#### Supplementary Text

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Regional specificity of the association between cortical folding and learning rate

Cortical folding develops mostly prenatally and during the first years of life <sup>1</sup>. Therefore, cortical developmental events preceding the late maturation phase could have an impact on cortical folding in pre-SMA/SMA and its relation to learning ability. To rule out factors that influenced the extent of cortical folding of the whole hemisphere, we re-examined the correlation between local cortical folding in pre-SMA/SMA and the learning rate ( $R^2 = 0.317$ , p < .001, Fig. 2B), this time adjusted for individual differences in the folding of the whole cortex (total folding index, see <sup>2</sup>). Although a higher total folding index was significantly correlated with a steeper learning rate ( $R^2 = 0.05$ , p = .041), the relationship between local cortical folding in pre-SMA/SMA and learning ability was only slightly reduced when we adjusted for total folding index (partial  $R^2 = 0.286$ , p < .001). This shows that the impact of cortical folding on learning is region-specific and relatively independent from hemispheric cortical folding.

Effect size estimation for interventions using correction for individual differences in folding

Ouantitative comparison of effect sizes suggests that performance-enhancing 17 psychological strategies (positive social comparative feedback  $[R^2 = 0.31 \text{ in }^3]$ ) or exercise 18 interventions (post-exercise physical activity  $[R^2 = 0.301 \text{ in }^4 \text{ or chronic physical activity}]$ 19  $[R^2 = 0.259 \text{ in }^5]$ ) can have as large an impact on stabilometer motor performance as cortical 20 folding in the pre-SMA/SMA ( $R^2 = 0.30$ ). Our previous study <sup>5</sup> showed that 2 weeks of 21 intense physical exercise prior to motor practice increases the learning rate in this postural 22 task. Since we included this subsample in our analysis (e.g., the green dots in Fig. 2C), we 23 were interested in the covariation pattern of cortical folding and previous physical exercise 24 effects on learning rate. Therefore, we reanalyzed the data and, first, replicated the positive 25 effect of prior intense physical activity on learning rate compared to the control condition 26 (two-tailed unpaired t-test, t(29) = 2.50, p = .022, mean difference = 0.12, Cohen's d = 27 0.898, 95% CI = 0.150, 1.632). When we adjusted for inter-individual differences in cortical 28 folding in pre-SMA/SMA, the magnitude of the exercise effect continued to increase (two-29 tailed unpaired t-test, t(29) = 3.29, p = .003, mean difference = 0.13, Cohen's d = 1.181, 30 95% CI = 0.406, 1.940), indicating independent effects of prior intense exercise and cortical 31 folding on learning rate. From a statistical perspective, this finding suggests that 32 consideration of individual predispositions in the brain can improve effect size estimation 33 for interventions targeting behavioral change. 34

#### The impact of motivation

Feedback and instructions from teachers or coaches can have a positive (or negative) 37 impact on motor performance and learning <sup>6</sup>. Modulating a learners' focus-of-attention or 38 motivation can be beneficial for skill acquisition and retention <sup>7</sup>. Although we controlled for 39 performance feedback as well as task instructions and the degree of practice exposure 40 throughout our experiments, individual differences in motivation may have had an impact 41 on motor performance. However, core motivational brain circuits are localized in the 42 midbrain, ventral striatum and orbitofrontal cortex <sup>8</sup> and neural activity in these areas adapts 43 during learning <sup>9</sup>. Future studies are required to disentangle the motivational contribution of 44 reward-related brain structures to motor learning of the stabilometer task. Of particular 45 interest in this context is whether individual differences in motivation share variance with 46 the effects of cortical folding on learning rate. The spatial localization of our present 47 neuroimaging result, however, indicates that cortical folding in pre-SMA/SMA supports 48 postural task-specific learning capabilities rather than general functions such as motivation. 49

# 50Relationship between cortical curvature and training-induced microstructural cortical51plasticity

The sample used for the main analysis consists of three subsamples (see 52 Supplementary Table 4) with slightly different research questions and study designs. 53 Respective research questions centered around balance-induced and exercise-induced brain 54 plasticity and the time course of training-related functional and structural brain changes. In 55 some of these studies, and thus also in the sample for our main analysis, training-related 56 changes in the left pre-SMA/SMA were identified. For example, in our original balance 57 training study<sup>10,11</sup>, we found changes in gray matter volume and functional connectivity in 58 the left pre-SMA/SMA. The pre-SMA/SMA was also observed in our most recent training 59 study<sup>12</sup> (n=26). Both data sets were part of the current sample for the main analysis (n=84). 60 A thorough comparison between cross-sectional measurements of cortical anatomy (e.g., 61 cortical folding) and training-induced changes in cortical structure requires homogenization 62 of different training protocols and different imaging methodologies between studies (partly 63 on different MRI scanners). This is crucial in order to identify common patterns of plasticity 64 65 across studies, which can then be compared with the baseline values of e.g. cortical folding. In the meantime, however, we can correlate effects within a subsample. Our most recent 66 study<sup>12</sup> provides the largest subsample (n=26) and a plasticity index (orientation dispersion 67 index, ODI) that changed with balance training and also correlated with inter-individual 68 differences in learning rate. While both measures individually correlated with learning rate 69 in this sub-sample (r = 0.59 for ODI change and r = 0.51 for cortical curvature), a 70 71 correlation analysis suggests a low correlation between ODI change and cortical folding in pre- SMA/SMA (r = .18). When we entered these two variables (change in ODI as well as 72 cortical curvature) in a linear regression model predicting learning rate, the results show that 73 both ODI change and cortical curvature represent two distinct mechanisms significantly 74 predicting individual differences in learning rate ( $\beta$ =.53, T=3.3, p=.004 for ODI change; 75  $\beta$ =.36, T=2.2, p=.036 for curvature). 76

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**Supplementary Table 1:** List of differing head coils, echo times, repetition times, flip angles, and inversion times of the T1-weighted MRI datasets used for the main analysis of learning rate (n=84, samples marked with \*) and the additional analysis of the first practice session (n=131).

| number of<br>subjects | head coil | echo time<br>(ms) | repetition<br>time (ms) | flip angle (°) | inversion<br>time (ms) |
|-----------------------|-----------|-------------------|-------------------------|----------------|------------------------|
| N=27*                 | 32 ch     | 3,46              | 1300                    | 10             | 650                    |
| N=26*                 | 64 ch     | 2,82              | 2500                    | 7              | 1100                   |
| N=31*                 | 32 ch     | 2,98              | 2300                    | 9              | 900                    |
| N=24                  | 32 ch     | 2,96              | 2300                    | 9              | 900                    |
| N=11                  | 12 ch     | 3.46              | 1300                    | 10             | 650                    |
| N=6                   | 12 ch     | 2.96              | 2300                    | 9              | 900                    |
| N=5                   | 32 ch     | 3,46              | 1300                    | 10             | 650                    |
| N=1                   | 12 ch     | 2,23              | 1300                    | 10             | 650                    |

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Supplementary Table 2: Cortical folding results from the main analysis (N=84) are not dependent on the use of different imaging protocols. Table shows multiple regression models containing cortical folding in pre-SMA/SMA, age, gender, body height, and TIV as predictors of learning rate (Model A). In model B, head coil (binary coded) was included as additional co-regressor, whereas models C considered the three imaging protocols as dummy coded variable (based on Suppl. Table 1, marked with\*). Partial regression statistics between cortical folding and learning rate are shown.

|                             | β    | Т     | р     |
|-----------------------------|------|-------|-------|
| Model A                     | ,398 | 4,988 | <,001 |
| Model B (+ head coil)       | ,402 | 4,988 | <,001 |
| Model C (+ MRI<br>protocol) | ,404 | 5,335 | <,001 |

Supplementary Table 3: Correlations between cortical folding and learning rate are
 consistent between the 32-ch (N=58) and 64-ch (N=26) data subsets of the main analysis
 (see supplementary table 1). We extracted cortical folding values in pre-SMA/SMA for
 these two subsets of participants. The residuals of learning rate and cortical folding (see
 Fig. 2 in the main manuscript) constituted the input for Chow's test (Chow, 1960), which
 tests the null hypothesis that two regression lines are equal (i.e. can be represented by one
 single regression line).

|                  | F-statistics | p-value |
|------------------|--------------|---------|
| Left pre-SMA/SMA | 0.071442     | 0.9311  |

**Supplementary Table 4:** Detailed characteristics of samples included in the main analysis (N=84) and the separate analyses of initial performance and short-term adaptations (N=131). The table includes sub-study, sub-sample, age (mean), number of female participants, body height (in cm) and mean TIV (in ml). Data was analyzed from several studies: <sup>a</sup> from Taubert et al. (2010)<sup>11</sup>, <sup>b</sup> unpublished data, <sup>c</sup> from Lehmann et al. (2023)<sup>12</sup>, <sup>d</sup> from Lehmann et al. (2020)<sup>5</sup>, e from Taubert et al. (2016)<sup>13</sup>.

| Whole<br>sample | Analysis  | Sub-<br>studies | Sub-groups        | Mean age | #females | Mean body<br>height (cm) | Mean TIV |
|-----------------|---|-----------------|-------------------|----------|----------|--------------------------|----------|
| N=131           | Main sample for<br>analysis of long-<br>term learning (6<br>practice sessions,<br>N=84)     | N=27            | N=14ª             | 25.8     | 6        | 172                      | 1481     |
|                 |   |                 | N=13 <sup>b</sup> | 22.8     | 4        | 177                      | 1569     |
|                 |   | N=26            | N=26℃             | 22.1     | 4        | 179                      | 1557     |
|                 |   | N=31            | N=16 <sup>d</sup> | 23.6     | 11       | 174                      | 1505     |
|                 |   |                 | N=15 <sup>d</sup> | 24.3     | 9        | 173                      | 1478     |
|                 | Additional sample<br>with short-term<br>learning data only<br>(1 practice<br>session, N=47) | N=24            | N=24 <sup>e</sup> | 25.9     | 12       | 174                      | 1443     |
|                 |   | N=11            | N=11 <sup>b</sup> | 26.3     | 6        | 173                      | 1502     |
|                 |   | N=6             | N=6 <sup>b</sup>  | 26.5     | 2        | 173                      | 1530     |
|                 |   | N=5             | N=5 <sup>b</sup>  | 27.4     | 2        | 173                      | 1476     |
|                 |   | N=1             | N=1 <sup>b</sup>  | 23.0     | 1        | 163                      | 1338     |



- 109 Supplementary Figure 1: Relationship between learning rate and initial level of
- 110 **performance.**
- 111 Scatter plot showing Spearman correlation between learning rate *n* across the six practice sessions
- and initial performance in session 1.
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|                       |                         | learning rate n | initial<br>performance a | age   | gender | height | TIV   | study1 | study2 | study3 |
|-----------------------|-------------------------|-----------------|--------------------------|-------|--------|--------|-------|--------|--------|--------|
| learning rate n       | correlation coefficient |                 |                          | -     | -      | -      |       |        |        |        |
|                       | N                       | 84              |                          |       |        |        |       |        |        |        |
| initial performance a | correlation coefficient | 657             |                          |       |        |        |       |        |        |        |
|                       | Sig. (2-sided)          | .000            |                          |       |        |        |       |        |        |        |
|                       | N                       | 84              | 84                       |       |        |        |       |        |        |        |
| ade                   | correlation coefficient | 170             | 037                      |       |        |        |       |        |        |        |
| -                     | Sig. (2-sided)          | .881            | .741                     |       |        |        |       |        |        |        |
|                       | N                       | 84              | 84                       | 84    |        |        |       |        |        |        |
| gender                | correlation coefficient | ,355            | -,207                    | ,043  |        |        |       |        |        |        |
|                       | Sig. (2-sided)          | ,001            | ,059                     | ,696  |        |        |       |        |        |        |
|                       | N                       | 84              | 84                       | 84    | 84     |        |       |        |        |        |
| height                | correlation coefficient | ,220            | -,210                    | -,118 | ,571   |        |       |        |        |        |
|                       | Sig. (2-sided)          | ,045            | ,055                     | ,284  | ,000   |        |       |        |        |        |
|                       | N                       | 84              | 84                       | 84    | 84     | 84     |       |        |        |        |
| TIV                   | correlation coefficient | ,230            | -,144                    | ,075  | ,633   | ,624   |       |        |        |        |
|                       | Sig. (2-sided)          | ,035            | ,191                     | ,497  | ,000   | ,000   |       |        |        |        |
|                       | N                       | 84              | 84                       | 84    | 84     | 84     | 84    |        |        |        |
| study1                | correlation coefficient | ,079            | -,045                    | ,276  |        | ,274   | ,155  |        |        |        |
|                       | Sig. (2-sided)          | ,441            | ,687                     | ,011  |        | ,012   | ,159  |        |        |        |
|                       | Ν                       | 84              | 84                       | 84    |        | 84     | 84    | 84     |        |        |
| study2                | correlation coefficient | -,194           | ,070                     | ,171  |        | -,107  | ,005  |        |        |        |
|                       | Sig. (2-sided)          | ,078            | ,526                     | ,121  |        | ,331   | ,966  |        |        |        |
|                       | Ν                       | 84              | 84                       | 84    |        | 84     | 84    |        | 84     |        |
| study3                | correlation coefficient | ,263            | -,025                    | ,099  |        | -,159  | -,153 |        |        |        |
|                       | Sig. (2-sided)          | ,016            | ,881                     | ,370  |        | ,149   | ,164  |        |        |        |
|                       | N                       | 84              | 84                       | 84    |        | 84     | 84    |        |        | 84     |

### Supplementary Figure 2: Correlation matrix including initial performance and learning rate as well as nuisance variables.

117 Correlation matrix with learning rate *n* and all nuisance variables used in the main analysis

118 (n=84). Coefficients indicate Pearson or Spearman correlations between two continuous variables

119 (e.g. age and body height) depending on data normal distribution as well as Point-biserial

120 correlation coefficients for a continuous and a categorial variable (e.g. gender and age).

121 Correlations between two categorical variables (e.g. gender and study 1) were omitted. Studies 1,

122 2 and 3 include datasets for the main analysis (N=84) indicated in supplementary table 1 with \*.



### 125 Supplementary Figure 3: Cortical folding and initial performance.

126 Correlation between local cortical folding and initial performance (mean performance across 15

trials in practice session 1) in 131 (left) and 84 (right) participants (see supplementary table 1). No

significant effects were found across the whole cortex. Color bar represents uncorrected p values.

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#### 131 Supplementary Figure 4: Cortical folding and early performance improvements.

132 Correlation between local cortical folding and early (short-term) learning within the first practice

session (N=131). No significant effects were found across the whole cortex. Color bar represents
 uncorrected p values.



## Supplementary Figure 5: Correlation coefficients r depicting positive relationships between cortical folding and learning rate.

- 139 Vertex-wise map of correlation coefficients (*r*) from the main exploratory analysis (Fig. 2). The
- 140 color bar represents r values depicting the positive correlation between local cortical folding and 141 learning rate n.
- 142



#### 144 Supplementary Figure 6: Reproducible effect of cortical folding on learning rate.

- 145 (A) Results of whole-brain regression of vertex-wise cortical curvature to learning rate using a
- second MRI scan of the same participants. Uncorrected results at p < .001 (left) and family-wise
- error-corrected results at p < .05 (right) were projected onto a template brain showing variations in sulcus depth. (B) Positive correlation of residual cortical curvature (in the cluster representing
- the FWE-corrected effect in the exploratory analysis [Fig. 2A]) and learning rate. (C) Subsample
- results in the three independent learning experiments. (D) Pearson correlation coefficients
- 151 between residualized cortical curvature and motor performance. Grey bars represent session-
- 152 specific performance controlled for initial performance in session 1 (i.e., residual gain) and black
- bars represent correlations with actual session-specific performance. \* indicate significant
- 154 correlations at p < .05.

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# Supplementary Figure 7: Study-specific sub-group analyses of cortical curvature effects on learning rate.

159 The five sub-groups were drawn from the three longitudinal training studies<sup>5,11,12</sup>. Sub-group 160 analyses of the effect of cortical curvature on learning rate n. The effect was consistent across

regions-of-interests (ROI) obtained from the original analysis (Fig. 2), its reproduction with

another MRI scan of the same participants as well as from a previous study in which postural

163 learning-induced grey matter changes were found in this region <sup>11</sup>. All correlation coefficients

164 were significant at p < .05.

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## Supplementary Figure 8: Statistical maps showing positive correlations between cortical folding and asymptotic (final) performance.

- 169 Correlation between local cortical folding and final performance in the sixth practice session
- 170 (N=84). Significant effects (corrected for multiple comparisons) were found in a cluster located in
- the left supramarginal gyrus (see red circle on the right; labeling according to the Desikan-
- 172 Killiany atlas). Note a trend level effect (left) in the cluster in left pre-SMA/SMA overlapping
- with the cortical folding effect on learning rate (Fig. 2). Color bar represents uncorrected p values.



## Supplementary Figure 9: Positive correlation between local gyrification index (Schaer et al., 2008) and learning rate.

178 Vertex-wise correlation between local gyrification index (lGI) and learning rate (N=84). While

179 local curvature characterizes local geometrical properties of the cortical surface (Fig. 2), the lGI is

180 sensitive to variations in surface area buried within cortical sulci. Note the positive correlation in

the caudal part of the left superior frontal gyrus spatially coincides with results from the local  $\frac{1}{2}$ 

182 curvature analysis (pre-SMA/SMA, Fig. 2). Color bar represents Z values.



184 Supplementary Figure 10: Effects of age and learning rate on intracortical microstructure.

Correlation analysis of study 2 (N = 26, yellow dots in Fig.2) for which we acquired additional quantitative MRI protocols. We analysed intracortical microstructural properties include myelinsensitive magnetization transfer (MT). Vertex-wise correlations with MT were calculated with learning rate n (A) and age (B). Exploratory statistical thresholds (p < .01 uncorrected at vertex-

level) were applied and thresholded *t* values are displayed. Color bar represents *t* values.

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# Supplementary Figure 11: The influence of initial performance differences on later achievements decreased with practice.

Pearson correlation between mean performance in session 1 and mean performance in sessions 2-6.



- Supplementary Figure 12: Performance decrements between successive practice sessions.
  The bars represent the difference in our primary behavioural outcome measure (time in target
- 200 zone in seconds) between the mean of the first two trials of the actual session (session n+1) and 201 the mean of the last two trials of the previous session (session n).
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