Auditory verbal hallucinations (AVHs) are one of the most common symptoms in schizophrenia. Current cognitive models suggest that auditory hallucinations are the result of defective self-monitoring (1–4). A feed-forward mechanism (5) has been proposed to explain self-monitoring of motor actions. In this model, motor commands that are needed to achieve a specific goal are identified and subsequently issued. Simultaneously, an efference copy is generated and transmitted through a corollary discharge mechanism to the sensory brain regions that are relevant to the planned act. The efference copy of the motor command serves to predict the sensory effects of the motor act. If the actual and predicted sensory feedback match, the actual sensory feedback will be attenuated. If a movement is externally controlled, there will be no efference copy that can attenuate the sensory information. Hence, the monitoring of motor acts can help to recognize whether a movement is self-initiated or externally generated. If generation of language and thought can be considered as a type of motor act (6–8), then monitoring of the verbal act may contribute to the distinction between self as source versus others as source. If this monitoring system is defective, verbal thoughts will not be recognized as being self-generated, leading to the experience of AVHs (8).

Background: Auditory verbal hallucinations in schizophrenia have been linked to defective monitoring of one’s own verbal thoughts. Previous studies have shown that patients with auditory verbal hallucinations show attenuated activation of brain regions involved with auditory processing during the monitoring of inner speech. However, there are no functional magnetic resonance imaging studies explicitly comparing the perception of external speech with internal speech in the same patients with schizophrenia. The present study investigated the functional neuroanatomy of inner and external speech in both patients with schizophrenia and healthy control subjects.

Methods: Fifteen patients with schizophrenia and 12 healthy control subjects were studied using functional magnetic resonance imaging while listening to sentences or imagining sentences.

Results: Significant interactions between group (control subjects vs. patients) and task (listening vs. inner speech) were seen for the left superior temporal gyrus, as well as regions within the cingulate gyrus.

Conclusions: Attenuated deactivation of the left superior temporal gyrus in schizophrenia patients during the processing of inner speech may reflect deficits in the forward models subserving self-monitoring.

Key Words: Auditory verbal hallucinations, fMRI, inner speech, schizophrenia, temporal lobe, verbal self-monitoring
Impairments in the auditory cortical processing and to exclude the increased demands associated with generating more complex stimuli such as imagery. To our knowledge, there are no fMRI studies explicitly comparing this in patients with schizophrenia.

In the present study, we used fMRI to investigate brain activation during the perception of internally generated speech and external speech in both patients with schizophrenia and matched healthy control subjects. Activation during the self-monitoring of inner speech was compared directly with activation during listening to externally generated speech, a task requiring no self-monitoring. We predicted that inner speech would lead to attenuation of temporal cortex activation in healthy control subjects as a consequence of corollary discharge, while the patients with schizophrenia would show less attenuated activation of the temporal cortex during inner speech trials, indicating faulty verbal self-monitoring. Since external sensory stimulation does not lead to an efference copy, we anticipated an interaction between task (listening vs. inner speech) and group (control subjects vs. patients) in the temporal cortex. As silent articulation is associated with activation of the left inferior frontal cortex, we also predicted that inner speech trials would be accompanied by an increased activation of this area compared with listening trials in both patients and the control subjects. The present study minimized the effects of acoustic scanner noise during the stimulus presentation through the use of a partially silent acquisition in which the auditory stimulus preceding the inner speech and listening prompts was presented in a partially silent gap followed by a period of continuous image acquisition (19).

**Methods and Materials**

**Subjects**

Fifteen male patients with a DSM-IV diagnosis of schizophrenia and right-handed, as assessed with the Annett Handedness Inventory (20), participated in the study. Patients were recruited through consultant and key worker recommendations and had all experienced prominent auditory hallucinations during exacerbations of their illness. Fourteen subjects were outpatients and one subject was an inpatient. All patients were receiving regular doses of antipsychotic medication. Mean age of the patients was 34.7 years (SD = 8.7). Mean duration of illness was 11.2 years (range 3–27). Mean score on the Positive and Negative Syndrome Scale (21) was 48.5 (SD = 16.5, range 30–83). Patients were matched for age, sex, and handedness to a control group. Twelve healthy right-handed, male control subjects were recruited through an advertisement in a city-wide newspaper. They did not suffer from psychiatric disorders and had no family history of psychiatric disorder. Mean age of the control subjects was 34.4 years (SD = 7.9). English was a first language for all subjects and all subjects had a minimum of 11 years of education. Exclusion criteria were any illicit drug use within the previous 6 months or any contraindications to magnetic resonance imaging (MRI) scanning. Potential subjects were assessed on their ability to perform the tasks (detailed below) outside the scanner, approximately 1 week before scanning. Subjects provided written informed consent, and ethical approval was provided by the Institute of Psychiatry and Maudsley National Health Services Trust.

**Tasks Performed During fMRI**

Subjects performed two active tasks, listening and inner speech, and there was one additional null baseline condition. Each of these tasks was administered over six counterbalanced blocks, each block comprising four listening trials, four inner speech trials, and four baseline trials, with the baseline trials consisting of a silent period equal in length to four paired stimuli (Figure 1).

Auditory stimuli used for the listening and inner speech conditions were 24 neutral sentences, spoken by an adult female native English speaker. During the listening trials, the auditory stimulus was followed by a visual cue prompting the subjects to listen to a second auditory stimulus identical to the first; during the inner speech trials, the auditory stimulus was followed by a visual cue prompting the subjects to covertly imagine repeating the sentence to themselves in their own voice and press a button with their right index finger once this was completed.

Each auditory stimulus was presented once during a listening trial and once during an inner speech trial. Sentences were presented via pneumatically driven earphones, incorporated within ear defenders, specifically designed for functional MRI (Quiet Muff 29 Earmuffs, Avotec, Jensen Beach, Florida). These reduced unattenuated noise from the scanner. As some sentences were longer than others, the duration of the stimuli varied from 1124 msec to 2862 msec, with an average length of 1971 msec. The auditory stimulus was followed by a 1000-msec period in which the visual prompt was presented, followed by the actual task (listening/inner speech). There was a gap before the onset of the next trial (intertrial interval: 14,728 msec). Image acquisi-
tion was performed during this nontrial interval. Total length of the task was 17 minutes and 39 seconds.

Image Acquisition

Gradient-echo echo-planar imaging data were acquired on a neuro-optimized GE Signa 1.5 Telsa system (General Electric, Milwaukee, Wisconsin) at the Maudsley Hospital, London. For complete details on methodologies, please see Section 1 in Supplement 1.

Image Analyses

Data were realigned (22) to minimize motion-related artefacts, smoothed using a Gaussian filter (full-width at half maximum 7.2 mm) and transformed into Talairach space (23). Task (listening vs. inner speech) and group (control subjects vs. patients) specific comparisons were carried. Second, the interaction between task and group was tested by subtracting the inner speech trials from the listening trials. Cluster-level statistics (22) corrected for multiple comparisons were thresholded at \( p < .009 \) to ensure less than one false-positive cluster per image. For complete details on methodologies, please see Section 2 in Supplement 1.

Results

Behavioral Data

The mean response time to complete the inner speech task was 2.62 seconds (SD = 1.59) for control subjects and 2.28 seconds (SD = .85) for patients. This difference was not statistically significant \( t(22) = .64, p = .53 \). On postscan debriefing, all subjects reported that they had been able to perform the tasks and thus data from all subjects were included in the analyses.

Imaging Data: Inner Speech Versus Listening

The main effect of inner speech was associated with greater activation in the left inferior frontal gyrus and anterior cingulate gyrus compared with listening (Figure 2A, Table 1). Listening was associated with greater activation in the right superior temporal gyrus, left transverse temporal gyrus, and right inferior parietal lobule across both groups.

Between-Group Differences in Activation

The main effect of group showed significant differences in the left and right occipital gyrus (Figure 2B, Table 1). Patients demonstrated greater bilateral activation of the occipital gyri during the active tasks compared with the control subjects, who showed greater activation during the baseline condition.

Interaction Between Task and Group

Significant interactions between group (control subjects vs. patients) and task (listening vs. inner speech) were seen for the left superior temporal gyrus, bilateral anterior cingulate gyrus, right hippocampus, and the left posterior cingulate gyrus (Figure 3A, Table 1). The plots from the peak sum of squares response in the left superior temporal gyrus of both patients and controls demonstrated activation during the listening task compared with baseline. However, during the generation of inner speech, the control subjects showed markedly attenuated activation, while this difference between inner speech and listening was less pronounced in the patients (Figure 3B).

Within the other regions with significant interactions, including the right hippocampus and regions within the cingulate gyrus, control subjects showed activation during inner speech compared with listening, while the patients showed a decrease in activation during the inner speech.

Table 1. Regions of Differential Activation

<table>
<thead>
<tr>
<th>Region of Activation (BA)</th>
<th>Coordinates</th>
<th>Cluster Size</th>
</tr>
</thead>
<tbody>
<tr>
<td>Effect of Task Inner Speech &gt; Listening</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Left inferior frontal gyrus (BA 44)</td>
<td>(-54, 4, 15)</td>
<td>33</td>
</tr>
<tr>
<td>Anterior cingulate gyrus (BA 32)</td>
<td>(4, 15, 42)</td>
<td>45</td>
</tr>
<tr>
<td>Inner Speech &lt; Listening</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Right superior temporal gyrus (BA 22)</td>
<td>(-58, -7, -2)</td>
<td>168</td>
</tr>
<tr>
<td>Left transverse temporal gyrus (BA 41)</td>
<td>(-58, -19, 9)</td>
<td>161</td>
</tr>
<tr>
<td>Right inferior parietal lobule (BA 40)</td>
<td>(33, -37, 53)</td>
<td>42</td>
</tr>
<tr>
<td>Effect of Group (Control Subjects &gt; Patients)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Left middle occipital gyrus (BA 18)</td>
<td>(-25, -81, 4)</td>
<td>59</td>
</tr>
<tr>
<td>Right middle occipital gyrus (BA 19)</td>
<td>(33, -74, 15)</td>
<td>33</td>
</tr>
<tr>
<td>Interaction</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Left superior temporal gyrus (BA 22)</td>
<td>(-54, -11, -2)</td>
<td>20</td>
</tr>
<tr>
<td>Left cingulate gyrus (BA 24)</td>
<td>(-11, -19, 42)</td>
<td>81</td>
</tr>
<tr>
<td>Right cingulate gyrus (BA 24)</td>
<td>(18, -15, 39)</td>
<td>221</td>
</tr>
<tr>
<td>Right hippocampus</td>
<td>(29, -37, -2)</td>
<td>69</td>
</tr>
<tr>
<td>Left posterior cingulate (BA 29)</td>
<td>(-11, -41, 9)</td>
<td>86</td>
</tr>
</tbody>
</table>

Thresholded at \( p < .009 \). Cluster size indicates total number of activated voxels in region. BA, Brodmann area.
The main finding was a significant interaction between group and task for the left superior temporal cortex. During the listening trials, the anticipated activation of the left superior temporal cortex was evident in control subjects and in schizophrenia patients, suggesting that listening to spoken sentences is not impaired in schizophrenia patients. Control subjects showed greater decrease in activation during inner speech compared with listening than patients. This provides evidence for defective self-monitoring of inner speech in schizophrenia patients. Failure to attenuate the activity in the temporal cortex may lead to the attribution of the verbal material as being of external origin, ultimately leading to auditory hallucinatory experiences.

The attenuation of temporal cortex activity is thought to be associated with corollary discharge from the inferior frontal cortex (24,25). Activation of the left inferior frontal cortex is thought to correspond to silent articulation (26–29). In line with previous studies (18,26), we found increased activation of the left inferior frontal gyrus during inner speech compared with the listening condition across groups.

Previous studies did not find clear differences between healthy control subjects and schizophrenia patients on an inner speech task (17,18). These studies used an inner speech task that required subjects to silently articulate a sentence in the form, “I like . . .” or “You are . . .,” appended with a single word that was presented to them in the auditory or visual modality. The task used in the present study differs from these early studies in requiring the subjects to repeat longer sentences that were presented to them aurally, placing higher demands on verbal self-monitoring. The same previous studies (17,18) did, however, find abnormal activation in the inferior frontal and temporal cortex during auditory verbal imagery, a task presumably requiring more self-monitoring than simple silent repetition. Thus, the findings of the present and previous studies suggest that when self-monitoring demands are absent or minimal, patients show no language network related abnormality, and it is only when the verbal material gets more complex and requires greater self-monitoring that the reduced attenuation of superior temporal cortex activation in patients with schizophrenia becomes apparent.

Studies on repetition priming in healthy control subjects show a decrease in activity in the left inferior prefrontal cortex and temporal cortex in response to repeated words in comparison with novel words (30). It has been suggested that repetition priming for semantic material may be reduced in patients (31). Since the auditory stimuli were repeated after presentation of a visual cue, it could be argued that the reduced attenuation in the patient group may be the result of reduced repetition priming. However, there were no significant group differences in activation of the left superior temporal gyrus during listening and there were no group differences in activation of the left inferior frontal gyrus. Furthermore, reaction times during the inner speech condition were not significantly shorter in control subjects, which may be expected in case of differential priming effects. The present study also differs considerably from the typical repetition priming designs in that the stimuli are more complex and the stimulus onset asynchrony is comparatively long (2124–3862 msec).

**Figure 3.** (A) Brain activation map for the interaction between the effects of task and group ($p = .009$, <1 false-positive cluster; blue: listening minus inner speech has higher peak SSQ in patients than in control subjects; yellow: listening minus inner speech has higher peak SSQ in control subjects than in patients). (B) SSQ plots for the interaction between task and group in the left superior temporal cortex. SSQ, sum of squares.
The present study found a main effect for the anterior cingulate gyrus, with inner speech trials eliciting more activation in this area than listening trials. There were significant interactions between group and task within the left and right anterior cingulate gyrus. Control subjects showed activation of these regions during inner speech and decreased activation when listening to external speech. Patients did not show this activation of the cingulate cortex during inner speech. The anterior cingulate has been implicated in divided attention (32) and competition monitoring (33), and patients with schizophrenia have been shown to display attenuated anterior cingulate activation during a cognitive inhibition task compared with control subjects (34). It has been suggested that the anterior cingulate may act as a top-down modulator of activity in the left superior temporal gyrus (35,36). This impaired modulation may be associated with external misattributions of inner speech (36).

The only between-group activations were in the occipital cortices; control subjects showed greater decreases in activation of the occipital cortex during the listening and inner speech tasks than the patients. Studies using fMRI have demonstrated that stimulation of one sensory modality can deactivate activation in other sensory regions (37,38). Functional MRI deactivation probably reflects inhibition of neural processes in task-irrelevant regions or regions that potentially interfere with the task at hand (39). This may indicate that schizophrenia patients show less cross-modal inhibition and may even engage some aspect of visual imagery upon hearing or generating the sentences. There were no group differences in the superior temporal gyrus during listening. Previous studies have found attenuated activation in response to listening to sentences (40,41); one reason for not replicating this finding may lie in the fact that the patients were not actively hallucinating during the imaging session. Earlier studies of inner speech per se have reported increased activation of the temporal cortex (42), but these differ significantly in including an active listening baseline rather than a null baseline as used in this study.

**Limitations**

Inner speech is a subjective phenomenon and as such cannot be easily objectively monitored. Training subjects on the task before scanning was done to minimize the influence of variation in performance. We also included an external measure of completion of the inner speech with an explicit button press, and there was no difference from control subjects in mean response time, suggesting that both groups performed the inner speech task in a similar manner. One further issue is related to the level of intrinsic, implicit, inner speech activity, which may be occurring during the baseline task; however, this is difficult to remove completely, as it occurs implicitly. The use of the identical baseline during the listening condition should go some way to eliminating any systematic effects due to this.

We selected patients on the basis of a prior history of auditory hallucinations, and the inclusion of a nonhallucinating group would have permitted the examination of the specificity of this finding with regard to hallucinations versus schizophrenia. Ford and Mathalon (43) conducted several studies using event-related potentials and electroencephalogram (EEG) gamma coherence as proxies of corollary discharge. These studies reported abnormalities in the corollary discharge system in patients with schizophrenia, without specificity to hallucinations. Other studies (44–46) using similar EEG paradigms have shown that the corollary discharge system is disrupted in patients, especially in those prone to AVHs. Even if inner speech deficits can be linked to AVHs, it remains to be seen whether these deficits can serve as the basis for all AVHs (47,48). Jones (48) argues that AVHs that involve voices attempting to regulate ongoing actions of the voice bearer may be consistent with inner speech based models, but other AVHs, e.g., those with a content clearly linked to previous traumatic experience, may be better explained by other cognitive models.

Intrinsic scanner noise poses a problem in all fMRI studies, especially in studies investigating auditory processing. The present study attempted to minimize scanner noise during the tasks by using a partially silent acquisition during the stimuli presentation. Nevertheless, scanner noise remains perceptible. Although scanner noise can significantly influence the blood oxygenation level-dependent response in the temporal cortex (19,49), the confounding effects of background noise should be considered as constant across conditions.

To summarize, our data indicate that when generating inner speech, schizophrenia patients show an attenuated deactivation in the left superior temporal gyrus, an area that has been implicated in verbal self-monitoring processes (18). This is consistent with the notion that auditory verbal hallucinations in schizophrenia may arise as a consequence of faulty predictive models underlying the monitoring of behaviors including inner speech (50,51).

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