Word repetition priming-induced oscillations in auditory cortex: a magnetoencephalography study

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Magnetoencephalography was used in a passive repetition priming paradigm. Words in two frequency bins (high/low) were presented to the participants auditorily. Participants' brain responses to these stimuli were analyzed using synthetic aperture magnetometry. The main finding of this study is that single-word repetition of low-frequency word pairs significantly attenuated the post-second word event-related desynchronization in the θ - α (5–15 Hz) bands, at 200-600 ms of post-second word stimulus onset. Peak significance between repeated high and low frequency words was evident at approximately 365-465 ms of posttarget onset. This finding has implications for: (i) the role of θ - α event-related desynchronization in lexical representation and access, (ii) the study of repetition suppression in the spectral-temporal domain, and (iii) the connection of neuronal repetition suppression

with behavioral effects of repetition priming. NeuroReport 00:000-000 © 2011 Wolters Kluwer Health | Lippincott Williams & Wilkins.

NeuroReport 2011, 00:000-000

Keywords: α -band, θ -band, auditory, event-related desynchronization, magnetoencephalography, repetition priming, speech, synthetic aperture magnetometry

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Received 26 July 2011 accepted 15 August 2011

Introduction

This study uses magnetoencephalography (MEG) to examine the brain's response to spoken words presented in a passive repetition priming paradigm. The primary goal is to use the neural response suppression associated with repetition – in this case, of spoken words – as a tool to probe the spectral-temporal characteristics of language processing in the brain.

The picture of language emerging from recent spectraltemporal studies of language reveals distinct stages of representation and computation across a number of distinct time-frequency intervals [1-4]. Among these, it appears that the neural representation and access of words is connected with θ - α event-related desynchronization (ERD) and with γ-band event-related synchronization (ERS) [4,5]. In this study, we analyzed the brain's response to repeated and nonrepeated auditorily presented words using synthetic aperture magnetometry (SAM), a beamforming approach to magnetic source imaging that has been successfully applied to spatiotemporal characterization of cortical oscillations without the need for time-locked averaging of the type that is performed in more tightly stimulus-locked responses [6].

One of the goals of this study is to provide a basic spectral-temporal picture of the repetition priming effect. In addition, this study is designed to identify the effects of priming in two different types of words. An important finding in the behavioral literature is that low-frequency words benefit from repetition more than high-frequency words, where the increased benefit is understood as

increased priming (i.e. a greater reduction in reaction time); this is the frequency attenuation effect [7,8]. On the basis of this, the words used in this study were divided into high and low lexical frequency categories.

This study used a passive repetition priming paradigm, and placed the emphasis on the neurobiological side of priming; in particular, on the phenomenon of response suppression. Suppressed responses associated with repetition have been reported in a number of different imaging techniques (for reviews see [9,10]), and it may in fact be affected by whether or not subjects perform active tasks [11]. Concentrating on EEG/MEG, repetition priming has been studied along two dimensions: in the temporal domain, with respect to evoked responses (the N400 in ERP, e.g. [12]; in MEG, [13-16]) and in the spectral-temporal domain, where suppressed activity in induced MEG has been found with repetition of words [17–19].

On the basis of earlier findings, which show θ – α ERD to be sensitive to lexicality [3,4], we hypothesized that repetition suppression with word stimuli would be manifested in this spectral range.

Methods

Eighteen healthy adult human participants (mean ± standard deviation age, 32 ± 9 years, eight male; selfreported right-handed) volunteered for the experimental procedure after giving their written informed consent. This study was approved by the Institutional Review Board of The Children's Hospital of Philadelphia, Pennsylvania, USA.

DOI: 10.1097/WNR.0b013e32834ca576

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system (Siemens, Erlangen, Germany).

Auditory cortex source localization was based on single equivalent current dipole modeling of word onset-evoked (M100) response. Raw MEG data preprocessing, bilateral auditory source modeling, and ERD/ERS analyses were carried out according to the procedures described in [4]. In each participant, dipole locations were used as virtual sensors for the SAM linearly constrained minimum variance beamformer. Differential ERD was computed using SAM applied to the raw MEG trial data band passed between 1.5 and 80 Hz (3.2-s epochs, 0.6-s prestimulus). Extending the epoch ensured the necessary length of baseline used to compute the signal covariance for each stimulus type at the virtual sensors. To visualize the activation induced by stimuli, time-frequency representation (TFR) plots (spectrograms) were computed as described in [4].

The time course of auditory oscillations in the TFR were characterized by sample-wise parametric testing to determine effects of repetition (identical vs. nonidentical) and word frequency (high vs. low) on induced activation. Poststimulus spectrotemporal blocks of interest were identified by the paired t-test (two tailed). On the basis of visual inspection of the distribution of significant spectrotemporal blocks, and on previous results reported in [4], we computed the mean power in the θ - α range. Power modulation was evaluated by repeated measures analysis of variance with factors hemisphere (left, right), repetition (identical, nonidentical), frequency (high, low), applied to power measurements following each stimulus (prime, target) for each participant. All reported significance levels for post-hoc tests were corrected for multiple comparisons using the Bonferroni adjustment.

Results

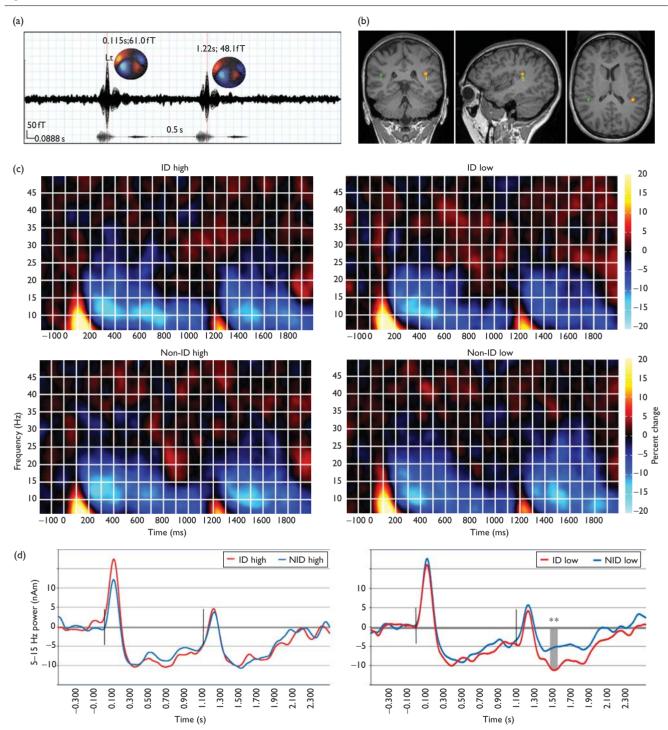
Fig. 1a and b show auditory-evoked neuromagnetic fields for the repetition priming paradigm, and M100 equivalent current dipole source localizations for a representative individual participant. Peak dipole field strengths of

stimuli onset responses showed a remarkable habituation effect between the first and second word presentation. Among the 18 participants, 13 had bilateral dipole fits accounting for 80-90% of variance in the fit interval. In five participants, single equivalent dipole fits were not possible for both hemispheres due to weak and or noisy responses; therefore, they were not considered in the analysis of word position effect on dipole field strengths. Word position-affected peak onset responses in the signal filtered between 1 and 40 Hz $[F_{(1.12)} = 31.718, P < 0.001;$ $\eta p^2 = 0.73$] such that responses to the second word $(14.64 \pm 1.79 \,\text{nAm})$ were significantly attenuated compared with those following the first word (22.49 ± 2.55 nAm). Fig. 1c shows the grand average, collapsed across participants and hemispheres, TFR plots of ERD/ERS in the two repetition conditions (identical/ nonidentical) for each word frequency bin (high-frequency words/low-frequency words). Generally, oscillatory activity was characterized by a dynamic pattern of ERS and ERD. The initial response was an evoked lowfrequency ERS at approximately 100-ms poststimulus onset, lasting approximately up to 200 ms following primes. Although dipole model peaks showed a habituation effect due to rapid stimulus repetition, the mean 1-40-Hz power in the interval of the ERS was not significantly affected by other factors (P > 0.05).

The evoked ERS was followed by prolonged differential ERD in the upper θ and α ranges. The onset of this lowfrequency ERD was approximately 200 ms after stimuli onsets; it lasted throughout the duration of the stimulus, with peak ERD approximately 400-ms poststimuli onsets. The TFRs indicated differential ERD that was modulated by word repetition and frequency (Fig. 1c). Samplewise interrogation of the ERD in the TFRs indicated that power modulation occurred in the θ - α frequency range between 5 and 15 Hz. The time course of this 5–15-Hz power modulation showed a maximal difference between repetition conditions at approximately 415 ms after target onset. As such, separate repeated measures analysis of variance of 5-15-Hz power were carried out in the interval 365-465 post stimuli - prime and targets - onsets. Critically, 5–15 Hz ERD power modulation in the interval 365-465 post prime stimulus (first word) onset was unaffected by factors repetition and frequency, (P > 0.05).

Results from the analysis of target (second word) induced 5–15 Hz ERD in the interval 365–465 post stimulus onset revealed a marked effect of word frequency $[F_{(1,17)}=9.965, P<0.01; \eta p^2=0.37]$. Post-hoc comparisons of cell means indicated a difference between high-frequency $(-11.40\pm1.30\,\text{nAm})$ compared with low-frequency $(-9.41\pm0.97\,\text{nAm})$ words, (P>0.05). Furthermore, 5–15-Hz power following targets was significantly affected by the factor repetition $[F_{(1,17)}=4.911, P<0.05; \eta p^2=0.22]$. Post-hoc comparisons showed repetition suppression, where repetition $(-9.44\pm1.07\,\text{nAm})$ resulted in an





(a) Word average-evoked response data. Cursors indicate maximal dipolar activity elicited by word onsets. (b) Auditory source localization results in a typical individual participant. (c) Time-frequency representation (TFR) plots for increases (yellow/red) and decreases (blue) in auditory cortical source power following high and low lexical frequency words for identical and nonidentical repetition conditions. The stimulus data were averaged over all trials, hemispheres, and participants (n=18) in units of percent change with respect to a 200-ms baseline. (d) Time course of grand-average mean 5-15-Hz activity defined by the spectral-temporal region of interest based on sample-wise t-tests of conditional TFRs. Activity is plotted throughout paired-word trials, and the shaded area (right panel) indicates interval where mean source power significantly differed for low-lexical frequency target stimuli across repetition conditions. **P<0.001 (uncorrected). Lt, left.

attenuation of the auditory ERD compared with nonidentical word pairings (-11.36 \pm 1.29 nAm), (P > 0.05). We show the grand averaged – collapsed across hemispheres and participants – time course of 5–15-Hz power for each frequency bin in the two repetition conditions in Fig. 1d. In this figure, a clear difference is discerned from the time course of low and high-frequency words between the two repetition conditions. Following targets, the ERD in the identical repetition condition for low-frequency words was markedly attenuated relative to low-frequency words when not repeated. Therefore, we carried out sample-wise paired t-tests (uncorrected) on the waveforms for low and highfrequency targets in both the repetition conditions. In Fig. 1d right panel, we highlighted in grey the region where low-frequency targets resulted in significantly (P < 0.001) attenuated ERD in the identical compared with the nonidentical condition. This difference was absent in the high-frequency waveforms. The significant difference in ERD attenuation for low-frequency words during identical repetition began at 355 ms, lasting approximately 70 ms after target stimulus onset.

Discussion

The main result of this study is the finding that θ - α band ERD following the target word in a repetition priming experiment is sensitive to word frequency. Although significant ERD is noted after the first (prime) word, as previously reported in single word studies [4], the magnitude of this ERD after the second (target) word is sensitive to repetition effects and to word frequency in the following way. Target word ERD is not significantly diminished in nonidentical word pairs, nor in paired (repeated) high-frequency words. It is, however, significantly diminished in the repeated word condition for lowfrequency words. This ERD modulation is statistically significant for approximately 70 ms, between 355 and 425 ms post target word onset. This finding appears to be bilateral, appearing in both left and right superior temporal gyrus responses.

A very slight and nonsignificant trend toward a brief period of differential ERD modulation (reduced ERD in repeated, paired, words) appears to be evident in the high-frequency words too, suggesting ERD suppression may be an indicator of repetition priming, and that the magnitude of such ERD suppression may be an index of word frequency.

The finding of θ - α band (5-15 Hz) suppression during repeated word trials is in agreement with activity attenuation results in the functional magnetic resonance imaging literature [10,20]. However, direct comparison of our results to electrophysiological studies is problematic as these studies were carried out in the visual domain and/or multimodal priming across visual and auditory domains focusing on evoked responses, which are time and phase locked to stimulus onset [21-24].

Taken together, the findings of this paper yield two primary results for further investigation. First, the primary findings center on the role of ERD following the presentation of words, and show that this activity is differentially affected by words of different types. This finding provides further evidence for the idea that θ - α ERD plays an important role in lexical representation and access. Second, our findings show θ – α ERD to be affected directly by repetition, further (i) strengthening the case for an important role of ERD in lexical processing and (ii) providing evidence about the nature of spectrotemporal repetition suppression. Moreover, the fact that repetition suppression is driven primarily by the low-frequency words resonates in suggestive ways with the behavior literature, and with the frequency attenuation effect (the finding that facilitation is enhanced with low-frequency words) in particular. In sum, although our findings in this domain do not allow for the formulation of specific hypotheses about the precise role of θ - α ERD, they show this activation to be sensitive to psycholinguistically significant properties of words in a way that provides a foundation for further study of linguistic computations in the brain.

Conclusion

The present findings show that θ – α ERD originating in auditory cortex does indeed show suppression in repeated versus nonrepeated words, and that this suppression is differentially affected by word frequency. Specifically 5–15 Hz of ERD elicited in auditory cortex approximately 200–600 ms after hearing a word is significantly attenuated in the context of priming by an identical word, when that word is of lower lexical frequency.

Acknowledgements

The authors thank Drs Suresh Muthukumaraswamy and Krish Singh for developing and sharing MatLab based beamformer source analysis software. They also thank Dr Maya Ravindranath for help with stimulus generation. This study was supported in part and funded by the National Institutes of Health Grant R01-DC008871 (T.R.) and also by a grant from the Nancy Lurie Marks Family Foundation (T.R.).

Conflicts of interest

Dr Roberts is a consultant for prism clinical imaging. Other authors have no disclosures.

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