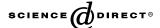


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Cerebral lateralization and general intelligence: Gender differences in a transcranial Doppler study

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Abstract

The present study evaluated cerebral lateralization during Raven's progressive matrices (RPM) paradigm in female and male subjects. Bilateral simultaneous transcranial Doppler (TCD) ultrasound was used to measure mean blood flow velocities (MBFV) in the right and left middle cerebral arteries (MCAs) in 24 (15 females and 9 males) right-handed normal subjects. The female subjects used a left hemisphere strategy, while males used a right hemisphere strategy to successfully solve RPM tasks. This implies that general intelligence is associated with neural systems within one hemisphere that are accessible to a variety of cognitive processes. © 2004 Elsevier Inc. All rights reserved.

Keywords: Cognition; Language; Cerebral blood flow; Ultrasound; Brain; Sex; Psychological tests; Laterality

1. Introduction

The determinants of human performance have been a subject of intense debate for over a century. Spearman in 1904, put forward the idea that measures of performance or success in diverse cognitive tests show a pattern of almost universal positive correlation (Jensen, 1987; Spearman, 1904, 1923, 1927). He postulated the hypothesis of a general intelligence or g factor making some contribution to success in diverse forms of cognitive activity. Subsequently, two contrasting postulates on cortical localization of intelligence were put forward. The first proposes that, processing of any task receives contribution from a large set of components or information-processing functions (Lashley, 1921, 1929; Satz, Strauss, & Whitaker, 1990; Thomson, 1951; Thurstone, 1938, 1940). Therefore, any two tasks are likely to share at least some components, giving rise to universal posi-

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tive correlation. This hypothesis assumes that, general ability is diffusely represented on the whole cortex, and injury to any region of the brain produces an intellectual decrement. Conversely, the second, so-called hierarchic hypothesis postulates that, in the course of evolution some area of the brain has gained a dominant role in sustaining general intelligence (Basso, De Renzi, Faglioni, Scotti, & Spinnler, 1973).

Snow, Kyllonen, and Marshalek (1984) have classified intelligence tests by task complexity, into tests at the center and tests at the periphery. Psychometric tests including RPM (Raven, 1938) and other complex reasoning tests were classified as tests at the center, while simpler tests were placed at the periphery. Applying this construct, irrespective of the views held by both hypotheses, the centrality of the RPM emerges in either case.

Furthermore, it has been suggested that RPM is a good test of intelligence, and should account for a great deal of the reasoning in other tests (Carpenter, Just, & Shell, 1990). It has been suggested that the neural substrate for intelligence lies within prefrontal cortex (Prabhakaran, Smith, Desmond, Glover, & Gabrieli, 1997), and

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post-rolandic structures (Basso et al., 1973). Recently, positron emission tomography studies have shown selective recruitment of lateral prefrontal cortex in one or both hemispheres (Duncan et al., 2000). Studies of cerebral lateralization of intelligence may implicate one hemisphere, or both, if general abilities were diffusely represented.

Cerebral lateralization could be studied using TCD (Bulla-Hellweg, Vollmer, Götzen, Skreczek, & Hartje, 1996; Cupini et al., 1996; Droste, Harders, & Rastogi, 1989; Evers, Dannert, Rodding, Rotter, & Ringelstein, 1999; Hartje, Ringelstein, Kistinger, Fabianek, & Willmes, 1994; Kelley et al., 1992; Markus & Boland, 1992; Njemanze, 1991, 1996, 2002, 2004; Njemanze, Gomez, & Horenstein, 1992; Rihs, Gutbrod, Steiger, Sturzenegger, & Mattle, 1995; Silvestrini, Troisi, Matteis, Razzano, & Caltagirone, 1994; Varnadore, Roberts, & McKinney, 1997; Vingerhoets & Stroobant, 1999). Studies with TCD have been cross-validated by functional MRI (Schmidt et al., 1999), and reproducibility assessed (Knecht et al., 1998).

The purpose of the present study is to evaluate cerebral lateralization during RPM tasks in females and males. It is postulated that general intelligence is associated with neural systems represented in one hemisphere that are accessible to a variety of cognitive processes. Consequently, unilateral representation of general intelligence will cause MBFV changes in ipsilateral MCA during performance of intelligence tasks. More specifically, the present study evaluates MBFV changes in both MCAs during performance of RPM tasks in both female and male subjects.

2. Materials and methods

2.1. Subjects

The studied group included 24 normal volunteers, 15 female and 9 male. Mean age was 25.08 ± 3.16 years (range 21–33); females (25.47 ± 3.5 years) did not differ from males (24.44 ± 2.5 years). All were right-handed as determined using the Edinburgh handedness inventory (mean laterality quotient was 100) (Oldfield, 1971). Subjects had no history of neurological or cardiovascular or respiratory diseases. None was under medication, including contraceptive pills for females or recreational drugs. All refrained from ingesting caffeine at least 24h prior to the study. The years of schooling ranged from 16 to 18. All subjects signed informed consent for the study. The Institutional Ethical Committee approved the study protocol.

2.2. Normal resting baseline measurements

Baseline data acquisition were made with subject in supine horizontal position with head inclined up at 30°. The

eyes were fixated on a cube target, projected onto a screen, placed 80 cm (from the nasal ridge) in front of the subject. The ears were partially closed with earplugs that also provided the base of the probe holder. All environmental noise including sound from the TCD instrument was excluded, and environmental luminance was kept constant for all participants. Measurements comprising a 60 s continuous train of velocity waveform envelopes were recorded with subject mute, still, and attention focused on the cube with no mental or manual task to perform.

2.3. Raven's progressive matrices test

Eight successive black and white slides of RPM tasks were arranged in increasing levels of difficulty and sequentially projected on a screen. Subjects were required to determine which of the 6–8 options best completes the matrix, so that the inter-relational rules among the elements between the row and columns were met. No verbal answers were required, instead subjects were asked to indicate their choice of option by slightly raising the corresponding finger during the inter-stimulus interval. The fingers were numbered sequentially from 1 to 8, starting with the right thumb as the first option to the left middle finger as option eight. An observer recorded the option signed by the subject during the 2s inter-stimulus interval, also used for task slide transition. All responses were transformed into Wrong or Correct ANSWER. Measurements comprising a 60s continuous train of velocity waveform envelopes were recorded during each task with subject mute, still and attention focused on the task.

2.4. Transcranial Doppler measurement

Transcranial Doppler scanning was performed by a method similar to that previously described elsewhere (Njemanze, 1991, 1996, 2002), using a bilateral simultaneous TCD instrument (Multi-Dop T, DWL, Sipplingen, Germany). MBFV in both MCAs was sampled in supine horizontal position using two 2 MHz transducers, with probes fixated on both temples using a probe holder—LAM-Rack (DWL, Sipplingen, Germany). The Doppler signals and a continuous train of flow velocity envelops were obtained at a depth of 50mm from the probe surface with the same gain and power settings for each subject. MBFV was recorded and averaged in 10s segments for each condition. Subjects were instructed to remain mute and motionless, and the probe-to-vessel angle remained constant throughout the data acquisition.

2.5. Other monitoring tests

Electrocardiographic monitoring and respiratory activity were recorded during experimental runs for control of any effects. Self-perceived anxiety levels by stan-

dardized questionnaire were recorded before and after data acquisition. Data acquisition were obtained under normal resting oscillations of these parameters. Operator monitoring of subjects' eye, voluntary, and involuntary movements provided a measure of control of these potential sources of artifacts. Motion-related artifacts causing deleterious interferences were marked and deleted from the analysis.

2.6. Calculations

Hemispheric advantage was assessed using laterality index (LI') expressed as LI' = (Right MBFV – Left MBFV /Right MBFV + Left MBFV) × 100. The hemisphere advantage was given by

LI = LI' paradigm -LI' baseline.

Positive LI suggests either a right lateralization or hemisphere advantage, while negative LI shows left lateralization. Zero LI may suggest no change from baseline or equal bilateral activation.

2.7. Methodological considerations

TCD measures MBFV in major intracranial vessels, which has been shown to correlate with blood flow estimates (Taylor, Short, Walker, & Traystman, 1990). TCD would characterize hemispheric differences in MBFV, but not changes within specific regions of the MCA territory. For example, if areas in the frontal, parietal, and temporal lobes were all crucial for intelligence (as measured by the RPM), activating any one area in the left or right hemisphere would change MBFV in the MCA main stem, and hence lateralization. The reason of course is that, the MCA supplies all these areas. However, the present study focuses on hemispheric advantage, therefore, the application of TCD is most appropriate. The measured MBFV in the main stem of the MCA could be considered as an averaged positive correlate of flow changes related to stimulus effect, suggesting a close correlation with neuronal activity. Moreover, cerebral blood flow velocity changes were elicited by neuronal activity observed during simultaneous TCD and video EEG recordings (Njemanze, Mirza, Rosenfeld, & Njemanze, 1992). Others have suggested a similar relationship among cerebral blood flow (CBF), metabolism and neuronal activity (Lueck et al., 1989; Petersen, Fox, Snyder, & Raichle, 1990; Reivich & Alavi, 1985). The MCA territory supplies about 80% of blood received by the cerebral hemispheres (Toole, 1984). In relation to site-specific functional localization, TCD could be considered a good test in series with other imaging modalities such as functional magnetic resonance imaging (fMRI) (Schmidt et al., 1999).

Potential sources of motion artifacts were handled by exclusion and cancelling out effects. In pre-experimental

control test runs, subtle indication with the finger did not cause a detectable rise in blood flow velocity in both MCAs. However, in the present study, recordings during inter-stimulus interval were marked and removed from the analysis. For male as well as female subjects, the use of right fingers for response was similar to the use of left (p > .05). This may suggest that any side-to-side effects of even 'internally planned' or mentally generated movements on data acquisition may have cancelled out. This assumes that the side of mentally generated movement is the same for ANSWER. A sample RPM task was administered in a practice session to eliminate ambiguity in instructions and reduced any possible role of anxiety due to novelty of task. No subject was aware that RPM measured general intelligence. However, all were motivated to solve the tasks to the best of their abilities.

Prior studies have shown that LI computations were independent of partial pressure of end-tidal carbon-dioxide, while the latter affects the sum of right and left MBFV (R + L) (Njemanze, 1991, 1996; Rumsey et al., 1987). The overall accuracy of R + L determines the validity of the measurements of hemispheric advantage in response to a task. It has been argued in the past that a right hemisphere competence may be revealed in non-response conditions (Milberg, Whitman, Rourke, & Glaros, 1981; Segalowitz & Cohen, 1989) due to absence of motor activity. The stimuli were designed to lack verbalizable features and excluded the possibility of interference from motor activity, by constrained control of subjects' response, until after each measurement.

2.8. Statistical analysis

Results were given as mean \pm SD where applicable. The LI scores were analyzed using multivariate analysis of variance (MANOVA). When applicable, group means were compared using t test. All statistical calculations were done using a statistical software package (Statistica, StatSoft, OK, USA).

3. Results

The data obtained with means \pm SD and Scheffé test probabilities for post hoc analysis for the different groups are shown in Table 1. The mean *Correct* score was $50.1 \pm 14\%$ for all subjects, and the mean for female subjects did not differ from male (p > .05). The design was a two-way MANOVA comprising: two levels of ANSWER (*Wrong* and *Correct*) and two levels of GENDER (*Female* and *Male*). There was a main effect of GENDER [F(1,1148) = 26.4, p < .0000003, MSe = 87.6]. There was no main effect of ANSWER (p > .05). There was a GENDER × ANSWER interaction [F(1,1148) = 3.8, p < .05]. As shown in Fig. 1, for *Correct* ANSWER, females were left

Answer	Gender	Laterality index (LI) means ± SD	Scheffé test probabilities for post hoc analysis interaction: 1×2				Valid N
			(1)	(2)	(3)	(4)	
Wrong	Female	$-0.22 \pm 1.7 (1)$		0.16	0.36	0.007	366
Wrong	Male	0.13 ± 1.9 (2)	0.16		0.002	0.7	216
Correct	Female	$-0.46 \pm 1.6 (3)$	0.36	0.002		0.000017	354
Correct	Male	$0.32 \pm 2.1 \ (4)$	0.007	0.7	0.000017		216
All groups		-0.12 ± 1.8					1152

Table 1
The mean ± SD and Scheffé test probabilities for post hoc analysis for the different groups

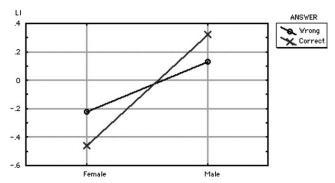


Fig. 1. Two way interaction of GENDER and ANSWER.

lateralized, while males were right lateralized. However, for *Wrong* ANSWER, there was a tendency towards bilateral hemispheric activation in both female and male subjects.

4. Discussion

Overall, these results demonstrate that for successful resolution of RPM tasks, females used a left hemisphere strategy while males used the right hemisphere. This implies that general intelligence is associated with neural systems within one hemisphere that are accessible to a variety of cognitive processes. These findings support the hierarchical hypothesis, and agree with results of recent studies, using positron emission tomography (Duncan et al., 2000) and fMRI (Prabhakaran et al., 1997). Similarly, the results are consistent with earlier studies in patients with lesions in the postrolandic structures (Basso et al., 1973).

In studies using RPM tasks, the areas of regional CBF activation found in young subjects (Esposito, Kirby, Van Horn, Ellmore, & Berman, 1999) comprised inferolateral temporal cortex including the fusiform gyrus bilaterally, and the middle temporal gyrus on the left, portions of the left medial temporal cortex including the parahippocampal gyrus, the left inferior parietal lobule, and the cerebellum. These anatomic structures (except for the cerebellum) derive major blood supply from the middle cerebral artery.

Gender differences in cognitive tasks have been a subject of intense debate. The Vandenberg mental rotation tests depicts rotation in three-dimensional space, and typically yields one of the largest established cognitive gender differences favoring males. Furthermore, processing in three dimensions is not a necessary condition for large sex difference on tests of mental rotation (Collins & Kimura, 1997). Conversely, superior performance by women on task requiring object location memory has challenged the traditional view that men excel on all spatial tasks (Kimura, 1996). The implications of the present findings of gender differential in RPM task could be better understood if a tentative hypothesis is advanced. It could be assumed that general intelligence is related to other lateralized hemispheric cognitive functions including some visuospatial skills. The finding that male subjects used a right hemisphere strategy may well explain the observation in numerous reports, that the average level of spatial talents usually requiring general intelligence is higher in male subjects than female (Buffery & Gray, 1972). This may suggest that, in males, ipsilateral right hemisphere localization of general intelligence facilitates visuospatial problem-solving. On the other hand, in females, transcallosal pathways are implicated in left hemisphere strategy of visuospatial processing. It is plausible that, reported gender-related hemispheric advantage is dependent on whether task processing strategies implicate either the intra-hemispheric or transcallosal pathways. Studies in commissurotomy patients might provide further insights on the implication of this model.

The finding of bi-hemispheric activation with *Wrong* ANSWER may suggest that, increasing level of task difficulty is associated with diverse pattern of neural activation due to broad sampling of all major cognitive functions. It could be postulated that, successful RPM problem-solving employs a discrete knowledge strategy (DKS), that selects neural pathways represented in one hemisphere. While unsuccessful outcome implicates a non-discrete knowledge strategy (nDKS).

RPM paradigm could be viewed as a working memory task. Others have argued that the necessity of keeping several conceptual formulations in mind during the RPM paradigm is by itself a working memory function (Carpenter et al., 1990). This suggests that the DKS model may have a correlate in mnemonic operations. In other words, DKS model may have a discrete knowledge base (DKB) of essential components needed for task resolution, while for nDKS, DKB is absent, and hence a 'global' or bi-hemispheric search occurs. Further studies with RPM in patients with memory deficits may provide useful insights.

The present study was based on the assumption that both neural and vascular brain structures share similar functional asymmetry. It has been suggested that neocortical vascularization which develops in conjunction with the nervous system and phylogenetic trends may share similar dominance characteristics (Njemanze, 1991). Recent evidence may support this view, for example, a receptor called neuropilin, is present both on neurons, where it is involved in axonal guidance, and on endothelial cells, where it aids angiogenesis (Chen, Chedotal, He, Goodman, & Tessier-Lavigne, 1997; He & Tessier-Levigne, 1997; Roush, 1998). Semaphorin III is another protein that has been identified and helps steer axons to their proper destination in the developing nervous system (He & Tessier-Levigne, 1997). It has been suggested that if neuropilin allows VEGF to act on neurons and semaphorins to act on endothelial cells, this might indicate an unexpected level of coordination between the developing nervous and circulatory systems (Roush, 1998).

The observed gender differences may suggest that sex hormones may be implicated in the above mechanisms. Moreover, there is growing evidence that early exposure to sex hormones has lasting effects on problem-solving behavior, and fluctuations in sex hormones in both men and women are associated with changes in cognitive pattern (Kimura, 1996). Furthermore, animal experiments have demonstrated that in the male rat at birth and in later life, the cortex of several areas in the posterior right hemisphere is significantly thicker than in corresponding areas on the left (Diamond, 1984). By contrast, in the female rat, there are several thicker areas in the left at the same periods. If the male rat is castrated at birth, many of the observed differences are not present at 90 days of age (Diamond, Dowling, & Johnson, 1981). Similar observations were made in females. In addition, ovarian hormones administered at birth alter the later pattern of cortical asymmetries (Geschwind & Galaburda, 1985).

Most of the postulates derived from the results of the present study are not aimed at resolving the outstanding issues in intelligence research, but is an attempt to further expand the scope of the discussion raised by results of prior studies (Basso et al., 1973; Duncan et al., 2000). It may appear simplistic to measure intelligence based on a single test. Raven's test is a good measure of intelligence and should account for a good deal of the reasoning in other tests (Carpenter et al., 1990). It may

still be that there is simplicity in the present approach. However, it is worthy to note, that even by this singular test, both cerebral lateralization for intelligence and gender differences were observed. It may be useful in future studies using TCD to evaluate if these results can be confirmed when a battery of intelligence tests is used. The present work suggests that TCD would be useful in the study of the diverse issues involved in intelligence research. TCD has high sensitivity (i.e., power to reveal subtle mental performance changes); and diagnosticity (i.e., capability to reveal the underlying processes that lead to these performance changes). The temporal superiority, portability, technical simplicity, cost-effectiveness, reliability, noninvasiveness, and ability to provide a global measure of cerebral lateralization are all added armamentarium that TCD provides to the study of brain cognitive functions.

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