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# Recovery of an injured corticoreticular pathway via transcallosal fibers in a patient with intracerebral hemorrhage

Sung Ho Jang<sup>1</sup> and Sang Seok Yeo<sup>2\*</sup>

## **Abstract**

**Background:** Several studies have reported on injury of the corticoreticular pathway (CRP), however, little is known about the mechanism for recovery of an injured CRP. We report on a patient with intracerebral hemorrhage (ICH) who showed recovery of an injured CRP via transcallosal fibers, which was demonstrated by diffusion tensor tractography (DTT).

**Case presentation:** A 67-year-old man presented with complete paralysis (Medical Research Council: 0/5) of the left extremities at the onset of a right putaminal hemorrhage. At six weeks after onset, he presented with more severe weakness of proximal joint muscles than distal joint muscles (right shoulder abductor; 2<sup>-</sup>, finger extensor; 3<sup>+</sup>, hip flexor; 2<sup>+</sup>, ankle dorsiflexor; 3). Although his right hemiplegia had recovered well, he consistently showed more severe proximal weakness (right shoulder abductor; 3, finger extensor; 4, hip flexor; 3<sup>+</sup>, ankle dorsiflexor; 4) until 16 weeks after onset. On both six- and 16-week DTTs, in the left (affected) hemisphere, the CRP showed severe narrowing with discontinuation of the anterior fibers at the corona radiata on six-week DTT, however, the discontinued anterior fibers of the CRP were connected to the right cerebral cortex via transcallosal fibers on 16-week DTT.

**Conclusion:** We demonstrated recovery of an injured CRP via transcallosal fibers in a patient with ICH. We believe that this might be a mechanism for recovery of an injured CRP.

**Keywords:** Corticoreticular pathway, Corticoreticulospinal tract, Transcallosal fibers, Diffusion tensor tractography, Stroke

# **Background**

The corticoreticulospinal tract, one of the extrapyramidal motor pathways, consists of the corticoreticular pathway (CRP) and the reticulospinal tract. This tract innervates proximal muscles of the extremities and axial muscles [1-3]. Exact evaluation of the CRP had been impossible in the live human brain. However, Diffusion tensor tractography (DTT), which is derived from diffusion tensor imaging (DTI), has enabled three-dimensional visualization and estimation of the CRP [4]. Several studies have reported on injury of the CRP in various brain pathologies, including intracerebral hemorrhage (ICH), cerebral infarct, and

In this study, we report on a patient with ICH who showed recovery of an injured CRP via transcallosal fibers, which was demonstrated by DTT.

## Case presentation

A 67-year-old right-handed man presented with complete paralysis of the right upper and lower extremities (Medical Research Council [MRC]: 0/5) at the onset of a spontaneous ICH (Table 1). At six weeks after ICH onset, he was admitted to the rehabilitation department of a university hospital after undergoing rehabilitation for three weeks at the previous university hospital. Brain MR images showed a hematoma in the left corona radiata and basal ganglia at six weeks after onset (Figure 1-A). He presented with severe weakness of the right upper and lower extremities

Full list of author information is available at the end of the article



traumatic brain injury [5-7]. However, knowledge regarding the mechanism for recovery of an injured CRP is limited [8].

<sup>\*</sup> Correspondence: eangbul@hanmail.net

<sup>&</sup>lt;sup>2</sup>Department of Physical Therapy, College of Health Sciences, Dankook University, 119, Dandae-ro, Dongnam-gu, Cheonan-si, Chungnam 330-714, Republic of Korea

Table 1 Longitudinal changes in motor function

Duration from onset		Onset	6 weeks	10 weeks	16 weeks
MRC	Shoulder abductor	0	2-	2	3
	Elbow flexor	0	3	3+	3+
	Finger flexor	0	3+	3+	4
	Finger extensor	0	3+	3+	4
	Hip flexor	0	2+	3	3+
	Knee extensor	0	3-	3+	4
	Ankle dorsiflexor	0	3	4	4
MEP	Latency		21.7	22.6	21.8
	Amplitude		300	800	2400
	ET(%)		95	95	80

MRC: Medical Research Council scale. 0; no contraction, 1; palpable contraction, but no visible movement, 2; movement without gravity, 3; movement against gravity, 4; movement against a resistance lower than the resistance overcome by the healthy side, 5; movement against a resistance equal to the maximum resistance overcome by the healthy side. MEP: motor evoked potential, ET: excitatory threshold.

with more severe weakness of proximal joint muscles than distal joint muscles (MRC: right shoulder abductor; 2<sup>-</sup>, finger extensors; 3<sup>+</sup>, hip flexor; 2<sup>+</sup>, ankle dorsiflexor; 3) (Table 1). He was administered comprehensive rehabilitative therapy, which included neurotropic drugs (dopaminergic drug [pramipexole, amantadine, and levodopa] and antidepressant [venlafaxine]), movement therapy, and neuromuscular electrical stimulation of the affected shoulder abductor, finger extensor, and ankle dorsiflexor [9,10]. Movement therapy was conducted with a focus on improvement of right upper and lower extremity functions, and was conducted five times per week in our physical and occupational therapy department. Although the weakness of his right side recovered well for 10 weeks until 16 weeks after onset, he consistently showed more severe proximal weakness (right shoulder abductor; 3, finger extensor; 4, hip flexor; 3<sup>+</sup>, ankle dorsiflexor; 4). As a result, he was able to perform some fine motor activities using the right hand and to walk independently at 16 weeks after onset. The patient provided informed consent, and the study protocol was approved by our institutional review board.

## Diffusion tensor tractography

DTI scanning was performed at six, 10 and 16 weeks after onset, using a multi-channel head coil on a 1.5-T Philips Gyroscan Intera (Philips, Best, Netherlands) with single-shot echo-planar imaging. For each of the 32 non-collinear diffusion-sensitizing gradients, we acquired 67 contiguous slices (matrix = 192 x 192, FOV = 240 mm, TR/TE = 10.726/76 ms, b =  $1000 \text{ mm}^2\text{s}^{-1}$ , NEX = 1, thickness = 2.5 mm). Three-dimensional reconstructions of fiber tracts were obtained using the DTI task card software (Philips Extended MR Work Space 2.6.3) (threshold fractional anisotropy = 0.15, angle =  $27^\circ$ ) [11]. The CRP was determined by selection of fibers passing through the

seed and target regions of interest (ROI). A seed ROI was placed on the reticular formation of the medulla. The target ROI was placed on the midbrain tegmentum [1,4,7].

On six-, 10- and 16-week DTTs, the CRPs in the right hemispheres originated from the premotor cortex and primary motor cortex, and descended through the known CRP pathway. By contrast, in the left (affected) hemisphere, the CRP showed severe narrowing with discontinuation of the fibers at the corona radiata on six-week DTT. On the other hand, the discontinued anterior fibers of the CRP were connected to the corpus callosum through the transcallosal fibers on 10-week DTT, and became thicker and connected to the right cerebral cortex via the corpus calllosum on 16-week DTT (Figure 1-B).

## Transcranial magnetic stimulation

Transcranial magnetic stimulation (TMS) was performed to demonstrate the recovery process of the corticospinal tract using a Magstim Novametrix 200 magnetic stimulator with a 9-cm mean diameter circular coil (Novametrix Inc). Cortical stimulation was performed with the coil held tangentially over the vertex. The left hemisphere was stimulated by a counterclockwise current and the right hemisphere was stimulated by a clockwise current. Motor-evoked potentials (MEPs) were obtained from both abductor pollicis brevis muscles in a relaxed state. The excitatory threshold (ET) was defined as the minimum stimulus required to elicit an MEP with a peak-to-peak amplitude of 50  $\mu$ V or greater, in two out of four attempts. Stimulation intensity was set at the ET plus 20% of the maximum stimulator output. Each site was stimulated three times at intervals with a minimum of 10 seconds, from which the shortest latency and the average peak to peak amplitudes were adopted.

On the six-week TMS study, an MEP with low amplitude was evoked (latency: 21.7 msec, amplitude: 0.3 mV ET: 95%), and on the 10-week and 16-week TMS study, the amplitude was gradually increased (10 weeks - latency: 22.8 msec, amplitude: 0.8 mV, ET: 95%; 16 weeks - latency: 21.8 msec, amplitude: 2.4 mV, ET: 80%).

# **Discussion**

In the current study, we attempted to demonstrate the recovery of an injured CRP by following up on the change of the injured CRP from six weeks to 10 weeks and 16 weeks during a 10-week rehabilitation period. The left injured CRP showed discontinuation in the anterior fibers at the corona radiata level with preservation of the smaller posterior portion on six-week DTT and the anterior discontinued fibers were connected to the corpus callosum through the transcallosal fibers on 10-week DTT, and became thicker and connected to the right (unaffected) cortex via transcallosal fibers on 16-week DTT. During the 10-week rehabilitation period, the patient showed

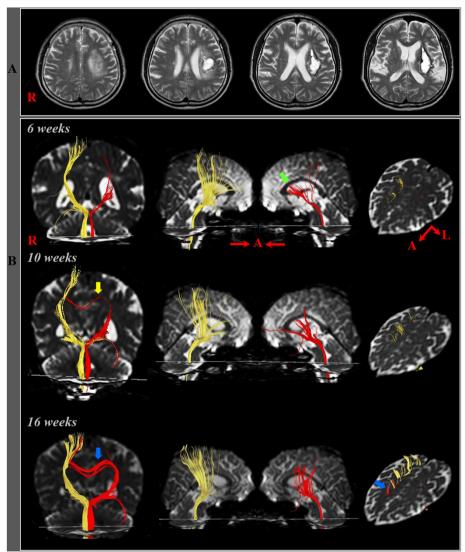


Figure 1 (A) T2-weighted MR images showed a hematoma in the left corona radiata and basal ganglia at six weeks after onset (B) Results of diffusion tensor tractography (DTT). On six-, 10- and 16-week DTTs, the CRPs in the right hemispheres originated from the premotor cortex and primary motor cortex, and descended through the known CRP pathway. By contrast, in the left (affected) hemisphere, the CRP showed severe narrowing with discontinuation of the anterior fibers (green arrow) on six-week DTT, however, the discontinued fibers of the CRP were connected to the corpus callosum through the transcallosal fibers (yellow arrow) on 10-week DTT, and became thicker and connected to the right cerebral cortex via the corpus calllosum (blue arrow) on 16-week DTT.

good motor recovery in both proximal and distal joint muscles (MRC: right shoulder abductor;  $2^- - > 3$ , finger extensor;  $3^+ - > 4$ , hip flexor;  $2^+ - > 3^+$ , ankle dorsiflexor;  $3^- > 4$ ). The motor recovery of distal joint muscles appears to be compatible with the increase of amplitude of MEP in the right hand muscle, which indicates an increase in the number of CST fibers [12,13]. The recovery of proximal joint muscles appears to be mainly attributed to the change of the left CRP: the anterior torn fibers of the left CRP at the corona radiata level were connected to the right (unaffected) cerebral cortex via transcallosal fibers. Therefore, we believe that this change of the injured CRP via transcallosal fibers might be a mechanism for recovery of an injured CRP.

To the best of our knowledge, recovery of an injured CRP has only been reported in one patient, using DTT [8]. In 2013, Yeo and Jang reported on a patient who presented with recovery of a discontinuation of the left CRP at the midbrain level at three weeks after recovery of an ICH through the normally existing pathway of the CRP to the left cerebral cortex at six weeks [8]. In their patient, the motor outcome in the proximal joint muscles (right shoulder flexor; 4, hip flexor; 4) at six weeks after onset was better than the motor outcome of our patient (the right shoulder abductor; 3, hip flexor; 3<sup>+</sup>) at 16 weeks after onset. This finding suggests that recovery via transcallosal fibers might accompany poorer motor

outcome than recovery via the normally existing neural pathway of the CRP. On the other hand, with regard to the CST, a few studies have reported on motor recovery by transcallosal fibers in stroke patients [14-16]. Consequently, to the best of our knowledge, our study is the first study to demonstrate motor recovery by transcallosal fibers of the CRP in a stroke patient.

#### Conclusion

We demonstrated recovery of an injured CRP via transcallosal fibers in a patient with ICH. We believe that this might be one of the mechanisms for recovery of an injured CRP. In addition, when comparing motor outcome with that of previous study [8], our results suggest that recovery via transcallosal fibers appeared to be associated with poorer motor outcome than recovery via the normally existing neural pathway of the CRP. However, this study is limited because it is a single case study. Conduct of further studies including larger case numbers is warranted. In addition, studies on rehabilitation strategies to induce or facilitate this recovery mechanism should be encouraged. Further studies on the other recovery mechanism following a CRP injury are also needed.

#### Consent

Written informed consent was obtained from the patient for publication of this case report and accompanying images. A copy of the written consent is available for review by the Editor-in-Chief of this journal.

#### Abbreviations

CRP: Corticoreticular pathway; DTI: Diffusion tensor imaging; DTT: Diffusion tensor tractography; ICH: Intracerebral hemorrhage; MRC: Medical research council; ROI: Regions of interest; TMS: Transcranial magnetic stimulation; MEPs: Motor-evoked potentials; ET: Excitatory threshold.

## Competing interests

The authors declare that they have no competing interests.

## Authors' contributions

SHJ: conceiving and designing the study, funding, data acquisition, manuscript development and manuscript writing. SSY: manuscript development, data acquisition, manuscript writing and manuscript authorization. Both authors read and approved the final manuscript.

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### **Author details**

<sup>1</sup>Department of Physical Medicine and Rehabilitation, College of Medicine, Yeungnam University 317–1, Daemyungdong, Namku, Taegu 705-717, Republic of Korea. <sup>2</sup>Department of Physical Therapy, College of Health Sciences, Dankook University, 119, Dandae-ro, Dongnam-gu, Cheonan-si, Chungnam 330-714, Republic of Korea.

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