

Morphometric Assessment of Corpus Callosum and Cerebral Hemispheres with Magnetic Resonance Imaging

Corpus Callosum ve Hemispherium Cerebri'lerin MRI ile Morfometrik Değerlendirmesi

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Objective: In this study, our purpose was to assess the relationship between handedness, footedness and the morphological differences of certain intracranial structures on MR images.

Material and Methods: 63 healthy male and 52 healthy female individuals were included in the study. In each subject, 16 measurements of intracranial structures were taken on MR images. Area of corpus callosum, also left and right cerebral hemispheres, length of CC, width of genu (r1), truncus (r2), isthmus (r3) and splenium of CC (r4) and width of corresponding cerebral hemisphere were measured.

Results: There was a statistically significant difference in R3 (width of the corresponding cerebral hemisphere to r3) distance between those using their right feet and those using their left feet. Angle of genu in right-handed people was greater in males than in females. Angle of genu in cases using the left foot was greater in females than in males for standing on one leg.

Conclusion: Morphometric assessment of CC with MR imaging related to handedness and footedness may be useful in demonstrating the relationship between callosal morphology, gender differences and extremity preference in neuroscience.

Key words: Corpus callosum; handedness; footedness; magnetic resonance imaging.

Amaç: Bu çalışmada amacımız, MR görüntülerinde çeşitli intrakranyal yapıların morfolojik farklılıkları ile el ve ayak tercih edilirliliği arasındaki ilişkiyi değerlendirmektir.

Gereç ve Yöntemler: Çalışma 63 erkek ve 52 kadın sağlıklı bireyi kapsıyordu. Her bir bireyde intrakranyal yapıların 16 ölçümü Manyetik Rezonans görüntülerinde yapıldı. Corpus callosum (CC), sol ve sağ hemispherium cerebri alanları, CC uzunluğu, CC'un genu (r1), truncus (r2), isthmus (r3) ve splenium (r4) kısımlarının genişliği ve bu kısımlara uyan hemispherium cerebri'lerin genişliğini ölçtük.

Bulgular: Sağ ile sol ayağını tercih edenler arasında R3 (r3'e uyan hemispherium cerebri'lerin genişliği) mesafesinde istatistiksel olarak anlamlı farklılık vardı. Sağ elini tercih edenlerde genu açısı erkeklerde kadınlardan daha büyüktü. Tek ayak üzerinde durma için sol ayağını kullanan vakalarda genu açısı kadınlarda erkeklerden daha büyüktü.

Sonuç: El ve ayak tercih edilirliliğine ilişkin MR görüntüleriyle CC'un morfometrik değerlendirmesi sinir biliminde CC'un morfolojisi, cinsiyet farklılıkları ve ekstremiter tercihi arasındaki ilişkiyi göstermede yararlı olabilir.

Anahtar sözcükler: Corpus callosum; el tercih edilirliliği; ayak tercih edilirliliği; manyetik rezonans görüntüleme.

INTRODUCTION

The corpus callosum (CC), composed of myelinated and unmyelinated axons, links the homotropic regions of the each cerebral hemisphere. Size variations in the CC presumably reflect differences in the size of axons that connect these regions.^[1]

The size of the CC corresponds to the number of the small and large myelinated fibres connecting the homologous cortical areas of the left and right hemispheres.^[2]

Investigations of effects of gender differences on the CC and cerebral hemispheres are important. Gender differences in gross corpus callosal neuroanatomy have been observed in several studies.^[3-7]

In recent studies, no consistent gender difference for callosal size has been observed. Additionally, the morphology of the CC may be influenced by handedness,^[8] and cerebral lateralization.^[2] The tasks were used in determining handedness (to write a letter legibly, to throw a ball to hit a target, hammering a nail, opening a door, erasing a blackboard, using an eraser on paper, combing the hair and cutting bread etc.) and footedness (kicking, hopping, jumping, first foot used when walking etc.) by some authors.^[9]

In this study, our purpose was to assess the handedness, footedness and the gender differences of the certain intracranial structures on MR images.

MATERIAL AND METHODS

This study was conducted on healthy males (n=63) and females (n=52). The age distribution of subjects was 10-84 years and the mean of age was 37.50±16.47. The subjects were divided into 3 groups. Group I consisted of those younger than 25 (n=33), Group II consisted of those whose age was between 26 and 40 (n=43), and Group III consisted of those older than 41 (n=39). The midsagittal and parasagittal MR images of 115 individuals were obtained from the Radiology Department of Faculty of Medicine, Gulhane Military Medical Academy. The age range was 10-84 years. The subjects did not have any neurological or psychiatric disorders and were not taking any drugs or chemical substances affecting the morphology of the brain. Additionally, they had no cerebral tumor or aneurysm. This study was performed between January 2007 and March 2008.

In each case, personal and family medical histories were noted. Subjects with major abnormalities like agenesis of CC were excluded. Also, any change in the handedness at any time, any injury to the limb in the past and incidence of left-handedness in the family or near relatives were recorded for each subject.

Hand and foot preferences were determined through 18 and 9 tasks respectively. The tasks used in determining handedness included opening a door, erasing a blackboard, using an eraser on paper, combing the hair,

carrying an object, lifting an object, holding scissors to cut paper, writing on a paper,^[8] cutting bread, throwing a stone, grasping an object, first hand used when swimming. Those for footedness were kicking, taking the first step forward and/or backward while walking, crossing one thigh over the other while sitting, foot tapping, riding on a two wheeler, standing on one leg,^[9] jumping, first foot used when stepping up and/or down, picking pebbles with toes. Furthermore, dominance in the eye and ear were ascertained by two tasks. Anyone preferring left hand for certain tasks (opening a door, erasing a blackboard, cutting bread, combing the hair, carrying an object, lifting an object, jumping, crossing one thigh over the other while sitting, foot tapping and standing on one leg) was assumed to be left-handed.

All sagittal images from lateral right to left through the hemispheres were examined in order to choose the image closest to the midline for each subject. Midsagittal images used in the analyses were those on which the pituitary, three lobules of the cerebellum, spinal cord, pons, and CC could be visualized.

On MR images, 16 measurements of intracranial structures were performed for each individual. Abbreviations used were as follows; r1, maximum dorsoventral width of genu of CC (the distance from the anterior-most point of the genu to the anterior-most point of the inner concavity of the anterior part of CC); R1, width of the corresponding cerebral hemisphere to r1; r2, minimum dorsoventral width of trunk of CC; R2, width of the corresponding cerebral hemisphere to r2; r3, minimum dorsoventral width of isthmus of CC; R3, width of the corresponding cerebral hemisphere to r3; r4, maximum dorsoventral width of splenium of CC; R4, width of the corresponding cerebral hemisphere to r4; L, anteroposterior length of CC (the distance from the anterior-most point of the genu to posterior-most point of the splenium) (Figure 1); X, angle of genu of CC (the anterior-most point of the genu was taken as the analysis point). Angle X was between the line originating from the analysis point and touching the dorsal-most point of the r2 and one tangential to top of the inner convexity of lower surface of the genu of CC); Y, Angle of splenium of CC (the posterior-most point of the splenium was taken as the analysis point. Angle Y was between the line originating from the analysis point and touching the dorsal-most point of the r3 and one tangential to top of the inner concavity of lower surface of the splenium of CC); IMMA, intracranial measurement of midsagittal area (inferior border of the intracranial area was defined with a line connecting the lip or the foramen magnum and the border of the prepontine cistern immediately posterior to the dens). Then we used the extra-cortical membrane, the optic chiasm and the pituitary stalk as a guide. These measurements are described by previous studies^[8,10] (Figure 2); ACC, midsagittal cross sectional area of CC (the CC boundary was circumscribed manu-

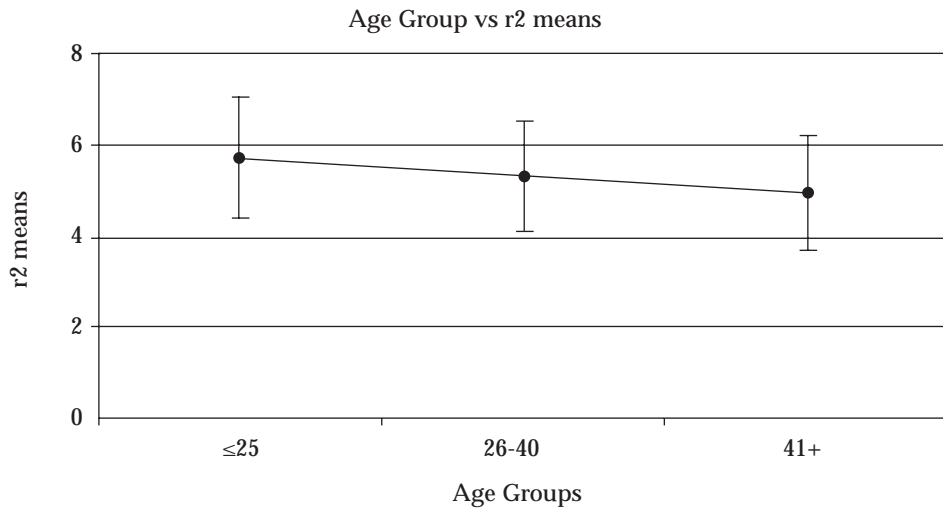


Figure 1. Relationship between age groups and means of r2 (means with SD).

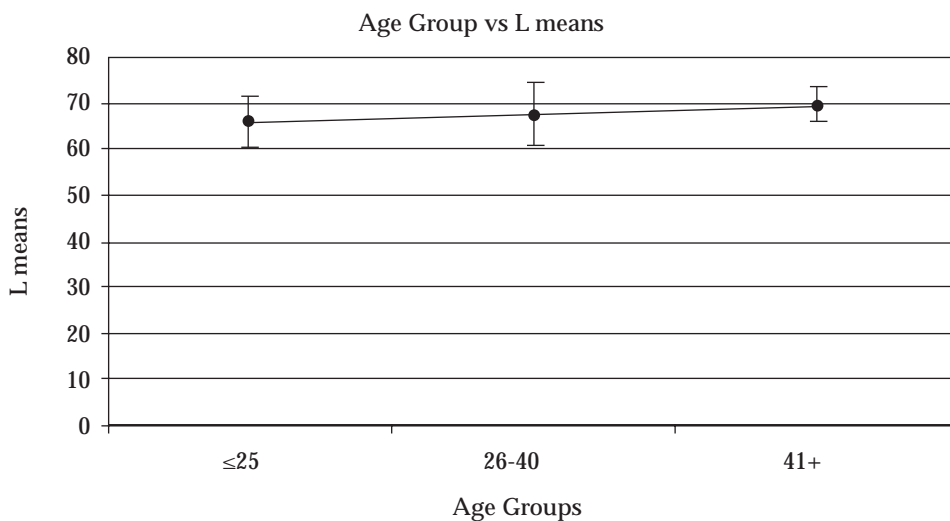


Figure 2. Relationship between age groups and means of L (means with SD).

ally with a trackball device); AH, area of hemisphere (midsagittal supratentorial intracranial area) (Figure 3); ARH, area of right hemisphere of brain (right parasagittal supratentorial intracranial area) and ALH, area of left hemisphere of brain (left parasagittal supratentorial intracranial area).

All patients were examined by 1.5 Tesla superconducting MR scanner (The New Intera Nova, Philips Medical Systems, Best, The Netherlands) using a standard quadrature head coil with version 9 software release. The system was equipped with magnetic field gradients capable of a maximum strength of 33 mT/m and maximum slew rate of 160 T/m/s. Axial and sagittal T1 weighted (583/15 ms, TR/TE, one excitation) spin-echo images were obtained by using 5 mm slice

thickness with a 1 mm intersection gap, 220x220 mm field of view (FOV) and 256x256 matrix size.

Statistical Analysis

MS-Excel 2003 and SPSS for Win. Ver. 15.00 (SPSS Inc., Chicago, IL., USA) and NCSS 2007 (Hintze J. (2006), NCSS, PASS, and GESS. NCSS, Kaysville, Utah, USA) software were used for statistical analysis. Distribution of variables was checked graphically and using the Shapiro-Wilk test. Descriptive statistics related to the variables are given as mean±standard deviation. Independent Sample t test (student's t test) was used for parametric variables and Mann-Whitney U test was used for non-parametric variables. For making comparisons according to age groups, ANOVA test was used for

parametric variables and Bonferroni post-hoc test was used for defining the source of difference. For non-parametric variables Kruskal-Wallis variance analysis was used and Bonferroni corrected Mann-Whitney U test was used for defining the source of difference. The relationship between age groups and r2 was analyzed using Pearson correlation and the relationship between age groups and L was analyzed using Spearman's (Rho) rank correlation. $p \leq 0.05$ was set as significant.

RESULTS

According to age groups, there were no statistically significant differences in r1, R1, R2, r3, R3, r4, R4, AH, ACC, IMMA, X, Y, ARH and ALH ($p > 0.05$). There was a statistically significant difference between Group I and Group III in r2 value ($p = 0.035$). There was a weak negative correlation (25.6%) between age and r2 value, which was statistically significant ($R^2 = -0.256$; $p = 0.006$) (Fig. 1). Similarly there was a statistically significant difference between Group I and Group III in L value ($Z = 2.623$; $p = 0.009$). There was a weak positive correlation (27%) between age and L value which was statistically significant ($Rho = 0.270$; $p = 0.004$) (Fig. 2).

Descriptive statistics of parameters are illustrated in Table 1. Gender comparative results in all parameters showed significant differences between males and females. Certain intracranial structures on MR images are

greater in males than in females; which is consistent with our results (Table 1). Reports of other researchers related to ACC are summarized in Table 2. In our study, ACC was $599.7 \pm 105 \text{ mm}^2$ in male, $573.6 \pm 129 \text{ mm}^2$ in female.

There was a statistically significant difference in R1 ($p = 0.004$), r2 ($p = 0.013$) and angle of genu ($p = 0.007$) distance between the genders (Fig. 3).

There was a statistically significant difference in the midsagittal area between right handed and left handed

Table 2. Values reported by researchers concerned with ACC (midsagittal)

ACC (mm ²)	Male Mean±SD	Female Mean±SD
... et al.	599.7±105	573.6±129
Allen et al. ^[14]	661±9	663±7
Kertesz et al. ^[16]	724*	716*
Clarke et al. ^[5]	540*	550*
Pujol et al. ^[19]	577±115	582±91
Hardan et al. ^[15]		671±100**
Mostofsky et al. ^[25]		627±73**
Oka et al. ^[18]		657±80**
Laissy et al. ^[17]		636**
Suganthi et al. ^[7]		700±84*

*Sd Not Presented
**No Gender Specified

Table 1. Descriptive statistics

Parameters	Male (n=63; 54.8%)			Female (n=52; 45.2%)			Test Value	P
	Min	Max	Mean±SD	Min	Max	Mean±SD		
r1 (mm)	5.80	14.50	10.59±1.73	4.80	15.30	10.34±2.18	t=0.695	0.489
R1 (mm)	28.80	53.70	39.57±5.01	27.20	48.00	35.95±4.82	Z=3.864	<0.001*
r2 (mm)	2.90	9.30	5.54±1.19	2.10	9.20	5.01±1.29	t=2.289	0.024*
R2 (mm)	33.30	51.20	42.26±3.79	31.80	50.00	40.73±3.51	t=2.231	0.028*
r3 (mm)	1.20	6.30	3.90±1.23	1.00	6.80	3.72±1.09	t=0.848	0.398
R3 (mm)	36.80	63.60	50.32±4.85	39.50	56.00	48.79±3.88	t=1.840	0.068
r4 (mm)	5.40	14.30	10.59±1.82	4.70	15.00	10.32±2.15	t=0.753	0.453
R4 (mm)	33.90	76.30	51.34±8.95	36.60	74.20	48.45±7.81	Z=1.899	0.058
L (mm)	56.60	81.60	67.73±4.69	38.50	84.00	67.49±6.73	Z=0.115	0.908
X°	29.70	116.10	74.06±19.99	34.50	112.80	69.61±16.37	t=1.290	0.200
Y°	63.60	146.50	104.92±18.05	70.00	141.30	103.68±16.75	t=0.378	0.706
IMMA (mm ²)	9418.60	15973.90	13896.93±1061.64	10533.60	15086.60	13106.20±814.64	Z=4.394	<0.001*
ACC (mm ²)	40.80	853.70	599.74±105.23	53.50	852.60	573.61±129.91	Z=0.753	0.451
AH (mm ²)	5419.20	10280.90	8895.89±805.47	6224.40	10198.20	8284.75±698.24	Z=4.355	<0.001*
ARH (mm ²)	6132.00	10666.80	8975.84±789.36	6304.30	9983.90	8402.55±725.30	t=4.020	<0.001*
ALH (mm ²)	6065.40	10643.40	8815.69±810.69	6803.20	9896.80	8225.13±636.28	t=4.276	<0.001*

*: Statistical significant

Table 1. r1 width of genu of CC, R1 width of the corresponding cerebral hemisphere to r1, r2 width of trunk of CC, R2 width of the corresponding cerebral hemisphere to r2, r3 width of isthmus of CC, R3 width of the corresponding cerebral hemisphere to r3, r4 width of splenium of CC, R4 width of the corresponding cerebral hemisphere to r4, L anteroposterior length of CC, X angle of genu of CC, Y angle of splenium of CC, ACC midsagittal cross sectional area of CC, ARH area of right hemisphere of brain, ALH area of left hemisphere of brain, IMMA intracranial measurement of midsagittal area of the brain.

subjects ($p=0.048$). Also, there was a statistically significant difference in $r3$ distance between those using their right hand and those using their left hand in cleaning the blackboard ($p=0.041$) (Fig. 4).

There was a statistically significant difference in $R3$ distance between those using their right feet and those using their left feet ($p=0.009$). Also, there was a statistically significant difference in $R1$ distance between those using their right feet and those using their left feet in jumping ($p=0.047$). Differences in other parameters related to gender and tasks are shown in Table 3.

Angle of genu in right-handed people was greater in males than in females (male, 79.34 ± 19.14 ; female, 64.38 ± 15.97). Angle of genu in cases using the left foot was greater in females than in males (male, 61.15 ± 19.45 ; female, 76.97 ± 12.11) for standing on one leg. In addition, angle of genu in cases using the left foot while using the right hand was greater in females than in males in our study (Fig. 5).

DISCUSSION

The CC is the largest fiber pathway of the brain in human and links the cerebral cortex of each hemisphere. The most conspicuous feature is the great commissure, the CC, a broad arched band in the floor of the central region of the longitudinal fissure. Its curved anterior part is the genu, continuous below with the rostrum and narrowing rapidly as it passes back to the upper end of the lamina terminalis. The genu continues above into the trunk, the main part of the commissure, which arches up and back to a thick, rounded posterior extremity, the splenium.^[11]

The findings regarding the morphometry of CC are important in the diagnosis of neurological diseases. It has been discussed that these measurements might be related to some diseases.^[12]

There are many studies where CC and cerebral hemisphere widths and areas have been measured on MR images.^[13-19]

A longer CC in males has been reported in two studies.^[7,20] However, most researchers have not identified sexual dimorphism in callosal length.^[14,21]

Several studies reported that the measurements of the anterior parts of the corpus callosum (rostrum, genu and anterior parts of the trunk) were decreased during normal aging.^[7,22-25] Takeda et al. reported that the widths of rostrum, body and splenium of the corpus callosum became thinner with age.^[25] Suganthy et al.^[7] reported that the length of the corpus callosum increased with age and regression equations for predicting age was derived from the length of the corpus callosum. In addition, they measured the average callosal length as 71.6 ± 4.7 mm. We found the average anteroposterior length of CC as

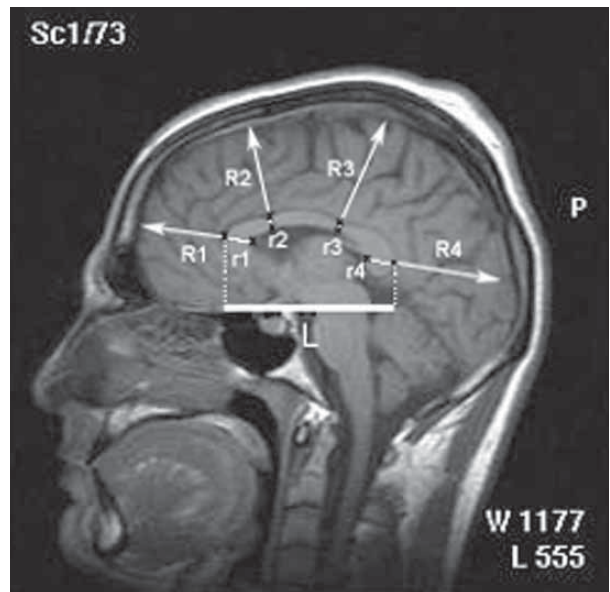


Figure 3. Mid-sagittal cross sectional image. Width of genu ($r1$); width of trunk ($r2$); width of isthmus ($r3$); width of splenium ($r3$), width of cerebral hemisphere of corresponding to $r1$, $r2$, $r3$, $r4$ ($R1$, $R2$, $R3$, $R4$), length of CC (L).



Figure 4. Mid-sagittal intracranial area of brain. Angle of genu (X); angle of splenium (Y); intracranial measurement of midsagittal area of brain (IMMA).

67.72 ± 4.69 mm in male, 67.48 ± 6.73 mm in female. We did not identify any differences in the length of CC between genders (Table 1).

Schmitt et al.^[20] found the area of CC as 782 ± 73 mm² and Peterson et al.^[26] as 785 ± 154 mm². These results were greater than that of other studies. Our results correspond to the results of other researchers in Table 2.

Denenberg et al.^[27] have evaluated the total area of CC. Regional widths were found to be sensitive to gender by handedness interactions in the anterior body, with right-handed females and left-handed males being larger. In the posterior body, males had wider callosa than females. A further analysis within the 'isthmus' region compared consistent and non-consistent right-handed males and females. Consistent right-handed males and both female groups had smaller callosa than non-consistent right-handed males.^[27]

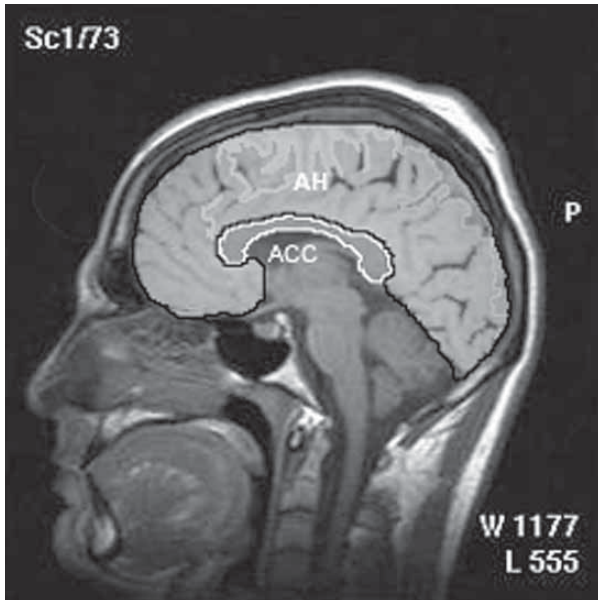


Figure 5. Mid-sagittal cross sectional area of corpus callosum. Midsagittal cross sectional area of CC (ACC); area of hemisphere (AH).

Mostofsky et al.^[28] have found the intracranial midsagittal area as $15900 \pm 1166 \text{ mm}^2$.^[13] Allen reported that values of the area of hemispherium (midsagittal area) were $9364 \pm 73 \text{ mm}^2$ in male, $8993 \pm 65 \text{ mm}^2$ in female. In our study, IMMA was $13896.93 \pm 1061.64 \text{ mm}^2$ in male, $13106.2 \pm 814.63 \text{ mm}^2$ in female and AH was $8895.89 \pm 805.47 \text{ mm}^2$ in male, $8284.74 \pm 698.24 \text{ mm}^2$ in female. The former results were consistent with ours.

Cowell et al.^[29] have assessed the area, axis length and widths of CC. They reported that emales did not attain maximum width until age 41-50 whereas males had peaked at 20 years and declined thereafter.

Although gender differences have been reported in the genu^[6], trunk^[30], splenium and the area of the CC in the literature^[12,21], in our study R1 and r2 were significantly different between the genders, being greater in the male (Table 1). Suganthy et al.^[7] have reported the width of splenium as $10.7 \pm 1.9 \text{ mm}$; in our study, r4 was $10.59 \pm 1.82 \text{ mm}$ in males and $10.32 \pm 2.15 \text{ mm}$ in females.

Oka et al.^[17] have reported gender differences in four specific angles of the CC. We found the angle of genu as $74.06 \pm 19.98^\circ$ in male, and $69.61 \pm 16.36^\circ$ in female. The average angle of genu in right-handed subjects was greater in males than in females. The average angle of genu among subjects using left foot for standing on one leg was greater in females than in males.

Dunham and Hopkins^[31] suggest that right-handed chimpanzees have a larger CC than left-handed chimpanzees. We did not find handedness differences for midsagittal cross sectional area of CC values in humans.

Table 3. Statistically significant differences in some parameters related to gender and tasks

Parameters	Groups	R1	r2	R2	r3	R3	R4	X
Gender	Male	▲	▲	⊗	⊗	⊗	⊗	▲
	Female	▼	▼	⊗	⊗	⊗	⊗	▼
Opening the door	Left	⊗	⊗	⊗	⊗	⊗	⊗	⊗
	Right	⊗	⊗	⊗	▲	⊗	⊗	⊗
Erasing a blackboard	Left	⊗	⊗	⊗	▼	⊗	⊗	⊗
	Right	▲	⊗	⊗	⊗	▲	⊗	⊗
Jumping	Left	▲	⊗	▲	⊗	⊗	⊗	⊗
	Right	▼	⊗	▼	⊗	▼	⊗	⊗
Cutting bread	Left	⊗	⊗	▲	⊗	⊗	⊗	⊗
	Right	⊗	⊗	▼	⊗	⊗	⊗	⊗
Combing the hair	Left	⊗	⊗	⊗	⊗	⊗	⊗	⊗
	Right	⊗	⊗	⊗	⊗	⊗	⊗	⊗
Carrying an object	Left	⊗	⊗	⊗	⊗	⊗	▲	⊗
	Right	⊗	⊗	⊗	⊗	⊗	▼	⊗
Lifting an object	Left	⊗	⊗	⊗	⊗	⊗	▲	⊗
	Right	⊗	⊗	⊗	⊗	⊗	▼	⊗

▲: Higher

▼: Lower

⊗: No differences

Witelson^[8] reported that the isthmus area was greater in left handed than right-handed subjects, whereas we did not find handedness differences for the width of isthmus of CC (r3) values.

Studies that have been performed to analyse the relationship between extremity preference and the structural differences of corpus callosum are cross-sectional. We suggest that the results of our cross-sectional study show that extremity preference may be a cause or a result of the structural differences of corpus callosum. Additionally, we consider that long term and ethically suitable cohort studies and visualisation techniques should be used to determine whether the structural differences of corpus callosum are a result or the cause of observed clinical signs.

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Conflict of Interest

No conflict of interest declared by the authors.

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