Predicting Conceptual Processing Capacity from Spontaneous Neuronal Activity of the Left Middle Temporal Gyrus

Citation

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Accessibility
Conceptual processing is a crucial brain function for humans. Past research using neuropsychological and task-based functional brain-imaging paradigms indicates that widely distributed brain regions are related to conceptual processing. Here, we explore the potential contribution of intrinsic or spontaneous brain activity to conceptual processing by examining whether resting-state functional magnetic resonance imaging (rs-fMRI) signals can account for individual differences in the conceptual processing efficiencies of healthy individuals. We acquired rs-fMRI and behavioral data on object conceptual processing tasks. We found that the regional amplitude of spontaneous low-frequency fluctuations in the blood oxygen level-dependent signal in the left (posterior) middle temporal gyrus (LMTG) was highly correlated with participants’ semantic processing efficiency. Furthermore, the strength of the functional connectivity between the LMTG and a series of brain regions—the left inferior frontal gyrus, bilateral anterior temporal lobe, bilateral medial temporal lobe, posterior cingulate gyrus, and ventromedial and dorsomedial prefrontal cortices—also significantly predicted conceptual behavior. The regional amplitude of low-frequency fluctuations and functionally relevant connectivity strengths of LMTG together accounted for 74% of individual variance in object conceptual performance. This semantic network, with the LMTG as its core component, largely overlaps with the regions reported in previous conceptual/semantic task-based fMRI studies. We conclude that the intrinsic or spontaneous activity of the human brain reflects the processing efficiency of the semantic system.

Introduction

Semantic memory is a system for the storage, retention, and recall of general conceptual knowledge about objects, people, facts, and beliefs that are unrelated to specific experiences (Tulving, 1972). This system serves as a foundation for various cognitive processes including language, object recognition and use, reasoning, and problem solving. Evidence from neuropsychological and task-based functional brain-imaging studies has shown that performing semantic/conceptual tasks (we do not intend to distinguish between semantics and conceptual knowledge here, and use these two terms interchangeably) implicates multiple areas in the temporal, frontal, and frontoparietal regions (Dronkers et al., 2004; Binder et al., 2009).

To identify how conceptual knowledge is maintained at rest in the absence of specific inputs or outputs, the current study explores the role of spontaneous brain activity in semantic memory using resting-state functional magnetic resonance imaging (rs-fMRI). We focus on object conceptual processing in relation to low-frequency (<0.08 Hz) fluctuations (LFFs) in the blood oxygen level-dependent (BOLD) signal at rest, given that these fluctuations are related to spontaneous neuronal activity (Logothetis et al., 2001; Raichle, 2006).

Previous investigations of the regional amplitude of the LFFs (ALFF) have found that the ALFF reflects physiological signals: the ALFF of gray matter is higher than that of the white matter (Biswal et al., 1995), the ALFF of the so-called “default mode network” (regions that are activate during resting state and deactivated during task performance) is higher than that of other regions (Zang et al., 2007), and individuals with cognitive brain disorders show abnormal ALFF in the regions critical for the corresponding cognitive processes (Zang et al., 2007; Hopstman et al., 2010). Healthy individuals’ resting-state ALFFs correlate with task-evoked BOLD responses and with participants’ behavioral measures (Mennes et al., 2011). Furthermore, high synchronization of LFFs between areas within the same neuroanatomical and/or functional systems has been reported (Biswal et al., 1995; Fox et al., 2005; Koyama et al., 2010). Significantly, the degree of such synchronization is associated with variability in healthy individuals’ cognitive processing ability (e.g., reading ability) (Hampson et al., 2006), performance improvements (e.g., visual-detection performance) (Lewisa et al., 2009), and personality traits (e.g., autistic traits) (Di Martino et al., 2009). Thus, it ap-
pears that regional activity amplitude and connectivity patterns of the LFFs are associated with aspects of cognitive functioning. In this study, we examined the extent to which intrinsic brain activity (both regional activity and functional connectivity patterns) at rest predicts individual variation in semantic processing capacity: (1) we performed a correlation analysis between participants’ performance on conceptual tasks and their regional ALFF to uncover potential core regions that could account for individual variation in semantic processing efficiency; (2) we defined such core regions as seeds and computed the functional connectivity strength between each seed and other voxels to reconstruct the brain network associated with the core regions, and then we assessed the predictive power of these connectivity strengths for subjects’ semantic performance. We found that regional ALFF in the left (posterior) middle temporal gyrus (LMTG) and its functional connectivity with a set of other regions strongly predicted individual variation in conceptual processing capacity.

Materials and Methods

Participants

Thirty-four healthy, right-handed college students (20 females; 22.5 ± 1.3 years old; range, 20 - 26 years old) with no history of neurological or psychiatric disorders were recruited from the campus of Beijing Normal University as paid participants, and written informed consents were obtained. This study was approved by the Institutional Review Board of the University as paid participants, and written informed consents were obtained. Subjects were free of neurological or psychiatric disorders, and all subjects should have no history of substance abuse or addiction.

Behavioral tests

To assess semantic processing ability, we focused on two commonly studied object categories: animals and artifacts. We selected three classification with a set of other regions strongly predicted individual variation in conceptual processing capacity.

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pass filter (0.01–0.08 Hz) was applied to reduce low-frequency drift and high-frequency noise. Finally, spatial smoothing (4 mm FWHM Gaussian kernel) was conducted to decrease spatial noise.

Because LFF is sensitive to signals in the gray matter, additional analyses were conducted on the gray matter mask generated using the following procedure. We included the voxels with a probability higher than 0.4 in the SPM5 template onto the gray matter mask. Given the signal distortion in cerebellum, we also excluded the cerebellar regions (radius = 6 mm) in the Automated Anatomical Labeling template (Tzourio-Mazoyer et al., 2002). In total, there were 36,272 voxels in the gray matter mask.

Regional analysis: ALFF calculation and ALFF-behavior analysis procedures. Following Zang et al. (2007), the ALFF value of each voxel in the brain was extracted as the sum of amplitudes within the low-frequency range (0.01–0.08 Hz). Specifically, a filtered time series was transformed to the frequency domain to obtain a power spectrum for each voxel. Next, the square root was calculated at each frequency of the power spectrum. The averaged square root across 0.01–0.08 Hz at each voxel was taken as the ALFF. We scaled the ALFF values for each participant for standardization purposes: the mean ALFF value within the gray matter mask of the participant was computed, and then the ALFF of each voxel was divided by this mean ALFF value.

We used two methods to approximate the semantic behavioral performance from the three semantic tasks. We examined whether regional spontaneous activity was related to semantic processing capacity using measures derived from both methods. The first approach (Method 1) was to obtain a composite score using the three semantic tasks and to conduct correlation analyses between this composite semantic score and the ALFF (for a similar rationale, see Schwartz et al., 2009). The second approach (Method 2) was to consider the three semantic tasks separately and then to explore areas shared by these tasks. In Method 1, we averaged the efficiency scores for the three semantic tasks to generate a semantic composite score for each participant, which was taken as an index of the participant’s semantic processing capacity. We performed a regression analysis to reveal the relationship between ALFF and semantic performance while partialling out the contribution of general response latency differences. Specifically, we regressed out the efficiency scores on the cued articulation and shape-matching control tasks from the semantic composite efficiency score and then computed the correlation coefficient between the ALFF and the residualized semantic composite score. In Method 2, we explored any potential overlap among the areas associated with different semantic tasks to identify the common region(s) for all three semantic tasks. To obtain a correlation map for a specific semantic task, we calculated the correlations between the ALFF and the residualized efficiency score in this specific task after regressing out the efficiency score in the corresponding control task. For the two naming tasks, the correlation map was obtained through a correlation between the ALFF and residualized efficiency scores in the naming task after regressing out the efficiency scores in the cued articulation task. Similarly, the correlation map for picture associative matching was obtained through a correlation between the ALFF and residualized efficiency scores in the picture associative-matching task after regressing out the efficiency scores in the shape-matching task. We then checked for the common regions for the three semantic tasks.

Network analysis: functional connectivity analysis and connectivity-behavior analysis procedures. On the basis of the results of regional ALFF analyses, we further explored whether any observed core region works in concert with other regions as a network for semantic processing. Our rationale was to first use the observed region(s) as seed(s) to perform functional connectivity analyses, mapping out the regions that function with the seeds as a network. Next, we examined whether or not any specific connections within this network were able to predict semantic behavior.

Before conducting functional connectivity analyses, six head motion parameters, white matter, and cerebrospinal signal were first regressed out. Functional connectivity analysis was performed using the Resting-State IMRI Data Analysis Toolkit (REST; http://www.restfmri.net). A spherical seed ROI (radius = 6 mm) was created, centering on the coordinates of each peak point identified in the regional ALFF-behavior correlation analysis. To obtain the functional connectivity map for each participant, we considered both of the two relevant measures: regression coefficient ($\beta$) and correlation coefficient ($r$). First, we calculated the mean time series of the seed ROI for each participant. For the same gray matter mask applied in the above regional analysis, we calculated the regression coefficients ($\beta$) and correlation coefficients ($r$) between the seed time series and other voxels to obtain a $\beta$ map and an $r$ map for each participant. Fisher z score transformations were conducted for the correlation coefficients ($r$) to generate a z-functional connectivity (FC) map for each participant. To identify the regions showing significant functional connectivity with the seed(s), we did one-sample t tests on these individual $\beta$ maps or z-FC maps to see whether they were significantly different from zero ($t > 6.43$; corrected $p < 0.01$; Bonferroni correction). For these regions, we performed correlation analyses between participants’ semantic performance and $\beta$ values or Fisher z scores while regressing out the peripheral response components reflected in the performance of the control tasks.

Validating the functional roles of the semantic network. To examine whether the regions found in the ALFF-FC-behavior analyses are specific to (object) conceptual processing or are more generally involved in other cognitive processes, we conducted the following ROI analyses. First, these regions and/or networks were defined as ROIs, and the corresponding ALFF/FC values within each ROI were averaged. Then correlation analyses were performed between averaged ALFF/FC values and participants’ behavioral indexes in the number judgment task.

Reconstructing the default mode network. We compared the obtained resting-state semantic network with the default mode network, which is active during the resting state, is deactivated by most cognitive tasks but not by semantic tasks (Raichle et al., 2001), and overlaps in part with semantic networks derived from task-based fMRI research (Binder et al., 2009). To identify the default mode network in our participant group, we performed functional connectivity analyses (both correlation coefficient and regression coefficient measures) based on one previously identified seed region located in posterior cingulate gyrus (PCG; radius = 6 mm; central coordinates, $-5$, $-49$, $40$) (Fox et al., 2005). Then, individuals’ correlation coefficients were transformed into z-FC maps by the Fisher z score transformation. One-sample t tests were then performed on these individual $\beta$ maps and z-FC maps ($t > 6.43$; corrected $p < 0.01$; Bonferroni correction).

Statistical analysis. AlphaSim was used to correct for multiple comparisons (originally in AFNI software and implemented in REST), and the corrected threshold was set at $p < 0.05$. More specifically, the threshold of regional ALFF-behavior analysis was the combination of a voxelwise $p$ value of $<0.05$ and a cluster size of $>58$ voxels (1566 mm$^3$). Because connectivity-behavior analyses were conducted on brain areas showing significant functional connectivity with the seeds, the threshold was a combination of a voxelwise $p$ value of $<0.05$ and a cluster size of $>45$ voxels (1215 mm$^3$) (small volume correction).

Results

Regional ALFF analysis and ALFF-behavior analysis

Participants’ mean response times, accuracies, and inverse efficiency values on the behavioral tasks are shown in Table 1.

To explore the association between semantic behavior and regional resting-state activity, we first correlated the averaged efficiency scores of the three semantic tasks (Method 1) with the ALFF value of each voxel across the whole brain. The ALFF of one region, the LMTG, was positively correlated with participants’ semantic performance ($t_{\text{peak}} = 0.68$; $r_{\text{cluster}} = 0.81$; $p$ values $<0.001$; $t_{\text{peak}}$ correlation coefficient between the ALFF in the peak coordinate and behavioral scores; $r_{\text{cluster}}$, correlation coefficient between the averaged ALFF within the significant region and the behavioral scores; Fig. 1A). Thus, the regional ALFF variance of LMTG accounted for 66% ($R^2$) of the variation in semantic task performance across individuals.

When we conducted separate analyses on the three semantic tasks (Method 2), the LMTG was the only brain region in which the ALFF was significantly positively correlated with (1) picture-naming performance with cued articulation performance regressed out ($t_{\text{peak}} = -60$, $-48$, $-3$; 2268 mm$^3$ (84 voxels); $r_{\text{peak}} =$
0.71; \( r_{\text{cluster}} = 0.73; p \text{ values} < 0.001 \) and (2) picture associative-matching performance regressed out (peak, \(-66, -45, -3\); 3375 mm\(^3\) (125 voxels); \( r_{\text{peak}} = 0.81; r_{\text{cluster}} = 0.59; p \text{ values} < 0.001 \). In the object sound-naming task, we found positive correlations in right superior temporal and inferior parietal lobes (peak, 54, -36, 24; 2511 mm\(^3\) (93 voxels); \( r_{\text{peak}} = 0.67; r_{\text{cluster}} = 0.67; p \text{ values} < 0.001 \). It should be noted that the particularly low levels of accuracy and large variance in performance in this task may have originated in part at the object sound perceptual/recognition phase. Since we did not include a control task for this phase of the object sound-naming task, we lowered the threshold to corrected \( p < 0.1 \) [a combination of a voxelwise \( p \) value of <0.05 and a cluster size of \( >1377 \text{ mm}^3 \) (51 voxels)]. We found that, with cued articulation regressed out, LMTG showed association with object sound-naming performance (peak, \(-66, -42, -6\); 1431 mm\(^3\) (53 voxels); \( r_{\text{peak}} = 0.59; r_{\text{cluster}} = 0.64; p \text{ values} < 0.001 \). Together, the findings obtained with the two methods converge in showing that the LMTG is relevant for semantic processing regardless of stimulus input and output modality.

### Functional connectivity analysis and connectivity-behavior analysis

To explore whether LMTG functions in concert with other brain regions for semantic processing, we performed a seed voxel correlation analysis between the LMTG region found with the semantic composite score (Method 1) and all other voxels in the brain. As seen in Figure 2, A and B, we found similar functional connectivity patterns using the regression coefficient and correlation coefficient measures. A wide range of brain regions showed significant functional connectivity with LMTG, including the bilateral temporal gyri, inferior occipital gyri, middle and inferior frontal gyri, precentral cortices, ventromedial and dorsomedial frontal cortices, angular gyri, and posterior cingulate gyri. We defined these regions as a large ROI mask and correlated each voxel value within this ROI with semantic composite scores across participants. For the regression coefficient approach, the strength of functional connectivity between LMTG and the following regions significantly predicted participants’ semantic performance: left inferior frontal gyrus (LIFG), bilateral lateral temporal lobe including anterior temporal lobe (ATL), left medial temporal lobe (LMTL), PCG, dorsomedial prefrontal cortex (DMPFC), and ventromedial prefrontal cortex (VMPFC) (Fig. 3; Table 2). When the correlation coefficients were considered, significant correlations were also observed between functional connectivity in most of these regions and LMTG and participants’ semantic performance (Table 2).

We performed ROI analyses to further examine whether the regional activity of these regions (in the regression coefficient results) correlated with semantic processing performance. First, the averaged ALFF value within each region was obtained. Then correlation analyses between participants’ semantic composite scores and averaged ALFF values for each region were performed. No region showed significant effects in these ROI analyses (\( r \text{ values} < 0.27; p \text{ values} > 0.12 \)).

### Combining regional- and connectivity-behavior analyses

To assess the joint contributions of the regional activity and connectivity of LMTG in predicting semantic behavior, we performed a multiple linear regression analysis. The dependent variable was participants’ semantic composite scores after regressing out the efficiency scores on the cued articulation and shape-matching control tasks. Independent variables included the ALFF values of LMTG and seven functional connectivity variables. The ALFF variable was obtained by averaging ALFF values within the LMTG region that was significant in the regional analysis (Method 1). The functional connectivity variables were extracted by averaging the regression coefficients within each observed region in the functional connectivity-behavior analyses. When all independent variables were simultaneously entered into the regression, the regression model explained 74% of the variation in participants’ semantic processing performance (\( R^2 = 0.74; F_{(8,25)} = 8.96; p < 0.001 \)). We also used a forward method to test how much independent contribution each of the two types of variables provided. When the ALFF variable was entered first and the 10 functional connectivity variables were entered in the second step, we found that the inclusion of the functional connectivity variables yielded a trend of improvement for the explanatory power of the regression model (\( R^2 = 0.66 \rightarrow 0.74 \)), but the effect was not significant (\( F_{(7,25)} = 1.18; p = 0.35 \)). When the functional connectivity variables were entered first, the addition of ALFF into the model yielded significant \( R^2 \) change (\( R^2 = 0.49 \rightarrow 0.74 \); \( F_{(1,25)} = 24.18; p < 0.001 \)). These results further highlight the central role of LMTG’s regional activity in predicting semantic performance.

### Table 1. Participants’ performance on behavioral tasks

<table>
<thead>
<tr>
<th>Tasks</th>
<th>Response time (ms) Mean</th>
<th>SD</th>
<th>Accuracy Mean</th>
<th>SD</th>
<th>Inverse efficiency Mean</th>
<th>SD</th>
</tr>
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<tr>
<td>Semantic tasks</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Object picture-naming</td>
<td>1082</td>
<td>152</td>
<td>94%</td>
<td>4%</td>
<td>1149</td>
<td>187</td>
</tr>
<tr>
<td>Object picture associative-matching</td>
<td>1608</td>
<td>243</td>
<td>91%</td>
<td>6%</td>
<td>1781</td>
<td>292</td>
</tr>
<tr>
<td>Object sound-naming</td>
<td>2857</td>
<td>506</td>
<td>76%</td>
<td>10%</td>
<td>3832</td>
<td>968</td>
</tr>
<tr>
<td>Other tasks</td>
<td></td>
<td></td>
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<td></td>
<td></td>
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<tr>
<td>Cued articulation</td>
<td>376</td>
<td>91</td>
<td>100%</td>
<td>0%</td>
<td>376</td>
<td>91</td>
</tr>
<tr>
<td>Shape matching</td>
<td>487</td>
<td>72</td>
<td>96%</td>
<td>4%</td>
<td>506</td>
<td>64</td>
</tr>
<tr>
<td>Number judgment</td>
<td>1316</td>
<td>310</td>
<td>93%</td>
<td>4%</td>
<td>1409</td>
<td>317</td>
</tr>
</tbody>
</table>

### Figure 1. Regional ALFF-behavior correlation analyses using composite semantic scores

A. Statistical map for the correlation between ALFF and semantic scores. The correlation value is indicated using the color scale to the top. B. Scatter plot shows the positive correlation between semantic performance and averaged ALFF in the region plotted in A, while the efficiency scores on the cued articulation and shape-matching control tasks were regressed out from the semantic composite efficiency score. Each dot represents data from one participant.
The functional specificity of the observed semantic network

The results of ROI analyses for the number judgment task are shown in Table 3. Within the LMTG network we obtained for the semantic tasks, we observed that the intrinsic brain activity (i.e., both regional activity and functional connectivity) was not significantly associated with the number-processing task. These results indicate that different domains of conceptual processing (object vs numbers) can be distinguished.

Default mode network and semantic network

As shown in Figure 4A, bilateral inferior parietal lobe, medial prefrontal cortex, superior frontal cortex, medial temporal lobe, and lateral temporal cortex had strong functional connectivity with PCG, and these regions constitute the default mode network. Figure 4B replots the data presented in Figures 3 and 4A, showing that the semantic network largely overlaps with the default mode network.
Regional activity imaging studies, the LMTG has been found to be activated by tasks that probe semantic processing through various modalities, including visual words and pictures (Vandenbergh et al., 1996), auditory words (Hickok and Poeppel, 2004), and tactile inputs (Stoeckel et al., 2003). Task-induced activity in the LMTG has also been observed for semantic information acquisition relative to baseline (Maguire and Frith, 2004). The essential role of the LMTG in word-level comprehension is also indicated by lesion-symptom mapping analysis on stroke patients (Bates et al., 2003; Dronkers et al., 2004; Schwartz et al., 2009) and voxel-based morphometric analyses on semantic dementia patients (Mummery et al., 2000). Turken and Dronkers (2011) showed previously that the LMTG has rich structural and spontaneous functional connectivity with other regions relevant for language comprehension, and they proposed that it plays a central role in comprehension.

In some task-based fMRI and lesion studies (Damasio et al., 1996; Martin et al., 1996), the LMTG has been shown to be more sensitive to certain semantic categories such as tools, while other studies have observed effects for a wide range of semantic categories (Rudrauf et al., 2008; Simmons et al., 2010), including verbs (Wills et al., 2011), suggesting that the effect reported here is not likely to be driven only by tool items. We nevertheless analyzed tool and animal items separately in our study, and found similar LMTG effects for the two categories (for tools, peak, −63, −45, −3; 3024 mm³ (112 voxels); r_peak = 0.64; r_cluster = 0.77; p values < 0.001; for animals, peak, −60, −48, −3; 3186 mm³ (118 voxels); r_peak = 0.63; r_cluster = 0.74; p values < 0.001). It is possible that the tool-specific effects reported in the literature originated from an LMTG region that was close to but different from ours (Simmons et al., 2010). We did show, however, that number judgment performance did not correlate with the observed object semantic LMTG network, but with parietal regions close to the classical number regions (Footnote a), suggesting that our technique has the potential to distinguish between conceptual domains.

Together, these findings constitute clear evidence for a critical role of the LMTG in semantic processing, at least for single object concepts (Lau et al., 2008). The present study further strengthens this view by showing the robust effect of the spontaneous activity of this region in predicting individual participants’ variation in semantic processing behavior.

LMTG as a core semantic region
The LMTG region identified here corresponds to the middle part of Brodmann’s area (BA) 21, extending to BA 37. In brain-imaging studies, the LMTG has been found to be activated by tasks that probe semantic processing through various modalities, including visual words and pictures (Vandenbergh et al., 1996), auditory words (Hickok and Poeppel, 2004), and tactile inputs (Stoeckel et al., 2003). Task-induced activity in the LMTG has been observed to be sensitive to certain semantic categories such as tools, while other regions have been found to be activated by the LMTG region identified here corresponds to the middle part of the regions that were reported in a previous meta-analysis of classical number regions (Footnote a), suggesting that our technique has the potential to distinguish between conceptual domains.

Regions for which connectivity with LMTG predicted semantic processing behavior
Among the brain regions showing resting-state synchronization with the LMTG, a subset of these regions further predicted variation in semantic processing behavior, although their contribution beyond the regional ALFF effect of the LMTG was not significant. Among these regions, the LIFG and left ATL have consistently been observed to have white matter fiber pathways...
(i.e., anatomical connectivity) projecting to the LMTG in diffusion tensor imaging (DTI) and anatomical dissection studies (Catani et al., 2005; Catani and Mesulam, 2008; Papagno et al., 2011; Turken and Dronkers, 2011). Importantly, these fiber bundles seem to be specifically engaged in semantic and not phonological tasks (Saur et al., 2008, 2010). Furthermore, LIFG, especially the specific region implicated in the current study (BA 47), is activated by a variety of semantic tasks (Fiez, 1997; Poldrack et al., 1999; Friederici et al., 2003). The involvement of the ATL in semantic processing is most strongly suggested by semantic dementia patients, who suffer from atrophy of the ATL accompanied by progressive semantic impairment (for review, see Patterson et al., 2007).

Using DTI, Gong et al. (2009) reported that white matter fiber bundles exist between the LMTG and PCG. PCG has been shown to be associated with various cognitive functions, including episodic retrieval (Wagner et al., 2005), self-monitoring (Vogt and Laureys, 2005), self-reflection (Johnson et al., 2002), and awareness (Vogt and Laureys, 2005). We speculate that semantic processing might induce episodic encoding and thus can lead to PCG engagement.

The anatomical connections between the other three regions (MTL, VMPFC, and DMPFC) and LMTG are far from clear. Neurophysiological, neuroimaging, and neuropsychological studies provide converging evidence that the MTL is critical for semantic processing, especially the acquisition and retrieval of semantic memory (for review, see Squire et al., 2004). VMPFC has been strongly linked to social cognition. Specifically, it has been ascribed a crucial role for the encoding of enduring social dispositions and interpersonal knowledge through the integration of social information over long stretches of time in multiple circumstances (Van Overwalle, 2009). Such processes may be shared by semantic memory, which also implicates the abstraction of consistent information from disparate instances of an object or event and the decontextualization of individual memories (Takashima et al., 2006). The role of DMPFC is less clear, but, as speculated by Binder et al. (2009), it might be involved in “self-guided retrieval” aspects of semantic processing. Lesions to this region have been found to induce symptoms of very little spontaneous speech or action despite the preserved ability to speak or act when prompted (Nagaratnam et al., 2004).

Finally, the resting-state semantic network observed here largely overlaps with the default mode network (Shulman et al., 1997; Raichle et al., 2001). It has been proposed that during its conscious resting state, this network reflects ongoing semantic processing, such as semantic knowledge retrieval and manipulation of represented knowledge for problem solving (Binder et al., 2009). Other researchers have linked the network’s activity to internally generated thought processes related to self-reflection (Gusnard et al., 2001) or mind wandering (Mason et al., 2007)—processes that implicate a critical role for semantics. Our results further reinforce the notion that the function of the default mode network involves semantic processing.

**Potential mechanisms for the rs-fMRI–behavior correlation**

The neuronal origin of rs-fMRI activity has been supported by many types of findings, including its correlation with electrophysiological recordings of neuronal firing (Nir et al., 2008), structural connectivity, slow cortical potentials, and the band-limited power of fast electrical activity (for review, see Zhang and Raichle, 2010). The correlation between rs-fMRI and behavioral efficiency reported here might be driven by either genetic and/or environmental variables. The influence of genetic factors on specific rs-fMRI patterns has been demonstrated (Glahn et al., 2010), and it is not implausible to assume that variation in genetic disposition may optimize brain regions/networks to acquire and process semantic knowledge most efficiently. On the other hand, training has also been shown to affect rs-fMRI patterns (Lewisa et al., 2009), and it is possible that the richness of semantic experience shapes the rs-fMRI patterns in relevant regions. Future studies should consider the relative contribution of these variables in determining rs-fMRI patterns.
In conclusion, we observed a network, with the LMTG as its core component, in which the resting-state activity predicts healthy individuals’ conceptual processing efficiency. This finding has multifaceted theoretical implications: (1) it further confirms the roles of these regions, especially the central role of LMTG, in object conceptual processing; (2) it reveals a potential new mechanism for the representation and “maintenance” of semantic memory (i.e., through resting-state fluctuations); and (3) the extensive overlap between the resting state of the semantic network and the default mode network reinforces previous claims that the function of the default mode network may involve semantic processing. Finally, our findings have important clinical implications, providing potential biomarkers for the early detection of conceptual knowledge deterioration.

References

Wei et al. • Predicting Conceptual Behaviors from Resting-State Activity


