the participants showed any clinical sign for intoxication, but future studies could still include
293 toxicological screenings to rule out any alcohol or drug use and employ SKID-II screenings to
294 determine whether activation is additionally influenced by personality factors. Finally, inclusion of
295 patients with other psychiatric disorders could determine whether our findings are specific for ADHD.

296

297 *Conclusion*

298 To our knowledge, this is the first fMRI study investigating the neural basis of impaired emotion
299 recognition in ADHD. Overall, our findings indicate that ADHD patients exhibit a general deficit in
300 emotion recognition which occurs irrespective of sensory modality or emotional category and relates
301 to hypoactivations in both an early (right posterior thalamus) and a late (right posterior STG/MTG)
302 brain region. In line with previous electrophysiological results activation in the late, but not the early
303 sensory processing region correlated with behavior pinpointing to the right posterior STG/MTG as a
304 potential neural correlate mediating behavioral consequences. (4000 words)

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The authors report no biomedical financial interests or potential conflicts of interest.

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Tables:**Table 1:** Demographic and psychometric data

	ADHD	HC	
	Mean \pm SD	Mean \pm SD	two-tailed p
Age	30.0 \pm 7.0	28.2 \pm 6.5	0.21
Educational years	15.9 \pm 3.1	16.6 \pm 2.4	0.25
	Mean \pm SEM	Mean \pm SEM	one-tailed p
WURS-K	43.2 \pm 2.0	8.3 \pm 0.9	1.5 x 10 ⁻²³
ADHD-SB	31.8 \pm 8.0	5.5 \pm 4.2	4.0 x 10 ⁻²⁹
BDI	7.0 \pm 1.1	1.8 \pm 0.3	8.7 x 10 ⁻³
MWT-B	101.3 \pm 1.6	107.4 \pm 1.9	9.4 x 10 ⁻³

ADHD: Attention deficit hyperactivity disorder, HC: Healthy Controls, SD: Standard Deviation, SEM:

Standard error of the mean

Table 2: Brain areas showing a significant audiovisual integration effect (AV-A \cap AV-V)

Anatomical Definition	MNI coordinates	Z score	cluster size
Right STG/MTG	48 -38 8	6.58	266
Bilateral medial SFG	6 56 34	6.50	253
Left hippocampus/amygdala	-22 -12 -14	6.42	57
Right posterior thalamus	14 -30 -2	6.38	28
Left STG/MTG	-58 -56 12	6.22	288
Right hippocampus/amygdala	22 -16 -14	6.04	47
Bilateral precuneus	0 -58 36	5.97	136
Bilateral gyrus rectus	2 46 -18	5.90	26
Right temporal pole	48 16 -32	5.81	56

STG: superior temporal gyrus, MTG: middle temporal gyrus, SFG: superior frontal gyrus, MNI:

Montreal Neurological Institute

Figure legends:

Figure 1: Task design. Stimuli, balanced for experimental condition (A, V, AV) and emotional category were presented across three imaging sessions with 60 trials each. Stimuli were presented in randomized order within sessions. Inter stimulus intervals ranged from 9-12 s and subjects had 6 s to choose a response. The classification task was performed on a circular scale with five emotional categories (“EROTIK”=eroticism/seduction, “FREUDE”=happiness, “EKEL”=disgust, “ÄRGER”=anger, “TRAUER”=sadness, “NEUTRAL”=neutral). In order to avoid possible laterality effects, the scale was flipped horizontally for half of the participants of both groups.

Figure 2: Emotion identification performance (unbiased hit rate \pm standard error of the mean). ADHD subjects exhibit poorer performance than healthy controls across all emotions (H: happy, E: erotic/seductive, D: disgusted, and A; angry) and modalities (right panel: unimodal acoustic, center panel: visual, and left panel: audiovisual).

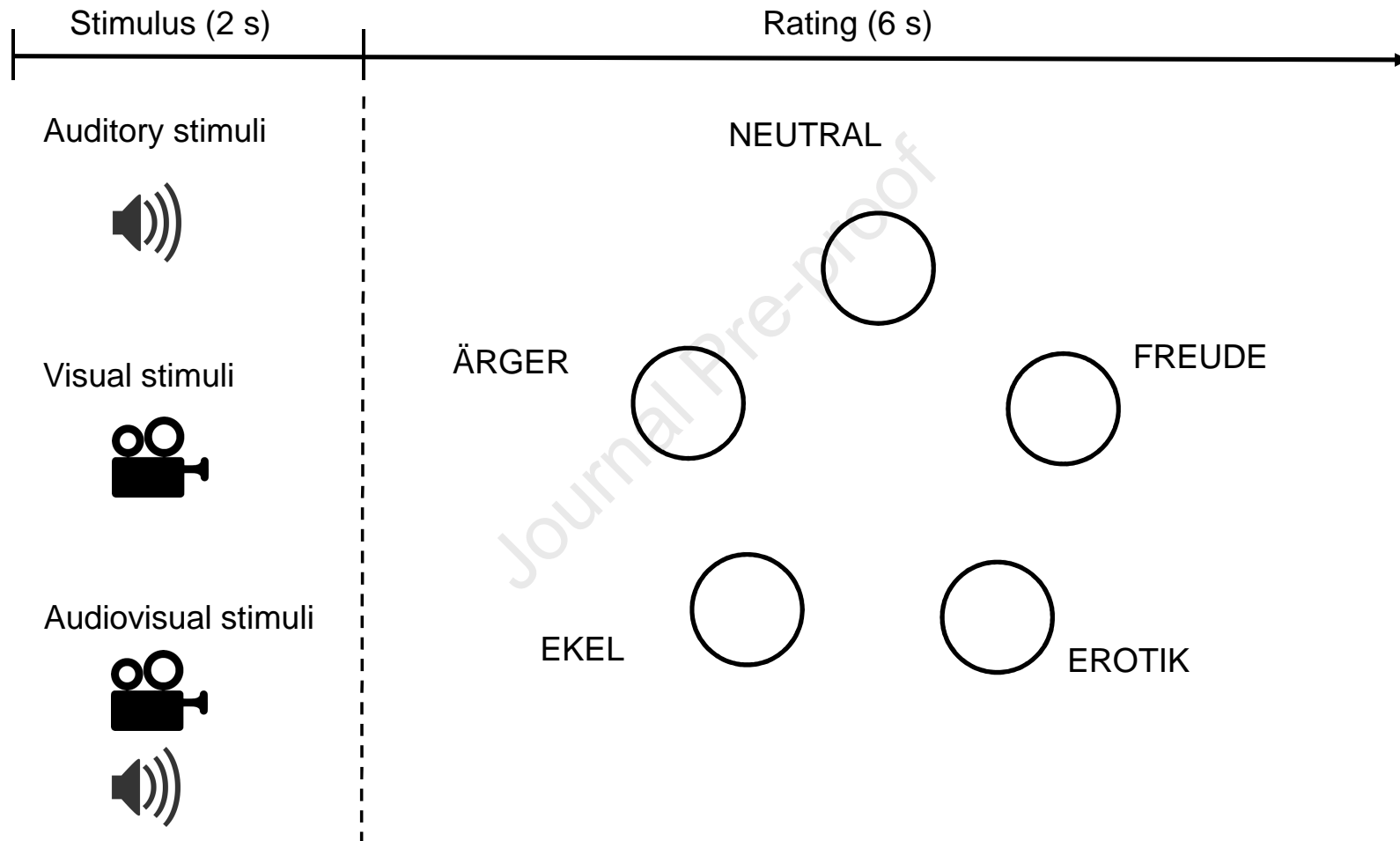
Figure 3: Conjunction analysis $AV - A \cap AV - V$. ADHD patients exhibited lower levels of activation in the right posterior STG/MTG (left panel) and right posterior thalamus (right panel). Activation levels in the right posterior STG/MTG were positively correlated with the emotion identification performance, most prominently within the auditory-visual condition. These effects were mostly driven by the performance of healthy controls (center panel).

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Emotion identification performance

