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Optimization of Transcutaneous Vagus Nerve Stimulation Using Functional MRI.

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Abstract

OBJECTIVE/HYPOTHESIS: Vagus nerve **stimulation** (VNS) is an established therapy for drug-resistant epilepsy, depression, and a number of other disorders. Transcutaneous **stimulation** of the **auricular** branch of the vagus nerve (tVNS) has been considered as a non-invasive alternative. Several **functional magnetic resonance** imaging (fMRI) studies on the effects of tVNS used different **stimulation** parameters and locations in the ear, which makes it difficult to determine the optimal tVNS methodology. The present study used fMRI to determine the most effective location for tVNS.

MATERIALS AND METHODS: Four **stimulation** locations in the ear were compared: the inner tragus, inferoposterior wall of the ear canal, cymba conchae, and earlobe (sham). Thirty-seven healthy subjects underwent two 6-min tVNS **stimulation** runs per electrode location (monophasic rectangular 500 μ s pulses, 25 Hz). General linear model was performed using SPM; region-of-interest analyses were performed for the brainstem areas.

RESULTS: Stimulation at the ear canal resulted in the weakest activation of the nucleus of solitary tract (NTS), the recipient of most afferent vagal projections, and of the locus coeruleus (LC), a brainstem nucleus that receives direct input from the NTS. **Stimulation** of the inner tragus and cymba conchae activated these two nuclei as compared to sham. However, ROI analysis showed that only **stimulation** of the cymba conchae produced a significantly stronger activation in both the NTS and LC than did the sham **stimulation**.

CONCLUSIONS: These findings suggest that tVNS at the cymba conchae properly activates the vagal pathway and results in its strongest activation, and thus may be the optimal location for tVNS therapies applied to the auricle.

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KEYWORDS: Auricular branch of vagus nerve; **functional magnetic resonance** imaging; locus coeruleus; nucleus of solitary tract; transcutaneous vagus nerve **stimulation**

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