1	Visual processing recruits the auditory cortices in prelingually deaf children
2	and influences cochlear implant outcomes
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17 Abstract

Objective: Although visual processing recruitment of the auditory cortices has been reported previously in prelingually deaf children who have a rapidly developing brain and no auditory processing, the visual processing recruitment of auditory cortices might be different in processing different visual stimuli and may affect cochlear implant (CI) outcomes.

Methods: Ten prelingually deaf children, 4–6 years old, were recruited for the study. Twenty prelingually deaf subjects, 4–6 years old with CIs for 1 year, were also recruited; 10 with well-performing CIs, 10 with poorly performing CIs. Ten age and sex-matched normal-hearing children were recruited as controls. Visual ('sound' photo (photograph with imaginative sound) and 'non-sound' photo (photograph without imaginative sound)) evoked potentials were measured in all subjects. P1 at Oz and N1 at the bilateral temporal-frontal areas (FC3 and FC4) were compared.

30 Results: N1 amplitudes were strongest in the deaf children, followed by those with 31 poorly performing CIs, controls and those with well-performing CIs. There was no 32 significant difference between controls and those with well-performing CIs. 'Sound' 33 photo stimuli evoked a stronger N1 than 'non-sound' photo stimuli. Further analysis 34 showed that only at FC4 in deaf subjects and those with poorly performing CIs were 35 the N1 responses to 'sound' photo stimuli stronger than those to 'non-sound' photo 36 stimuli. No significant difference was found for the FC3 and FC4 areas. No 37 significant difference was found in N1 latencies and P1 amplitudes or latencies.

38 Conclusions: The results indicate enhanced visual recruitment of the auditory
39 cortices in prelingually deaf children. Additionally, the decrement in visual
40 recruitment of auditory cortices was related to good CI outcomes.

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41 **(Keywords)** prelingual deafness, cochlear implant, cross-modal plasticity

43 Introduction

44 It is generally accepted that one sense can benefit from the deprivation of another 45 (1). This is observed in both blind and deaf individuals (2,3). In the prelingually deaf 46 the auditory cortex can respond to visual stimuli, indicating cross-modal recruitment 47 of auditory cortex by visual stimuli, known as cross-modal reorganization (4,5). 48 Neuroimaging studies using functional Magnetic Resonance Imaging (fMRI) and 49 magnetoencephalography (MEG) reveal that visual stimuli such as a moving dot 50 pattern, can activate certain regions of the auditory cortex (Brodmann's areas 42 and 51 22) in prelingually deaf participants (4,6,7). In addition, some event-related potential 52 (ERP) studies found larger ERP amplitudes and a greater anterior distribution of N1 53 components in deaf individuals when they processed the visual stimulus of an 54 isoluminant color change (8).

55 The proposed mechanism behind this cross-modal reorganization is that long-56 term visual stimuli can lead to specialization of auditory cortex with engagement of 57 specialized neural networks for hearing and language tasks. The evidence obtained 58 from animal research and related literature review has also indicated the presence of 59 visual cross-modal reorganization of auditory cortex in animal models (9-11). The 60 presence of a visual-auditory modality in early life offers opportunities for change in 61 individual behavior and audiological rehabilitation (12). A recent systematic review 62 (12) of deaf induced cortical change showed that behavioral changes were 63 accompanied by a reorganization of multisensory areas, ranging from higher order 64 cortex to early cortical areas, highlighting cross-modal interactions as a fundamental 65 feature of brain organization and cognitive processing. It was considered that the auditory cortex might reorganize to mediate other functions, for example vision, in 66 67 areas of the superior temporal sulcus, just caudal to the primary auditory cortex with the result that deaf people show greater recruitment when processing visual, tactile orsigned stimuli than normal hearing individuals (12).

70 Cochlear implants (CIs) have been widely used as an effective intervention tool 71 for profound hearing impairment in children (13). Recent studies have indicated that 72 CI effect on neuroplasticity of the central auditory system occurs only when adequate 73 stimulation is delivered during a sensitive period in early childhood (14-17). Sharma 74 and Dorman (2006) examined P1 latency in 245 congenitally deaf children fitted with 75 CIs using evoked cortical potentials. They found that children had normal P1 latencies 76 if they received their CIs before the age of 3.5 years, whereas after this time children 77 showed abnormal or highly variable and delayed cortical response latencies (12,18). 78 In Sharma et al. (19), significantly delayed cortical P1 responses generated from 79 auditory thalamic and cortical areas were also found in children with CIs.

80 Cortical activity and visual cross-modal effects on the auditory cortex have been 81 reported to play a role in CI outcomes. Lee et al. (2007) found hypometabolism in the 82 temporal lobes of prelingually deaf children, speech scores post CI positively 83 associated with enhanced metabolic activity in the prefontal cortex which contributes 84 to auditory processing, and decreased metabolic activity in Heschle's gyrus which 85 contributes to visual processing (20). Sandmann et al. (21) used parametrically 86 modulated reversing checkerboard images to examine the initial stages of visual 87 processing and confirmed visual take-over in the auditory cortex of CI users. In 88 addition, the extent of visual processing in auditory cortices in postlingually deaf 89 subjects was negatively related to CI outcomes (21).

Further evidence has suggested that many factors are associated with plasticity and CI outcomes in prelingually deaf individuals, such as the age at which the CI was received, cognitive abilities, family environment, etiology, and speech-language

93 therapy. Of these factors, age at implantation contributes for most in terms of CI 94 outcome in prelingually deaf children (16), i.e. younger age children with CI would 95 achieve better speech outcomes. However, in Schramm et al. (22), although their 96 results showed CI patients with prelinguistic deafness achieved significantly better speech understanding using phonetically balanced monosyllabic words, there was a 97 98 wide range of performance across patients. They found that some older prelingually 99 deaf children with CI also performed well in speech communication (22). They 100 suggested this may be due to the various extent of visual cross-modal impact on the 101 auditory cortex. Because of uncertainty in the status of auditory cortex plasticity 102 without auditory stimuli before cochlear implantation, the effectiveness of CI 103 outcomes is unlikely to be predicted for CI candidates, particularly for prelingually 104 deaf children.

Recently, visual evoked potentials (VEPs) have been used to investigate visual-105 106 auditory cross-modality in patients with CIs. Visually evoked fronto-temporal N1 107 responses were reported to be related to visual processing in the auditory cortex (23-108 25). Kristi et al. (2011) reported that in postlingually deaf subjects, the higher N1 109 VEP responses in the right temporal lobe in children with a CI was related to poor 110 speech perception (25). Moreover, different visual stimuli, 'sound' photo vs. 'non-111 sound' photo, have been reported to produce different N1 responses in the fronto-112 temporal area; i.e., 'sound' photo stimuli evoked stronger N1 responses than 'non-113 sound' photo stimuli in normal (26).

To our best knowledge, N1 VEP response to 'sound' or 'non-sound' photo stimuli in prelingually deaf children still remains unclear. Moreover, the relationship of visual processing recruitment of auditory cortices and auditory outcomes in prelingually deaf children with CIs is unknown. Therefore, in the present study, we

examined the extent to which visual processing recruitment of auditory cortices occurred in prelingually deaf children with CIs. In addition, the relationship between the visual processing recruitment and auditory performance in these children was explored.

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123 Materials and Methods

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Participants

125 Ten prelingually deaf children bilaterally profound hearing loss were recruited 126 from special education schools for the deaf as the deaf group. There were five boys 127 and five girls, aged between 4 and 6 years (mean age and SD: 4.4 ± 0.7 years). Twenty 128 prelingually deaf children fitted with a CI to the right side for at least one year were 129 also recruited. The CIs fitted in this group of patients included: 10 MEDEL 130 SONATAti100, 3 Cochlear Freedom (CI24RE), 7 Advanced Bionics (AB) HiRes 120. 131 On the basis of their Category of Auditory Performance (CAP) score (22), they were 132 divided into two groups. Ten subjects (4 boys and 6 girls, mean age 4.6±0.90 years 133 old, range 3–6 years old) with CAP scores better than 5 were assigned to the CI good 134 performer group, the remaining 10 (4 boys and 6 girls, mean age 4.4±1.0 years old, 135 range 3–6 years old) with CAP scores less or equal to 5 were in the CI poor performer 136 group (14). Ten age and sex matched normal-hearing children were recruited as the 137 control group. Table 1 provides detailed demographic information, together with communication mode (i.e., using sign language or oral communication) and socio-138 139 economic status.

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Table 1 near here

Ethical approval was obtained from the Institutional Review Board at Sun Yatsen Memorial Hospital at Sun Yat-sen University. Detailed information was provided to the parents and, written consent obtained before proceeding with the study.

144

Visual stimuli

145 One 'sound' photo (i.e., a photograph with imaginative sound) and one 'non-146 sound' photo (i.e., a photograph without imaginative sound) were presented as visual 147 stimuli in a similar way to the study of Proverbio (26). The photographs were chosen 148 to ensure that most of the children were familiar with the images and understood their 149 meaning. Figure 1 shows the experimental block design, which consisted of an 150 intermittent stimulus mode using 'sound' photo and 'non-sound' photo stimuli. For 151 the 'sound' photo stimulus experiment, it consisted of 85 trials of 'sound' photo 152 stimuli, and 15 trials of 'non-sound' photo stimuli as deviant stimuli. In contrast, for the 'non-sound' photo stimulus experiment, it consisted of 85 trials of 'non-sound' 153 154 photo stimuli, and 15 trials of 'sound' photo stimuli as deviant stimuli. As shown in 155 Figure 1, each stimulus was presented for 1 second, followed by one blank screen 156 (1.7–1.9 seconds in duration) as the inter-stimulus. To make sure that the participants 157 concentrated on the stimuli, one novel that consisted of 15 photographs was presented after 5-10 trials and the children were asked to press a button while the deviant 158 159 photograph present.

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Figure 1 near here

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• VEPs measurement

162 ERPs were recorded from 128 scalp electrodes (Dense Array EEG System with 163 HydroCel Geodesic Sensor Nets (EGI, OR, USA)). After installation of the 128-164 channel electrophysiological cap, the test took place in a soundproof and electrically 165 shielded room. Each participant was asked to sit on a comfortable chair approximately 166 100 cm away from the 19-inch high-resolution VGA computer screen on which the 167 visual stimuli were presented. The participants were instructed to watch the screen 168 throughout the entire experiment, avoiding/minimizing body and eye movements. The 169 impedance for each electrode was kept below 40 k Ω during the experiment (17).

170 The ERP responses were recorded continuously using Net Station 4.3 (EGI, 171 USA) and analyzed off-line. The ERP signals were digitally filtered with a band-pass 172 of 0.1–30 Hz and signals with a segment of 700 ms, including 100 ms of pre-stimulus 173 baseline were collected. Any signal with an electro-oculography amplitude exceeding 174 $75 \,\mu V$ was excluded as an artifact likely caused by eye movements or eye blinks. An 175 amplitude exceeding 75 μ V at any electrode site was defined as a poor channel. If 176 there were six or more poor channels in a segment, then this segment was excluded as 177 a bad segment. If fewer than six poor channels were present, the segment was 178 considered valid and each poor channel was replaced with the average value obtained 179 from its surrounding channels. The response waveforms evoked by the visual stimuli 180 were obtained by averaging all valid segments. All responses at individual electrodes 181 were referred to the average reference (27). The baseline was corrected according to 182 the mean amplitude over the 100-ms pre-stimulus level.

All responses evoked by using either the 'sound' photo or 'non-sound' photo stimuli were recorded and averaged, respectively. **Figure 2** shows an example of ERP recordings obtained from an individual. The small-group average regions of interest were also analyzed (**Figure 3**). The N1 (the first negative response) at both FC3 (the left frontal-temporal area) and FC4 (the right frontal-temporal area) as well as the P1 (positive response occurring at approximately 170 ms) at Oz (the occipital area) were analyzed.

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Figure 2 near here

191 **Figure 3 near here** 192 **Statistical Analysis** 193 Multifactorial repeated-measures ANOVAs were performed for the ERP data 194 analysis. The within factors were the stimulus categories ('sound' photo and 'non-195 sound' photo) and electrode sites (FC3 for the left side and FC4 for the right side), 196 and the between factors were groups (deaf, poor CI performers, good CI performers, 197 poor CI performers, and Controls). The alpha inflation caused by multiple 198 comparisons was corrected using Greenhouse-Geisser corrections. The post-hoc 199 Tukey's test was also used for multiple comparisons. 200 201 **Results** 202 Clear N1 responses at both FC3 and FC4 were found in all groups. In addition, a P1 response was found at Oz. Figure 3 shows an example of the N1 and P1 responses 203 204 obtained from children in four groups when using 'sound' photo and 'non-sound' 205 photo stimuli. 206 A 3-way RM-ANOVA was used with one between-subject factor (groups: Deaf, 207 Poor CI performers, good CI performers, and Control) and two within-subject factors 208 (stimuli: 'sound' and 'non-sound'; electrode sites: FC3 and FC4) for N1 amplitudes and latencies. Additionally, a 2-way RM-ANOVA was used with one between-subject 209 210 factor (group: Deaf, poor CI performers, Good CI performers, and control) and one 211 within-subject factor (stimulus: 'sound' and 'non-sound') for P1 amplitudes and 212 latencies. 213 Significant effects were obtained for group (F=44.747, p<0.001) and stimulus 214 ('sound' photo > 'non-sound' photo, F=17.282, p<0.001) referring to N1 amplitudes

215 (Figure 4). Group *stimulus* site interaction effects were also found to be significant

216 (F=5.483, p=0.003). No significant main effect was found for electrode sites of FC3 217 and FC4 (F=0.013, p=0.909).

218 A pairwise comparison found that N1 amplitudes in the deaf group were 219 significantly larger than in the poor CI performers, good CI performers and control groups (P=0.008, p <0.001, and p<0.001, respectively). N1 amplitudes in the poor CI 220 221 performers group were significantly larger than those in the good CI performers and 222 normal groups (p < 0.001 and p < 0.001, respectively). No significant difference was 223 found between the control and good CI performers groups (p=0.893). 224 Figure 4 near here 225 When comparing the effect of different stimuli, 'sound' photo evoked stronger 226 responses than 'non-sound' photo at FC4 in the deaf and poor CI performers groups 227 (F=8.82, p=0.005 and F=23.17, p<0.001, respectively) (Figure 5), but not in the good 228 CI performers and control groups. 229 Figure 5 near here

With respect to N1 latencies, the main effects were obtained for electrode sites (FC4 149.3 vs. 142.8 FC3, F=7.538, p=0.009) and stimuli ('sound' photo 148.9 vs. 143.2 'non-sound photo, F=10.787, p=0.002). No significant main effect was found for the variable group (F=0.781, p=0.512). In addition, group*stimulus, group*site and stimulus*site interactions were not significant (F=2.409, p=0.083; F=0.879, p=0.461; and F=1.454, p=0.236, respectively).

With respect to P1 latencies and amplitudes, no significant main effect was found for the variable group (F=0.781, p=0.512 for latency; F=2.409, p=0.083 for amplitude). In addition, the group*stimulus interaction was not significant (F=2.409, p=0.083; F=0.879, p=0.461, and F=1.454, p=0.236, respectively).

241 **Discussion**

242 The present study examined visual processing recruitment of auditory cortex in 243 prelingually deaf children with and without CIs in comparison to hearing controls. 244 'Sound' and 'non-sound' photos were used as the visual stimuli for VEP 245 measurements. The advantage of using images associated with sounds is enhancement 246 of visual activation of auditory cortex. Previous studies have shown a significantly 247 larger P1 amplitude at the occipital midline in adults with mild-moderate hearing loss 248 than controls when using a kind of visual stimulus called 'high contrast sinusoidal 249 concentric grating' (28). Consequently, they suggested that visual enhancement in the 250 occipital area is likely to be associated with better visual sensitivity in people with 251 hearing impairment. Moreover, by using 'sound' photo and 'non-sound' photo stimuli, 252 Proverbio et al. (26) found different ERP responses, i.e., strong N1 response in the 253 frontal area and weak response in the occipital area, when compared with using visual 254 motion stimuli, i.e., a strong N1 response in the occipital area and a weak response in 255 the frontal area (2,25). Further comparison showed that the N1 response evoked by 256 using the 'sound' photo was even greater than using 'non-sound' photo in the frontal 257 area, which can be used as an indicator of auditory cortical recruitment by 'sound' photo visual stimuli. 258

In the present study, the prelingually deaf children without CIs had significantly greater N1 VEP amplitudes in response to the visual stimuli (both 'sound' and 'nonsound' photo stimuli) than the children with CIs and controls. Further analysis showed that N1 amplitudes were largest in the deaf children, followed by those with poorly performing CIs, controls and those with well-performing CIs, whilst there was no significant difference between controls and those with well-performing CIs. However, Buckley et al. (25), reported that only N1 VEP amplitudes from the right temporal lobe were negatively related to speech perception in prelingually deaf children with CIs when they used the stimuli of moving visual gradients located in a square pattern on a gray background with still pictures of cartoon characters. Differences in the stimulus category of the two studies may be responsible for the discrepancy between the two outcomes (25,26,29). Buckley et al. (25) used a vision motion stimulus in the peripheral visual field, while in the present study, we presented the stimuli centrally, which produced bilateral N1 response enhancement.

273 Furthermore, as shown in Figure 4, children who used a CI had lower N1 VEP 274 amplitudes than deaf children, while those with well-performing CIs had lower N1 275 amplitudes than poor CI performers and similar N1 amplitudes as children with 276 normal hearing. Although recruitment of auditory cortices evoked by the visual 277 system to process the visual photos were found in deaf children with CI, the present 278 result implies that there is a negative relationship between the process and CI outcomes. As indicated previously, visual cross-modal take-over has been 279 280 demonstrated in postlingually deaf adults, which is related to the auditory 281 performance of the patients after receiving a CI (30,31). The adaption process after a 282 CI procedure may indicate a reversal of auditory functional take-over, while 283 insufficient adaptation to the new input may be reflected by residual signs of visual 284 take-over (31,32). In the present study, the positive relationship between the 285 decrement of the N1 amplitudes and CI outcomes may demonstrate the reversal of 286 auditory functional take-over. Further studies are needed to determine the relationship between decrement of N1 amplitude and the auditory performances in deaf children. 287

The other interesting finding obtained from the present study is that 'sound' photo evoked greater N1 amplitude compared to 'non-sound' photo, which is consistent with the findings of Proverbio et al. (26). However responses evoked by

291 using 'sound' photo were greater than using 'non-sound' photo only at FC4 in the deaf 292 and poor CI performers, but not in the good CI performers and controls. Buckley et al. 293 (25) found that the amplitudes of N1 VEP responses in the right temporal area were 294 negatively related to the speech performances of the CI patients. It is considered that 295 the left and right temporal lobes play different roles in processing auditory 296 information. The right lobe mainly participates in speech perception tasks in subjects with normal hearing and varies according to the degree of residual hearing. Right 297 298 temporal lobe structures can be recruited for speech perception processing if the 299 speech signal is degraded (33) and seems to be important for underlying meaning in 300 message extraction (34). However, the left temporal lobe mainly processes fine 301 structures of speech signals (35). In addition, several studies with deaf individuals and 302 CI users have shown that the effect of deprivation-induced cross-modal plasticity has 303 primarily been localized to the right hemisphere (4,31,35-37), either because the right 304 hemisphere is more susceptible to reorganizational changes compared with the left 305 hemisphere (37) or because the right hemisphere is more involved in the processing of 306 sounds with low complexity (38).

307 It is noteworthy that the present results were only obtained from the participants 308 with a CI on the right side. Although bilateral CIs are generally recommended for children with bilateral sever to profound hearing impairment, due to their 309 310 affordability, a majority of the suitable candidates were only fitted with a CI 311 unilaterally. It is interesting to investigate the similarity or significant difference in 312 terms of the effects on visual processing recruits the auditory cortices in comparison 313 of children with a unilateral CI (on either right ear or left ear) and those with bilateral 314 CIs in the future study.

315

316 Conclusions

317 The influence of visual processing recruitment of the auditory cortices is evident as 318 there were stronger N1 VEP responses in prelingually deaf children and there were 319 decrements in this recruitment in children with a CI. The recruitment decrement was 320 related to good CI outcomes. Consideration of the bilateral N1 response to the visual 321 stimuli, and also the difference in the frontal response to the 'sound' photo and 'nonsound' photo in prelingually deaf children without and with CI, the 'sound and non-322 323 sound' indicates that photos are feasible for the studying of visual recruitment of 324 auditory cortex. Further exploration and follow-up studies to determine visual impacts 325 on auditory cortices and their influence on auditory outcomes with a CI are needed.

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330 **References**

- 331
- Kujala T, Alho K, Naatanen R. Cross-modal reorganization of human cortical functions. *Trends in neurosciences* 2000;23:115-20.
- 334 2. Doucet ME, Bergeron F, Lassonde Met al. Cross-modal reorganization and
 335 speech perception in cochlear implant users. *Brain : a journal of neurology*336 2006;129:3376-83.
- 337 3. Sadato N, Okada T, Honda Met al. Cross-modal integration and plastic
 338 changes revealed by lip movement, random-dot motion and sign languages in
 339 the hearing and deaf. *Cerebral cortex* 2005;15:1113-22.
- Finney EM, Fine I, Dobkins KR. Visual stimuli activate auditory cortex in the
 deaf. *Nature neuroscience* 2001;4:1171-3.
- 5. Fine I, Finney EM, Boynton GMet al. Comparing the effects of auditory deprivation and sign language within the auditory and visual cortex. *J Cogn Neurosci* 2005;17:1621-37.
- Bavelier D, Neville HJ. Cross-modal plasticity: Where and how? *Nature reviews. Neuroscience* 2002;3:443-52.
- Finney EM, Clementz BA, Hickok Get al. Visual stimuli activate auditory
 cortex in deaf subjects: evidence from MEG. *Neuroreport* 2003;14:1425-7.
- 349 8. Armstrong BA, Neville HJ, Hillyard SAet al. Auditory deprivation affects
 350 processing of motion, but not color. *Brain research. Cognitive brain research*351 2002;14:422-34.
- 352 9. Kral A, Sharma A. Developmental neuroplasticity after cochlear implantation.
 353 *Trends in neurosciences* 2012;35:111-22.
- Land R, Baumhoff P, Tillein Jet al. Cross-Modal Plasticity in Higher-Order
 Auditory Cortex of Congenitally Deaf Cats Does Not Limit Auditory
 Responsiveness to Cochlear Implants. *The Journal of neuroscience : the official journal of the Society for Neuroscience* 2016;36:6175-85.
- Kok MA, Chabot N, Lomber SG. Cross-modal reorganization of cortical afferents to dorsal auditory cortex following early- and late-onset deafness. *The Journal of comparative neurology* 2014;522:654-75.
- 361 12. Bavelier D, Dye MW, Hauser PC. Do deaf individuals see better? *Trends in cognitive sciences* 2006;10:512-8.
- 363 13. Ponton CW, Don M, Eggermont JJet al. Maturation of human cortical auditory
 364 function: Differences between normal-hearing children and children with
 365 cochlear implants. *Ear and hearing* 1996;17:430-7.
- 366 14. Giraud AL, Price CJ, Graham JMet al. Cross-modal plasticity underpins
 367 language recovery after cochlear implantation. *Neuron* 2001;30:657-63.
- Pantev C, Paraskevopoulos E, Kuchenbuch Aet al. Musical expertise is related
 to neuroplastic changes of multisensory nature within the auditory cortex. *The European journal of neuroscience* 2015;41:709-17.
- Sharma A, Campbell J, Cardon G. Developmental and cross-modal plasticity
 in deafness: evidence from the P1 and N1 event related potentials in cochlear
 implanted children. *International journal of psychophysiology : official journal of the International Organization of Psychophysiology* 2015;95:13544.
- 17. Liang M, Zhang X, Chen Tet al. Evaluation of auditory cortical development
 in the early stages of post cochlear implantation using mismatch negativity
 measurement. Otology & neurotology : official publication of the American

- 379 Otological Society, American Neurotology Society [and] European Academy
 380 of Otology and Neurotology 2014;35:e7-14.
- 381 18. Sharma A, Dorman MF. Central auditory development in children with
 382 cochlear implants: clinical implications. *Advances in oto-rhino-laryngology*383 2006;64:66-88.
- 384 19. Sharma A, Martin K, Roland Pet al. P1 latency as a biomarker for central auditory development in children with hearing impairment. *Journal of the American Academy of Audiology* 2005;16:564-73.
- 20. Lee HJ, Giraud AL, Kang Eet al. Cortical activity at rest predicts cochlear
 implantation outcome. *Cerebral cortex* 2007;17:909-17.
- 389 21. Sandmann P, Dillier N, Eichele Tet al. Visual activation of auditory cortex
 390 reflects maladaptive plasticity in cochlear implant users. *Brain : a journal of*391 *neurology* 2012;135:555-68.
- 392 22. Schramm D, Fitzpatrick E, Seguin C. Cochlear implantation for adolescents
 393 and adults with prelinguistic deafness. *Otology & neurotology : official*394 *publication of the American Otological Society, American Neurotology Society*395 [and] European Academy of Otology and Neurotology 2002;23:698-703.
- Stropahl M, Chen LC, Debener S. Cortical reorganization in postlingually deaf
 cochlear implant users: Intra-modal and cross-modal considerations. *Hearing research* 2016.
- Bottari D, Heimler B, Caclin Aet al. Visual change detection recruits auditory
 cortices in early deafness. *NeuroImage* 2014;94:172-84.
- 401 25. Buckley KA, Tobey EA. Cross-modal plasticity and speech perception in pre-402 and postlingually deaf cochlear implant users. *Ear and hearing* 2011;32:2-15.
- 403 26. Proverbio AM, D'Aniello GE, Adorni Ret al. When a photograph can be heard:
 404 vision activates the auditory cortex within 110 ms. *Sci Rep* 2011;1:54.
- 405 27. Jung J, Morlet D, Mercier Bet al. Mismatch negativity (MMN) in multiple
 406 sclerosis: an event-related potentials study in 46 patients. *Clinical*407 *neurophysiology : official journal of the International Federation of Clinical*408 *Neurophysiology* 2006;117:85-93.
- 409 28. Campbell J, Sharma A. Cross-modal re-organization in adults with early stage
 410 hearing loss. *PloS one* 2014;9:e90594.
- 411 29. Näätänen R, Winkler I. The concept of auditory stimulus representation in cognitive neuroscience. *Psychological bulletin* 1999;125:826-59.
- 413 30. Strelnikov K, Rouger J, Demonet JFet al. Visual activity predicts auditory
 414 recovery from deafness after adult cochlear implantation. *Brain : a journal of*415 *neurology* 2013;136:3682-95.
- 416 31. Lee DS, Lee JS, Oh SHet al. Cross-modal plasticity and cochlear implants.
 417 *Nature* 2001;409:149-50.
- 418 32. Liikkanen LA, Tiitinen H, Alku Pet al. The right-hemispheric auditory cortex
 419 in humans is sensitive to degraded speech sounds. *Neuroreport* 2007;18:601-5.
- 420 33. Meyer M, Alter K, Friederici ADet al. FMRI reveals brain regions mediating
 421 slow prosodic modulations in spoken sentences. *Human brain mapping*422 2002;17:73-88.
- 423 34. Friederici AD, Alter K. Lateralization of auditory language functions: a
 424 dynamic dual pathway model. *Brain and language* 2004;89:267-76.
- 425 35. Lyness CR, Woll B, Campbell Ret al. How does visual language affect
 426 crossmodal plasticity and cochlear implant success? *Neuroscience and*427 *biobehavioral reviews* 2013;37:2621-30.
- 428 36. Rouger J, Lagleyre S, Demonet JFet al. Evolution of crossmodal

- reorganization of the voice area in cochlear-implanted deaf patients. *Human brain mapping* 2012;33:1929-40.
- 431 37. Lazard DS, Lee HJ, Truy Eet al. Bilateral reorganization of posterior temporal
 432 cortices in post-lingual deafness and its relation to cochlear implant outcome.
 433 *Human brain mapping* 2013;34:1208-19.
- 434 38. Hine J, Debener S. Late auditory evoked potentials asymmetry revisited.
 435 *Clinical neurophysiology : official journal of the International Federation of* 436 *Clinical Neurophysiology* 2007;118:1274-85.