

## Writing and drawing tilts after right hemisphere stroke are signs of a wrong verticality representation

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1	Writing and drawing tilts after right hemisphere stroke are signs of a wrong
2	verticality representation
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1 2 3		
4 5 6 7 8 9	1	Abstract
10 11	2	Background
12 13 14	3	Many signs of spatial dysgraphia and drawing errors after right hemispheric stroke (RHS) have
15 16	4	been attributed to spatial neglect or impaired sensory feedback. Counterclockwise (contralesional)
17 18 19	5	tilts of graphomotor productions remained to be explained.
20 21 22	6	Objective
23 24	7	To test whether graphomotor tilts stem from a tilted representation of verticality transposed to the
25 26 27 28	8	top/bottom axis of the sheet of paper, from data of the DOBRAS cohort.
29 30	9	Methods
31 32	10	Handwriting and drawing orientations were measured from the writing of three lines and the
33 34 35	11	drawing of the Gainotti Figure (house roof lines). Verticality perception was measured with the
36 37	12	visual (VV) and postural (PV) verticals. Severity of extra-body (EBN) and body (BN) neglect were
38 39 40 41	13	each quantified with composite scores (battery of 8 tests).
42 43	14	Results
44 45	15	We analyzed data of 133 individuals: 64 healthy individuals (median age 63 years) and 69 with a
46 47 49	16	subacute RHS (median age 68 years). With respect to normal cut-offs (writing -5.4°; drawing -
40 49 50	17	8.1°), 26/69 (38%) individuals with RHS showed at least one graphomotor tilt (median[IQR] of
51 52	18	writing tilt: -6.8°[-9.7;-1.7]; drawing tilt: -10.9°[-17.6;-6.4]). Compared to individuals without
53 54	19	graphomotor tilts, those having both writing and drawing tilts showed greater contralesional tilts in
55 56 57	20	verticality perception (VV: -1.4°[-4;0.6] vs -7.9°[-11.5;-7.5]; PV: -1°[-2.4;0.2] vs -8°[-9;-5.4], Ps
57 58 59	21	≤.001) and more severe spatial neglect (EBN: 4.2[1.8;9.3] vs 16.6[10.2;20.4]; BN: 22.7[17;28.2]
60 61 62 63	22	vs 37.8[35.9;39.7], <i>P</i> s ≤.01). Composite graphomotor z-scores on writing and drawing correlated
64 65		1

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23	both with verticality estimates (VV+PV, $r_s$ =.46, $p < 10^{-4}$ ) and spatial neglect scores (EBN+BN, $r_s$ =-
24	.36, <i>p</i> <.01).

25 Conclusion

26 Contralesional tilts of writing and drawing after RHS are primary related to a tilted representation 27 of verticality, and secondary related to spatial neglect. They are easy to detect and could be 28 considered as a first step to perform early, before conventional tests of verticality perception.

29 Keywords

Right hemisphere stroke; graphomotor tilts; spatial dysgraphia; verticality representation; spatial
 neglect

32 Abbreviations

33 a.u.: arbitrary unit

34 BN: body neglect

2 35 CBS: Catherine Bergego Scale

4 36 DOBRAS: Determinants of Balance Recovery After Stroke

<sup>6</sup> 37 EBN: extra-body neglect

<sup>o</sup> 38 RHS: right hemispheric stroke

1 39 SCP: Scale for Contraversive Pushing

3 40 VP: postural vertical

<sup>5</sup> 41 VV: visual vertical

## Introduction

Manifestations of spatial dysgraphia and drawing errors after right hemispheric stroke (RHS) have revealed the contribution of the right hemisphere to the spatial organization of graphomotor behaviors, such as handwriting and drawing [1–6]. Spatial dysgraphia refers to various handwriting difficulties distorting the spatial layout of writing. It encompasses letter or word omission/iteration, overwriting, writing only in the right margin, and sloping lines [1,4,6,7]. Spatial dysgraphia has been mainly associated with damage to the right parietal lobe [1,4,5], as well as to subcortical grey and white matters [8,9]. Drawing errors typically comprise line omission/iteration, perspective loss or abnormal location, wrong proportions, and tilted orientation [2,3,6]. This constructional disorder has been frequently associated with damage to the right temporo-parietal lobes and adjacent white matter [10–13].

The brain areas involved in spatial dysgraphia and drawing errors explain why pioneer clinical observations reported a strong association between these graphomotor errors and spatial neglect [3,4,6,14]. This association is causal for lateralized graphomotor errors such as left omissions and right-page preference, plainly explained by attentional/representational deficits underlying the left-right gradient characterizing spatial neglect [15]. In contrast, spatial neglect does not easily explain - and was found dissociated from - non-lateralized errors such as letter duplication in the writing (e.g., [7]) or altered spatial relations in the drawing (e.g., [3]). These graphomotor errors could be rather explained by impaired visual/kinaesthetic feedbacks [6,7] or impaired spatial remapping [12].

But, none of the above-mentioned mechanisms hold for the counterclockwise tilts of graphomotor productions, which are frequent and remain to be explained. We hypothesized that they might result from a contralesional tilt in the representation of the vertical, transposed to the sheet of paper referring to the top and bottom. A recent single case study [5] supports this hypothesis, that we tested here, in a substantial series of right-handed individuals examined at 

the early subacute stage after an RHS. We investigated the associations between contralesional
graphomotor tilts, verticality estimates, and other relevant clinical deficits such as spatial neglect.

## 69 Methods

Data and analysis codes of this study are available on reasonable request from the corresponding
author. The reporting follows STROBE recommendations [16].

72 Study design and regulatory considerations

This study was performed in compliance with the Declaration of Helsinki and regulatory rules for human research in France. It was registered at the national committee for informatics and freedom (Commission Internationale Informatique et Liberté, no. 2014874 v1). All individuals were informed orally and in writing about the study and gave their informed consent. All participants, controls and with RHS, were blinded to the study hypotheses.

Writing and drawing productions of individuals with RHS were analyzed from data of the DOBRAS cohort (Determinants of Balance Recovery After Stroke), approved by our institutional review board (CHU Grenoble Alpes), which validated the ClinicalTrials.gov registration (NCT03203109). According to French law, eligible individuals were informed about the DOBRAS study (orally and in writing) and use of their data; those who did not want to participate signed an opposition form. Studies of the DOBRAS cohort focus on the recovery of balance and gait disorders after a first-ever hemispheric stroke, particularly in relation to spatial cognition deficits, such as impaired verticality perception and spatial neglect (for the first results, see [17-19]). Writing and drawing tasks were part of the routine assessment of spatial neglect.

87 The recruitment of healthy controls, selected to be matched to individuals with RHS, was 88 approved by the ethical review board of Grenoble-Alpes University (CERGA-2019–04–09–1).

They were recruited from the relatives of authors and from the relatives of patients hospitalized in our ward.

Their writing and drawing data have partly been reported in a previous methodological study intending to identify reliable criteria to measure writing and drawing orientations [20]. The goal of the present study was to understand graphomotor tilts after RHS, by identifying their clinical determinants.

#### **Participants**

We extracted from the DOBRAS database only the observations for right-handed individuals (Edinburgh inventory score  $\geq 0.4$ ) with a first-ever RHS, to focus on individuals potentially presenting spatial representation disorders, spatial dysgraphia, and drawing errors [3,6,21]. All individuals were able to write and draw with their non-paretic right upper limb. Among the three evaluations performed at 30, 60, and 90 days (D30, D60, D90) after the RHS, we selected the earliest and most complete evaluation in terms of writing, drawing, and other spatial cognition tests.

Exclusion criteria common to all DOBRAS studies were recurrent or complicated stroke, unstable medical condition, previous disability interfering with balance and verticality perception, psychiatric problems, or dementia (e.g., [18]). Additional exclusion criteria for this ancillary study were severe comprehensive impairment (due to aphasia, even if rare after RHS in right-handers), not French-speaking (as a native language), limb apraxia (Test of Upper Limb Apraxia score < 9; [22]), and altered ability to remain seated (Postural Assessment Scale for Stroke score < 8; [23]; see flowchart in Fig. 1).

We planned to recruit about 60 individuals with RHS to build multivariate models and test the effects of impaired verticality perception and spatial neglect on graphomotor tilts against each 2 other.

Healthy controls were recruited to be matched to the individuals with RHS in manual laterality (right-handers), age, sex ratio, and education level (1 = primary school certificate, 2 = ninth-grade certificate ["brevet"], 3 = bachelor's degree). They were asked about their medical history, had to not have neurological disorders, and had to have preserved abilities to draw and write legibly. No specific clinical examination or complementary investigation was performed.

#### 118 Writing and drawing tests: orientation criteria

For writing, the material was the French battery for neglect assessment (*Batterie d'évaluation de la négligence*; [24]; see Fig. 2a). Individuals were asked to manually write personal details on three lines on the blank sheet of paper in portrait format. For drawing, the material was the Gainotti Figure, representing a landscape with four trees and one house oriented vertically and horizontally [25]; see Fig. 2b). The Gainotti Figure was printed on the top of a page in landscape format, and individuals were instructed to manually copy it just below. For both tasks, the neuropsychologist checked that the sheet of paper was centered on the individual's trunk but never gave instructions regarding the disposition, orientation, or speed of writing and drawing.

To measure writing and drawing orientation criteria, all graphomotor productions were first digitized. Then, a home-made electronic protractor was used to compute the orientation of segments (e.g., the drawing of one of the house roof lines) defined from landmarks (e.g., extremities of the drawn segment) positioned by the operator (RL). The accuracy was set to 0.1°. RL was blinded to data from tests of verticality perception and spatial neglect. More details are given in our recent methodological paper [21], which showed that among various graphomotor criteria tested, the two best were the mean tilt from the horizontal of the three written lines ("writing orientation", Fig. 2a) and of the two drawn roof lines of the house ("drawing orientation", Fig. 2b). These two graphomotor criteria have excellent clinimetric properties and are the most suited to detect and quantify contralesional (i.e., counter-clockwise) tilts in individuals with a RHS [20]. Of
note, the "horizontal" orientation refers here to the width of the sheet of paper. By convention,
negative signs indicated contralesional orientations.

For both writing and drawing criteria, we determined the thresholds of abnormal contralesional tilt from data distribution (mean  $\pm 2$  SD) obtained in the 64 healthy controls. These distributions were approximatively Gaussian according to visual inspection and Kolmogorov-Smirnov tests (writing: p = .64; drawing: p = .10). These thresholds were [-5.4°; 1.7°] for the written lines and [-8.1°; 2.2°] for the drawn roof lines. With regards to ipsilesional graphomotor tilts, we found that only 2/69 (3%) individuals had a writing tilt > 1.7° (of a small magnitude: 3.1° and 3.4°) and 0/69 had a drawing tilt >  $2.2^{\circ}$ . These two individuals with a marginal ipsilesional writing tilt (without contralesional drawing tilt) had normal VV (-0.7°, -1.8°) and normal/quasi-normal PV (1°, -2.8°), and no spatial neglect. Because we planned to analyze whether contralesional graphomotor tilts were related to contralesional VV/PV tilts, we pooled these two individuals with those who had a normal graphomotor orientation, and a similar clinical profile without bias in verticality representation or spatial neglect (see Table 1 in Results).

We also computed a composite "graphomotor z-score" to serve as a robust index of the global magnitude of writing and drawing tilts in individuals with RHS. For this, their writing and drawing orientations were each z-scored from the means and standard deviations obtained in healthy controls, then averaged.

#### 155 Clinical deficits

156 Impaired verticality representation

The representation of verticality was assessed with two well-validated measures, the visual
 vertical (VV) and postural vertical (PV), described in detail elsewhere [26].

VV was measured in darkness by binocular visual adjustment of the orientation of a luminous

line (visual angle =  $5.7^{\circ} \times 0.1^{\circ}$ ) presented on a computer screen at eye level, about 1.5 m from the nasion. In case of hemianopia, the computer screen was adequately translated toward the non-hemianopic side. Individuals were seated with their head and trunk maintained upright and were asked to adjust the line to the vertical by giving verbal instructions to the examiner. The starting orientation of the line, randomly ranging from  $\pm 5^{\circ}$  to  $\pm 30^{\circ}$ , was balanced to the right or left following a sequence fixed across individuals.

PV was assessed in darkness with the wheel test (see [26]). Individuals were seated, restrained in a wheel-like device that was slowly and manually turned by the examiner and were asked to adjust their body to the vertical by giving verbal instructions to the examiner. The starting orientation of their body, ranging from  $\pm 15^{\circ}$  to  $\pm 45^{\circ}$ , was balanced to the right or left following a sequence fixed across individuals.

For both tasks, individuals performed 10 trials. All measures started after 2 min of darkness. The examiner never gave a time limit or feedback. Data from all 10 trials were averaged to calculate individual mean VV and PV orientations. In addition, we calculated "transmodal" vertical estimates (means of VV and PV) to get a unique measure of verticality representation (i.e., independent from a specific vestibular or somaesthetic modality of gravity sensing). According to standards, negative and positive VV or PV values corresponded to contralesional (i.e., counterclockwise) and ipsilesional (i.e., clockwise) orientations, respectively. VV and PV tilts were diagnosed according to well-established ranges of normality (-2.5 to 2.5°; [26,27]).

#### 9 Spatial neglect

Extra-body neglect (EBN) and body neglect (BN) were evaluated by trained neuropsychologists and occupational therapists with a battery of eight well-validated tests routinely used in clinical practice [24]. EBN was evaluated with (1) the Bells cancellation test [28], (2) the overlapping figure test [29], (3) the reading test [24], and (4) four EBN items of the Catherine Bergego Scale (CBS; [30]). BN was evaluated with (5) the Bisiach test [31], (6) the Fluff test [32], (7) the Comb and

Razor test [33,34], and (8) four BN items of the CBS [30]. Participants completed these tests with
 their sound right upper limb (see [18,19] for more details about these eight tests).

Spatial neglect was diagnosed from at least two abnormal test results, whatever the spatial neglect domain. We also evaluated EBN and BN severities by calculating composite scores. For this, raw scores for all eight neglect tests were converted to percentages based on their respective ranges (0% = no neglect, 100% = maximal neglect). These percentages were then averaged to calculate EBN and BN scores, as well as a "global" spatial neglect score (mean of EBN and BN scores). More details about these composite scores are provided in Appendix A.

#### 193 Disability and other deficits

Gait and disability were evaluated with the modified Fugl-Meyer Assessment gait score [35], ranging from 0 arbitrary unit (a.u.) = no gait, to 6 a.u. (= normal gait) and the Functional Independence Measure [36], ranging from 18 a.u. (= very severe disability, a.u.) to 126 a.u. (= total independence). Lateropulsion, which refers to an abnormal postural behavior against gravity, was evaluated with the Scale for Contraversive Pushing (SCP; [37]), ranging from 0 a.u. (= upright body orientation, a.u.) to 6 a.u. (= active pushing of the body toward the contralesional side). Lateropulsion was considered moderate or severe with an SCP score > 0.5 [18, 27]. These three scales are detailed in previous DOBRAS reports (e.g.,[18,26]). Hemianopia was manually evaluated by trained physicians.

#### 3 Statistics

Descriptive statistics are first presented for each graphomotor criterion, both for orientation magnitude and prevalence of abnormal tilt. Data were analyzed with non-parametric tests, as residual plots from parametric tests revealed assumption violations and outlying data.

To identify the main determinants of contralesional graphomotor orientation, we first compared the clinical profile of three groups of individuals with RHS: (1) those having normal graphomotor

orientation or light ipsilesional tilt, (2) those having a contralesional writing *or* drawing tilt, (3) those
 having both contralesional writing *and* drawing tilts. Differences between groups were tested with
 Kruskal-Wallis (omnibus) tests for continuous measures, and with chi-squared tests for categorical
 measures.

Then, we conducted two-tailed Spearman correlations and Pearson and Filon' z tests to test whether the graphomotor z-scores were more associated with the transmodal vertical estimates (mean of VV and PV) than with the global spatial neglect scores (mean of EBN and BN).

Continuous data are presented as median [Q1; Q3] and categorical data as number (%). Standardized effect sizes were calculated by using the  $r_s$  (rho) value of the Spearman test, the  $\sqrt{\eta^2}$  value of the Kruskal-Wallis test, the r value of the Mann-Whitney test, and the Cramer's w of the chi-squared test. Effect sizes > 0.1, 0.3, or 0.5 were considered small, medium, or large, respectively [38]. The significance level was set at *p* < .05 and Bonferroni-corrected when needed. All data analyses were conducted with R 4.0.3 [39].

## Results

#### Participants

A total of 133 individuals took part in this study (see flowchart in Fig. 1): 69 with RHS and 64 healthy controls, matched in age (median 68 years [62; 73] vs 63 years [59; 71]; Mann-Whitney, p = .24, r = 0.10) and sex ratio (n = 43/69 [62%] vs 35/64 [55%] males;  $\chi 2$ , p = .47, w = 0.06). With regards to education levels, no difference was found between individuals with RHS and healthy controls ( $\chi 2$ , p = .10, w = 0.19): level 1, 27/69 [39%] vs 14/64 [22%]; level 2, 18 [26%] vs 21 [33%]; level 3, 24 [35%] vs 29 [45%]). Clinical data were collected mostly at D30 (n = 48, 70%), sometimes at D60 (n = 13, 19%), and more rarely at D90 (n = 8, 12%). The stroke was mainly a cerebral infarct (n = 61, 88%). The individual disability was moderate, with a median FIM score of 102 [75; 114]; the median mFMA gait score was 5 [2; 6], indicating that half was able to walk without

technical aid; the median SCP score was 0.25 [0; 1.5], indicating that most of them were either
upright or showed mild lateral body tilt with respect from gravity.

#### 6 Missing data

Only 9/4200 (0.21 %) data points were missing in the whole dataset. For clinical deficits, 3 (4%) PV tests and 2 (3%) VV tests were missing because of participant uneasiness or dizziness in the dark. Two (3%) other PV tests were missing because of technical problems or the inability of a participant to sit in the device because of a pelvic fracture. For the drawing test, roof lines were not drawn by 2 (3%) individuals. Data imputation was not required given these few missing data.

#### 42 Distribution of graphomotor orientations in participants with RHS

Figures 3a and 3b present data distribution for each graphomotor criterion. Orientations were mostly contralesional for both criteria: -1.8° [-4.5; -0.6] for the writing and -4.4° [-8.3; -2.5] for the drawing. The frequency of contralesional (abnormal) tilts were comparable for the writing (n =15/69, 22%) and the drawing (n = 18/69, 26%). Overall, 43/69 (62%) individuals did not show any contralesional graphomotor tilt, 19/69 (28%) individuals showed a contralesional tilt for at least one criterion (writing tilt = -2.6° [-8; -1.1]; drawing tilt = -9.6° [-15.1; -4.6]), and 7/69 (10%) a contralesional tilt bearing on both criteria (writing tilt = -9.5° [-13.1; -7.4]; drawing tilt = -17.6° [-23.7; -11.2]). Writing and drawing orientations were nonetheless not significantly correlated (Spearman test,  $r_s = .20$ , p = .10). As highlighted in Fig. 3, there were cases of dissociations: 8/69 (12%) individuals showed an isolated contralesional writing tilt, whereas 13/69 (19%) showed an isolated contralesional drawing tilt.

Figure 3c shows data distribution for the graphomotor z-scores, representing the mean severity of writing and drawing tilts. Important variations were noted: range = -8.3 to 1.5 SD, median = -0.4

#### 6 SD [-2; 0.3]).

#### 7 Verticality perception, spatial neglect, and hemianopia

For VV, abnormal contralesional tilts were found in 34/67 (51%) individuals. In others, VV was either normal in 25/67 (37%) individuals or more rarely subjected to a light ipsilesional tilt (*n* = 8/67, 12%). For PV, abnormal tilts were found in 27/64 (42%) individuals, always toward the contralesional side, otherwise was normal in 37/64 (58%). Contralesional tilts were transmodal (VV and PV) in 20/64 (31%) individuals. See Appendix B for details on associations and dissociations.

Spatial neglect was diagnosed in 41/69 (59%) individuals, with 29 (42%) individuals having EBN, 36 (52%) having BN, and 24 (35%) having both. Left hemianopia was diagnosed in 23/69 (33%) individuals.

#### 7 Determinants of graphomotor tilts

Table 1 shows the clinical profile of individuals with RHS according to their graphomotor orientations (normal graphomotor orientation or light ipsilesional tilt [n = 43/69, 62%], contralesional writing or drawing tilt [n = 19, 28%], contralesional writing and drawing tilts [n = 7, 10%]). As for verticality estimates, the severity and frequency of contralesional VV and PV tilts gradually increased with the number of contralesional graphomotor tilts (all *p*-values  $\leq$  .01, moderate effect sizes). Whereas VV and PV estimates were in normal ranges (-2.5 to +2.5°) in individuals without graphomotor tilts, they raised to -8° (about 6 SD) in individuals with writing and drawing tilts. These latter had 2 to 4 times more tilts in verticality perception. Congruently, they also showed greater lateropulsion. The severity and frequency of spatial neglect (EBN and BN) also increased with the number of contralesional graphomotor tilts. This was specially the case for EBN (*p*-values  $\leq$  .01, moderate effect sizes), found in 28% of individuals without graphomotor tilts,

and in 100% of individuals with writing and drawing tilts (Table 1). In contrast, the three graphomotor groups did not differ for hemianopia and demographic features.

Figure 4 shows that graphomotor z-scores were moderately correlated with transmodal vertical estimates ( $r_s = .46$ ,  $p < 10^{-4}$ ) and global spatial neglect scores ( $r_s = -.36$ , p < .01), without significant difference between these two correlations (Pearson and Filon's z test on absolute values, z = 0.85, p = .39).

Overall, these results showed the importance of spatial deficits, tilted verticality representation and spatial neglect, in contralesional graphomotor tilts of individuals with RHS.

## Discussion

Our main objective was to test whether contralesional tilts of writing and drawing after an RHS were associated with a tilted verticality representation, transposed to the top/bottom axis of the sheet of paper. For this aim, individuals with a subacute RHS were asked to write three lines and to draw the Gainotti Figure in the framework of clinical routine practice. Two reliable criteria of graphomotor orientation were extracted: the three lines for the writing, and the two roof lines for the drawing. Our results reveal that tilted writing and drawing orientations were primary correlated to an impaired verticality representation, and secondary correlated to spatial neglect severity. Congruently, individuals with a contralesional writing and drawing tilt differed from others by more severe and frequent contralesional VV and PV tilts, greater lateropulsion, and more severe spatial neglect. These findings support the hypothesis that spatial cognition deficits, and notably tilts in verticality representation, may manifest in the orientation of graphomotor productions after an RHS.

# Tilted verticality representation: the primary determinant of writing and drawing tilts

Altogether, our results show that contralesional graphomotor tilts were associated to all measures of verticality estimates (visual, postural, and transmodal). This suggests that writing and drawing tilts are not related to a specific graviceptive modality (vestibular or somaesthetic) but rather to a higher-order representation of verticality. Of note, the VV mainly tests the processing of gravityrelated *vestibular* information (conveyed by the otolith organs; [27,40]), whereas the PV mainly tests the processing of gravity-related *somesthetic* information (conveyed by proprioceptive, tactile, and visceral receptors; [41,42]). These two sources of information would be integrated into and pondered by the multisensory thalamo-vestibular network to represent the vertical [26,43,44].

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<sup>4</sup> 312 The RHS could disturb these multisensory mechanisms, leading to abnormal contralesional tilts not only in VV and PV estimates, but also in graphomotor tasks involving an implicit representation of the "upright". Of interest, very severe tilts in verticality representation might even be able to manifest in various graphomotor tasks. Indeed, the most severe tilts of VV and VP were found in individuals having both writing and drawing tilts, two tasks that yet differ in terms of instructions, materials, and mechanisms (generation [writing] vs copy [drawing]).

Is the association between graphomotor tilts and tilted verticality representation causal? Experimental investigations in healthy individuals have established that verticality representation is used for various sensorimotor tasks such as body orientation, catching, grasping, or pointing movements (for recent reviews see [45,46]). The brain might also use a representation of verticality during the execution of graphomotor tasks involving spatial orientation skills in 2D. This view is supported by Jolly et al. (2020), who recently showed that a short-term modulation of verticality perception in the patient JW (PV, -10° before vs 0.5° after) was congruently associated with a significant diminution of his writing tilts (-10° before vs. -6° after).

In the current series, the association between verticality estimates and graphomotor tilts was less strong than that reported by Jolly et al. (2020) in their case-study (i.e., moderate vs. strong effect sizes). This finding may be explained first by the multi-determined nature of graphomotor behaviors. Writing and drawing involve a large range of cognitive operations, encompassing spatial attention, spatial remapping, coordinate transformation, object recognition, executive functions, and motor control [1-3,6]. The disruption of one of these mechanisms after an RHS could induce graphomotor tilts on their own, independent of an impaired verticality representation. This could be notably the case in individuals showing a dissociation between writing and drawing tilts (about 30%). For example, an impaired spatial remapping might disturb writing and drawing orientations in a non-systematic way, depending on the trajectory of the saccadic eye movements. The Gainotti task requires numerous up and down saccades between the model and the drawing space, whereas the writing task requires horizontal saccades between the letters and words. An

impaired motor control might also influence the writing and drawing in different ways, as these tasks require different motor planification and motor commands. Second, the moderate effect sizes we observed could also be explained by the use of the paper sheet's borders, or drawing model, as a visual allocentric frame of reference. Indeed, individuals with an impaired integration of gravity-related vestibular or somaesthetic cues showed increased reliance on visual cues to estimate verticality [47,48]. Despite a severe tilt in verticality representation, individuals with an RHS using this allocentric strategy might compensate for their deficit in such conditions and exhibit upright graphomotor orientations. This allocentric strategy could have been used especially in the drawing task, as the Gainotti Figure provides numerous visual landmarks.

#### Spatial neglect: a secondary determinant of writing and drawing tilts?

Contrary to our expectations, we found that graphomotor tilts were also moderately related to spatial neglect severity and frequency. Although spatial neglect is an important determinant of many signs of spatial dysgraphia or drawing errors [1,4,6], how it would induce graphomotor tilts remains still difficult to conceive.

The more plausible interpretation is that the association between graphomotor tilts and spatial neglect is mediated by concomitant tilts in verticality representation. Indeed, clinical studies have increasingly suggested that an impaired representation of verticality is a key component of spatial neglect [17,19,21,49], and it has been recently shown that transmodal (visual and postural) vertical tilts [19] and lateropulsion [17] constituted very strong proxy of spatial neglect. This mediation hypothesis is also supported by a recent case study, reporting a dissociated recovery between spatial neglect and contralesional graphomotor tilts [5].

359 Might spatial neglect induce graphomotor tilts because of a distortion between the 360 representation of the left and right hemispaces? A possible mechanism could involve the line 361 breaking, more and more shifted to the right in individuals with spatial neglect [1,4], which results

in a left margin tilt. If the orthogonality between the margin (vertical coordinate) and writing lines (horizontal coordinate) is respected in these individuals [5,21], this line breaking should also indirectly tilt the writing lines upward. But such a mechanism cannot explain the tilts of the drawing rooflines. Thus, our data and the recent literature rather suggest that spatial neglect plays a role through a bias in verticality representation.

#### Clinical implications

In individuals showing a contralesional tilt in both writing and drawing tasks, VV and VP tilts were peculiarly frequent (> 80%) and severe (median tilt >  $-7^{\circ}$ ), and associated with a severe lateropulsion (median SCP = 3). Therefore, these graphomotor tilts, which are easy to search early even in patients with a severe stroke and bedridden, should be more systematically detected in individuals with a RHS [24].

More indirectly, our results might also suggest a role of spatial cognition disorders in 3D on non-lateralized visuo-constructive deficits, which remain to be better understood. We speculate that the combination of and impaired verticality representation and a representational distortion on the left-right axis (spatial neglect), possibly with an impaired motor control, might alter assembling segments to draw or build figures in 2D and 3D, leading to the so-called constructional apraxia.

#### 79 Limitations

A first limitation is the observational nature of the writing and drawing data collected from the DOBRAS cohort. Hence, we could not verify that the sheet of paper was always perfectly aligned with the trunk and head of the participant, although neuropsychologists carefully monitored the positions of the sheet and the patient's body. Likewise, we could not investigate other potential

mechanisms of graphomotor tilts, such as feedback-related errors [6,7] and impaired spatial remapping [9,12,50], which still needs to be studied. Second, we did not control for the potential effect of ocular torsion on graphomotor tilts, although ocular torsion is infrequent after hemispheric stroke [27] and should not alter our findings. Finally, we did not investigate the representation of "horizontality", which is known to be, as with VV, tilted counterclockwise after stroke [21]. Nevertheless, clinical and experimental studies suggest that the VV and visual horizontal are approximately orthogonal [21].

### Conclusions

Our study supports the hypothesis that contralesional tilts of writing and drawing after RHS are related to a tilted representation of verticality transposed to the top/bottom axis of the sheet of paper. This may explain why these graphomotor tilts are observed in individuals with spatial neglect, who are often subject to a wrong representation of verticality. These graphomotor tilts should be more systematically searched, which can be done early after the stroke, even in bedridden individuals having a severe RHS. Their existence may alert about the existence of a wrong verticality representation which will require specific rehabilitation.

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## **CRediT** authorship contribution statement

Rémi Lafitte: Writing – original draft, Investigation, Data curation, Formal analysis. Flora Diaine: Writing – original draft, Investigation, Data curation, Formal analysis. Shenhao Dai: Writing – original draft, Investigation, Data curation, Formal analysis. Olivier Carré: Software. Eve Dupierrix: Writing - review & editing, Conceptualization. Caroline Jolly: Writing - review & editing, Methodology, Conceptualization. Céline Piscicelli: Writing – review & editing, Investigation, Conceptualization, Methodology. Dominic Pérennou: Writing - original draft, Conceptualization, Methodology, Supervision, Funding acquisition.

## **Declaration of competing interests**

None.

## **Data availability**

The dataset analyzed during the current study is protected and not available due to data privacy

laws, but is available from the corresponding author on reasonable request.

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## **Figure Legends**

Fig 1. Flowchart of participants in the study.

DOBRAS: Determinants of Balance and Recovery After Stroke; PASS: Postural Assessment Scale for Stroke

Fig 2. Measures of writing and drawing orientations with satisfactory clinimetric properties [20].
(a) Writing criterion. (b) Drawing criterion. For both (a) and (b), the yellow segments defined the horizontal and vertical axes of the sheet. Of note, the angle values extracted by the software had to be reversed.

Fig 3. Distribution of (a) writing line orientations, (b) drawing roof line orientations, and (c) graphomotor z- scores. Distributions are ranked by decreasing order of writing line orientations.
The horizontal dashed lines represent the ranges of normality for each graphomotor criterion.
Positive and negative values refer to clockwise and counterclockwise orientations, respectively.

Fig. 4. Effects of transmodal verticality estimates and global spatial neglect scores on the global

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<sup>4</sup> 576	magnitude (z-scores) of contralesional graphomotor tilts.

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Table	1
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Response variable	Normal orientation or ipsilesional tilt (n = 43)	Contralesional writing or drawing tilt ( <i>n</i> = 19)	Contralesional writing and drawing tilt (n = 7)	p*	Effect size**
VV (°)***	-1.4 [-4; 0.6]	-6.4 [-9.2; -1.4]	-7.9 [-11.5; -7.5]	< .001	0.47
VV (contralesional tilt)	15/43 (35%)	13/19 (68%)	6/7 (86%)	.005	0.40
PV (°)***	-1 [-2.4; 0.2]	-3.5 [-5.3; -2]	-8 [-9; -5.4]	.002	0.41
PV (contralesional tilt)	10/43 (23%)	11/19 (58%)	6/7 (86%)	< .001	0.46
SCP (a.u.)	0 [0; 0.6]	0.5 [0; 3]	3 [1.8; 3.6]	< .001	0.45
EBN score (%)	4.2 [1.8; 9.3]	10 [4; 22]	16.6 [10.2; 20.4]	.004	0.37
EBN (yes)	12/43 (28%)	10/19 (53%)	7/7 (100%)	< .001	0.45
BN score (%)	22.7 [17; 28.2]	31.3 [18.8; 37.2]	37.8 [35.9; 39.7]	.0038	0.37
BN (yes)	19/43 (44%)	11/19 (58%)	6/7 (86%)	.11	0.26
Hemianopia (yes)	11/43 (26%)	8/19 (42%)	4/7 (57%)	.16	0.23
Education (1/2/3)	14 (33%) / 13 (30%) / 16 (37%)	9 (47%) / 4 (21%) / 6 (32%)	4 (57%) /1 (14%) /2 (29%)	.66	0.19
Sex (male)	27/43 (63%)	12/19 (63%)	4/7 (57%)	.96	0.04
Age (years)	66.5 [55; 72.2]	68 [63; 71]	69 [63.5; 76]	.53	0.00

**Table 1.** Univariate analyses of possible indicators determining the clinical profile of individuals showing contralesional graphomotor tilt.

Data are median [Q1; Q3] unless otherwise indicated.

\* The alpha level was set to .0038 because of multiple tests (.05/13). Significant p values are shown in bold.

\*\*  $\sqrt{\eta^2}$  from the Kruskal-Wallis test for continuous variables and Cramer's *w* from the chi-squared test for categorical variables. Effect sizes  $\ge 0.1$ , 0.3, or 0.5 were considered small, medium, or large, respectively. \*\*\* For VV and PV, statistical analyses involved 67 and 64 individuals, respectively, because of missing data.

a.u.: arbitrary unit, VV: visual vertical, PV: postural vertical, SCP: Scale for Contraversive Pushing, EBN: extra-body neglect, BN: body neglect















Appendix A

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