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Alternative Tinnitus Management Techniques Developed from Volitional Control Over the Activity of the Auditory Cortex

Matthew S. Sherwood, PhD

Emily E. Diller, MS; Subhashini Ganapathy, PhD;
Jeremy T. Nelson, PhD; Jason G. Parker, PhD



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Tinnitus

- Lots of Debate
- Difficult to Study
 - Subjective
 - Highly Variable
- Various Models
 - Bottom-up Deafferentation
 - Top-Down Deficiencies
- Critical Gap in Treatment Technologies/Therapies

Our Therapeutic Idea

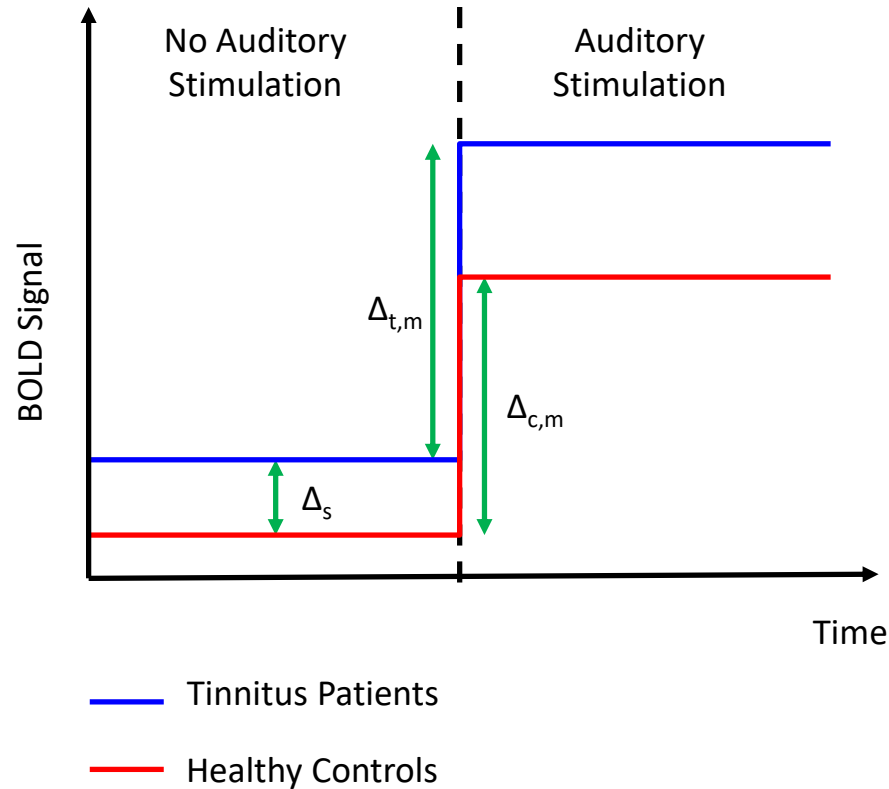
Phase 1

- Train control over the activity of the primary auditory cortex using functional MRI neurofeedback
- Assess changes in behavioral and neural measures
- Determine control methods resulting in greatest behavioral/neural effects

Phase 2

- Develop a simple application to train control methods in the absence of neurofeedback
- Determine changes in behavioral and neural measures
- Compare effects with functional MRI neurofeedback

Findings from Functional Neuroimaging



- $\Delta_{t,m} - \Delta_{c,m} > 0$: Mean signal change between auditory stimulation and no stimulation is elevated in tinnitus patients^{1,2}
- Δ_s : elevated steady-state metabolism^{3,4}
 - Should result in increased BOLD signal and CBF
- Attentional, emotional and auditory network components are altered^{5,6}

¹ Gu et al. (2010). Journal of Neurophysiology, 104(6), 3361-3370.

² Seydell-Greenwald et al. (2012). Brain Research, 1485, 22-39.

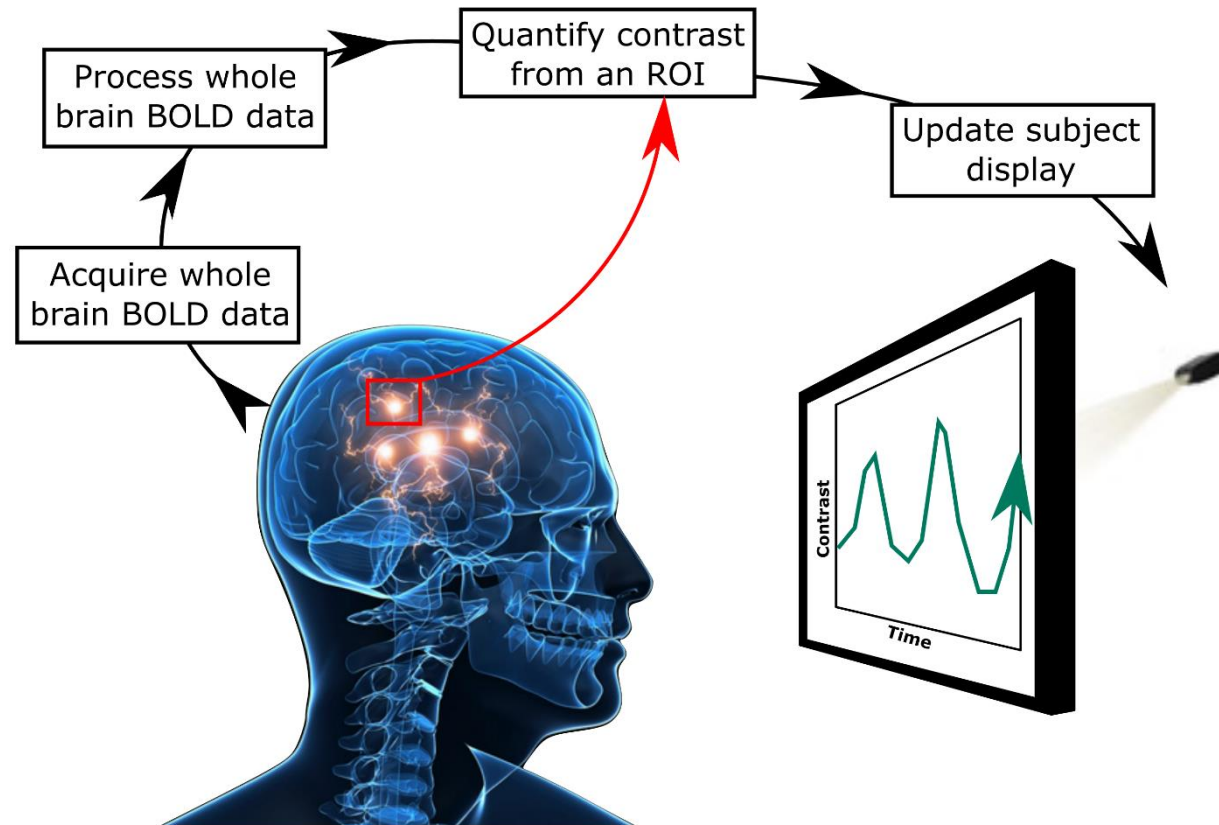
³ Wang et al. (2001). Chinese Medical Journal, 114(8), 848-851.

⁴ Schecklmann et al. (2013). Human Brain Mapping, 34(1), 233-240.

⁵ Kim et al. (2012). International Journal of Audiology, 51(5), 413-417.

⁶ Maudoux et al. (2012). PLoS ONE, 7(5), e36222.

Real-time fMRI Neurofeedback



Phase 1: Experimental Methods

27 Healthy Participants

- Written informed consent obtained prior to any experimental procedures

Grouping

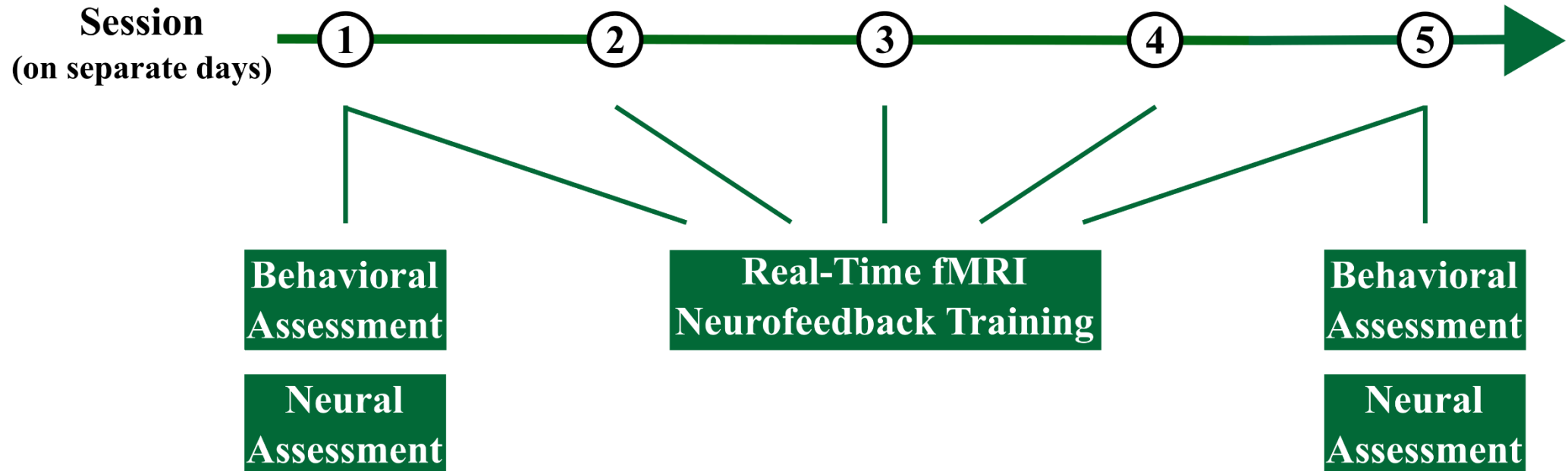
Experimental Group (EXP)

- $n = 18$ (mean age 23.2 ± 1.1 years, 11 males)
- Supplied real neurofeedback from Primary Auditory Cortex (A1)

Control Group (CON)

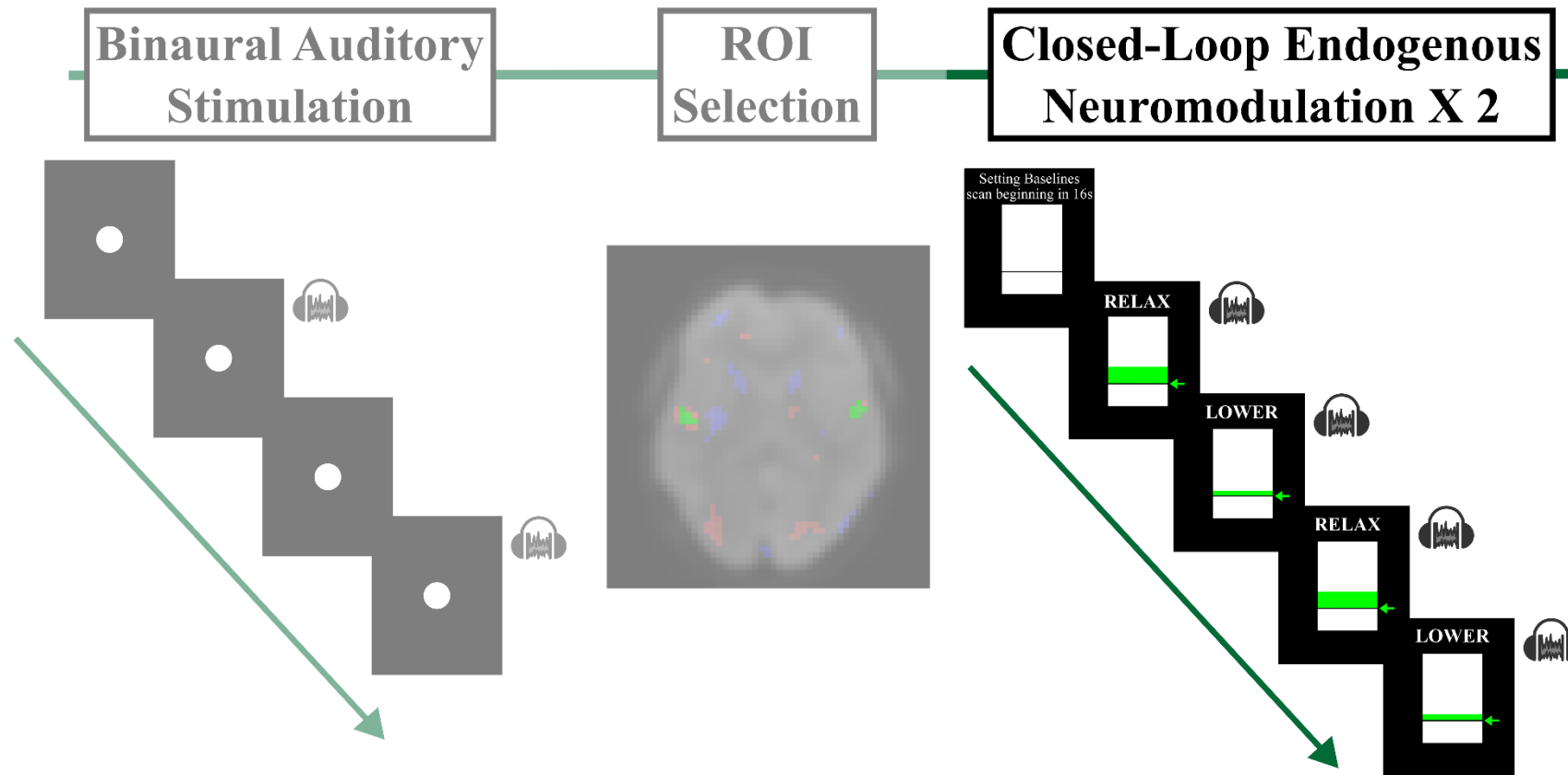
- $n = 9$ (mean age 24.4 ± 2.5 years, 4 males)
- Supplied sham neurofeedback

Phase 1: Experimental Methods (cont.)

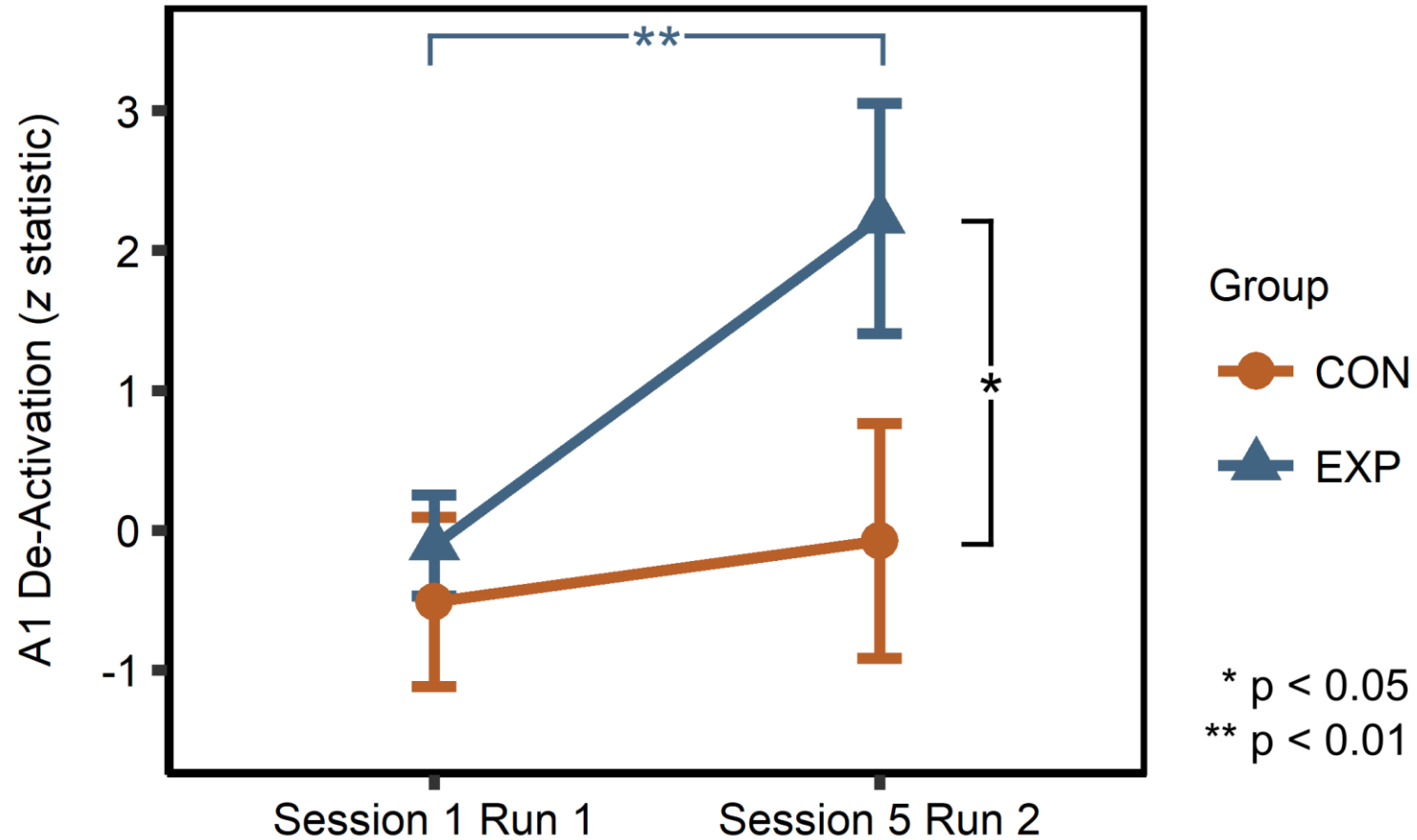


Experimental Methods (cont.)

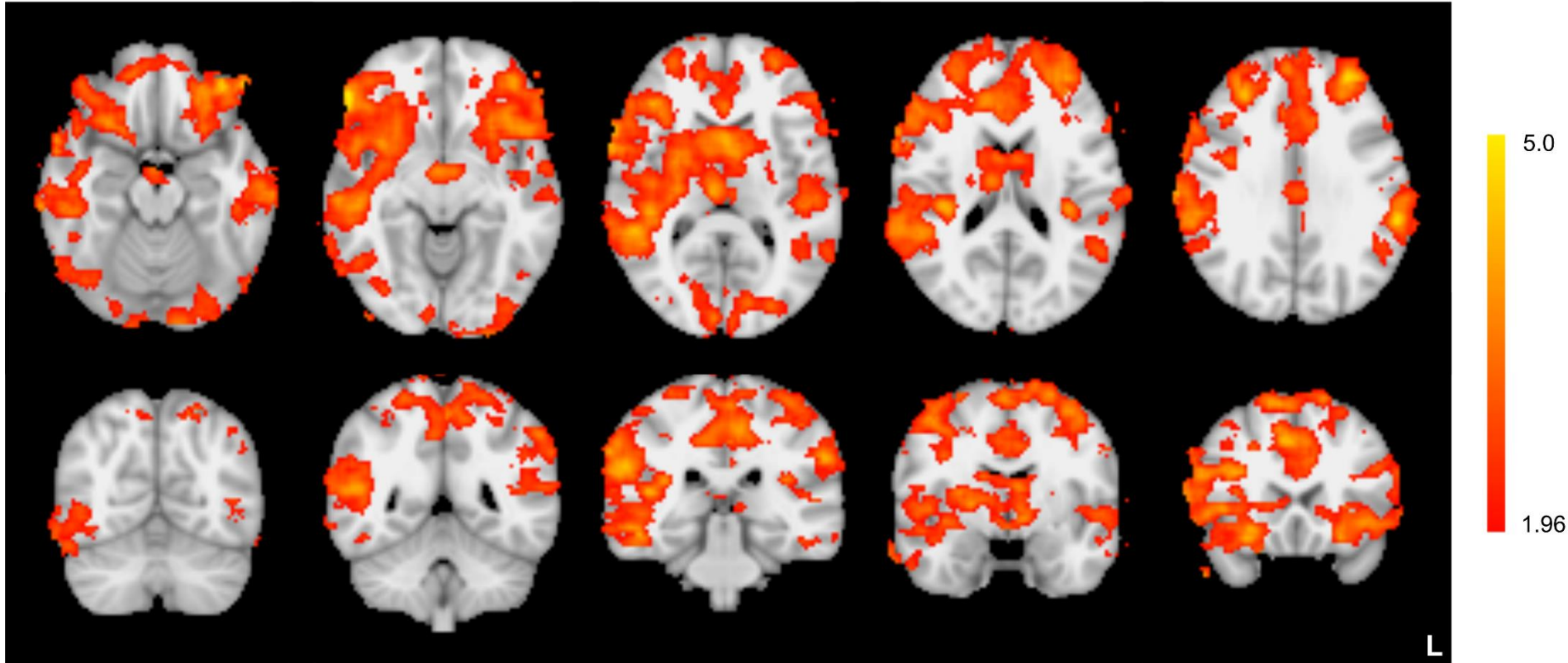
Neurofeedback Training



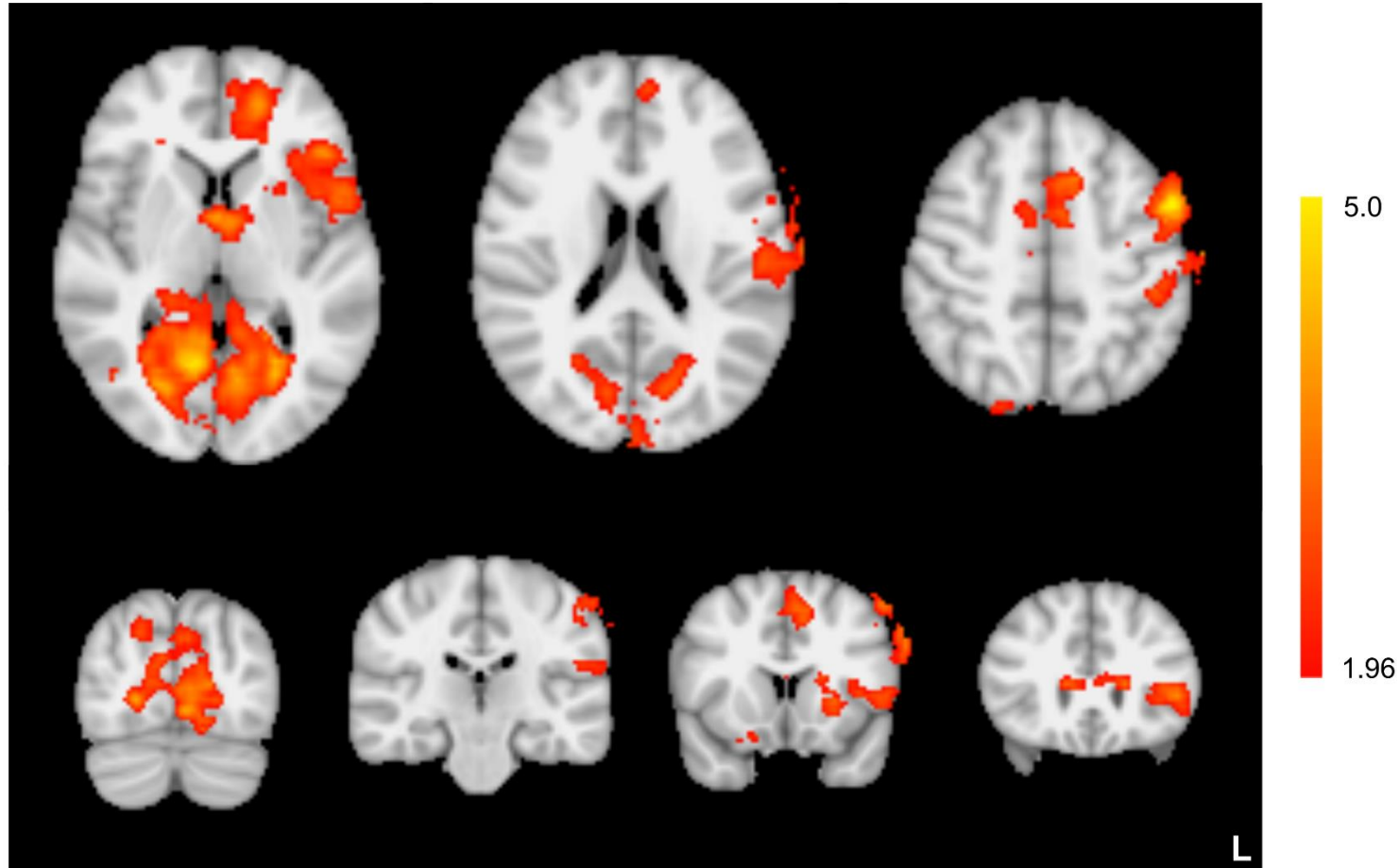
Phase 1: Results



Phase 1: Results (cont.)



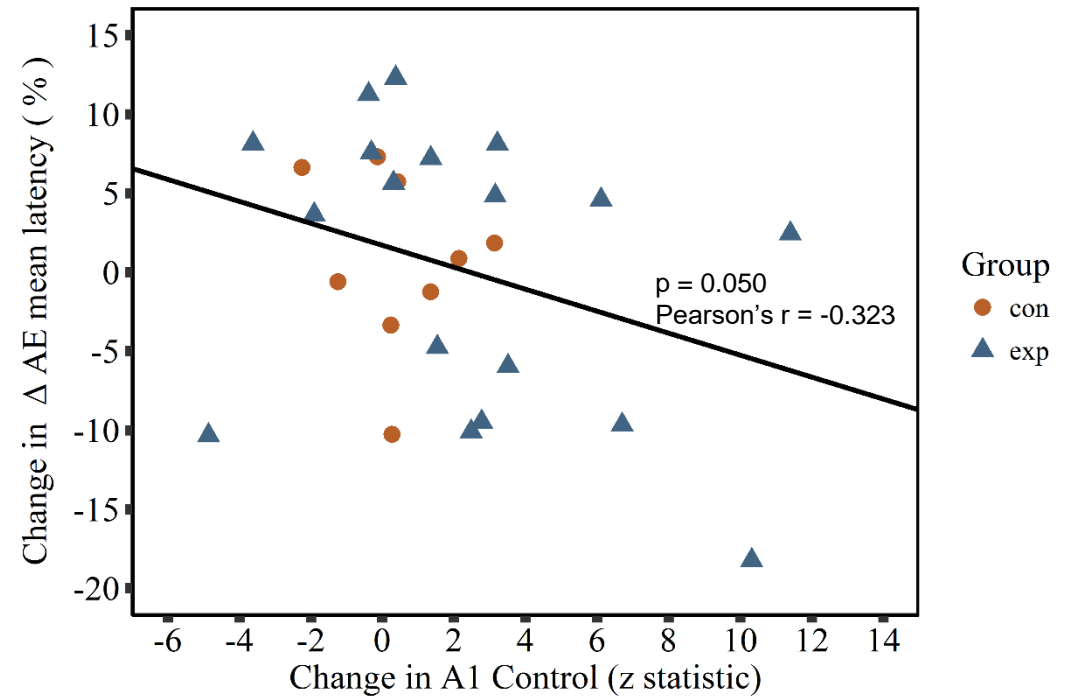
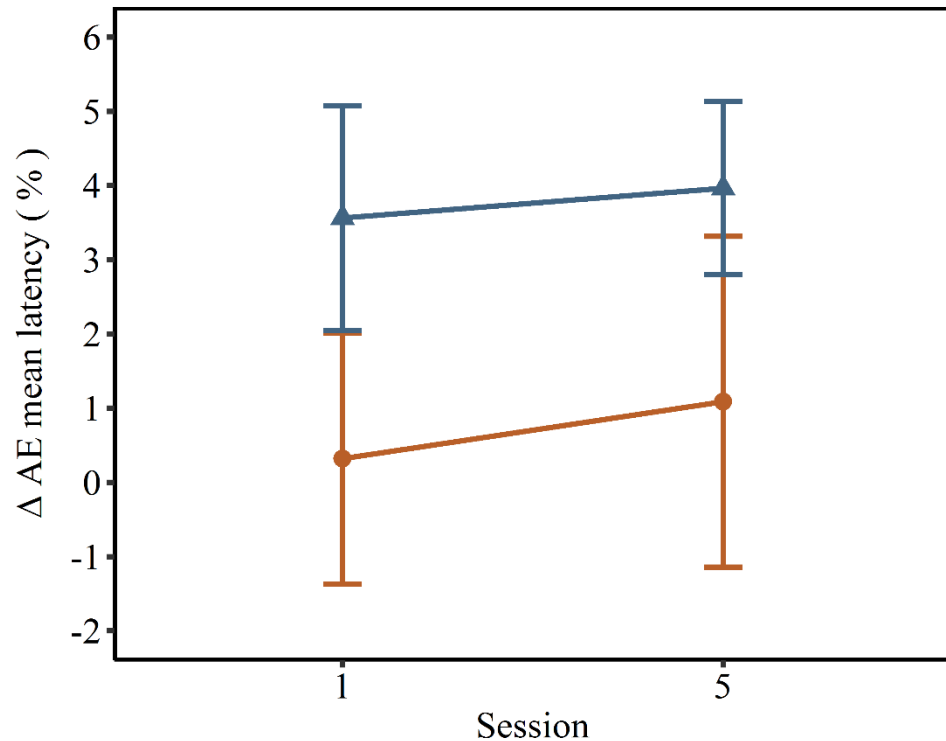
Phase 1: Results (cont.)



Phase 1: Results (cont.)

Attention to Emotion Task

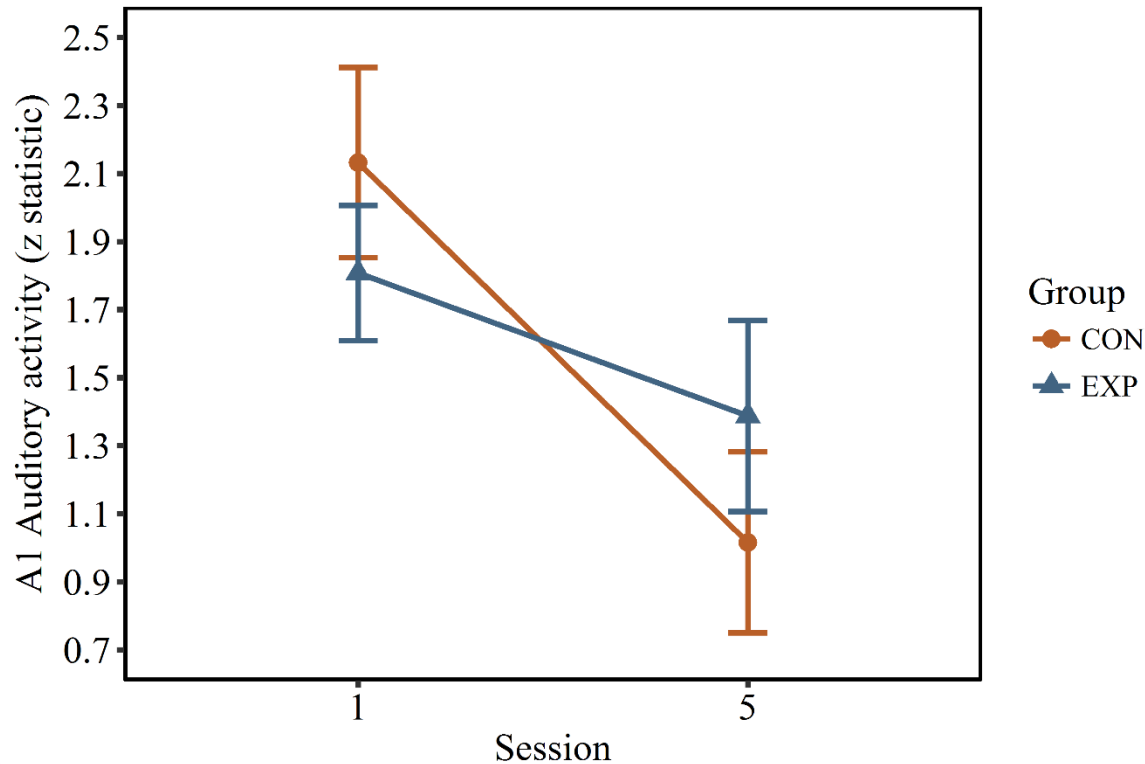
- Assessed impact of emotion on attention
- Calculated the percent change in latency between emotional and neutral trials



Phase 1: Results (cont.)

A1 Response to Auditory Stimulation

- Assessed A1 activation from binaural continuous noise
- Performed a 2x2x2 repeated measures ANOVA

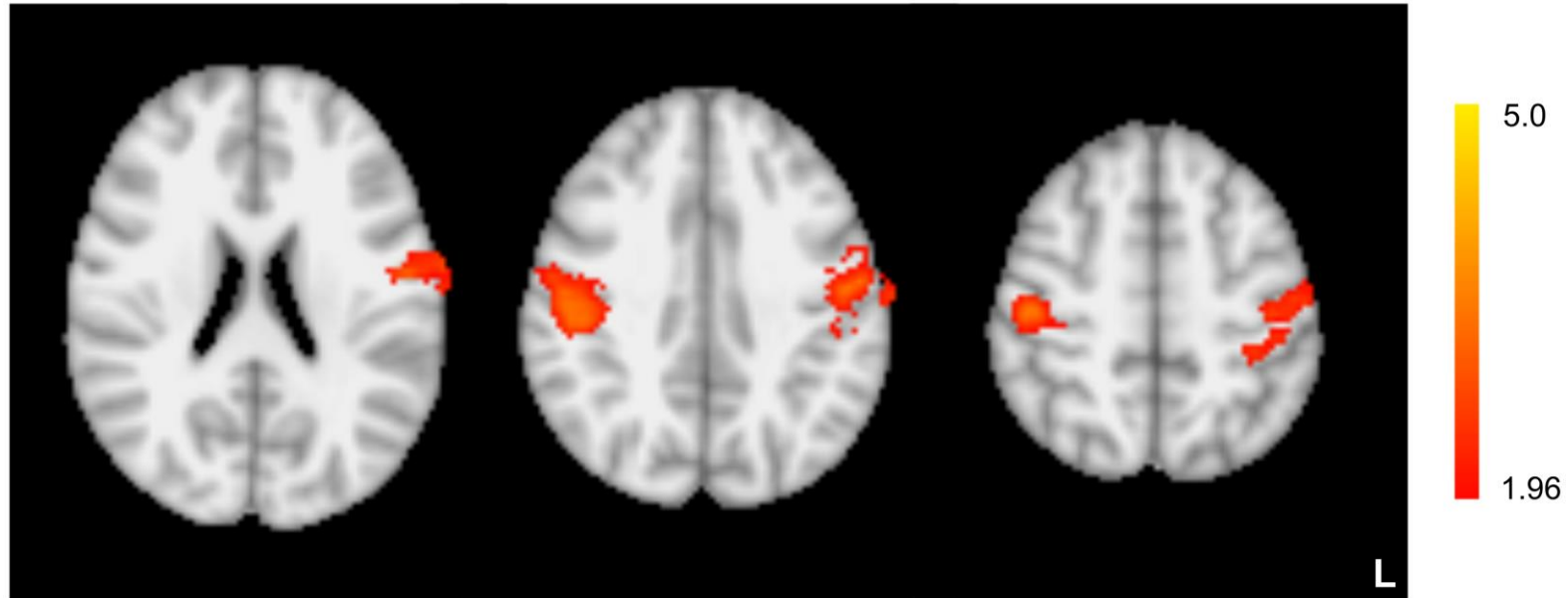


Source	Sig. (one-tailed)
Intercept	.000
Group	.478
Error	

Factor	Sig. (one-tailed)
Session	0.0115
Session * Group	0.142
Hemisphere	0.2205
Hemisphere * Group	0.1635
Session * Hemisphere	0.0975
Session * Hemisphere * Group	0.217

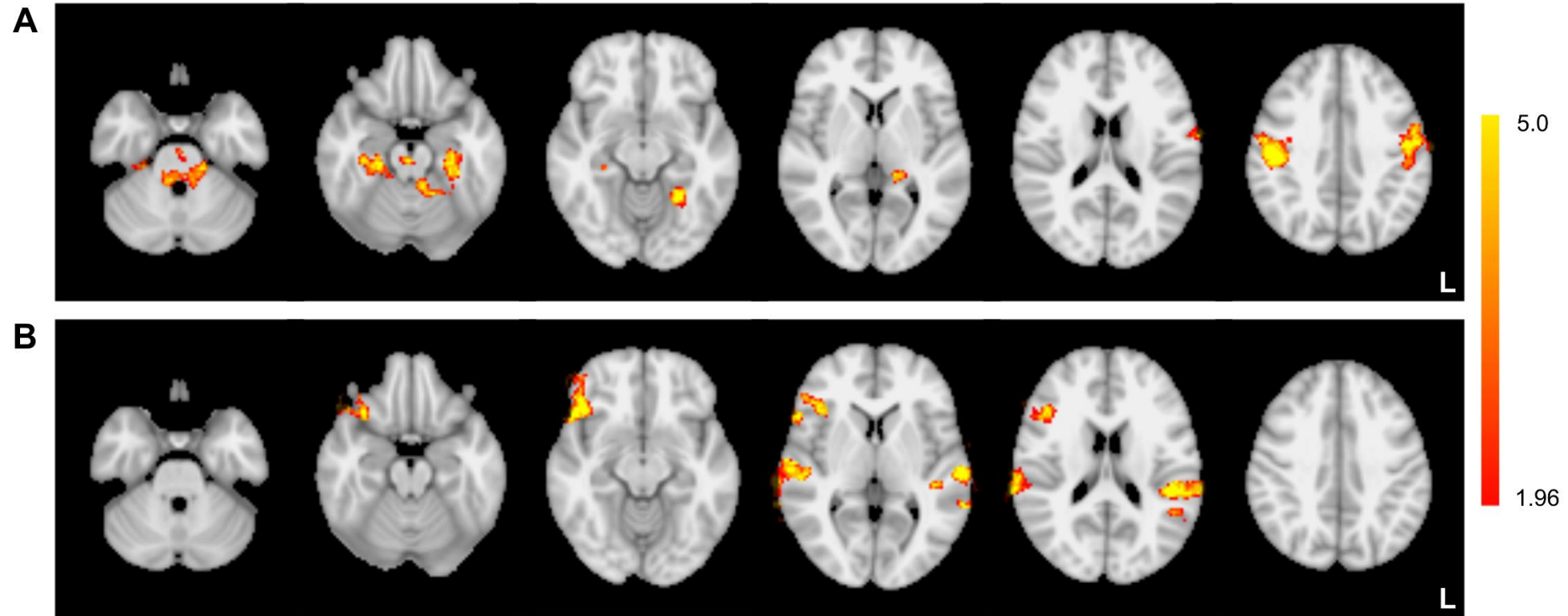
Phase 1: Results (cont.)

Response to Auditory Stimulation



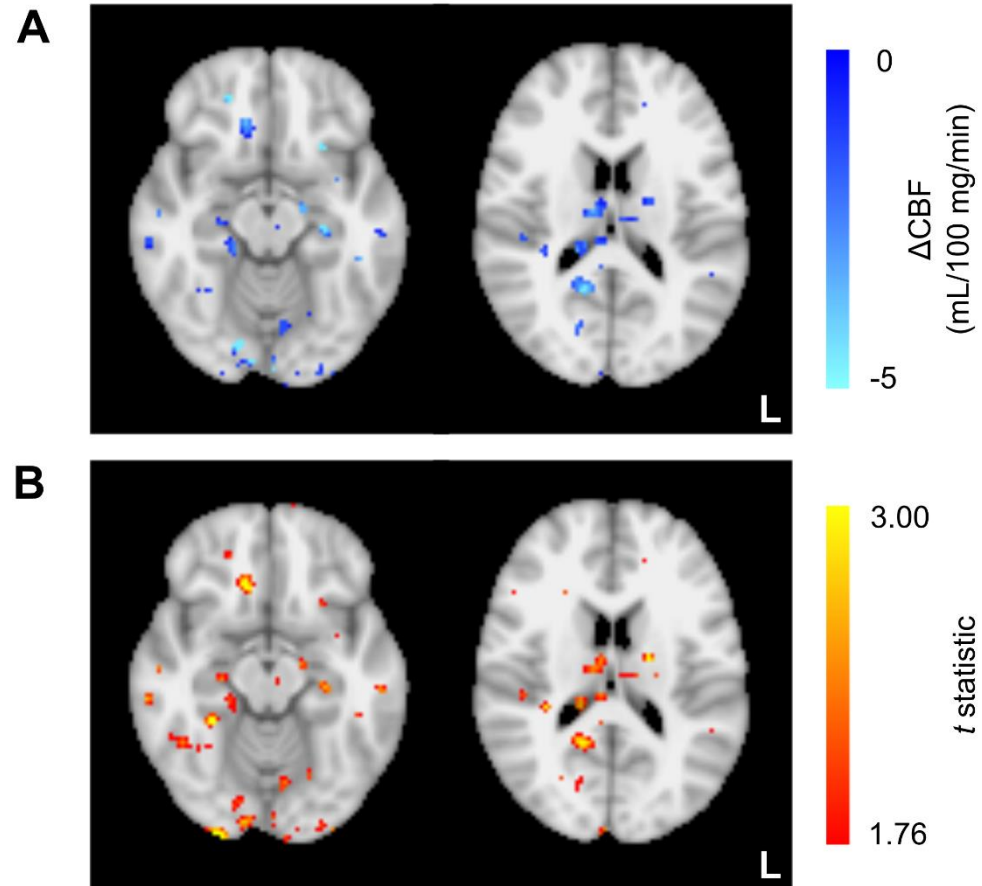
Phase 1: Results (cont.)

Response to Auditory Stimulation



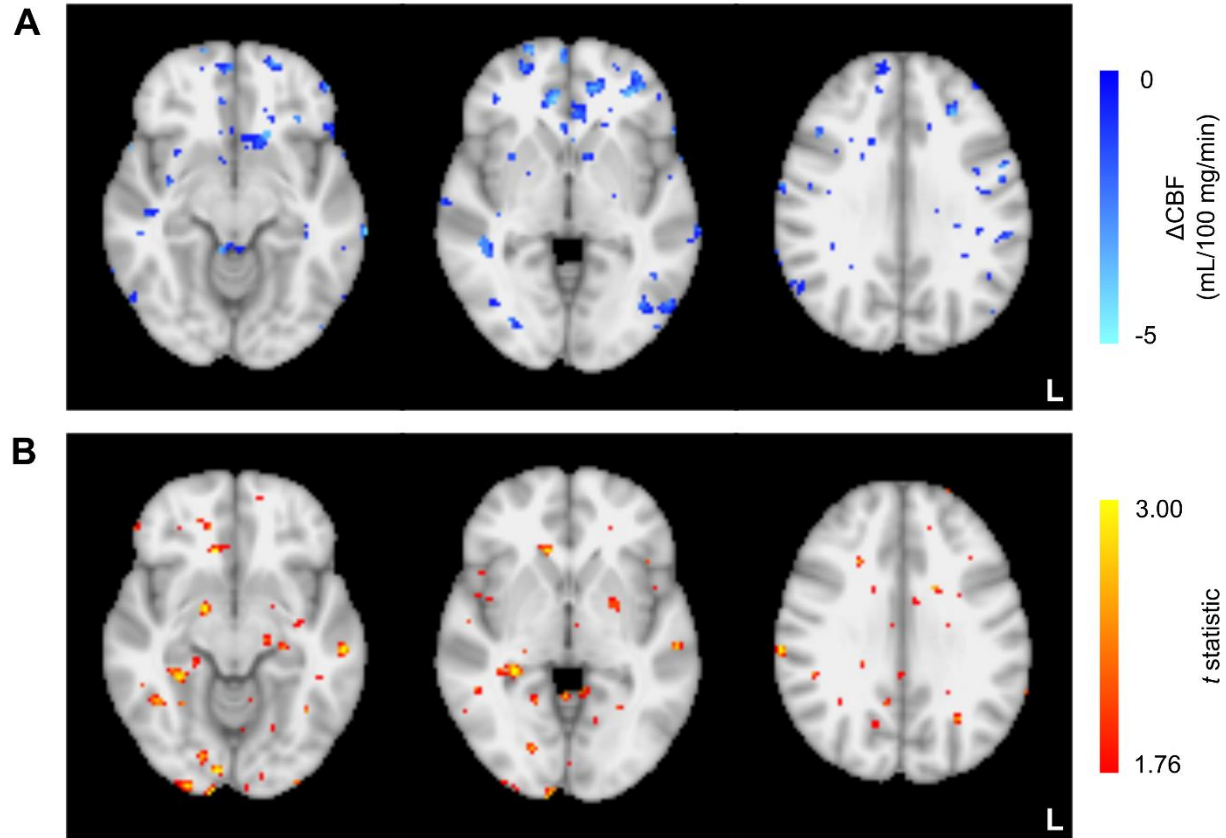
Phase 1: Results (cont.)

Steady-State Perfusion



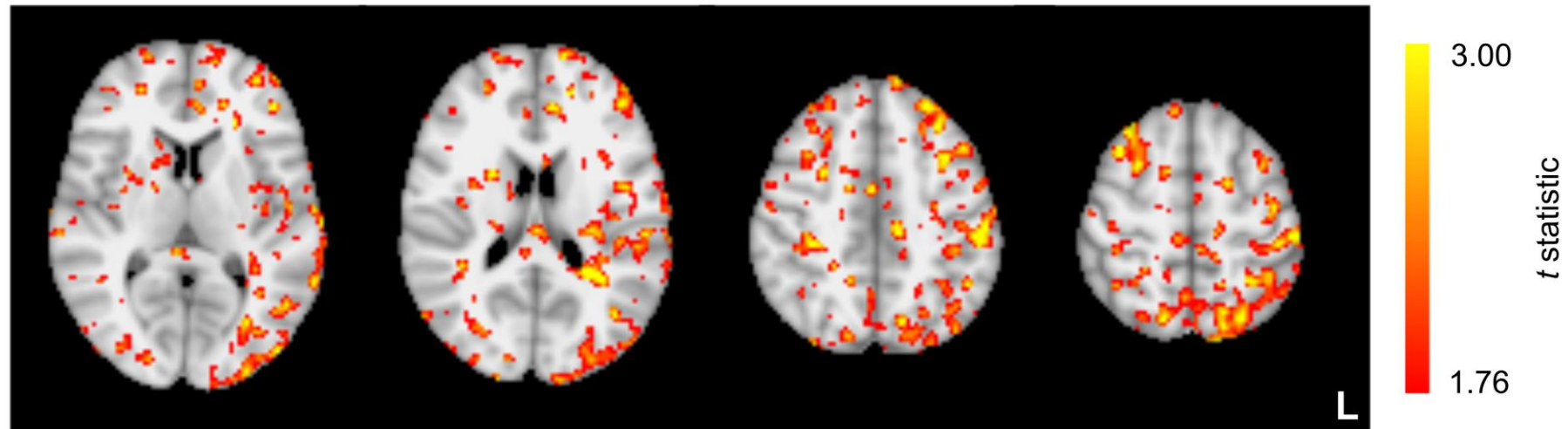
Phase 1: Results (cont.)

Steady-State Perfusion



Phase 1: Results (cont.)

Steady-State Perfusion



Phase 1: Discussion

- Previously, self-regulation via fMRI-NFT of the following have been implicated:
 - Activated cortical volume in A1 and the secondary auditory cortex¹
 - A1 activation magnitude²
- This work indicates self-regulation over A1 deactivation magnitude is achievable using fMRI-NFT in the presence of auditory stimulation

¹ Yoo et al. (2006). NeuroReport, 17(12), 1273-1278.

² Haller et al. (2010). European Radiology, 20(3), 696-703.

Phase 1: Discussion (cont.)

- Enhanced emotional response to auditory stimulation has been reported in tinnitus patients^{1,2}
 - Improved control over A1 led to a decreased effect of emotionally-charged stimuli on attention
- Tinnitus patients exhibit an elevated response in A1 activation to auditory stimuli^{3,4}
 - Attempting volitional control over A1 was found to decrease the A1 activation arising from auditory stimuli
- Tinnitus patients exhibit elevated steady-state metabolism in A1
 - Observed increased perfusion in middle temporal gyrus and superior temporal gyrus

¹ Wunderlich et al. (2010). *Audiology and Neurotology*, 15(3), 137-148.

² Golm et al. (2013). *Hearing Research*, 295, 87-99.

³ Gu et al. (2010). *Journal of Neurophysiology*, 104(6), 3361-3370.

⁴ Seydell-Greenwald et al. (2012). *Brain Research*, 1485, 22-39.

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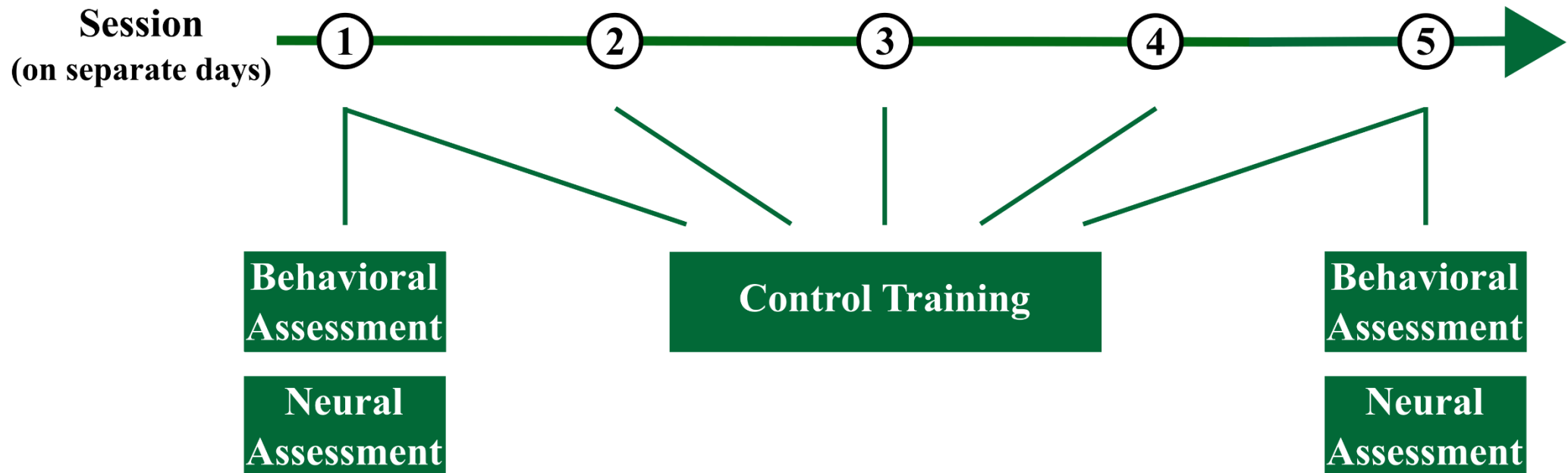
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⁴ Seydell-Greenwald et al. (2012). *Brain Research*, 1485, 22-39.

Phase 2: Experimental Methods



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