# White matter plasticity in the cerebellum of elite basketball athletes

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**Abstract:** Recent neuroimaging studies indicate that learning a novel motor skill induces plastic changes in the brain structures of both gray matter (GM) and white matter (WM) that are associated with a specific practice. We previously reported an increased volume of vermian lobules VI–VII (declive, folium, and tuber) in elite basketball athletes who require coordination for dribbling and shooting a ball, which awakened the central role of the cerebellum in motor coordination. However, the precise factor contributing to the increased volume was not determined. In the present study, we compared the volumes of the GM and WM in the sub-regions of the cerebellar vermis based on manual voxel analysis with the ImageJ program. We found significantly larger WM volumes of vermian lobules VI–VII (declive, folium, and tuber) in elite basketball athletes in response to long-term intensive motor learning. We suggest that the larger WM volumes of this region in elite basketball athletes represent a motor learning-induced plastic change, and that the WM of this region likely plays a critical role in coordination. This finding will contribute to gaining a deeper understanding of motor learning-evoked WM plasticity.

Key words: Coordination, Motor learning, Magnetic resonance imaging, Vermis

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# Introduction

Advances in brain imaging have provided new insights into the relationship between motor learning and brain structure. Imaging investigations have revealed that motor learning and training can evoke plastic changes in brain structures that are associated with distinct training demands, and such learninginduced structural alterations have been demonstrated to arise in both the gray matter (GM) and white matter (WM) in the human brain.

Longitudinal investigations focused on motor learning have revealed structural plasticity in specific WM regions.

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For example, Juggling training increased fractional anisotropy (FA), a measure of the directionality of water diffusion to measure the WM microstructure, in the WM underlying the intraparietal sulcus [1]. Furthermore, bilateral upper extremity motor skill training increased FA in the posterior and anterior limbs of the internal capsule, the corona radiata, and the body of the corpus callosum [2], and complex visuomotor rotation training was found to result in increase in FA below the primary motor cortex [3]. Conversely, learning of a complex whole-body balance task decreased FA in prefrontal WM regions [4].

In our previous study, we found that elite basketball athletes showed larger volumes of vermian lobule VI–VII (declive, folium, and tuber) in the cerebellum after longterm intensive training. This finding indicated the important role of the cerebellum in motor learning and coordination [5]. However, we did not determine whether GM or WM contributed to the increased volume of the cerebellar vermis.

In this study, we further examined whether the increased

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volume of this region induced by extensive motor learning in elite basketball athletes is due to a larger GM or WM volume. The results of this study are expected to lead to a better understanding of learning-induced structural plasticity in the human brain. In addition, this study design is advantageous compared to pooling large datasets from elite athletes.

# **Materials and Methods**

### Subjects

Nineteen male elite basketball athletes group (BG) involved in national and international competitions were recruited from several university teams in South Korea through direct visiting interviews. Members of the BG with heights over 190 cm were excluded from this investigation to reduce the height variation among subjects. Twenty healthy male subjects who did not engage in any sports-related activity were selected as a control group (CG) through advertisements on Korea University web pages and in the local community newspapers.

Before magnetic resonance imaging (MRI) scanning, we obtained information about alcohol consumption and the type of alcohol consumed, as well as the use of nicotine and other drugs through a questionnaire. Neurological examination was performed by a neurologist at Korea University Medical Center, and any subjects with potential neurological impairments were excluded from the study. All subjects had healthy general medical and neurological profiles, and none of the subjects presented postural and vision problems.

The duration of training and practice time per day were checked in the BG, and body heights and weights were measured in all subjects. Table 1 presents the average age, height, and weight of all right-handed subjects. Written informed consent was acquired from all the subjects, and all procedures were approved by the Institutional Review Board of Korea University.

### **MRI** acquisition

MRI was executed on a 1.5-tesla Magnetom vision instru-

Table 1	Physical	characteristics	of subjects
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	BG (n=19)	CG (n=20)	P-value			
Age (yr)	20.31±1.05	21.60±2.68	0.058			
Height (cm)	$181.89 \pm 4.40$	180.70±3.11	0.333			
Weight (kg)	80.52±4.36	70.52±7.25	< 0.001***			
Practice hours per day	6.36±1.06	N/A	-			
Years of training	8.15±2.69	N/A	-			

Values are means±SD. BG, elite basketball athletes group; CG, control group; N/A, not applicable. \*\*\**P*<0.001.

ment (Siemens, Erlangen, Germany) using  $T_1$ -weighted magnetization prepared rapid acquisition gradient echo sequence (repetition time, 9.7 milliseconds; echo time, 4 milliseconds; flip angle, 12 degrees; slice thickness, continuous 2.0 mm; field of view, 250 mm; number of slices, 90; and matrix, 256×256). After the acquisition of magnetic resonance (MR) images, the Digital Imaging and Communications in Medicine (DICOM) files were transferred to an IBM compatible PC.

### Volumetry

The region of interest (ROI) areas of the cerebellar vermis was manually delineated on two sagittal slices of MR images (the mid-sagittal slice and the parasagittal slice in which the cerebral aqueduct became continuous with the fourth ventricle) using the ImageJ program (National Institutes of Health, Bethesda, MD, USA) to measure the volumes of the ROI's for GM and WM (Fig. 1). The numbers of pixels of marked region of each ROI were calculated by histogram analysis and then copied to an Excel sheet to distinguish between GM and WM based on gray-scale density (Fig. 1). A pixel density value from 26 to 100 was regarded as GM, and values greater than 105 were regarded as WM. The GM and WM volumes for all ROIs were calculated by multiplying the total numbers of pixels of each ROI by the voxel size ( $0.89 \times$  $0.89 \times 2.00$  mm).

Vermian V1 included lobules I–V (the lingual, centralis, and culmen), vermian V2 included lobules VI and VII (the declive, folium, and tuber), and vermian V3 comprised lobules VIII–X (the pyramis, uvula, and nodulus). All ROIs were independently drawn for each subject by two trained raters that were strictly blinded to the subjects' identity and assignment, and were checked by a neuroradiologist. Interrater reliability was tested using two-tailed Pearson correlation. Inter-rater correlation coefficients were over 0.90 for all ROIs. We normalized these ROIs by calculating the relative GM and WM ROI volumes for each subject as a percentage of their intracranial volume in order to minimize inter-subject variability.

## Statistical analysis

Statistical analyses were performed with the SPSS software package version 12.0 (SPSS Inc., Chicago, IL, USA). The Student's t test was performed to determine the effect of athletic proficiency on the absolute and relative GM and WM volumes of the ROIs. All analyses were two-tailed, and a P<0.05 was considered as statistically significant. All data are

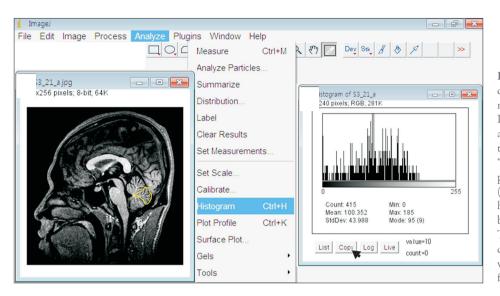


Fig. 1. Edited screen capture obtained during pixel counting to analyze gray matter and white matter with the ImageJ program. The region of interest area was marked using the free drawing tool in sagittal view (left image). The "Analyze-Histogram" menu gives the pixel density value as a separate window (right image). A valley observed in the histogram represents the cut-off point between gray matter and white matter. The detailed value can be copied by clicking the "copy" button (arrow) and was then pasted into an Excel program for further analyses.

Table 2. Absolute volumes (cm<sup>3</sup>) of each region of interest

	BG (n=19)	CG (n=20)	P-value
V1GM (cm <sup>3</sup> )	0.762±0.223	0.911±0.334	0.108
V1WM (cm <sup>3</sup> )	$1.032 \pm 0.313$	0.913±0.281	0.220
V2GM (cm <sup>3</sup> )	$0.576 \pm 0.277$	0.563±0.237	0.878
V2WM (cm <sup>3</sup> )	0.550±0.211	0.419±0.196	0.053
V3GM (cm <sup>3</sup> )	0.591±0.301	0.674±0.290	0.384
V3WM (cm <sup>3</sup> )	0.579±0.269	0.510±0.283	0.441

Values are means±SD. BG, elite basketball athletes group; CG, control group; V1GM, absolute volume of gray matter in vermian lobules I–V (lingual, centralis, and culmen); V1WM, absolute volume of white matter in vermian lobules I–V (lingual, centralis, and culmen); V2GM, absolute volume of gray matter in vermian lobules VI–VII (declive, folium, and tuber); V2WM, absolute volume of white matter in vermian lobules VI–VII (declive, folium, and tuber); V3GM, absolute volume of gray matter in vermian lobules VIII–X (pyramis, uvula, and nodulus); V3WM, absolute volume of white matter in vermian lobules VIII–X (pyramis, uvula, and nodulus).

represented as mean±SD.

# Results

Comparison of the absolute volumes of each ROI between the BG and CG showed no significant differences (Table 2). Comparison of the relative volumes of each ROI between the BG and CG showed significant differences in the relative WM volumes of V2 (Table 3). The relative GM volume of the V1, relative GM volume of the V2, relative GM volume of the V3, relative WM volume of the V1, and relative WM volume of the V3 showed no group differences.

### Table 3. Relative volumes (%) of each region of interest

	BG (n=19)	CG (n=20)	P-value
rV1GM (%)	$0.048 \pm 0.013$	0.058±0.022	0.111
rV1WM (%)	$0.066 \pm 0.020$	0.057±0.016	0.142
rV2GM (%)	$0.036 \pm 0.017$	$0.035 \pm 0.015$	0.906
rV2WM (%)	$0.035 \pm 0.014$	$0.026 \pm 0.011$	0.033*
rV3GM (%)	$0.037 \pm 0.018$	0.043±0.019	0.368
rV3WM (%)	0.037±0.017	0.031±0.017	0.336

Values are means±SD. BG, elite basketball athletes group; CG, control group; rV1GM, relative volume of gray matter in vermian lobules I–V (lingual, centralis, and culmen); rV1WM, relative volume of white matter in vermian lobules I–V (lingual, centralis, and culmen); rV2GM, relative volume of gray matter in vermian lobules VI–VII (declive, folium, and tuber); rV2WM, relative volume of white matter in vermian lobules VI–VII (declive, folium, and tuber); rV3GM, relative volume of gray matter in vermian lobules VI–VII (declive, folium, and tuber); rV3GM, relative volume of gray matter in vermian lobules VIII–X (pyramis, uvula, and nodulus); *rV*3WM, relative volume of white matter in vermian lobules VIII–X (pyramis, uvula, and nodulus). *\*P*<0.05.

# Discussion

In the present study, we identified larger WM volumes of the vermian lobules VI and VII (the declive, folium, and tuber) in the cerebellums of elite basketball athletes compared to those of controls. We suggest that the larger volume of vermian lobule VI–VII observed in elite basketball athletes in our previous study [5] was driven mainly by the WM of this region. The results of our study suggest that the structural alterations of WM in this region appear to have been induced by long-term extensive motor learning, and that the WM of this region might play an essential role in coordination.

Elite athletes usually begin sports training in early child-

hood, and tend to practice several hours per day intensively. Therefore, highly trained elite athletes could provide a good model to investigate training-induced structural plasticity in the human brain. A number of studies using MRI have revealed structural differences in the GM and WM of multiple brain regions in response to a practiced task between elite athletes and control subjects, which might reflect the effects of long-term motor learning [6-10]. However, inconsistent results have been reported for both GM and WM structures.

In GM structures, elite judo players had a larger GM volume in the frontal lobe which is related to motor planning and execution, as well as in regions of the prefrontal cortex, which is related to working memory and cognitive processes [6]. In addition, professional and low-handicap golfers had larger GM volumes in the fronto-parietal network, including premotor and parietal areas [7], and badminton athletes showed a greater GM concentration in the right anterior and posterior lobes of the cerebellum [8]. However, decreased GM volumes were observed in the left premotor cortex, supplementary motor area, putamen, and the superior frontal gyrus of professional female ballet dancers [9]. With respect to WM structure, world-class gymnasts showed higher FA values in the bilateral corticospinal track [10]. By contrast, professional and low-handicap golfers showed a lower WM volume and FA values in the vicinity of the corticospinal tract at the level of the internal and external capsule and in the parietal operculum [7], and professional ballet dancers showed a smaller WM volume in the corticospinal tracts, internal capsules, corpus callosum, and left anterior cingulum and lower FA values in the left and right premotor cortex [9].

Some longitudinal studies have shown a cause-and-effect relationship with respect to motor learning such as juggling training, bilateral upper extremity training, visuomotor rotation training, and balance training, and their direct influence on WM structure in the adult human brain [1-4]. Moreover, inter-individual variation in the WM microstructure is related to performance variation in a bimanual coordination task [11]. The cerebellum plays an important role in motor learning [12, 13], and motor learning-induced structural plasticity in the WM of the cerebellum has also been reported. MRI analyses of adult Japanese macaque monkeys showed an increased WM volume in the cerebellum after tool-use training [14]. Pianists and string players, as examples of individuals with specific motor expertise, showed a plastic change of WM in the cerebellum [15]. Indeed, FA in the WM of the cerebellum positively correlates with learning scores of a visuo-motor

task [16]. Motor coordination was shown to be involved in individual differences in WM integrity in the cerebellum in karate black belts groups [17]. These findings indicate that structural properties of WM in the cerebellum are relevant to motor learning.

WM, which lies under the GM cortex, consists of millions of axons that link neurons in different regions into functional circuits. The white color originates from the myelin that covers the axons. Myelin is crucial for the conduction of electrical impulses, and its deficit can damage neuronal transmissions [18]. Changes in WM, including the number of axons, diameter of axons, myelin thickness, axon branching, axon trajectories, and myelination, influence the speed of impulse propagation [19]. Thus, improving the velocity or synchrony of impulse conduction could be an essential feature of motor learning, and could in turn influence the improvement of performance with motor learning [19, 20].

Previous studies revealed that experience or training could change the WM structure. An enriched environment increases the number of unmyelinated and myelinated axons, as well as oligodendrocytes that form myelin in rats [21-26], and the size of the corpus callosum in rhesus monkeys [27]. Environmental effects on the WM structure have also been demonstrated in the human brain. Early experience increases the WM structure in the internal capsule and frontal lobes in newborn infants [28]. Tool use training induces extension of the axonal branches in the cerebral cortex of monkeys [29], and the amount of electrical activity within an axon could regulate its degree of myelination [30, 31].

In this study we attempted to match the age, height, and weight as much as possible because these variables could affect the cerebellar volume. There was significant difference in weight between BG and CG. Cerebellar volume significantly affected by body height [32]. Therefore, it is not likely that the weight variable affected the result of this study.

The results of our study, combined with the previous demonstration of WM alterations in the cerebellum of musicians [15] and tool-use trained monkeys [14], suggest that extensive motor learning may induce morphological alterations in the WM of the cerebellum. Basketball involves complex motor skills, which require bimanual and eye-hand coordination for dribbling and shooting a ball. Thus, participation in a basketball competition at a high level of competition requires very high motor proficiency. These skills are central to basketball achievement and are therefore extensively and specifically trained. It seems that elite basketball athletes have highly developed motor skills, and such long-term extensive motor learning would evoke structural plasticity in the WM of vermian lobules VI–VII in the cerebellum. Such WM plasticity in this region of athletes may be crucial to support the increased functional demands for athletic achievement. Extensive motor learning may induce neural activity, which in turn may increase the number of axons, axon diameter, thickness of myelin, axon branching, axon trajectories, and myelination in this region. These microscopic changes might be reflected in the WM of this region in elite basketball athletes to improve the speed or synchrony of impulse transmission to support increased motor demands, including coordination and balance control [19, 20].

However, T1-weighted MRI has relatively low sensitivity, because MRI cannot reflect the structures of WM in the human brain as diffusion tensor imaging. Therefore, the findings of WM volume differences must be interpreted with caution. In addition, it is technically difficult to apply our MR images acquired from non-isometric voxels to a voxelbased morphometry technique based on statistical parametric mapping. Further studies are required to compare imaging data with the post-mortem histological measures that underlie these macroscopic finding in elite athletes, in order to more clearly establish the anatomical basis of these motor learning-induced morphological alterations in the human brain. In addition, the connectivity of the cerebellum with the red nucleus and cerebral cortex also requires further study to clarify this issue.

In conclusion, we demonstrated that elite basketball athletes have a larger WM volume of the vermian lobules VI and VII (the declive, folium, and tuber) in the cerebellum compared to control subjects, which might reflect a refinement of the connectivity in this region. This result suggests that long-term intensive motor learning evokes WM plasticity in this region of the cerebellum, and that the WM of this region plays a critical role in coordination based on motor learning. The findings of brain plasticity induced by motor learning could contribute to the development of neuro-rehabilitation programs.

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# References

- 1. Scholz J, Klein MC, Behrens TE, Johansen-Berg H. Training induces changes in white-matter architecture. Nat Neurosci 2009;12:1370-1.
- 2. Wang X, Casadio M, Weber KA 2nd, Mussa-Ivaldi FA, Parrish TB. White matter microstructure changes induced by motor skill learning utilizing a body machine interface. Neuroimage 2013; 88C:32-40.
- Landi SM, Baguear F, Della-Maggiore V. One week of motor adaptation induces structural changes in primary motor cortex that predict long-term memory one year later. J Neurosci 2011; 31:11808-13.
- 4. Taubert M, Draganski B, Anwander A, Müller K, Horstmann A, Villringer A, Ragert P. Dynamic properties of human brain structure: learning-related changes in cortical areas and associated fiber connections. J Neurosci 2010;30:11670-7.
- 5. Park IS, Lee KJ, Han JW, Lee NJ, Lee WT, Park KA, Rhyu IJ. Experience-dependent plasticity of cerebellar vermis in basketball players. Cerebellum 2009;8:334-9.
- Jacini WF, Cannonieri GC, Fernandes PT, Bonilha L, Cendes F, Li LM. Can exercise shape your brain? Cortical differences associated with judo practice. J Sci Med Sport 2009;12:688-90.
- 7. Jancke L, Koeneke S, Hoppe A, Rominger C, Hänggi J. The architecture of the golfer's brain. PLoS One 2009;4:e4785.
- Di X, Zhu S, Jin H, Wang P, Ye Z, Zhou K, Zhuo Y, Rao H. Altered resting brain function and structure in professional badminton players. Brain Connect 2012;2:225-33.
- 9. Hänggi J, Koeneke S, Bezzola L, Jäncke L. Structural neuroplasticity in the sensorimotor network of professional female ballet dancers. Hum Brain Mapp 2010;31:1196-206.
- Wang B, Fan Y, Lu M, Li S, Song Z, Peng X, Zhang R, Lin Q, He Y, Wang J, Huang R. Brain anatomical networks in world class gymnasts: a DTI tractography study. Neuroimage 2013;65:476-87.
- Johansen-Berg H, Della-Maggiore V, Behrens TE, Smith SM, Paus T. Integrity of white matter in the corpus callosum correlates with bimanual co-ordination skills. Neuroimage 2007;36 Suppl 2:T16-21.
- Ito M. Mechanisms of motor learning in the cerebellum. Brain Res 2000;886:237-45.
- Thach WT. What is the role of the cerebellum in motor learning and cognition? Trends Cogn Sci 1998;2:331-7.
- Quallo MM, Price CJ, Ueno K, Asamizuya T, Cheng K, Lemon RN, Iriki A. Gray and white matter changes associated with tooluse learning in macaque monkeys. Proc Natl Acad Sci U S A 2009;106:18379-84.
- 15. Abdul-Kareem IA, Stancak A, Parkes LM, Al-Ameen M, Alghamdi J, Aldhafeeri FM, Embleton K, Morris D, Sluming V. Plasticity of the superior and middle cerebellar peduncles in musicians revealed by quantitative analysis of volume and

number of streamlines based on diffusion tensor tractography. Cerebellum 2011;10:611-23.

- Tomassini V, Jbabdi S, Kincses ZT, Bosnell R, Douaud G, Pozzilli C, Matthews PM, Johansen-Berg H. Structural and functional bases for individual differences in motor learning. Hum Brain Mapp 2011;32:494-508.
- Roberts RE, Bain PG, Day BL, Husain M. Individual differences in expert motor coordination associated with white matter microstructure in the cerebellum. Cereb Cortex 2013;23:2282-92.
- Fields RD. Neuroscience. Change in the brain's white matter. Science 2010;330:768-9.
- Zatorre RJ, Fields RD, Johansen-Berg H. Plasticity in gray and white: neuroimaging changes in brain structure during learning. Nat Neurosci 2012;15:528-36.
- 20. Fields RD. White matter in learning, cognition and psychiatric disorders. Trends Neurosci 2008;31:361-70.
- 21. Juraska JM, Kopcik JR. Sex and environmental influences on the size and ultrastructure of the rat corpus callosum. Brain Res 1988;450:1-8.
- 22. Markham JA, Greenough WT. Experience-driven brain plasticity: beyond the synapse. Neuron Glia Biol 2004;1:351-63.
- 23. Markham JA, Herting MM, Luszpak AE, Juraska JM, Greenough WT. Myelination of the corpus callosum in male and female rats following complex environment housing during adulthood. Brain Res 2009;1288:9-17.
- 24. Sirevaag AM, Greenough WT. Differential rearing effects on rat visual cortex synapses. III. Neuronal and glial nuclei, boutons, dendrites, and capillaries. Brain Res 1987;424:320-32.
- 25. Szeligo F, Leblond CP. Response of the three main types of

glial cells of cortex and corpus callosum in rats handled during suckling or exposed to enriched, control and impoverished environments following weaning. J Comp Neurol 1977;172:247-63.

- 26. Zhao YY, Shi XY, Qiu X, Lu W, Yang S, Li C, Chen L, Zhang L, Cheng GH, Tang Y. Enriched environment increases the myelinated nerve fibers of aged rat corpus callosum. Anat Rec (Hoboken) 2012;295:999-1005.
- 27. Sanchez I, Hassinger L, Paskevich PA, Shine HD, Nixon RA. Oligodendroglia regulate the regional expansion of axon caliber and local accumulation of neurofilaments during development independently of myelin formation. J Neurosci 1996;16:5095-105.
- Als H, Duffy FH, McAnulty GB, Rivkin MJ, Vajapeyam S, Mulkern RV, Warfield SK, Huppi PS, Butler SC, Conneman N, Fischer C, Eichenwald EC. Early experience alters brain function and structure. Pediatrics 2004;113:846-57.
- 29. Hihara S, Notoya T, Tanaka M, Ichinose S, Ojima H, Obayashi S, Fujii N, Iriki A. Extension of corticocortical afferents into the anterior bank of the intraparietal sulcus by tool-use training in adult monkeys. Neuropsychologia 2006;44:2636-46.
- 30. Demerens C, Stankoff B, Logak M, Anglade P, Allinquant B, Couraud F, Zalc B, Lubetzki C. Induction of myelination in the central nervous system by electrical activity. Proc Natl Acad Sci U S A 1996;93:9887-92.
- Ishibashi T, Dakin KA, Stevens B, Lee PR, Kozlov SV, Stewart CL, Fields RD. Astrocytes promote myelination in response to electrical impulses. Neuron 2006;49:823-32.
- Hutchinson S, Lee LH, Gaab N, Schlaug G. Cerebellar volume of musicians. Cereb Cortex 2003;13:943-9.