# Retrosplenial Amnesia without Topographic Disorientation Caused by a Lesion in the Nondominant Hemisphere

Shinichiro Maeshima, MD, PhD,\* Aiko Osawa, MD, PhD,\* Fumitaka Yamane, MD, PhD,† Tomoyuki Yoshihara, MD, PhD,† Ryuzaburo Kanazawa, MD, PhD,† and Shoichiro Ishihara, MD, PhD†

We report the case of a 68-year-old right-handed man who was admitted to our hospital because of sudden onset of headache. On admission, he presented with left homonymous hemianopsia, disorientation, and recent memory disturbance; however, he had normal remote memory and digit span. He was able to recall the room layout of his house and describe the route from the nearest station to his home on a map. However, at the hospital, he sometimes lost his way because of amnesia. Computed tomography (CT) and magnetic resonance imaging revealed a subcortical hematoma in the right occipital forceps and the parietal lobe, involving the cingulate isthmus. Single-photon emission CT imaging showed reduced perfusion not only in the retrosplenial region but also in the right thalamus. These findings suggested that the retrosplenial amnesia might have been caused by the interruption of hippocampal input into the anterior thalamus—retrosplenial region.

© 2014 by National Stroke Association

#### Introduction

Lesions of the limbic system, particularly the hippocampus or anterior thalamus, are well known to cause memory impairment. Retrosplenial cortex also plays an important role in the human memory by connecting the hippocampus with other structures involved with memory, including the anterior thalami, septal nuclei, and mammillary bodies.<sup>1</sup> Amnesia, resulting from lesions involving the retrosplenial region, has been attributed to the damage of this vital link. Valenstein et al<sup>1</sup> reported a case

1052-3057/\$ - see front matter

of retrograde and anterograde amnesia after hemorrhage because of an arteriovenous malformation situated close to the retrosplenium of the corpus callosum. They termed this amnesia "retrosplenial amnesia." Lesions in the left hemisphere are common; however, lesions in the right hemisphere are rare.<sup>2,3</sup> A lesion in the same location in the right hemisphere as that in the left hemisphere produces topographical disorientation.<sup>4,5</sup> We report a case of amnesia without topographical disorientation because of intracerebral hemorrhage in the right retrosplenial region.

#### **Case Report**

A 68-year-old man with an educational history of 9 years was admitted to our hospital. He was completely right handed with a laterality quotient of +100 when formally assessed by the Edinburgh Handedness Inventory,<sup>6</sup> and all members of his immediate family are right handed. He presented with a 3-day history of headache and had developed untidy behavior and memory disturbance over a couple of days. He had no neurological deficits except left hemianopsia and amnesia.

From the \*Department of Rehabilitation Medicine, Saitama Medical University International Medical Center, Hidaka, Saitama, Japan; and †Department of Endovascular Neurosurgery, Saitama Medical University International Medical Center, Hidaka, Saitama, Japan.

Received January 3, 2013; revision received March 11, 2013; accepted March 19, 2013.

Address correspondence to Shinichiro Maeshima, MD, PhD, Department of Rehabilitation Medicine II, School of Medicine, Fujita Health University, 424-1 Oodori-cho, Tsu, Mie 514-1295, Japan. E-mail: maeshima@fujita-hu.ac.jp.

<sup>© 2014</sup> by National Stroke Association

http://dx.doi.org/10.1016/j.jstrokecerebrovasdis.2013.03.026

Table 1.	Results	of	neuropsychological tests	5
----------	---------	----	--------------------------	---

	3 d	6 wk	Cutoff score
Mini-mental state examination (/30)	21	22	23/24
Digit span (forward/backward)	4/3	4/3	
Auditory verbal learning test (/15)			
Immediate	1, 5, 7, 8, 7	4, 5, 6, 5, 6	
Recognition	12	13	
Delayed	0	4	
Raven's progressive matrices (/36)	17	23	24/25
Frontal Assessment Battery (/18)	11	16	12/13
Trail-Making Test			
Α	2'15"	221"	
В	Not done	3'23"	
Wechsler Adult Intelligence Scale, Third Edition			*
Verbal IQ	72	80	
Performance IQ	57	72	
Full IQ	62	74	
Verbal comprehension	76	82	
Visual perception	54	75	
Working memory	65	76	
Process speed	66	75	
Wechsler Memory Scale-Revised			*
Verbal memory	70	87	
Visual memory	55	88	
General memory	61	86	
Concentration/attention	72	78	
Delayed recall	59	71	
The Rivermead Behavioral Memory			
Test			
Screening score (/12)	4	5	5/6
Profile score (/24)	9	15	15/16
Behavioral Assessment of Dysexecutive	44	69	
Syndrome			
Age matched score (classification)	Impaired	Impaired	

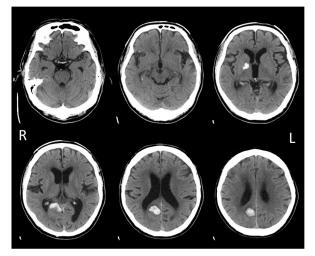
Abbreviation: IQ, intelligence quotient.

\*The median of Wechsler Scale is centered at 100, with an SD of 15.

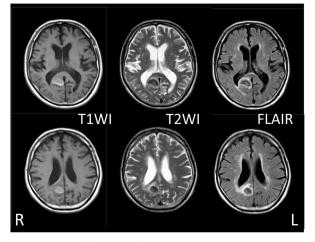
He was alert and cooperative to a series of neuropsychological examinations, although he was in an amnesic state, with disorientation and recent memory loss.

Retrograde episodic memory concerning both personal and public remote events was nearly intact, but he had anterograde amnesia. For instance, he had no memory of whether he had received his meals and repeatedly asked for them. He even kept forgetting the most recent conversations. He did not show confabulation. His speech was fluent and not aphasic. Objective naming, letter reading, verbal comprehension, and dictation were normal. He showed constructional apraxia, but ideomotor or ideational apraxia and unilateral spatial neglect were not noted. He had no signs of callosal disconnection such as apraxia or tactile anomia in his left hand. He could recall the room layout of his house and describe the route from the nearest station to his home on a map. However, he sometimes lost his way at the hospital because of amnesia without topographic disorientation.

Table 1 shows the results of neuropsychological tests in this patient. On the Wechsler Adult Intelligence Scale, Third Edition, his verbal intelligence quotient (IQ) was 72; performance IQ, 57; and full IQ, 62. His verbal memory was 70 and visual memory was 55 on the revised Wechsler Memory Scale. The score on the Frontal Assessment Battery was 11/18, and the word fluency test where the patient was required to produce exemplars from given categories resulted in scores of 9, 3, and 5 for the categories "animal," "fruit," and "vehicle," respectively, and where he was required to produce words that begin with given letters resulted in scores of 2, 5, and 3 for the letters "shi," "i," and "re," respectively. The level of executive functioning was classified as "impaired" on the basis of the scores of Behavioral Assessment of the Dysexecutive Function, that is, overall profile, 5, and age-corrected score, 44 (impaired).



CT scan on admission



MRI 6 days from the admission

**Figure 1.** CT scan on admission (upper) and magnetic resonance imaging tomography 6 days after the admission (lower). A subcortical hematoma was evident between the right forceps occipitalis and the parietal lobe, involving the right cingulate isthmus. A high-density area in CT scan was evident in right globus pallidus without pathologic calcification. Abbreviations: CT, computed tomography; FLAIR, Fluid attenuated inversion recovery; L, left; MRI, magnetic resonance imaging; R, right; T1WI, T1 weighted images; T2WI, T2 weighted images.

#### Neuroradiological Findings

Computed tomography (CT) and magnetic resonance imaging (MRI) revealed a subcortical hematoma  $(2 \times 2.5 \times 2 \text{ cm})$  between the right forceps occipitalis and the parietal lobe, involving the right cingulate isthmus (Fig 1). Single-photon emission CT imaging using Technetium-99m–labeled ethyl cysteinate dimer performed 3 days from the onset of stroke revealed reduced perfusion not only in the retrosplenial region but also in the right thalamus (Fig 2, A).

### Clinical Course

The patient was treated medically and visual field defect resolved within several days. He was discharged from the hospital 10 days after the onset. He was able to accurately recognize familiar buildings and find his way. The amnesia gradually improved, but memory impairments were detected at 6 weeks after the onset by using neuropsychological tests. Single-photon emission CT performed 45 days after onset showed an improvement of cerebral blood flow in the right thalamus (Fig 2, B).

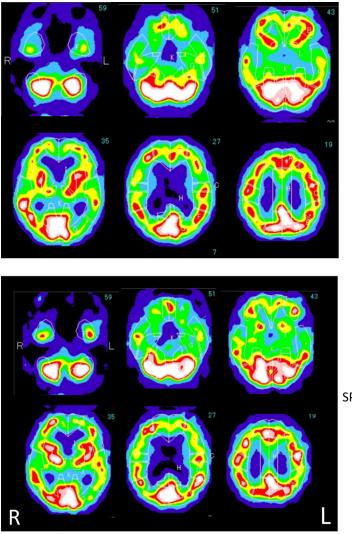
## Discussion

The sequential memory of our patient was intact, and he could recall prehospital events very well, indicating the absence of retrograde amnesia. However, he could not learn the disease name, patient room number, and the primary physician's name, suggesting mainly anterograde amnesia.

Damage to the retrosplenial region has been reported to be sufficient to induce anterograde amnesia.<sup>1-3,7-10</sup> Several etiologies of the tumor, infarction, and hemorrhage have been reported in patients with retrosplenial amnesia, but lesions in the right hemisphere are less common.

Regarding dysmnesia, the lesions in the left hemisphere are known to mainly impair verbal memory and those in the right hemisphere mainly impair visual spatial memory.<sup>11</sup> In our case, although the lesions were in the right hemisphere, both visual and verbal memories were impaired (visual > verbal). These findings suggested that the laterality of memory was not as distinct as that of language or visuospatial cognition, which was not limited to retrosplenial amnesia. In fact, some cases have been reported in which both the verbal and visual memories were impaired by the lesions in the right hemisphere.<sup>11</sup> Some experimental studies have also shown that anterograde disorientation can be caused by lesions on either side.<sup>12</sup> Thus, in our case, it seemed that the memory function was not completely lateralized congenitally, and the lesions in the right hemisphere caused damage to both verbal and visuospatial memory.

The isthmus of the cingulate gyrus connects the visual cortex in the occipital lobe with such limbic structures as the hippocampus, anterior cingulate gyrus, and anterior thalamus.<sup>13,14</sup> Lesions in the retrosplenial cortex may disrupt the connections between the anterior thalamus and hippocampus and thus cause memory disturbance. In the present case, amnesia improved only slightly with an increase in cerebral blood flow in the right thalamus. We suspected that the retrosplenial amnesia was caused by the interruption of hippocampal input into the anterior thalamus. Because the retrosplenial cortex itself is vital for memory, the remote effect might alter the impact of anterior thalamic lesions. These findings provide novel insights into the functional pathology of diencephalic amnesia and might be useful to determine the causes of posterior cingulate hypoactivity and show how distal changes in plasticity could contribute to



SPECT 3 days

Figure 2. SPECT imaging performed 3 days from stroke onset at around the revealed reduced perfusion not only in the retrosplenial region but also in the right thalamus (upper). SPECT performed 45 days after onset showed an improvement of cerebral blood flow in the right thalamus (lower). Abbreviations: L, left; R, right; SPECT, single-photon emission computed tomography. (Color version of figure is available online.)

# SPECT 45 days

diaschisis.<sup>15</sup> Therefore, the clinical course of retrosplenial amnesia might have a good prognosis.

Topographical disorientations are usually caused by the lesions in the retrosplenial region.<sup>5</sup> Our patient could not remember the name of the previously unknown hospital ward or the layout of the hospital and often could not find the toilet located several meters away from his room. However, he could accurately recognize familiar buildings and identify them on the map. Furthermore, he could describe the route from the nearest station to his home and could draw the map. The anterograde amnesia might have been caused by the retrosplenial lesion. We, therefore, suspected that his impaired ability to remember unfamiliar landscape was because of anterograde amnesia and not because of topographic disorder.

### References

1. Valenstein E, Bower D, Verfaellie M, et al. Retrosplenial amnesia. Brain 1987;110:1631-1646.

- 2. Yasuda Y, Watanabe T, Tanaka H, et al. Amnesia following infarction in the right retrosplenial region. Clin Neurol Neurosurg 1997;99:102-105.
- 3. Masuo O, Maeshima S, Kubo K, et al. A case of amnestic syndrome caused by a subcortical haematoma in the right occipital lobe. Brain Inj 1999;13:213-216.
- 4. Cammalleri R, Gangitano M, D'Amelio M, et al. Transient topographical amnesia and cingulate cortex damage: a case report. Neuropsychologia 1996;34:321-326.
- Takahashi N, Kawamura M, Shiota J, et al. Pure topographic disorientation due to right retrosplenial lesion. Neurology 1997;49:464-469.
- 6. Oldfield RC. The assessment and analysis of handedness: the Edinburgh inventory. Neuropsychologia 1971;9:97-113.
- Rudge P, Warrington EK. Selective impairment of memory and visual perception in splenial tumours. Brain 1991;114:349-360.
- 8. Maguire EA. The retrosplenial contribution to human navigation: a review of lesion and neuroimaging findings. Scand J Psychol 2001;42:225-238.
- Maeshima S, Ozaki F, Masuo O, et al. Memory impairment and spatial disorientation following a left retrosplenial lesion. J Clin Neurosci 2001;8:450-451.
- 10. Osawa A, Maeshima S, Kubo K, et al. Neuropsychological deficits associated with a tumour in the posterior

corpus callosum: a report of two cases. Brain Inj 2006; 20:673-676.

- 11. Aggleton JP. Understanding retrosplenial amnesia: insights from animal studies. Neuropsychologia 2010; 48:2328-2338.
- 12. Olton DS. Strategies for the development of animal models of human memory impairments. Ann N Y Acad Sci 1985;444:113-121.
- 13. Garden DL, Massey PV, Caruana DA, et al. Anterior thalamic lesions stop synaptic plasticity in retrosplenial cor-

tex slices: expanding the pathology of diencephalic amnesia. Brain 2009;132:1847-1857.

- 14. Rosene DL, van Hoesen GW. Hippocampal efferents reach widespread areas of cerebral cortex and amygdala in the rhesus monkey. Science 1977;198: 315-317.
- Irle E, Markowitsch HJ. Connections of the hippocampal formation, mammillary bodies, anterior thalamus and cingulate cortex. Exp Brain Res 1982; 47:79-94.