Development of selective verbal memory impairment secondary to a left thalamic infarct: a longitudinal case study

J M Schott, S J Crutch, N C Fox, E K Warrington

A 68 year old man suffered an acute dysphasic episode with persistent memory disturbance while taking part as a control in a longitudinal magnetic resonance imaging (MRI) study. A small new left thalamic infarct involving the mammillo-thalamic tract could be demonstrated on volumetric MRI, coinciding with the development of a selective verbal memory impairment. This suggests that lateralisation of cognitive processing of visual and verbal material exists at the thalamic as well as the cortical level. High resolution volumetric MRI may be helpful in demonstrating small subcortical infarcts that may not be seen using computed tomography or conventional MRI.

Cerebrovascular disease may lead to memory impairment because of involvement of cortical or subcortical structures. It has long been recognised that discrete focal thalamic lesions may be associated with acute and persistent memory dysfunction.1 We report the case of a control subject taking part in a prospective longitudinal magnetic resonance imaging (MRI) and neuropsychology study who suffered a left thalamic stroke during the course of follow up. We were able to demonstrate the appearance of this lesion on MRI, and the coincident onset of new focal memory impairment.

CASE REPORT
ANC, a 68 year old right handed man, volunteered and consented to take part as a control subject in a longitudinal MRI study at the Institute of Neurology, London. He was seen for his first assessment in February 2001 when he was well, with no cognitive symptoms. He was a retired management consultant, whose past medical history included a hip replacement, mild renal impairment, and benign prostatic hypertrophy, for which he was taking finasteride. He did not smoke and he drank less than 21 units of alcohol a week.

Examination revealed hypertension (blood pressure 170/110), a body mass index of 28 kg/m², and was otherwise unremarkable. He scored 29/30 on a mini-mental state examination (MMSE).2 Baseline neuropsychology was undertaken (see below) and a volumetric MRI brain scan acquired. Over the following nine months he remained well and had five further volumetric MRI scans as part of the study. There was no evidence of sustained hypertension. He was started on orlistat during this time by his general practitioner to help weight control. He scored 30/30 on a mini-mental state examination (MMSE) with no cognitive symptoms. He was a retired management consultant and referred him to the local neurology service.

One month later he attended for his final study visit. He and his wife reported that his cognition had improved, but that although his ability to recognise faces was unimpaired, he had persistent difficulties recalling people’s names. He had also become more reliant on his diary. On examination, the MMSE was 26/30, the blood pressure was 130/85, and a neurological examination was normal.

Imaging
All volumetric imaging was done on the same 1.5 Tesla Signa unit, using a spoiled gradient echo technique with the same imaging parameters (TR/TE/TI/Theta 17/4.2/450/20, field of view 24 × 18 cm, 256 × 192 image matrix). All scans were reported by an expert neuroradiologist. The first scan revealed evidence of mild small vessel disease especially affecting the pallidum, with normal hippocampi and no evidence of global or regional cerebral atrophy. The next five scans (over the following nine months) revealed no significant atrophy and no new ischaemic lesions. The final scan, one month after the acute event, revealed a discrete new infarct in the left thalamus, involving the medial thalamic nuclei and interrupting the mammillo-thalamic tract (fig 1).

Neuropsychology
ANC was assessed at the start and end of the study using a standard battery of neuropsychological tests. A summary of these results is shown in table 1. At the start of the study, he performed in the superior range on tests of verbal and non-verbal intelligence, recognition memory, reading, and picture naming. Calculation was in the average range, and visuo-perceptual and visuospatial skills were satisfactory.

Reassessment was carried out at one year, a month after the acute event. The only change noted was a subtle decline in naming skills on the graded naming test; nonetheless, his score still fell within the superior range (90th centile). In the light of his persistent memory complaint, additional tests of word retrieval skills and episodic memory were administered.

Language skills
Two further stringent graded difficulty naming tests were attempted. ANC was able to name 22 of 30 objects and 28 of 30 animals (> 50th centile and > 90th centile, respectively) from the McKenna category specific names test.3 On a comparable proper noun retrieval test (historical figures, countries, buildings),4 his performance was extremely competent (25/30).

His good verbal comprehension was demonstrated on a test of knowledge of synonyms5 (concrete words: 22/25, > 50th
centile; abstract words: 25/25, > 90th centile). He also expressed himself fluently using a wide vocabulary. Thus at this stage there was no evidence of a dysphasic syndrome.

Episodic memory
The routine neuropsychological battery contained only the easy recognition memory test, on which ANC scored at ceiling. He was also tested on the standard recognition memory test, which allows visual and verbal memory to be assessed independently with an identical test design for each component. He scored 41/50 on the visual section of the test (faces: 50th centile). By contrast, on the verbal section of the test he scored only 36/50 (words: 5th centile). This discrepancy of five points represents a selective verbal memory deficit (p < 0.02).

On a demanding test of visual recognition memory—the topographical recognition memory test—he scored 26/30 (75th centile).

His performance was also impaired on a measure of verbal recall: he scored at the 5th centile on the Camden paired associate learning test (CPALT).

In an attempt to document this recall deficit in more detail, ANC was assessed with part of a longer famous faces test previously used by Cipolotti et al to provide evidence of retrograde amnesia in their patient vc. This test consists of 39 monochrome photographs of people from the 1990s famous in the fields of politics, entertainment, sport, and because of particular newsworthy events. The subject was first requested to recall the name of each individual orally. Subsequently, each photograph was re-presented with a choice of three names in a forced choice recognition paradigm—the target name alongside two equally famous distractor personalities. His scores were compared with those of 20 age matched control subjects tested by Cipolotti et al. He was able to recall only 21% of names correctly, significantly fewer than the control subjects (50%: z = 2.0, p < 0.02, two tailed test). However, no such difference was found using the forced choice measure (ANC 92%, controls 85%). These findings corroborate the evidence from the anterograde memory tests, providing further evidence for a selective verbal memory deficit.

DISCUSSION
Using longitudinal imaging in this individual, we have been able to demonstrate the appearance of a new discrete left sided thalamic infarct. At follow up, both ANC and his wife reported memory deficits occurring during the period over which the lesion appeared. Although he scored at ceiling on the easy recognition memory test at both the start and end of the study, his new reported memory impairment prompted us to undertake more detailed testing at follow up. Clear focal cognitive deficits were determined at this assessment, which we conclude are highly likely to be the result of the localised thalamic infarct.

Thalamic lesions may produce a wide range of neuropsychological deficits. The memory dysfunction associated with thalamic lesions appears to be best correlated with disruption of the mamillo-thalamic tract, as seen in this case. The role of the medial thalamic nuclei in memory dysfunction is less clear (for a review, see Van Der Werf et al). We have shown that ANC developed a selective episodic memory impairment for verbal material with preservation of visual memory—as demonstrated by his performances on the recognition memory test, the Camden paired associate learning test, and the topographical recognition memory test—and his poor recall of names of famous faces despite good recognition. There was no evidence of semantic memory impairment, as evidenced by the high score on the graded naming test. Thalamic lateralisation (where dominant lesions result in verbal memory impairment, and non-dominant lesions in visual memory) could be tested in a future study to determine the possible role of thalamic lateralisation in this case.

Table 1: Neuropsychological raw scores and centile rankings at the initial and one year assessments

<table>
<thead>
<tr>
<th>Neuropsychological tests</th>
<th>Assessment 1 (Feb 2001)</th>
<th>Assessment 2 (Feb 2002)</th>
</tr>
</thead>
<tbody>
<tr>
<td>The Wechsler abbreviated scale of intelligence:</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Vocabulary (raw score)</td>
<td>77</td>
<td>76</td>
</tr>
<tr>
<td>Matrix reasoning (raw score)</td>
<td>28</td>
<td>29</td>
</tr>
<tr>
<td>Predicted full scale IQ</td>
<td>134 (99%ile)</td>
<td>135 (99%ile)</td>
</tr>
<tr>
<td>National adult reading test:</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Number of errors</td>
<td>5</td>
<td>4</td>
</tr>
<tr>
<td>(Predicted IQ)</td>
<td>(124)</td>
<td></td>
</tr>
<tr>
<td>Easy recognition memory test:</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Words</td>
<td>25/25 (&gt;50%ile)</td>
<td>25/25 (&gt;50%ile)</td>
</tr>
<tr>
<td>Faces</td>
<td>25/25 (&gt;50%ile)</td>
<td>25/25 (&gt;50%ile)</td>
</tr>
<tr>
<td>Graded difficulty arithmetic test</td>
<td>14/24 (&gt;50%ile)</td>
<td>14/24 (&gt;50%ile)</td>
</tr>
<tr>
<td>Graded naming test</td>
<td>29/30 (&gt;99%ile)</td>
<td>25/30 (&gt;90%ile)</td>
</tr>
<tr>
<td>Visual object and space perception battery:</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Silhouettes</td>
<td>21/30 (&gt;25%ile)</td>
<td>21/30 (&gt;25%ile)</td>
</tr>
<tr>
<td>Number location</td>
<td>8/10 (&gt;5%ile)</td>
<td>10/10 (&gt;50%ile)</td>
</tr>
</tbody>
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%ile, centile.
memory problems) has been proposed by several investigators,\textsuperscript{1,16} although others have failed to demonstrate a consistent effect.\textsuperscript{17} Our findings support the hypothesis that lateralisation of cognitive processing of visual and verbal material exists at the thalamic as well as at the cortical level. While lesions throughout the dominant hemisphere can cause a selective impairment of verbal memory,\textsuperscript{10,11} it is rare for such lesions to be demonstrated longitudinally.

Small thalamic or other subcortical lesions may produce defects of memory or other cognitive functions. It is likely that many of these lesions will not be visualised using computed tomography, and may even be missed by conventional MRI. The possibility of a thalamic lesion should be considered in patients presenting with new persistent memory dysfunction,\textsuperscript{18} and may even be missed by conventional MRI. In these cases, small lesions may be demonstrated using the higher resolution afforded by volumetric MRI.

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