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## Research report

## A brain for numbers

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## ABSTRACT

Healthy human brains come equipped with several circuits that contribute to number processing. Nature and nurture interact to produce a unique combination of core skills and more sophisticated abilities, by building on a handful of auxiliary routes (e.g., verbal language, body knowledge and visuospatial attention). Transcranial magnetic stimulation (TMS) studies on number processing will be here succinctly reviewed, in light of their most stimulating and challenging contributions. New research directions will be pointed out, that might enhance their theoretical impact.

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*The great glory of mathematics is its durative nature; that is one of humankind's longest conversations; that it never finishes by answering some questions and taking a bow.* (Barry Mazur (2003) – “Imagining Numbers”, p. 225)

Number processing is an essential part of our culture (Bynner and Parsons, 1997). Numbers are used for counting, measuring, comparing, ordering, identifying objects, calculating. Even people who do not deal professionally with numbers use their number skills in everyday life, and acquired impairment of numerical skills is a severe handicap in working life (Butterworth, 1999). Severe difficulties in learning about numbers (dyscalculia) are probably as widespread as disorders of literacy development (dyslexia), and the best prevalence estimates for each lie between 3.6 and 6.5% (Butterworth, 2005).

A deeper understanding of normal numerical skills is a necessary requirement for studying the impairment itself. TMS studies in mathematical cognition are aimed precisely to clarify the relation between the mental operations involved in normal number processing and specific brain circuits.

Within mathematical cognition, certain topics have attracted more attention (e.g., core quantity representation, the relation between finger pointing and counting or between numbers and spatial attention) probably also due to the fact that they are methodologically easier to investigate with TMS than others (e.g., complex arithmetic). Existing studies have met their objectives with mixed success and different protocols (see Table 1 for an overview), revealing the necessity of different cortical regions for number skills. Going beyond the virtual lesion metaphor, moreover, is perhaps the most intriguing open challenge in this field.

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**Table 1 – Summary of reviewed TMS studies including stimulation parameters and results**

| Study                       | TMS parameters |                                 |                                                                      | Sites and behavioural effects                                                                                                                   |                                                      |                                             |                                        |                                                                                                                                      |
|-----------------------------|----------------|---------------------------------|----------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------|---------------------------------------------|----------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------|
|                             | Frequency      | Intensity                       | Duration                                                             | Stimulation sites                                                                                                                               | Localization method                                  | Critical area                               | Task                                   | TMS effect                                                                                                                           |
| Göbel et al. (2001) Exp 1   | 10 Hz          | 105% of active MT <sup>a</sup>  | rTMS delivered pseudo-randomly, 500 ms, 100 ms before stimulus onset | PPL: X = $\pm 42$ , Y = $-58$ , Z = 52                                                                                                          | Functional and Brainsight™                           | Left PPL                                    | Comparison with 65                     | Slower RTs for numbers larger than and close to 65                                                                                   |
| Exp 2                       |                |                                 |                                                                      | SMG: X = $\pm 52$ , Y = $-46$ , Z = 44                                                                                                          |                                                      |                                             |                                        | No effect                                                                                                                            |
| Exp 3                       |                |                                 |                                                                      | PPL                                                                                                                                             |                                                      |                                             | Comparison with 65 (reversed response) | Generalized slowing of RTs                                                                                                           |
| Sandrini et al. (2004)      | 15 Hz          | 110% of resting MT <sup>a</sup> | 225 ms from stimulus onset                                           | 1 cm lateral to CP3 vs 1 cm lateral to CP4 (SMG) vs sham X = $\pm 48$ , Y = $-47$ , Z = 52                                                      | 10/20 EEG system and Softaxic                        | Left SMG                                    | Comparison of two digits               | Generalized slowing of RTs                                                                                                           |
| Oliveri et al. (2004)       | 1 Hz offline   | 90% of resting MT <sup>b</sup>  | 5 min followed by 30–60 min rest before the next session             | P3 vs P4 vs baseline                                                                                                                            | 10/20 EEG system                                     | P4                                          | Comparison of two numerical intervals  | Elimination of pseudoneglect                                                                                                         |
| Rusconi et al. (2005) Exp 1 | 1 Hz offline   | 90% resting MT <sup>a</sup>     | 10 min, 20–25 min pause in between blocks                            | AG vs SMG                                                                                                                                       | Brainsight™                                          | Left and Right AG                           | Finger gnosis task                     | Symmetric interference with the finger schema task from left AG; asymmetric (contralateral > ipsilateral) interference from right AG |
| Exp 2                       | 10 Hz          | 60% <sup>c</sup>                | rTMS delivered pseudo-randomly, 500 ms from stimulus onset           | AG                                                                                                                                              |                                                      | Left AG                                     | Magnitude matching + arithmetic prime  | Slower RTs in trials with unrelated primes                                                                                           |
| Andres et al. (2005)        | Single pulse   | 130% of resting MT <sup>a</sup> | Pulses at 150, 200, 250 ms after stimulus onset                      | P3 vs P4 vs P3+P4 (bilateral stimulation) vs sham                                                                                               | 10/20 EEG and TMS navigator (Noirhomme et al., 2004) | P3, P4 and P3+P4                            | Comparison with 5                      | Slower RTs to numbers close to 5 with TMS on P3 and P3+P4; slower RTs to far numbers with TMS on P3+P4                               |
| Göbel et al. (2006a)        | 5 Hz           | 110% of active MT <sup>a</sup>  | rTMS delivered pseudo-randomly, 1000 ms from stimulus offset         | AG/adjacent posterior part of the IPS left: X = $-34$ , Y = $-62$ , Z = 50; right: X = 38, Y = $-65$ , Z = 48; vs OCC X = 1, Y = $-98$ , Z = 18 | Functional and Brainsight™                           | Right AG/adjacent posterior part of the IPS | Mental number bisection                | Reduction of pseudoneglect                                                                                                           |

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