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Functional brain correlates of writing acquisition / ECRITAPP

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Project-ANR-14-CE30-0013 (S. Kandel, 2014-2019)



-Filan



french word « galop » written by a child (left) and by an adult (right)

77 participants included 3 groups: CE2 (N = 24), CM2 (N = 25) and Adults (N=28) Inclusion: standardized graphomotor, spelling and reading tests

LOCALIZER RUN

Write Letters, words, or draw loops, tablet recordings

TR= 957 ms, voxel size= 2.5 mm3, multiband factor= 4, slices= 56

335 volumes

4 DICTATION RUNS

Write regular and irregular words to dictation, tablet recordings

TR= 957 ms, voxel size= 2.5 mm3, multiband factor= 4, slices= 56

440 volumes per run



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Palmis et al. Dev. Sci. 2021

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T2 MRI

Voxel

size=

1mm3

T1 MRI

Voxel

size=

1mm3

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440 volumes per run

Developmental effects of spelling regularity: Palmis, Fabiani et al., in prep.

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Morphometric analysis: Cachia, Dupont et al., in prep.

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Sarah Palmis's PhD

Postdoctoral researchers



Sarah Palmis, PhD Postdoctoral Fellow



Our main goal at the Advances in Brain and Child Development Research Laboratory is to improve the lives and well-being of children with neurodevelopment disorders. We perform clinical and neuroscientific investigations that will help us to better understand the underlying mechanisms of function and dysfunction in individuals with brain-based disorders from birth to early adulthood.

Sarah joined the ABCD research laboratory as a Postdoctoral Fellow in February 2021. She completed a PhD in Neurosciences at Université de Aix-Marseille. During her PhD she developed an expertise in functional MRI technique as a proxy to study neural and motor development. During her postdoctoral training, Sarah is interested in using a variety of quantitative structural MRI techniques to characterize brain growth in a variety of clinical populations. In her free time, she enjoys doing sport in particular basketball, reading, listening to music, and cooking.

HANDWRITING PROCESSES



Relationship between orthographic and motor processes during handwriting

Rapp and McCloskey, 2017

HANDWRITING PROCESSES



Rapp and McCloskey, 2017

Serial vs parallel flow of processing?

Do orthographic processes affect motor processes?

How does this develop?

Assessment of the effect of psycholinguistic variables on parameters of motor execution during written word production

Delattre and Bonin, 2006; Kandel et al., 2006; Kandel and Perret, 2015; Damian and Freeman, 2008; Baus et al., 2013; Sausset et al., 2013; Planton et al., 2017b; Pinet et al., 2017; Roux et al., 2013; Scaltritti et al., 2016; 2017; etc...

HANDWRITING ACQUISITION



Palmis et al, Cogn. Neuropsychol., 2017

HANDWRITING ACQUISITION

Orthographic processes



Motor processes

"Developmental studies support the idea that at ages 9–10 word writing starts to be regulated by orthographic knowledge" (Kandel and Perret, 2015)



THE WRITING NETWORK

- Left inferior frontal gyrus
- Left fusiform gyrus
- Left superior parietal lobule
- Left superior frontal gyrus
- Right Cerebellum

Regions coding orthographic information

Regions coding motor aspects of writing



Purcell et al., 2011; Planton et al., 2013; Rapp et al., 2016, etc...

Outline of the talk:

1- Quick overview of previous studies

2- Developmental effects of spelling regularity on writing behavior and brain activation

 \rightarrow The aim: show you the results and discuss questions raised by statistical analyses

3- If time: Morphometric analisys of OTS and ACC: early cerebral constraints on writing development

1- Quick overview of previous studies

The impact of spelling regularity on handwriting: relationship between orthographic and motor processes in adults (study 1)

Palmis, Velay, Fabiani, Nazarian, Anton, Habib, Kandel, Longcamp, Cortex, 2019

STUDY 1: BACKGROUND

Spelling inconsistent words: a <u>conflict</u> between the output of the two routes



Rapp and McCloskey, 2017

STUDY 1: DESIGN and BEHAVIORAL RESULTS

Effects of the presence of an irregularity and its position (Roux et al., 2013)NATUREPHARAONCAFARD







STUDY 1 : RESULTS

IFG

Beta value 0-

-2-

IRB

IRF Irregularity



IRB

IRF

Irregularity

REG

Orthographic processes are still active during motor execution

PREVIOUS STUDY IN ADULTS



Orthographic processes are still active during motor execution

Orthographic processes influence motor processes

Study 2: Comparing the writing network of adults and children (study 2)

Palmis et al. Dev. Sci. 2021

STUDY 2 : DESIGN

Block design: write series of letters, words and draw loops

Subsample of 23 adults (aged 19 to 40, mean 24.91) and 42 children (aged 8 to 11, mean 8.88)

We pooled CE2 and CM2 in a single children group



STUDY 2 : RESULTS



- Writing network also activated in children
- More activation clusters in children
- Activation in anterior/frontal regions

STUDY 2 : RESULTS



 Writing network also activated in children

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- More activation clusters in children
- Activation in anterior/frontal regions
- Group differences: Frontal regions (ant insulas, ACC);
 Left Fusiform Gyrus ; Cerebellum (right anterior lobe) ;

Study 3: Developmental effects of spelling regularity on writing behavior and brain activation





Palmis, Fabiani et al, in prep.

How does the relationship between orthographic and motor processes of handwriting evolve between middle childhood and adulthood ?

Children writing is less automated than adults writing: <u>Age 9 is critical</u> because it is the period in which grapho-motor skills start being automatic (Halsband & Lange, 2006; Mojet, 1991)

In adults, orthographic and motor processes occur in parallel during handwriting

H1 = In children, orthographic and motor processes should be more independent and sequential (Olive, 2014; Kandel and Perret, 2015)

H2 = Lexical processes strongly impact graphomotor processes during writing acquisition but this influence diminishes at some point in development (Afonso et al., 2018)

STUDY 3: DESIGN



- Writing-to-dictation task on a MRI-compatible tablet
- 2 conditions : regular words (REG) and words irregular <u>at the end (IR)</u>
- 100 words : 50 REG (<u>aut</u>ruche, <u>avoine</u>, <u>ban</u>ane, etc)
 50 IR (automne, avocat, bandit, etc)
- Recording behavioural and fMRI data

STUDY 3: BEHAVIOURAL DATA



Writing Latency

Total duration

Duration of the first 3 letters

STUDY 3: BEHAVIOURAL DATA



STUDY 3: BEHAVIORAL RESULTS



(rather a good point for the fMRI statistical models)

Interaction (5th gr vs adults)

Interaction (5th gr vs adults)

+ Similar effects on writing size

STUDY 3: fMRI DATA

fMRI data preprocessing: fMRIprep

- Calculation of the IQMs with MRIqc and descriptive stats (Julien)
- Exclusion of the sessions where more than 25% of the data points have FD values > 0.9 mm (27 sessions, including 2 participants (children))
- Systematic exploration of the IQMs before and after preprocessing (fMRIprep)
- fMRIprep: fMRIPrep 20.0.6 (Esteban, Markiewicz, et al. (2018); Esteban, Blair, et al. (2018); RRID:SCR_016216) tpl-MNI152NLin2009cAsym_space-MNI_res-01_T1w spatially smoothed (FWHM 5 mm) with FSL

Confounds: 24 motion regressors + 26 nuisance regressors (22+1 WM, 22+1 CSF) + motion censoring (Frames that exceeded a threshold of 0.5 mm FD or 1.5 standardised DVARS were annotated as motion outliers)

STUDY 3: fMRI DATA

fMRI data modeling - Main model

5 regressors of interest :

- Regular words writing



- Irregular words writing
- Rest
- Auditory stimulus
- Empty trials

!! Participant excluded if nb sessions < 3 out of 4</pre>

 $17/24/26 \rightarrow 10/20/26$ participants per group

	CONDITION	DURATION	ONSET
ed	REG	Effective total writing duration Or mean writing duration	Onset of stim + latency Or mean latency
ut	IR	Effective total writing duration Or mean writing duration	Onset of stim + latency Or mean latency
	Rest	10.27s	Onset of stim
6	Stim	0	Onset of stim
qr	Empty	0	Onset of stim



STUDY 3: fMRI DATA

fMRI data modeling - Other models

Auditory Stimulus

5 regressors of interest :

- Regular words Auditory stim

Irregular words Auditory stim

All categories (except empty trials)

- Rest

-

- Auditory stimulus
- Empty trials

For all 3 models: The number of sessions was matched between groups

Writing: correct trials

8 regressors of interest :

- Correct Regular words
- Correct Irregular words
- Misspelled Regular words
- Misspelled Irregular words
- Rest
- Auditory stimulus
- Empty trials
- Other trials



Second level models: flexible factorial designs







STUDY 3: fMRI RESULTS



STUDY 3: RESULTS

Massive main effect of regularity during (Palmis et al., 2019)

Auditory stimulus

Writing response









STUDY 3: RESULTS

Distribution of irregularity processing in adults (red) and 5th Graders (Green) during writing



Irregularity is processed moslty in the writing network of adults

More prefrontal distribution in 5th graders

p < .005 unc.
STUDY 3: RESULTS

« Orthographic regions »: L FuG and L IFG







group

Adults

CE2

STUDY 3: RESULTS

« Motor regions »: R Cb and L dPM





STUDY 3: RESULTS

ACC : interaction between group and regularity during writing \rightarrow selective effect in 5th graders



Interaction group X condition: whole brain





ACC + Caudate Nucleus + dorsolateral PFC: conflict monitoring network





MNI Pediatric template

Spatial Normalization: choice of template

Weng et al., 2015 the norm index of the affine transformation matrix, i.e., the SFN, characterizes the difference between a template and a native image and differs significantly across subjects

 \rightarrow include the SFN as a covariate in group-wise statistics?

Extra models: FIR



STUDY 3: CONCLUSIONS

Summary of the findings

- Coupled behavioral and brain evidence that orthographic aspects of handwriting are still active during writing, and cascade over the motor components in both expert adults and children
- Writing acquisition is mediated by massive behavioral changes, and by differential involvement of several parts of the writing network in adults and children
- Similar behavioral effects but different neural underpinning in 5th graders vs adults
- Possible confounds: writing in the MRI scanner and absence of visual feedback, data quality differences between groups, amount of data, precision of behavioral data, and choice of template for spatial norm.

STUDY 4: MORPHOMETRIC ANALYSIS:

Relationship between folding patterns in ACC and OTS and writing skills: Early cerebral constraints on writing acquisition (Arnaud Cachia, LaPsyDE, Paris with Olivier and Guillaume)



Figure 1. A-Brain activations during writing in a group of adults and children (8-11 y.o., Palmis et al., 2021). The ACC and fusiform Gyri are circled. B- Sulcal variability in the ACC (Borst et al., 2014). C- Sulcal Variability in the OTS (Borst et al., 2016)

Cortex cingulaire antérieur (ACC)

Patterns du CCA et contrôle cognitif : plus efficient lorsque les motifs des sillons sont asymétriques (Cachia *et al.*, 2014; Borst et al. 2014; Tissier et al. 2018)



L'effet de l'asymétrie du CCA sur l'efficience du contrôle cognitif chez des enfants (Cachia et al., 2014)

Sillon occipito-temporal (OTS)

Patterns de l'OTS et lecture : plus performante lorsque les motifs des sillons sont discontinus en postL (Borst et al., 2016 ; Cachia et al., 2018)



L'effet de la morphologie de l'OTS sur la compétence de lecture chez des lettrés et des illettrés (Cachia et al.,2018)

Structural MRI data processing

- A partir des images anatomiques T1 et T2→ reconstitution de l'interface matière grise – matière blanche en un maillage constitué d'environ 60 000 sommets (freesurfer)
- Ces images tridimensionnelles ont ensuite été importées dans le logiciel BrainVISA.
- Les motifs des sillons ont été déterminés visuellement « en aveugle » et labélisés manuellement par 2 des co-auteurs à partir de la reconstruction 3D des plis corticaux (Cachia et al., 2014).
- CCA : « simple » ou « double parallèle » basé sur la présence ou l'absence d'un sillon paracingulaire (PCS)
- OTS: continuous or interrupted we identified whether OTS interruption was located in the posterior part of the sulcus hosting the VWFA or anterior

Analyse IRM : Identification des motifs sulcaux CCA



Visualisation de deux hémisphères (gauche et droit) – En jaune : sillon cingulaire ; en bleu : sillon paracingulaire

Preliminary results

- R Package Lavaan : SEM to test the influence of sulcal patterns of OTS and ACC on graphomotor, orthographic and reading scores, without latent variable
- → Replication of the results of Cachia et al., 2018, with significant effect of left posterior interruption on reading scores (nb words read per minute for adults, and reading accuracy for children)
- → Finding of a combined effect of ACC asymetry and right OTS interruption on graphomotor performance (writing speed and quality, larger effects for children)
- \rightarrow No effect of sulcal patterns on orthographic scores





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