

Published in final edited form as:

J Neurosci. 2011 March 2; 31(9): 3400–3406. doi:10.1523/JNEUROSCI.3296-10.2011.

Common or redundant neural circuits for duration processing across audition and touch

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Abstract

Certain features of objects or events can be represented by more than a single sensory system such as roughness of a surface (sight, sound and touch), the location of a speaker (audition and sight) or the rhythm or duration of an event (by all three major sensory systems). Thus, these properties can be said to be sensory-independent or amodal. A key question is whether common multisensory cortical regions process these amodal features, or does each sensory-system contain its own specialized region(s) for processing common features? We tackled this issue by investigating simple duration detection mechanisms across audition and touch, specifically because fine duration discriminations are possible in both systems. The mismatch negativity (MMN) component of the human event-related potential provides a sensitive metric of duration-processing and it has been elicited independently during both auditory and somatosensory investigations. Employing high-density electroencephalographic recordings in conjunction with intracranial subdural recordings, we asked whether fine duration discriminations, represented by the MMN, were generated in the same cortical regions regardless of the sensory modality being probed. Scalp-recordings pointed to statistically distinct MMN topographies across senses, implying differential underlying cortical generator configurations. Intracranial recordings confirmed these non-invasive findings, showing generators of the auditory MMN along the superior temporal gyrus with no evidence of a somatosensory MMN in this region, whereas a robust somatosensory MMN was recorded from post-central gyrus in the absence of an auditory MMN. The current data clearly argue against a common circuitry account for amodal duration processing.

Keywords

Somatosensory; Auditory; duration processing; mismatch negativity; multisensory; cross-modal

Introduction

The separate sensory epithelia are specialized for very different energy sources, such that light can have no impact on the hair-cells along the basilar membrane, and it is nonsensical, outside of poetry, to talk of tickling the photoreceptors of the retina. There are, however, also properties of objects that can be explored and represented by more than a single sensory system. We can readily perceive the frequency of a vibration through both hearing and touch, the speed of a moving bus through vision and audition, and the duration of an event through all three of these primary senses. In this way, these latter properties can be said to be amodal or sensory-independent qualia of a given object or event. This notion of amodal properties gives rise to an obvious question when one considers the neural processes responsible for their analysis. If these features are sensory-independent, are they then analyzed by a network of multisensory regions such that all sensory systems feed into and rely upon multisensory amodal systems for the analysis of common properties. This is an attractive thesis in that it furnishes what is potentially an economical solution. The alternative, whereby each sensory system has its own specific system for analyzing such properties, seems to require unnecessary duplication of function. However, it is also possible that duration is an elemental feature that is best extracted at the level of sensory registration. Of course, a complication with the simple dichotomy outlined is that not all senses are created equal when it comes to analyzing a given amodal feature. Audition is extremely sensitive to temporal properties with a great deal higher resolution than say vision. Vision is sensitive to the spatial location of objects whereas audition is a poor substitute. Touch is more appropriate for texture discrimination than vision. Thus, if a given sensory system has considerably superior tuning for a given property, one could imagine a scenario whereby more local cortical real estate would be sequestered for a largely unisensory analysis of that property. In this scenario, the sensory system with lower sensitivity might be expected to rely on its own unisensory circuitry, perhaps with ancillary input from the regions responsible for analysis within the superior sensory system.

The current study was designed to assess this notion of common circuitry, with an eye to adjudicating between the possibilities outlined above. We chose to begin our investigations of this issue with a study of duration processing across the auditory and somatosensory systems for two main reasons. First, our previous work in multisensory integration has shown intimate links between these two sensory systems, which share common neural circuitry during very early sensory processing (e.g. (Foxy *et al.*, 2000, 2002, 2005; Schroeder & Foxy, 2002, 2005; Kayser *et al.*, 2005; Murray *et al.*, 2005)). Second, duration is an amodal property for which both sensory systems have a good degree of sensitivity and thus, neither system is so much more sensitive than the other as to render the comparison blatantly imbalanced. Our dependent measure was the mismatch negativity (MMN), an event related potential (ERP) component of the auditory and somatosensory systems (Näätänen, 1992; Kekoni *et al.*, 1997), that has been shown to index automatic change detection for duration (Näätänen, 1992; Akatsuka *et al.*, 2005). Here using both high-density electrical mapping of scalp-recorded potentials and intracranial electro-corticographic (eCOG) recordings, we ask if the MMN to auditory and somatosensory duration change is generated in the same cortical regions or if each sensory system contains its own specialized duration processing mechanism.

Methods

Participants

8 participants (5 male) ranging in age from 22-33 years (mean 26.3) with normal hearing completed the experiment for a modest fee of \$12 per hour. All participants were right handed and reported normal hearing and no known neurological deficits. All participants

gave written informed consent, and all procedures were approved by the ethical review board of The City College of New York. Ethical guidelines were in accordance with the Declaration of Helsinki.

Apparatus

Tones were presented to the right ear via headphones (Sennheiser HD600). Somatosensory stimulation was presented via a Low Cost Piezo Systems Linear amplifier (Piezo Systems Inc.) to the index finger of the right hand. All experiments were carried out in a darkened, acoustically and electrically shielded room.

Mismatch task

The MMN component is a robust metric for studying pre-attentive processing of stimulus properties for both the auditory and somatosensory systems. While recording the electroencephalogram (EEG) from participants, the mismatch evoked potential is elicited automatically by deviations in an otherwise repetitive stream of stimuli (Näätänen, 1992; Naatanen *et al.*, 2007). The auditory mismatch (aMMN) has been extensively studied for several auditory stimulus properties; such as duration (Näätänen, 1992; De Sanctis *et al.*, 2009), frequency (Sams *et al.*, 1985), and complex stimuli (Saint-Amour *et al.*, 2007). The amplitude of the aMMN component reflects participant's ability to discriminate between standard and deviant stimuli (Amenedo & Escera, 2000; Sanctis *et al.*, 2009). The somatosensory mismatch (sMMN), in comparison, has not been nearly as well characterized. It has, however, been similarly elicited for many different stimulus properties such as a duration (Akatsuka *et al.*, 2005; Spackman *et al.*, 2007), frequency (Kekoni *et al.*, 1997), and location (Shinozaki *et al.*, 1998). As with the aMMN, the sMMN component correlates with the ability to discriminate between standard and deviant stimuli (Spackman *et al.*, 2007). The MMN is elicited by the infrequent occurrence of a “deviant” stimulus within the context of a frequently occurring “standard” stimulus. It is usually measured by subtracting the response to the deviant from the response to the standard. For the auditory MMN this results in a relative negativity about 100 ms following the onset of deviance (which is delayed with respect to stimulus onset for a duration deviant) that is focused over fronto-central scalp areas (e.g., Naatanen, 2002). For the somatosensory MMN the timing and scalp distribution has been shown to be quite similar, but the initial negative deflection is also followed by a relative positivity (see e.g. Spackman *et al.*, 2007). Here participants viewed a silent movie with subtitles during stimulus delivery and were told to ignore the auditory and somatosensory stimuli. In the aMMN condition, the standard stimuli ($p=0.75$) were 200Hz tones of 50ms duration and the deviant stimuli ($p=0.25$) were 200Hz tones of 100ms duration. In the sMMN condition, the standard stimuli were vibro-tactile stimuli of 200Hz of 50ms duration and the deviant stimuli were vibro-tactile stimuli of 200Hz of 100ms duration. All stimuli were sinusoids convolved with a trapezoid such that there was a 5ms rise at the onset and a 5ms ramp at the offset. A separate behavioral testing session in 7 additional subjects who did not serve in the main MMN experiment, indicated that there were no differences in the ability to detect the auditory and the somatosensory duration deviants ($t(6)=-0.667$, $p=0.52$), with mean detection rates of 93.5% in both conditions.

There were two blocked conditions: aMMN and sMMN. There were a total of 1600 stimuli with 400 deviants per block. The order of presentation was counterbalanced across participants. The standard and deviant were delivered in a pseudo random order to ensure that deviants were never presented sequentially. The inter stimulus interval was fixed at 1000ms.

EEG Recording and Analysis

Brain activity was recorded using a Biosemi System 168 channel EEG system (BioSemi, Amsterdam, Netherlands). The data were recorded at 512Hz and band-pass filtered from 2 to 45Hz (24 dB/octave). The data were analyzed offline using the Matlab programming language (Mathworks, Natick, MA). Epochs of 600ms with 100ms pre-stimulus baseline were extracted from the continuous data. We applied an automatic artifact rejection criterion of $\pm 75\mu\text{V}$ across all electrodes in the array. Trials with more than six artifact channels were rejected. In trials with less than six such channels, we interpolated any remaining bad channels using a nearest neighbor spline (Perrin *et al.*, 1987; Perrin *et al.*, 1989). The data were re-referenced to the average of all channels (Michel *et al.*, 2004; Nunez & Srinivasan, 2006). The average accepted trials per condition were ~300 deviants and ~900 standards. The mismatch waveforms were classically obtained by subtracting the standard evoked potential from the deviant evoked potential.

Mismatch-responses

To test for the presence of the MMN, mean amplitude measurements were obtained in a 40ms window centered at the group-mean peak latency for the largest negative minimum component between 100ms-200ms for both the aMMN and the sMMN, as well as the following largest negative minimum component of the aMMN and largest positive maximum component of the sMMN from 200ms-300ms. From the literature, the electrode cluster surrounding the midline frontal site, what we refer to as Fz, was identified as a region of interest for both the somatosensory and auditory MMN. Since we recorded with a high-density montage, what we refer to as the Fz site hereafter indicates the average of the five nearest electrodes sites, which gives a better signal-to-noise ratio and is a more representative waveform of the fronto-central region than that given by a single electrode site.

Topographical Analysis

In line with our primary question, the key analysis strategy involved a determination of whether common circuitry underlies both the aMMN and sMMN. For this, we used the Topographical ANOVA (TANOVA), as implemented in the Cartool software, to statistically test for possible topographical differences between the sMMN and aMMN using global dissimilarity and non-parametric randomized testing (Lehmann & Skrandies, 1980). Global dissimilarity is an index of configuration differences between two scalp distributions, independent of their strength as the data were normalized using the global field power. For each subject and time point, the global dissimilarity indexes a single value, which varies between 0 and 2 (0, homogeneity; 2, inversion of topography). To create an empiric probability distribution against which the global dissimilarity can be tested for statistical significance, the Monte Carlo MANOVA was applied (for a more detailed description see (Manly, 1991)). To control for Type I errors a period of statistical significance was only plotted and considered as significant if an alpha criterion of 0.05 or less was obtained for at least 11 consecutive sample points (~21ms) (e.g. Foxe & Simpson, 2002; Guthrie & Buchwald, 1991).

Dipole Source Analysis

To estimate the location of the intra-cranial generators of the scalp recorded grand mean MMNs, we also performed source modeling using brain electric source analysis (BESA 5.1.8, MEGIS Software GmbH, Gräfelfing, Germany; Scherg and Von Cramon, 1985). Results of this procedure are reported in the supplementary materials only.

Intracranial data

An additional dataset was collected from a patient (female, age 48) who was undergoing invasive intracranial mapping for intractable epilepsy following an earlier right anterior temporal lobe resection. Intracranial EEG recordings were obtained using a multi-array grid of 48 linear contacts (6 rows \times 8 columns, 10mm inter-contact spacing), which was placed over the auditory cortex and surrounding regions of the right hemisphere and from a single linear strip of 6 contacts (1 row \times 6 columns, 10mm inter-contact spacing), which was placed over somatosensory cortex of the right hemisphere.

The precise anatomical location of each electrode contact was determined by co-registration of the post-op CT scan on pre and post-op anatomical Magnetic resonance imaging (MRI) and then normalized into the Montreal Neurological Institute (MNI) space (using SPM8 developed by Wellcome Department of Imaging Neuroscience, UCL). The pre-op MRI was used for its accurate anatomical information, the post-op CT scan provided undistorted placement of the electrode contacts, and the post-op MRI allowed assessment of the quality of the entire co-registration process since it includes both electrodes and anatomical information. In Supplementary Figure 3, for the purpose of illustrating activity and relative electrode locations the grid is depicted on a template brain.

Subdural electrodes are highly sensitive to local field potentials generated within $\sim 4.0 \text{ mm}^2$ area and much less sensitive to distant activity (Allison *et al.*, 1999; Lachaux *et al.*, 2005; Molholm *et al.*, 2006; Fiebelkorn *et al.*, 2010), which allows for improved localization of underlying current sources relative to scalp-recorded EEG. As the electrodes were located over the right-side of the brain, the tones and somatosensory stimulation were presented to the left middle finger and left ear. Brain activity was recorded using a Brain Amp system (Brain Products GmbH, Munich, Germany). Recordings were digitized online at 1000Hz and bandpass-filtered offline from 2 to 45 Hz (24db/octave). Epochs of 600ms with 100ms pre-stimulus baseline were extracted from the continuous data. The data were analyzed offline using a $\pm 350\mu\text{V}$ artifact rejection criterion. Statistical analyses of intracranial data were identical to those for scalp data except that they were based on the variance measured across single trials of the current source density rather than the variance of the evoked potential measured across participants.

Current Source Density

Current source density (CSD) profiles were calculated using either a five point formula, Eqn (1), if the electrode was on a grid, or a three point formula, Eqn (2), if the electrode was on a strip to estimate the second spatial derivative of the voltage axis. CSD profiles provide an index of the location, direction, and density of transmembrane current flow, the first-order neuronal response to synaptic input (Nicholson & Freeman, 1975)

$$\text{CSD}_{i,j} = 4 \times V_{i,j} - V_{i+1,j} - V_{i-1,j} - V_{i,j+1} - V_{i,j-1} \quad (1)$$

$$\text{CSD}_{i,j} = 2 \times V_{i,j} - V_{i+1,j} - V_{i-1,j} \quad (2)$$

Where $V_{i,j}$ denotes the recorded field potential at the i^{th} row and j^{th} column in the electrode grid. CSD profiles for electrode contacts located on the border of the grid or the ends of the strip were not calculated because of the lack of nearest neighbors, and no such electrodes were used in our analysis.

Results

Scalp Recordings

Inspection of the grand mean ERP-difference waveforms, deviant minus standard showed clear mismatch responses (black line) for both the auditory and somatosensory conditions over the midline fronto-central scalp (Fz) (Figure 1). For both sensory systems there were two separable phases of the mismatch response. The auditory mismatch had a pair of successive negative peaks centered at approximately ~155ms and ~235ms respectively. The somatosensory mismatch had an early negative component at ~147ms, followed by a positive component that peaked at ~235ms.

To statistically verify the presence of the components at the two phases, each participant's standard and deviant waveforms were averaged over ~40ms centered at each group-mean component peak and submitted to a t-test. The auditory standard and deviant were found to be statistically different for both components (121-161ms: $t_{df=7}=-3.161$ $p<0.05$; 217-256ms: $t_{df=7}=-4.0515$ $p<0.005$), and the somatosensory standard and deviant were statistically different at both components (128-168ms: $t_{df=7}=-2.522$, $p<0.05$; 214-254ms: $t_{df=7}=8.945$, $p<0.005$).

The scalp topographic maps representing the potential distributions of the grand mean somatosensory and auditory mismatches for both phases are shown in Figure 2. The first auditory mismatch component has a frontal left-lateralized negative distribution, while the somatosensory has a fronto-central negative distribution. The second components for the somatosensory and auditory mismatch have very different topographic distributions. The auditory topographic map has a slightly left lateralized fronto-central negative distribution while the somatosensory “mismatch” topographic distribution has a frontal positive distribution.

To verify the differences in topographies of the somatosensory and auditory mismatches a TANOVA analysis was conducted (Figure 2). There were three clear periods of statistical difference between the topographies of the auditory and somatosensory mismatch responses. The first two periods aligned with the latencies of the two components revealed during the analysis over the frontal scalp (Fz) for both sensory modalities, at ~105-135ms followed by ~200-250ms. The third period was ~275-325ms, which coincided with a later third positive peak of the aMMN, centered at ~300ms. Another possibility is that an amodal processor might not necessarily be engaged at precisely the same time by the auditory and somatosensory stimuli. We therefore performed an additional test between the topographies of the late aMMN and the early sMMN. This too revealed statistically significant differences between the auditory and somatosensory MMN topographies (not depicted).

Intracranial Recordings

All contacts were examined for the presence of an MMN (see supplementary Figure S3 for more extensive representation of the responses, across 22 of the contact sites). For the sMMN, the response was clearly focused along a small part of the strip over the post-central gyrus and was not seen in any of the grid contacts (see supplementary material). PcG₁ (see Figure 3, and Figure S3) had the largest negative and positive sMMNs and was used in our analyses; these responses appeared to invert polarity in channel PcG₂ (Figure 3, and Figure S3), which was also used in our analyses. An aMMN was not seen in any of the strip contacts. For the aMMN, the response was somewhat more extensive due to the greater coverage over relevant sites, and was seen in contacts over the STG. The early negativity was of greatest amplitude in STG₁, and the later negativity in STG₂ (see Figure 3, and Figure S3), the two contacts used from the grid for our analyses. Figure 3 shows current source density waveforms from the 4 selected subdural electrodes for both the

somatosensory and auditory mismatch. Two adjacent electrodes (STG₁ and STG₂) were positioned along the Sylvian Fissure over inferior frontal gyrus (IFG) and the superior temporal gyrus (STG). Two adjacent electrodes (PcG₁ and PcG₂) were positioned over the Postcentral Gyrus.

Inspection of the mean auditory CSD difference-waveforms averaged over trials at electrodes STG₁ and STG₂ showed a clear aMMN response. The mean somatosensory CSD difference-waveforms at the same electrodes did not show evidence of an sMMN response. The reverse was the case for PcG₁ and PcG₂ which showed clear sMMN responses but did not show aMMN responses. In the auditory condition the mismatch waveforms at STG₁ show a clear polarity inversion at STG₂ indicating a local generator. The aMMN waveform at STG₁ and STG₂ showed polarity inverted components centered at approximately 176ms followed by a mirrored components at approximately 295ms. For the somatosensory stimulation there were no discernible MMN components and no obvious inversions at STG₁ and STG₂. The sMMN response at PcG₁, on the other hand, show a clear polarity inversion at PcG₂ indicating a local generator of the sMMN. The sMMN at PcG₁ and PcG₂ had inverted components centered at approximately 110ms and 226ms. The aMMN at PcG₁ and PcG₂ had no discernible peaks and exhibited no obvious polarity inversions.

To statistically verify the presence of the MMN responses, individual trials were averaged over a 40ms interval that was centered on the peak amplitude of the average MMN waveform. The standard and deviant trials were sorted and submitted to an unpaired t-test. The auditory standard and deviant were found to be statistically different for both components at STG₁ (156-196ms: $t_{df=827}=-8.753$, $p<0.0001$; 276-316ms: $t_{df=827}=5.96$, $p<0.0001$) and STG₂ (156-196ms: $t_{df=827}=6.25$, $p<0.0001$; 242-282ms: $t_{df=827}=6.70$, $p<0.0001$), and the somatosensory standard and deviant were not statistically different at STG₁ (156-196ms: $p=0.18$; 276-316ms: $p=0.256$) and STG₂ (156-196ms: $p=0.7839$; 242-282ms: $p=0.9$). The somatosensory standard and deviant were found to be statistically different for both components at PcG₁ (101-141ms: $t_{df=864}=-6.67$, $p<0.0001$; 207-247ms: $t_{df=864}=6.18$, $p<0.0001$) and PcG₂ (95-135ms: $t_{df=864}=-4.27$, $p<0.0001$; 206-246ms: $t_{df=864}=-3.03$, $p<0.0005$), and the auditory standard and deviant were not statistically different at both components at PcG₁ (101-141ms: $p=0.87$; 207-247ms: $p=0.588$), and only the first component was statistically different at PcG₂ (95-135ms: $t_{df=827}=2.59$, $p<0.005$; 206-246ms: $p=0.22$). While this latter finding appears to suggest that there is cross-sensory MMN activity in PcG₂, observation of the waveforms themselves suggests otherwise, since there is no evidence of an auditory evoked potential in this site to the auditory standard and the deviant waveform shows no obvious evoked componentry either. Further, follow-up running t-tests indicated that the onset of differences between the standard and deviant responses actually preceded the onset of physical deviance, making it clear that this apparent effect could not be related to the duration difference.

Discussion

In launching this investigation, we theorized that there was a strong case for suspecting that the auditory and somatosensory systems might share processing resources for amodal features, since work on auditory-somatosensory integration processes had already identified considerable common circuitry. The auditory and somatosensory systems have been shown to have strong neurophysiological links (Foxe *et al.*, 2000, 2002, 2005; Schroeder *et al.*, 2001, 2004; Lutkenhoner *et al.*, 2002; Schroeder & Foxe, 2002, 2005; Fu *et al.*, 2003; Kayser *et al.*, 2005; Murray *et al.*, 2005), making them ideal candidates in which to investigate the processing of amodal stimulus properties. For example, multi-contact linear array electrode recordings in macaques showed that both the somatosensory and auditory systems feed directly into layer IV, the afferent input layer, of the caudomedial belt area

(CM) with identical timing (see Schroeder & Foxe, 2002), and this multisensory region is just a single synapse from primary auditory cortex (A1). Using functional Magnetic Resonance Imaging (fMRI) Foxe *et al.*, (2002) found that tactile stimulation elicited activation in a very similar region in human auditory cortex, a finding since replicated and extended using fMRI in macaques (Kayser *et al.*, 2005). A key question becomes what the functional role of these hierarchically early multisensory regions might be. We reasoned that one possibility might be basic feature detection for properties common to both sensory systems, in this case simple duration processing. The MMN response was therefore employed to test whether the generators of duration change detection for somatosensory and auditory cues might rely on common multisensory circuitry. Previous work had pointed to the gross similarities between the sMMN and the aMMN (Spackmann *et al.* 2010), but a common circuitry account had not been directly tested. Here, using high-density mapping we found that both the morphology and the topographical distributions of the duration MMN were highly distinct across sensory systems. The sMMN topography at 235ms showed a clear frontal positive distribution, while the aMMN topography at the same timing showed a distinctly different fronto-central negative distribution. Further supporting separate underlying neural generators, source modeling solutions indicated that the dominant neural generators resided in or close to auditory and somatosensory cortical regions respectively for the aMMN and sMMN (see supplementary data). As such, these results in conjunction with the intracranial recordings make it clear that the somatosensory and auditory MMNs to duration deviants are generated in separate cortical regions, and we find no evidence for shared common amodal processing regions for duration detection.

General overview of aMMN and sMMN

The aMMN responses and topographies presented in this paper are similar to other duration aMMNs reported in the literature (Näätänen, 1992; Amenedo & Escera, 2000; De Sanctis *et al.*, 2009). The aMMN response over frontal midline scalp had two negative peaks followed by a positive peak. The second and larger amplitude negative peak at ~235ms is most commonly defined as the duration aMMN, and here has a topography that is entirely typical of an auditory duration MMN (Doeller *et al.*, 2003; De Sanctis *et al.*, 2009). The intracranial data localized the generator of the aMMN to the superior temporal gyrus (STG) and along the Sylvian fissure, which is consistent with findings in other studies (Dittmann-Balcar *et al.*, 2001; Schall *et al.*, 2003; Edwards *et al.*, 2005; Molholm *et al.*, 2005; Rinne *et al.*, 2005).

The sMMN response over the frontal midline scalp is also representative of what has been shown in previous literature (Kekoni *et al.*, 1997; Shinozaki *et al.*, 1998; Spackman *et al.*, 2007, 2010). The sMMN waveform has two phases: an earlier negative peak at ~145ms followed by a positive peak at ~235ms. Since this is the first study, that we are aware of, to use a high-density montage to assess the sMMN, a detailed assessment and comparison of its topographical distribution to that of the aMMN was possible. The early phase of the sMMN had a negative fronto-central distribution that evolved into a later positive frontal distribution. From the intracranial recordings, the sMMN was localized over the postcentral gyrus, with very similar timing to that seen in the scalp recordings, consistent with other intracranial recordings of the sMMN (Spackman *et al.*, 2010).

Comparison of the Auditory and Somatosensory Mismatch

To the best of our knowledge only one paper has previously looked at the sMMN and aMMN using a within-subjects design (Restuccia *et al.*, 2007). These authors recorded scalp EEG data from patients with unilateral cerebellar damage while conducting a location mismatch paradigm, where the standard and deviant were vibro-tactile stimulation of the first and fifth fingers respectively. When the ipsilateral hand to the damaged cerebellum was stimulated, an abnormal sMMN was elicited, whereas stimulation on the contralateral hand

elicited a normal sMMN. In a control condition, two of the patients were also tested using an auditory MMN paradigm where the deviant feature was a frequency change and it was found that their aMMN was unaffected by the cerebellar lesion, regardless of side of presentation. Thus, the data point to some dissociation of the generators of the MMN across these two modalities. However, since these authors were not specifically investigating the processing of amodal stimulus features, they did not equate the tasks across modalities, using different feature dimensions as deviants in both (location versus frequency). An amodal system would be expected to operate on a single feature and so it is difficult to draw any direct conclusions regarding this issue based on their data. Here we made a direct comparison of the aMMN and sMMN for the common feature of duration.

While our results clearly demonstrate that sensory-specific cortical areas code for duration processing, the possibility certainly remains that these regions might interact under multisensory stimulation conditions, or indeed that additional amodal regions might be activated for multisensory inputs. Some support for this thesis comes from a recent Transcranial Magnetic Stimulation (TMS) study (Bolognini et al., 2010). In this study, a single TMS pulse was delivered over auditory processing regions along the STG at varying intervals (60, 120 and 180 ms) after the presentation of tactile stimuli upon which participants were required to perform temporal discriminations. It was shown that TMS at 180 ms over STG resulted in impairment of temporal discrimination, suggesting that auditory regions on the STG were involved in cross-sensory tactile temporal processing. These results make a compelling case for interactions between unisensory auditory and tactile duration processing regions. We suggest that our finding of anatomically distinct automatic duration processing for auditory and tactile information makes the case for duration being processed and represented in sensory specific areas under conditions of unisensory stimulation, but that these areas may well be expected to interact when concurrently engaged by multisensory inputs. Under this account, one could posit that the TMS manipulation of Bolognini and colleagues resulted in co-activation of auditory and tactile regions, and that it was this co-activation that lead to interference in tactile duration discriminations. Certainly there is known inter-connectivity between the auditory and somatosensory systems (see Smiley and Falchier, 2009), and there are numerous examples of the bidirectional influence of these sensory systems on perception and performance (e.g. Lappe et al., 2008; Jousmaki & Hari, 1998; Guest et al., 2002; Guest & Spence, 2003; Foxe, 2009; Sperdin et al., 2009; Yau et al., 2009). The highly similar timing (~235ms) of the primary aMMN and sMMN peak responses seen here also lends further credence to the possibility of auditory and tactile duration information interacting under multisensory stimulation conditions, but this remains to be directly tested.

Despite a clear dissociation of the areas involved in auditory and somatosensory duration processing as indexed by the MMN, these data do not rule out the possibility that amodal duration representations come online during the active performance of a duration task, or that there might be amodal processing that feeds into sensory specific areas. For example, Moberget et al (2008) found that patients with cerebellar degeneration showed a selective auditory deficit in the duration-MMN (relative to frequency and location MMNs), and it is possible that the role of the cerebellum in the processing of temporal information is amodal. It is also possible that there might be other metrics of unisensory duration processing that would reveal an amodal representation. However, the MMN provides a solid metric of duration processing for which there are no alternatives that we are aware of, and the loci of MMN generation is commonly considered to reflect where the “deviant” feature is processed (Molholm et al., 2005). It is also the case that the auditory-MMN is influenced by multisensory inputs (Saint-Amour et al., 2007), and thus it seems an excellent candidate for measuring a potentially amodal process.

Conclusion

We tested whether processing of the amodal property of duration might rely on common circuitry across audition and touch, given that there is a large emerging body of anatomical and physiological work pointing to hierarchically early common multisensory regions that process inputs from both of these sensory systems. Instead, data from both scalp and subdural recordings point to clearly separate generators for simple duration detections across these two sensory systems, strongly implicating a sensory-specific account.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

Acknowledgments

This study was supported by a grant from the U.S. National Institute of Mental Health (NIMH) to J.J.F. and S.M. (MH85322). Dr. Mercier received additional support from a post-doctoral fellowship award from the Swiss National Science Foundation (PBELP3-123067). The Cartool software (<http://brainmapping.unige.ch/Cartool.htm>) was programmed by Denis Brunet of the Functional Brain Mapping Laboratory, Geneva, Switzerland, and is supported by the Center for Biomedical Imaging (CIBM) of Geneva and Lausanne. The authors would like to thank our colleague Dr. Pierfilippo De Sanctis for helpful discussions.

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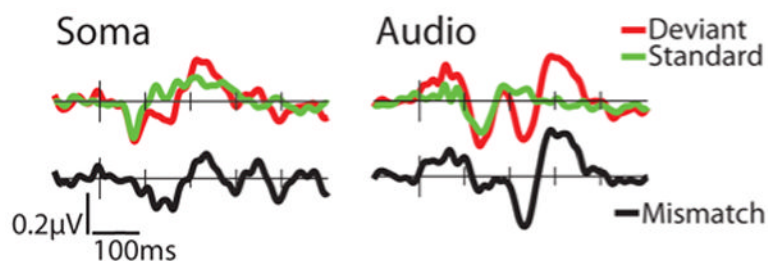


Figure 1.

Grand Mean ERPs for the somatosensory and auditory conditions at the Fronto-central site, Fz, for deviant, 50ms (red line), standard 100ms (green line) and the subtraction waveform, Mismatch (black line). The times indicate the midpoint latencies of the 40ms windows over which the average amplitude of the MMNs were measured.

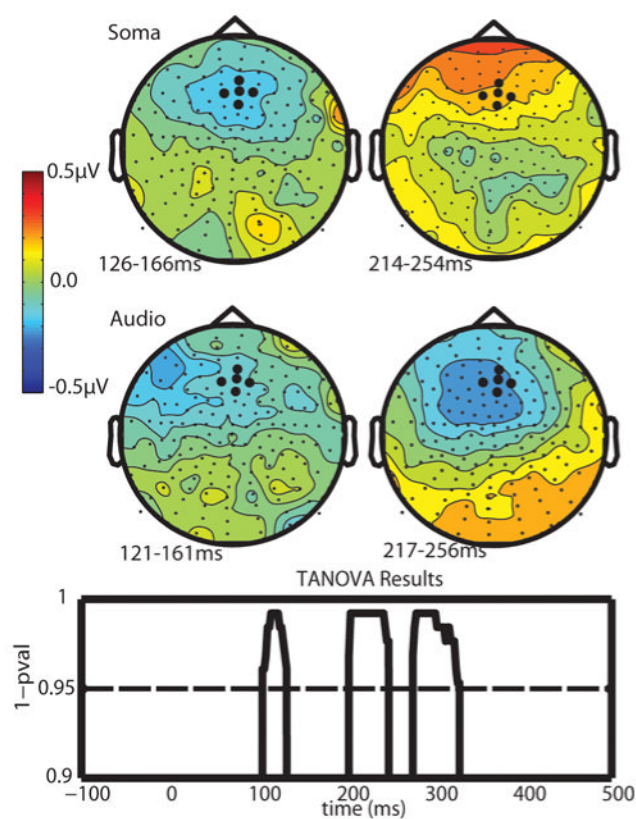
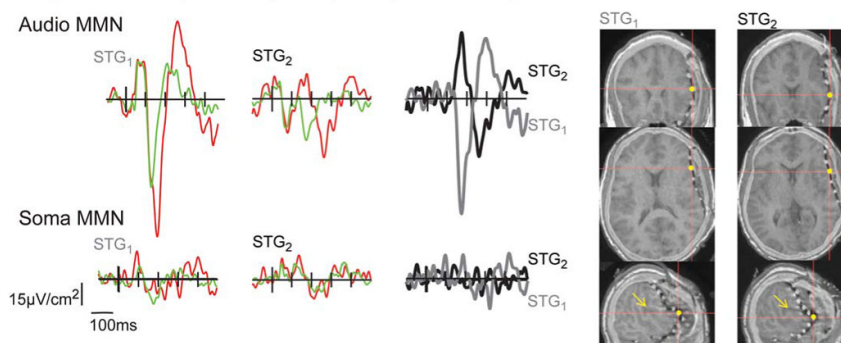


Figure 2.

Topographical plots of the grand average Mismatch waves for each modality (row) and each phase (column). The MMN is plotted at peak latency for each modality at the two different phases. The TANOVA tested for topographical differences at each time point between the sMMN and the aMMN.

Superior Temporal Gyrus (STG) - Auditory Cortex



Postcentral Gyrus (PcG) - Somatosensory Cortex

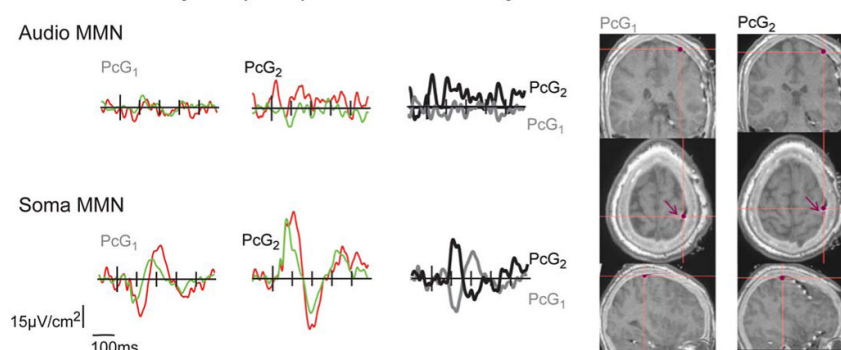


Figure 3.

Grand mean Current Source Density, CSD, at intracranial electrodes sites deviant (red line), standard (green line) and the subtraction waveform, MMN (black and grey line). The yellow and magenta dots represent the electrode locations on the MRI-CTs. The yellow and magenta arrows indicate the Sylvian fissure and the post-central gyrus, respectively.