Revisiting the Stimulation-Rate-Dependent Pattern Mismatch Negativity

Due Date

2	Marc Pabst
3	Affiliation
4	Course Title
5	Professor Name

7 Abstract

How does the brain process and represent successive sound in close temporal proximity? By investigating mismatch negativity (MMN) components, prior research (Sussman & Gumenyuk, 2005; Sussman, Ritter & Vaughan, 1998) has suggested that temporal proximity plays an 10 important role in how sounds are represented in auditory memory. Here, we investigate how 11 predictability affects the election of mismatch negativity components in auditory sequences 12 consisting of two tones (frequent tone A = 440 Hz, rare tone B = 494 Hz, fixed SOA 100 ms). 13 In the predictable condition, tones are presented in a fixed order whereas in the unpredictable 14 condition, standards and deviants are presented in a pseudo-random order. We expect to find 15 that B tones in the unpredictable condition will elicit a significant MMN while B tones in the 16 predictable conditions will not. A repeating five-tone pattern was presented at several 17 stimulus rates (200, 400, 600, and 00 ms onset-to-onset) to determine at what temporal 18 proximity the five-tone repeating unit would be represented in memory. The mismatch 19 negativity component of event-related brain potentials was used to index how the sounds were 20 organized in memory when participants had no task with the sounds. Only at the 200-ms 21 onset-to-onset pace was the five-tone sequence unitized in memory. At presentation rates of 22 400 ms and above, the regularity (a different frequency tone occurred every fifth tone) was not 23 detected and mismatch negativity was elicited by these tones in the sequence. The results 24 show that temporal proximity plays a role in unitizing successive sounds in auditory memory. 25 These results also suggest that global relationships between successive sounds are represented 26 at the level of auditory cortices.

Revisiting the Stimulation-Rate-Dependent Pattern Mismatch

29 Negativity

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44 Introduction

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- Introducing oddball paradigm - The auditory oddball paradigm is well-established type of
   experimental designs extensivly used in event related potential (ERP) studies. In the most
   basic form of this paradigm, subjects are presented with a series of similar tones or sounds
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   (so-called standards), interrupted by rare tones or sounds that differ in at least one feature
   (deviants) from the more frequent ones. Since it is assumed that the brain constantly makes
   predictions about future sensory impressions and deviating auditory events must violate these
   predictions, these rare sounds play an important role in understanding prediction and
   expectation in the human brain. Different measures have been used to quantify differenced in
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   processing between standard and deviant events,
          - introducing MMN - One of the best-studied approaches to measure these differences
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   in processing is known as the missmatch negativity (MMN) component, obtained by
   subtracting the reponse to deviant events from the response to standard events. Negativity is
   strongest in the fronto-temporal area of the scalp with a peak latency ranging from 100 to 250
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   ms after stimulus onset The eliction MMN is not restricted to the reptition of physically
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   identical stimuli but can also be observed when deviant events are of complex nature,
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   e.g. when abstract auditory regularities are violated (Paavilainen, 2013). The regularities can
   come in the form of relationships between two Saarinen et al. (1992) or multiple tones (Alain
   et al., 1994; Nordby et al., 1988; Schröger et al., 1996) a
          - introducing sussmans study - Sussman et al. (1998) presented participants with a
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   sequency of frequent pure tones and rare pitch deviants. Tones were arranged in a predictable
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   five-tone pattern consisting of four standard tones and one deviant
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   (i.e. A-A-A-B-A-A-A-B, "-" indicating silence between the tones). ERPs to A and B
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   tones were compared for rapid (SOA of 100 ms) and slow (SOA of 1200 ms) stimulation rates.
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   For the 100 ms SOA, they also included a control condition in which tone order was
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   pseudo-random (e.g. A-A-A-B-A-B-A-A) without altering deviant probability (p_B = 20\%).
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   MMNs were only elicted if tone presentation was slow and predictable or fast and random. In
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   a subsquent study, Sussman & Gumenyuk (2005) used the same pattern at different SOAs
   (200 ms, 400 ms, and 800 ms). Simmilarly to their prevous study, grouped presentation at 400
   ms and 800 ms SOA elicted a MMN, while at a stimulution rate of 200 ms such evidence was
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absent. Sussman et al. attributed this observation to sensory memory limitations. Only when auditory memory accommodates enough repetitions of the five-tone pattern, tones could be integrated into a coherent representation allowing for accurate predictions of deviant tones (explaing the absence of MMNs. They further argued that while this must be the case for fast presentation rates with SOAs up to 200 ms, for longer SOAs pattern durations would be too long ans thus ecceed sensory memory capacity. The main weakness in their study is that they ma

- scharf muller - In a recent in-class replication study, (???). found that

82 Methods and Materials

83 Data Acquisition

84 Participants

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100 ms Presentation Rate Twenty-three psychology undergraduate students (2 males, 85 average age 22.6 yrs., SD = 5.57, range 18 - 42 yrs.) were recruited at the Institute of 86 Psychology at the University of Leipzig. All participants reported good general health, normal 87 hearing and had normal or corrected-to-normal vision. Written informed consent was obtained 88 before the experiment. One-third (34.8%) of participants spent time enaging in musical 89 activities at time of survey, while 8.7% had no prior experience in music training. Handedness 90 was assected using a modified version of the Edinburgh Handedness Inventory (Oldfield, 1971, 91 see appendix). A majority (00%) of parcicipants favored the right hand. Participants were 92 blinded in respect to the purpose of the experiment and received course credit in 93 compensation. 94

150 ms Presentation Rate Twenty healthy participants (0 males, average age 00.0 yrs., SD = 0.00, range 00 - 00 yrs.) were recruited. Participants gave informed consent and reported normal hearing and corrected or corrected-to-normal vision. All participants were naive regarding the purpose of the experiment and were compensated in cource credit or money. 00 participants (00%) had received musical training in the last 5 years before the experiment while 00 (00%) reported no musical experiance. In addition, participants reported if streaming occurred during the presentation of the tones.

oz Stimuli and Stimulis Delivery

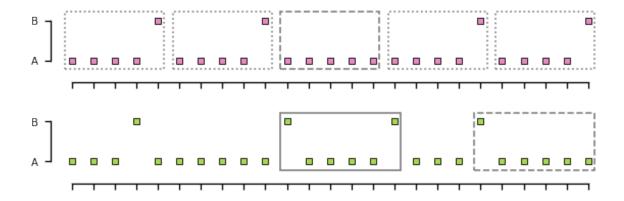


Figure 1. Tones of two different frequencies (A=440 Hz, B=449 Hz) were presented in two blocked conditions: In the "predictable" condition (top half), tones followed a simple pattern in which a single B-tone followed four A-tones. Some designated B-tones were replaced by A-tones ("pattern deviants"). In the "random" condition (lower half), tones were presented in a pseudo-random fashion ()

Participants where seated in a comfortable chair in a sound-insulated cabin. The 103 experimental setup was practically the same as the one used ny Sussman, but instead of 104 reading a book, subjects were asked to focus their attention on a previously selected movie. 105 Movies were presented with subtitles but without sound. Commercially available software 106 (MATLAB R2014a; The MathWorks Inc, Natick, MA) in conjunction with the Psychophysics 107 Toolbox extension (version 3.0.12, Brainard, 1997; Kleiner et al., 2007) was used to control 108 stimulus presentation. Stimuli consisted of pure sinusoidal tones with a duration of 50 ms 109 (including a 10 ms cosine on/off ramp), presented isochronously at a stimulation onsets 110 asynchrony (SOA) of 100 ms for study 1 and 150 ms for study 2. Overall, a total of 40 blocks 111 containing a mixture of frequent 440 Hz tones ("A" tones) and infrequent 449 Hz tones ("B" 112 tones) were delivered binaurally using Sennheiser XY headphones. In one half of the blocks, 113 tones were presented in pseudo-random order (e.g. A-A-A-B-A-B-A), "random" condition), 114 while in the remaining block tone presentation followed a simple pattern in which a 115 five-tone-sequence of four frequent tones and one infrequent tone (i.e. A-A-A-B) was 116 repeated cyclically ("predictable" condition). The ratio of frequent and infrequent tones was 117 10% for both conditions. Within the predictable condition, 10% of designated (infrequent) B 118 tones were replaced by A tones, resulting in sporadic five-tone sequences consisting solely of A 119

tones (i.e. A-A-A-A), thus violating the predictability rule. To assure comparability of local histories between tones in both conditions, randomly arranged tones were interspersed with sequences mimicking aforementioned patterns from the predictable condition (B-A-A-A-B and B-A-A-A-A) in the random condition. A grand total of 2000 tones in study 1 and 4000 tones in study 2 were delivered to each participant.

125 Data Acquisition

Electrophysiological data was recorded from active silver-silver-chloride (Ag-AqCl) electrodes 126 using an ActiveTwo amplifier system (BioSemi B.V., Amsterdam, The Netherlands). Acquisition was monitored online to ensure optimal data quality. A total of 39 channels were 128 obtained using a 32-electrode-cap and 7 external electrodes. Scalp electrode locations 129 conformed to the international 10-20 system. Horizontal and vertical eye movement was 130 obtained using two bipolar configurations with electrodes placed around the lateral canthi of 131 the eyes and above and below the right eye. Additionally, electrodes were placed on the tip of 132 the nose and at the left and right mastoid sites. Data was sampled at 512 Hz and on-line 133 filtered at 1000 Hz. 134

135 Analysis Pipeline

Data prepossessing was implemented using a custom pipeline based on the MNE Python software package (Gramfort, 2013) using Python 3.7. All computations were carried out on a cluster operated by the University Computation Center of the University of Leipzig. Code used in thesis is publicly available at https://github.com/marcpabst/xmas-oddballmatch.

First, EEG data was subjected to the ZapLine procedure (de Cheveigné, 2020) to
remove line noise contamination. A fivefold detection procedure as described by
Bigdely-Shamlo et al. (2015) was then used to detect and subsequently interpolate bad
channels. This specifically included the detection of channels thain contain prolonged
segments with verry small values (i.e. flat channels), the exclusion of channels based on robust
standard deviation (deviation criterion), unusually pronounced high-frequency noise (noisiness
criterion), and the removal of channels that were poorly predicted by nearby channels
(correlation criterion and predictability criterion). Channels considered bad by one or more of
these methods were removed and interpolated using spherical splines (Perrin et al., 1989).

Electrode locations for interpolations were informed by the BESA Spherical Head Model.

For Independent Component Analysis (ICA), data 1-Hz-high-pass filtered (134th order 150 hamming-windowed FIR) was applied prior to ICA (Winkler et al., 2015). To further reduce 151 artifacts, Artifact Subspace Reconstruction (ASR, Mullen et al., 2015) was used to identify parts of the data with unusual characteristics (bursts) which were subsequently removed. ICA 153 was then carried out using the *Picard* algorithm (Ablin et al., 2018, 2017) on PCA-whitened 154 data. To avoid rank-deficiency when extracting components from data with one or more 155 interpolated channels, PCA was also used for dimensionality reduction. The EEGLAB 156 (version 2020.0, Delorme & Makeig, 2004) software package and the IClabel plugin (version 157 1.2.6, Pion-Tonachini et al., 2019) were used to automatically classify estimated components. 158 Only components clearly classified (i.e. confidence above 50%) as resulting from either eye 159 movement, muscular, or heartbeat activity were zeroed-out before applying the mixing matrix to unfiltered data.

In line with recommendations from Widmann et al. (2015) and de Cheveigné & Nelken 162 (2019), a ORDER finite impulse response (FIR) bandpass filter from 0.1 Hz to 40 Hz (Hamming window, 0.1 Hz lower bandwith, 4 Hz upper bandwidth, 0.0194 passband ripple, 164 and 53 dB stopband attenuation). Continuous data was epoched into 400 ms long segments 165 around stimulus onsets. Epochs included a 100 ms pre-stimulus interval. No baseline 166 correction was applied. Segments exceding a peak-to-peak voltage difference of 100 μ V were 167 removed. No data set meet the pre-registrated exclusion criterion stated of less than 100 trials 168 per condition, thus data from all participants (20 for 100 ms presentation rate and 23 for 150 169 ms presentation rate) was analysed. 170

171 Statistical Analysis

72 **MMN**

The dependent variable for analysing missmatch response was calculated by averaging
amplitudes in a time window extedning ±25 ms around the maximum negativity obtained by
subtracting the mean ERP timecourse following the (expected) deviant event from the ERP
following the (expected) standard event. To obtain mean amplitudes, ERPs to 4th position A
tones (A-A-A-A-X, boldface indicates the tone of interest) and B tones (A-A-A-B) were

averaged seperatly for both the *random* and the *predictable condition*. For the *random* condition, only tones that were part of a sequence matching the patterns in the *predictable* condition were included.

In accordance with the original analysis by Sussman & Gumenyuk (2005), mean amplitudes for frontocentral electrodes (FZ, F3, F4, FC1, and FC2) and the two mastoid positions (M1 and M2) were averaged separately. Then, for both SOAs, independent three-way repeated measures analyses of variance with factors condition (factors predcitable and random), stimulus type (factors A tone and B tone), electrode locations (levels fronto-central and mastoids), and all possible interactions were calculated.

187 Results

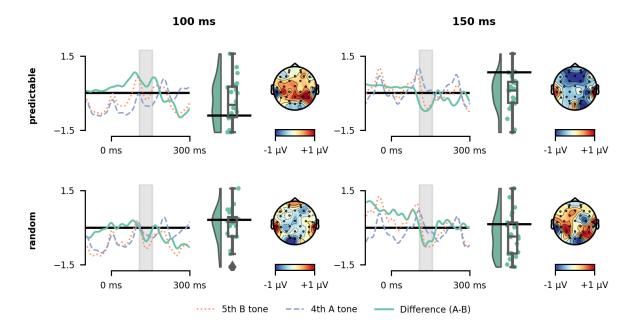


Figure 2. ERP grand averages (pooled FZ, F3, F4, FC1, and FC2 electrode locations) for an SOA of 100 ms (left) and 150 ms (right), for A tones (A-A-A-A-X, blue dashed lines) and B tones (A-A-A-B, orange dashed line) and their difference (B - A, green solid line). Upper panels show ERPs for tones presented in a predcitable pattern (predcitable condition) while lower panels show ERPs for tones presented in pseudo-random order (random condition). Shaded area marks MMN latency window (110 ms to 160 ms) used to calculate the distribution of amplitude differences across participants (middle of each panel) and the difference of topographic maps averaged over the same interval (right of each panel).

Grand averages of event-related potentials (ERP) at pooled FZ, F3, F4, FC1, and FC2 188 electrode locations to A tones (A-A-A-A-X), B tones (A-A-A-B), and their difference (B 189 tone minus A tone) are displayed in Figure X for both 100 ms (left panel) and 150 ms (right 190 panel) stimulus onset asynchronies. Top half of each panel shows ERPs in the predictable 191 condition while lower half depicts ERPs in the random condition. For both presentation rates, 192 clear rythms matching the presentation frequency of 10 Hz (100 ms) and respectively 6.667 Hz 193 (150 ms) are seen as a result from substantial overlap of neighboring tones. Panels also show 194 the distribution of mean amplitude differences in the MMN latency window (as defined above, 195 110 ms to 160 ms after stimulus onset) across participants and the difference of sclap 196 topogrphies averaged over the same interval. Simmilarly, waveforms and mean amplitude 197

difference distributions at pooled mastoid sites are shown in Figure X.

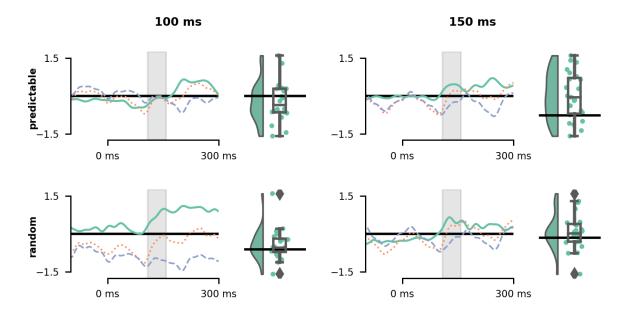


Figure 3. ERP grand averages (pooled M1, M2 electrode locations) for an SOA of 100 ms (left) and 150 ms (right), for A tones (A-A-A-A-X, blue dashed lines) and B tones (A-A-A-B, orange dashed line) and their difference (B - A, green solid line). Upper panels show ERPs for tones presented in a predcitable pattern (predcitable condition) while lower panels show ERPs for tones presented in pseudo-random order (random condition). Shaded area marks MMN latency window (110 ms to 160 ms) used to calculate the distribution of amplitude differences across participants.

Evoked responses to A and B tones were compared by calculating mean amplitudes in 199 the MMN latency window. Mean amplitudes in the MMN latency window and their standard 200 deviantions (SD) for all conditions are shown in Table X. Descriptively, mean amplitudes at 201 pooled fronto-central electrode locations were more negative for randomly presented B tones 202 than for randomly presented A tones, regardless of tone presentation rate (100 ms: 203 $\Delta M = -0.358 \,\mu V$; 150 ms: $\Delta M = -0.555 \,\mu V$) This also held true for tones presented in a 204 predictable fashion, but for the slower of the two presentation rates only ($\Delta M = -0.582 \,\mu V$)). 205 In contrast, when predcitable tone patterns occured at a faster 100 ms rate, B tones elicted 206 descriptively more positive responses than A tones ($\Delta M = 0.383 \,\mu V$). Descriptive comparison 207 of evoked responses from pooled left and right mostoids revealed that pseudo-randomly 208 presented B tones were more positive in the MMN latency window than A tones (100-ms-SOA: 209 $\Delta M = 0.746 \,\mu\text{V}$, 150-ms-SOA: $\Delta M = 0.510 \,\mu\text{V}$). A simmilar observation could be made for 210

precitable B tones compared to the preceding A tones at a SOA of 150 ms ($\Delta M = 0.399 \,\mu V$)) but not for the faster presentation rate ($\Delta M = -0.132 \,\mu V$).

SOA	Condition	${\bf Stimulus Type}$	Mean	\mathbf{SD}	Mean	SD
100	predictable	A	-0.431	1.23	-0.052	1.51
		В	-0.0477	1.22	-0.184	1.56
	random	A	-0.225	1.82	-1.04	2.64
		В	-0.583	2.16	-0.296	3.23
150	predictable	A	0.25	0.967	-0.349	1.19
		В	-0.331	1.09	0.0492	1.33
	random	A	0.0233	1.75	-0.292	1.64
		В	-0.531	1.82	0.218	2.38

Statistical analyses provided support for these findings. For the 100 ms stimulation rate, the three-way ANOVA yielded a significant three-way interaction effect (condition x stimulus type x electrode locations; F(1,19) = 7.53, p = 0.0130) but failed to reveal main effects for factors stimulus type (F(1,19) = 1.05, p = 0.3180), condition (F(1,19) = 0.83, p = 0.3730), and electrode locations (F(1,19) = 0.04, p = 0.8520). In contrast, for tones presented at a SOA of 150 ms only the two-way interaction term stimulus type x electrode locations had a significant effect (F(1,22) = 20.76, p = 0.0002). Mean amplitudes in the MMN latency window however did not differ for factors stimulus type (F(1,22) = 0.32, p = 0.5790), electrode locations ().

Two-way ANOVAs (condition x stimulus type) were carried out seperatly for pooled fronto-central and mostoid electrode locations. For 100 ms tone presentation rate, the condition x stimulus type interaction only revealed a significant effect for the fronto-central electrode cluster (F(1,19)=16.75, p=0.0006) but not for pooled mastoid sites (F(1,19)=2.37, p=0.1410) indicating that the three-way interaction effect condition x stimulus type x electrode is indeed driven by the amplitude differences in te fronto-central electrode locations. Contrary to this, for the 150 ms presentation rate, main effects for stimulus type were significant for both fronto-central and mastoid sites, suggesting that there was both a MMN at fronto-central locations as well as a polarity-reversal at the mastoid electrodes.

Table 1

Results of the 3-way ANOVA (condition x stimulus x electrode) for repeated measures conducted on the mean ERP-amplitudes (time window 111 - 161 ms) at electrode Fz (upper section). The significant interaction between the three factors included was further analyzed by 2-way ANOVAS (stimulus x electrode) conducted separately for the random condition (middle section) and the predictable condition (lower section).

	Effect	DFn	DFd	\mathbf{F}	p	p<.05	ges
	Condition	1	19	0.831	0.373		0.008
	StimulusType	1	19	1.05	0.318		0.002
α	Electrode	1	19	0.036	0.852		0.000331
100 ms	Condition x StimulusType	1	19	0.051	0.823		7.55e-05
\vdash	Condition x Electrode	1	19	0.763	0.393		0.002
	StimulusType x Electrode	1	19	0.797	0.383		0.001
	Condition x Stimulus Type x Electrode	1	19	7.53	0.013	*	0.01
	Condition	1	22	0.08	0.78		0.000263
	StimulusType	1	22	0.317	0.579		0.000339
\mathbf{z}	Electrode	1	22	0.035	0.854		0.000301
150 ms	Condition x StimulusType	1	22	0.16	0.693		0.000124
	Condition x Electrode	1	22	1.13	0.299		0.003
	StimulusType x Electrode	1	22	20.8	0.000155	*	0.026
	Condition x Stimulus Type x Electrode	1	22	0.053	0.819		4.63e-05

Figure X shows EEG waveform averages for five-tone sequences (A-A-A-B)

presented in a *predictable* (top panel) and *random* contexts (lower panel).

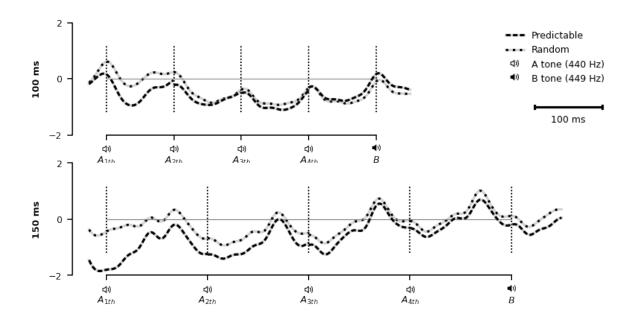


Figure 4. EEG waveforms for five-tone sequences presented in an predictable context (dotted line) and pseudo-random condition (dashed line) for 100 ms presentation rate (top panel) and 150 ms presentation rate (lower pabel). Vertical lines indicate tone onset.

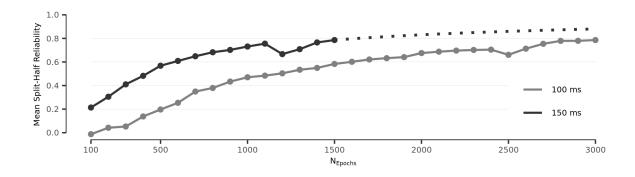


Figure 5. EEG waveforms for five-tone sequences presented in an predictable context (dotted line) and pseudo-random condition (dashed line) for 100 ms presentation rate (top panel) and 150 ms presentation rate (lower pabel). Vertical lines indicate tone onset.

Table 2

Results of the 3-way ANOVA (condition x stimulus x electrode) for repeated measures conducted on the mean ERP-amplitudes (time window 111 - 161 ms) at electrode Fz (upper section). The significant interaction between the three factors included was further analyzed by 2-way ANOVAS (stimulus x electrode) conducted separately for the random condition (middle section) and the predictable condition (lower section).

		Effect	DFn	DFd	\mathbf{F}	p	p<.05	ges
	Frontal	Condition	1	19	0.16	.694		0.003
		StimulusType	1	19	0.006	.938		1.5e-05
100 ms		Condition x StimulusType	1	19	16.7	<.001	*	0.013
100	Mastoids	Condition	1	19	1.28	.272		0.014
		StimulusType	1	19	1.21	.285		0.004
		Condition x StimulusType	1	19	2.37	.141		0.009
	Frontal	Condition	1	22	0.947	.341		0.006
		StimulusType	1	22	22.7	<.001	*	0.038
150 ms		Condition x StimulusType	1	22	0.028	.868		2.2e-05
	Mastoids	Condition	1	22	0.206	.655		0.001
		StimulusType	1	22	6.56	.018	*	0.018
		Condition x StimulusType	1	22	0.122	.730		0.00028

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