

**Localization Patterns of Speech and Language Errors during Direct Electrical Brain Stimulation: A Systematic Review, Neurosurgical Review, K.E. (Ellen) Collée, A. (Arnaud) J.P.E. Vincent, E. (Evy) Visch-Brink<sup>1</sup>, E. (Elke) De Witte<sup>1</sup>, C. (Clemens) M.F. Dirven, D. (Djaina) D. Satoer**

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#### **Supplementary Information 1: Search strings**

Embase.com

('surgery'/de OR surgery:Ink OR 'surgical technique'/de OR 'neurosurgery'/exp OR 'cancer surgery'/de OR 'intraoperative period'/de OR 'brain mapping'/de OR 'electrostimulation'/de OR 'neuromonitoring'/de OR 'glioma'/exp OR 'brain tumor'/exp OR (map\* OR language\*-monitor\* OR neuromonitor\* OR neuro-monitor\* OR craniotom\* OR surg\* OR neurosurg\* OR operat\* OR intraoperat\* OR anesthes\* OR resect\* OR electrostimulation\* OR electro-stimulation\* OR glioma\* OR glioblastoma\* OR astrocytoma\* OR oligodendrogioma\* OR ((intracranial\* OR brain\*) NEAR/3 (neoplas\* OR tumor\* OR tumour\* OR cancer\*)):ab,ti,kw) AND ('wakefulness'/de OR (awake\* OR waking\* OR wake\*):ab,ti,kw) AND ('speech disorder'/exp OR 'perseveration'/de OR 'language ability'/de OR 'language disability'/de OR 'linguistics'/exp OR 'language processing'/de OR (aphasi\* OR paraphasi\* OR dysphasi\* OR perseveration\* OR anomia\* OR apraxia\* OR dysarthria\* OR semantic\* OR neologism\* OR linguistic\* OR ((language\* OR speech\*) NEAR/3 (error\* OR arrest\* OR perform\* OR mistake\* OR disturbance\* OR abilit\* OR disabilit\* OR dysfunction\* OR disorder\* OR process\*)):ab,ti,kw)

Medline Ovid

(surgery.fs. OR Neurosurgery/ OR exp Neurosurgical Procedures/ OR Intraoperative Period/ OR Brain Mapping/ OR exp Glioma/ OR exp Brain Neoplasms/ OR (map\* OR language\*-monitor\* OR neuromonitor\* OR neuro-monitor\* OR craniotom\* OR surg\* OR neurosurg\* OR operat\* OR intraoperat\* OR anesthes\* OR resect\* OR electrostimulation\* OR electro-stimulation\* OR glioma\* OR glioblastoma\* OR astrocytoma\* OR oligodendrogioma\* OR ((intracranial\* OR brain\*) ADJ3 (neoplas\* OR tumor\* OR tumour\* OR cancer\*))).ab,ti,kf.) AND (Wakefulness/ OR (awake\* OR waking\* OR wake\*).ab,ti,kf.) AND (exp Speech Disorders/ OR Language Disorders/ OR Anomia/ OR Linguistics/ OR (aphasi\* OR paraphasi\* OR dysphasi\* OR perseveration\* OR anomia\* OR apraxia\* OR dysarthria\* OR semantic\* OR neologism\* OR linguistic\* OR ((language\* OR speech\*) ADJ3 (error\* OR arrest\* OR perform\* OR mistake\* OR disturbance\* OR abilit\* OR disabilit\* OR dysfunction\* OR disorder\* OR process\*))).ab,ti,kf.)

Web of Science

TS=(((map\* OR language\*-monitor\* OR neuromonitor\* OR neuro-monitor\* OR craniotom\* OR surg\* OR neurosurg\* OR operat\* OR intraoperat\* OR anesthes\* OR resect\* OR electrostimulation\* OR electro-stimulation\* OR glioma\* OR glioblastoma\* OR astrocytoma\* OR oligodendrogioma\* OR ((intracranial\* OR brain\*) NEAR/2 (neoplas\* OR tumor\* OR tumour\* OR cancer\*)))) AND ((awake\* OR waking\* OR wake\*)) AND ((aphasi\* OR paraphasi\* OR dysphasi\* OR perseveration\* OR anomia\* OR apraxia\* OR dysarthria\* OR semantic\* OR neologism\* OR linguistic\* OR ((language\* OR speech\*) NEAR/2 (error\* OR arrest\* OR perform\* OR mistake\* OR disturbance\* OR abilit\* OR disabilit\* OR dysfunction\* OR disorder\* OR process\*))))))

Cochrane Central Register of Controlled Trials

((map\* OR (language\* NEXT/1 monitor\*) OR neuromonitor\* OR neuro-monitor\* OR craniotom\* OR surg\* OR neurosurg\* OR operat\* OR intraoperat\* OR anesthes\* OR resect\* OR electrostimulation\* OR electro-stimulation\* OR glioma\* OR glioblastoma\* OR astrocytoma\* OR oligodendrogioma\* OR ((intracranial\* OR brain\*) NEAR/3 (neoplas\* OR tumor\* OR tumour\* OR cancer\*)):ab,ti,kw) AND ((awake\* OR waking\* OR wake\*):ab,ti,kw) AND ((aphasi\* OR paraphasi\* OR dysphasi\* OR perseveration\* OR anomia\* OR apraxia\* OR dysarthria\* OR semantic\* OR neologism\* OR linguistic\* OR ((language\* OR speech\*))

NEAR/3 (error\* OR arrest\* OR perform\* OR mistake\* OR disturbance\* OR abilit\* OR disabilit\* OR dysfunction\* OR disorder\* OR process\*)):ab,ti,kw)

Google Scholar

craniotomy|mapping|surgery|neurosurgery|intraoperative|resection|resected|electrostimulation|glioma|glio  
blastoma|astrocytoma|oligodendrogioma awake|waking|wakefulness  
aphasia|paraphasia|dysphasialanguage|speech errors|performance|disorders|processing"

Medline (Ovid)

Neurosurgery/

Exp Brain Mapping/

Exp Glioma/

Wakefulness/

Exp Speech Disorders/

**Supplementary Information 2: Speech and language errors taken from the articles categorized by type (in bold)**

No	Speech and language errors	No	Speech and language errors
1	<b>Speech arrest</b>		Dyslexia
2	<b>Dysarthria/Anarthria</b>		Reading deficits/disturbances/impairment
	Dysarthria		Reading arrest
	Articulatory difficulty		Delayed (comprehensive) reading
	Anarthria	8	<b>Speech initiation difficulties</b>
3	<b>Semantic errors</b>		Delayed speech initiation
	Semantic speech error/paraphasia		Initiation difficulty
	Semantic disturbance/error		SMA aphasia
	Semantic disorder/deficit/aphasia		Difficulty with sentence completion
	Semantic association disturbance/error		Prolonged inter-word durations
	Non-verbal semantic processing problem		Reduction of spontaneous speech
	Semantic jargon aphasic language	9	<b>Production errors</b>
	Semantic paraphasias in writing		Slurred speech
	Semantic comprehension error		Speech delay
4	<b>Phonemic errors</b>		Hesitation
	Phonemic speech error/paraphasia		Vocalization
	Phonemic disturbance		Slow speech
	Phonological paraphasia		Stammering
	Phonological processing/disturbance		Stuttering
	Phonemic paraphasias in writing	10	<b>Anomia/word finding difficulties</b>
5	<b>(Morpho-)syntactic errors</b>		Anomia
	Syntactic disorders		Naming delay/delayed word retrieval
	Syntactic gender error		Word finding/searching/retrieval difficulties
	Morphological over regularization (verbs)		Circumlocutions
	Inflection errors	11	<b>Perseveration</b>
6	<b>Comprehension errors</b>	12	<b>Writing errors</b>
	Comprehension difficulty		Writing arrest
	Impaired comprehension		Alterations of letter shapes in writing
	Auditory comprehension difficulty/error		Writing drift
	Word deafness		Spelling errors
7	<b>Reading errors</b>	13	<b>Verbal apraxia</b>
	Alexia	14	<b>Irrelevant paraphasia</b>

No = number

### Supplementary Information 3. Explanation of the structure of the data

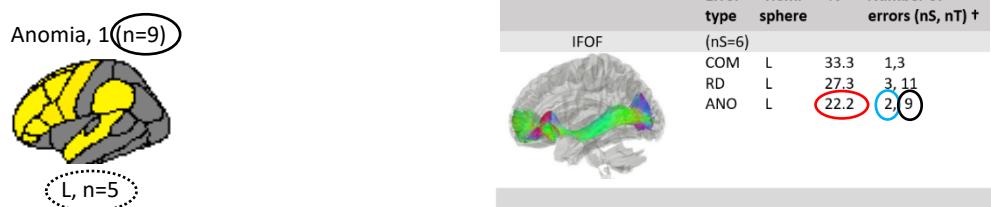
#### A. Second analyses: calculation

For each data set (except for the excluded 4) and each paraphasia type separately, the percentage of occurrence per location was calculated based on all occurrences of that paraphasia (cortically and subcortically). Example: in data set 1, speech arrest occurred 6 times in the PrG. In total, 10 speech arrests were found in this data set (cortically and subcortically). Based on this total, the occurrence of speech arrest in the PrG is calculated:  $((6/10)*100=60\%$ . Note: these numbers are for illustrative purposes only and do not reflect existing data.

Data set 1			Data set 2		Data set 3		
Speech arrest			Speech arrest		Speech arrest		
C/S	Location	n	Percentage	n	Percentage	n	Percentage
C	PrG	6	$(6/10)*100=60.0\%$	5	$(5/7)*100=71.4\%$	1	$(1/6)*100=16.6\%$
C	MTG	2	$(2/10)*100=20.0\%$	1	$(1/7)*100=14.3\%$	3	$(3/6)*100=50.0\%$
S	FAT	1	$(1/10)*100=10.0\%$	1	$(1/7)*100=14.3\%$	0	$(0/6)*100=00.0\%$
S	WM below x	1	$(1/10)*100=10.0\%$	0	$(0/7)*100=00.0\%$	2	$(2/6)*100=33.3\%$
<b>10=100%</b>			<b>7=100%</b>		<b>6=100%</b>		
 Anomia			 Anomia		 Anomia		
C	SFG	3	$(3/9)*100=33.3\%$	3	$(3/9)*100=33.3\%$	1	$(1/8)*100=12.5\%$
C	MTG	2	$(2/9)*100=22.2\%$	2	$(2/9)*100=22.2\%$	0	$(0/8)*100=00.0\%$
S	WM under x	2	$(2/9)*100=22.2\%$	2	$(2/9)*100=22.2\%$	6	$(6/8)*100=75.0\%$
S	IFOF	2	$(2/9)*100=22.2\%$	2	$(2/9)*100=22.2\%$	1	$(1/8)*100=12.5\%$
<b>9=100%</b>			<b>9=100%</b>		<b>8=100%</b>		

#### B. Second analyses: visualization

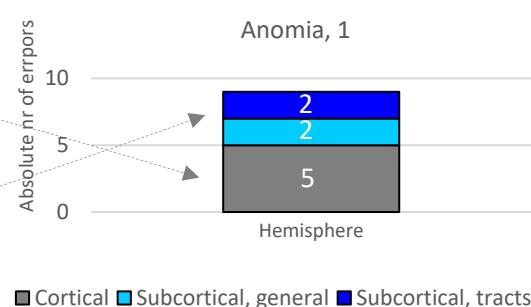
The calculated percentages (see A) were used to compute cortical (Figure 3, 4) and subcortical (Figure 5) plots. A separate cortical plot was computed for each data set and paraphasia type (three plots per paraphasia type). Example: in data set 1, anomia occurred 9 times in total (cortically and subcortically, see black matching circle in A and B) and 5 times at the cortical level (dashed black circle in A and B). Anomia occurred twice at the IFOF (blue circle in A and B), which corresponds to 22.2% (red circle in A and B). Note: even though one plot visualized either cortical or subcortical areas, the used totals and percentages were based on BOTH levels (see B).



#### C. Third analyses

It was calculated how often each paraphasia type occurred cortically and subcortically per data set. A division was made between general subcortical areas and tracts. For example, anomia from data set 1 (see A, copied below) occurred  $(3+2)=5$  times cortically and  $(2+2)=4$  times subcortically, of which 2 times at the general level and 2 times at the tract level (see Figure 7 for actual plot). In total, anomia occurred 9 times in this data set, which can be seen in the right plot on the y-axis (absolute number of errors). Each data set and paraphasia type was seen as a subset (plot below was seen as 1 subset). The subset below contained more cortical (5) than subcortical errors (4). Additionally, it contained subcortical general and tract locations.

Anomia		
C	SFG	$\circlearrowleft$ =5
C	MTG	$\circlearrowleft$ =2
S	WM under x	$\circlearrowleft$ =2
S	IFOF	$\circlearrowleft$ =2 +
<b>9=100%</b>		



#### D. Summary

- The percentages/totals are based on cortical and subcortical locations, even though one plot visualised either cortical or subcortical locations.
- The percentages from the cortical plots and subcortical plot do not add up to 100%, since 1) some cortical areas are unplotable with the DKT-atlas (see 2.4.4.) and are thus not shown in the plot and 2) only the subcortical tracts and not the general subcortical areas are displayed in Figure 5 (see 2.4.2.).
- Note: for analyses 1, a comparable structure was followed. However, the frequencies of occurrence per location (cortical and subcortical) were combined for all data sets and paraphasia types (see Supplementary Materials 4 for DKT-atlas compatible cortical frequencies). Additionally, these percentages were only visualized in a cortical plot (see Figure 2) and not in a subcortical or division plot, while the totals were again based on cortical and subcortical data.

**Supplementary Information 4: The brain locations in the current data and the converted brain locations compatible with the (lateral) DKT-atlas.**

<b>Brain locations (our data)</b>	<b>Converted brain locations (DKT-atlas)</b>
Broca's area	Pars opercularis, pars triangularis
Wernicke's area	Superior temporal gyrus
Inferior frontal gyrus	Pars opercularis , pars orbitalis, pars triangularis
Pars opercularis	Pars opercularis
Premotor cortex, motor cortex, precentral gyrus,	Precentral
SMA	
Frontal lobe	Precentral, caudal middle frontal, rostral middle frontal, superior frontal, pars opercularis, triangularis, pars orbitalis, lateral orbitofrontalis
Occipital lobe	Lateral occipital
Temporal lobe	Superior temporal, middle temporal, inferior temporal, transverse temporal, banks of the superior temporal sulcus
Parietal lobe	Postcentral, superior parietal, supramarginal, inferior parietal
Prefrontal cortex	Superior frontal, rostral , caudal middle frontal, pars opercularis/orbitalis/triangularis
Angular gyrus, between angular and supramarginal gyrus	Supramarginal gyrus

**Supplementary Information 5: All articles included in the systematic review “Localization Patterns of Language Errors during Direct Electrical Brain Stimulation: A Systematic Review”**

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- Altieri, R., Melcarne, A., Junemann, C., Zeppa, P., Zenga, F., Garbossa, D., . . . Barbagallo, G. (2019). Inferior Fronto-Occipital fascicle anatomy in brain tumor surgeries: From anatomy lab to surgical theater. *J Clin Neurosci*, 68, 290-294. doi:10.1016/j.jocn.2019.07.039
- Barone, F., Alberio, N., Iacopino, D. G., Giammalva, G. R., D'arrigo, C., Tagnese, W., . . . Maugeri, R. (2018). Brain mapping as helpful tool in brain glioma surgical treatment—Toward the “perfect surgery”? *Brain Sci*, 8(11). doi:10.3390/brainsci8110192
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- Benzagmout, M., Gatignol, P., & Duffau, H. (2007). Resection of world health organization grade II gliomas involving Broca's area: Methodological and functional considerations. *NEUROSURGERY*, 61(4), 741-752. doi:10.1227/01.Neu.0000298902.69473.77
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- Bresson, D., Madadaki, C., Poisson, I., Habas, C., & Mandonnet, E. (2013). Functionally tailored transcortical approach of deep-seated lesions: An alternative to the transulcal approach? A technical case report. *Neurochirurgie*, 59(4-5), 159-162. doi:10.1016/j.neuchi.2013.08.002
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- Chan-Seng, E., Moritz-Gasser, S., & Duffau, H. (2014). Awake mapping for low-grade gliomas involving the left sagittal stratum: Anatomofunctional and surgical considerations: Clinical article. *J Neurosurg*, 120(5), 1069-1077. doi:10.3171/2014.1.Jns132015
- Chernoff, B. L., Sims, M. H., Smith, S. O., Pilcher, W. H., & Mahon, B. Z. (2019). Direct electrical stimulation of the left frontal aslant tract disrupts sentence planning without affecting articulation. *CogN*, 36(3-4), 178-192.
- Corriveau, F., de Schotten, M. T., Poisson, I., Froelich, S., Descoteaux, M., Rheault, F., & Mandonnet, E. (2019). Dissociating motor–speech from lexico-semantic systems in the left frontal lobe: insight from a series of 17 awake intraoperative mappings in glioma patients. *Brain Struct Funct*, 224(3), 1151-1165. doi:10.1007/s00429-019-01827-7
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- De Witte, E., Satoer, D., Robert, E., Colle, H., Verheyen, S., Visch-Brink, E., & Mariën, P. (2015). The Dutch Linguistic Intraoperative Protocol: A valid linguistic approach to awake brain surgery. *BRAIN LANG*, 140, 35-48. doi:10.1016/j.bandl.2014.10.011
- Duffau, H. (2009). A personal consecutive series of surgically treated 51 cases of insular WHO Grade II glioma: Advances and limitations - Clinical article. *J Neurosurg*, 110(4), 696-708.

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- the right pars opercularis with electrical stimulation frees the song: Case report. *J Neurosurg*, 123(6), 1401-1404. doi:10.3171/2014.12.Jns141829
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