

**Kertas Asli/Original Article**

**Activation Characteristics of the Primary Motor (M1) and Supplementary Motor (SMA) Areas during Robust Unilateral Finger Tapping Task**  
[Ciri Pengaktifan bagi Kawasan Motor Primer (M1) dan Motor Tambahan (SMA) Semasa Tugas Menepik Jari Unilateral Secara Robus]

AHMAD NAZLIM YUSOFF, MAZLYFARINA MOHAMAD, KHAIRIAH ABDUL HAMID, AINI ISMAFAIRUS ABD HAMID, HANANI ABDUL MANAN & MOHD HARITH HASHIM

ABSTRACT

*This study investigated the functional specialisation characteristics of brain in multiple right-hand dominant subjects pertaining to the activation of the cerebral motor cortices evoked by unilateral finger tapping, especially in primary motor (M1) and supplementary motor (SMA) areas. This multiple-subject study used unilateral ( $UNI_{right}$  and  $UNI_{left}$ ) self-paced tapping of hand fingers to activate the M1 and SMA. Brain activation characteristics were analysed using statistical parametric mapping (SPM). Activation for  $UNI_{right}$  and  $UNI_{left}$  showed the involvement of contralateral and ipsilateral M1 and SMA. A larger activation area but with a lower percentage of signal change (PSC) were observed in the left M1 due to the control on  $UNI_{right}$  (4164 voxels at  $\alpha = 0.001$ ,  $PSC = 1.650$ ) as compared to the right M1 due to the control on  $UNI_{left}$  (2012 voxels at  $\alpha = 0.001$ ,  $PSC = 2.377$ ). This is due to the influence of the tapping rate effects which is greater than what could be produced by the average effects of the dominant and sub-dominant hands. The significantly higher PSC value observed in the right M1 ( $p < 0.05$ ) is due to a higher control demand used by the brain in coordinating the tapping of the sub-dominant fingers. The findings obtained from this study showed strong evidence of the existence of brain functional specialisation and could be used as baseline references in determining the most probable motor pathways in a sample of subjects.*

*Keywords: Finger tapping; random-effects analysis; statistical parametric mapping*

ABSTRAK

*Kajian ini menyelidiki ciri pengkhususan kefungsi otak bagi subjek berbilang dominan tangan kanan yang merujuk kepada pengaktifan pada korteks motor serebrum yang dicetus oleh tepikan jari secara unilateral terutama dalam kawasan motor primer (M1) dan motor tambahan (SMA). Kajian subjek berbilang ini menggunakan tepikan jari tangan secara unilateral ( $UNI_{kanan}$  dan  $UNI_{kiri}$ ) rentak sendiri untuk mengaktifkan M1 dan SMA. Ciri pengaktifan otak dianalisis menggunakan pemetaan statistik berparameter (SPM). Pengaktifan untuk  $UNI_{kanan}$  dan  $UNI_{kiri}$  menunjukkan penglibatan M1 dan SMA secara kontralateral dan ipsilateral. Kawasan pengaktifan yang besar tetapi dengan perubahan perubahan isyarat (PPI) yang kecil diperhatikan pada M1 kiri yang mengawal  $UNI_{kanan}$  (4164 voksel pada  $\alpha = 0.001$ ,  $PSC = 1.650$ ) berbanding M1 kanan yang mengawal  $UNI_{kiri}$  (2012 voksel pada  $\alpha = 0.001$ ,  $PSC = 2.377$ ). Ini adalah berpunca daripada pengaruh kesan kadar tepikan yang lebih besar daripada yang boleh dihasilkan oleh kesan purata bagi tangan dominan dan subdominan. Nilai PPI yang lebih tinggi secara bererti ( $p < 0.05$ ) yang diperhatikan pada M1 kanan adalah disebabkan oleh keperluan kawalan yang lebih tinggi yang digunakan oleh otak dalam mengkoordinasi tepikan jari sub-dominan. Penemuan yang diperolehi daripada kajian ini menunjukkan bukti kukuh kewujudan pengkhususan kefungsi otak dan boleh digunakan sebagai rujukan garis pangkal dalam menentukan laluan motor paling mungkin dalam suatu sampel subjek.*

*Kata kunci: Tepikan jari; analisis kesan rawak; pemetaan statistik berparameter*

INTRODUCTION

In spite of a vast number of research conducted in studying how uni- and bilateral motor action are coordinated by the brain (Grefkes et al. 2008; Kasess et al. 2008; Walsh et al. 2008), questions still arise about the exact mechanism underlying the existence of activation clusters in the contralateral as well as in the ipsilateral regions and their functional relationships. These are pertaining, in particular,

to the height and spatial extent of activation of the activated areas and their connectivity, not only in one hemisphere but also with the ones on the opposite hemisphere.

In a novel works on motor activation and network in humans, Walsh et al. (2008) reported that the dominant hemisphere is responsible in initiating the control over bilateral movement. They also discovered that bilateral activation is not the sum of the right and left unilateral activation from which it was later indicated that the left and right unimanual

movements differ significantly in terms of the activation of and connectivity between the areas involved.

It has been established that the primary motor area (M1) in the precentral gyrus (PCG) and the supplementary motor area (SMA) in the medial dorsal wall are involved in movement preparation and execution of motor action (Toga & Thompson 2000). The understanding of how these areas interact in normal people is important so that the study on plasticity or reorganisation of brain function in motor-impaired patients can be precisely conducted. Such decisions are of great practical relevance in diagnosing the function or wellness of motor areas for pre or post treatment or surgery.

Previous finger tapping studies (Lutz et al. 2005; Aramaki et al. 2006; Grefkes et al. 2008) relied on systematically constrained instruction visually or verbally given to the subjects. The externally triggered stimuli evoked responses not only in motor areas but also in areas related to vision and hearing, which in turn complicate the study of connectivity between motor areas. These were not considered in those studies. Furthermore, externally triggered methods are lacking in robustness and discount generalization. Therefore, in order to exclude areas not related to motor function and to impose robustness, this study are conducted in which the subjects are instructed to perform self-paced finger tapping at moderate tapping force and speed.

This study is a continuation of our previous work on a single subject (Ahmad Nazlim Yusoff et al. 2006c) and multiple subjects (Ahmad Nazlim Yusoff et al. 2010). In this study, the brain functional specialisation was investigated on multiple subjects with regards to the activation in the cerebral motor cortices evoked by finger tapping which was robustly done by the subjects. First, group analyses were conducted by means of random (RFX) effects analysis and inferences based on the group responses were made onto the whole subject. Secondly, conjunction analysis was performed to search for common activated areas among the subjects. Thirdly, the group's percentage of signal change in the ROIs, in particular the left and right M1 and SMA were computed. Finally, a conclusion is made with regards to the activation characteristics in the contralateral and ipsilateral regions.

## MATERIALS AND METHODS

Functional magnetic resonance imaging (fMRI) examinations were performed on 16 right-handed healthy male and female subjects. The subjects were conveniently sampled based on Desmond & Glover (2002). They suggested that for a liberal significant level of 0.05, about 12 subjects were required to achieve 80% power at the single voxel level for typical activation. In anticipating for non activated brain regions, 16 subjects were selected. The subjects were given informed consent and screening forms as required by the Medical Research Ethical Committee of the Universiti Kebangsaan Malaysia (UKM). The subjects were interviewed on their health condition prior to the scanning session and were confirmed to be healthy. Prior to the fMRI scans, the subjects' handedness is tested using

the Edinburgh handedness inventory (Oldfield 1971). All subjects were confirmed to be right-handed.

Functional magnetic resonance imaging (fMRI) scans were conducted in the Department of Radiology, Universiti Kebangsaan Malaysia Hospital. Functional images were acquired using a 1.5 tesla magnetic resonance imaging (MRI) system (Siemens Magnetom Vision VB33G) equipped with functional imaging option, echo planar imaging (EPI) capabilities and a radiofrequency (RF) head coil used for signal transmission and reception. The imaging parameters for the structural (T1) and functional (T2\*) scans have been described elsewhere (Ahmad Nazlim Yusoff et al. 2006c).

The subjects were instructed on how to perform the motor activation task and were allowed to practice prior to the scanning. The subjects had to press all four fingers against the thumb beginning with the thumb-index finger contact and proceeding to the other fingers in sequence which would then begin anew with contact between thumb and index finger. This study used a robust self-paced finger movement. The tapping of the fingers would approximately be two times in one second (using an intermediate force between too soft and too hard). A six-cycle active-rest paradigm which was alternately cued between active and rest was used with each cycle consists of 10 series of measurements during active state and 10 series of measurements during resting state. The tapping of the fingers were done unilaterally ( $UNI_{left}$  or  $UNI_{right}$ ), see Ahmad Nazlim Yusoff et al. (2006c) for details.

All the functional (T2\*-weighted) and structural (T1-weighted) images were sent to Universiti Kebangsaan Malaysia Hospital (HUKM) MedWeb and were later retrieved in the Functional Image Processing Laboratory (FIPL), Diagnostic Imaging & Radiotherapy Programme, Faculty of Allied Health Sciences, UKM Kuala Lumpur for further analyses. Image analyses were performed using a personal computer (PC) with a high processing speed and large data storage. The MATLAB 7.4 – R2006a (Mathworks Inc., Natick, MA, USA) and Statistical Parametric Mapping (SPM5) (Functional Imaging Laboratory, Wellcome Department of Imaging Neuroscience, Institute of Neurology, University College of London) software packages were used for that purposes. Activated voxels were identified by the general linear model (GLM) by estimating the parameters of the model and by deriving the appropriate test statistic ( $T$  statistic) at every voxel. Statistical inferences were finally obtained on the basis of SPM and the Gaussian random field theory (Brett et al. 2004; Friston 2004). The inferences were made using the  $T$ -statistic at uncorrected ( $\alpha = 0.001$ ) significant level, whereas for the analysis of conjunction, the significant level is taken at  $\alpha = 0.1$ . Steps taken in data analyses using SPM have been completely described in various similar studies (Ahmad Nazlim Yusoff et al. 2005, 2006a, 2006b & 2006c).

The region of interest (ROI) analyses were performed in order to compare the response of the brain due to lateralisation (left and right) and task ( $UNI_{right}$  and  $UNI_{left}$ ). The ROIs were the activation clusters obtained from the subjects' activation map defined using automated anatomical labelling (AAL) (Tzourio-Mazoyer et al. 2002)

and WFU Pick Atlas (Maldjian et al. 2003) at  $\alpha = 0.1$ . The two selected ROIs were bilateral precentral gyrus (PCG) and supplementary motor area (SMA). Small volume correction was performed within the predefined ROIs (Worsley et al. 1996). Group's fixed-effects (FFX) percentage of signal change (PSC) relative to the baseline for all ROIs was extracted from a 4-mm radius sphere with the peak coordinates as the centre using MarsBar toolbox for SPM (Matthew Brett et al. 2002).

## RESULTS

Demographical data for all the subjects are depicted in Table 1. The mean age and its standard deviation for the subjects was  $22.31 \pm 2.65$  years old. Four (25%) male and (75%) female subjects participated in this study.

TABLE 1. Demographical data for all subjects

Subject	Gender	Age	Race	Handedness
S1	Female	24	M	Right
S2	Male	23	C	Right
S3	Female	19	M	Right
S4	Female	22	M	Right
S5	Female	23	M	Right
S6	Male	24	M	Right
S7	Male	24	C	Right
S8	Female	26	M	Right
S9	Female	23	M	Right
S10	Female	19	M	Right
S11	Female	28	M	Right
S12	Female	19	M	Right
S13	Male	22	C	Right
S14	Female	20	C	Right
S15	Female	22	M	Right
S16	Female	19	C	Right

Figure 1 is the statistical parametric maps (SPMs) obtained from random-effects (RFX) analysis showing contralateral and ipsilateral brain activations due to (a)  $UNI_{right}$  and (b)  $UNI_{left}$ . The crossing of the hair-line indicates the point of maximum intensity which occurred at  $(-32, -22, 50)$  and  $(38, -20, 62)$  in the left and right hemispheres, respectively. In order to illustrate several ipsilateral region, Figure 1 was taken at  $\alpha = 0.01$ . Some RFX statistical data, MNI coordinates at the point of maximum intensity in each respective cluster and the anatomical areas in which the maxima in the brain activation due to  $UNI_{right}$  and  $UNI_{left}$  occur are summarised in Table 2.

For  $UNI_{right}$ , seven significant clusters survive a height threshold of uncorrected  $\alpha = 0.001$  and a spatial threshold of 50 voxels. This is due to the fact that the other clusters are believed to be generated by factors not included in the experimental paradigm such as aliased biorhythm and mild responses of the brain during the experiment. There is a total of 4164 activated voxels ( $t > 3.73$ ) in the main cluster which covers parts of the left post and precentral gyrii and left SMA. The eight highest peaks are at Talairach-MNI coordinates of  $(-32, -22, 50)$ ,  $(-42, -18, 52)$ ,  $(-42, -24, 52)$ ,  $(-26, -16, 64)$ ,  $(-36, -12, 64)$ ,  $(-42, -36, 46)$ ,  $(-46, -16, 48)$  and  $(-6, 6, 48)$ . The results indicate that 29.8% of the main cluster is in the left BA6 (27.8% activated), 9.7% of cluster is in the left BA2 (44.6% activated), 7.8% of cluster is in the left BA3b (50.9% activated) and 7.1% of cluster is in the left BA4p (52.5% activated).

For  $UNI_{left}$ , 5 significant clusters survive the uncorrected height threshold of  $\alpha = 0.001$ . The main activation cluster in the precentral gyrus consists of 5 maxima. Their Talairach-MNI coordinates are  $(38, -20, 62)$ ,  $(42, -24, 62)$ ,  $(36, -32, 60)$ ,  $(46, -16, 58)$  and  $(48, -22, 58)$ . A number of 2012 voxels are activated ( $t > 3.73$ ); 35.0% of cluster is in the right BA6 (16.0% activated), 13.5% of cluster is in the right BA1 (32.7% activated), 10.2% of cluster is in

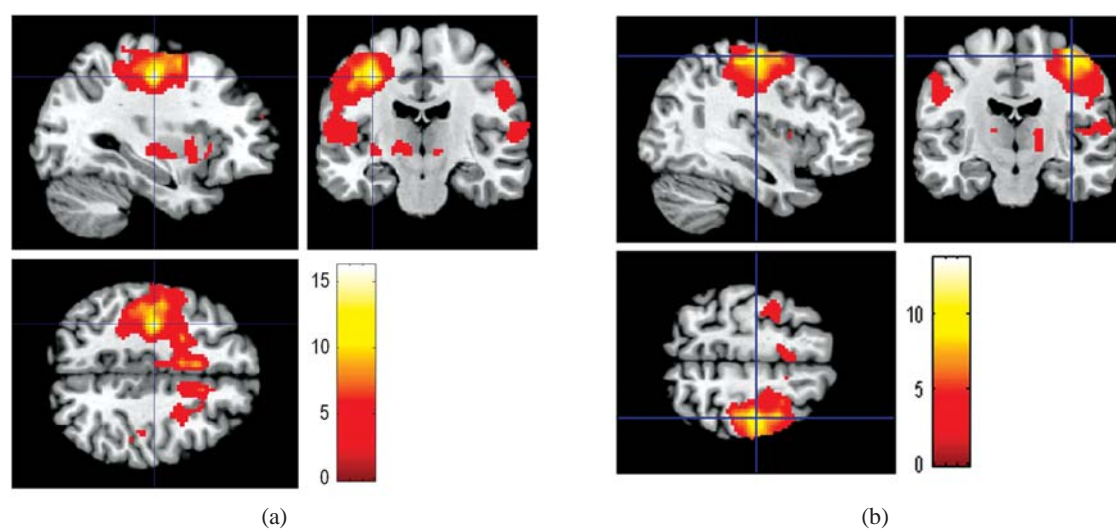


FIGURE 1. Statistical parametric maps (SPMs) obtained from random-effects (RFX) analysis ( $n = 16$ ,  $t > 2.60$ ,  $p < 0.01$  uncorrected) showing brain activation due to (a)  $UNI_{right}$  and (b)  $UNI_{left}$  overlaid onto structural brain images. Color codes represent increasing  $t$  value from red to white

TABLE 2. Statistical data, Tailarach-MNI coordinates ( $x, y, z$ ) and the respective anatomical areas obtained from  $UNI_{right}$  and  $UNI_{left}$  by means of RFX on 16 subjects at  $\alpha = 0.001$

Movement	Set-level		Cluster-level		Voxel-level		$x, y, z$ (mm)			Anatomical area
	$P_{uncorrected}$	Cluster	$p_{uncorrected}$	No. of activated voxel	$p_{uncorrected}$	$t$ -value				
$UNI_{right}$	< 0.001	1	< 0.001	4164	< 0.001	16.33	-32	-22	50	Left postcentral gyrus
					< 0.001	10.80	-42	-24	52	Left postcentral gyrus
					< 0.001	10.44	-26	-16	64	Left precentral gyrus
		2		< 0.001	9.02	60	10	16	Right precentral gyrus	
				< 0.001	6.82	56	6	32	Right precentral gyrus	
				< 0.001	5.28	60	6	4	Right Rolandic Operculum	
		3		< 0.001	7.85	10	10	52	Right supplementary motor area	
				< 0.001	6.91	8	-4	52	Right supplementary motor area	
				< 0.001	6.85	8	8	44	Right middle cingulate cortex	
		4		< 0.001	7.74	-50	-2	6	Left Rolandic operculum	
				< 0.001	4.80	-38	8	-6	Left insula lobe	
		5		< 0.001	7.54	38	-8	60	Right precentral gyrus	
				< 0.001	5.68	26	-8	60	Right superior frontal gyrus	
		6		< 0.001	7.29	60	-12	42	Right postcentral gyrus	
< 0.001	6.40		60	-14	34	Right postcentral gyrus				
7	< 0.001	5.56	54	-22	38	Right postcentral gyrus				
	0.023	5.37	-12	-20	4	Left thalamus				
$UNI_{left}$	0.760	1	< 0.001	2012	< 0.001	13.81	38	-20	62	Right precentral gyrus
			< 0.001		10.14	36	-32	60	Right postcentral gyrus	
			< 0.001		9.64	46	-16	58	Right precentral gyrus	
		2	0.091	73	< 0.001	6.70	-56	4	30	Left precentral gyrus
			< 0.001	4.14	-52	0	38	Left precentral gyrus		
		3	0.134	56	< 0.001	4.51	-4	4	50	Left supplementary motor area
		4	0.636	6	< 0.001	4.75	-30	4	56	Left middle frontal gyrus
		5	0.330	23	< 0.001	4.263	-40	-10	58	Left precentral gyrus

the BA3b (22.5% activated) and 8.7% of cluster is in the right BA4a (15.2% activated).

The results obtained from the analysis of conjunction on the present  $UNI_{right}$  and  $UNI_{left}$  datasets indicate that all subjects show common activation areas in the primary motor area. For  $UNI_{right}$ , 3 activation clusters are detected in the left postcentral gyrus and precentral gyrus. The main cluster which has 64 activated voxels ( $t > 1.28$ ) with the point of maximum activation at  $(-34, -22, 54)$ , shows that 55.1% of cluster is in the left BA4p (6.3% activated), 25.6% of cluster in the left BA4a (1.3% activated), 10.7% of cluster in the left BA6 (0.2% activated) and 8.6% of cluster is in the left BA3b (0.9% activated).

For  $UNI_{left}$ , the analysis of conjunction at significant level of  $\alpha = 0.1$ , reveals 1 cluster of activation which is in the right precentral gyrus. The cluster consists of 95 activated voxels ( $t > 1.28$ ) and has 5 maxima with the highest two at  $(36, -20, 62)$  and  $(40, -14, 56)$ . 89.9% of the cluster is in the right BA6 (1.9% activated), 8.2% is in the right BA4a (0.7% activated), 0.5% of cluster is in the right BA4p (0.1% activated) and 0.3% of cluster is in the right BA3b (0.1% activated).

Figure 2 is the plot of adjusted fitted responses and the corresponding error term calculated at the group peak

coordinates for all subjects in (a) left M1 during  $UNI_{right}$  and (b) right M1 during  $UNI_{left}$ . Subjects' responses at peak coordinates are well fitted into the model but with a large variability between subjects. It can be seen that the M1 peak intensity is roughly higher during  $UNI_{left}$  than during  $UNI_{right}$ . The percentage of change in signal intensity (PSC) that had occurred in the left and right M1 and SMA are tabulated in Table 3 for  $UNI_{right}$  and  $UNI_{left}$ . For  $UNI_{right}$ , M1(L) and SMA(L) show higher PSC values as compared to the ipsilateral M1(R) and SMA(R). Similarly, the PSC values for  $UNI_{left}$  are higher in M1(R) and SMA(R) as compared to the ipsilateral M1(L) and SMA(L). The PSC results for M1 are in good agreement with the fitted responses mentioned earlier. As opposed to number of activated voxels, the tapping of the left hand fingers generate higher signal change in M1(L) than in M1(R) during the tapping of right hand fingers. The effect is however incomparable in SMA.

## DISCUSSION

Based on Figure 1 and Table 2, it is quite interesting to see that the left side of the brain (triggered by the tapping of



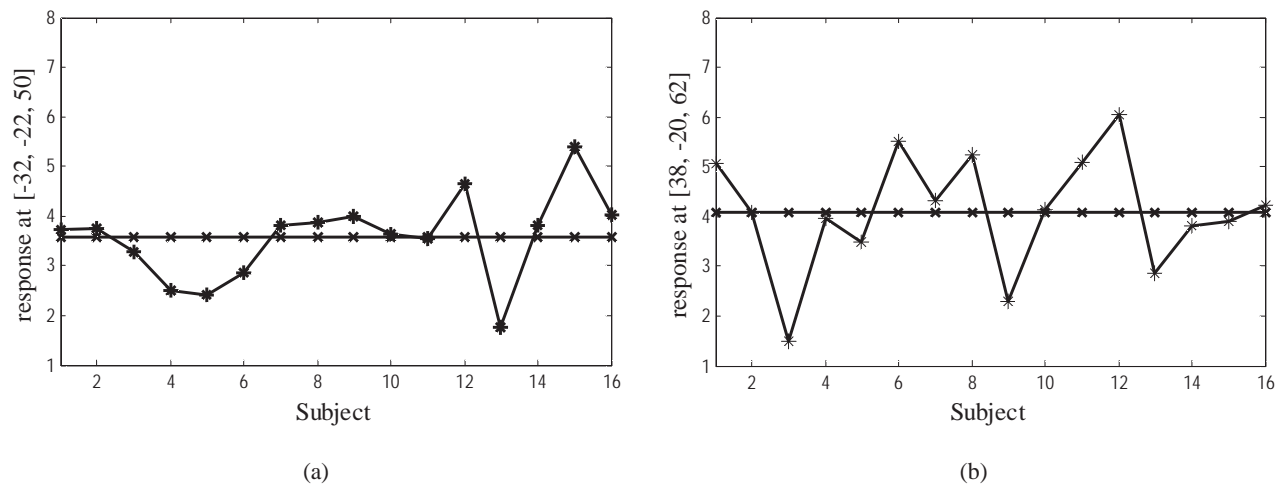


FIGURE 2. Adjusted fitted responses (x) and the error (\*) term calculated at the group peak coordinates for all subjects in (a) left M1 during  $UNI_{right}$  and (b) right M1 during  $UNI_{left}$ . The response at peak coordinates in the left M1 during  $UNI_{right}$  shows a higher intensity

TABLE 3. Percentage of signal change of the peak coordinates of the right and left M1 and SMA for  $UNI_{right}$  and  $UNI_{left}$

	Percentage of signal change/%			
	M1(L)	M1(R)	SMA(L)	SMA(R)
$UNI_{right}$	1.650 (-32 -20 52)	0.633 (36 -10 62)	0.860 (-6 -2 54)	0.670 (8 6 52)
$UNI_{left}$	0.713 (-36 -8 56)	2.377 (40 -20 66)	0.739 (-6 4 52)	0.793 (10 4 50)

the right hand fingers) shows a larger number of activated voxels and higher activation intensity as compared to the right side of the brain (triggered by the left-hand finger tapping), as opposed to our previous study on a single male subject (Ahmad Nazlim Yusoff et al. 2006c), despite the fact that all the subjects are right handed. This shows the reliability of multiple subject analyses in making inference over a population. Moreover, group results indicate the existence of ipsilaterality accompanying the expected contralaterality. The analyses conducted were focused on two anatomical regions that are known to be involved in controlling motor movement which are the primary motor cortex in the precentral gyrus (will be named as M1) and SMA which is also known to be involved in planning complex movements and in coordinating movements involving both hands (Walsh et al. 2008). The pre-motor cortex (PMC) is not included in the present study due to the inconsistency of the activation in the respective pre-motor area for all subjects, which resulted in lack of activation in group results. This could be due to the nature of task done by the subjects that does not involve the integration of sensory information which is one of the functions of PMC (Grefkes et al. 2008). M1 and SMA were found to be activated at different significant level in all participating subjects but the coordinates of the activation peaks differ by a few millimeters from subject to subject.

The typicality of the effects of the right and left unilateral tapping of fingers in all subjects was investigated using conjunction analysis. Conjunction analysis, as described by Friston (2004) provides a way to locate common features of functional anatomy between subjects under the same experimental condition. The results obtained from the analyses of conjunction on the present  $UNI_{right}$  and  $UNI_{left}$  datasets indicate that all subjects show common activation areas in precentral gyrus (M1). However, the SPM results generated at significant level of  $\alpha = 0.1$  indicate significant activation only at voxel level. Both the set and cluster level inferences about the activation clusters revealed insignificant brain activation.

In our previous study on a single right-handed male subject (Ahmad Nazlim Yusoff 2006c), the activated motor areas in the right hemisphere due to  $UNI_{left}$  showed a higher signal intensity and larger activation area as compared to that in the left hemisphere due to  $UNI_{right}$ . The findings obtained from our single subject study are in good agreement with a multiple subject fMRI study on unilateral and bilateral sequential movement in right-handers (Jäncke 1998). They found that the right hemisphere showed more activation than the left hemisphere in both unilateral and bilateral task at two tapping frequencies. They also concluded that faster movement rates will cause higher activation both in terms of signal intensity and number of activated voxel, the so called "rate effects." Their interpretations are that right-handers expend more effort to perform with their non-preferred hand. A stronger activation pattern in the right hemisphere is the result of trying to perform with a system that is slightly less competent with the implication that the more skilled and competent system will expend less effort and will therefore provide a weaker activation. As for the rate effects, they concluded that faster movement involves the recruitment of more motor units and will therefore activate a greater

number of voxels. Their findings were later reconfirmed in Lutz et al. (2005).

However, in this study and in separate study on seven right-handed female subjects (Ahmad Nazlim Yusoff et al. 2010), the average responses obtained from FFX and RFX indicate higher height (signal intensity) and spatial (activation area) extent of activation in the left hemisphere for both unilateral and bilateral types of finger tapping. As mentioned earlier, this study used a robust self-paced finger tapping. Prior to the fMRI scan, the subjects were told that they need to tap their fingers two times in one second using an intermediate force between too soft and too hard. However, since all the subjects are right-hand dominant, there would be a tendency for the subjects to tap their preferred hand fingers faster than their non-preferred hand fingers, resulting in the rate effects. Based on the interpretation given above, it seemed that the influence of the rate effects is greater than the effects that would be produced by the average effects of the dominant and sub-dominant hand, hence greater activation in the left hemisphere. A larger activation area could also be due to the tendency of these right-handers to press their fingers harder against the thumb using their dominant hand fingers, whereby a larger force will activate a larger area with higher intensity. Interestingly, in contrast to the spatial extent of activation, the PSC for M1 obtained in this study is higher in the right hemisphere (due to  $UNI_{left}$ ) as compared to the PSC measured in the left hemisphere (due to  $UNI_{right}$ ), (Table 3). This finding is in contrast to the number of activated voxels which is higher for  $UNI_{right}$  as compared to  $UNI_{left}$ . PSC is defined as the relative signal change within a cytoarchitectonic area evoked by the different experimental conditions, which reflects the involvement of that particular area in a specific task (Eickhoff et al. 2005). It is simply the ratio between the condition-specific signal change and the mean signal during the session. In relation to the discussion above, it can be assumed that tapping rate does not influence the height extent of activation as it does on the spatial extent of activation. As a result, the higher PSC observed in the right hemisphere is due only to a higher control demand used by the brain in coordinating the tapping of the sub-dominant hand fingers.

The results depicted in Figure 1, Table 2 and Table 3 clearly revealed significant activated areas in the opposite hemisphere to the contralateral hemisphere. For  $UNI_{right}$ , ipsilateral activation occurs in the right postcentral gyrus, right Rolandic operculum, right precentral gyrus, right middle frontal gyrus, right superior frontal gyrus, right SMA and right middle cingulate gyrus. For  $UNI_{left}$ , the ipsilateral areas are left precentral gyrus, left SMA and left middle frontal gyrus. The existence of ipsilateral activation in motor cortex has been widely reported and discussed (Grefkes et al. 2008; Walsh et al. 2008; Newton et al. 2005). It shows evidence of involvement of ipsilateral areas in coordinating motor movement. One of the observed effects related to ipsilateral activation is inhibition whereby increased neuronal activation in motor area of one hemisphere

suppresses neuronal activity of the same area in the opposite hemisphere. Inhibitory has been shown to be either in terms of activated volume or percentage of signal change (Newton et al. 2005). Inhibition is not observable in this study since tapping style is kept constant. However, as can be seen from Figure 1 and Table 3, ipsilaterality did occur in both M1 and SMA and the effects are asymmetrical and these shows possible evidence of inhibitory of activation in the ipsilateral areas.

## CONCLUSION

The results obtained from this multiple-subject study on right-handed male and female subjects showed that the observed brain activation for  $UNI_{right}$  and  $UNI_{left}$  fulfill contralaterality behavior of motor coordination. Brain activations are also presented in the ipsilateral regions indicating important roles of brain regions that are located on the same side of movement. Dominant hand has been found to produce stronger tapping rate effects (larger activation area) for this group of right handers as compared to subdominant hand and the influence is greater than what would be produced by the average effects of the dominant and sub-dominant hand. However, the higher percentage of signal change observed in the right M1 that controls  $UNI_{left}$  is due to a higher control demand used by the brain in coordinating the tapping of the sub-dominant hand fingers.

## ACKNOWLEDGEMENT

The authors would like to thank Sa'don Samian, the MRI Technologist of the Universiti Kebangsaan Malaysia Hospital (HUKM), for the assistance in the scanning and the Department of Radiology, Universiti Kebangsaan Malaysia Hospital for the permission to use the MRI scanner. The authors were also indebted to Professor Karl J. Friston and the functional imaging group of the University College of London for valuable discussions on experimental methods and data analyses. This work is supported by the research grants IRPA 09-02-02-0119EA296, the Ministry of Science, Technology and Innovation of Malaysia and and UKM-GUP-SK-07-20-205, Universiti Kebangsaan Malaysia.

## REFERENCES

- Ahmad Nazlim Yusoff, Mohd Harith Hashim, Mohd Mahadir Ayob & Iskandar Kassim. 2005. Pengimejan resonans magnet kefungsiian: Pemerolehan, analisis dan pentafsiran data (Functional magnetic resonance imaging: Data acquisition, analyses and interpretation). *Mal. J. Health Sci.* 3(2): 19-37.
- Ahmad Nazlim Yusoff, Mohd Harith Hashim, Mohd Mahadir Ayob & Iskandar Kassim. 2006a. Analisis data pengimejan resonans magnet kefungsiian: Prapemprosesan ruang menggunakan kaedah pemetaan statistik berparameter (Functional magnetic resonance data analyses: Spatial pre

- processing using statistical parametric mapping method). *Mal. J. Health Sci.* 4(1): 21-36.
- Ahmad Nazlim Yusoff, Mohd Harith Hashim, Mohd Mahadir Ayob & Iskandar Kassim. 2006b. Pengaktifan otak akibat gerakan jari bagi subjek dominan tangan kanan dan kiri (Brain activation evoked by finger movement for right- and left-hand dominant subjects). *Mal. J. Health Sci.* 4(2): 63-83.
- Ahmad Nazlim Yusoff, Mohd Harith Hashim, Mohd Mahadir Ayob, Iskandar Kassim, Nur Hartini Mohd Taib & Wan Ahmad Kamil Wan Abdullah. 2006c. Functional specialisation and connectivity in cerebral motor cortices: A single subject study using fMRI and Statistical Parametric Mapping. *Mal. J. Med. Health Sci.* 2(2): 37-60.
- Ahmad Nazlim Yusoff, Mazlyfarina Mohamad, Aini Ismafairus Abd Hamid, Wan Ahmad Kamil Wan Abdullah, Mohd Harith Hashim & Nurul Zafirah Zulkifli. 2010. Functional Specialisation and Effective Connectivity in Cerebral Motor Cortices: An fMRI Study on Seven Right Handed Female Subjects. *Mal. J. Med. Health Sci.* 6(2): 71-92.
- Aramaki, Y., Honda, M. & Sadato, N. 2006. Suppression of the non-dominant motor cortex during bimanual symmetric finger movement: A functional magnetic resonance imaging study. *Neuroscience* 141: 2147-2153.
- Brett, M., Penny, W. & Kiebel, S. 2004. An introduction to random field theory. In *Human Brain Function*, edited by R.S.J. Frackowiak, K.J. Friston, C.D. Frith, R.J. Dolan, C.J. Price, S. Zeki, J. Ashburner and W.D. Penny by Amsterdam: Elsevier Academic Press.
- Desmond, J.E. & Glover, G.H. 2002. Estimating sample size in functional MRI (fMRI) neuroimaging studies: Statistical power analyses. *J. Neurosci. Methods* 118: 115-128.
- Eickhoff, S.B., Stephan, K.E., Mohlberg, H., Grefkes, C., Fink, G.R., Amunts, K. & Zilles, K. 2005. A new SPM toolbox for combining probabilistic cytoarchitectonic maps and functional imaging data. *NeuroImage* 25: 1325-1335.
- Friston, K.J. 2004. Experimental design and statistical parametric mapping. In: *Human Brain Function*, edited by Frackowiak R.S.J., Friston, K.J., Frith, C.D., Dolan, R.J., Price, C.J., Zeki, S., Ashburner, J. & Penny, W.D. Amsterdam: Elsevier Academic Press.
- Grefkes, C., Eickhoff, S.B., Nowak, D.A., Dafotakis, M. & Fink, G.R. 2008. Dynamic intra- and interhemispheric interactions during unilateral and bilateral hand movements assessed with fMRI and DCM. *NeuroImage* 41: 1382-1394.
- Jäncke, L., Peters, M., Schlaug, G., Posse, S., Steinmetz, H. & Müller-Gärtner, H. -W. 1998. Differential magnetic resonance signal change in human sensorimotor cortex to finger movements of different rate of the dominant and subdominant hand. *Cogn. Brain. Res.* 6: 279-284.
- Kasess, C. H., Windischberger, C., Cunnington, R., Lanzenberger, R., Pezawas, L. & Moser, E. 2008. The suppressive influence of SMA on M1 in motor imagery revealed by fMRI and dynamic causal modeling. *NeuroImage* 40: 828-837.
- Lutz, K., Koeneke, S., Wüstenberg, T. & Jäncke, L. 2005. Assymetry of cortical activation during maximum and convenient tapping speed. *Neuroscience lett.* 373: 61-66.
- Maldjian, J.A., Laurienti, P.J., Kraft, R.A. & Burdette, J.H. 2003. An automated method for neuroanatomic and cytoarchitectonic atlas-based interrogation of fMRI data sets. *Neuroimage* 19(3): 1233-1239.
- Matthew Brett, Jean-Luc Anton, Romain Valabregue, Jean-Baptiste Poline. 2002. Region of interest analysis using an SPM toolbox. Proceedings of the 8th International Conference on Functional Mapping of the Human Brain; 2002 Jun 2-6: Sendai Japan. Available in CD-ROM in *NeuroImage*, 16(2).
- Newton, J. M., Sunderland, A. & Gowland, P.A. 2005. fMRI signal decrease in ipsilateral primary motor cortex during unilateral hand movements are related to duration and side of movement. *NeuroImage* 24: 1080-1087.
- Oldfield, R. 1971. The assessment and analysis of handedness: the Edinburgh inventory. *Neuropsychologia* 9: 97-113.
- Tzourio-Mazoyer, N., Landeau, B., Papathanassiou, D., Crivello, F., Etard, O., Delcroix, N., Mazoyer, B. & Joliot, M. 2002. Automated anatomical labeling of activations in SPM using a macroscopic anatomical parcellation of the MNI MRI single-subject brain. *Neuroimage* 15: 273-289.
- Toga, A.W. & Thompson, P.M. 2000. An introduction to maps and atlases of the brain. In *Brain Mapping: The Systems*, edited by A.W. Toga and J.C. Mazziotta. San Diego, U.S.A.: Academic Press.
- Walsh, R.R., Small, S.L., Chen, E.E. & Solodkin, A. 2008. Network activation during bimanual movements in humans. *NeuroImage* 43: 540-553.
- Worsley, K., Marrett, S., Neelin, P., Vandal, A., Friston, K. & Evans, A. 1996. A unified statistical approach for determining significant signals in images of cerebral activation. *Human Brain Mapping* 4: 58-73.

Ahmad Nazlim Yusoff  
 Mazlyfarina Mohamad  
 Khairiah Abdul Hamid  
 Aini Ismafairus Abd Hamid  
 Mohd Harith Hashim  
 Functional Image Processing Laboratory (FIPL)  
 Diagnostic Imaging & Radiotherapy Program  
 Faculty of Allied Health Sciences  
 Universiti Kebangsaan Malaysia  
 Jalan Raja Muda Abdul Aziz  
 50300 Kuala Lumpur

Hanani Abdul Manan  
 Medical Imaging Program  
 Masterskill University College of Health Sciences  
 43000 Cheras, Selangor

Corresponding author: Ahmad Nazlim Yusoff  
 Email address: nazlim@medic.ukm.my  
 Tel: +603-92897257; Fax: +603-92897915

Received: October 2009  
 Accepted for publication: May 2010