

Primo corso di perfezionamento:
Neurologia Cognitiva

 International School
of Neurological Sciences
Sezione di Neuro-oncologia, Epilessia e Nutrizione

International School University
ISOLA SAN SERVOLO - VENEZIA

27/28
Aprile 2018

 Regione Lombardia  Fondazione I.R.C.C.S. Istituto Neurologico Carlo Besta  IRE Roma FISS  International School of Neurological Sciences
Sezione di Neuro-oncologia, Epilessia e Nutrizione



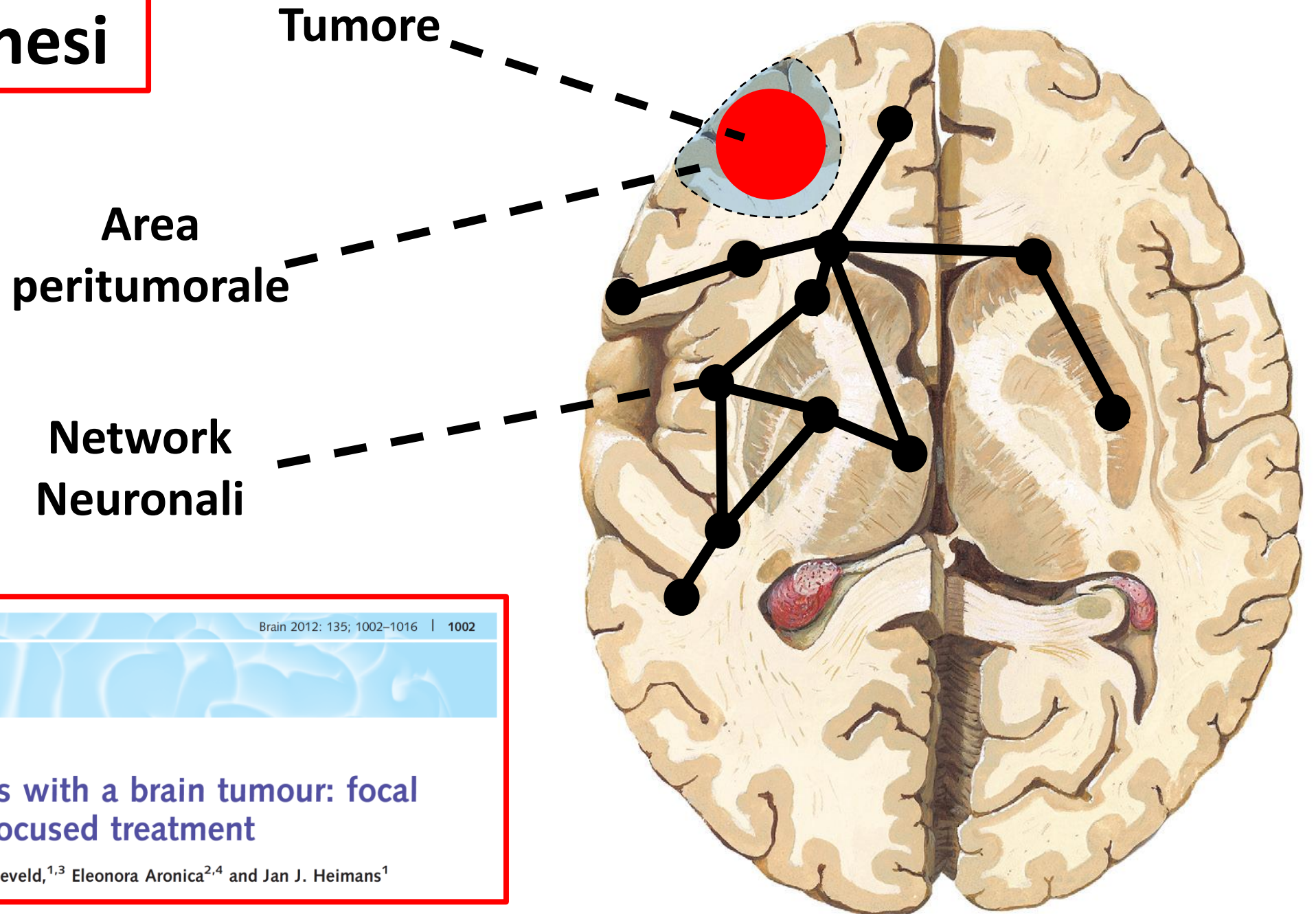
Epilessia e tumori cerebrali
Nuovi approcci chirurgici: quale margine
per un risparmio cognitivo
Vincenzo Esposito e molti altri
Dipartimento di Neuroscienze «Giampaolo Cantore»
I.R.C.C.S. Neuromed - Pozzilli (IS)

NEUROMED
I.R.C.C.S.  ISTITUTO
NEUROLOGICO
MEDITERRANEO



 **SAPIENZA**
UNIVERSITÀ DI ROMA

Epilettogenesi



doi:10.1093/brain/awr310

Brain 2012; 135; 1002–1016 | 1002

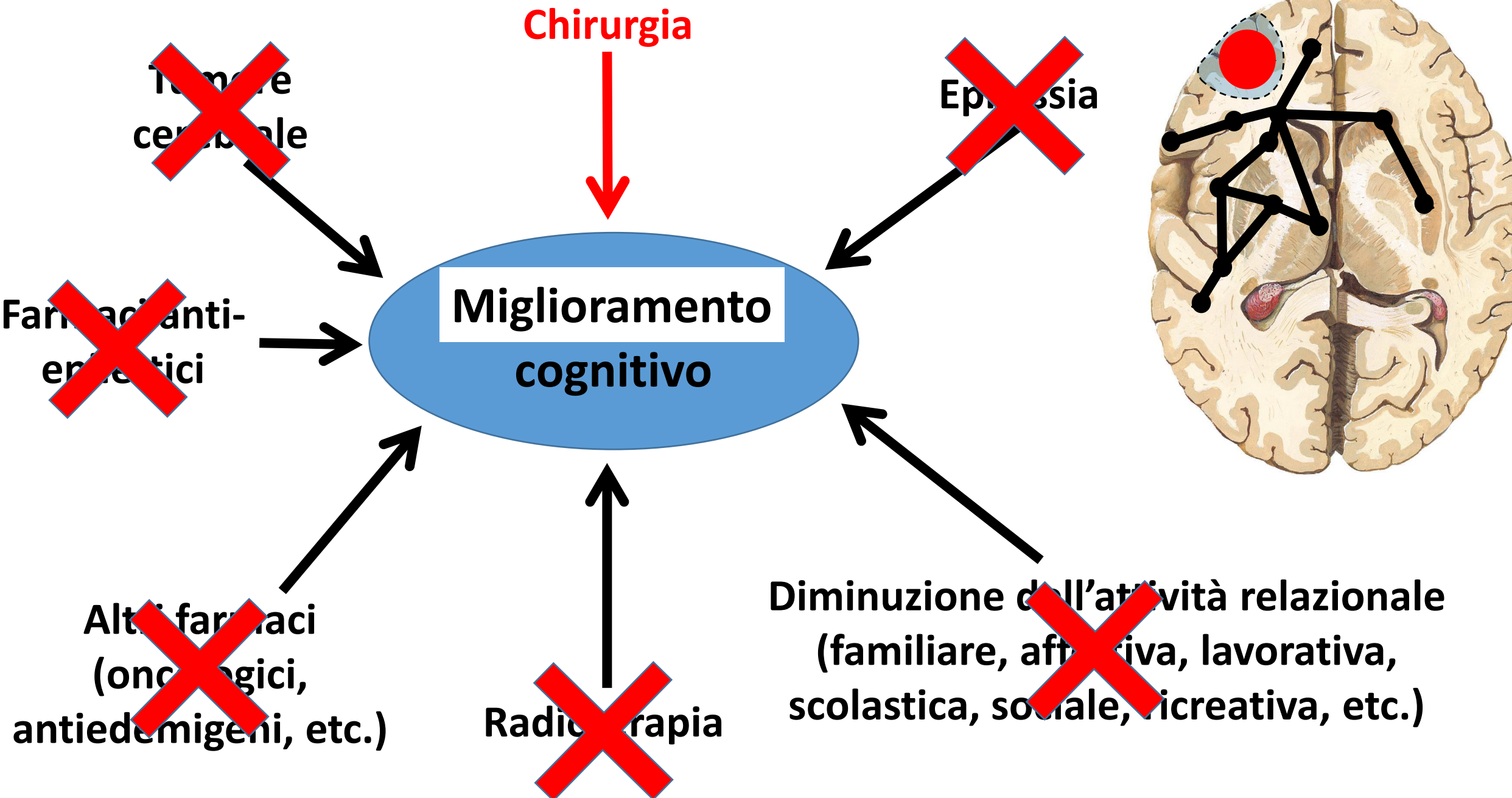
BRAIN

A JOURNAL OF NEUROLOGY

REVIEW ARTICLE

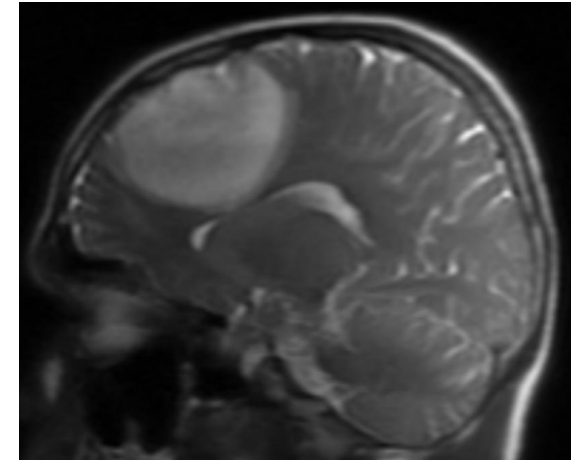
Epilepsy in patients with a brain tumour: focal epilepsy requires focused treatment

Marjolein de Groot,^{1,2} Jaap C. Reijneveld,^{1,3} Eleonora Aronica^{2,4} and Jan J. Heimans¹

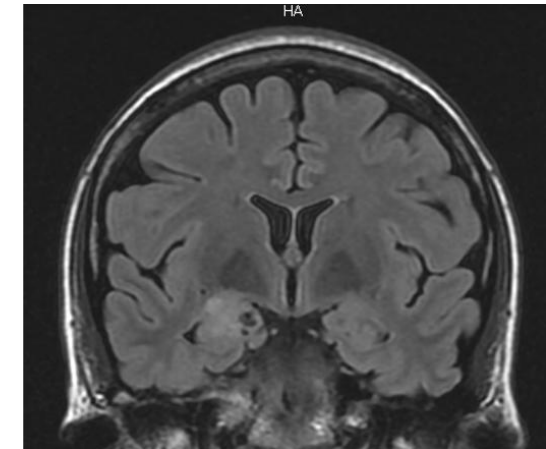


2 tipi di tumori associati ad epilessia

Gliomi: evolutività oncologica, localizzazioni varie. In genere non necessitano di valutazione in un centro dedicato alla Chirurgia dell'Epilessia. L'asportazione radicale del tumore ha un'ottima prognosi sull'epilessia.



LEAT (Long-Term Epilepsy Associated Tumors) o epileptomi: tumori intrinsecamente epilettogeni tipici della giovane età, quasi sempre nel lobo temporale (circa 80%), con scarsa evolutività oncologica. Valutazione in un centro per la Chirurgia dell'Epilessia (non infrequentemente l'asportazione va estesa oltre il tumore per risolvere l'epilessia).



Acta Neuropathol (2014) 128:39–54
DOI 10.1007/s00401-014-1288-9

REVIEW

A neuropathology-based approach to epilepsy surgery in brain tumors and proposal for a new terminology use for long-term epilepsy-associated brain tumors

Ingmar Blumcke · Eleonora Aronica · Horst Urbach ·
Andreas Alexopoulos · Jorge A. Gonzalez-Martinez

1 – Gliomi cerebrali

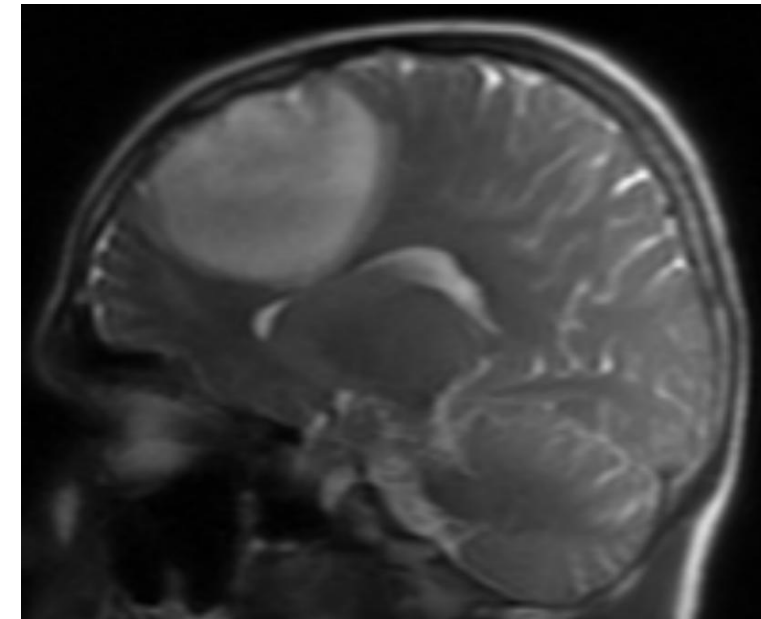
Tumori con evolutività oncologica

Localizzazioni varie.

In genere non vengono valutati in un centro dedicato alla Chirurgia dell'Epilessia

- le considerazioni oncologiche prevalgono di solito su quelle epilettologiche
- la storia di epilessia è in genere di breve durata (epilettogenesi secondaria rara)
- alta probabilità che il tumore coincida con la zona epilettogena

L'asportazione radicale del tumore ha un'ottima prognosi sull'epilessia.



European Association for Neuro-Oncology (EANO) guideline on the diagnosis and treatment of adult astrocytic and oligodendroglial gliomas



Michael Weller, Martin van den Bent, Jörg C Tonn, Roger Stupp, Matthias Preusser, Elizabeth Cohen-Jonathan-Moyal, Roger Henriksson, Emilie Le Rhun, Carmen Balana, Olivier Chinot, Martin Bendszus, Jaap C Reijneveld, Frederick Dhermain, Pim French, Christine Marosi, Colin Watts, Ingela Oberg, Geoffrey Pilkington, Brigitta G Baumert, Martin J B Taphoorn, Monika Hegi, Manfred Westphal, Guido Reifenberger, Riccardo Soffetti, Wolfgang Wick, for the European Association for Neuro-Oncology (EANO) Task Force on Gliomas

- 4300 nuovi casi di tumore cerebrale/anno
- Incidenza dei gliomi (incluso il glioblastoma): 6 casi/100000/anno
- Causa del 7%/anno dei decessi per tumore in popolazione < 70 anni

Guidelines on management of low-grade gliomas: report of an EFNS–EANO* Task Force

R. Soffietti^a, B.G. Baumert^b, L. Bello^c, A. von Deimling^d, H. Duffau^e, M. Frénay^f, W. Grisold^g, R. Grant^h, F. Grausⁱ, K. Hoang-Xuan^j, M. Klein^k, B. Melin^l, J. Rees^m, T. Siegalⁿ, A. Smits^o, R. Stupp^p and W. Wick^q

- Surgery is **necessary to provide tissue** for distinguishing between the histologic types, grading the malignancy and assessing the molecular status of tumors. Moreover, there are scenarios that pose problems of differential diagnosis between LGGs and non-neoplastic lesions (demyelination, inflammation or infection), and thus histological verification is mandatory.
- Total resection **improves seizure control**
- Total/near total resection **decreases the incidence of recurrence and the risk of malignant transformation** and improves progression free survival and overall survival
- Total resection is achieved in no more than 36% of patients
- **The risk of deferring surgery includes managing at a later timepoint a larger tumor, which may have undergone anaplastic transformation**

C. Watts^{*†}, S.J. Price^{*}, T. Santarius^{*}

** University of Cambridge, Department of Clinical Neurosciences, Division of Neurosurgery, Addenbrooke's Hospital, Cambridge, UK*

† Department of Clinical Neurosciences, Cambridge Centre for Brain Repair, University of Cambridge, Cambridge, UK

”The **goal of surgery** is to obtain a **histological diagnosis** and, where possible, **remove as much of the tumour as possible without damaging the adjacent healthy brain tissue.**“

Resezione chirurgica

Scopi

Controllo della malattia
Risoluzione delle crisi epilettiche

- Rispetto delle funzioni corticali e sottocorticali
- Rispetto delle strutture anatomiche profonde
- Vascolarizzazione > complicanze ischemiche post-operatorie

Limiti

Corteccia motoria
Aree del linguaggio
Aree visive
Insula
Corpo calloso
Nuclei della base
...

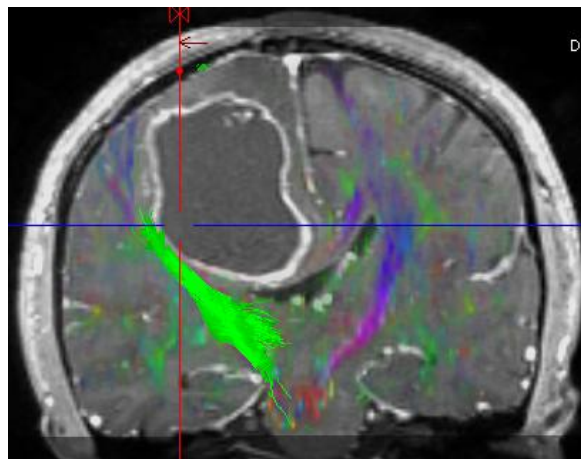
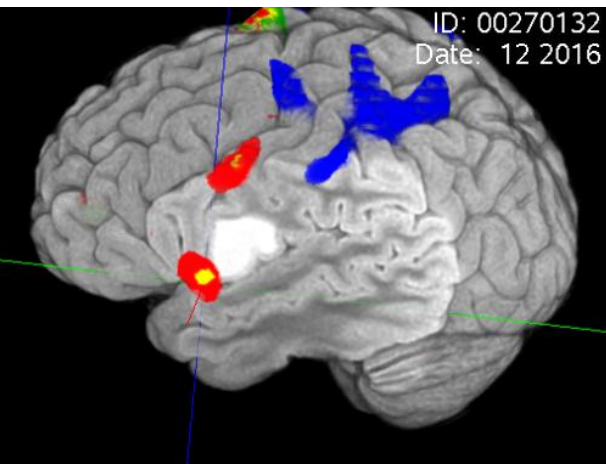
Tratto piramidale
Radiazioni ottiche
IFOF
Fascicolo Arcuato
...

Asportazione massimale: minimizzare le complicanze

Pre-operatorio:



- **Valutazione neuropsicologica estesa**
- RM funzionale
- Trattografia
- RMN 3D



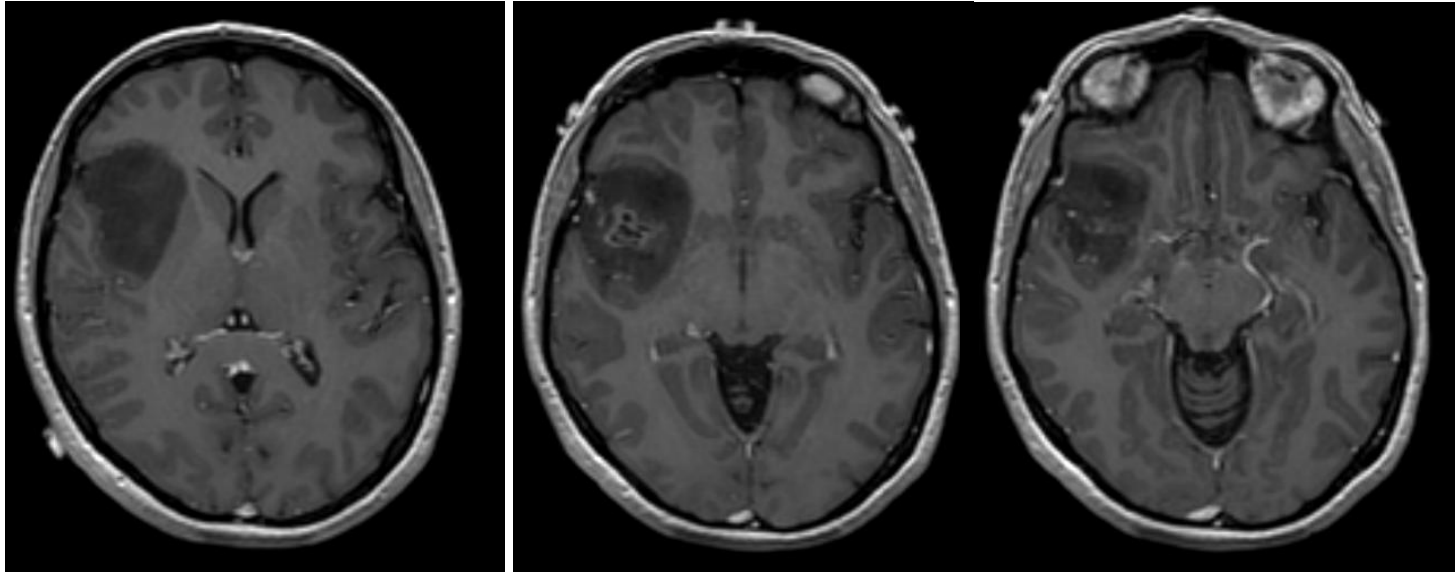
Intra-operatorio:

- Monitoraggio neurofisiologico
- Ricostruzione corticale RMN 3D
- Ecografia intraoperatoria neuronavigata
- Awake surgery (casi selezionati)
- **CUSA-radar** (CUSA con stimolazione sottocorticale continua)
- **Microscopio ad alto ingrandimento, dissezione subpiaie**
- Chirurgia in 2 tempi
- **Esperienza!!**

S. N. 42 y.o., F

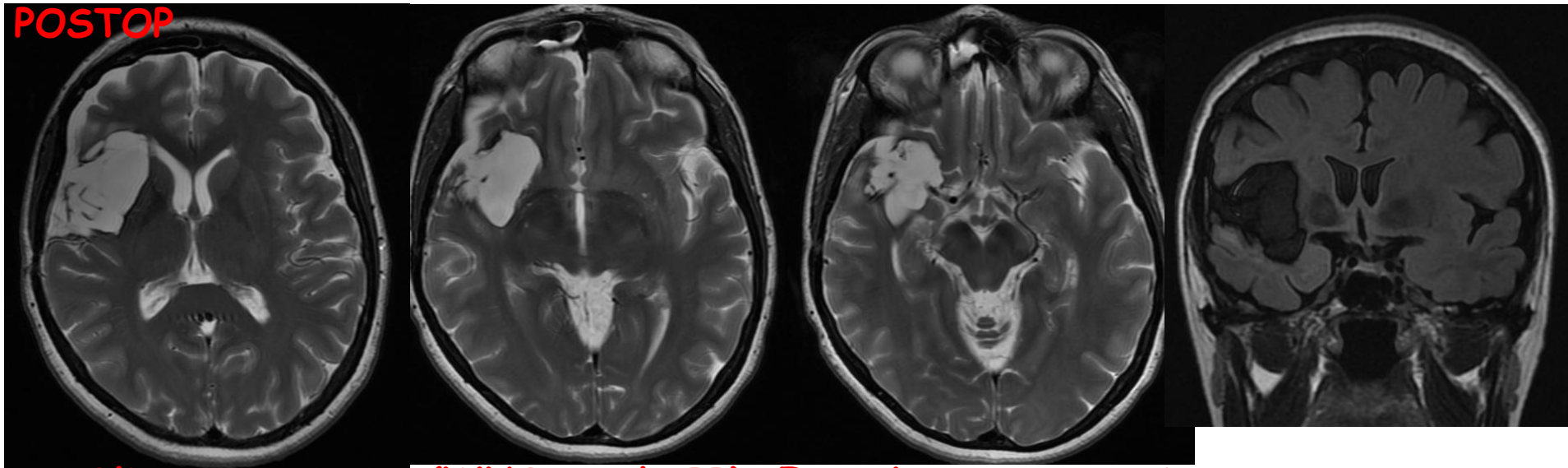
Partial seizures

PREOP



Operata
26.11.2011

POSTOP

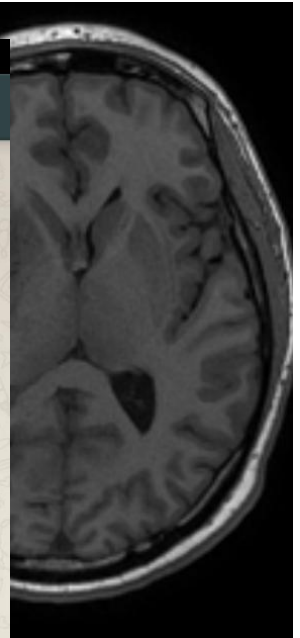
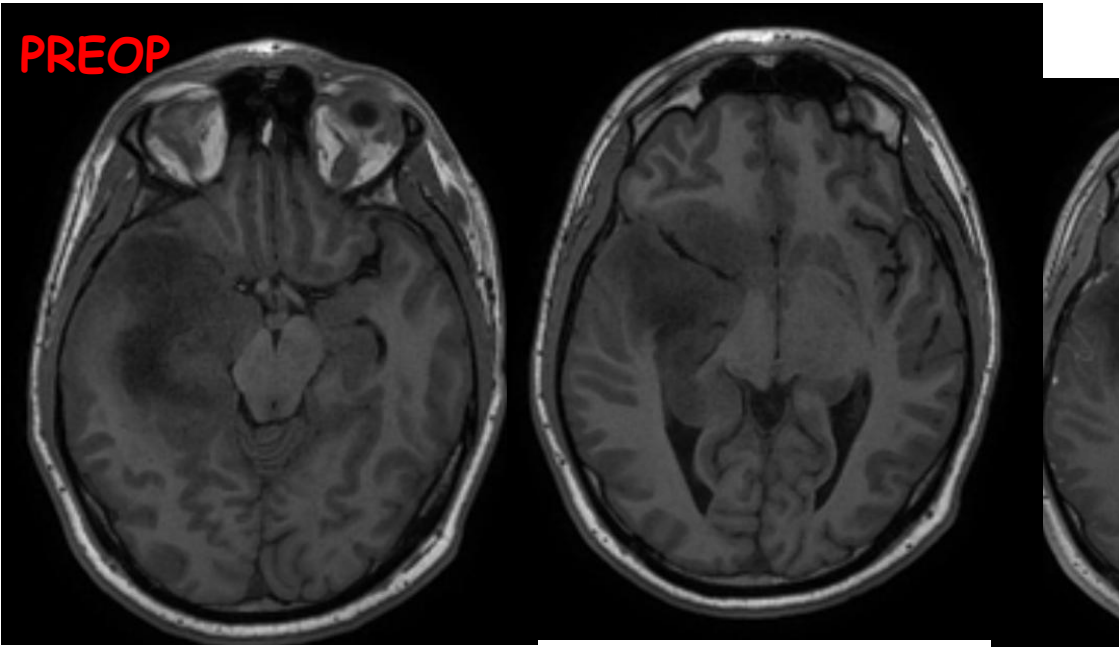


Oligoastrocytoma (WHO grade II), F.U. 6 years, no seizure

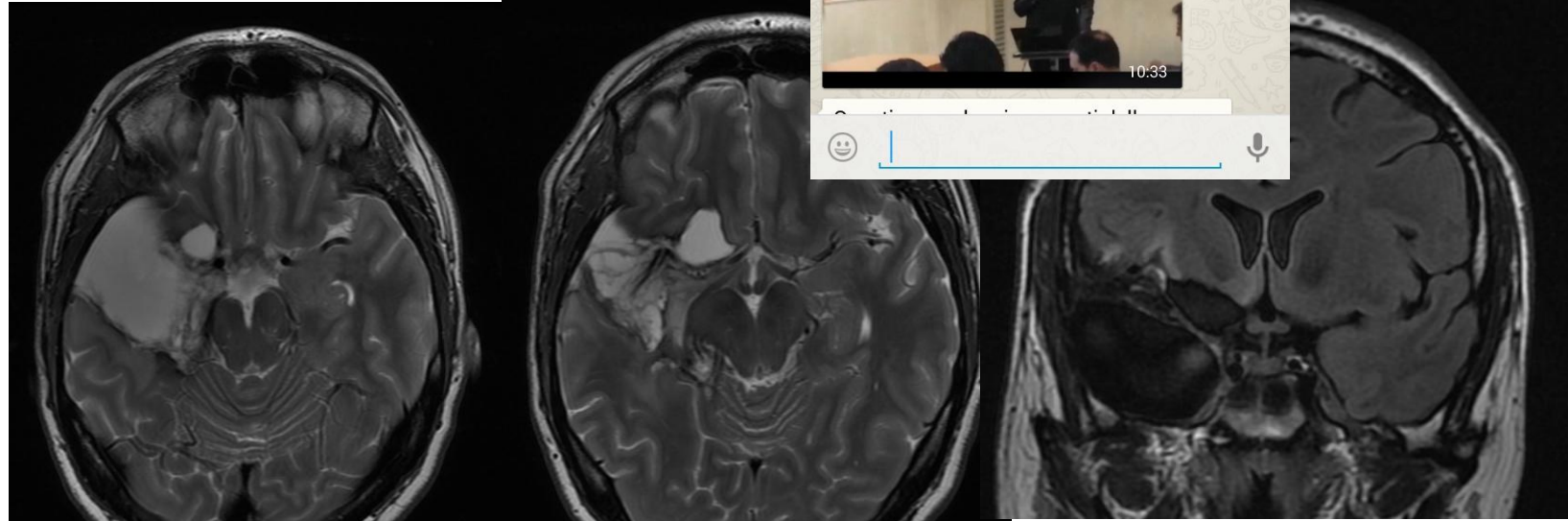
L.S. 28 y.o., M

Seizures

PREOP



POSTOP



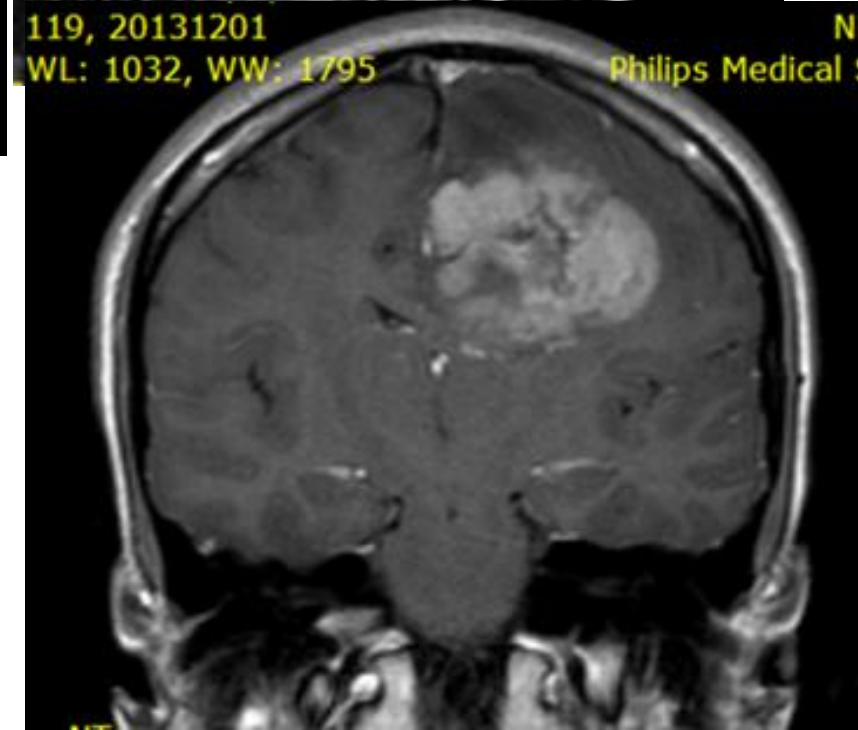
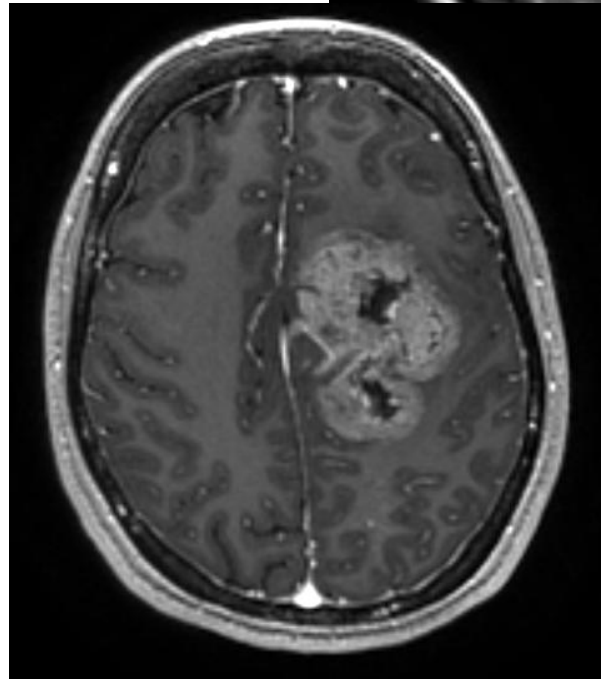
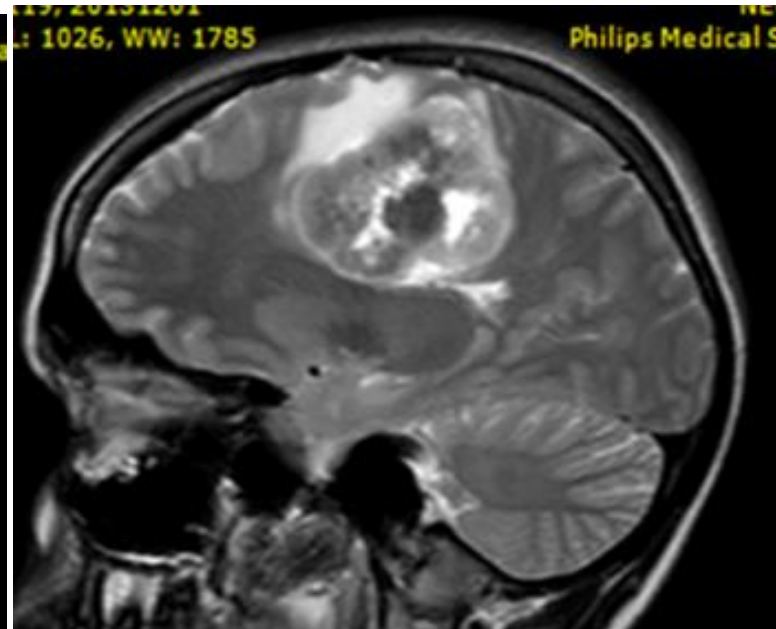
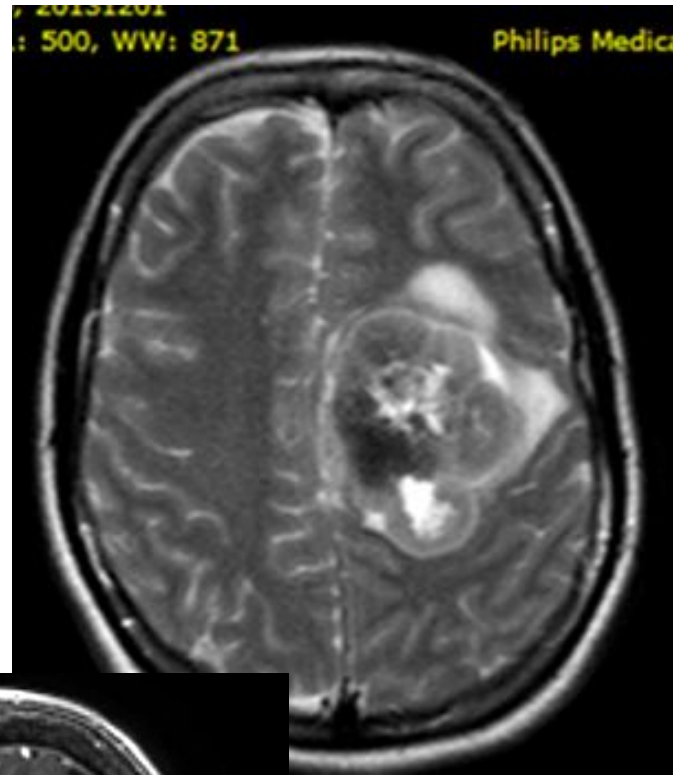
Op.
19.02.2014

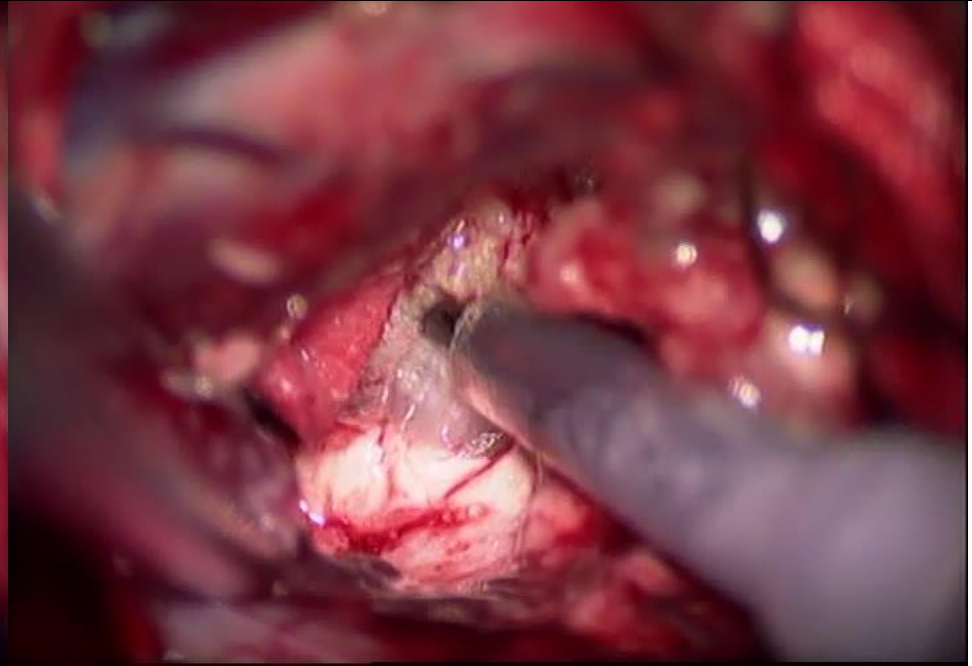
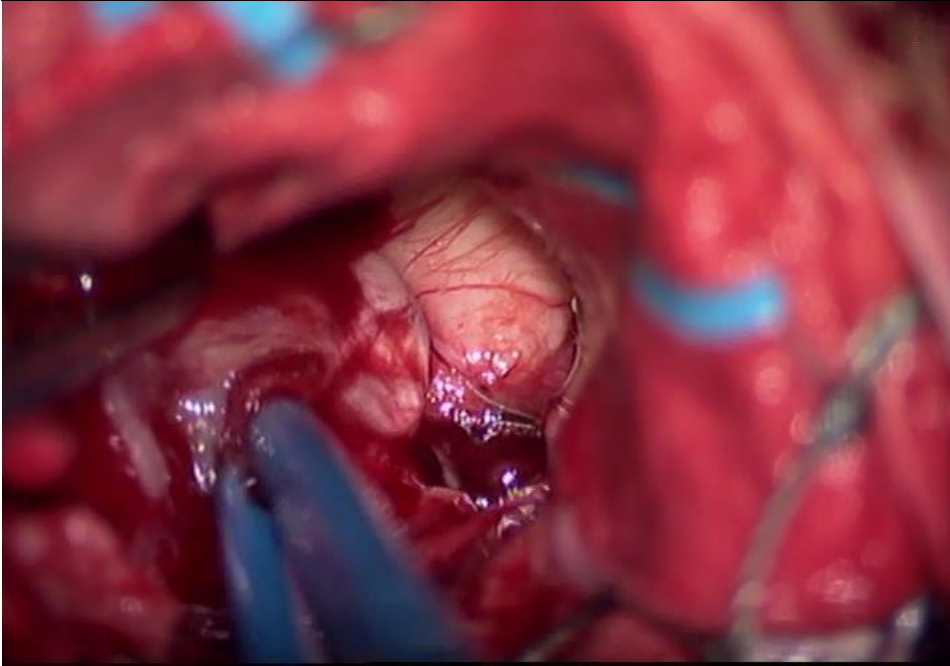
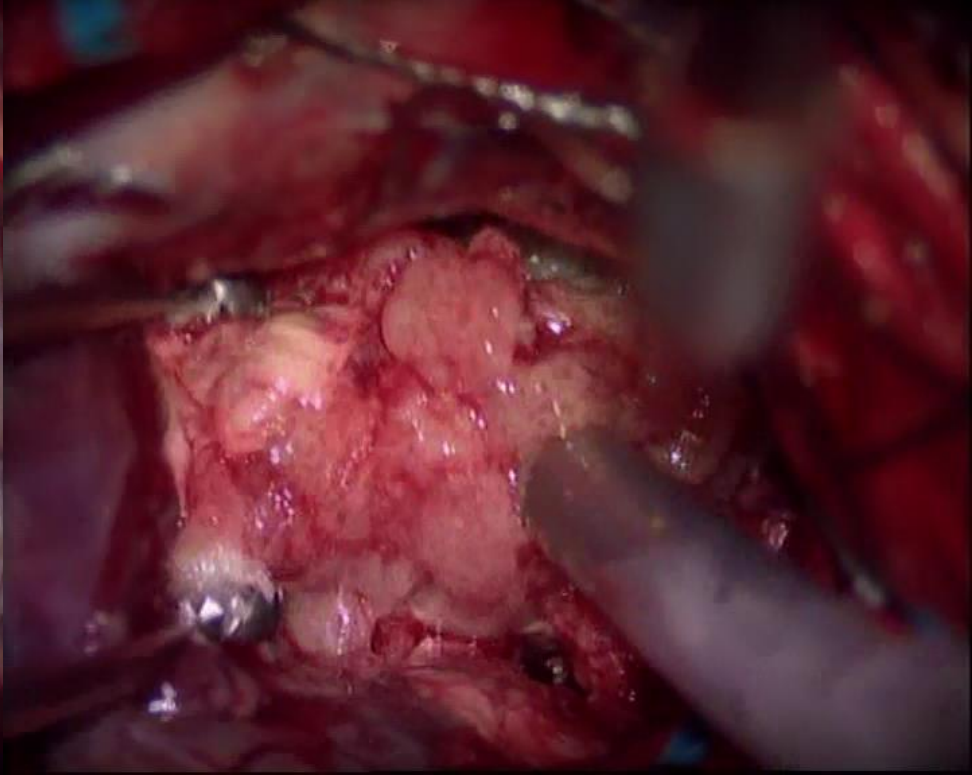
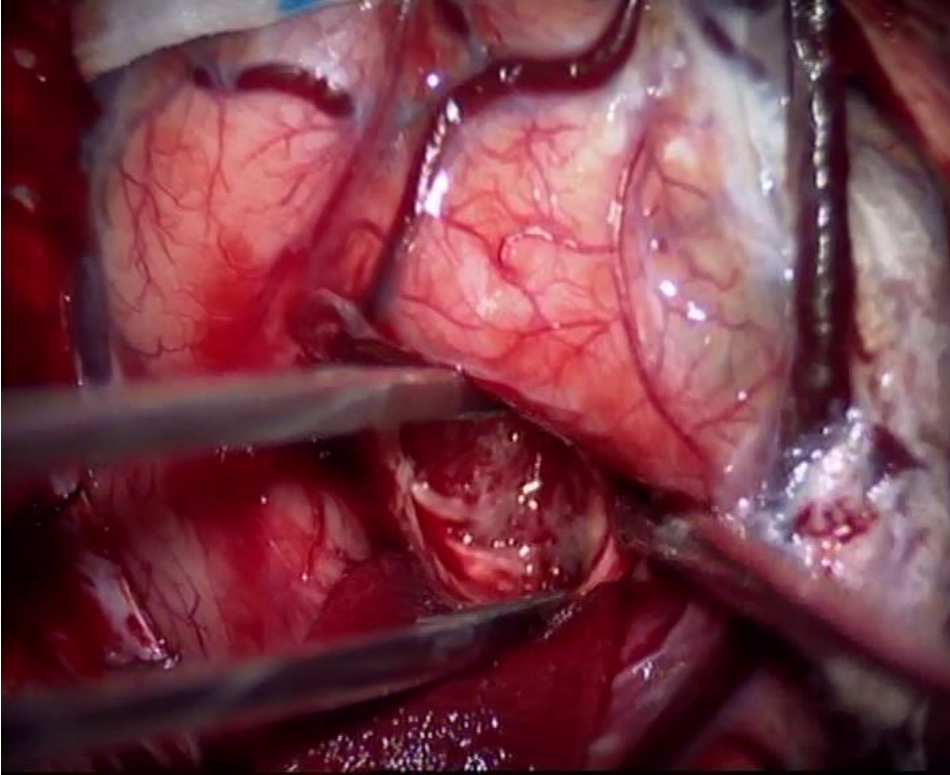
OLIGOASTROCITOMA (WHO grade II), 4 years f.u, no seizures

S.D.P. 18 y.o., F

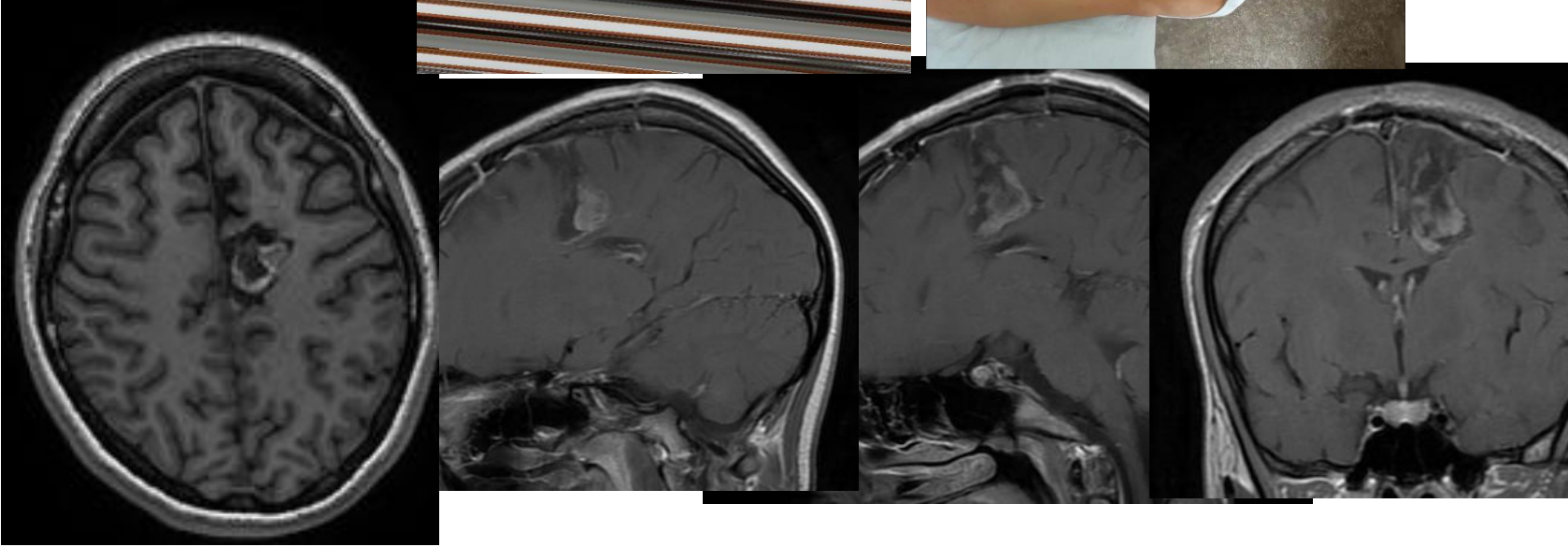
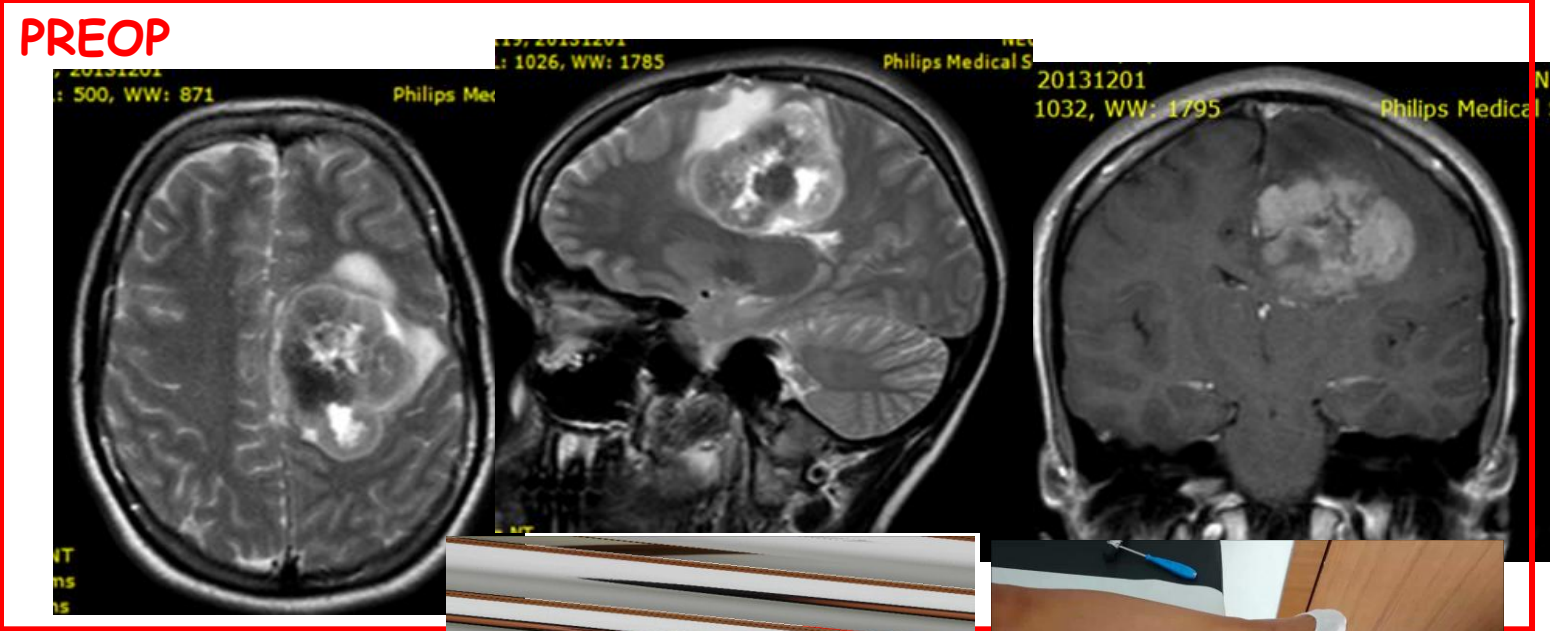
Right hemiparesis and focal motor seizures

PREOP

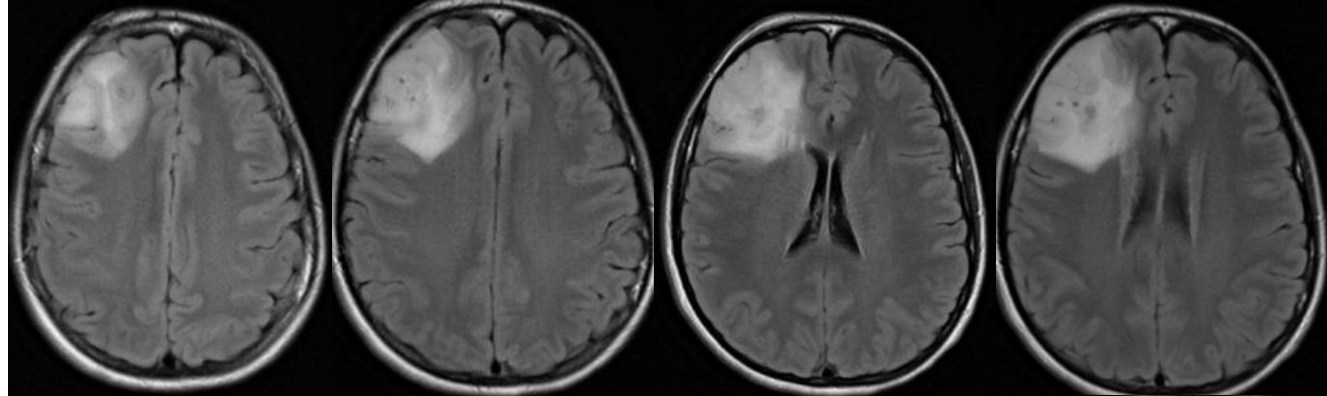




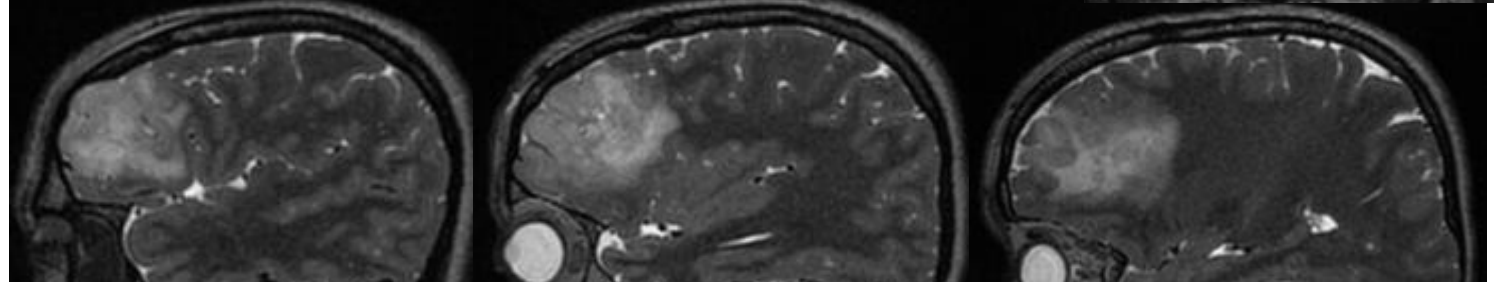
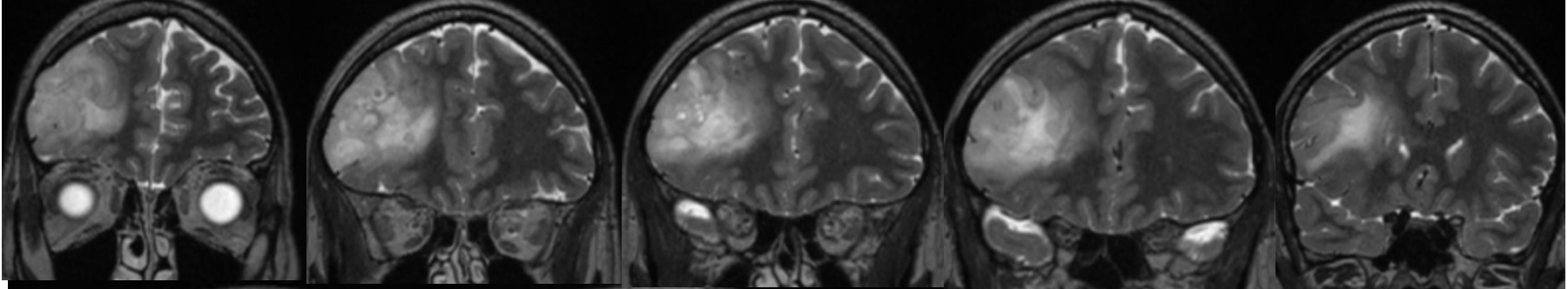
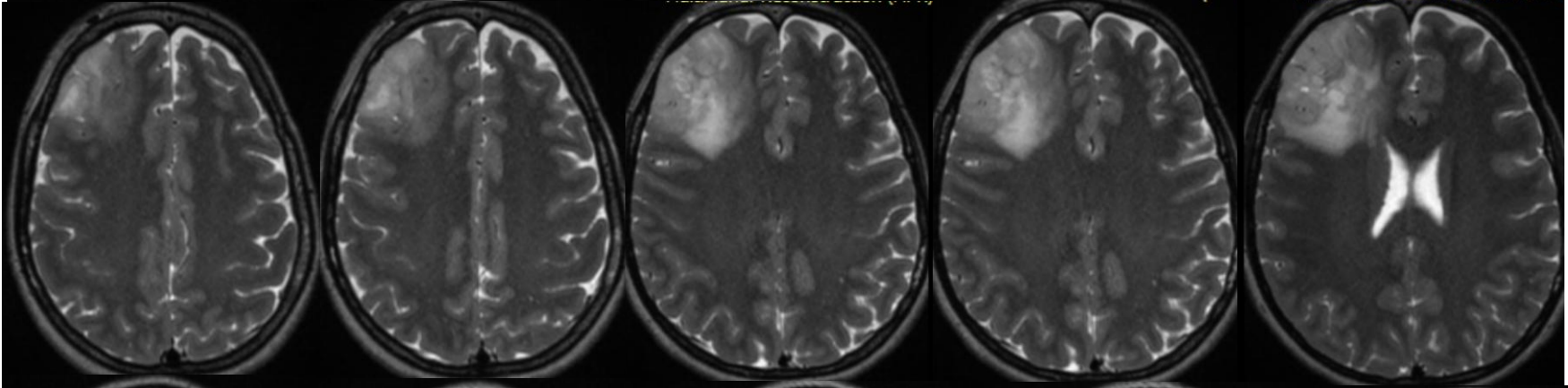
PREOP

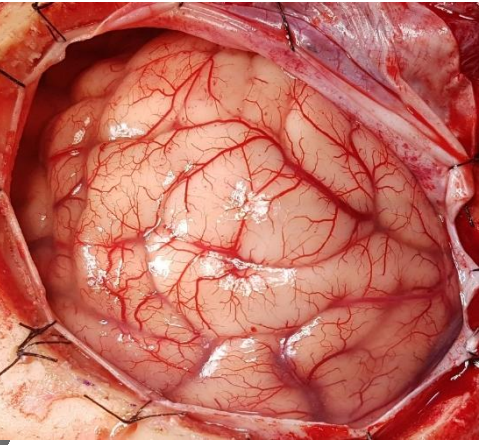
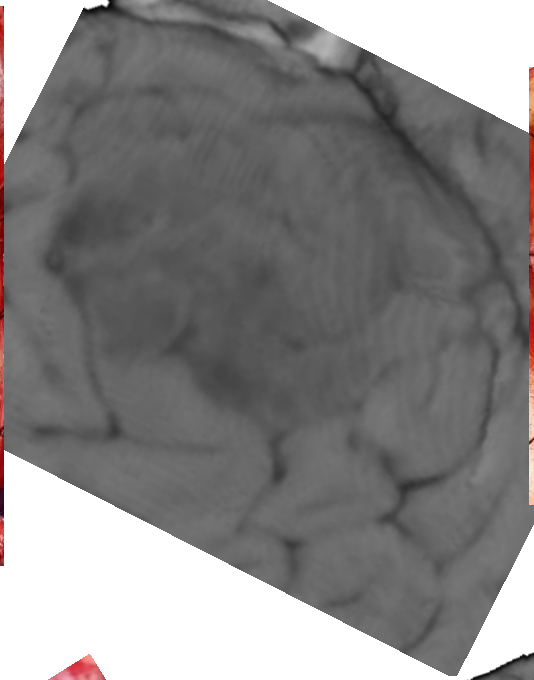
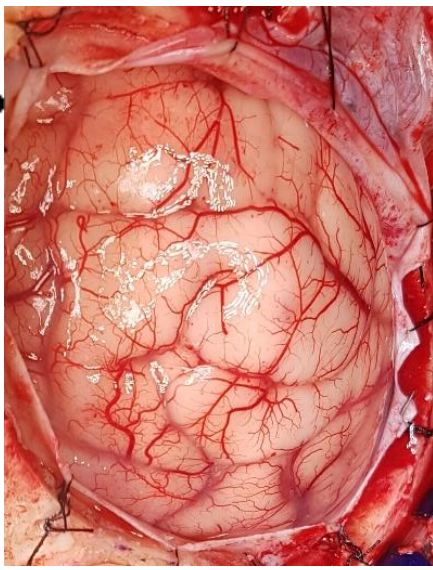
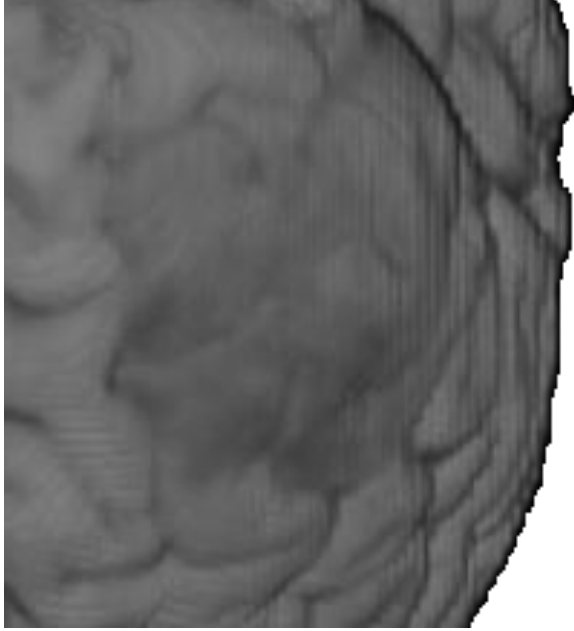


Histology: Astroblastoma, 4 years f.u., no seizures

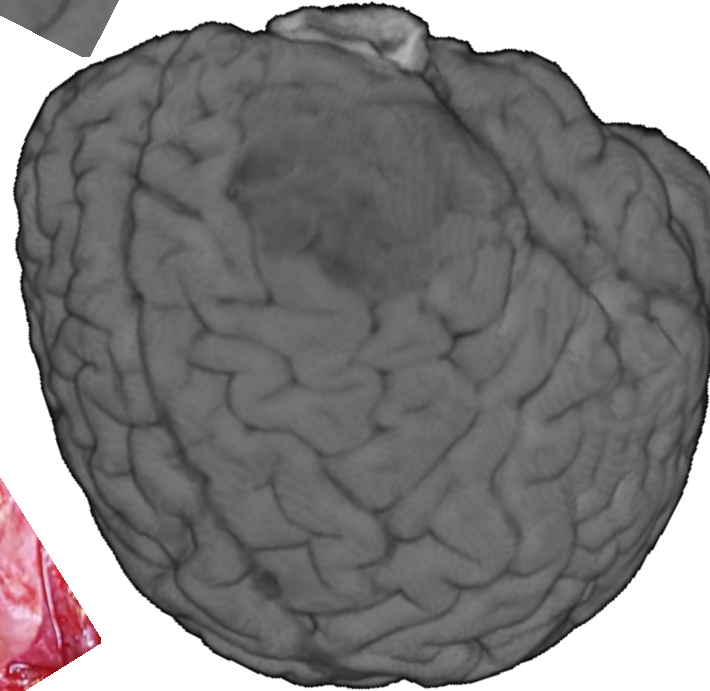
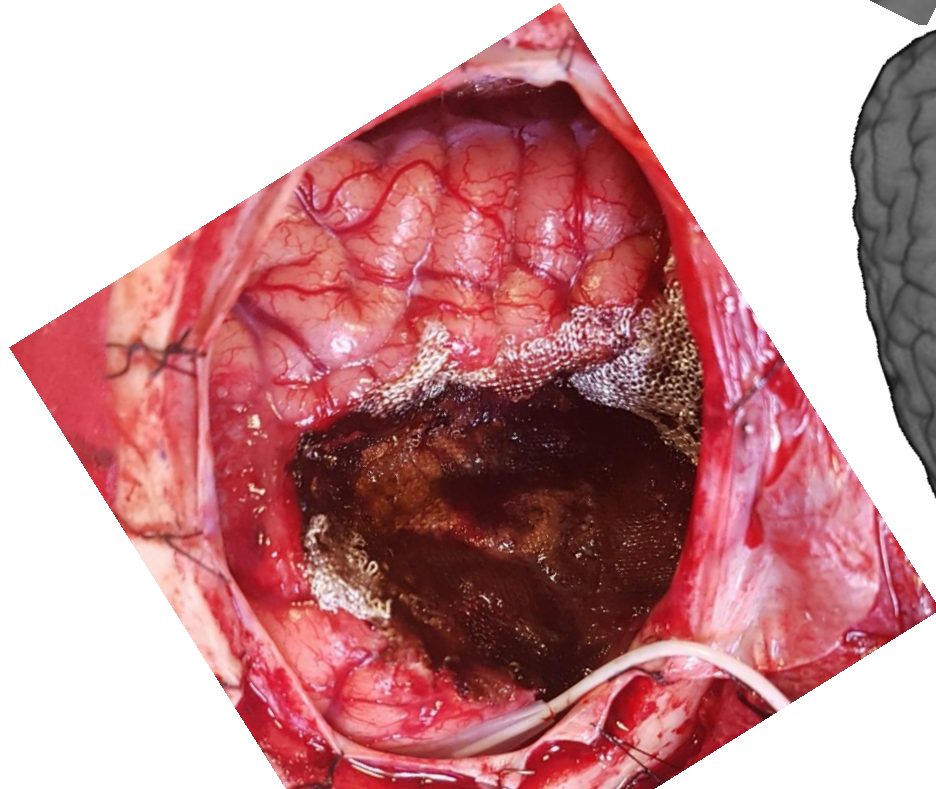
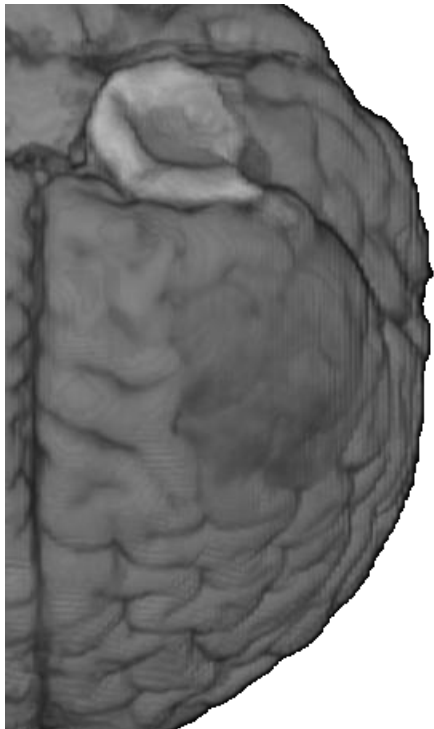


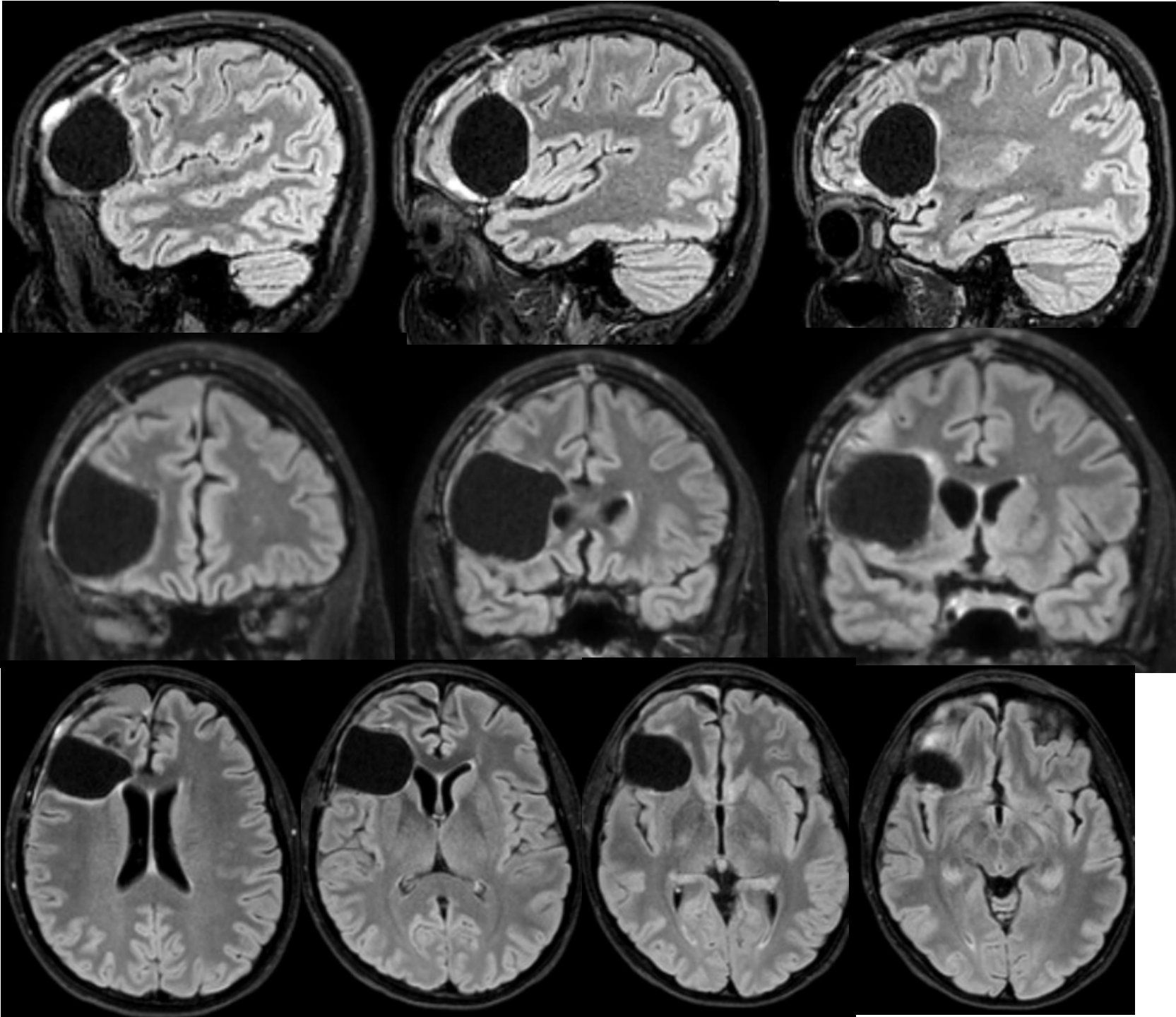
F.M., 36 a., f
2 crisi generalizzate





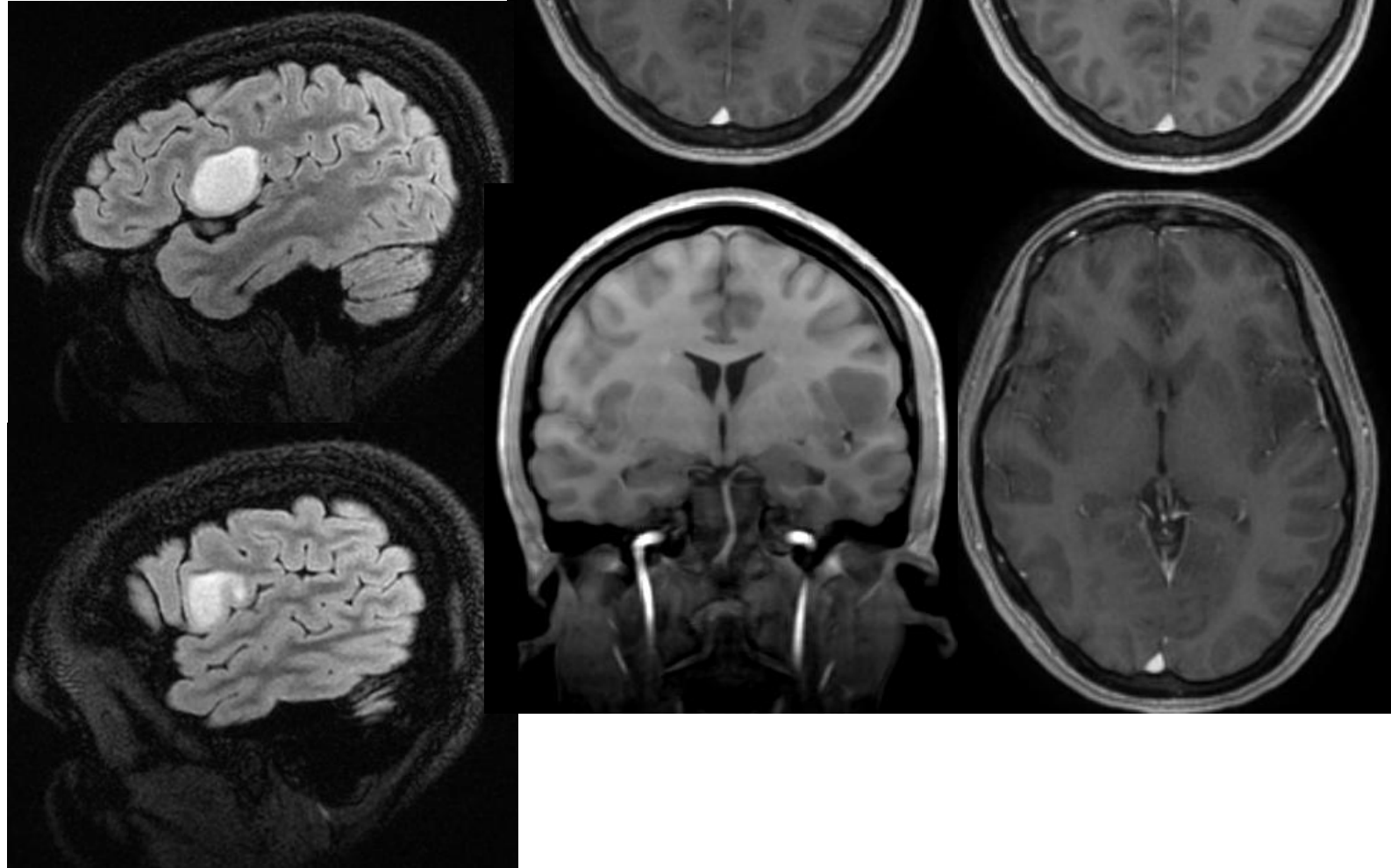
RMN 3D
Tecnica
subpiale

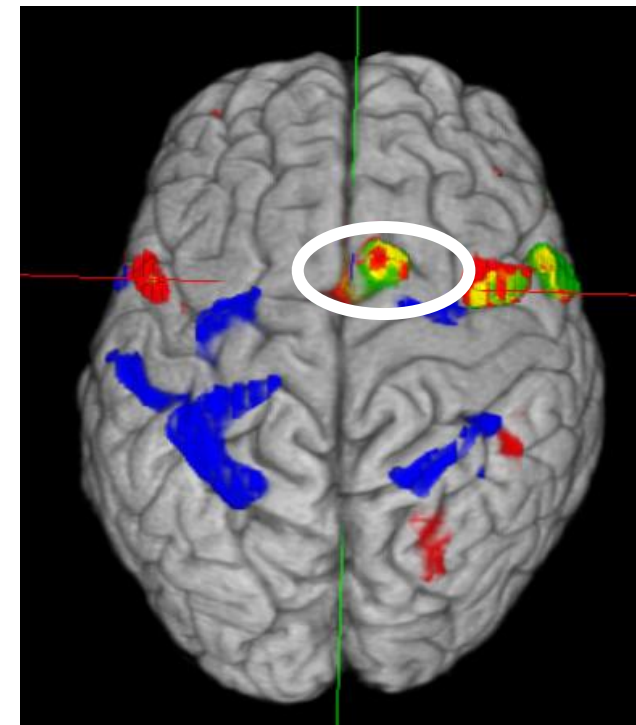
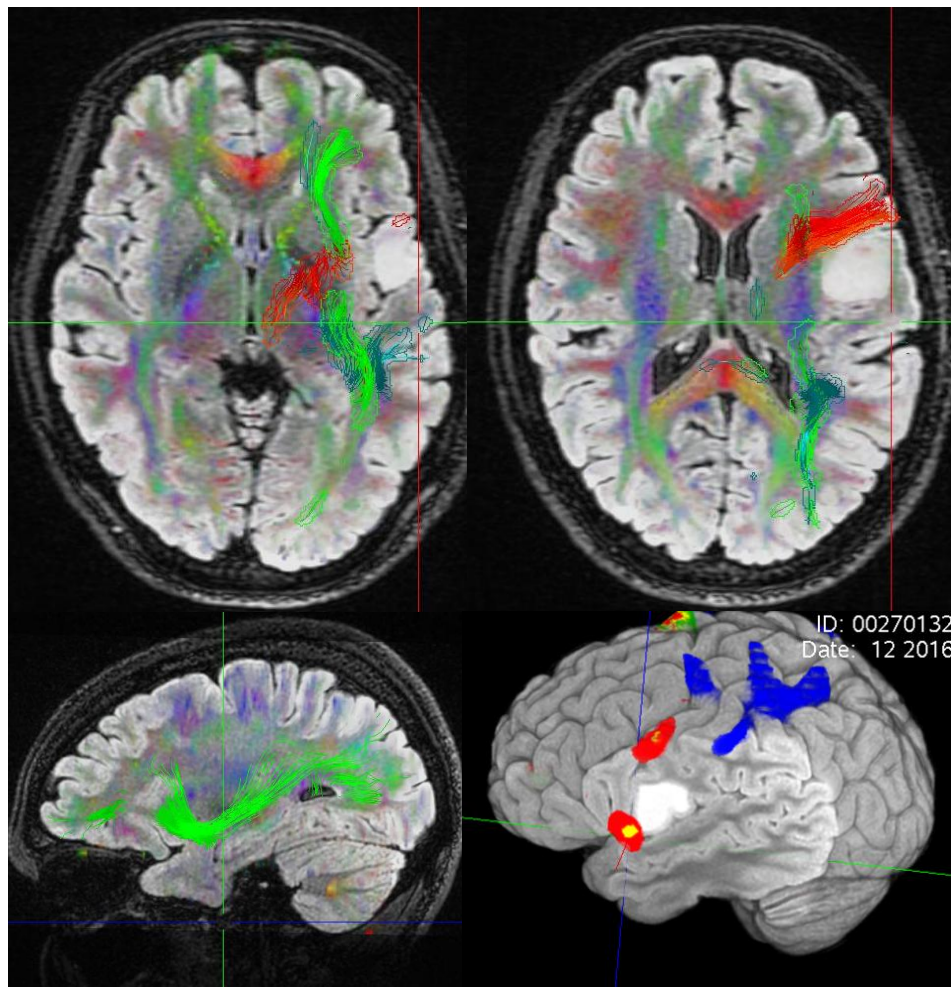




Oligodendroglioma,
WHO Grade II
NO seizures

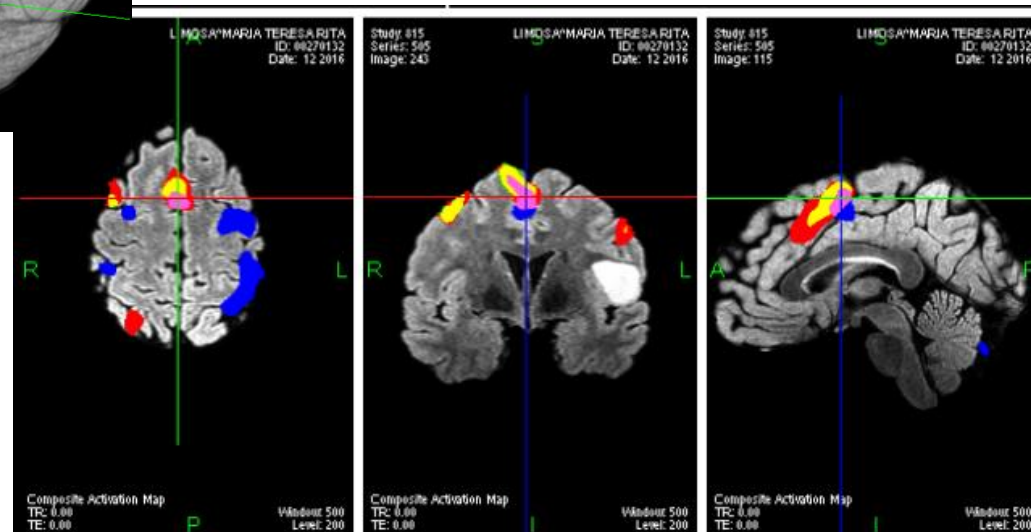
- F, 28 anni
- Crisi epilettiche
- **Ambidestra**



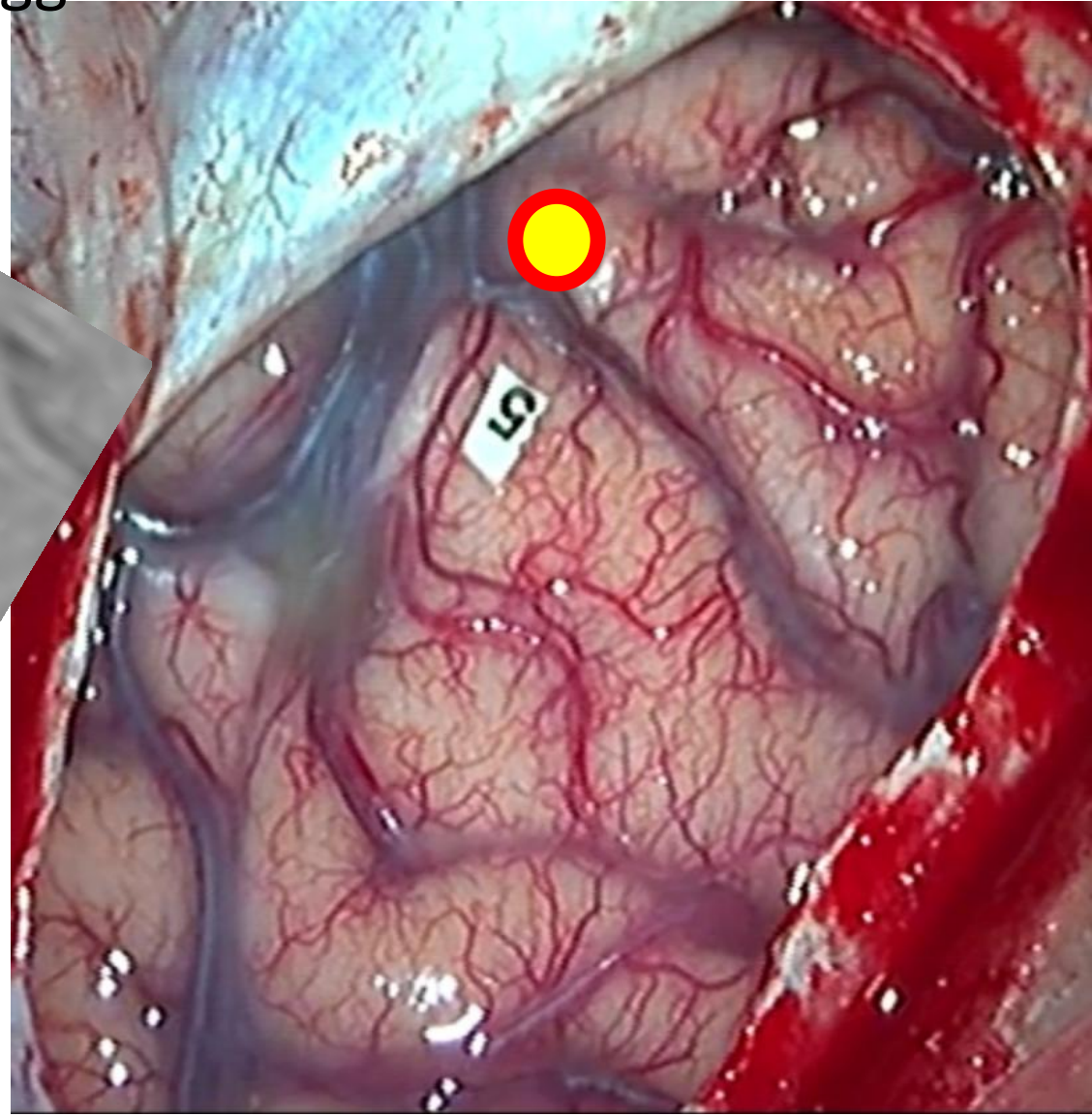
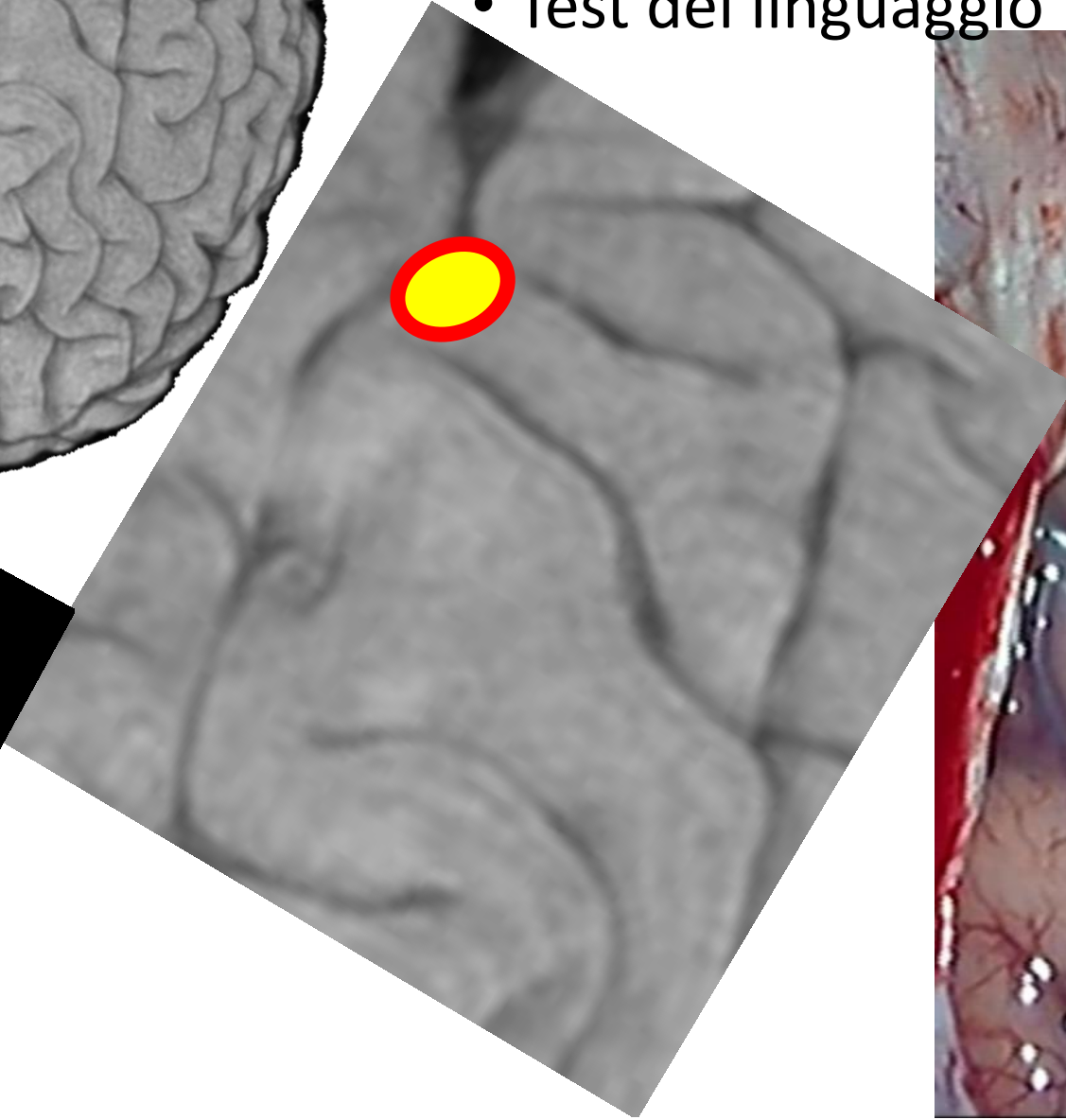
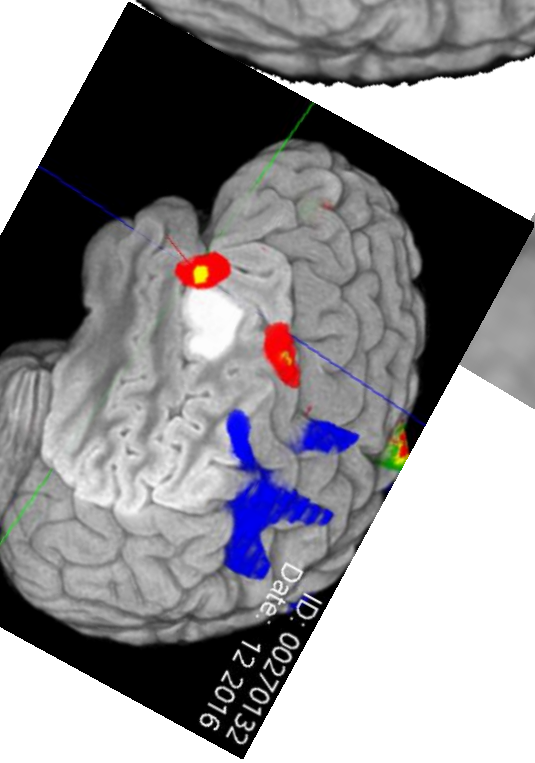
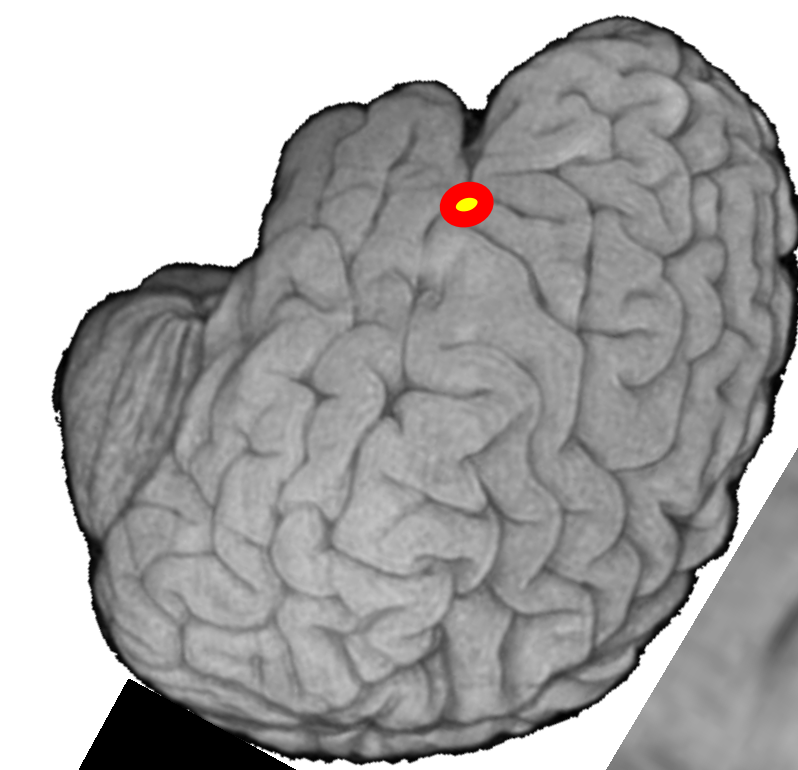


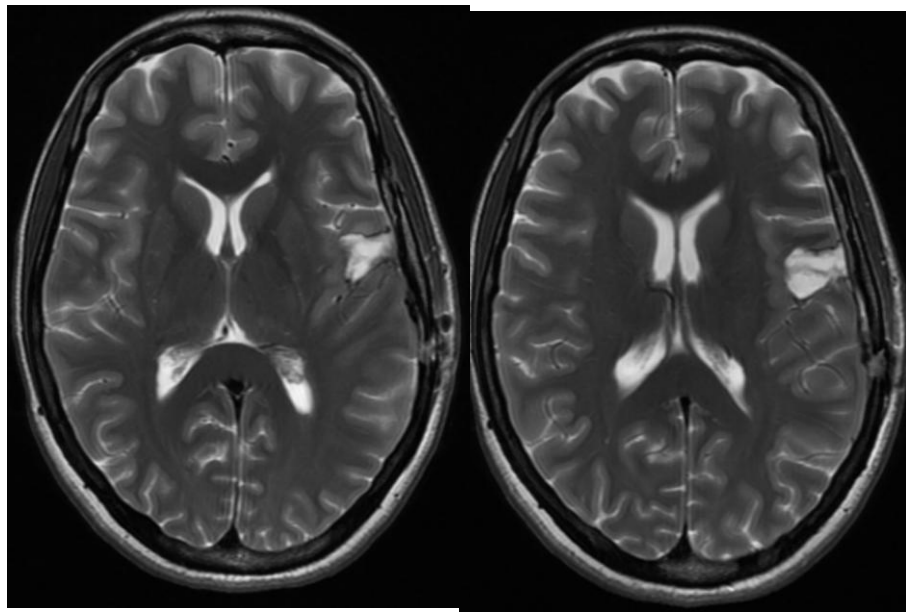
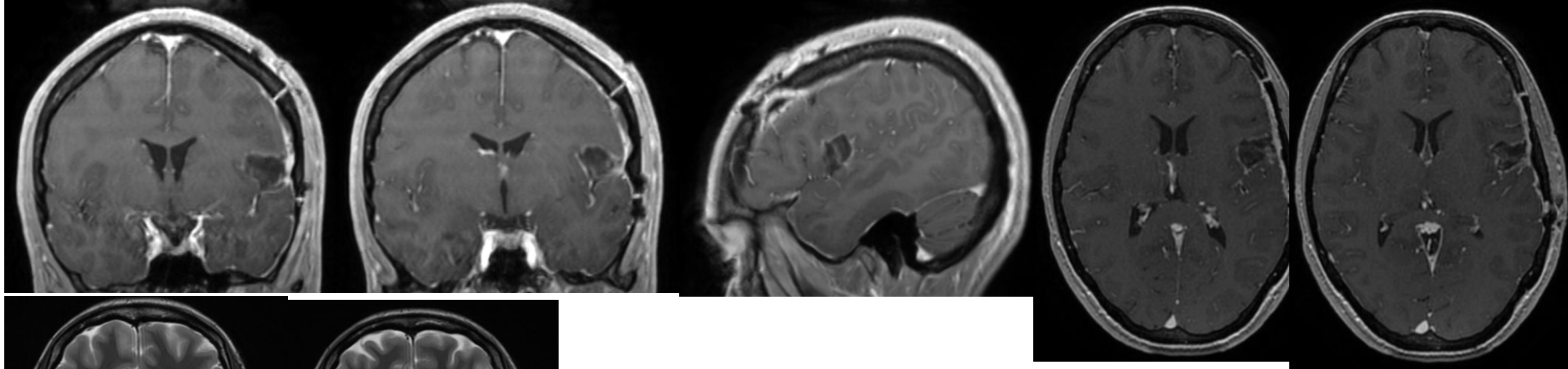
L'fMRI evidenzia una chiara
dominanza emisferica destra.

- 1) Finger tapping
- 2) **Fluenza Semantica**
- 3) **Fluenza Fonemica**
- Finger tapping + Fluenza semantica
- Finger tapping + Fluenza semantica
+ fluenza fonemica



- Awake surgery
- RMN 3D – Tecnica subpiale
- Test del linguaggio



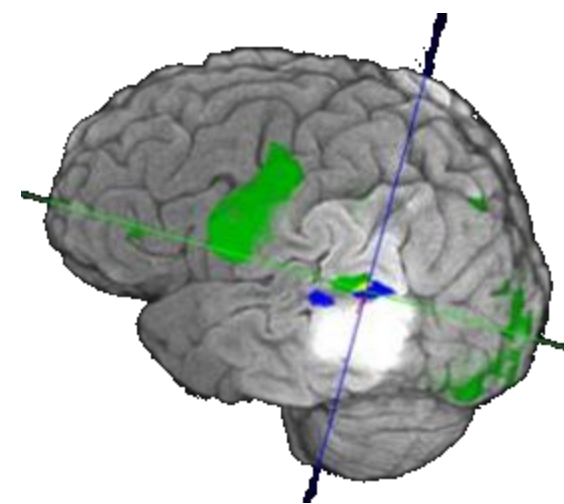
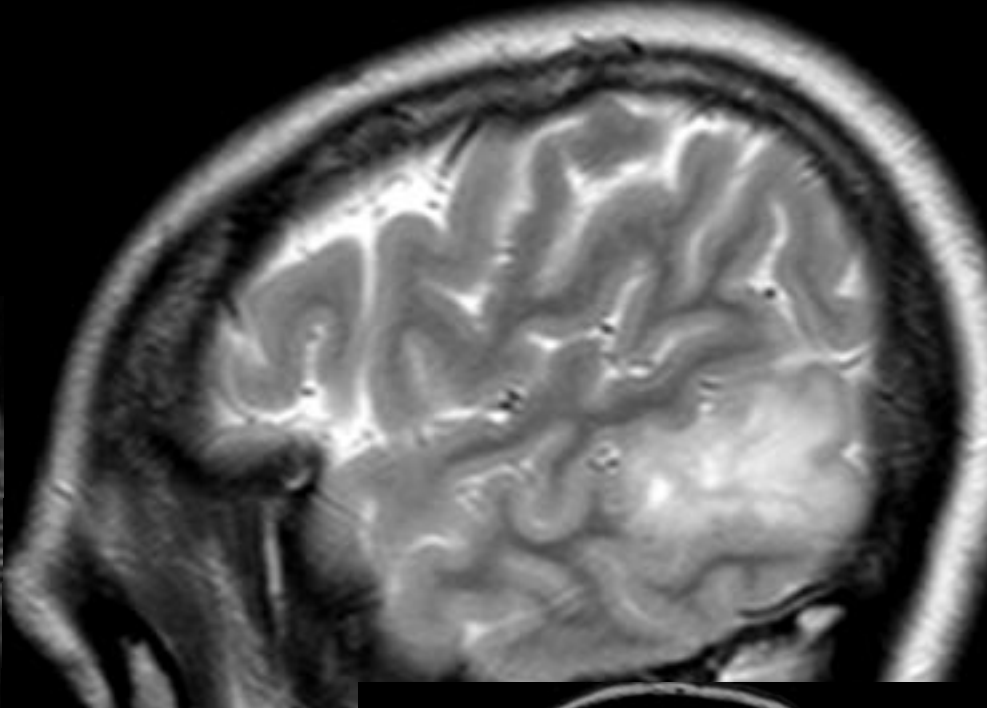
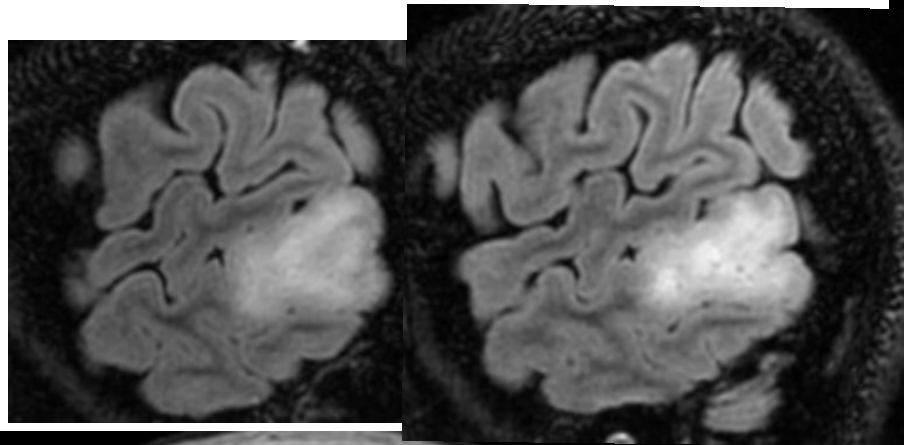


Astrocitoma diffuso grado II WHO

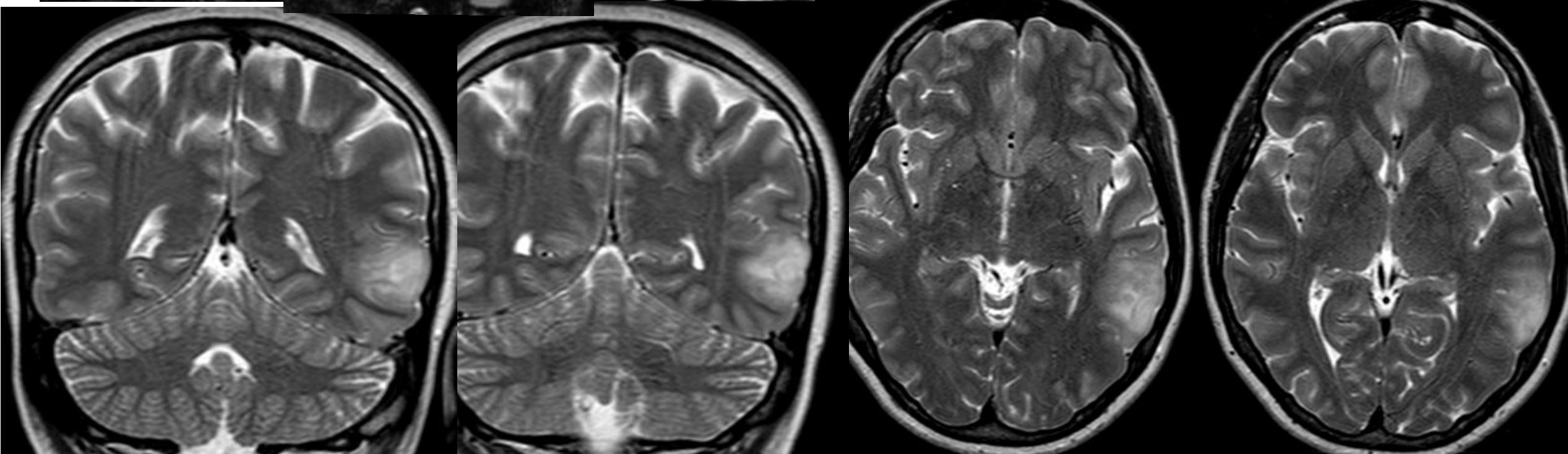
Non crisi

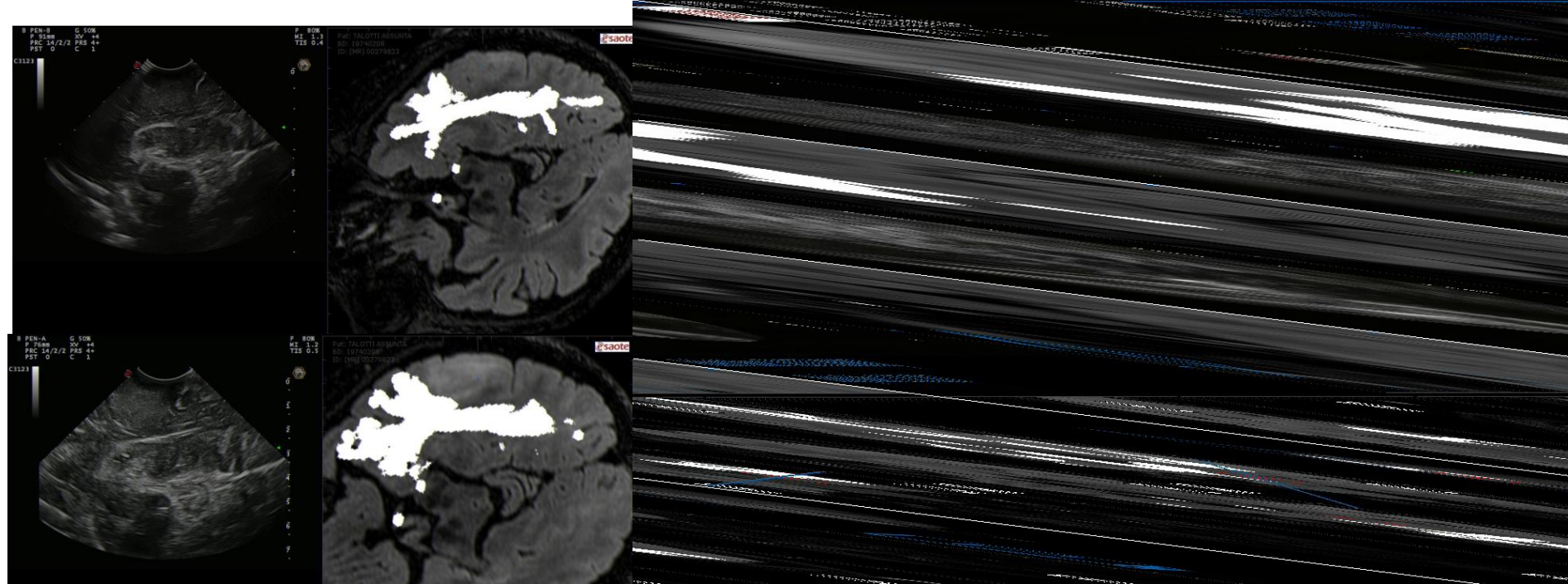


- F, 43 anni
- Crisi epilettiche

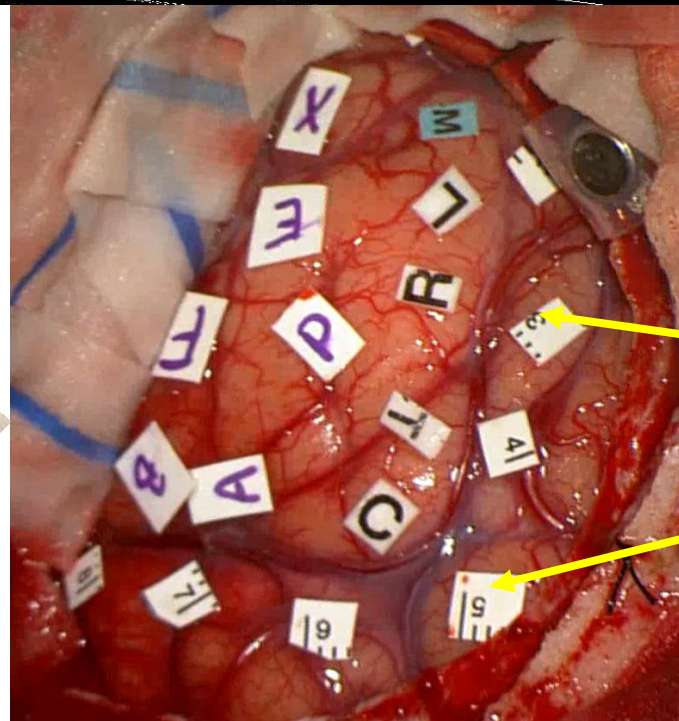
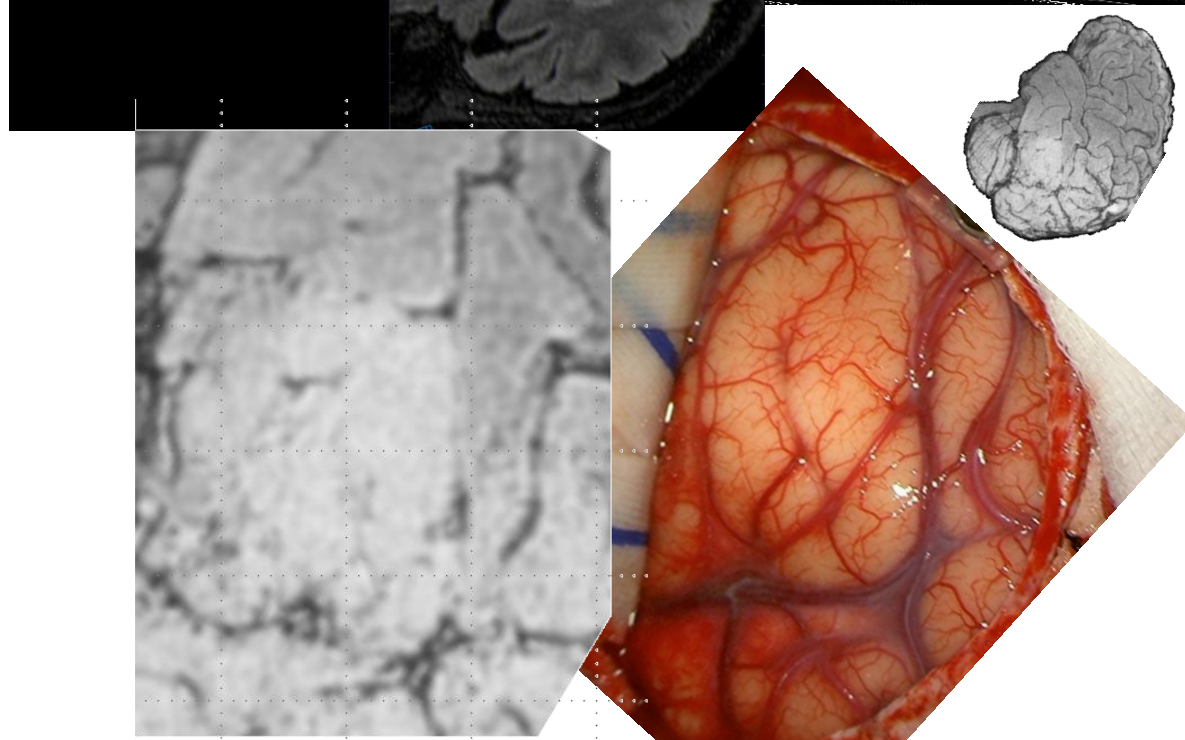


RMN funzionale

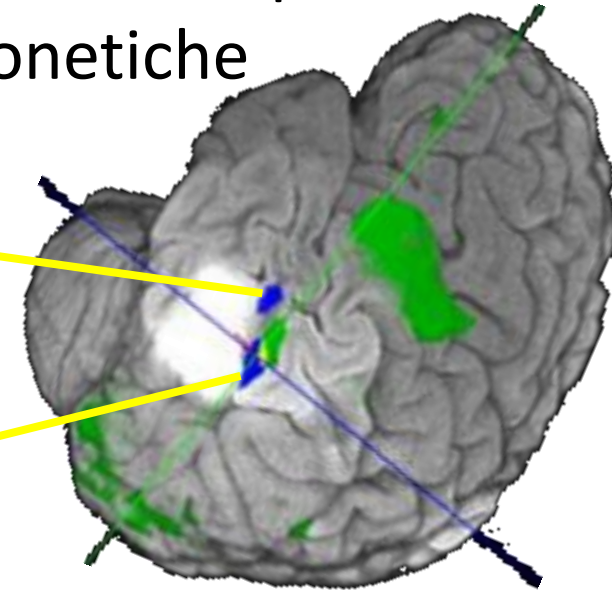


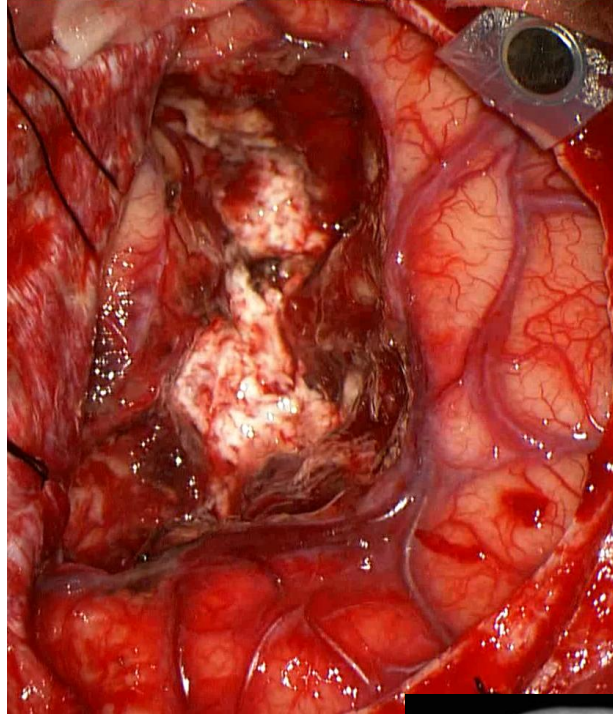
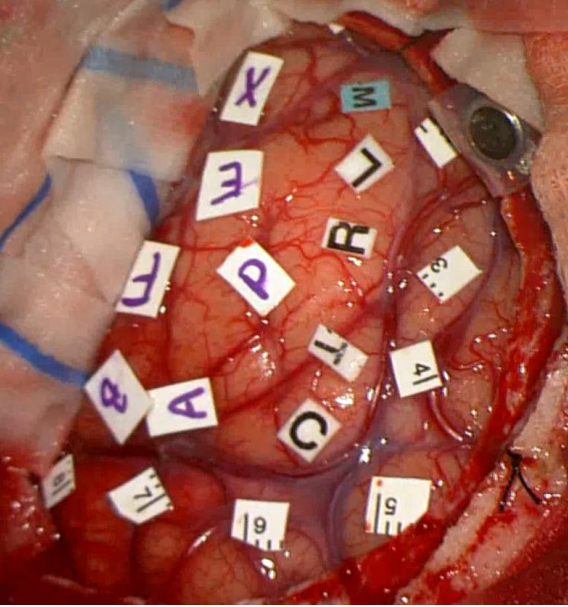


Awake surgery

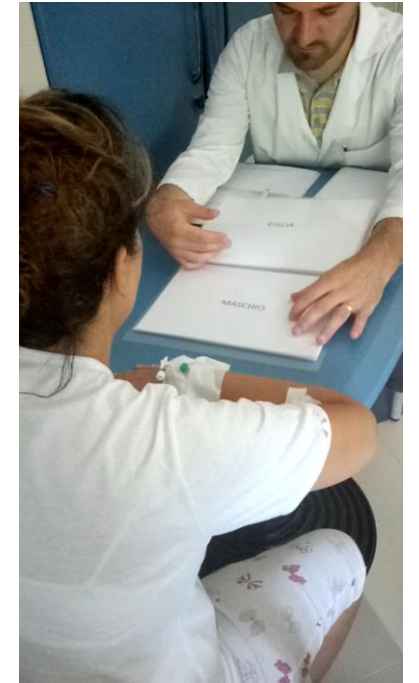


Anomie e parafasia fonetiche

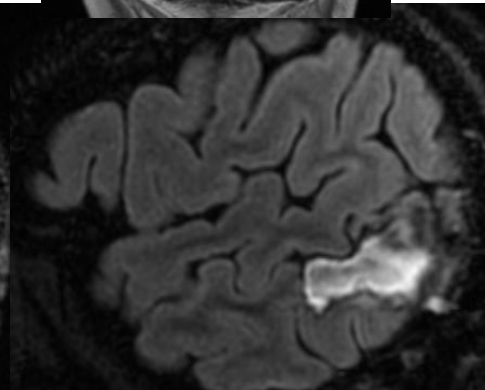
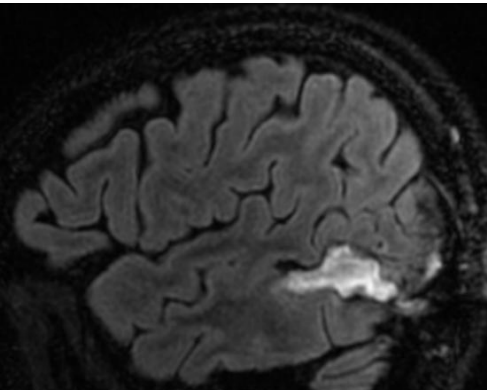
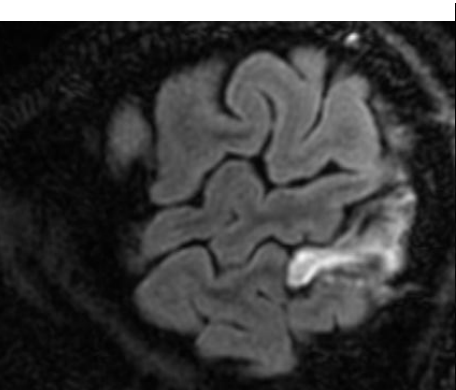
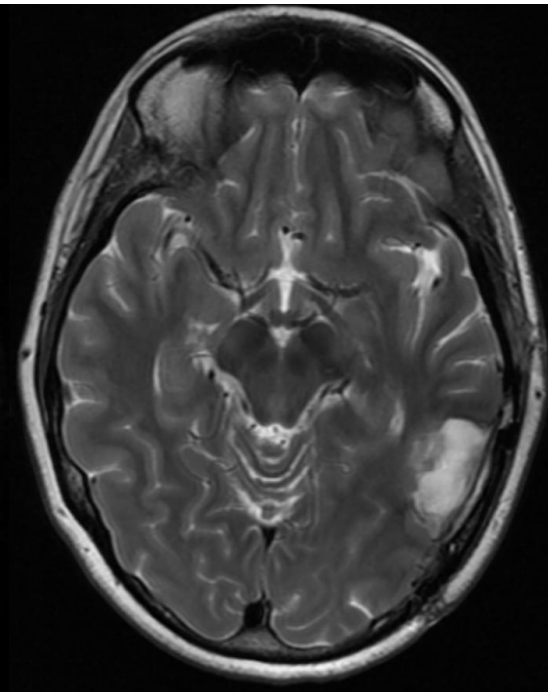
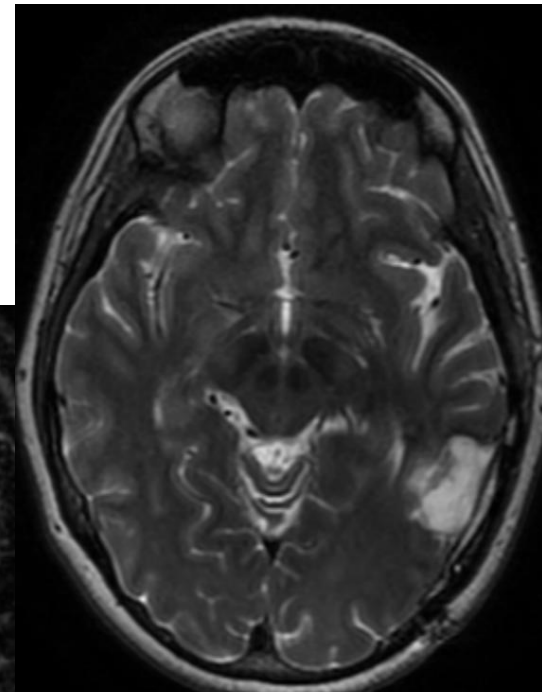
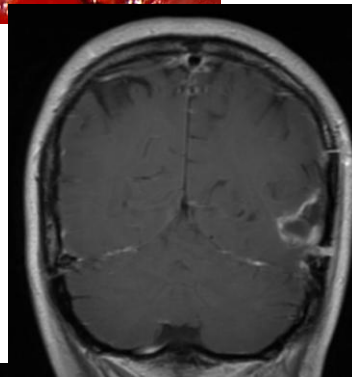


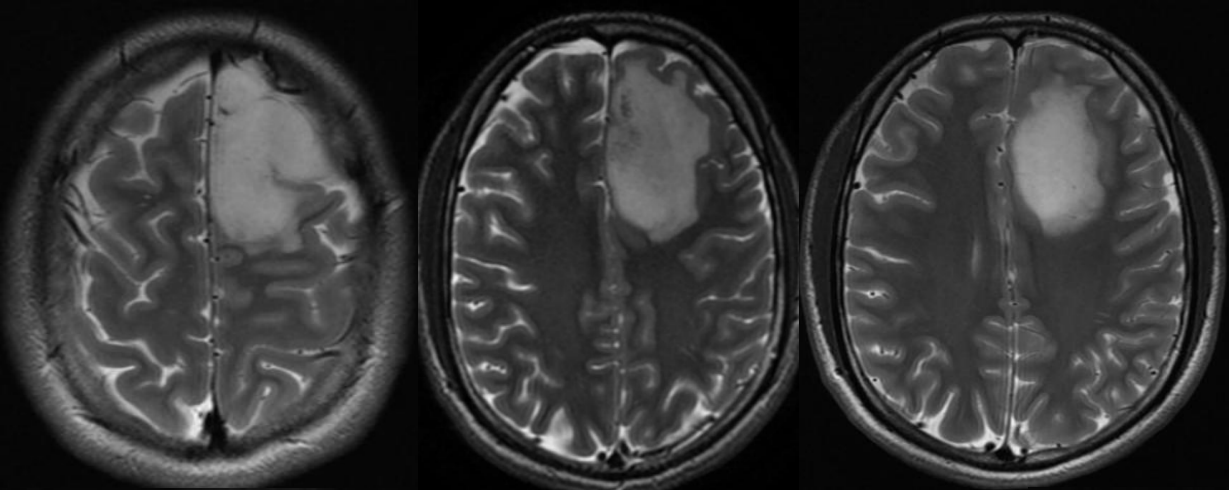


Oligodendroglioma
II grado WHO
Non crisi

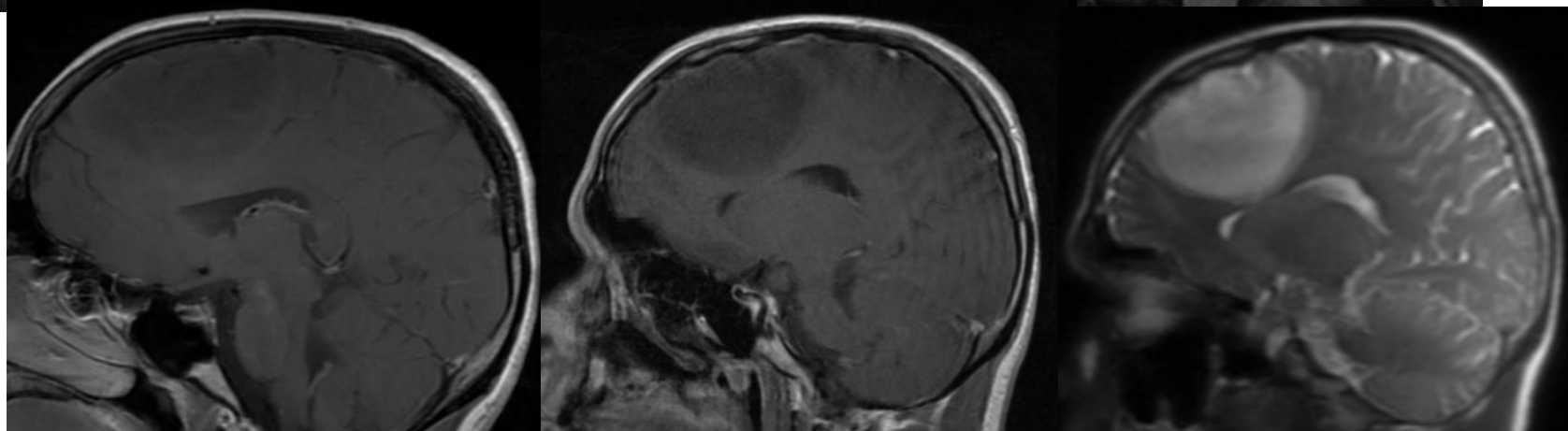
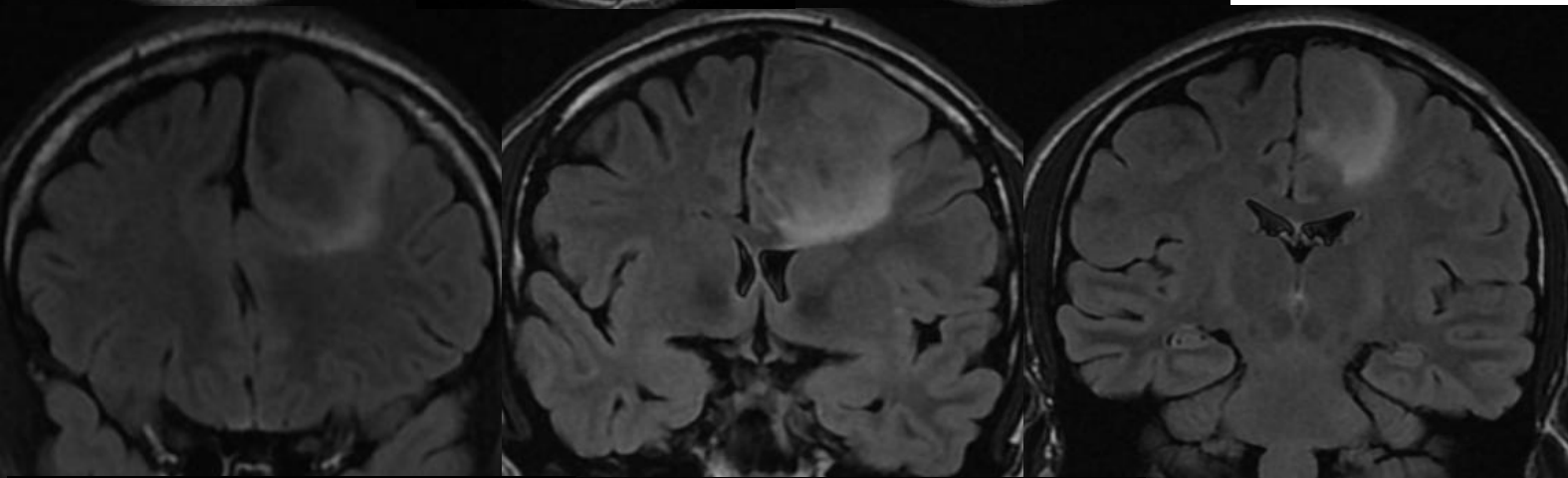


POST-OPERATORIO





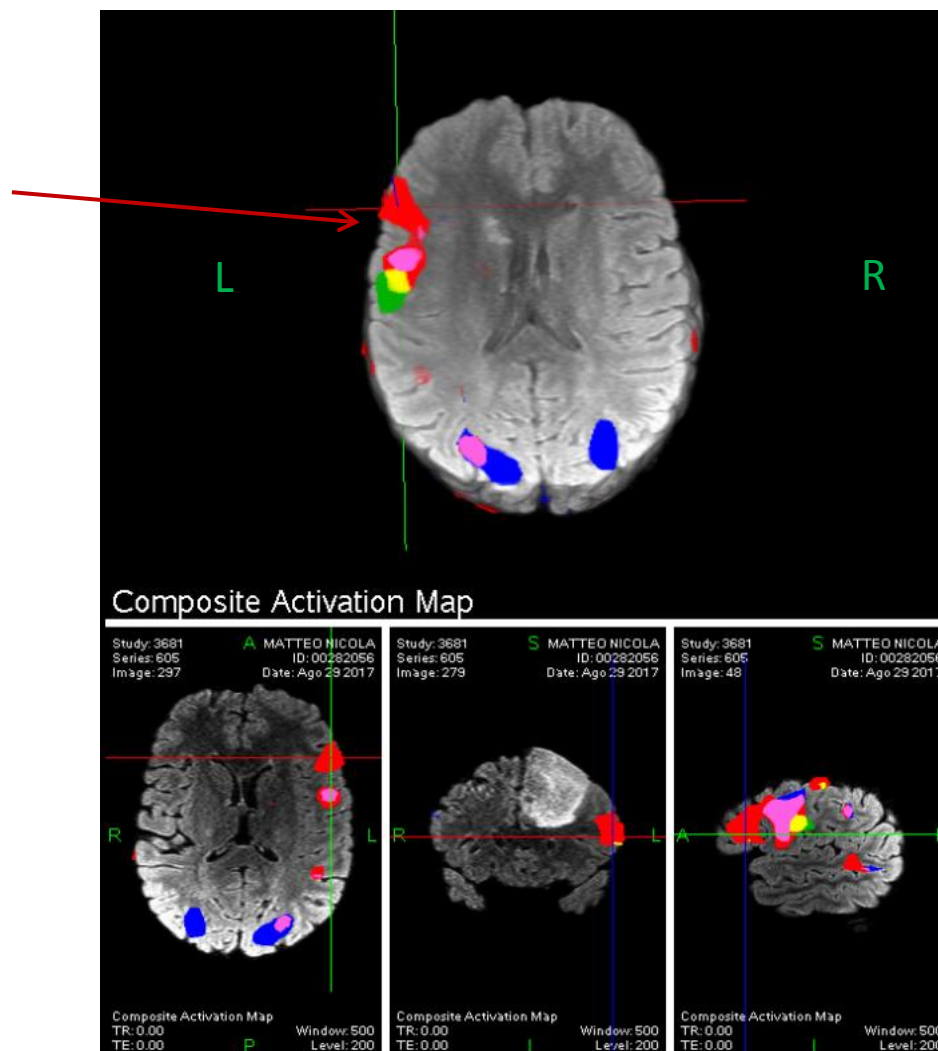
- M, 20 anni
- Crisi epilettiche generalizzate
- Non deficit neurologici
- Mancino



Awake surgery?

Mapping funzionale del linguaggio: paziente mancino

**Dominanza
emisferica
sinistra**



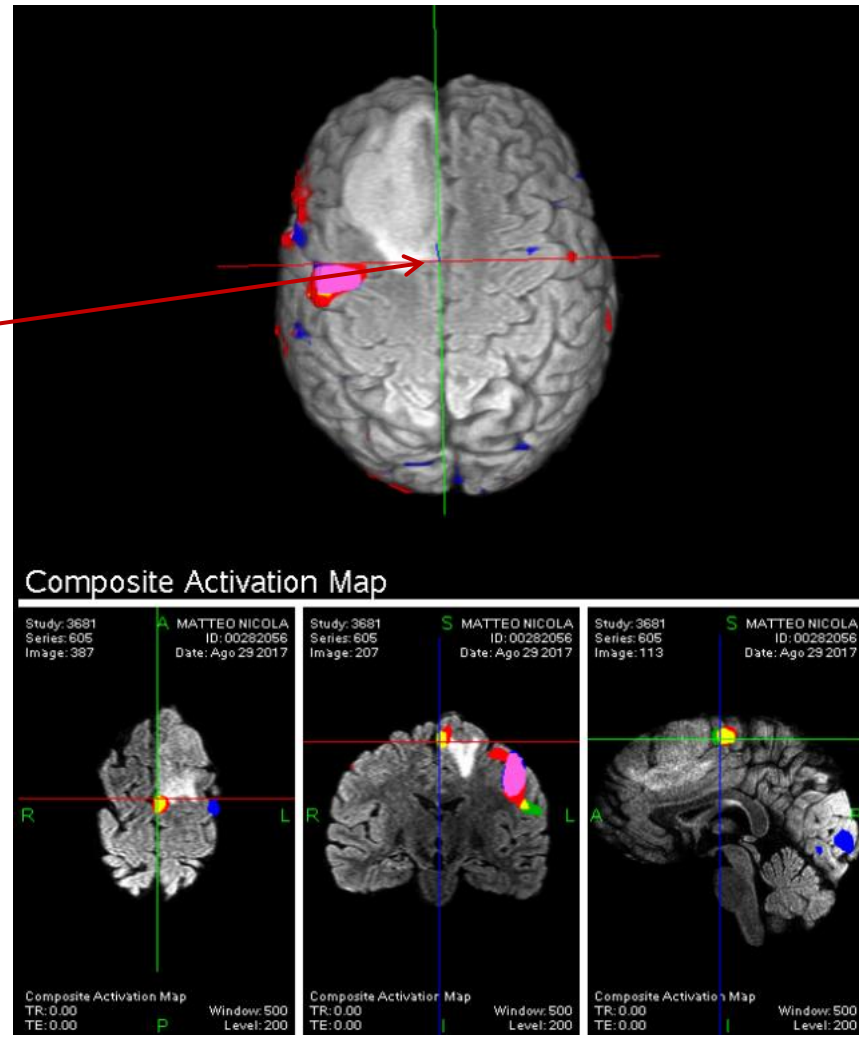
Mapping funzionale del linguaggio: individuazione delle aree critiche

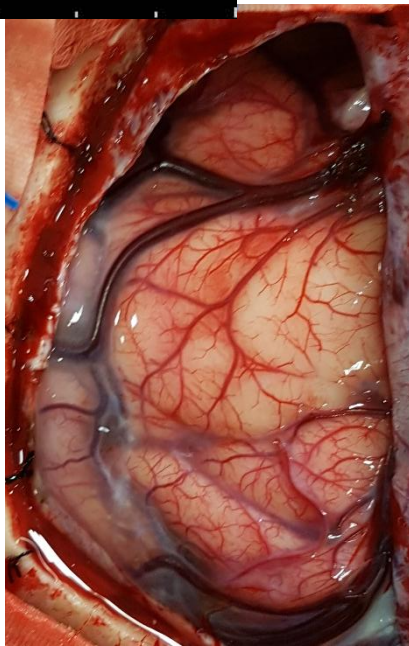
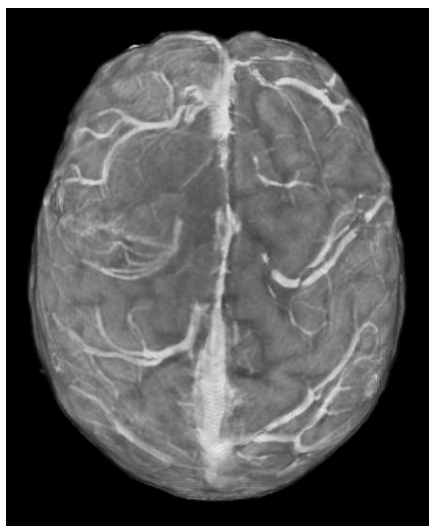
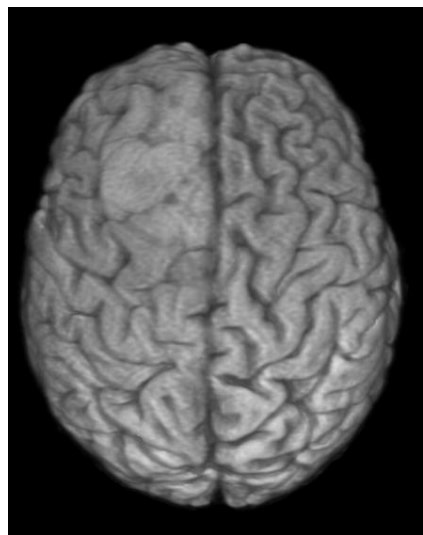
**Area critica
(pre-SMA)**

**Valutazione
neuro-
psicologica
estesa**

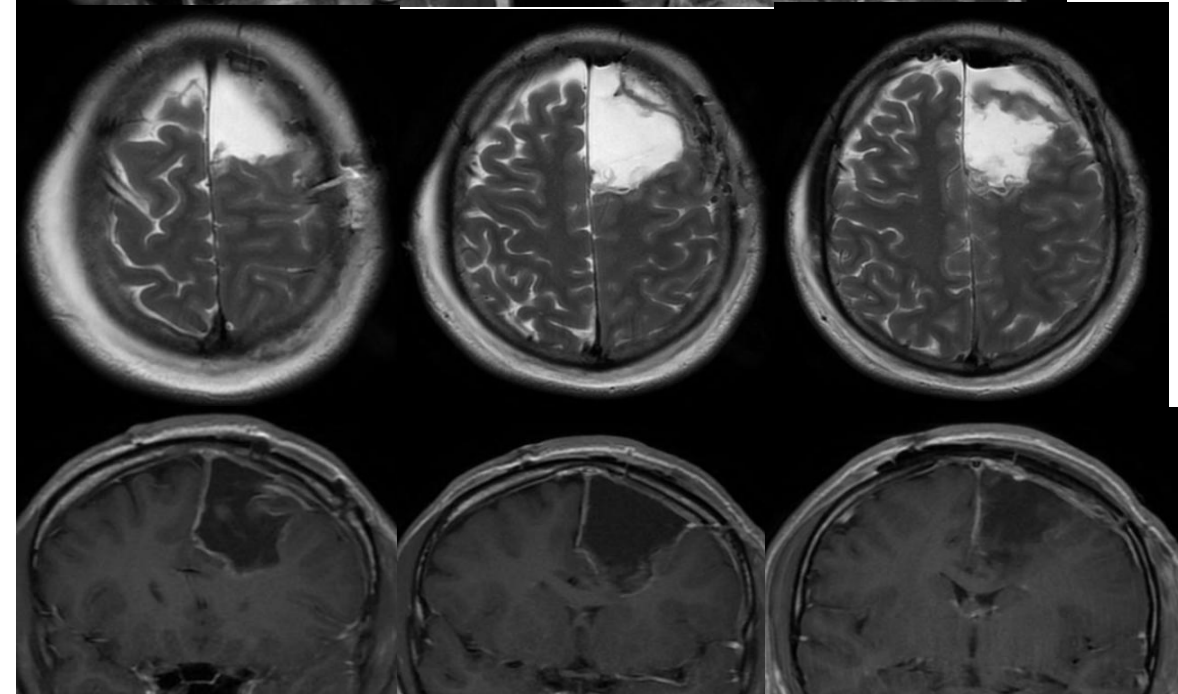
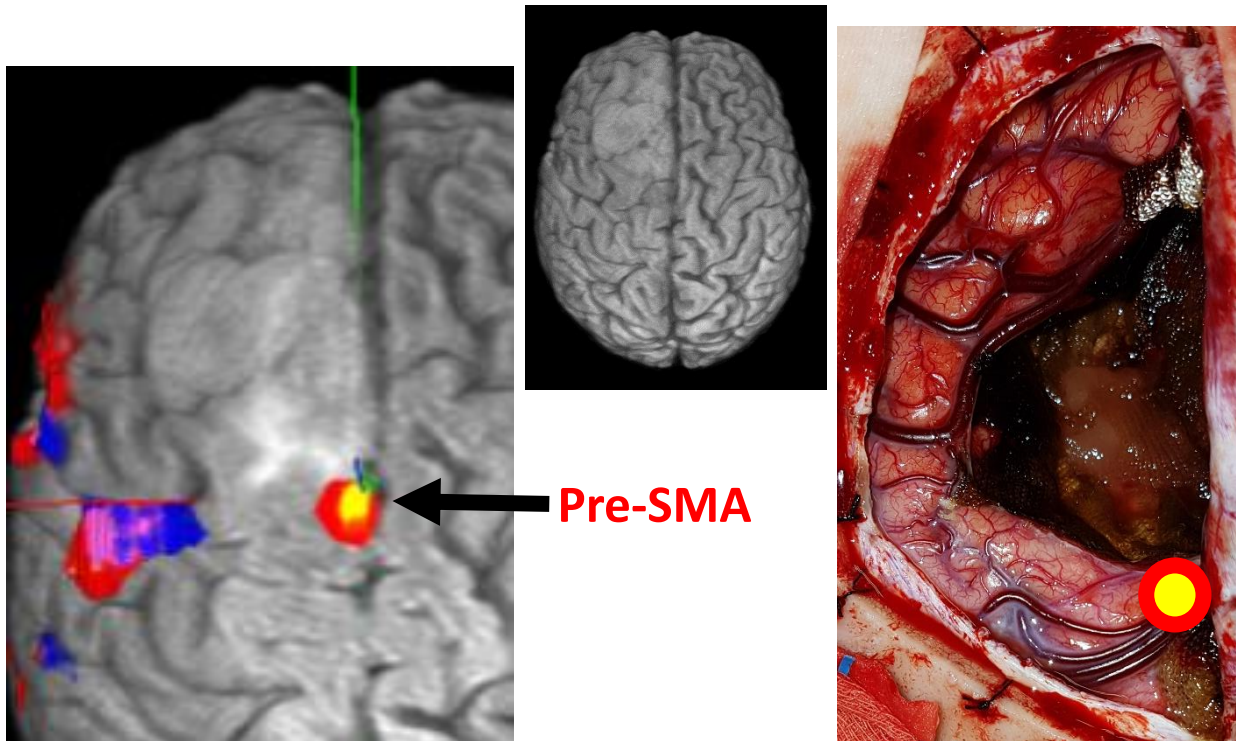
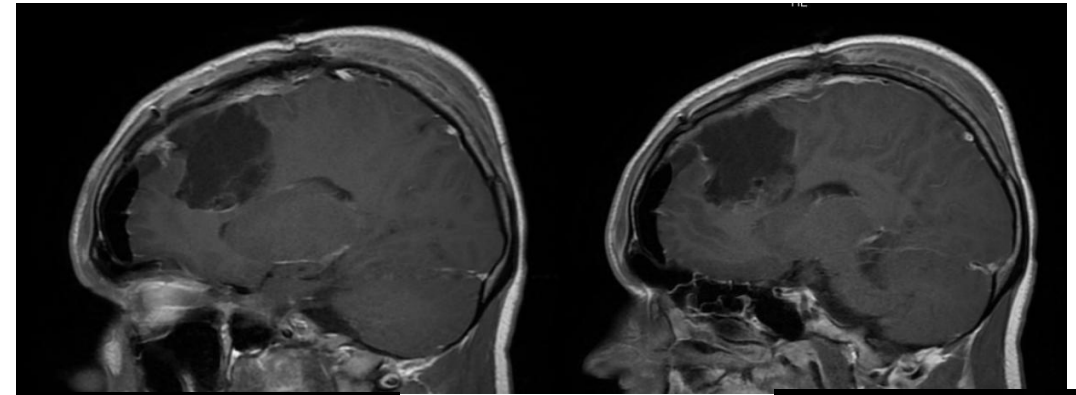
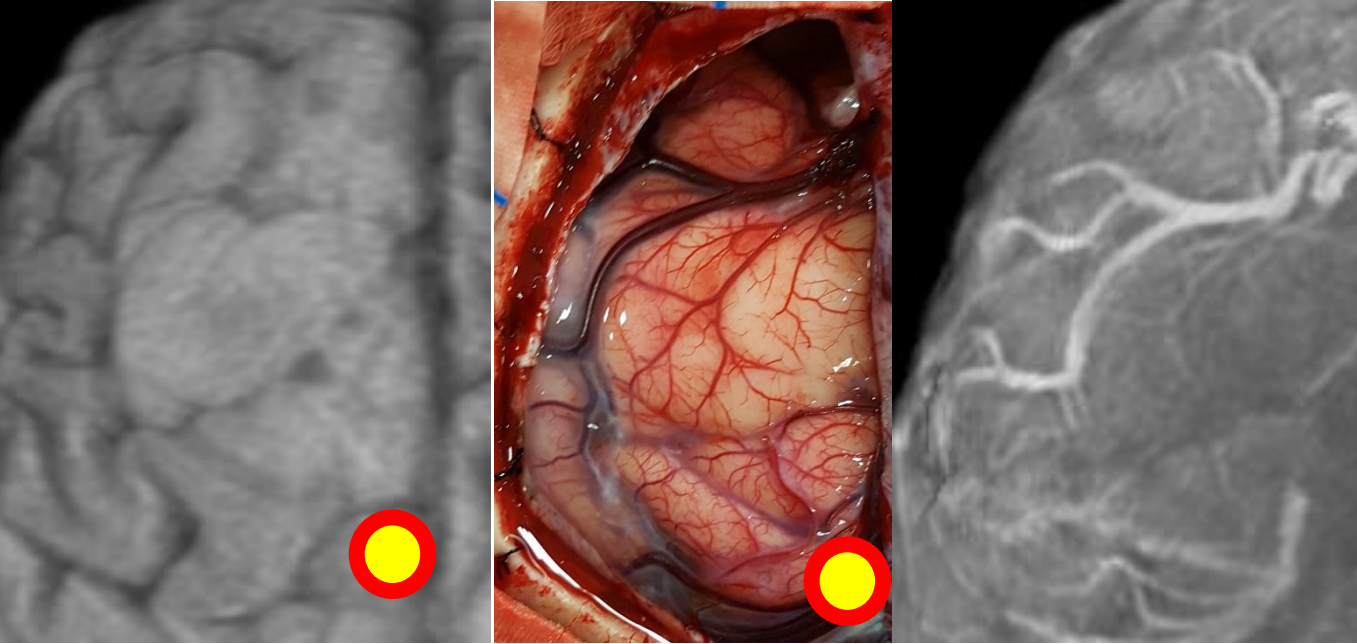
Awake
surgery?

NO



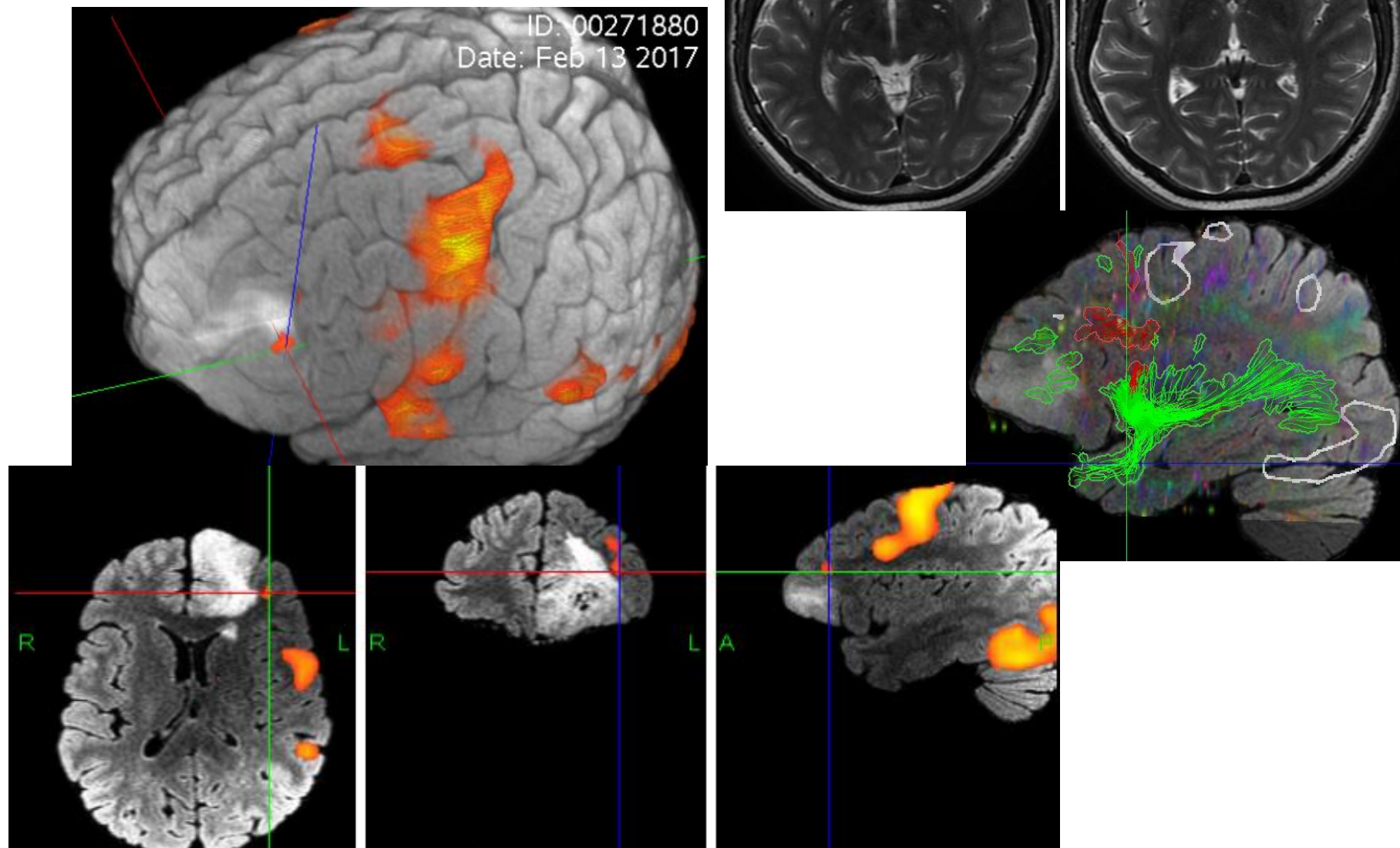


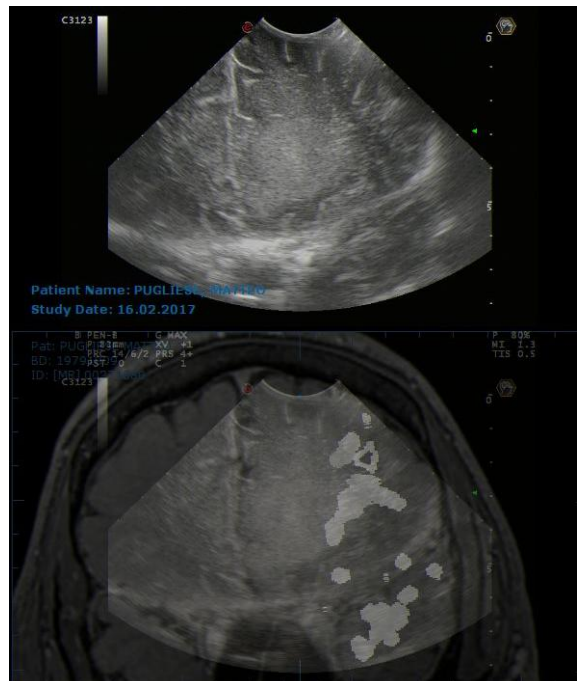
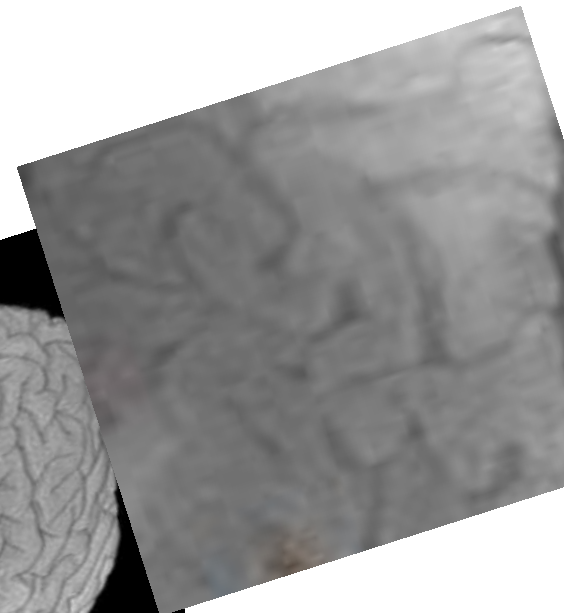
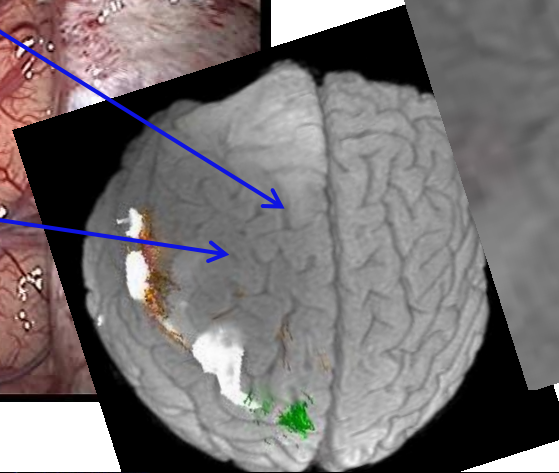
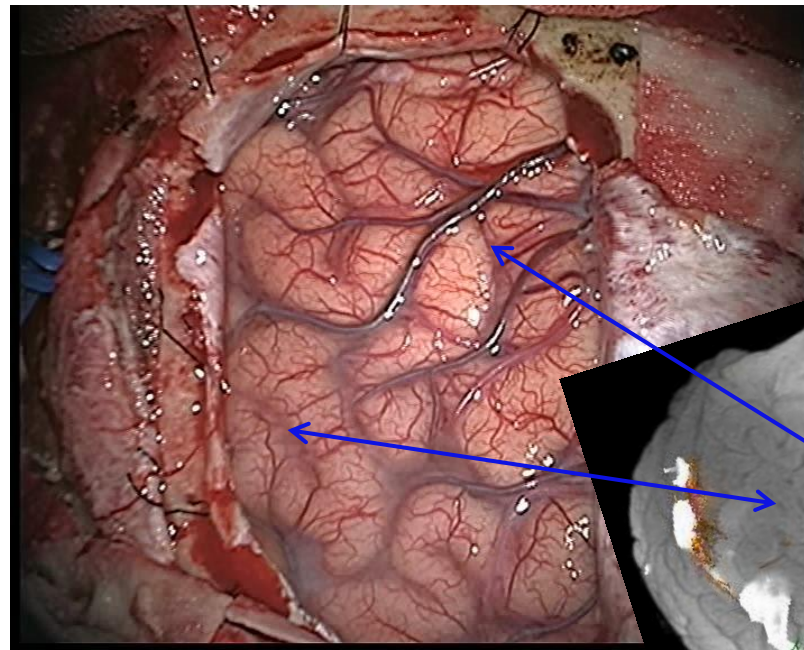
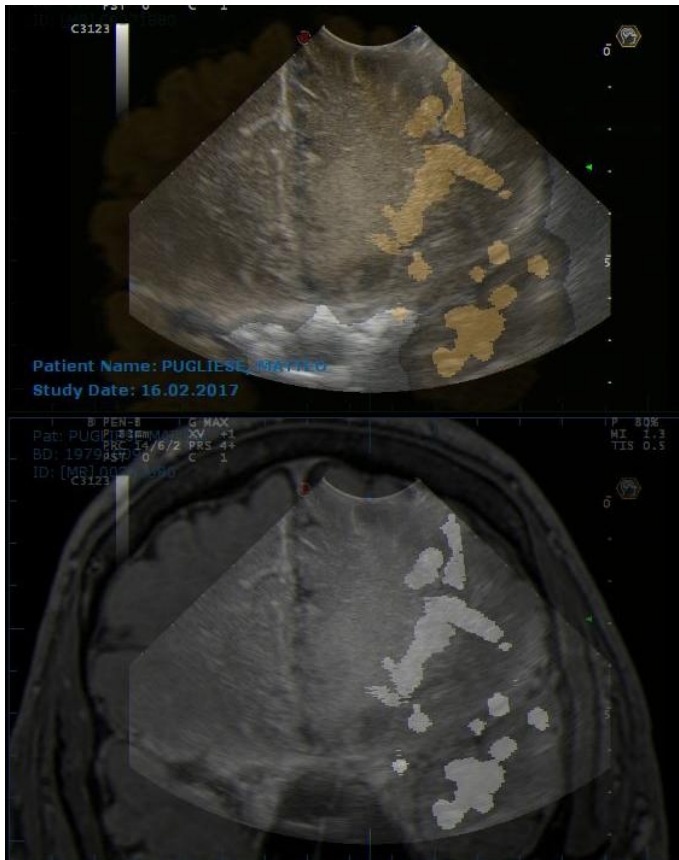
Astrocitoma diffuso II grado WHO Non crisi



Nessun disturbo del linguaggio

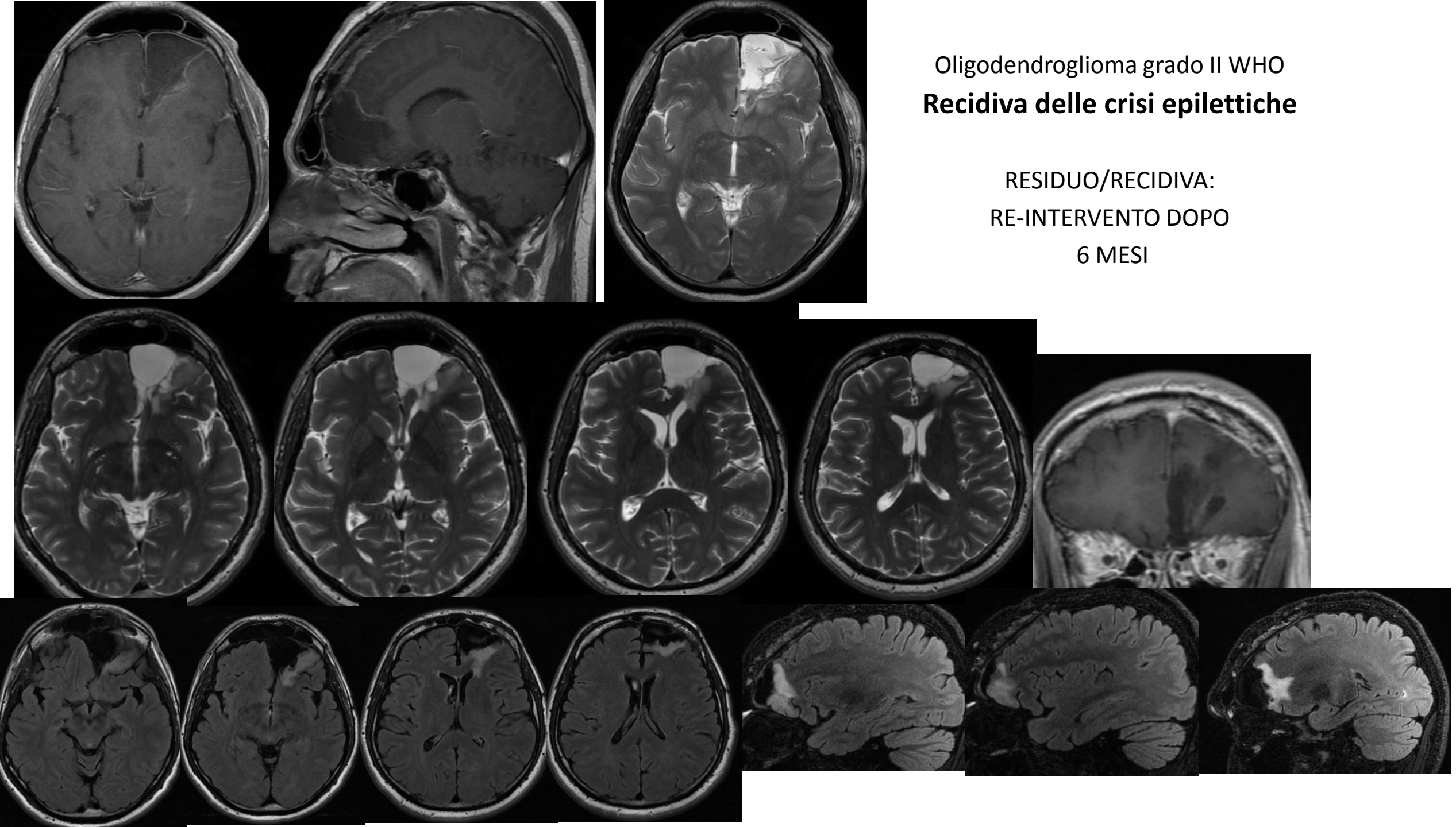
- M, 37 anni
- Esordio con crisi epilettica generalizzata, poi ripetutesi

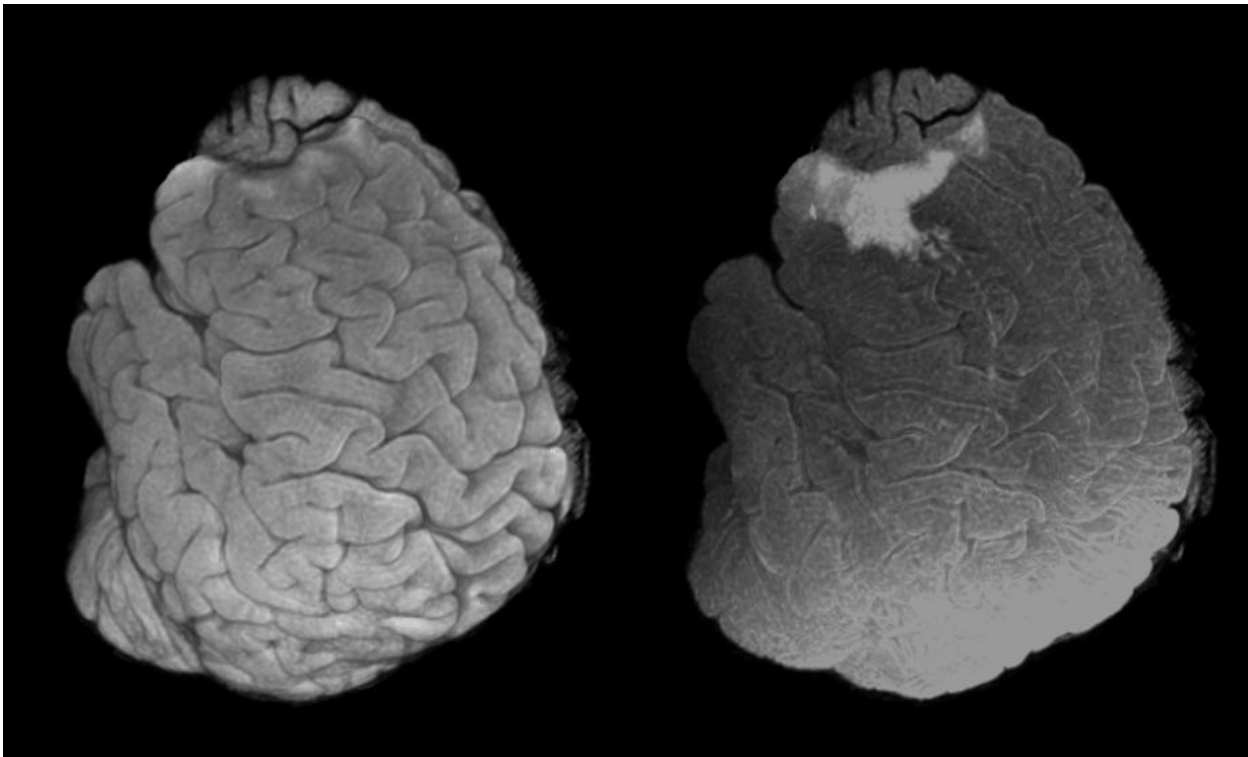
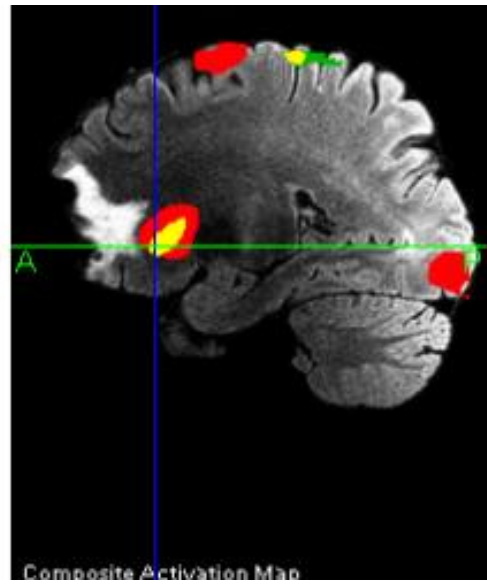
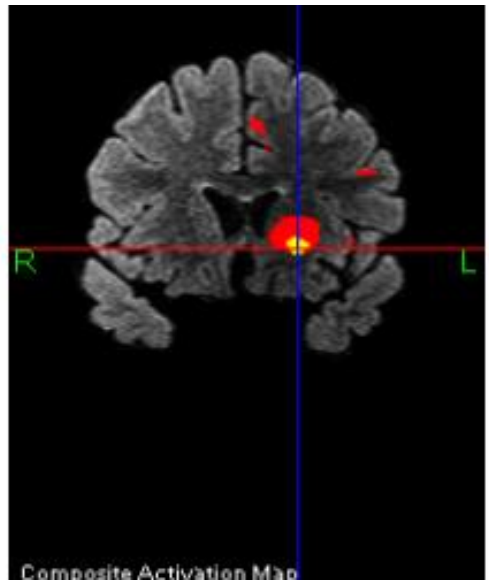
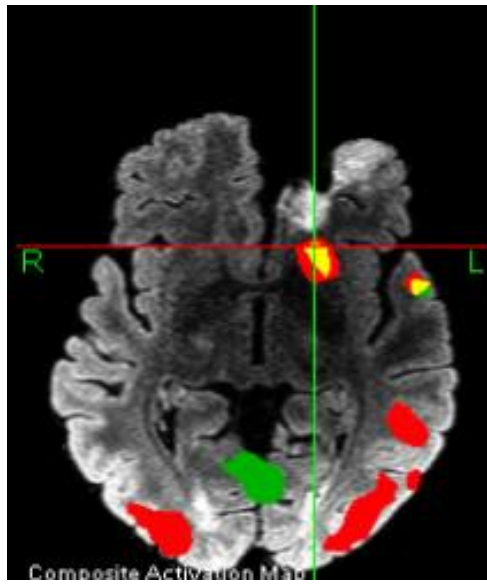


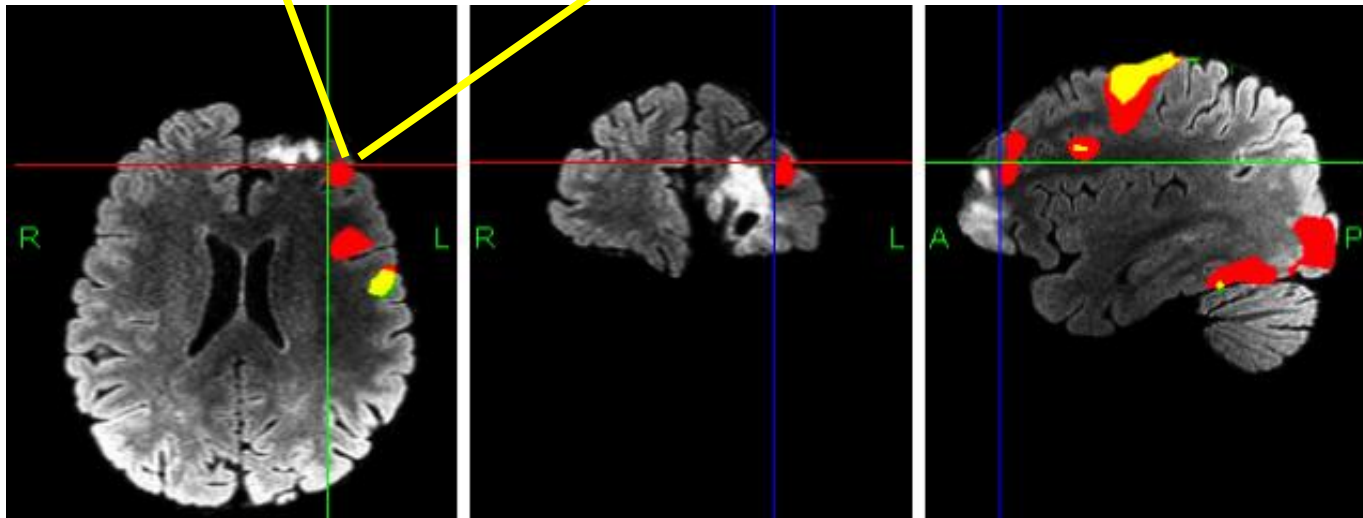
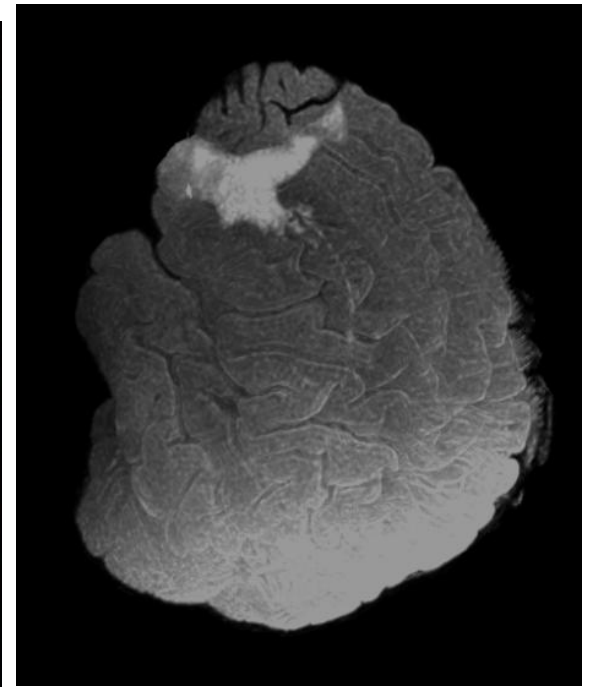
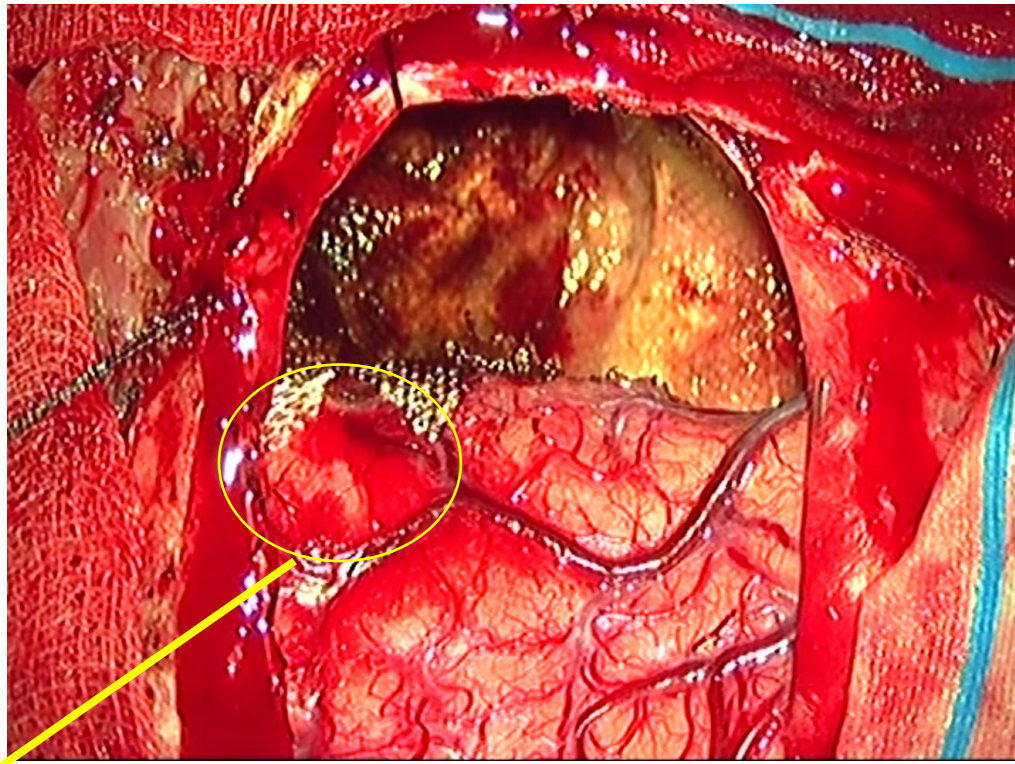
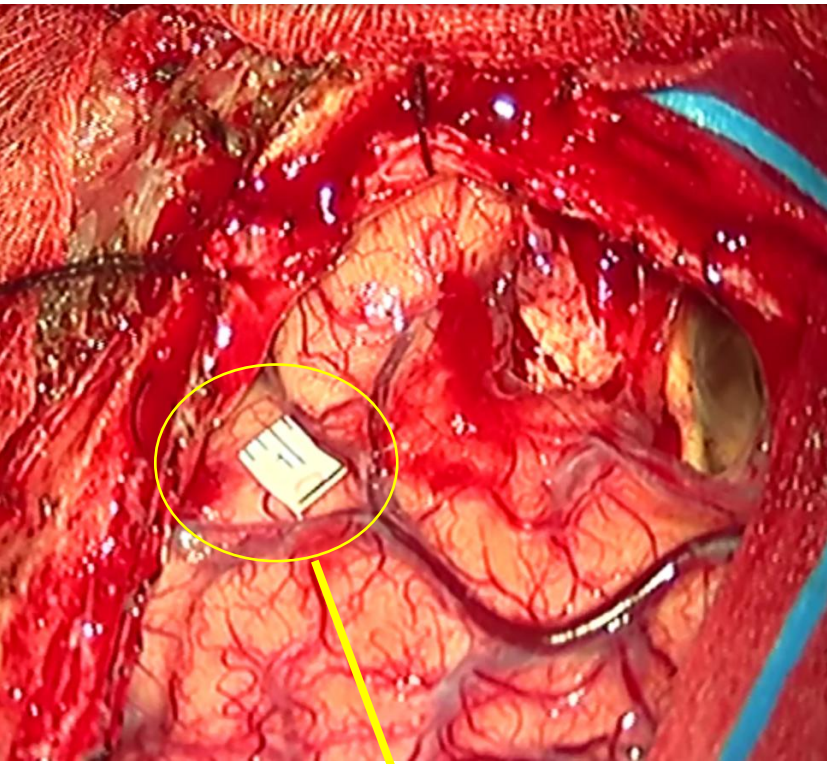


Oligodendroglioma grado II WHO
Recidiva delle crisi epilettiche

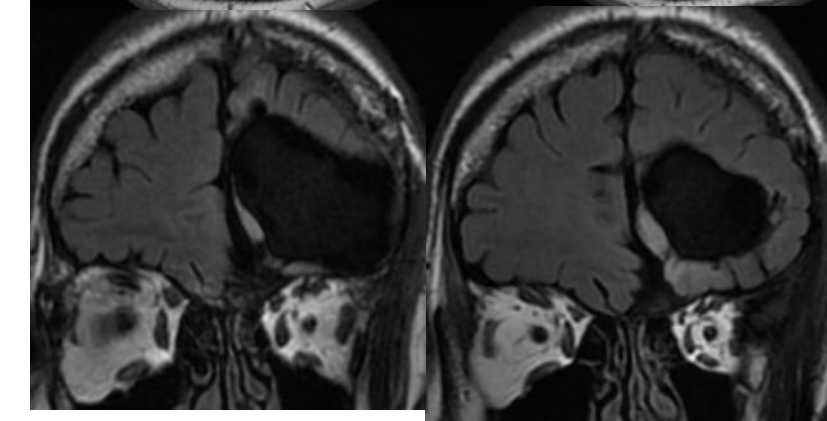
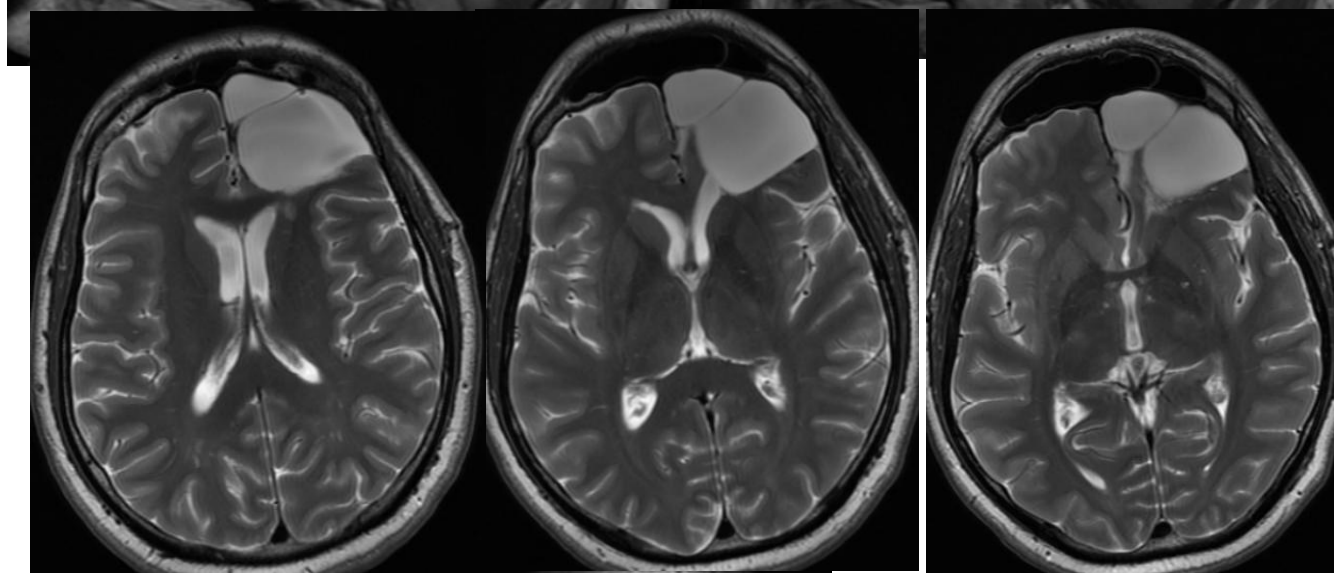
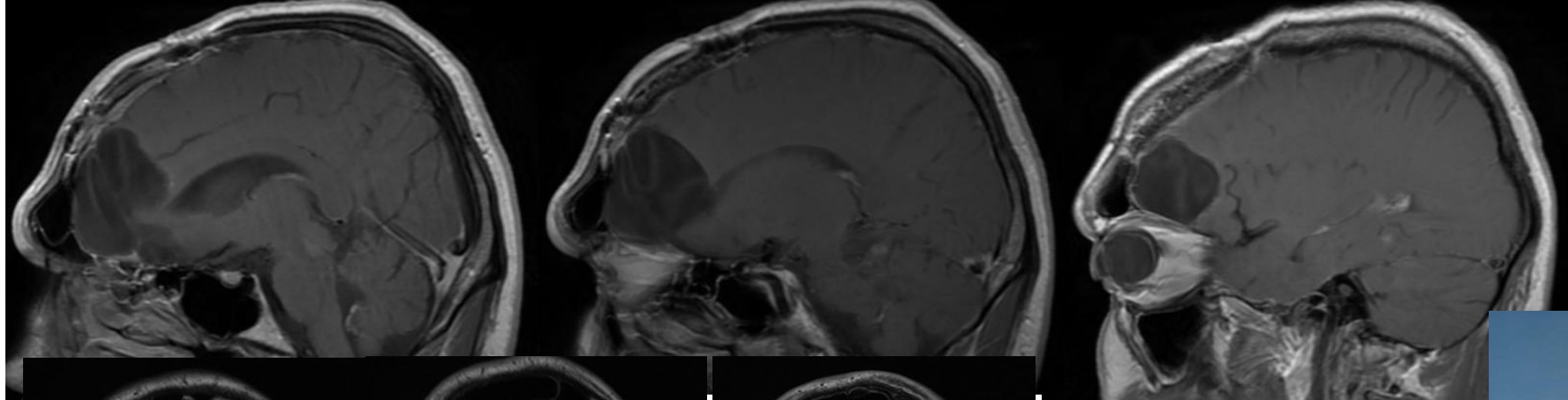
RESIDUO/RECIDIVA:
RE-INTERVENTO DOPO
6 MESI



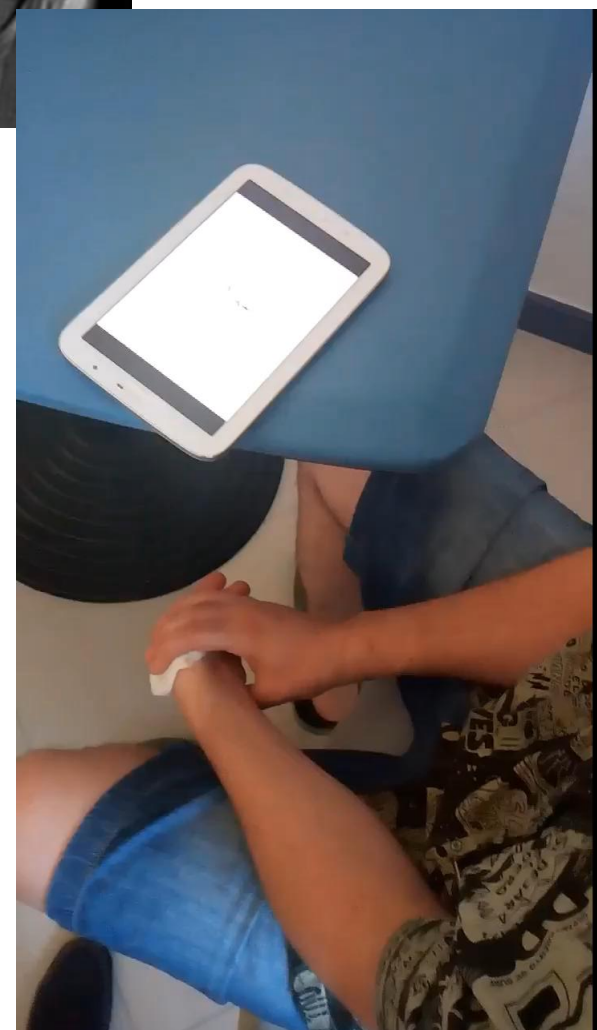




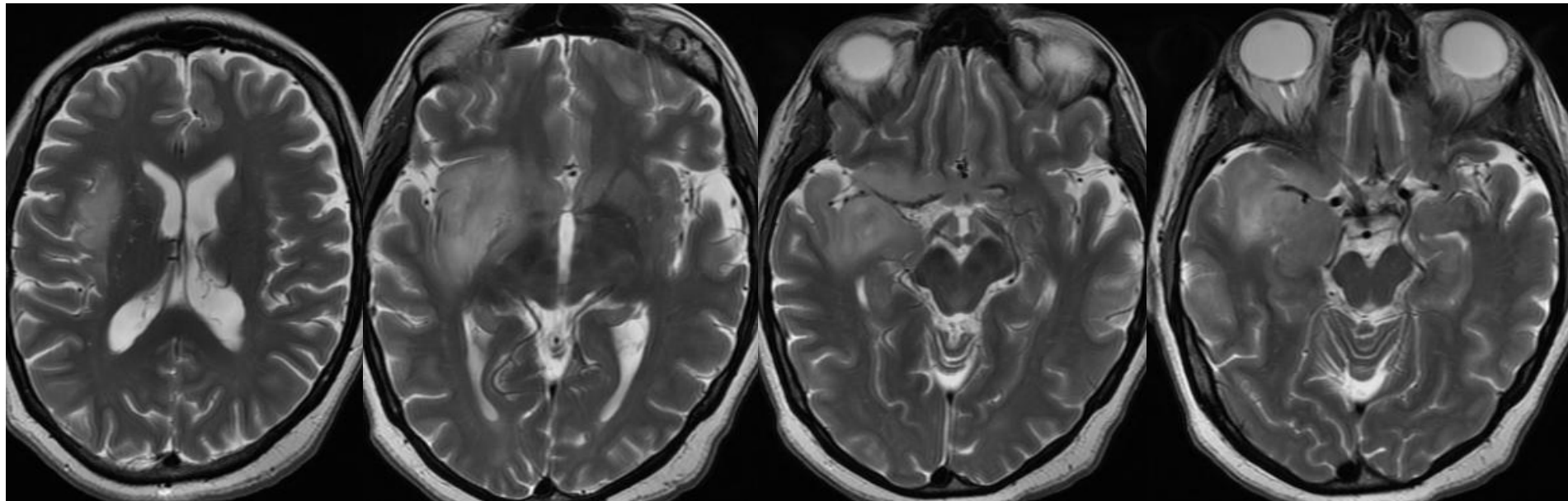
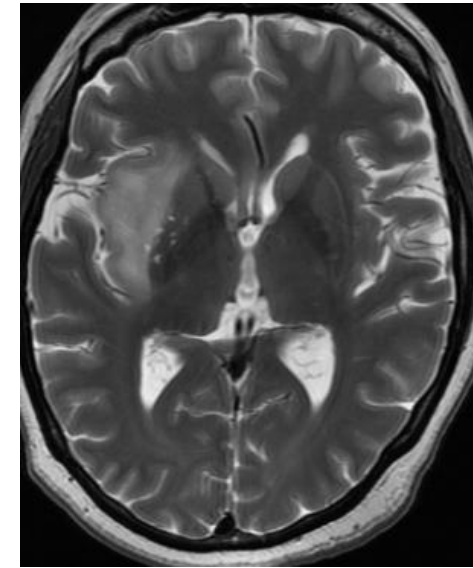
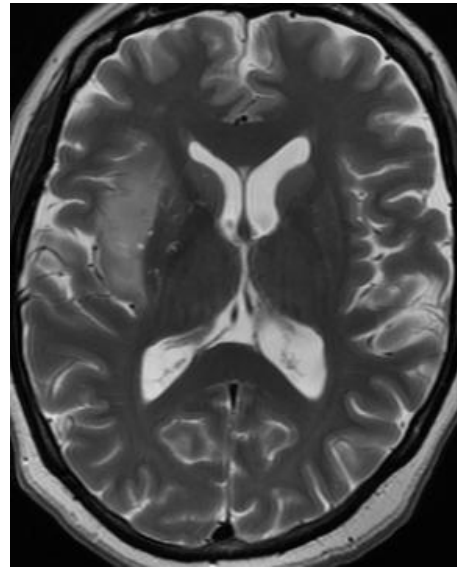
Reintervento in
Awake surgery

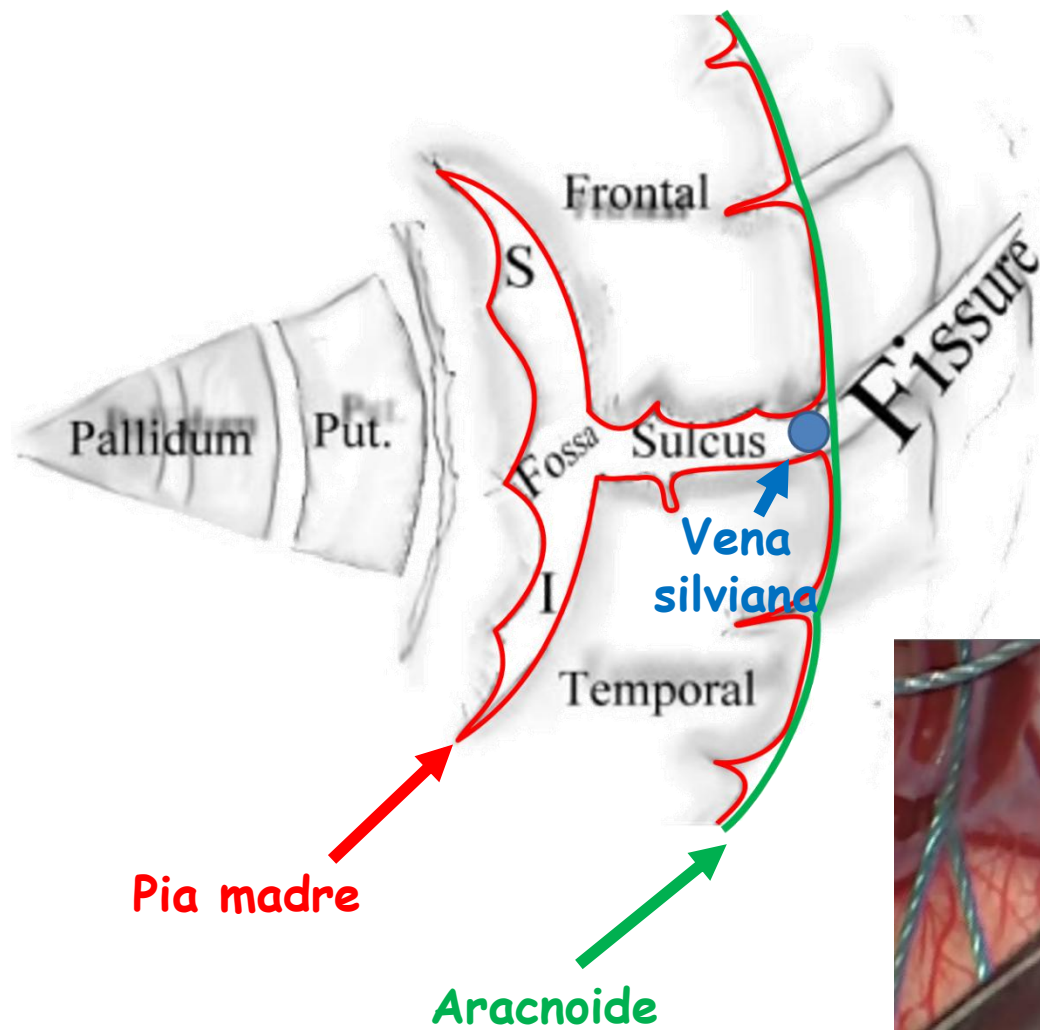


Scomparsa delle crisi epilettiche

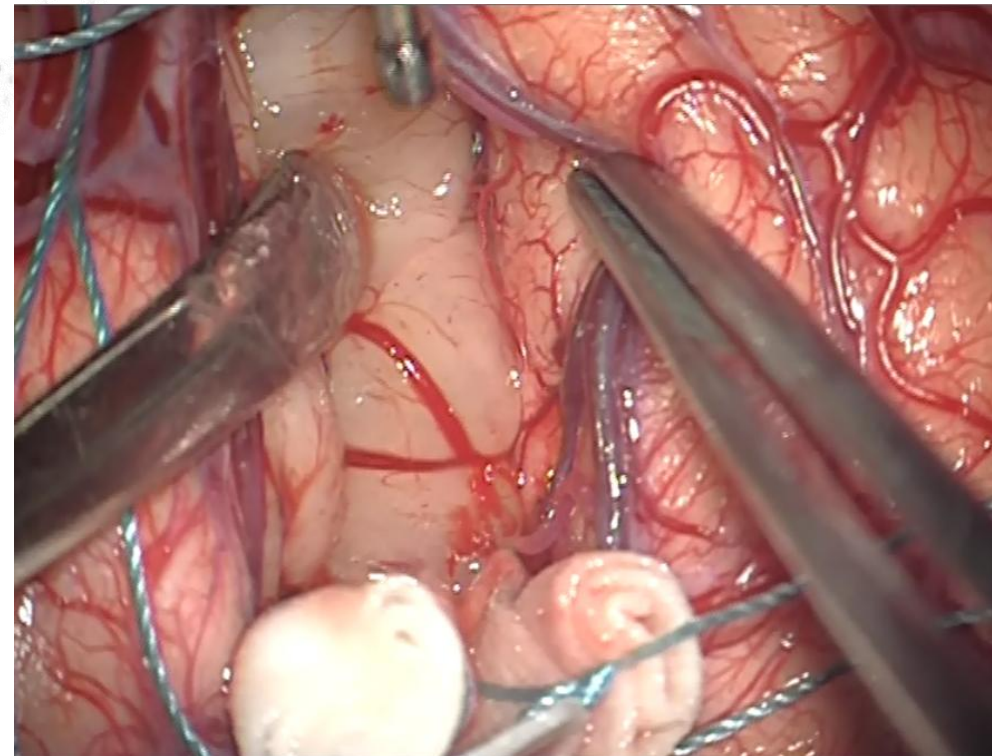
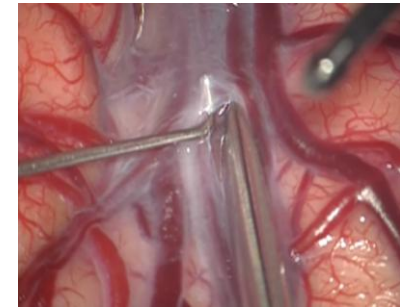


- F, 49 anni, impiegata
- Storia di crisi epilettiche notturne

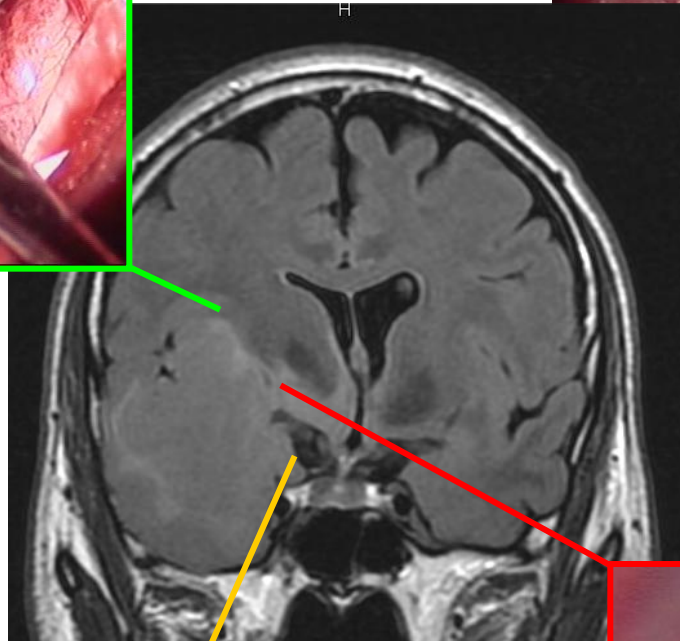
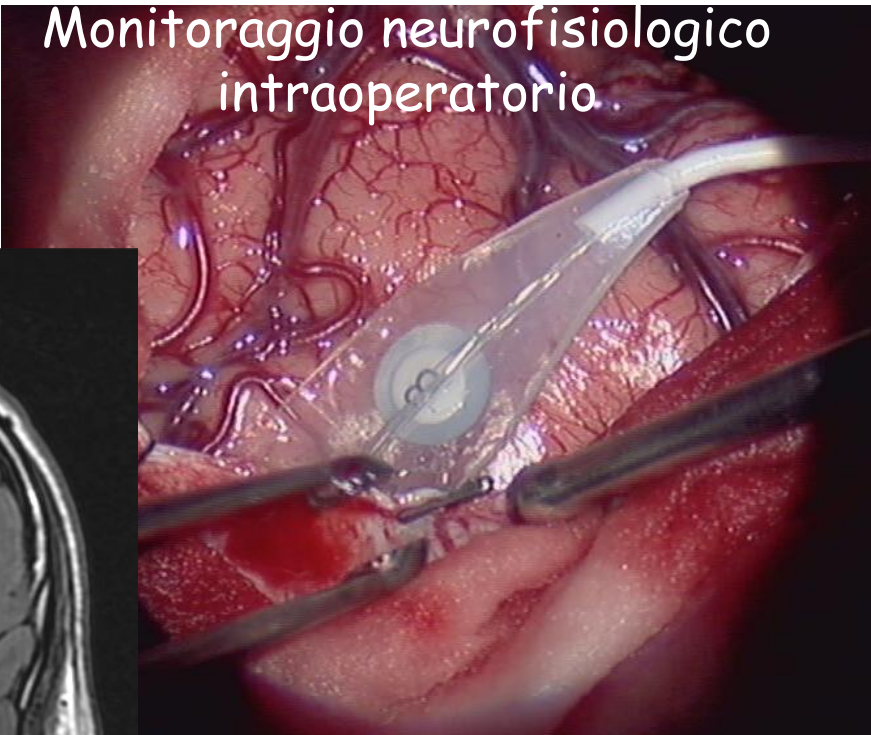
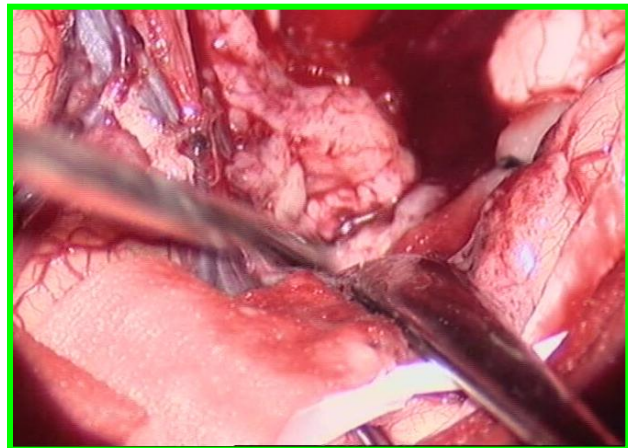




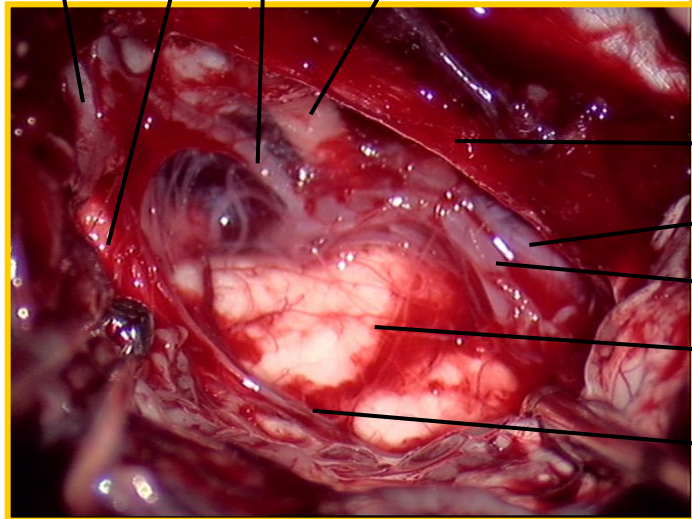
- Microchirurgia ad alto ingrandimento
- Dissezione subpiaie



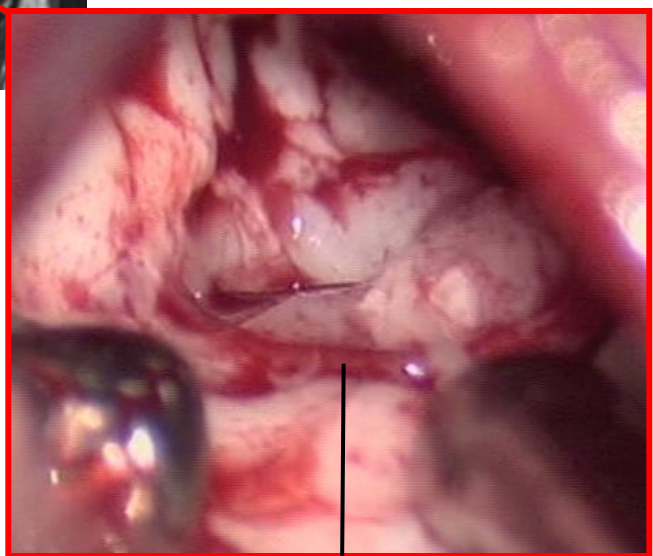
Monitoraggio neurofisiologico
intraoperatorio



ACI
PCoMA
Opt.T.
3rd nc

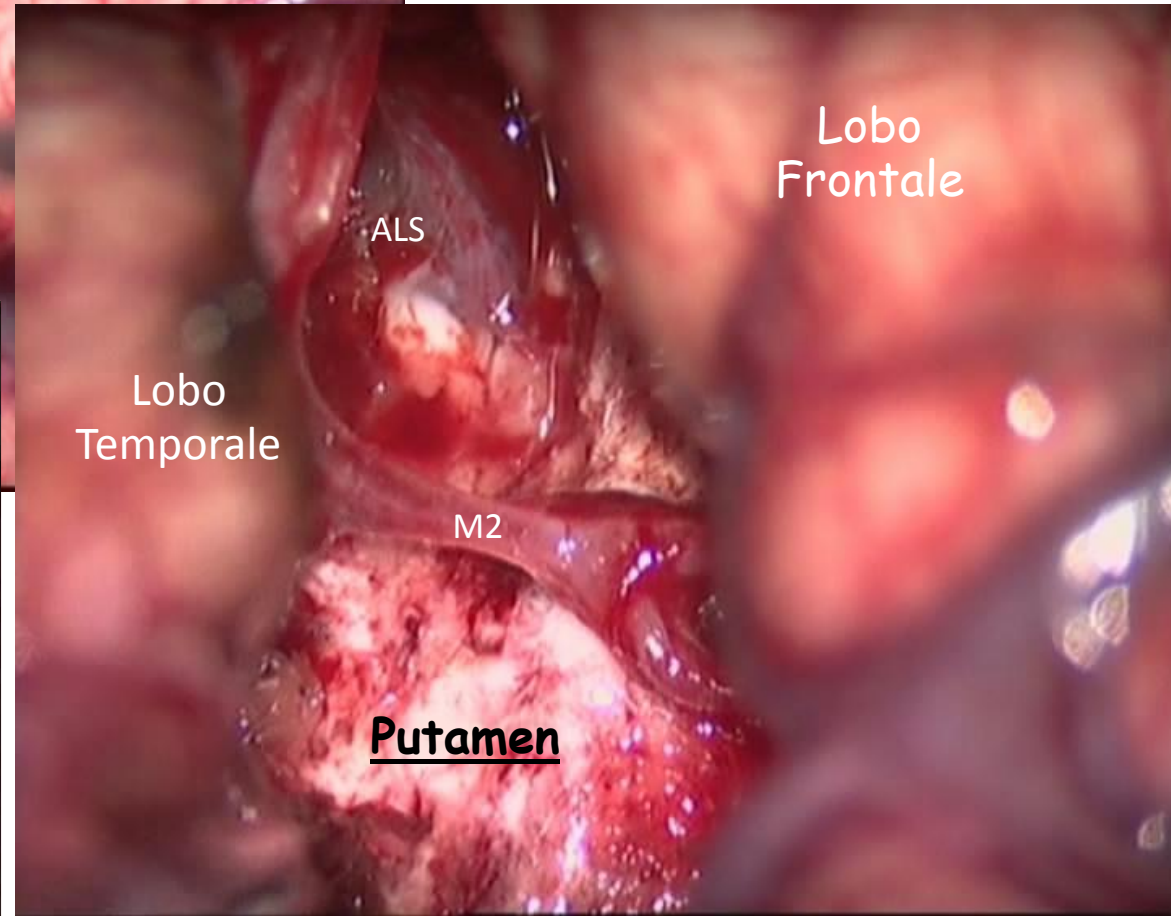
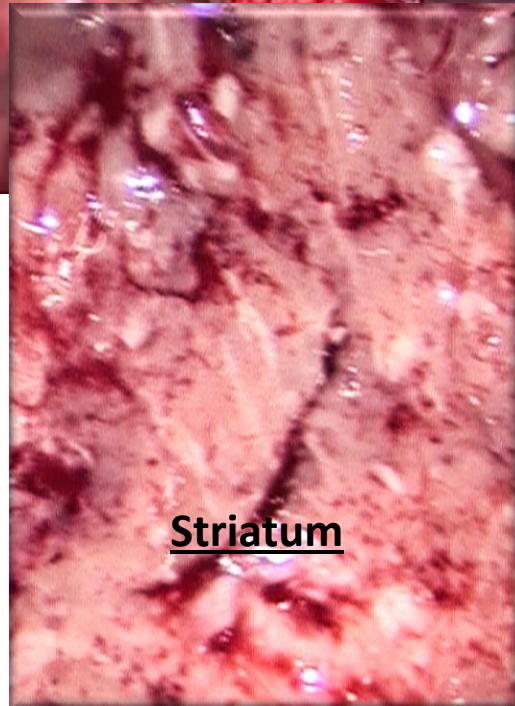
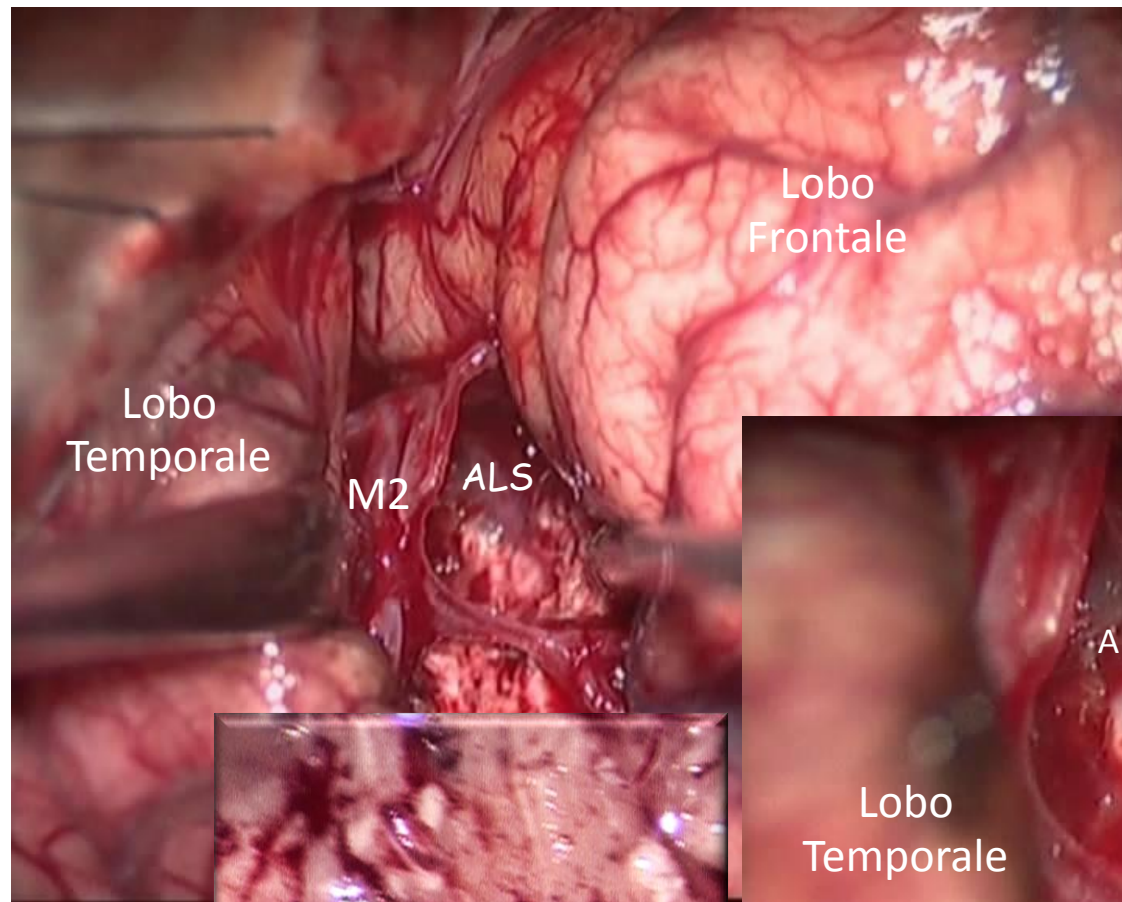


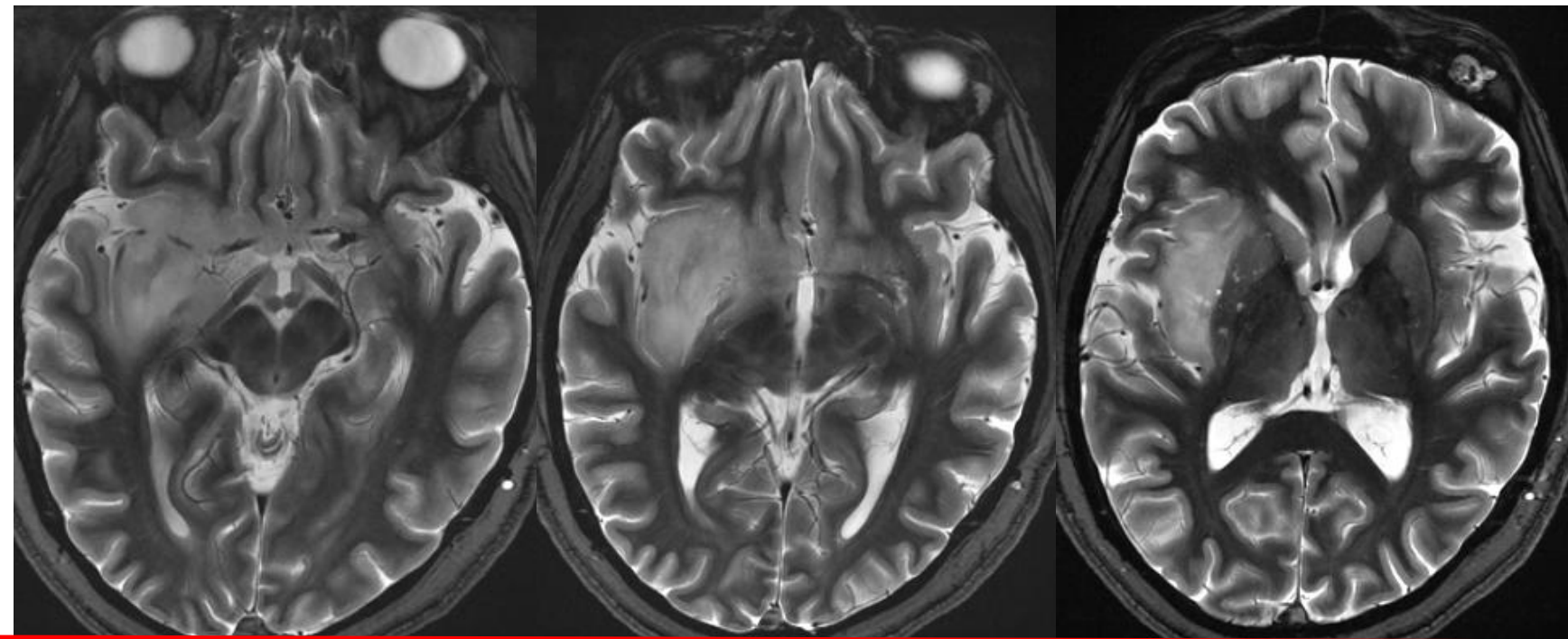
Tentorium
S.C.A.
P.C.A.
Brain Stem
A.Cho.A.



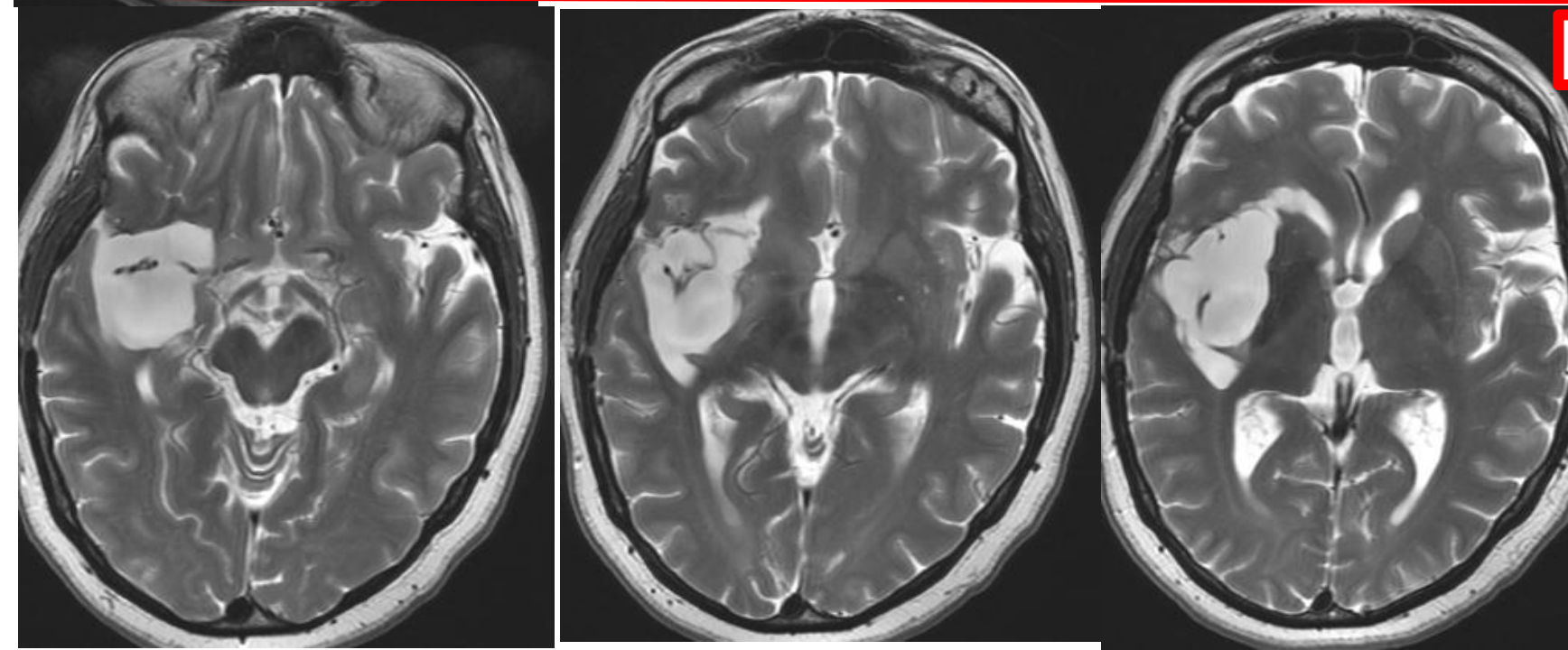
Arterie lenticolo-striate

Microchirurgia ad alto ingrandimento





Preop



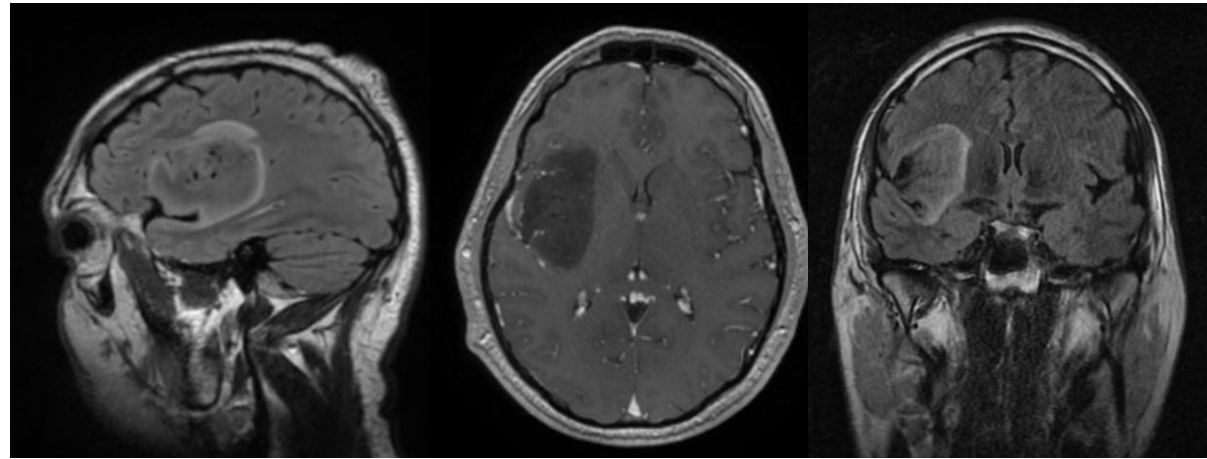
Postop

Oligodendroglioma Grado II WHO
F.U. 3 anni **Scomparsa delle crisi**

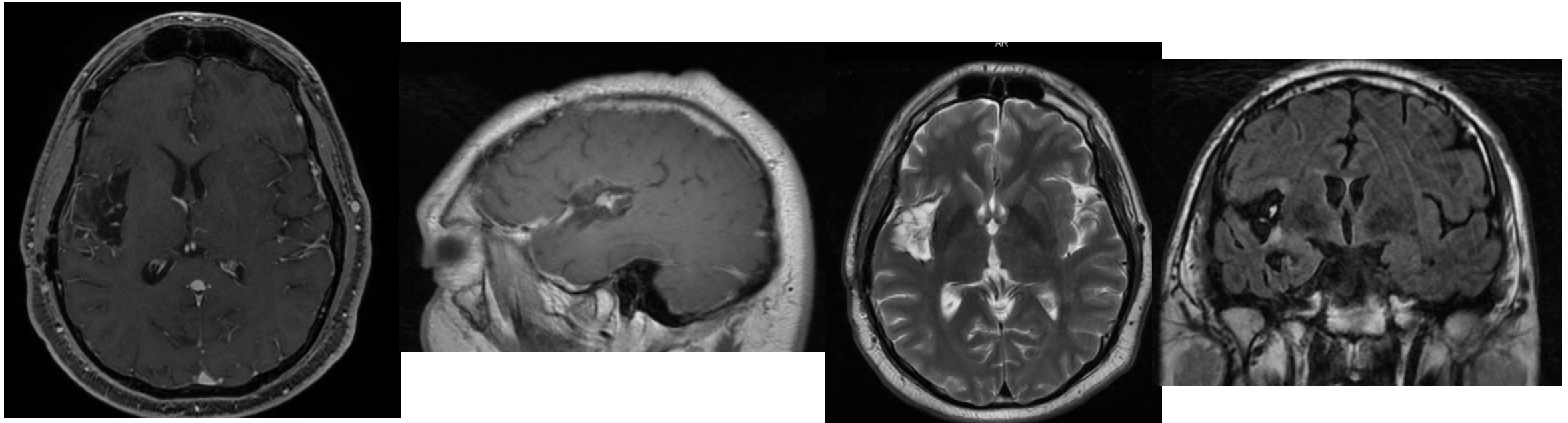


- M, 31 anni
- Esordio con crisi epilettiche
- Neoplasia insulare destra

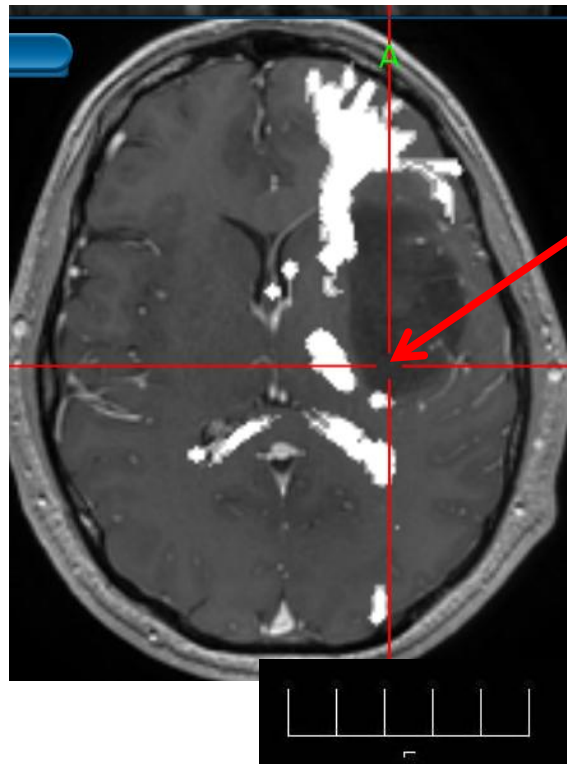
PRE-OPERATORIO



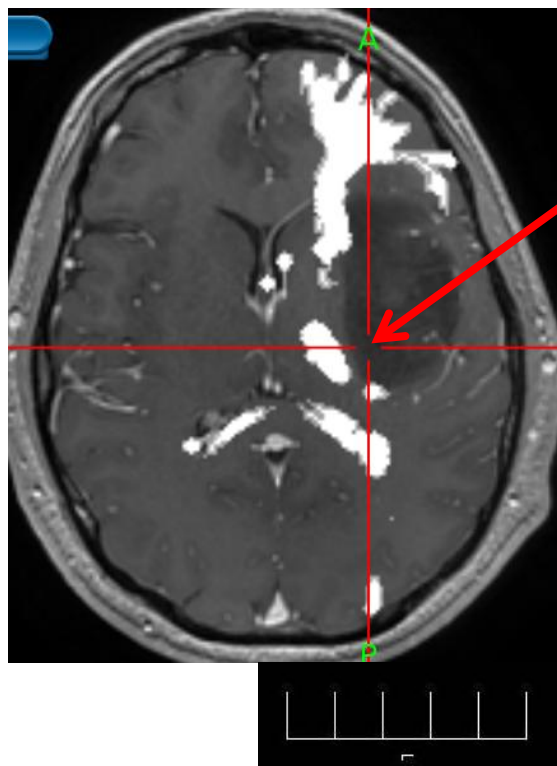
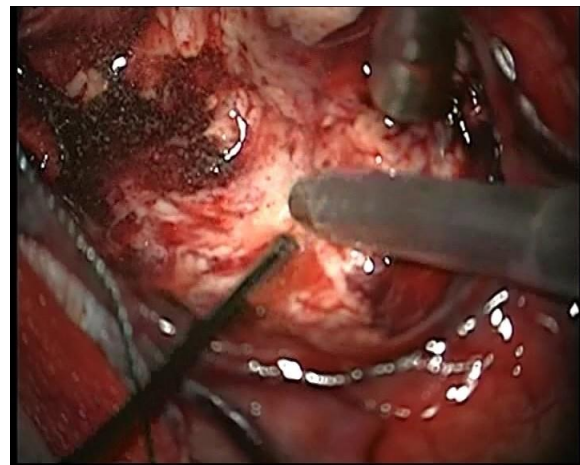
POST-OPERATORIO



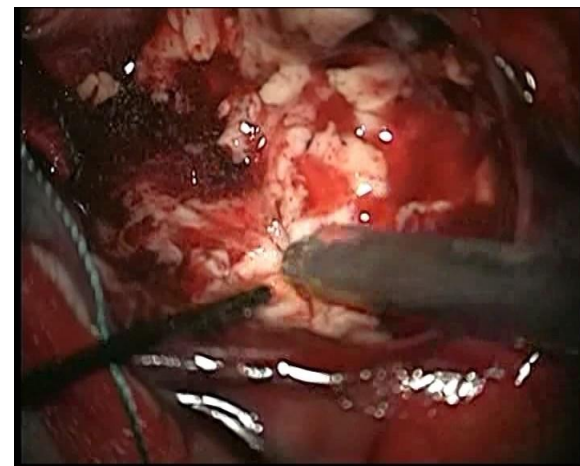
- Astrocitoma diffuso II grado WHO
- **Scomparsa delle crisi**



7 mA



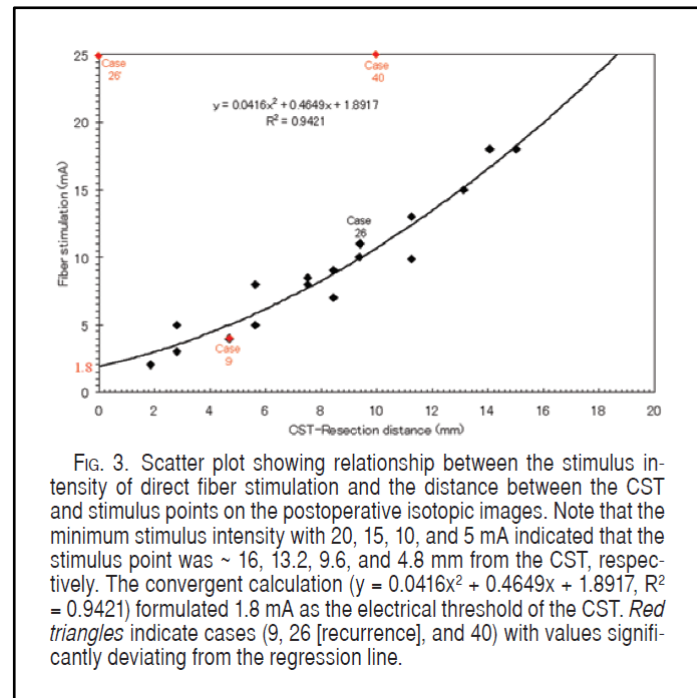
4 mA



The motor-evoked potential threshold evaluated by tractography and electrical stimulation

Clinical article

**KYOUSUKE KAMADA, M.D., PH.D.,¹ TOMOKI TODO, M.D., PH.D.,¹ TAKAHIRO OTA, M.D.,¹
KENJI INO, PH.D.,² YOSHITAKA MASUTANI, PH.D.,² SHIGEKI AOKI, M.D., PH.D.,²
FUMIYA TAKEUCHI, PH.D.,³ KENSUKE KAWAI, M.D., PH.D.,¹
AND NOBUHITO SAITO, M.D., PH.D.¹**



Neurosurgery 71 [ONS Suppl 1]: ons104-ons115, 2012

Low-Threshold Monopolar Motor Mapping for Resection of Primary Motor Cortex Tumors

Kathleen Seidel, MD
Jürgen Beck, MD, PhD
Lennart Stieglitz, MD
Philippe Schucht, MD
Andreas Raabe, MD, PhD

Department of Neurosurgery, Inselspital, Bern University Hospital, Bern, Switzerland

BACKGROUND: Microsurgery within eloquent cortex is a controversial approach because of the high risk of permanent neurological deficit. Few data exist showing the relationship between the mapping stimulation intensity required for eliciting a muscle motor evoked potential and the distance to the motor neurons; furthermore, the motor threshold at which no deficit occurs remains to be defined.
OBJECTIVE: To evaluate the safety of low threshold motor evoked potential mapping for tumor resection close to the primary motor cortex.
METHODS: Fourteen patients undergoing tumor surgery were included. Motor threshold was defined as the stimulation intensity that elicited motor evoked potentials

J Neurosurg 118:287-296, 2013

The warning-sign hierarchy between quantitative subcortical motor mapping and continuous motor evoked potential monitoring during resection of supratentorial brain tumors

Clinical article

KATHLEEN SEIDEL, M.D.,¹ JÜRGEN BECK, M.D., PH.D.,¹ LENNART STIEGLITZ, M.D.,¹ PHILIPPE SCHUCHT, M.D.,¹ AND ANDREAS RAABE, M.D., PH.D.¹

¹Department of Neurosurgery, Inselspital, Bern University Hospital, Bern, Switzerland

J Neurosurg 120:1015-1024, 2014

Continuous dynamic mapping of the corticospinal tract during surgery of motor eloquent brain tumors: evaluation of a new method

Clinical article

ANDREAS RAABE, M.D., JÜRGEN BECK, M.D., PHILIPPE SCHUCHT, M.D., AND KATHLEEN SEIDEL, M.D.

Department of Neurosurgery, Inselspital, Bern University Hospital, Bern, Switzerland

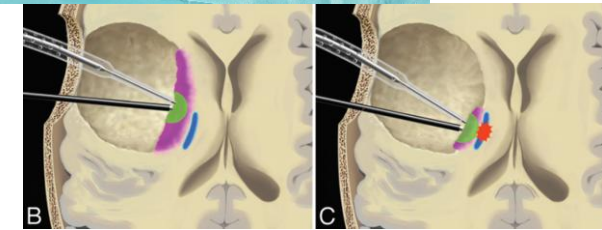
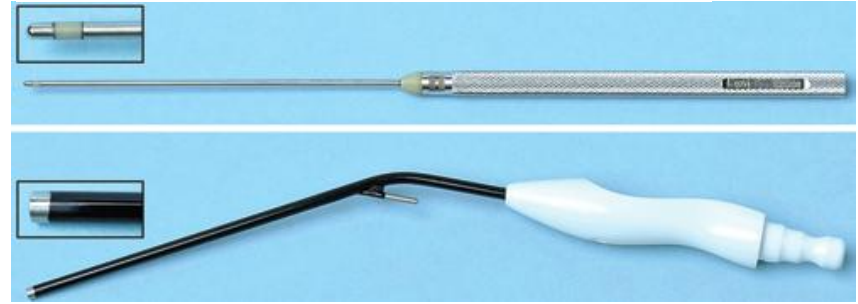
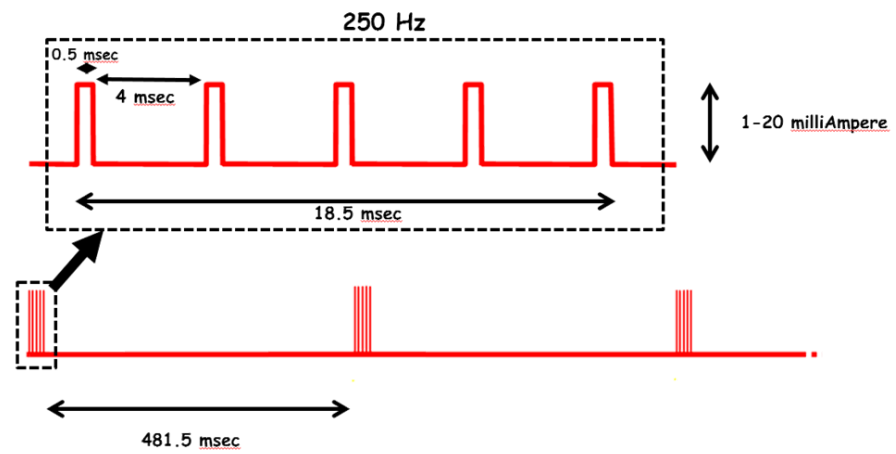
Neurosurg Focus 37 (6):E16, 2014

Intraoperative monopolar mapping during 5-ALA-guided resections of glioblastomas adjacent to motor eloquent areas: evaluation of resection rates and neurological outcome

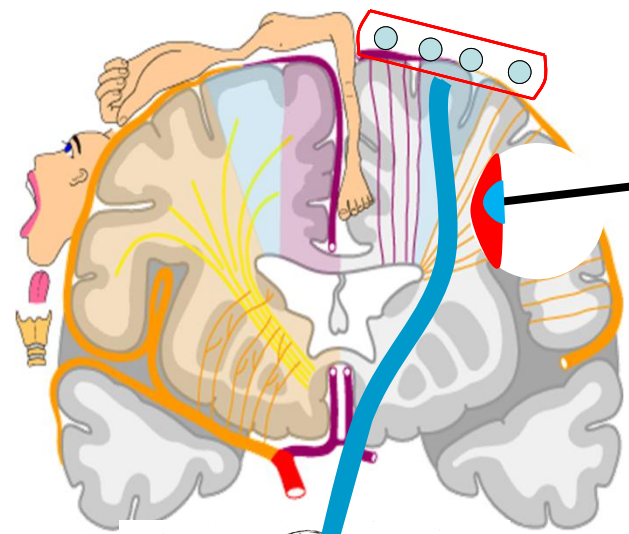
PHILIPPE SCHUCHT, M.D.,¹ KATHLEEN SEIDEL, M.D.,¹ JÜRGEN BECK, M.D.,¹ MICHAEL MUREK, M.D.,¹ ASTRID JILCH, M.D.,¹ ROLAND WIEST, M.D.,² CHRISTIAN FUNG, M.D.,¹ AND ANDREAS RAABE, M.D.¹

Departments of ¹Neurosurgery and ²Neuroradiology, Inselspital, Bern University Hospital, Bern, Switzerland

Monopolar high-frequency motor mapping Train-of-five stimuli

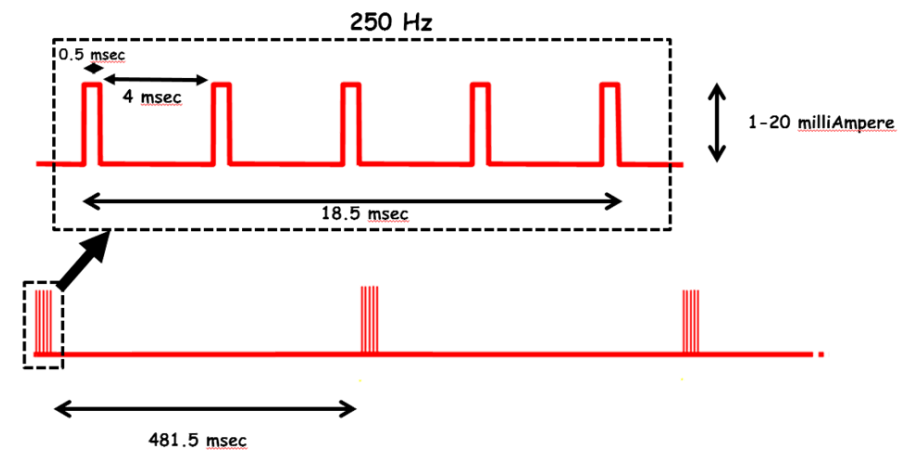
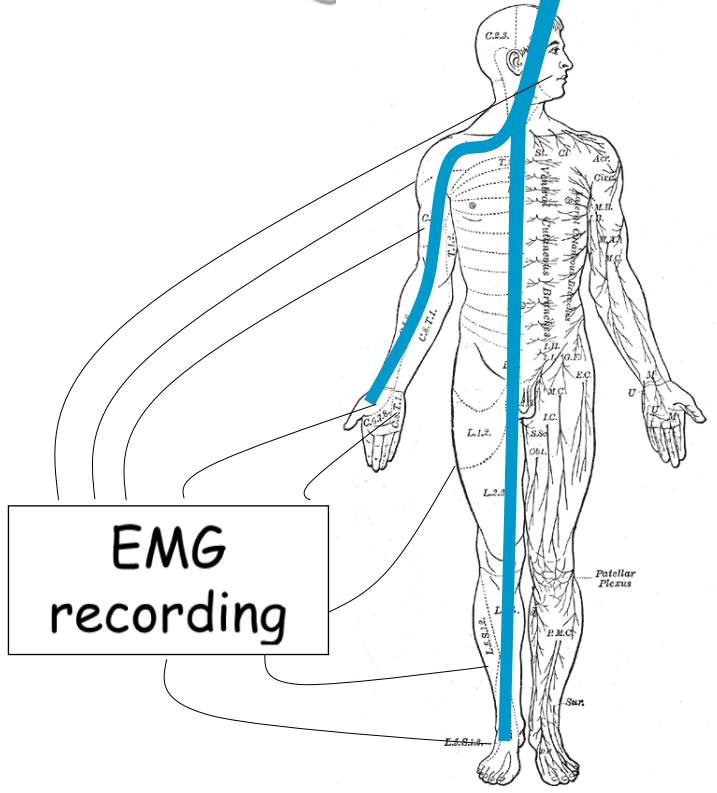


Motor Evoked Potentials

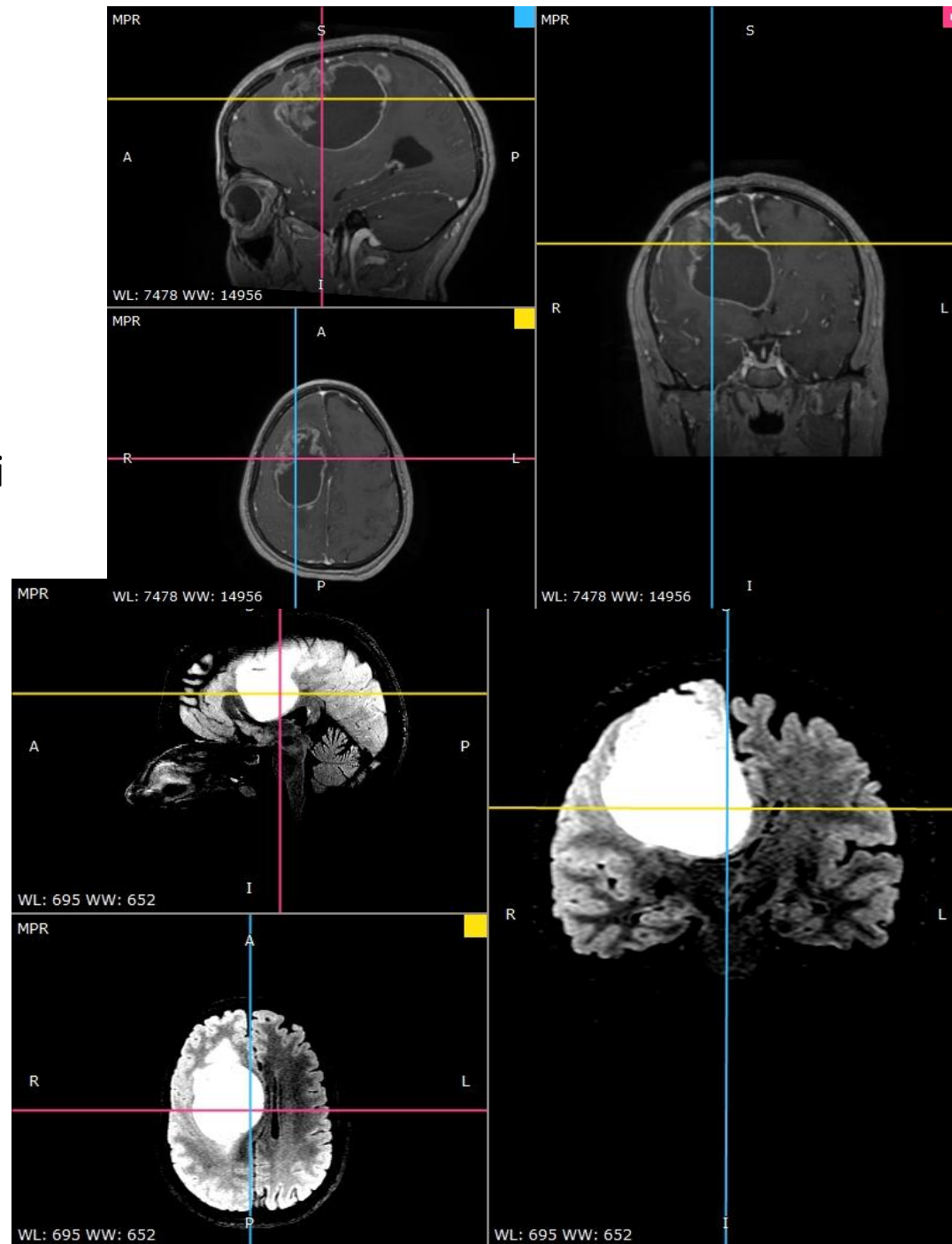
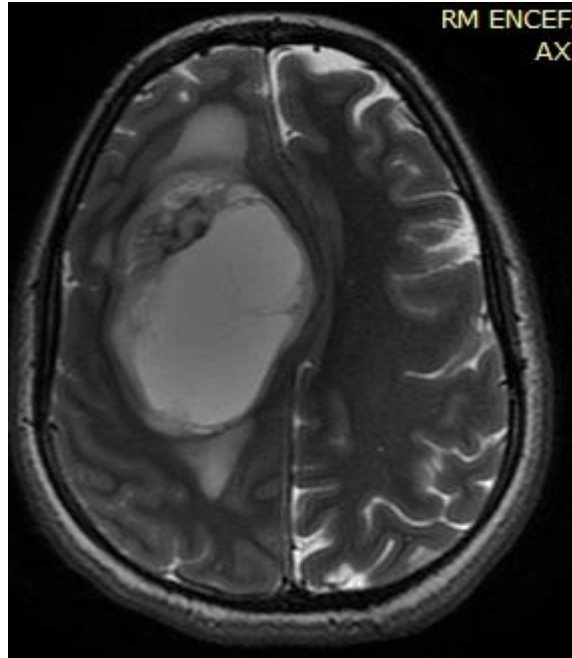


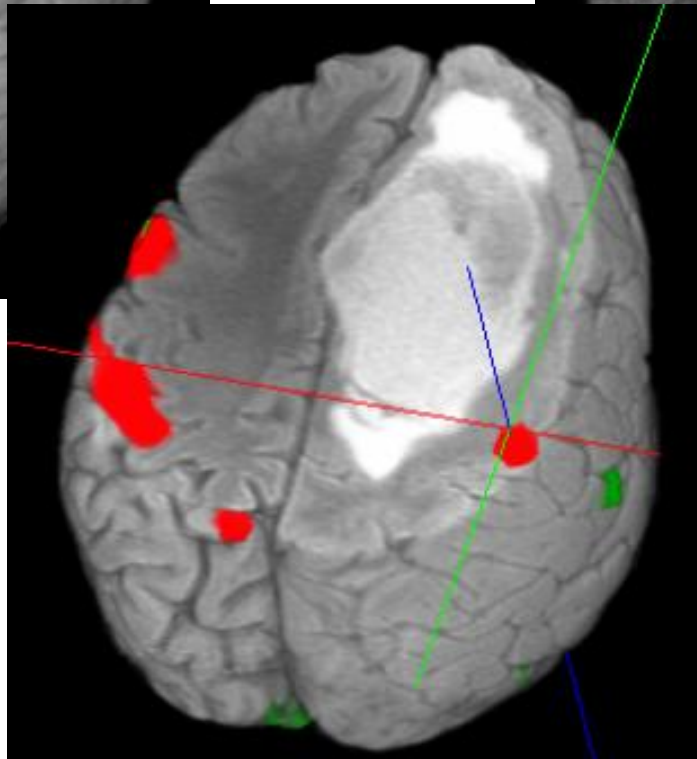
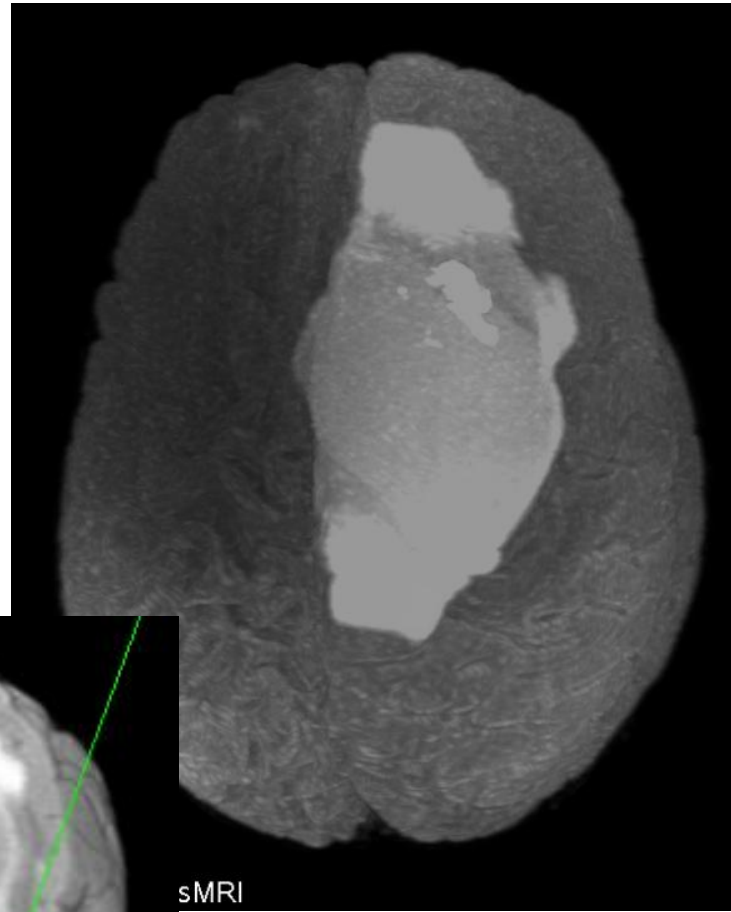
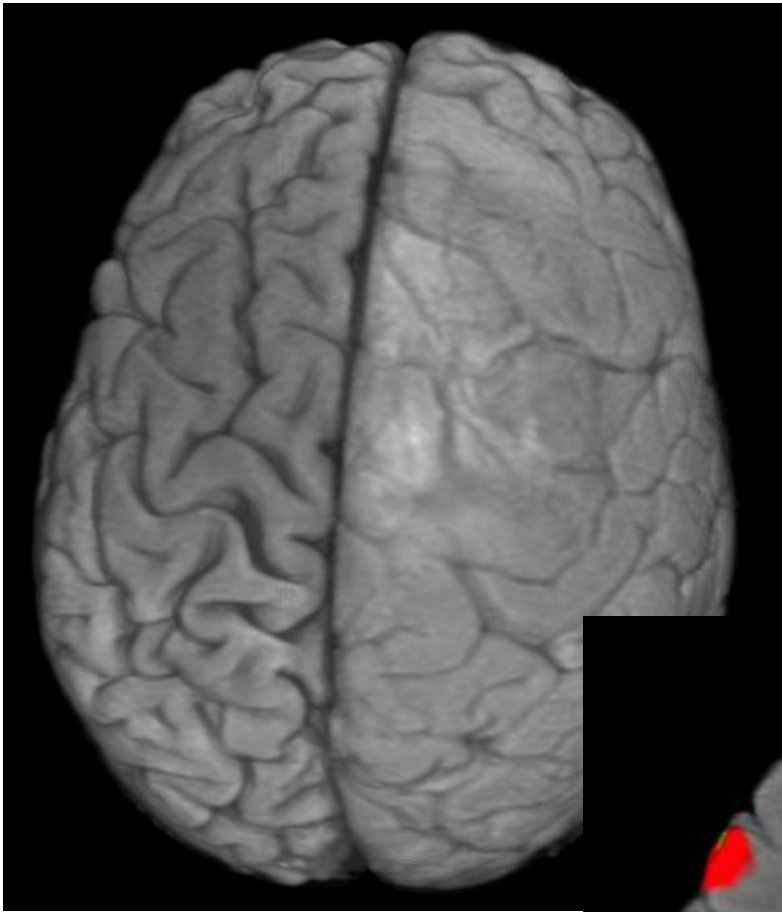
Continuous subcortical mapping

Monopolar high-frequency motor mapping
Train-of-five stimuli

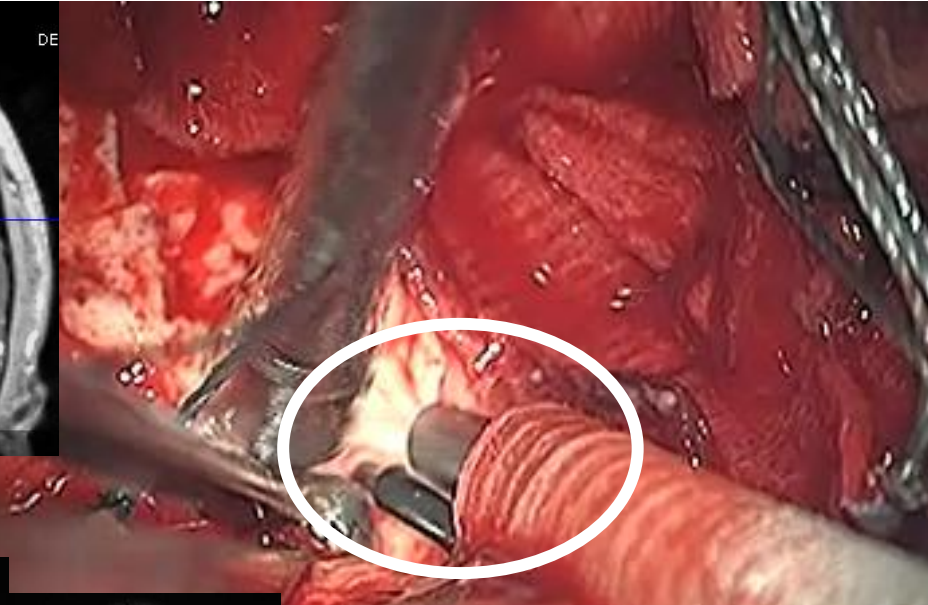
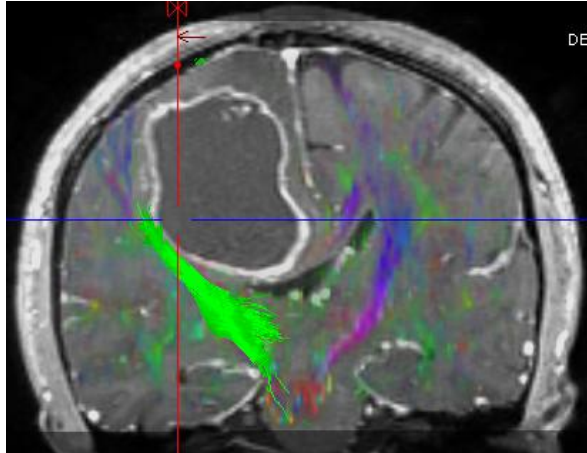
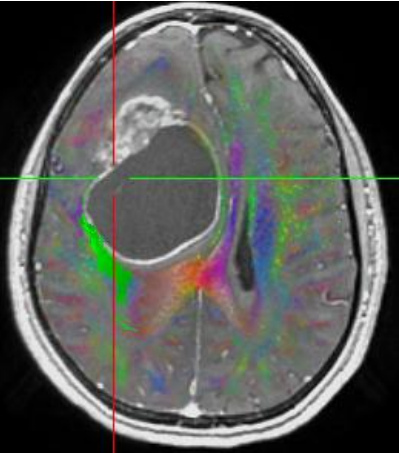


- F, 26 anni
- Già operata 2 anni prima presso altro istituto, asportazione parziale
- Astrocitoma diffuso grado II WHO
- EON: Emiparesi sinistra e crisi epilettiche

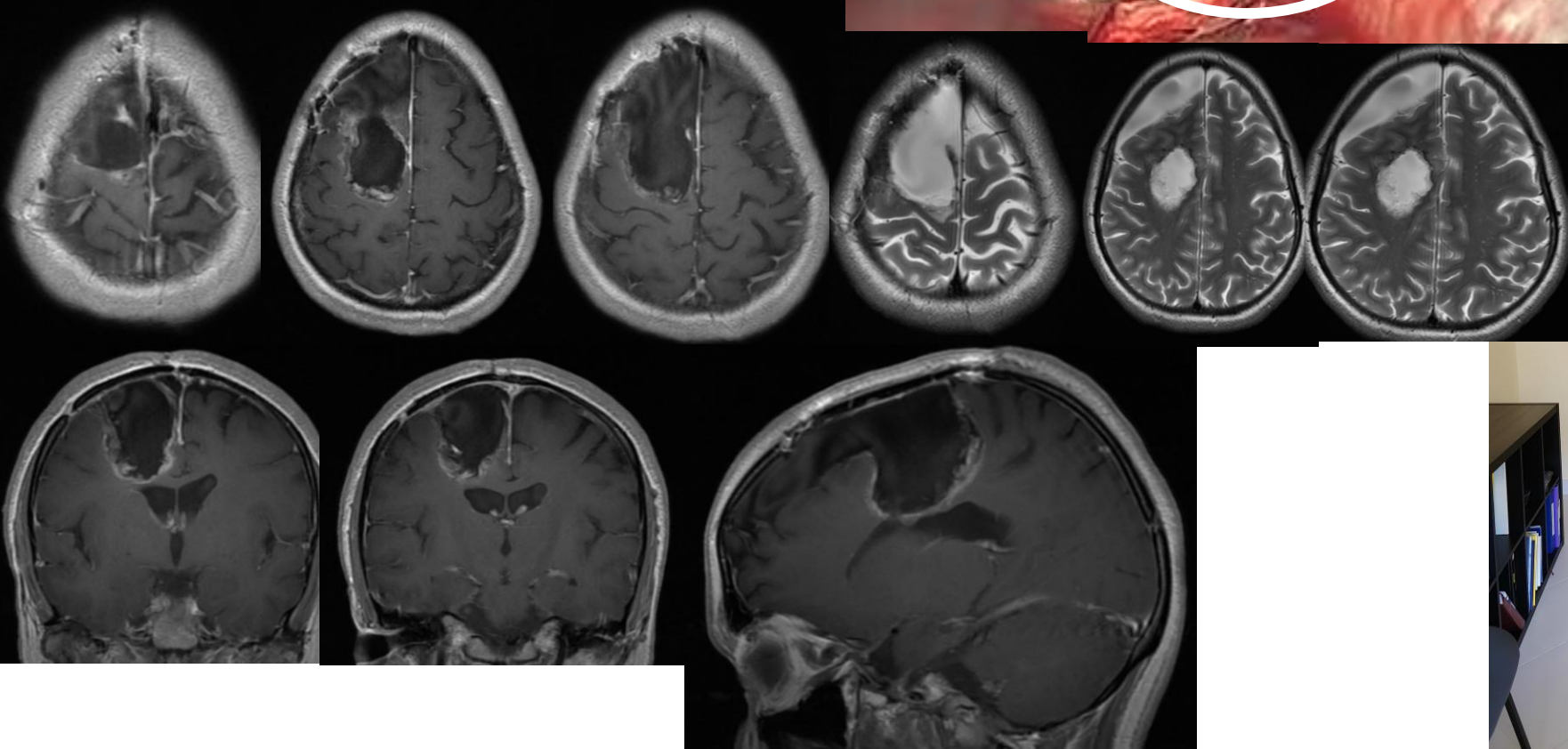




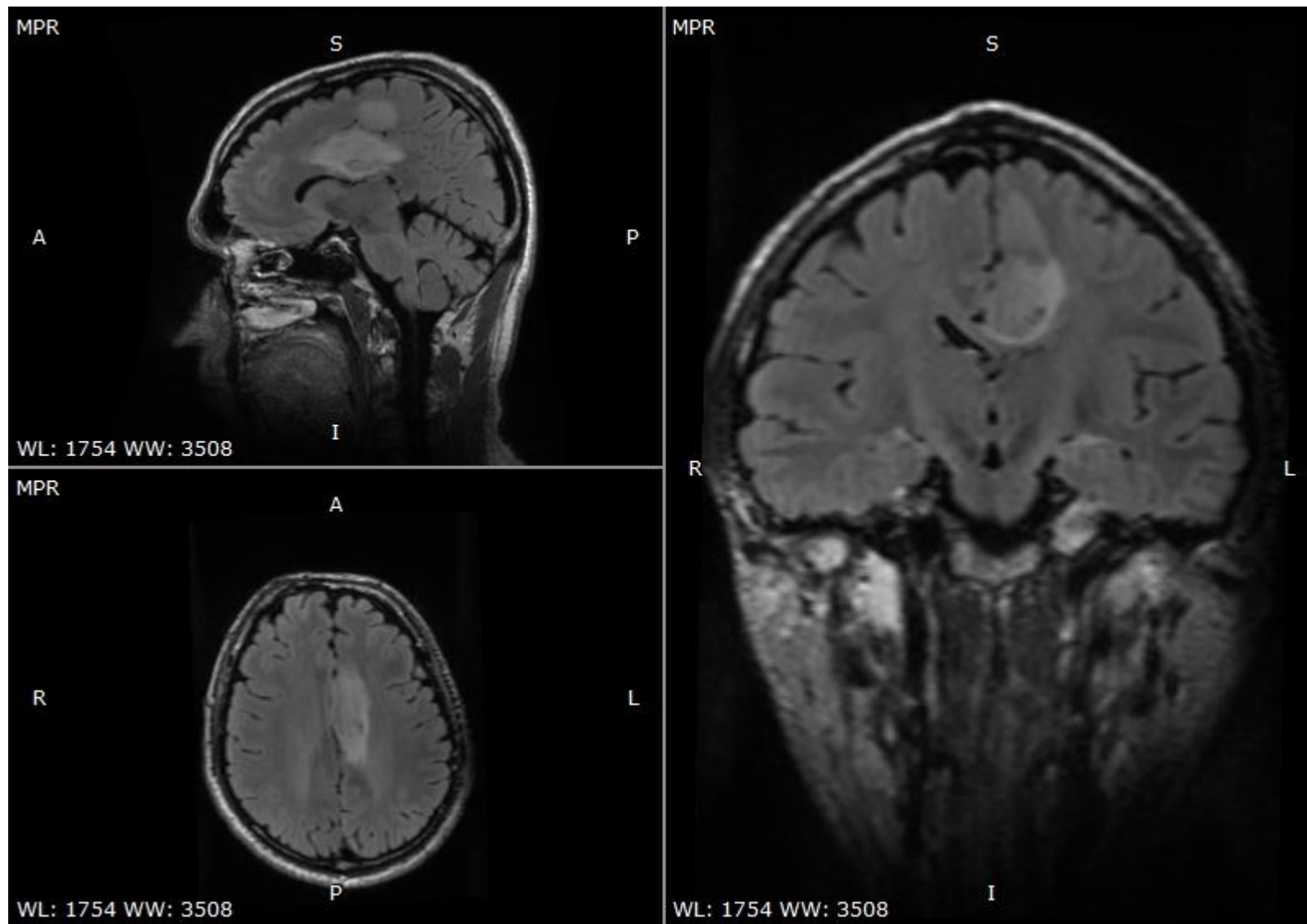
sMRI

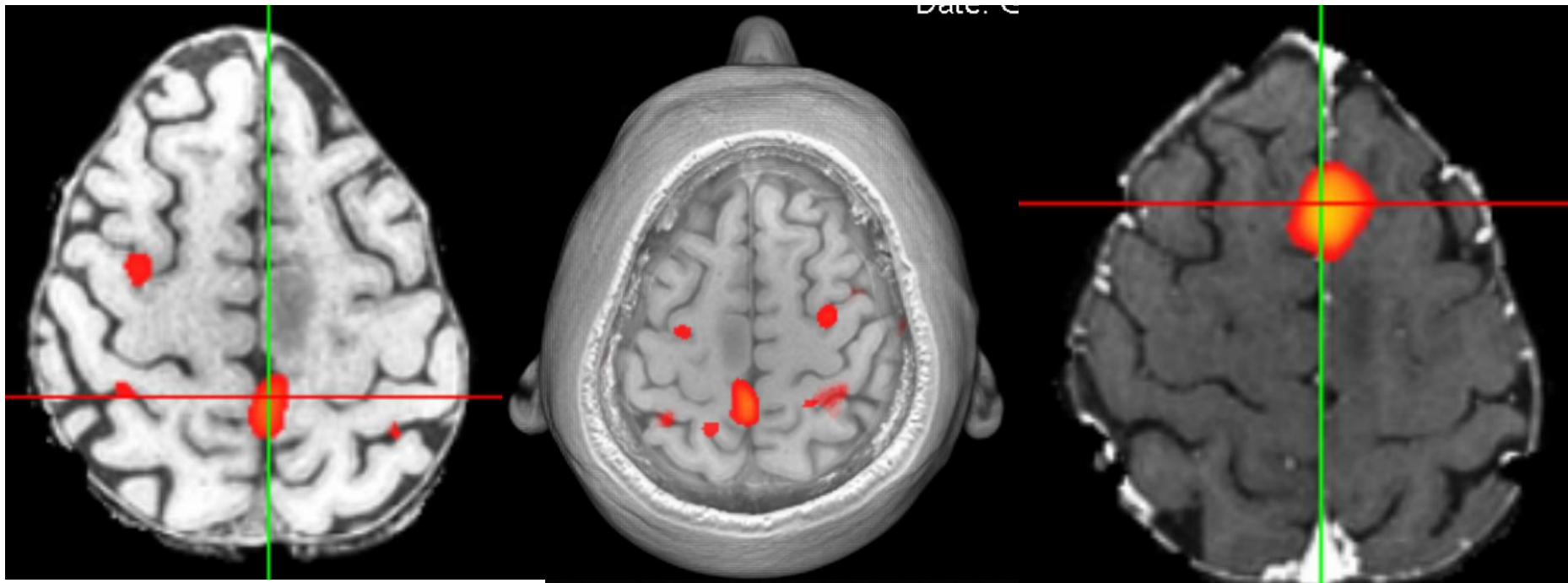


- Peeling del fascio piramidale con **tecnica CUSA – radar** fino a 1 mA
- Transitoria emiparesi sinistra post-operatoria con successivo recupero
- Regressione delle crisi
- **Glioblastoma (grado IV WHO)**

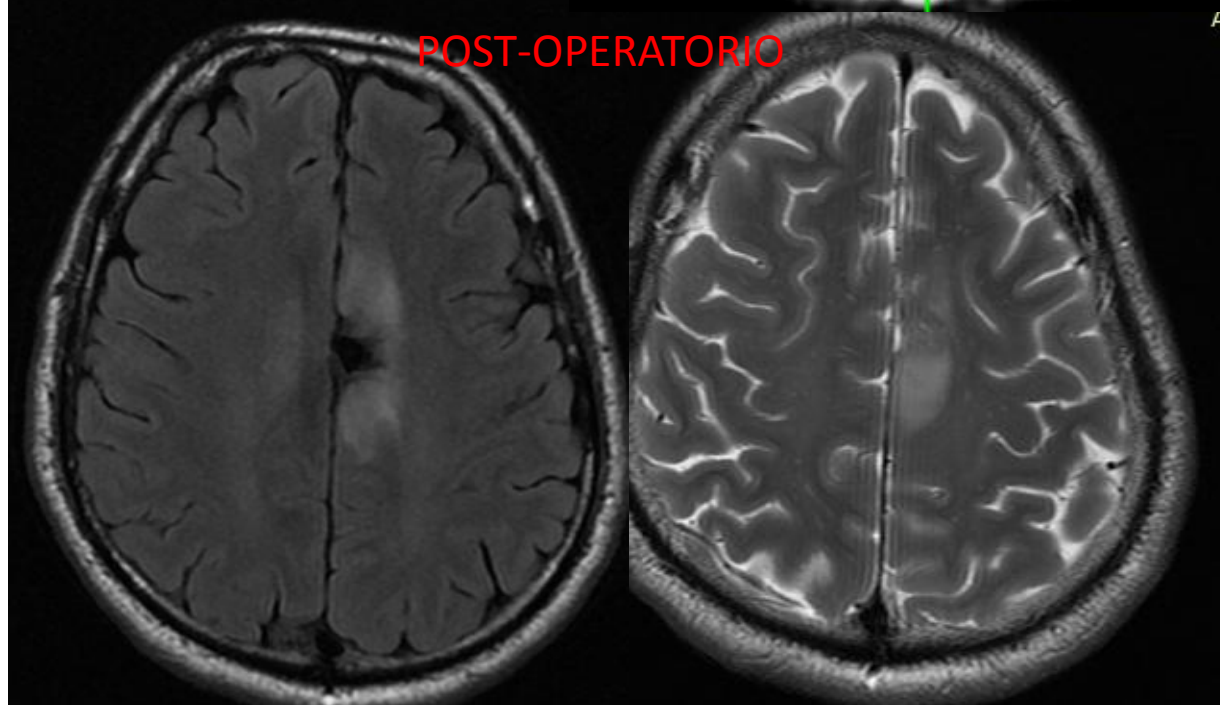


- M, 46 anni
- Esordio con crisi epilettiche

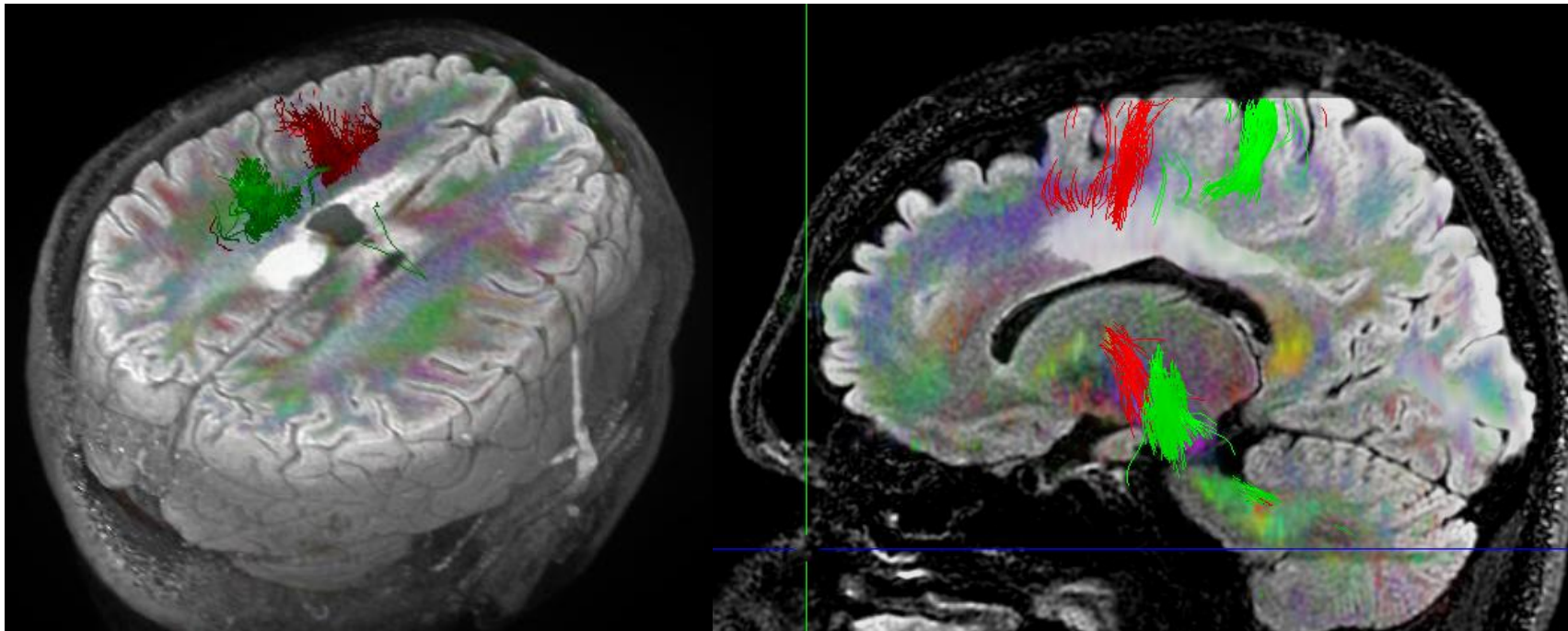




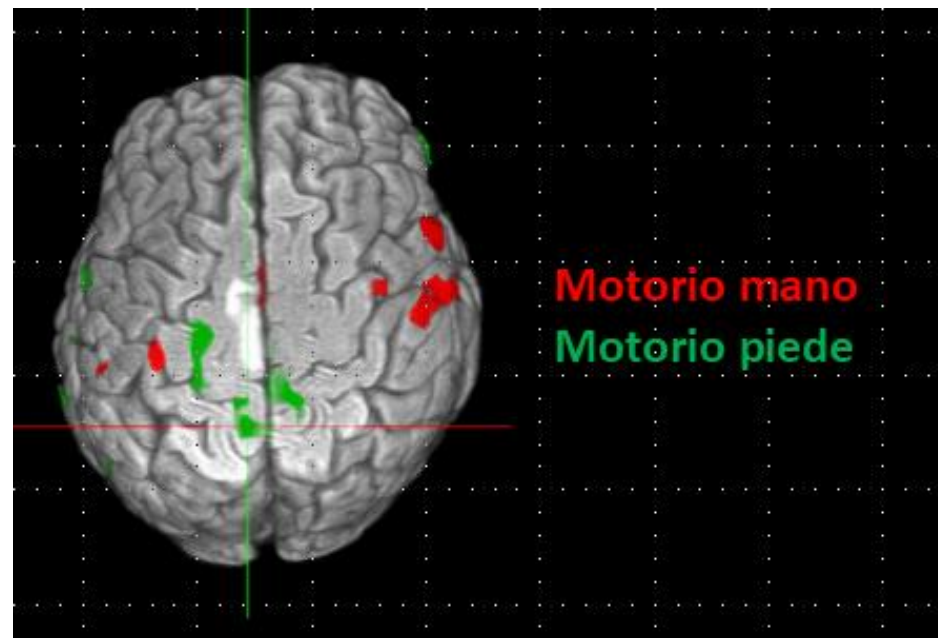
POST-OPERATORIO

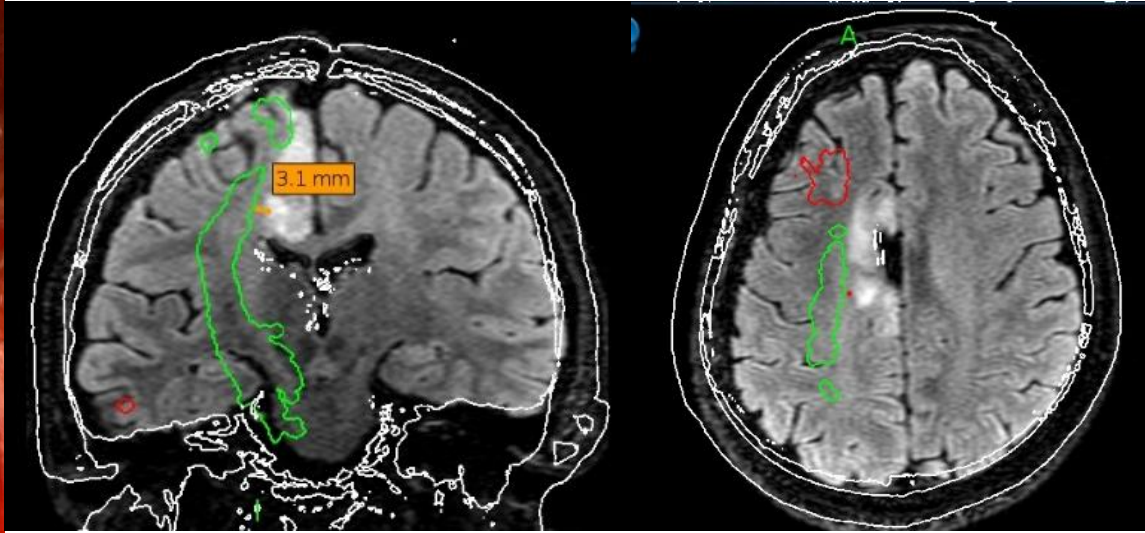
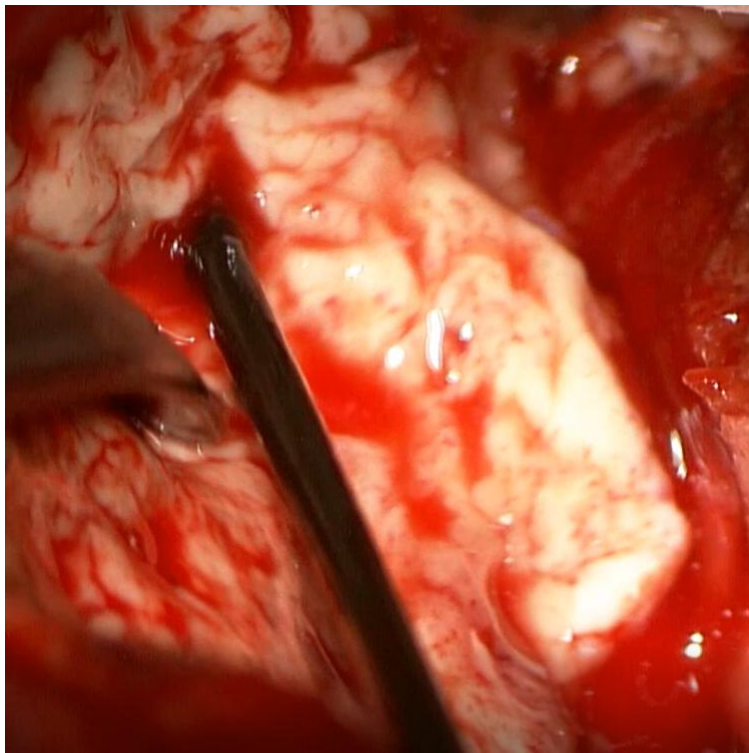


- Asportazione parziale
- Astrocitoma II grado WHO
- Emiparesi destra transitoria ed afasia con completo recupero
- Persistenza delle crisi epilettiche

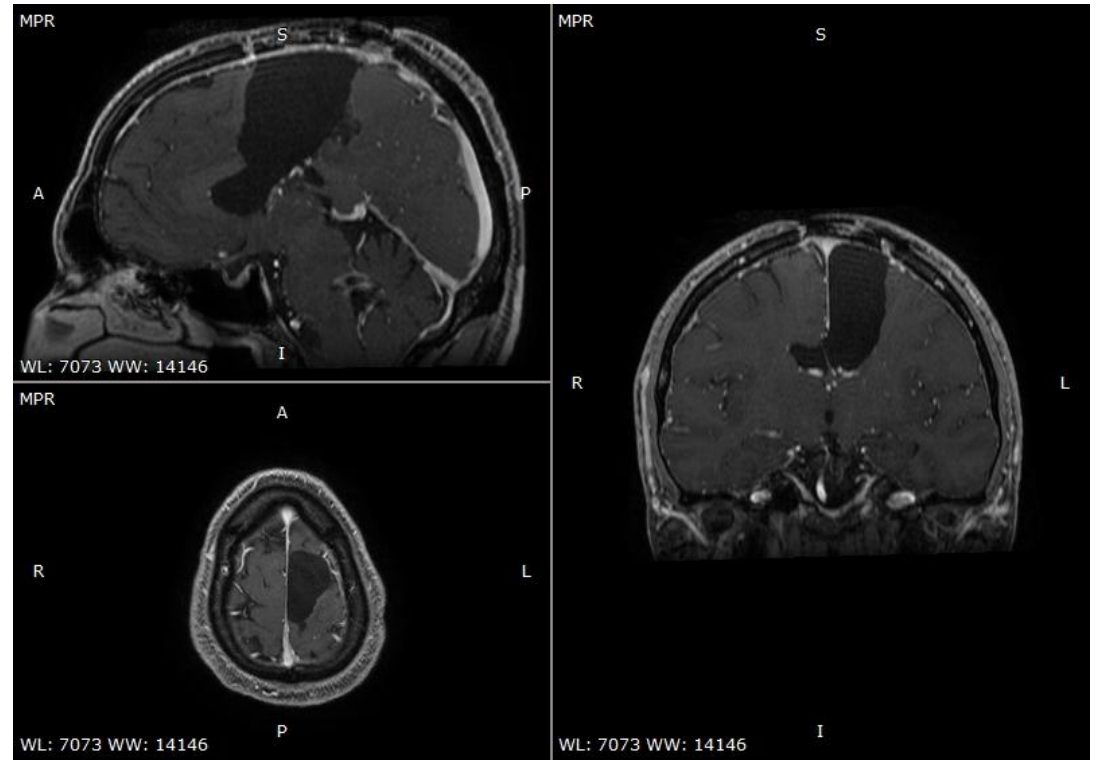


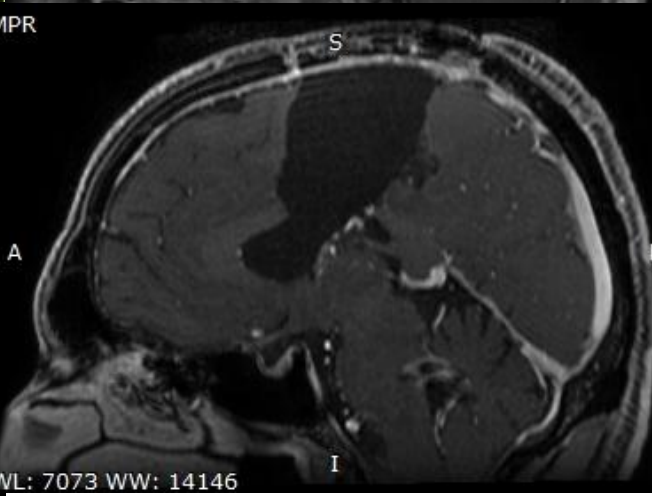
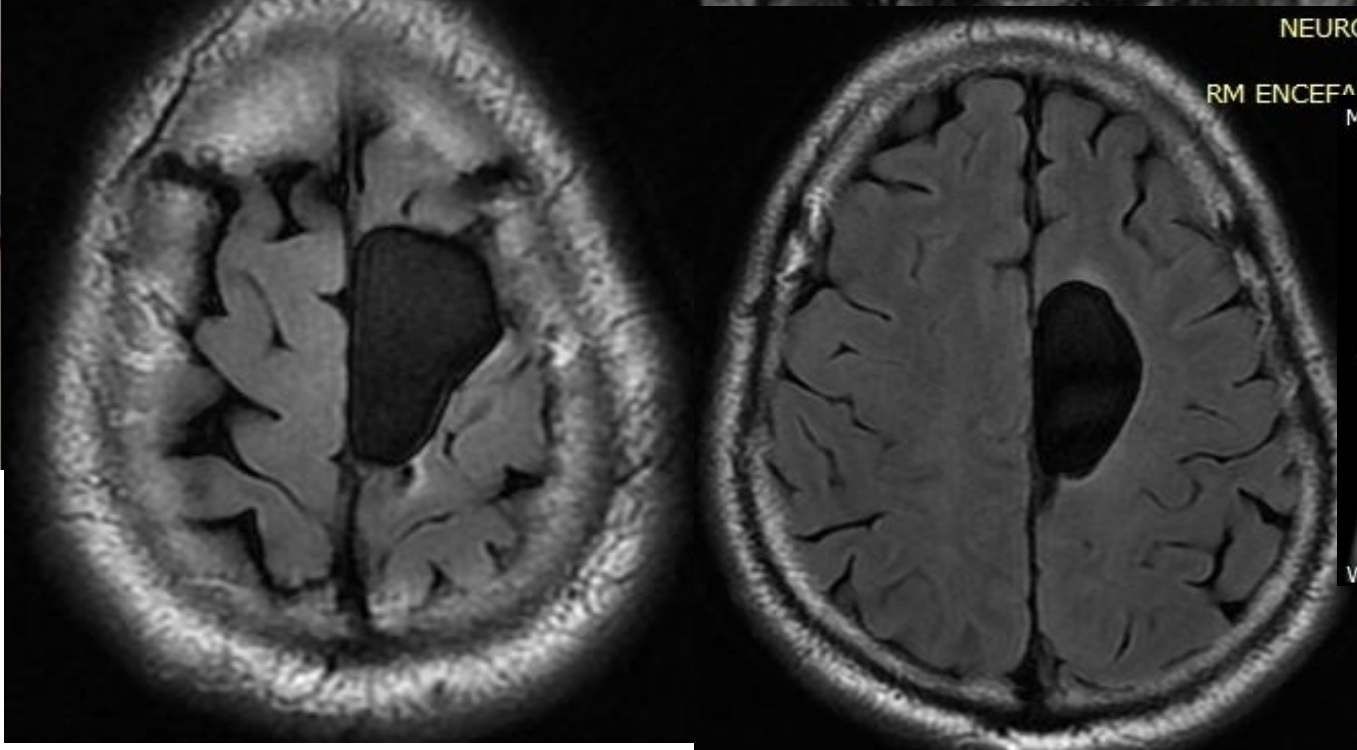
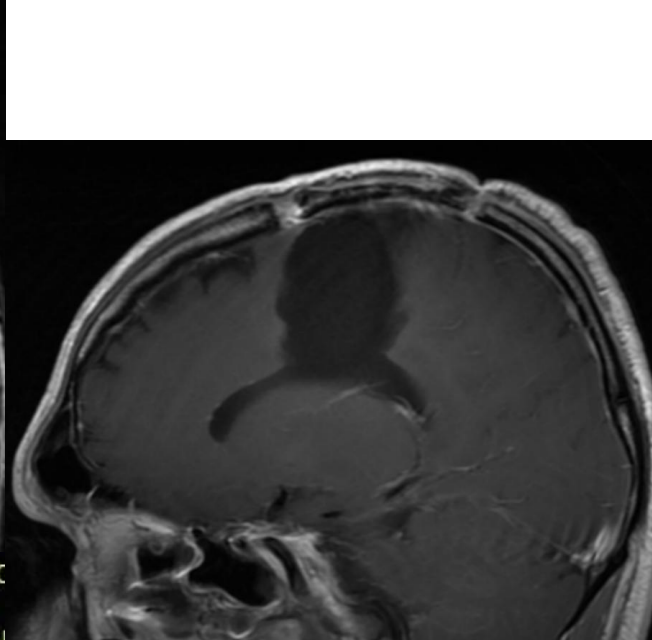
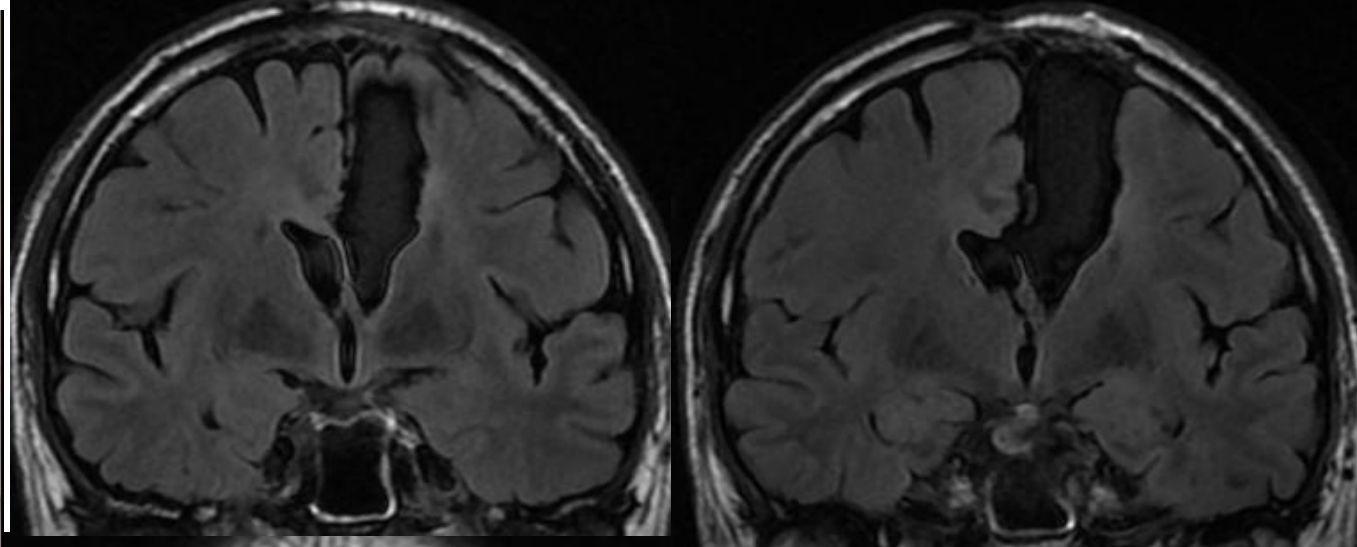
- Aumento volumetrico del residuo tumorale a distanza di 1 anno
- EON: negativo
- **Reintervento**





- Peeling del fascio piramidale con **tecnica CUSA-radar**: fino a 2 mA
- Transitoria emiparesi destra post-operatoria con successivo completo recupero





Scomparsa delle crisi

Integrazione Neuropsicologia-Navigazione DTI-RMN funzionale

Casistica consecutiva settembre 2015 - settembre 2017

N = 82

Asportazione	<p>GTR=59 (72%)</p> <p>Subtot=16 (20%)</p> <p>Parziale/subtot=7(8)%</p>
Clinica postop	<p>Nuovi deficit = 4 (5%)</p> <p>Peggioramento = 26 (31%)</p> <p>Invariato = 41 (47%)</p> <p>Miglioramento = 11 (12%)</p>
Clinica a 1 mese	<p>Nuovi deficit = 4 (5%)</p> <p>Peggioramento = 13 (16%)</p> <p>Invariato = 55 (67%)</p> <p>Miglioramento = 10 (12%)</p>

**Scomparsa delle
crisi epilettiche
nei pazienti con
asportazione
totale:
80%**

2 - LEAT (Long-Term Epilepsy Associated Tumors) o epileptomi

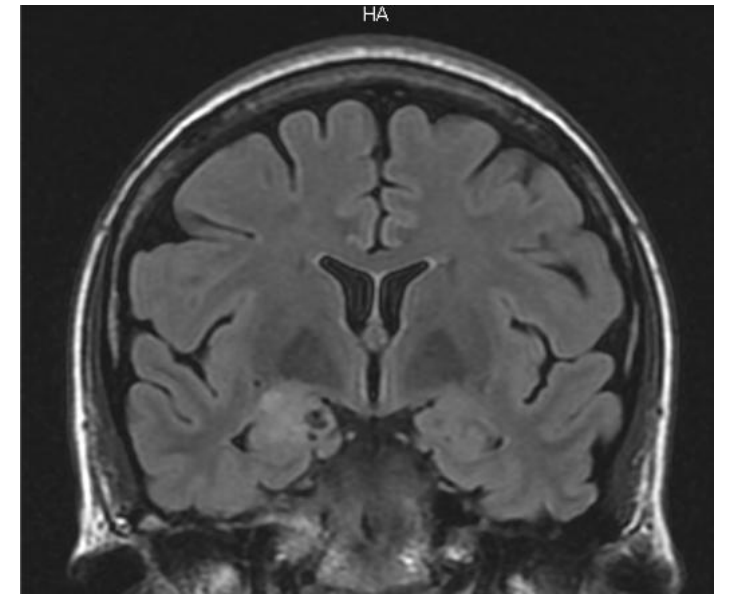
Scarsa evolutività oncologica nella gran maggioranza dei casi

Tumori intrinsecamente epilettogeni

Tipici della giovane età

Quasi sempre nel lobo temporale (circa 80%)

Valutazione in un centro per la Chirurgia dell'Epilessia (non infrequentemente l'asportazione va estesa oltre il tumore per risolvere l'epilessia).

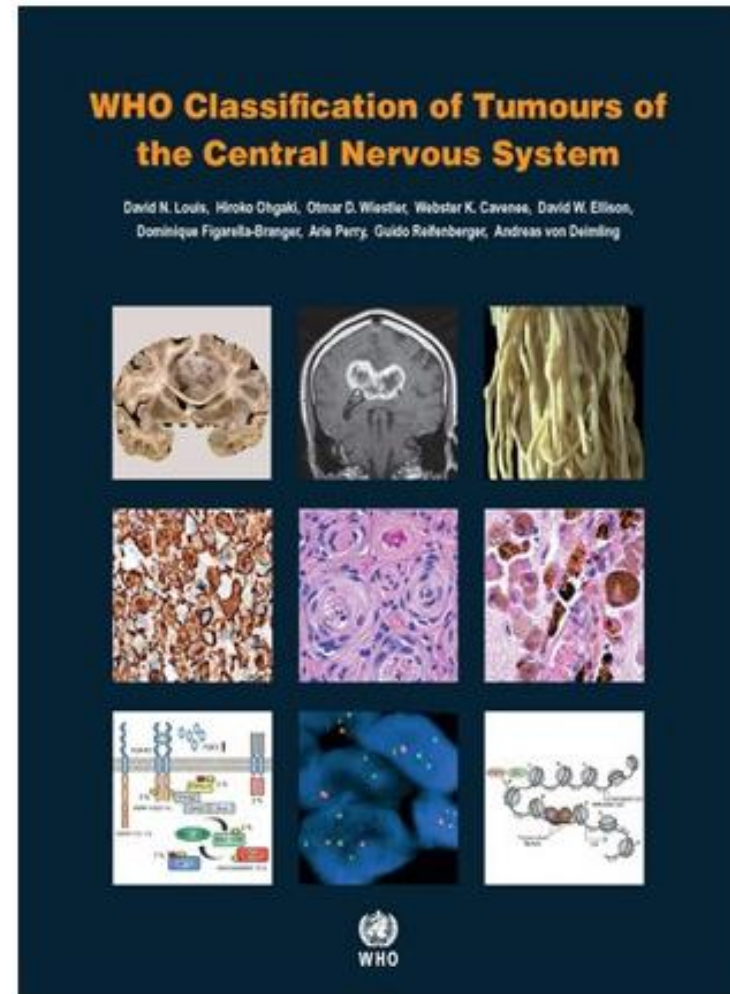


Neuronal and mixed neuronal-glial tumours (WHO Classification)

~ 1,3% DI TUTTI I TUMORI CEREBRALI

Neuronal and mixed neuronal-glial tumours

→ Dysembryoplastic neuroepithelial tumour	9413/0
→ Gangliocytoma	9492/0
→ Ganglioglioma	9505/1
Anaplastic ganglioglioma	9505/3
Dysplastic cerebellar gangliocytoma (Lhermitte–Duclos disease)	9493/0
Desmoplastic infantile astrocytoma and ganglioglioma	9412/1
Papillary glioneuronal tumour	9509/1
Rosette-forming glioneuronal tumour	9509/1
<i>Diffuse leptomeningeal glioneuronal tumour</i>	
Central neurocytoma	9506/1
Extraventricular neurocytoma	9506/1
Cerebellar liponeurocytoma	9506/1
Paraganglioma	8693/1

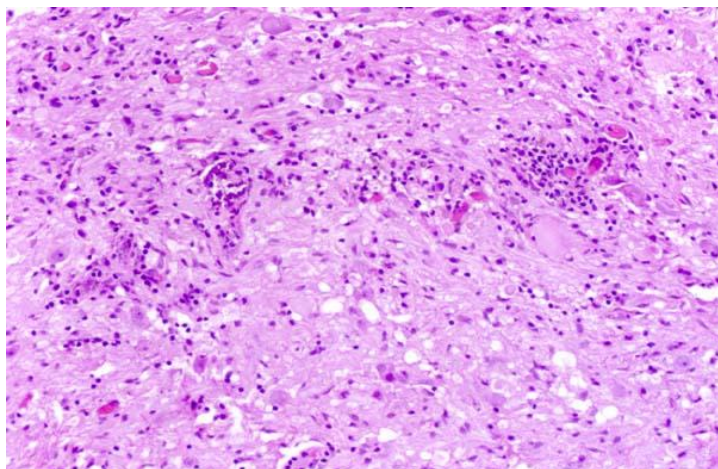


Tumori glioneuronali

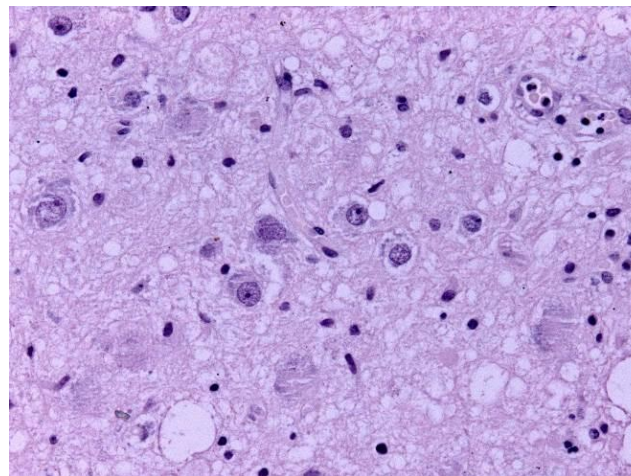
Componente neuronale, o neuronale-gliale

Lenta crescita (grado I o II)

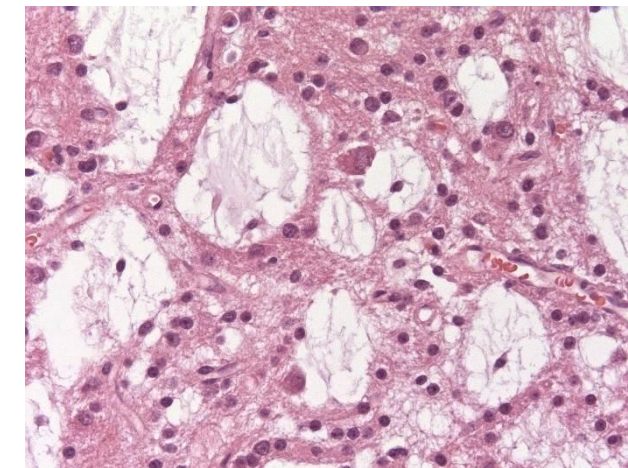
Spesso epilettogeni



Ganglioglioma



Gangliocitoma



**Disembrioneuroepitelioma
(DNET)**

Grado I

Grado I-IV ←

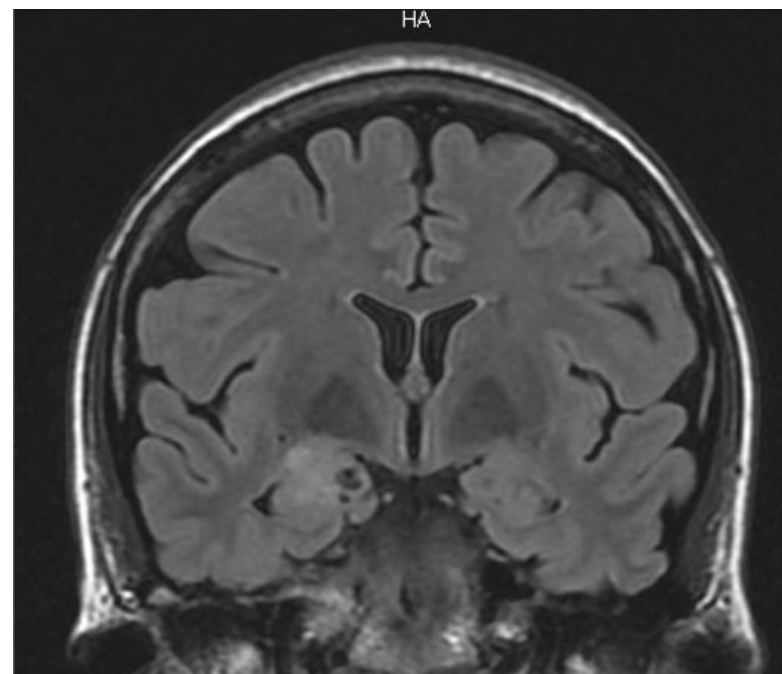
GANGLIOGLIOMI GANGLIOCITOMI DISEMBRIONEUROEPITELIOMI (DNT)

Grado I (nella
maggior parte dei
casi)

PREVALENTE LOCALIZZAZIONE NEL
LOBO TEMPORALE

REGIONI
TEMPORO-MESIALI

Luyken C, Blümcke I, Fimmers R,
Urbach H, Wiestler O, Schramm J
Cancer (2004)



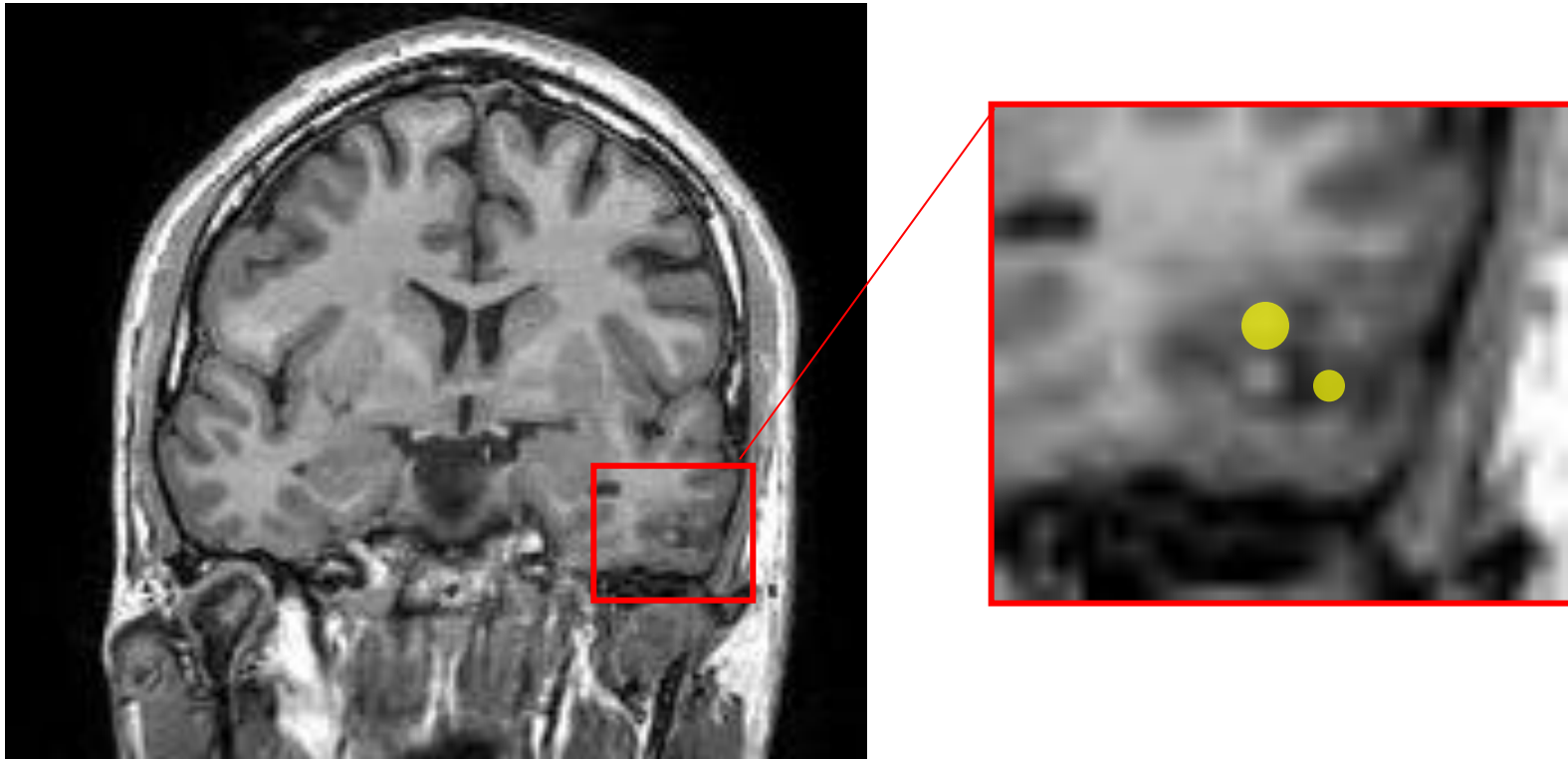
EPILESSIA INTRATTABILE → 90-100%

Rudà R, Trevisan E, Soffietti R (2010) Epilepsy and brain tumors. Curr Opin Oncol
22(6):611-620

TUMORI GLIONEURONALI ED EPILESSIA

MECCANISMI FISIOPATOLOGICI

**ATTIVITA' EPILETTOGENA DELLE COMPONENTI
NEURONALI DEL TUMORE**



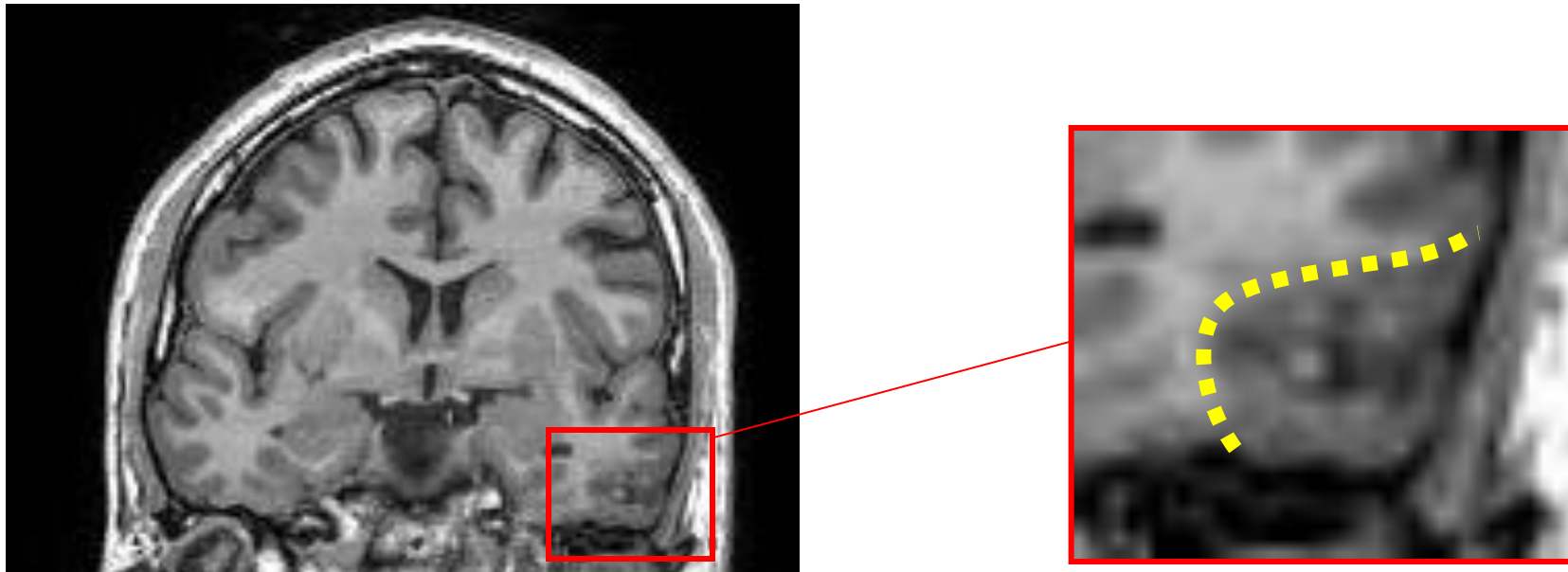
Shamji MF, Fric-Shamji EC, Benoit BG (2009) Brain tumors and epilepsy: pathophysiology of peritumoral changes. *Neurosurg Rev* 32(3):275-285

TUMORI GLIONEURONALI ED EPILESSIA

MECCANISMI FISIOPATOLOGICI

ALTERAZIONI BIOCHIMICHE E STRUTTURALI NEL PARENCHIMA CEREBRALE ADIACENTE ("ZONA DI COLLISIONE")

- ✓ ALTERAZIONE DEI LIVELLI DI NEUROTRASMETTITORI
- ✓ REAZIONI INFIAMMATORIE/DANNO IPOSSICO
- ✓ ACCUMULO DI CELLULE MICROGLIALI

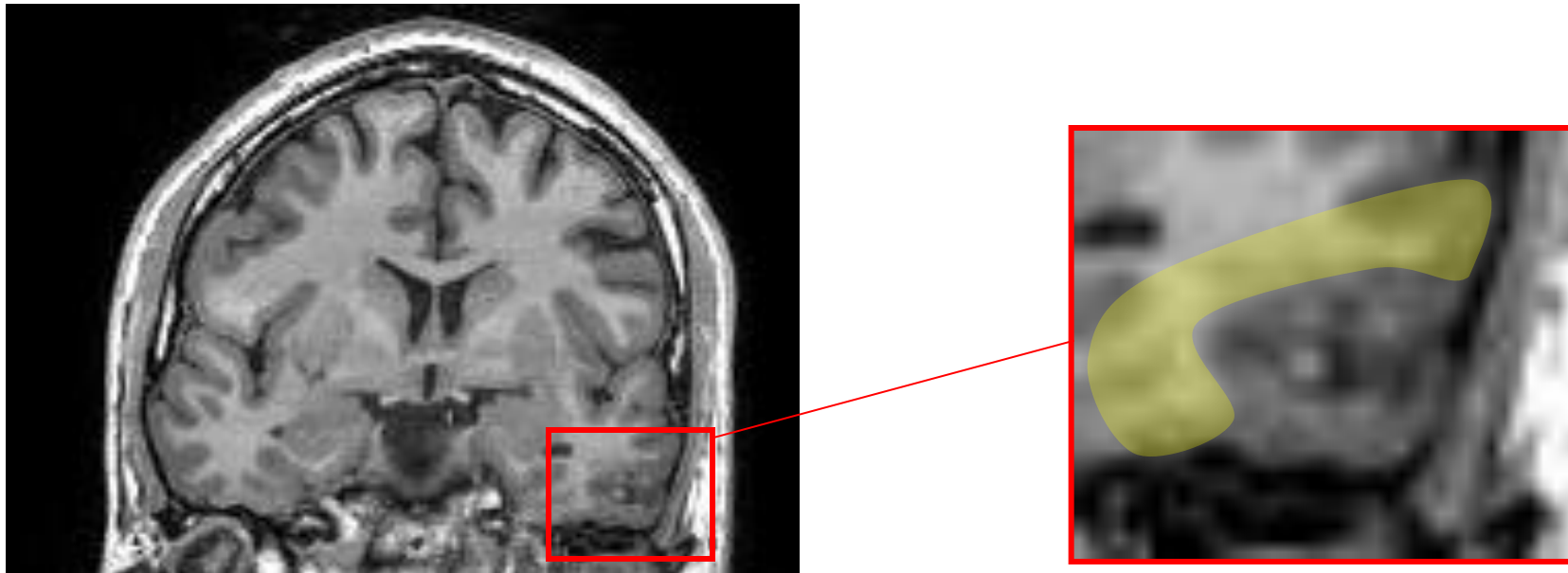


Aronica E, Leenstra S, van Veelen CWM, et al (2001): Glioneuronal tumors and medically intractable epilepsy: a clinical study with long-term follow-up of seizure outcome after surgery. *Epilepsy Res* 43(3):179-191

TUMORI GLIONEURONALI ED EPILESSIA

MECCANISMI FISIOPATOLOGICI

CONCOMITANTE **DISPLASIA PERITUMORALE** (DOUBLE PATHOLOGY)



Rajneesh KF, Binder DK (2009) Tumor-associated epilepsy. Neurosurg Focus 27(2):E4

TUMORI GLIONEURONALI ED EPILESSIA

STRATEGIE CHIRURGICHE

Epilepsia, 53(1):51–57, 2012
doi: 10.1111/j.1528-1167.2011.03269.x

FULL-LENGTH ORIGINAL RESEARCH

Factors associated with seizure freedom in the surgical resection of glioneuronal tumors

Dario J. Englot, Mitchel S. Berger, Nicholas M. Barbaro, and Edward F. Chang

Department of Neurological Surgery, University of California, San Francisco, California, U.S.A.

Table 1. Studies included in analysis

Alexiou et al. (2009)	Lee et al. (2000)
Aronica et al. (2001)	Lombardi et al. (1997)
Bauer et al. (2007)	Luyken et al. (2003)
Benifla et al. (2006)	Minkin et al. (2008)
Bilginer et al. (2009)	Morioka et al. (2007)
Cataltepe et al. (2005)	Morris et al. (1998)
Chan et al. (2006)	Nolan et al. (2004)
Chang et al. (2010)	Ogiwara et al. (2010)
Choi et al. (2004)	Panda et al. (2005)
Devaux et al. (1997)	Park et al. (2008)
Drake et al. (1987)	Pilcher et al. (1993)
Giulioni et al. (2005)	Radhakrishnan et al. (2006)
Giulioni et al. (2006)	Raymond et al. (1995)
Giulioni et al. (2009)	Sandberg et al. (2005)
Jooma et al. (1995)	Sharma et al. (2009)
Kameyama et al. (2001)	Tran et al. (1997)
Khajavi et al. (1994)	Wennberg et al. (1999)
Khajavi et al. (1999)	Zaatreh et al. (2003)
Kim et al. (1995)	Zentner et al. (1997)
Kirkpatrick et al. (1993)	

39 STUDI

TOT. 910 PAZIENTI

Table 2. Seizure outcomes stratified across factors of interest

	Engel I	Engel II-IV	χ^2	p-value
Patient age				
<18 years old	215 (83)	44 (17)	0.53	0.49
≥18 years old	104 (80)	26 (20)		
Tumor location				
Temporal	519 (81)	124 (19)	1.51	0.229
Extratemporal	155 (77)	47 (23)		
Pathologic diagnosis				
Ganglioglioma	409 (78)	119 (23)	3.41	0.067
DNET	315 (83)	67 (18)		
Seizure control ^a				
Controlled	46 (84)	9 (16)	0.45	0.601
Refractory	589 (80)	148 (20)		
Seizure semiology				
Partial only	181 (87)	27 (13)	13.00	<0.001 ^b
Generalized/mixed	147 (73)	55 (27)		
Duration of epilepsy				
≤1 year	63 (97)	2 (3)	13.71	<0.001 ^b
>1 year	246 (77)	74 (23)		
Extent of resection				
Gross-total	552 (87)	80 (13)	79.60	<0.001 ^b
Subtotal	88 (55)	72 (45)		
Intraoperative ECoG				
Used	124 (84)	23 (16)	1.97	0.200
Not used	441 (79)	116 (21)		
Total	724 (80)	186 (20)		

Number of patients (%) across all studies that were seizure-free (Engel class I) or continued to have seizures (Engel class II-IV) in each group postoperatively.

^aSeizures medically controlled or refractory preoperatively.

^bSignificant value (p < 0.02).

Table 3. Seizure outcomes stratified by extent of resection and location in temporal lobe tumors

	Engel I	Engel II-IV	OR (95% CI)	p-value
(A) Extent of resection ^a				
Subtotal	28 (37)	47 (63)	0.16 (0.09-0.29)	<0.001 ^b
lesionectomy				
Gross-total	178 (78)	49 (22)	1 [Reference]	-
lesionectomy (GTR)				
GTR + hippocampectomy	64 (94)	4 (6)	4.40 (1.53-12.69)	<0.01 ^b
GTR + corticectomy	38 (95)	2 (5)	5.23 (1.21-22.45)	0.01 ^b
GTR + hippocampectomy + corticectomy	146 (90)	16 (10)	2.51 (1.37-4.60)	<0.01 ^b
(B) Location in temporal lobe				
Mesial temporal lobe	121 (82)	27 (18)	1 [Reference]	-
Lateral temporal lobe	58 (81)	14 (19)	0.92 (0.45-1.89)	0.85
Mixed	15 (65)	8 (35)	0.42 (0.16-1.09)	0.09
Not specified	260 (79)	69 (21)	0.84 (0.51-1.38)	0.29
Total	454 (79)	118 (20)		

Data shown are number of patients (%).

^aNote: Ns not equal to data in Table 2, as only temporal lobe tumors with extent of resection data are included here.

^bSignificant value (p < 0.02).

Fattori prognostici sull'andamento postoperatorio dell'epilessia

Favorevoli

- Durata dell'epilessia uguale o inferiore a un anno
- Lesionectomia totale
- Resezioni chirurgiche estese (amigdalo-ippocampectomia e/o corticectomia) nelle lesioni del lobo temporale

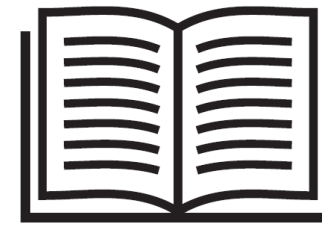
Sfavorevoli

- Crisi epilettiche preoperatorie con generalizzazione secondaria




I fattori critici sul risultato epilettologico sono:

- **intervento precoce**
- **resezione totale**

FULL-LENGTH ORIGINAL RESEARCH



Epilepsy surgery of “low grade epilepsy associated neuroepithelial tumors”: A retrospective nationwide Italian study

¹Marco Giulioni, ^{2,3}Gianluca Marucci , ⁴Veronica Pelliccia, ⁴Francesca Gozzo, ⁵Carmen Barba , ⁶Giuseppe Didato, ⁶Flavio Villani, ⁷Giancarlo Di Gennaro, ⁷Pier Paolo Quarato, ^{7,8}Vincenzo Esposito, ⁹Alessandro Consales, ^{1,10}Matteo Martinoni, ¹Gianfranco Vornetti, ¹¹Corrado Zenesini, ¹²Carlo Efisio Marras, ¹³Nicola Specchio, ¹³Luca De Palma, ¹⁴Raffaele Rocchi, ¹⁵Flavio Giordano , ¹⁶Giovanni Tringali, ¹⁷Paolo Nozza, ¹⁸Gabriella Colicchio, ^{19,20}Guido Rubboli , ⁴Giorgio Lo Russo, ^{5,21}Renzo Guerrini, ^{20,22}Paolo Tinuper, ⁴Francesco Cardinale, and ⁴Massimo Cossu, On behalf of the Commission for Epilepsy Surgery of the Italian League Against Epilepsy

Epilepsia, **(*) :1–10, 2017
doi: 10.1111/epi.13866

339 consecutive patients with LEATs who underwent surgery between January 2009 and June 2015. Epilepsy surgery of LEATs led to a favorable seizure outcome in **88%** of drug-resistant patients and in **98%** of drug-responsive patients
Younger age at surgery, temporal resection site, and complete tumor removal are predictors of a favorable seizure outcome in refractory epilepsy
A **timely surgical treatment**, oriented to optimize epileptologic, neuropsychological, and oncologic outcome should be nowadays warranted

NEUROMED 2002-2017

103 pazienti

Tipo	N.	Età Media	Temporale	Epilessia	Durata Epil.
Gangliogliomi	70	29	52	53	16
DNT	21	22	14	18	10
Gangliocitomi	12	39	6	5	13
Tot.	103	30	72 (69%)	76 (74%)	13

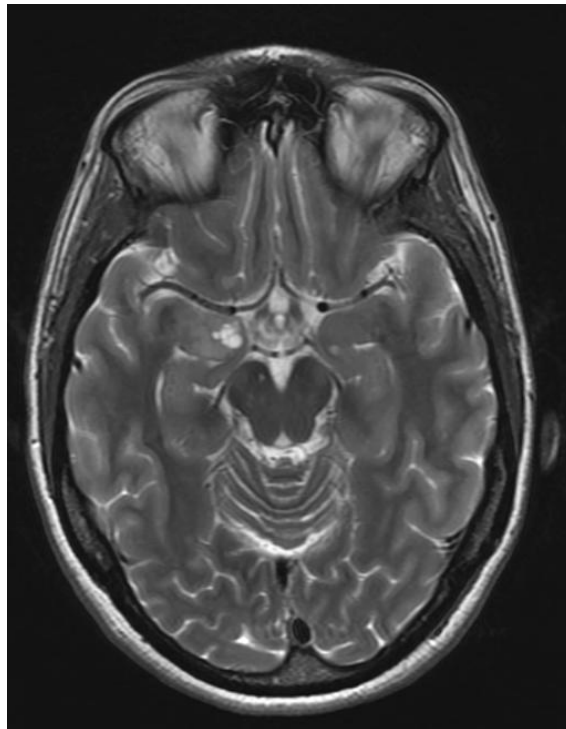
GANGLIOGLIOMA	I WHO*	65
GANGLIOGLIOMA	III WHO	5
DNT	I WHO	21
GANGLIOCITOMA	I WHO	12
RESEZIONE TOTALE		103

*2 Reinterventi per recidiva, 2^a diagnosi: ganglioglioma grado I

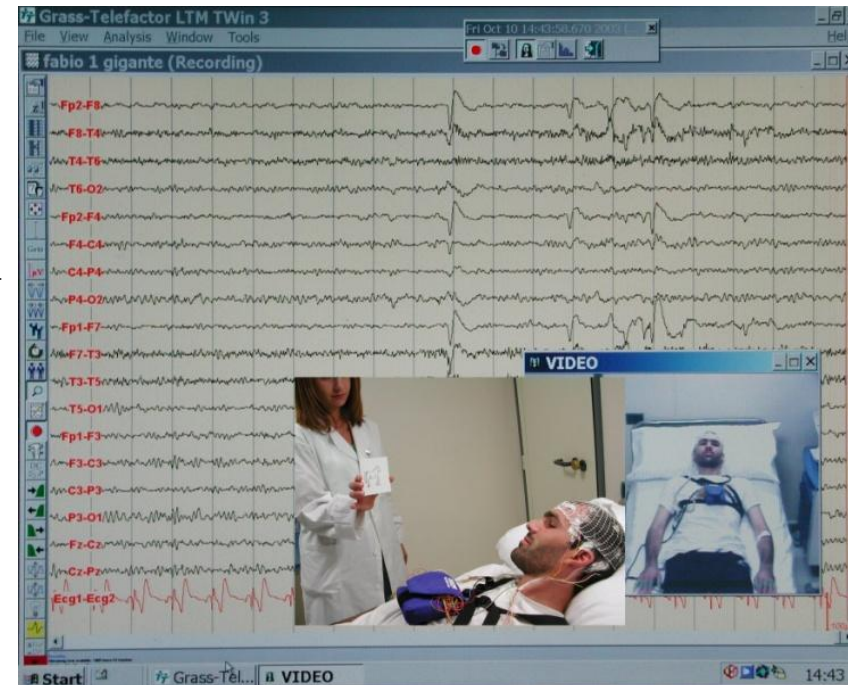
- Morbidità Permanente 1 emiparesi brachio-cruale lieve (1 %)
- Mortalità 0
- Engel I
(completa assenza di crisi) **85%**

OBBLIGATORIE NEI PAZIENTI CON EPILESSIA FARMACORESISTENTE

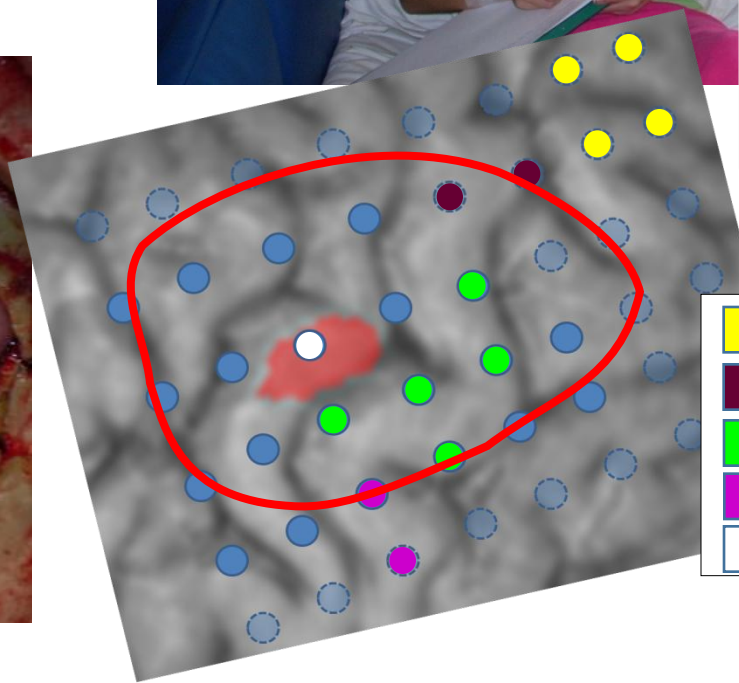
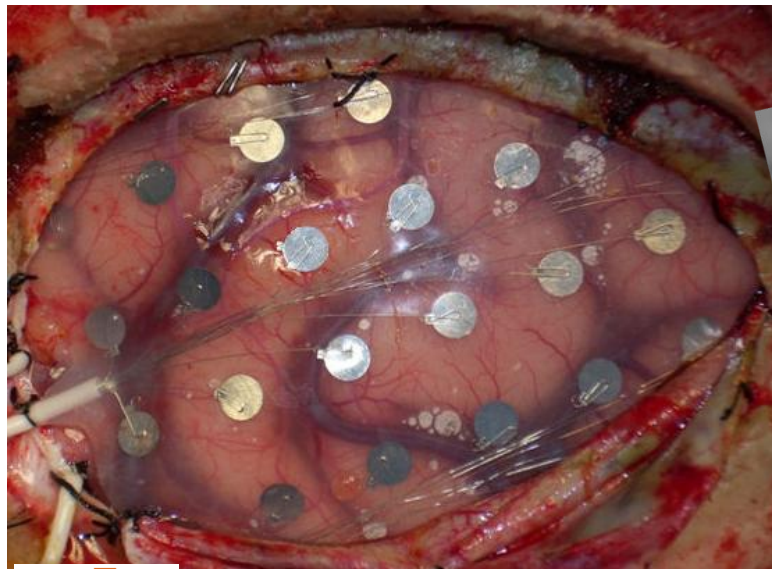
Neuroimaging



Video-EEG



Valutazione neuropsicologica



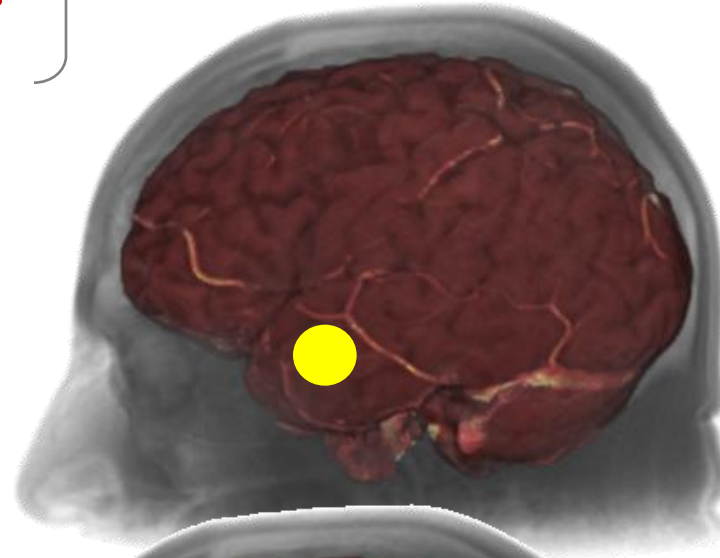
- Arresto del linguaggio
- Movim. Faccia
- Movim. Mano
- Biceps brachialis
- Zona Epilettogena

PROCEDURE CHIRURGICHE EFFETTUATE

GANGLIOGLIOMI
DNT
GANGLIOCITOMI

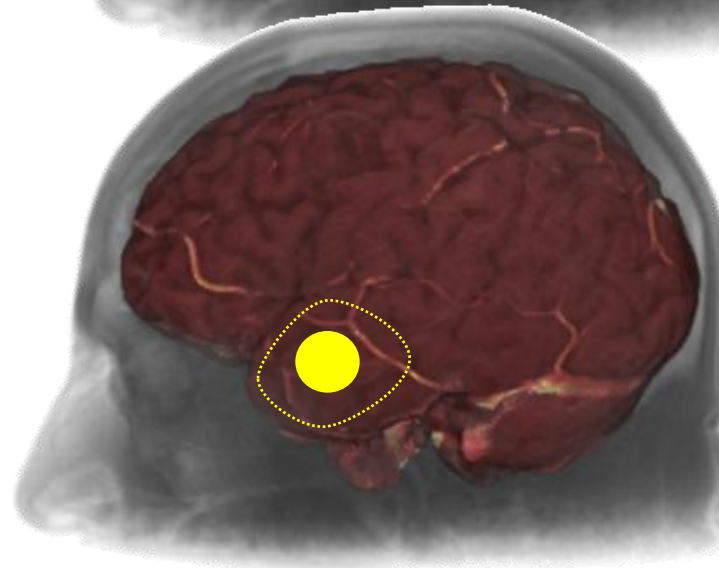
Pazienti epilettici

LESIONECTOMIA



60%

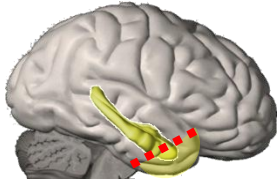
**LESIONECTOMIA
+ CORTECTOMIA**



40%

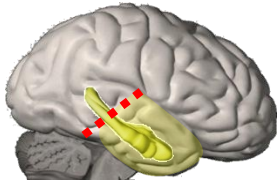
LOBO TEMPORALE

(TOT 58 pazienti)



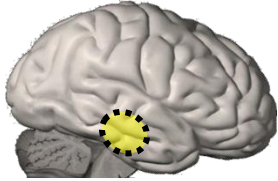
Lobectomie Temporo-Mesiali
Anteriori

18 (31%)



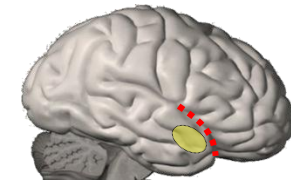
Lobectomie Temporali Estese

4 (7%)



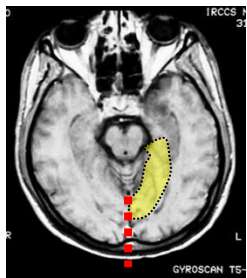
Lesionectomia
Lesioni laterali e basali

20 (35%)



Lesionectomia (**APPROCCIO TRANSILVIANO**)
Lesioni mesiali

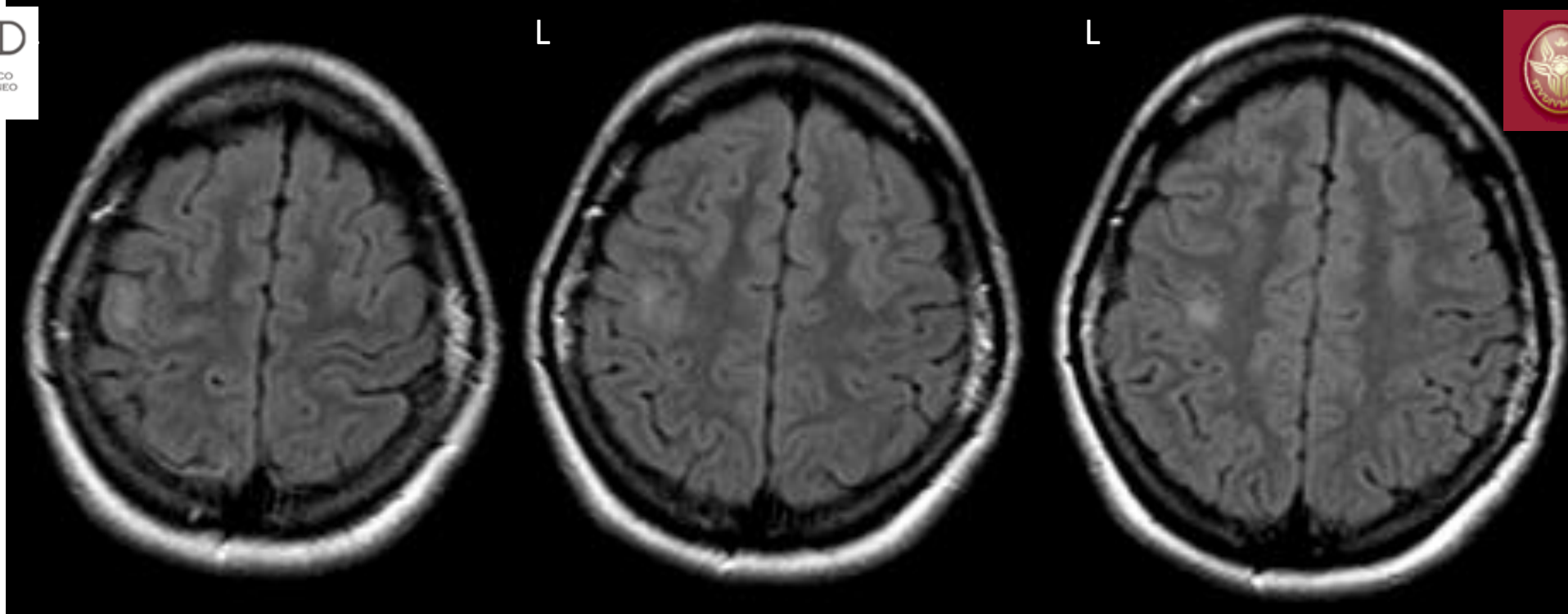
13 (22%)



Lesionectomia (**APPROCCIO INTEREMISFERICO**)

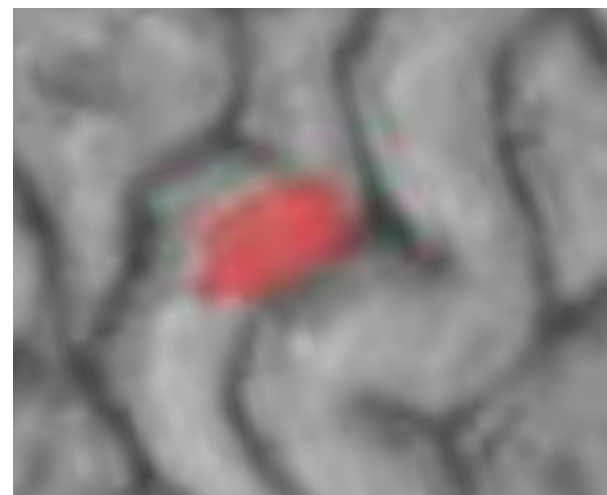
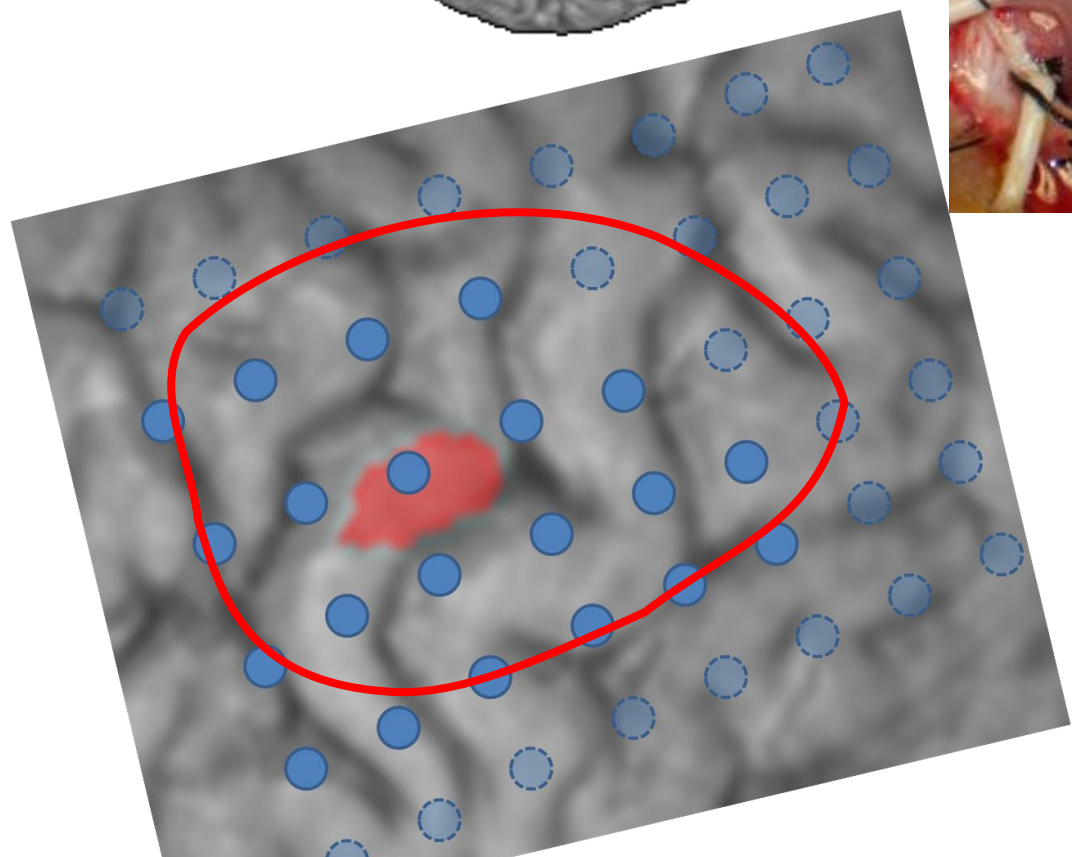
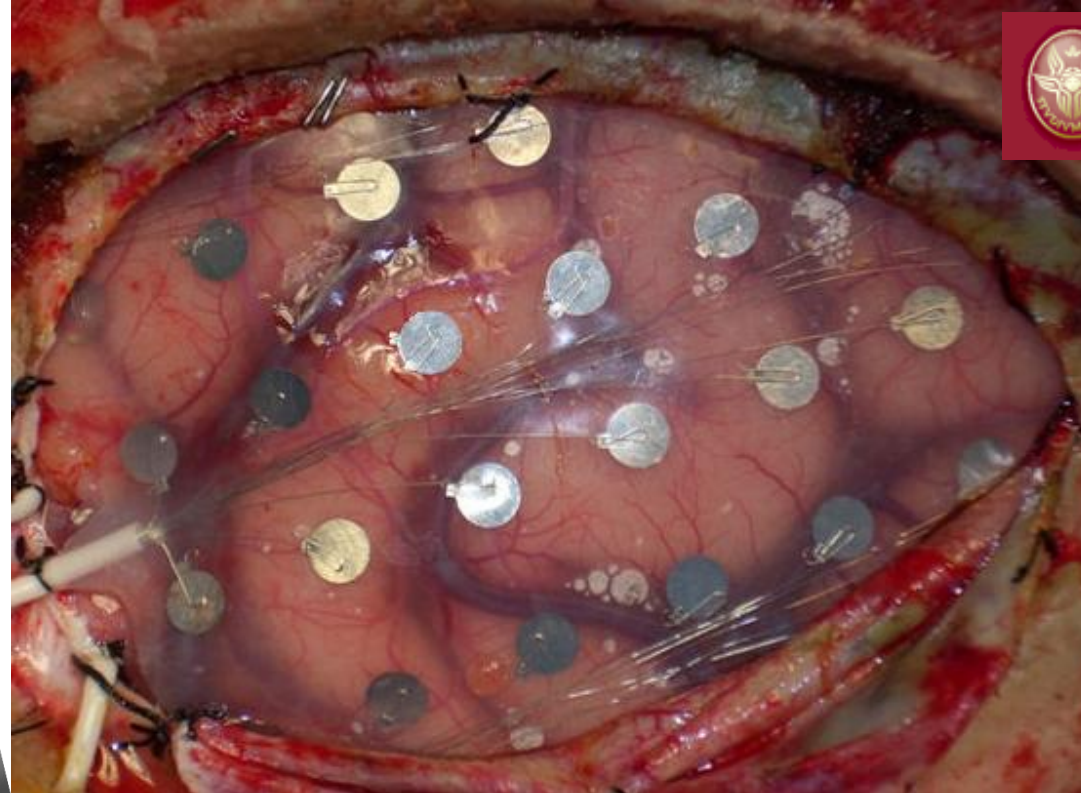
Lesioni mesiali temporo-occipitali

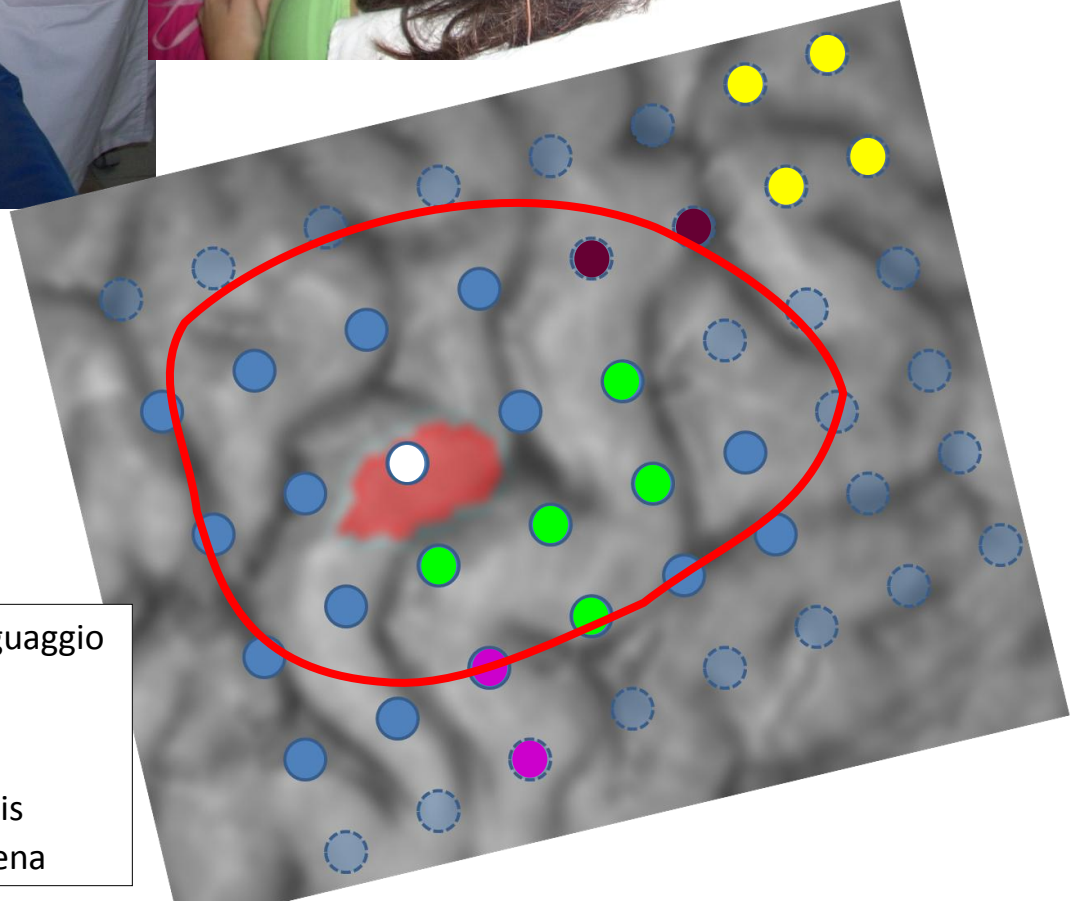
3 (5%)








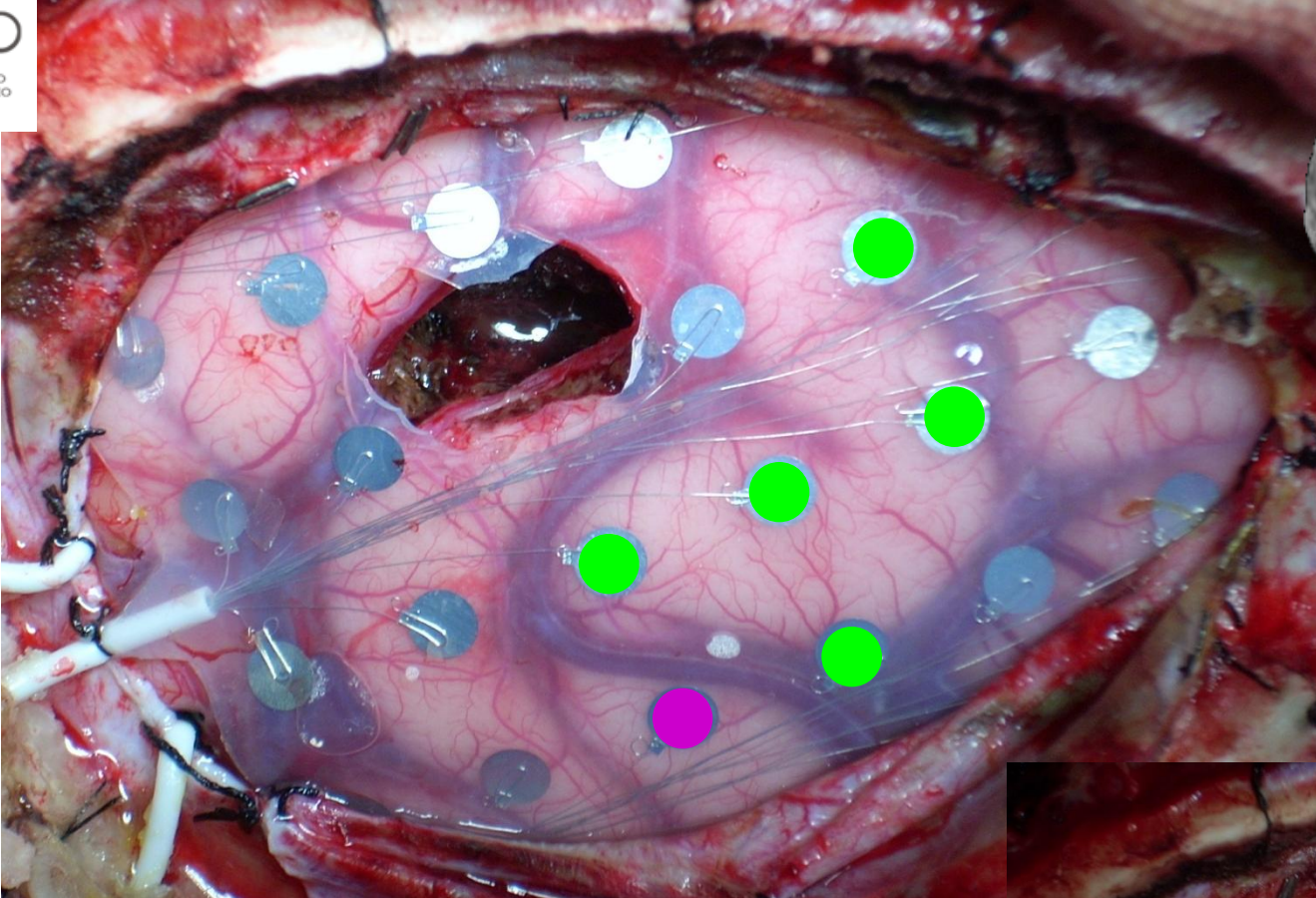
M.T. , 23 anni, ♀
Da 2 anni crisi motorie alla mano
destra
Fino a 100 crisi al giorno !!



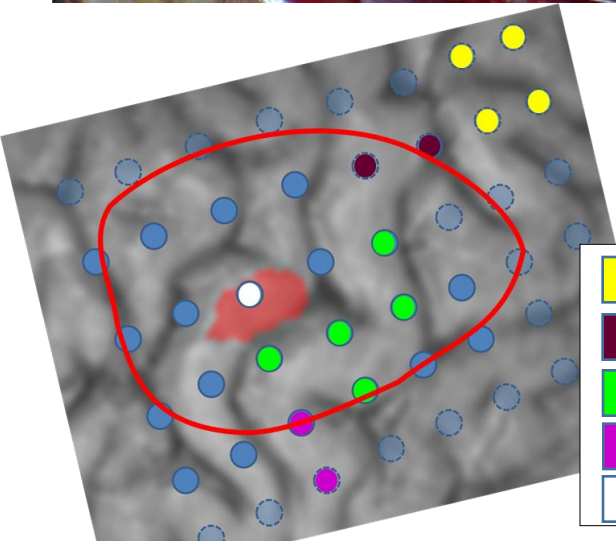
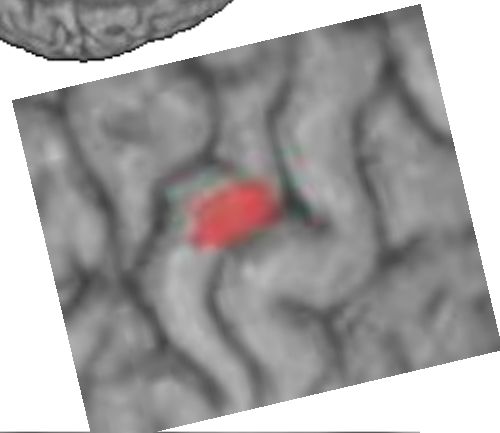




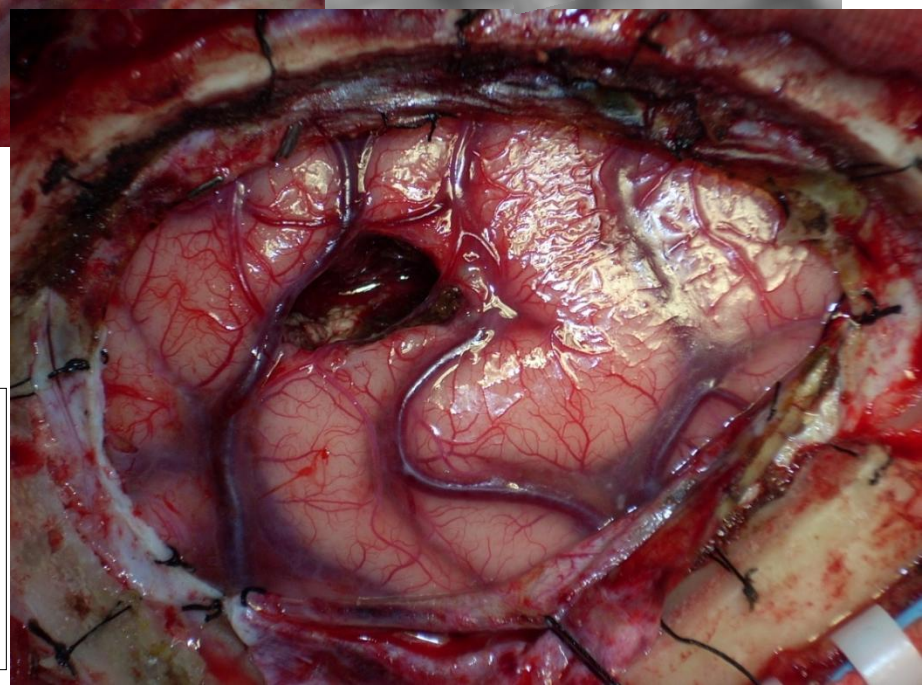
- | | |
|---|------------------------|
|  | Arresto del linguaggio |
|  | Movim. Faccia |
|  | Movim. Mano |
|  | Biceps brachialis |
|  | Zona Epilettogena |

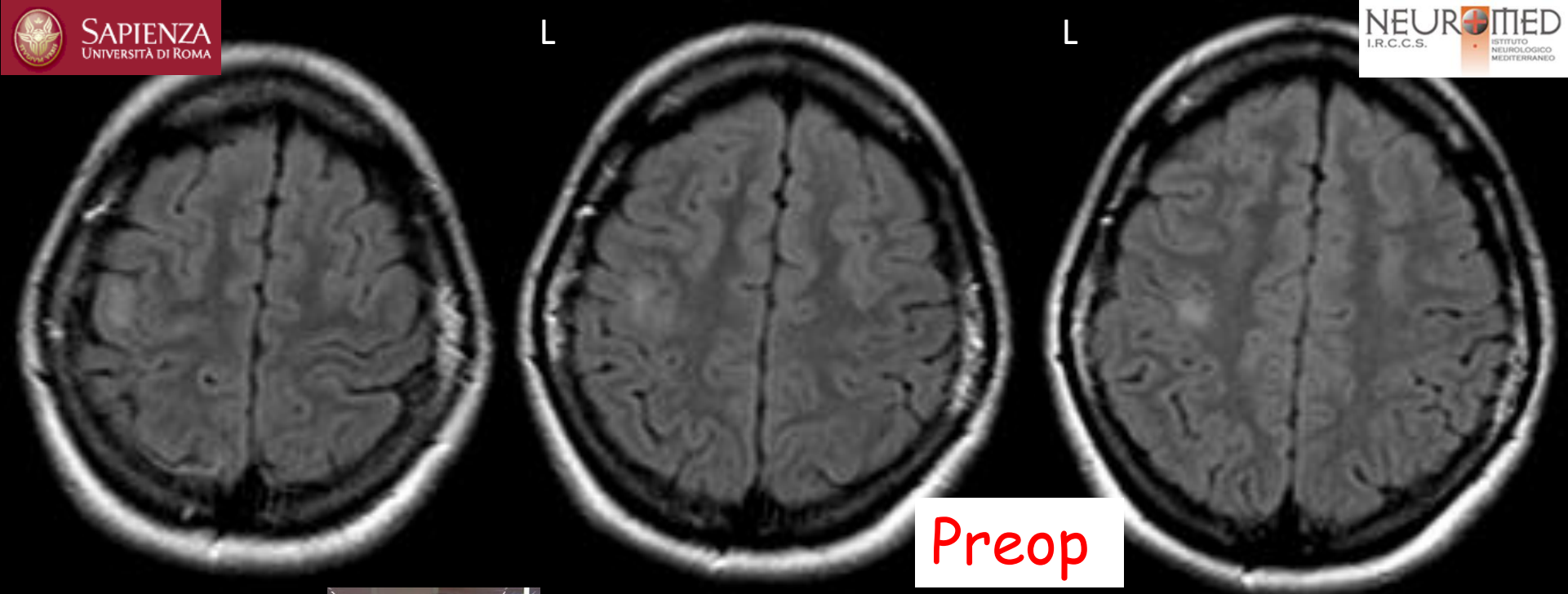


Operata
Novembre 2007

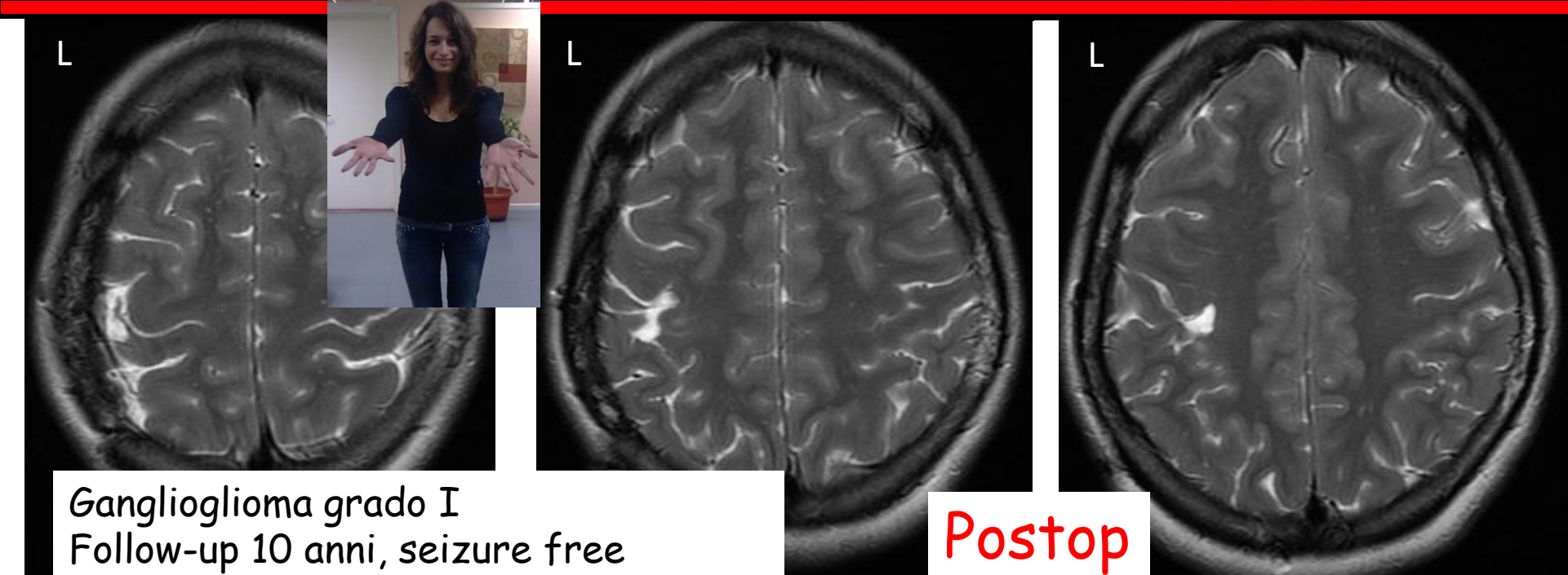


- Arresto del linguaggio
- Movim. Faccia
- Movim. Mano
- Biceps brachialis
- Zona Epilettogena





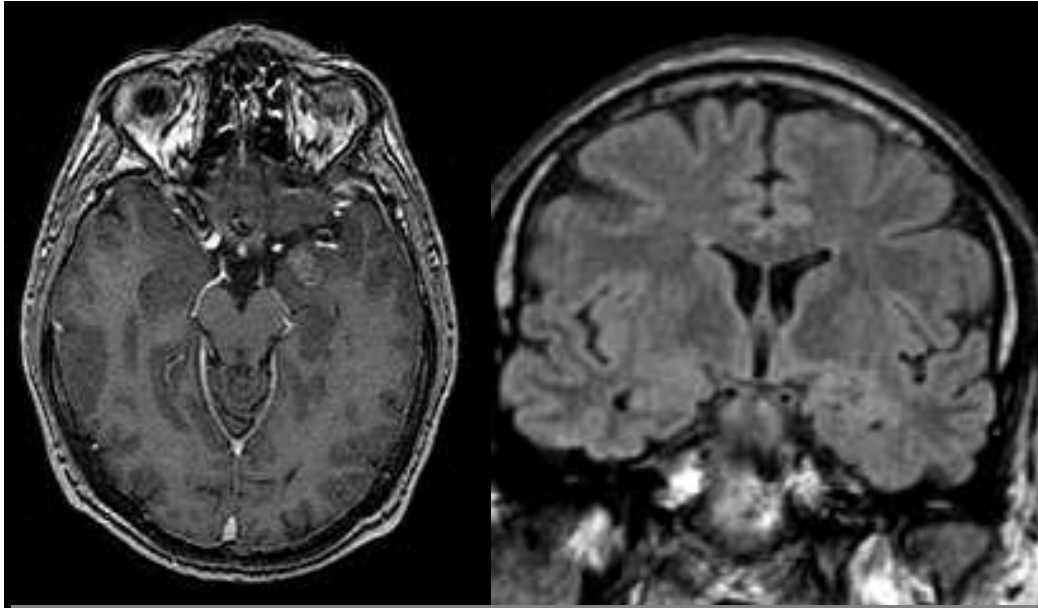
Preop



Postop

Ganglioglioma grado I
Follow-up 10 anni, seizure free





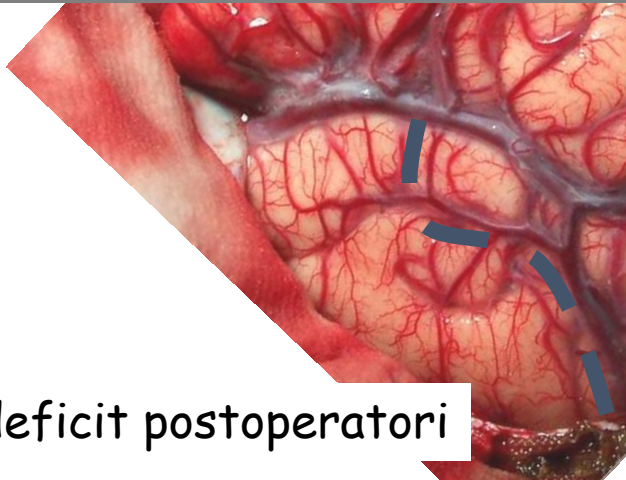
m 41 aa, destrimane

Epilessia farmaco-resistente
(durata 19 aa)

Crisi parziali complesse

Frequenza plurisettimanale

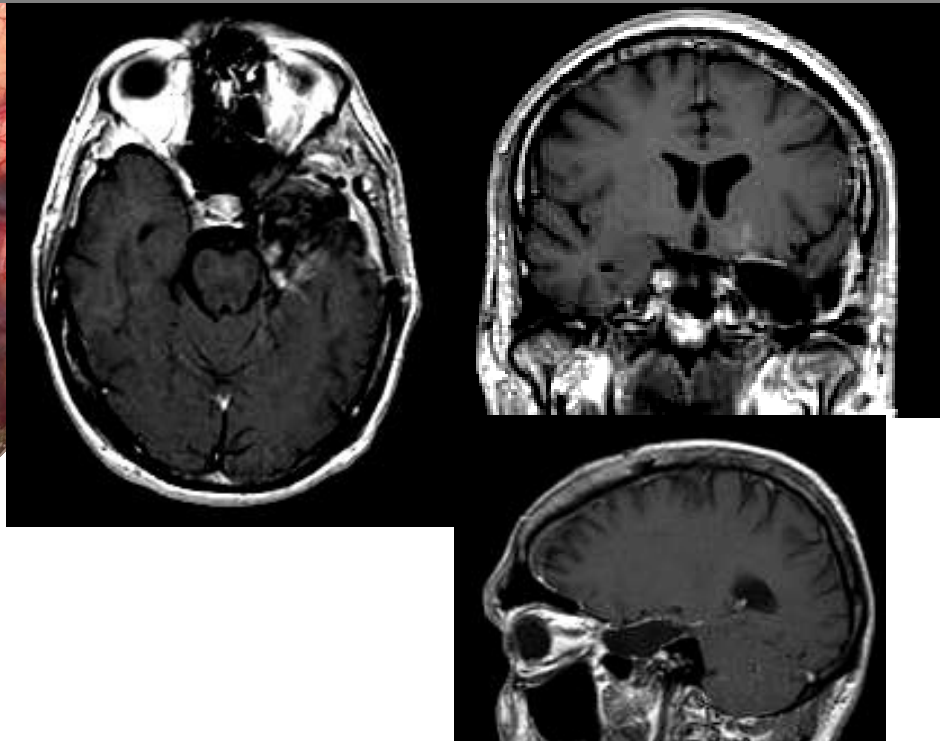
Video-EEG: pattern
elettroclinico latero-mesiale



No deficit postoperatori

Libero da crisi (Engel I)
follow-up 5 aa

Ganglioglioma (I WHO)

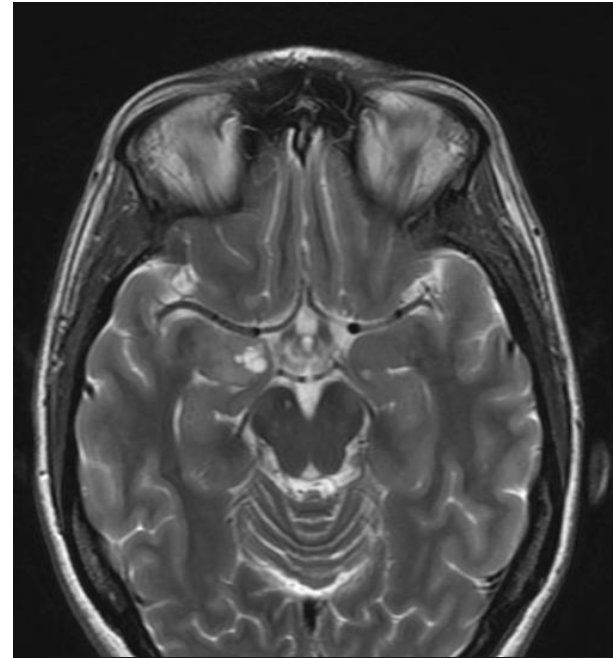
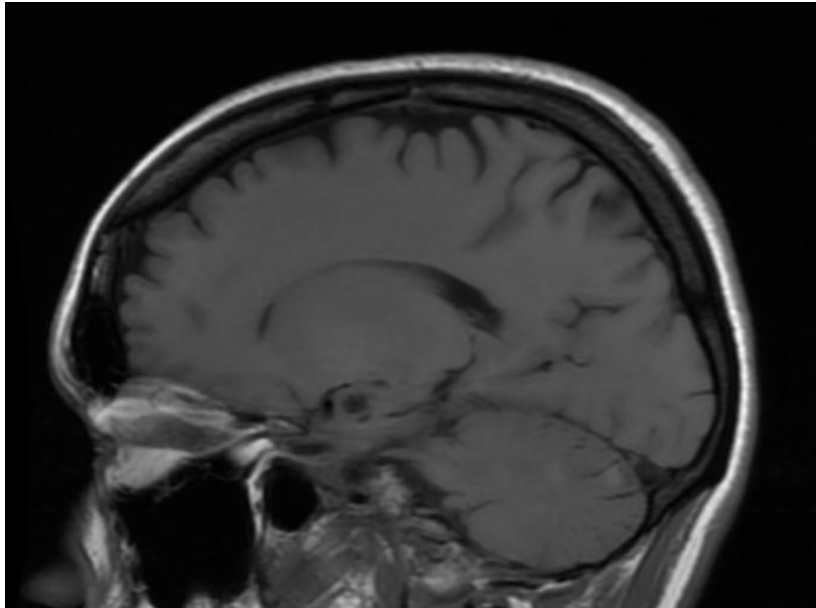


f 39 aa, mancina corretta
Epilessia farmacoresistente
(durata 15 aa)

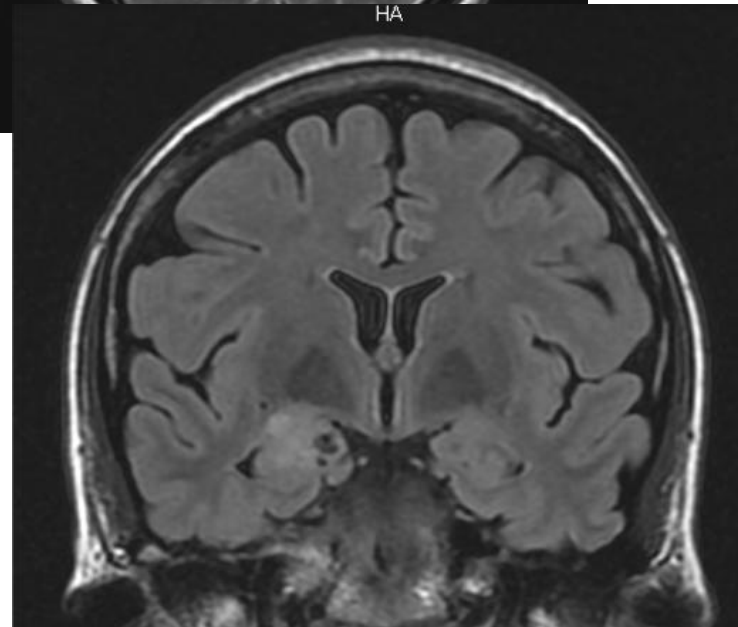
Crisi parziali semplici

Frequenza plurimensile

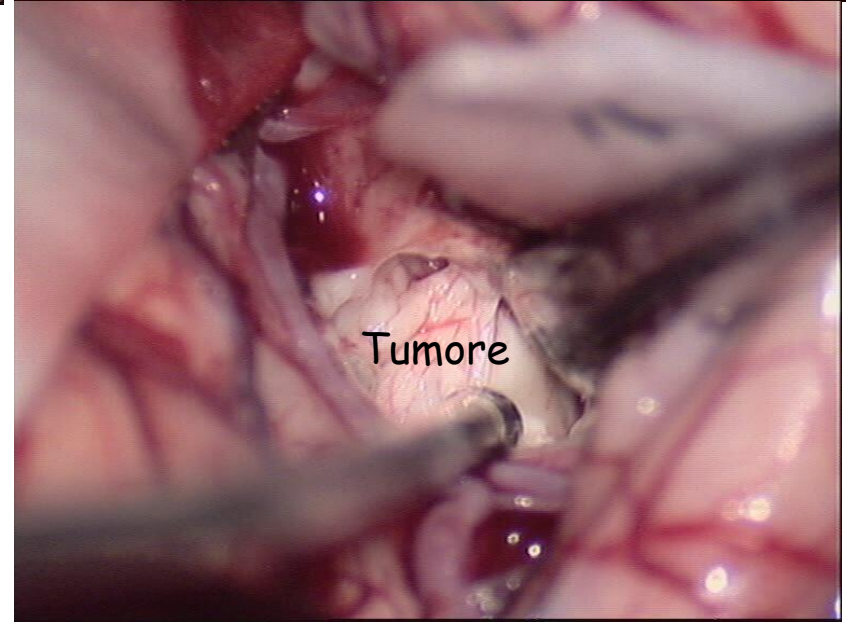
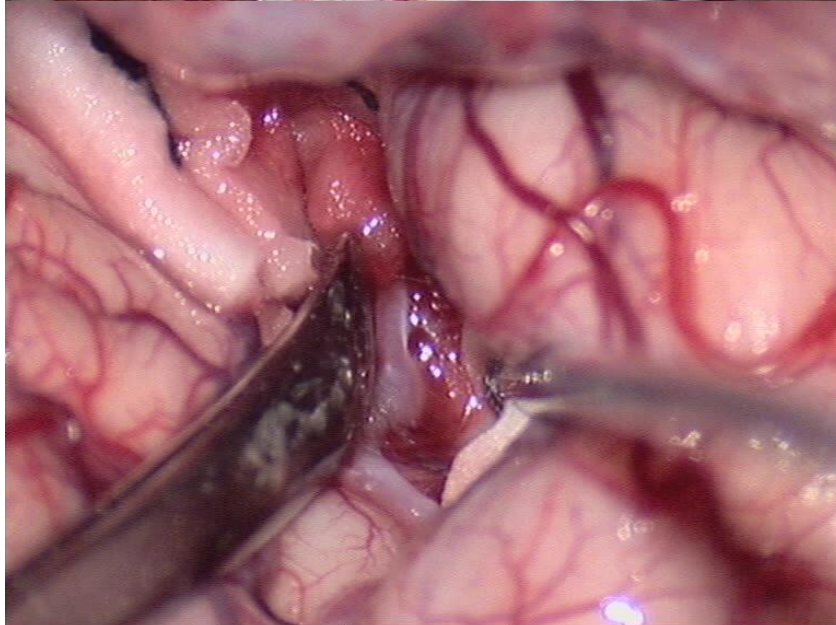
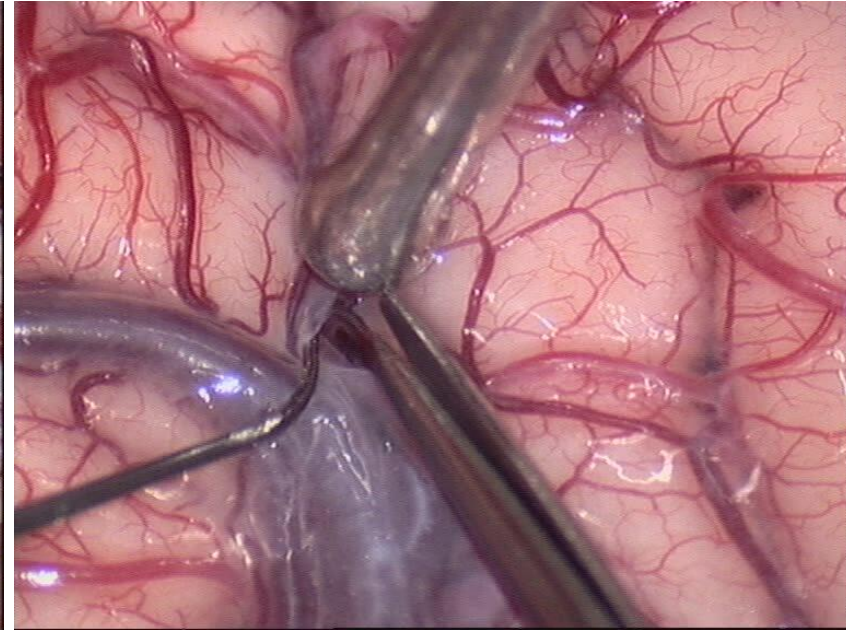
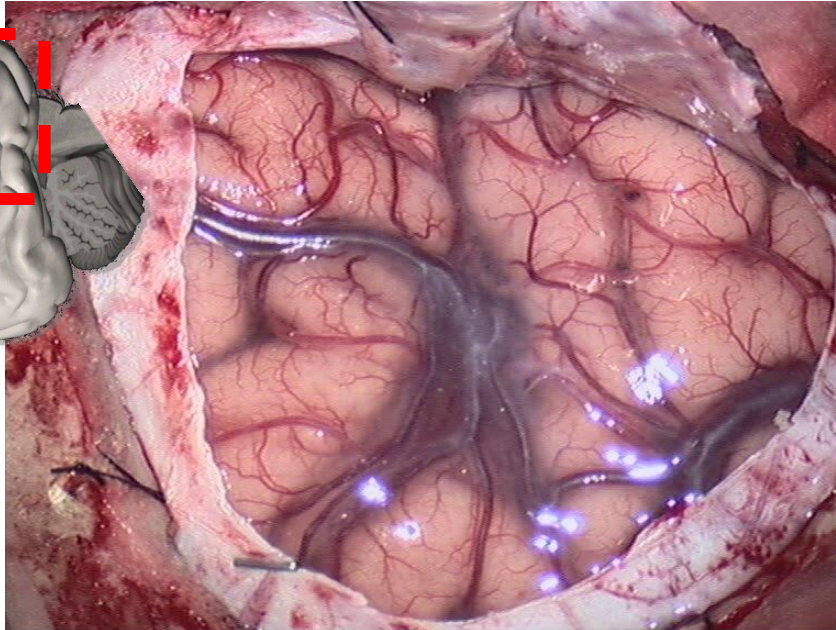
Video-EEG: pattern
elettroclinico mesiale



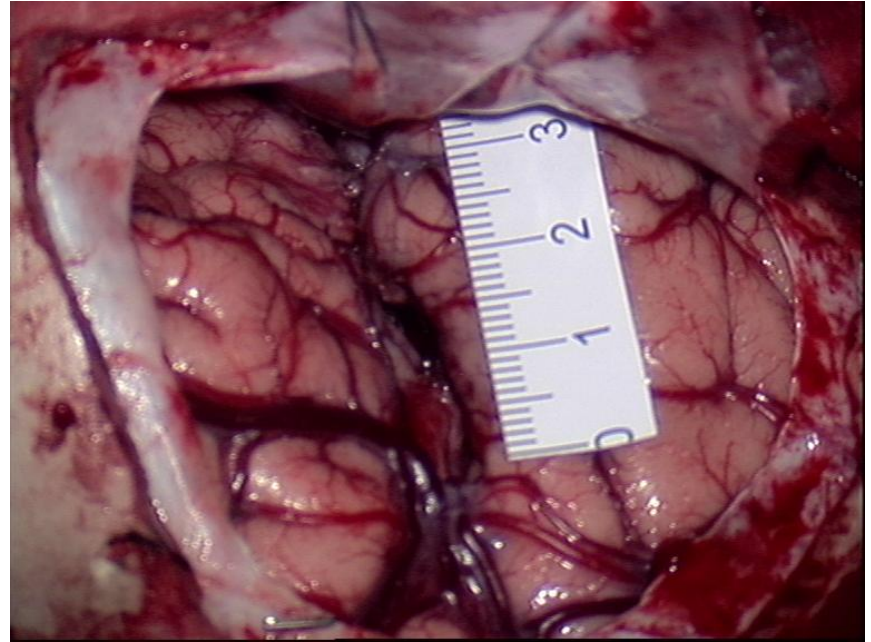
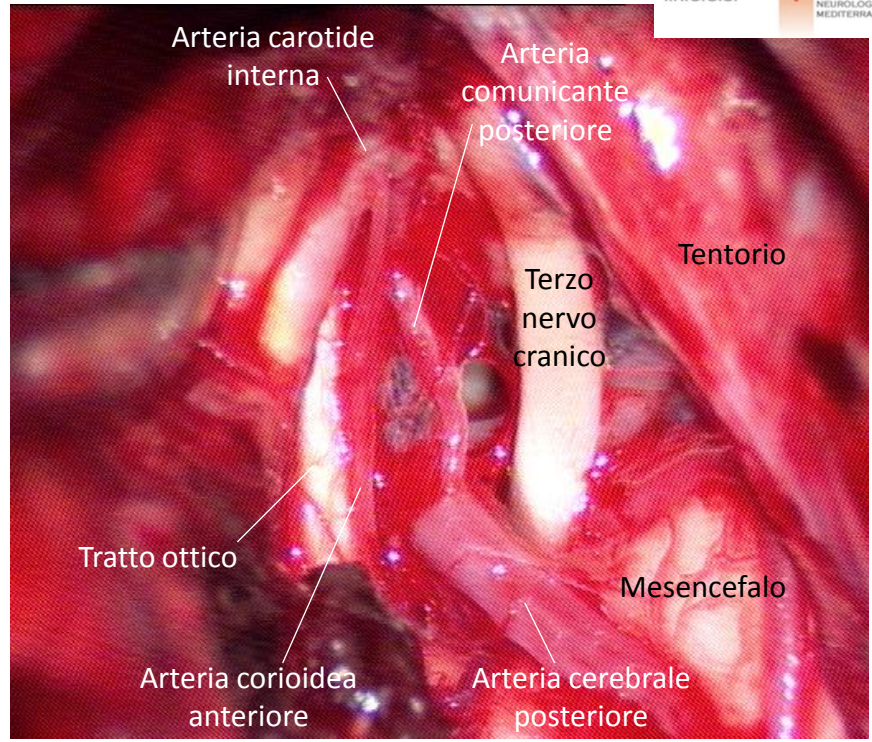
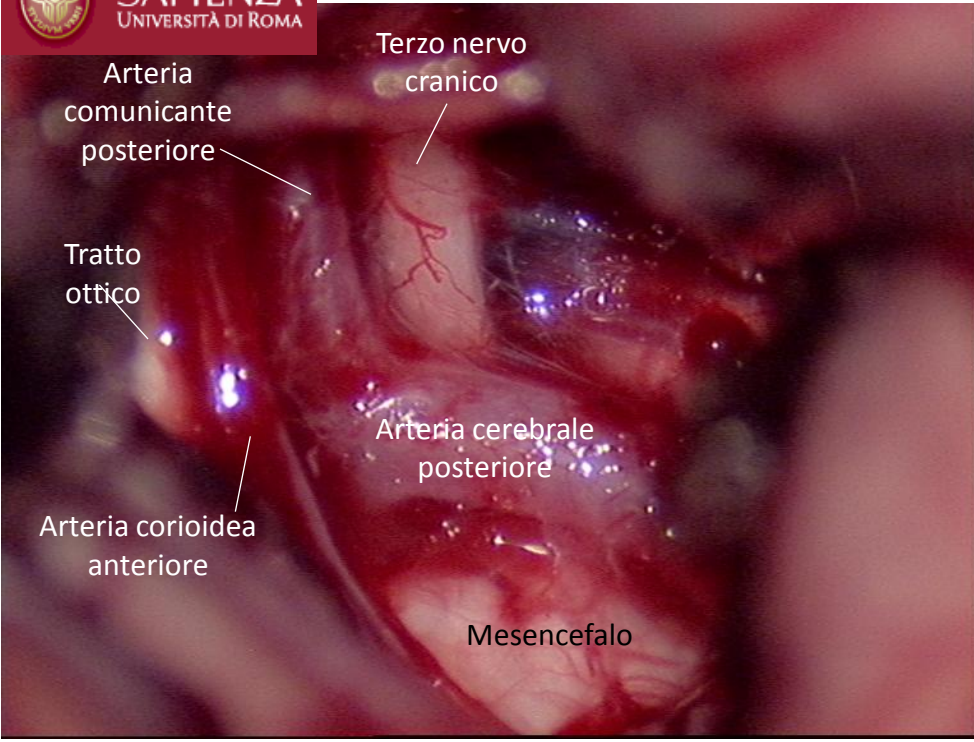
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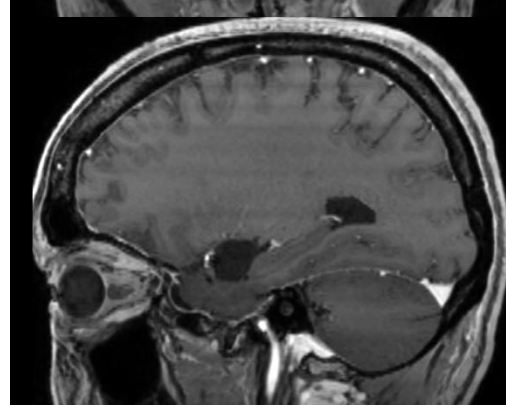
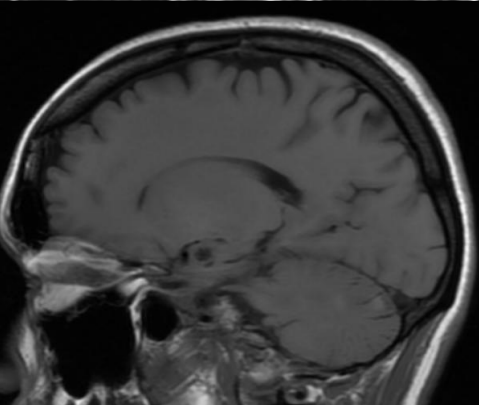
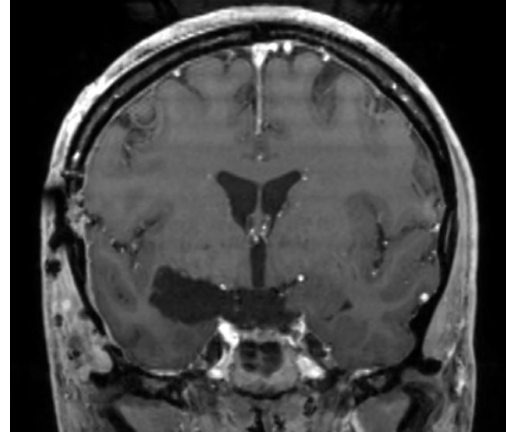
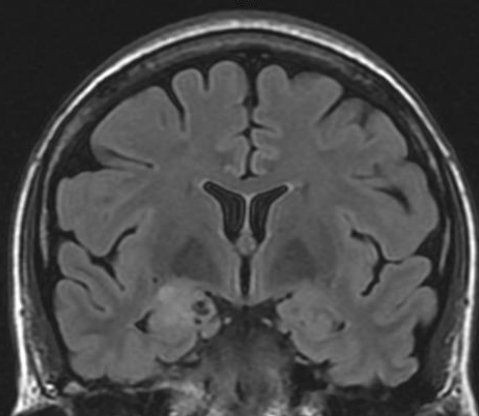
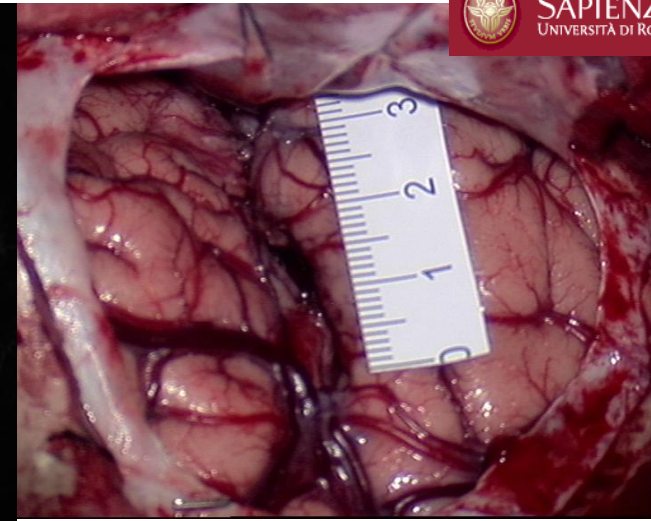
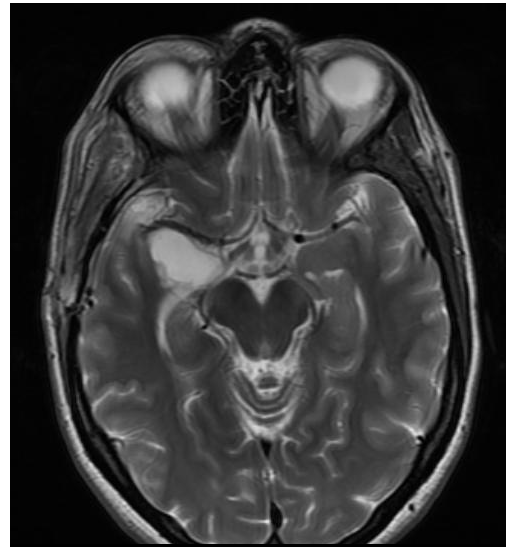
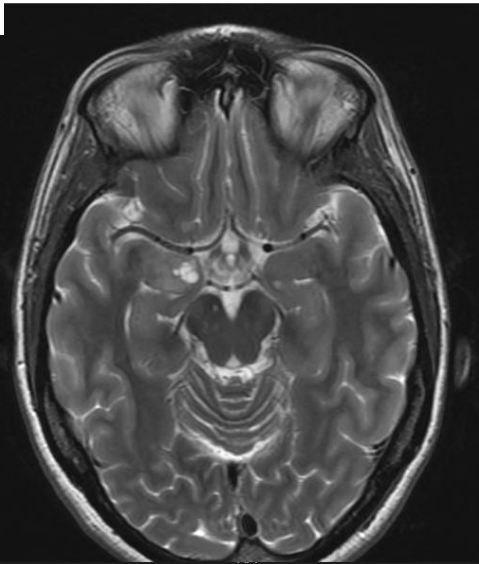


Via Trans-silviana



Tumore



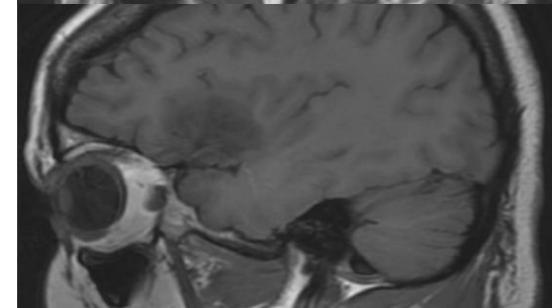
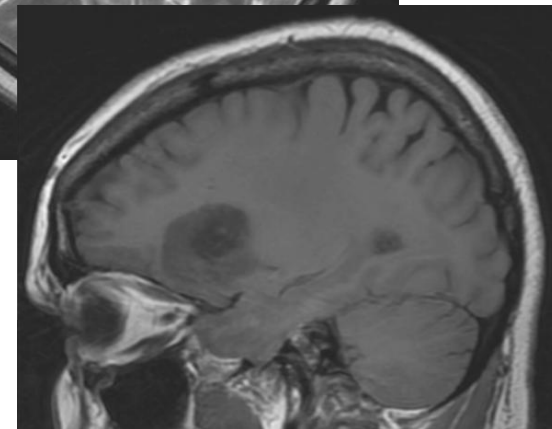
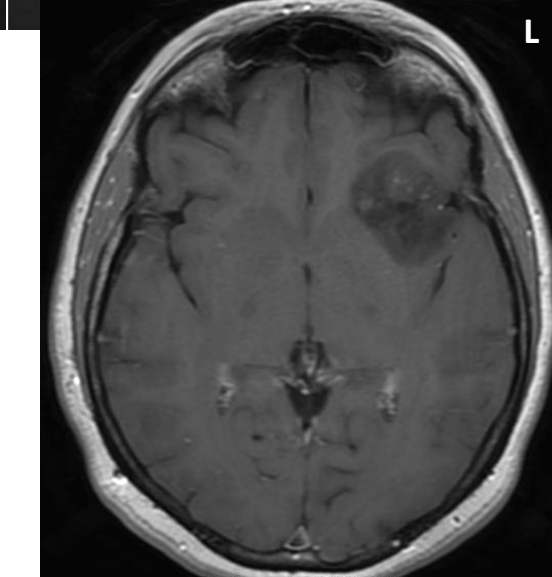
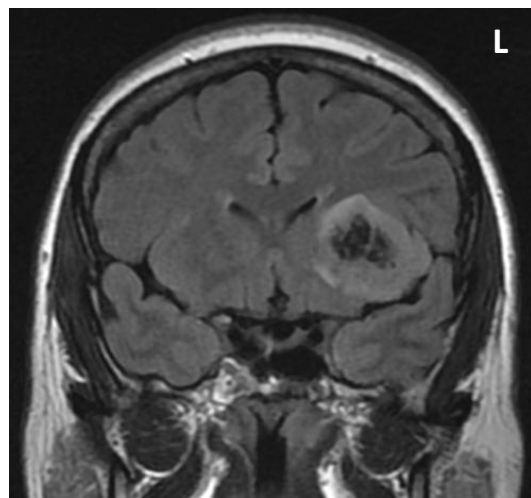
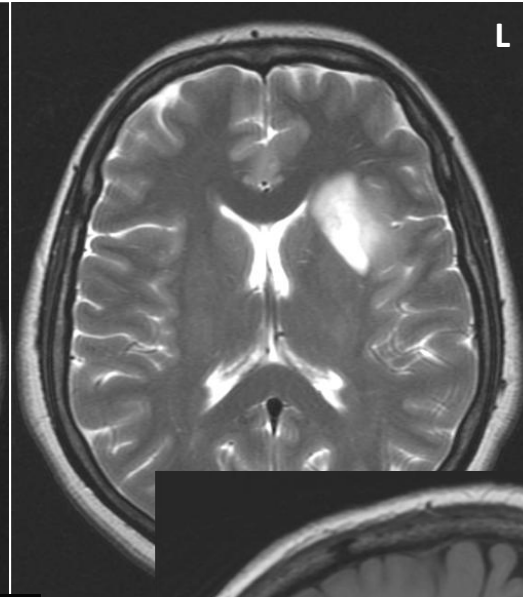
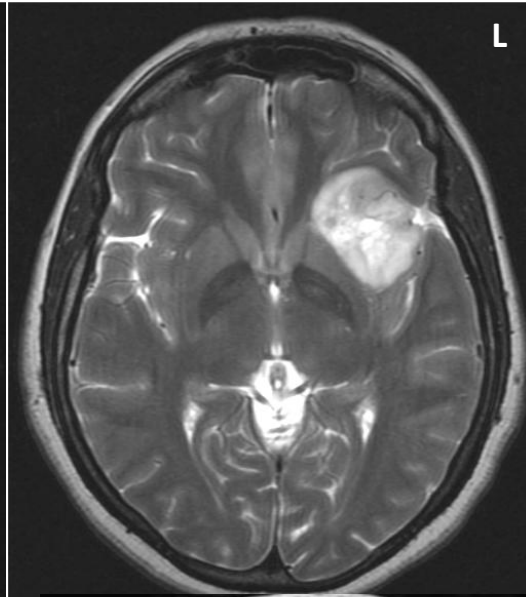
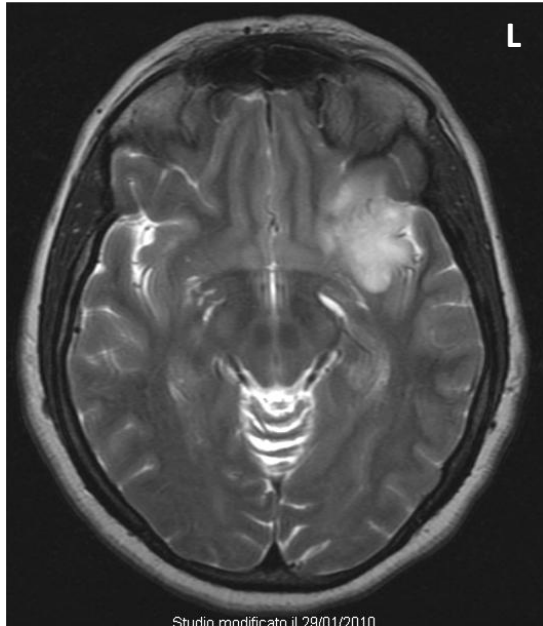


DNT (I WHO)

Libera da crisi (Engel I)
Follow-up 6 aa

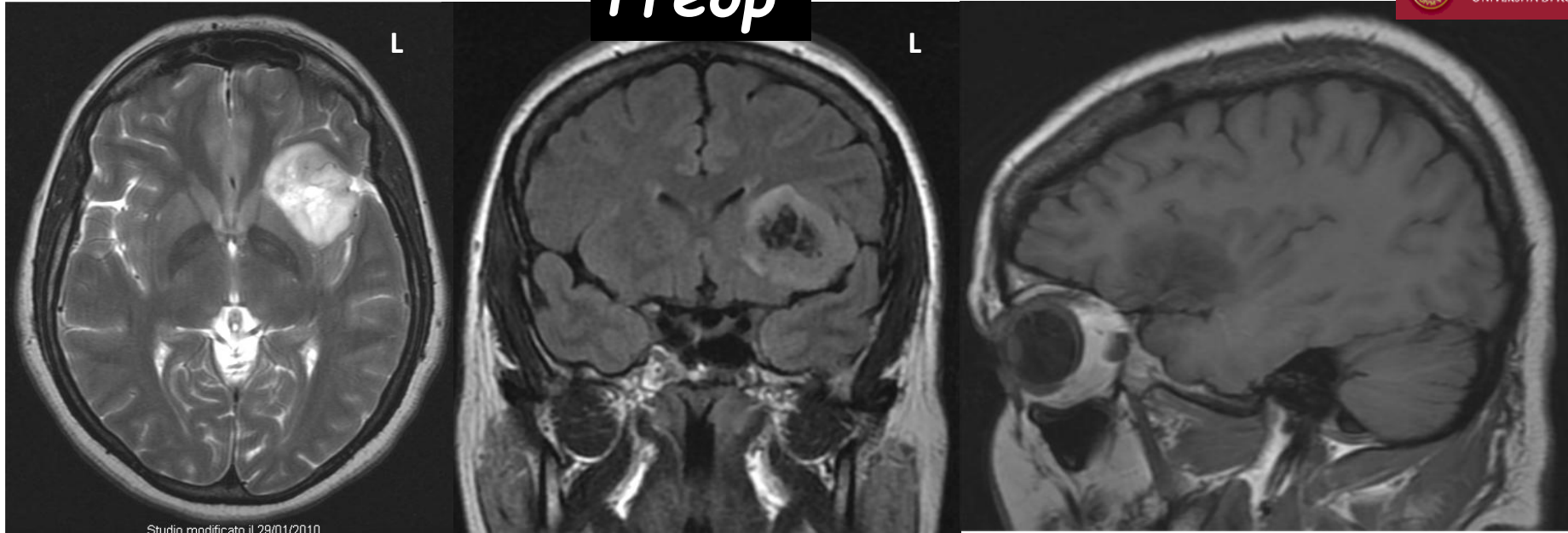
G.R., f, 38 a. Crisi epilettiche

Esame obiettivo neurologico normale

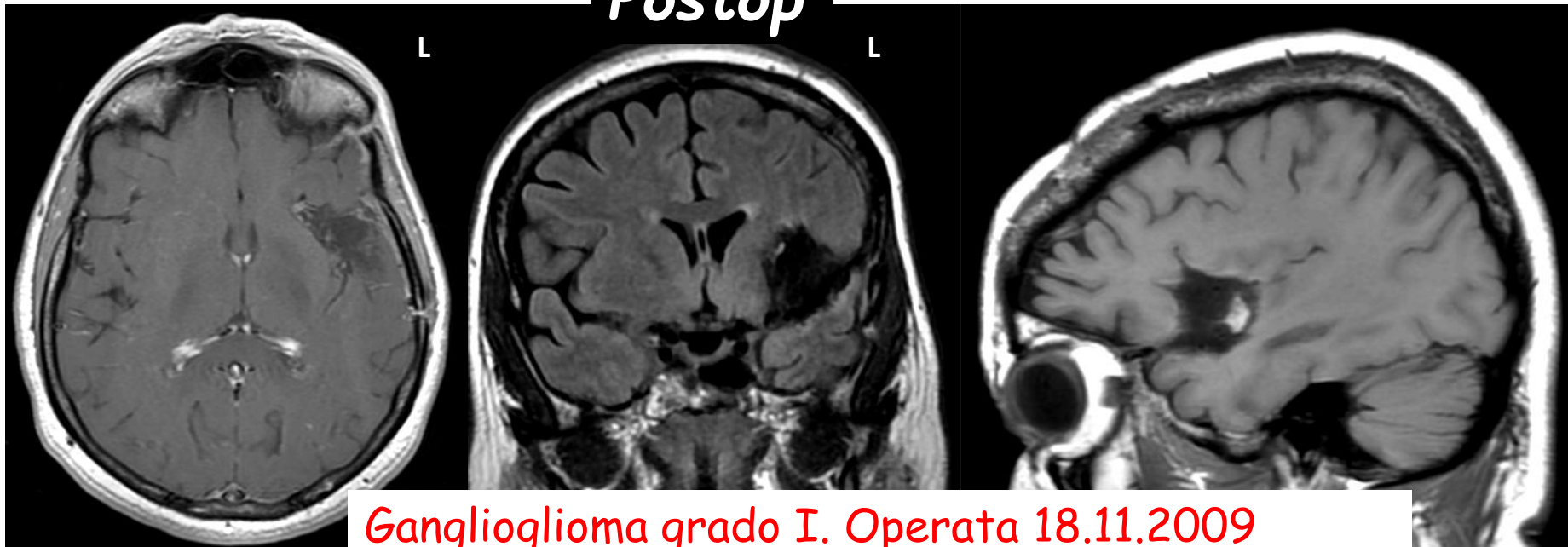


Novembre 2009

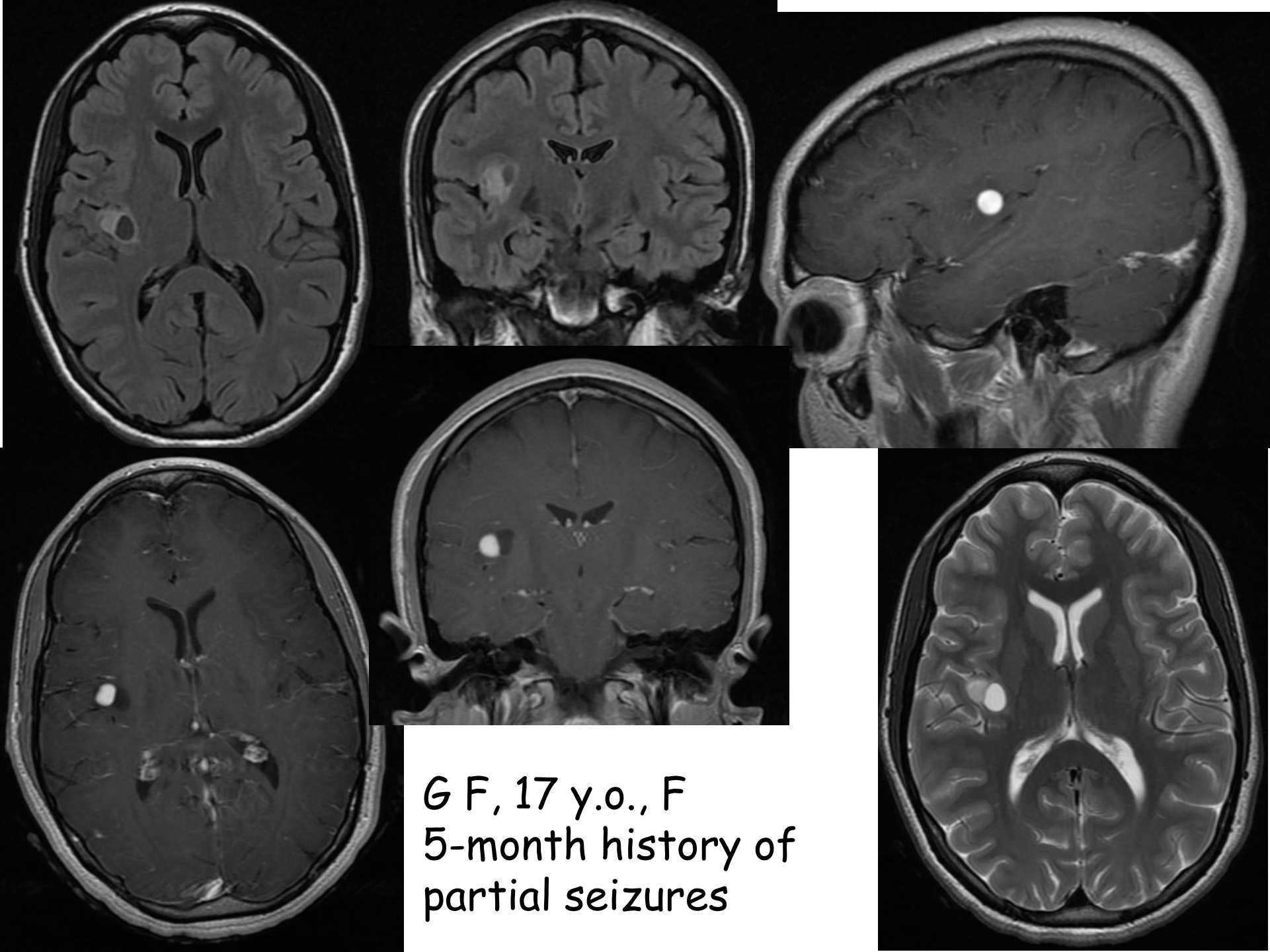
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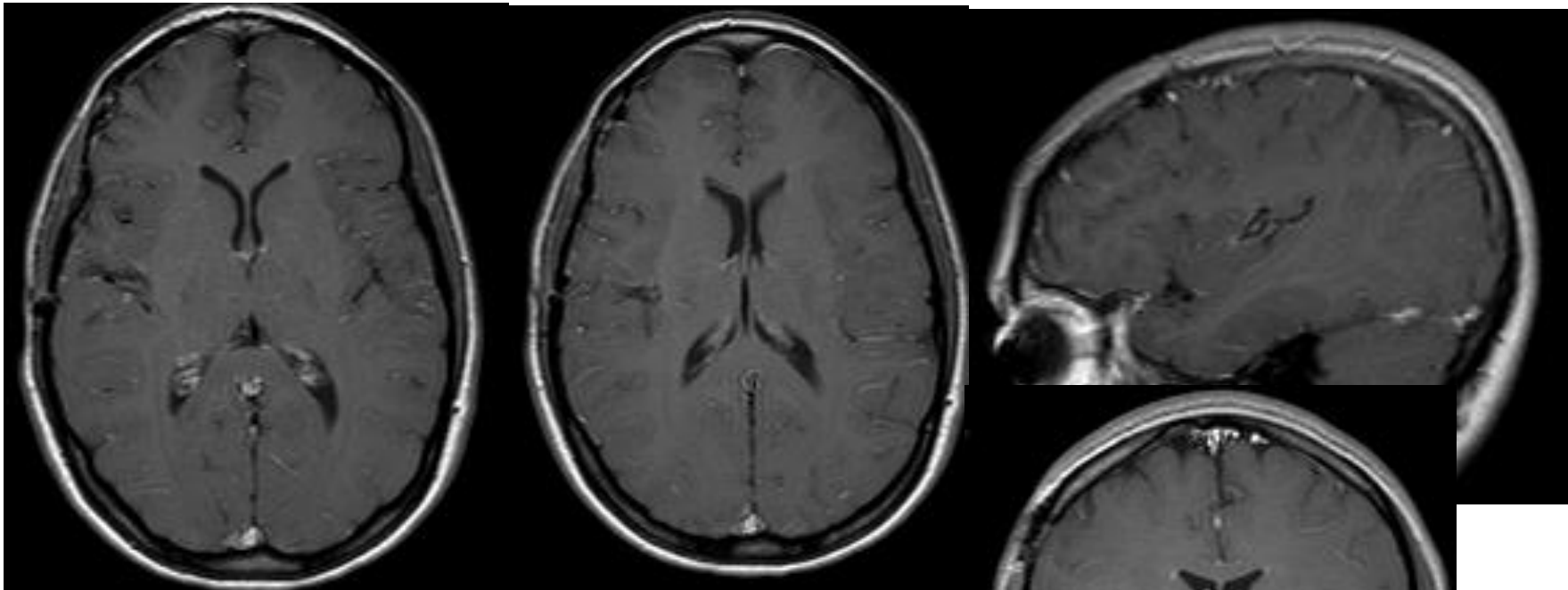
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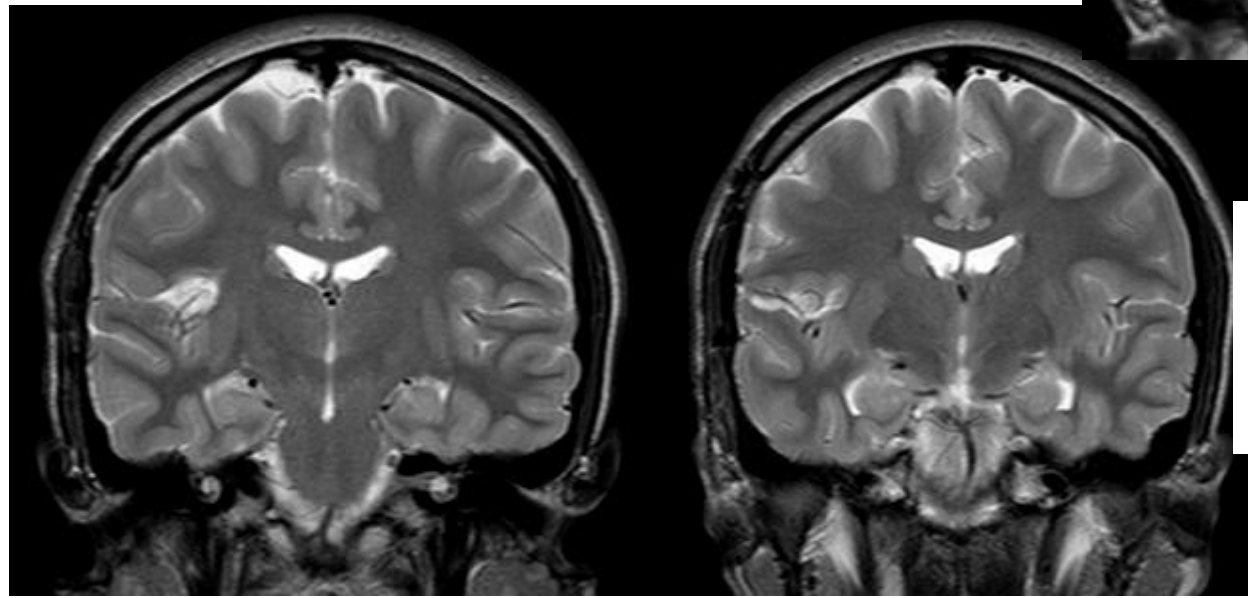
Ganglioglioma grado I. Operata 18.11.2009



G F, 17 y.o., F
5-month history of
partial seizures



Ganglioglioma, WHO grade I
4 years F.U., no seizures

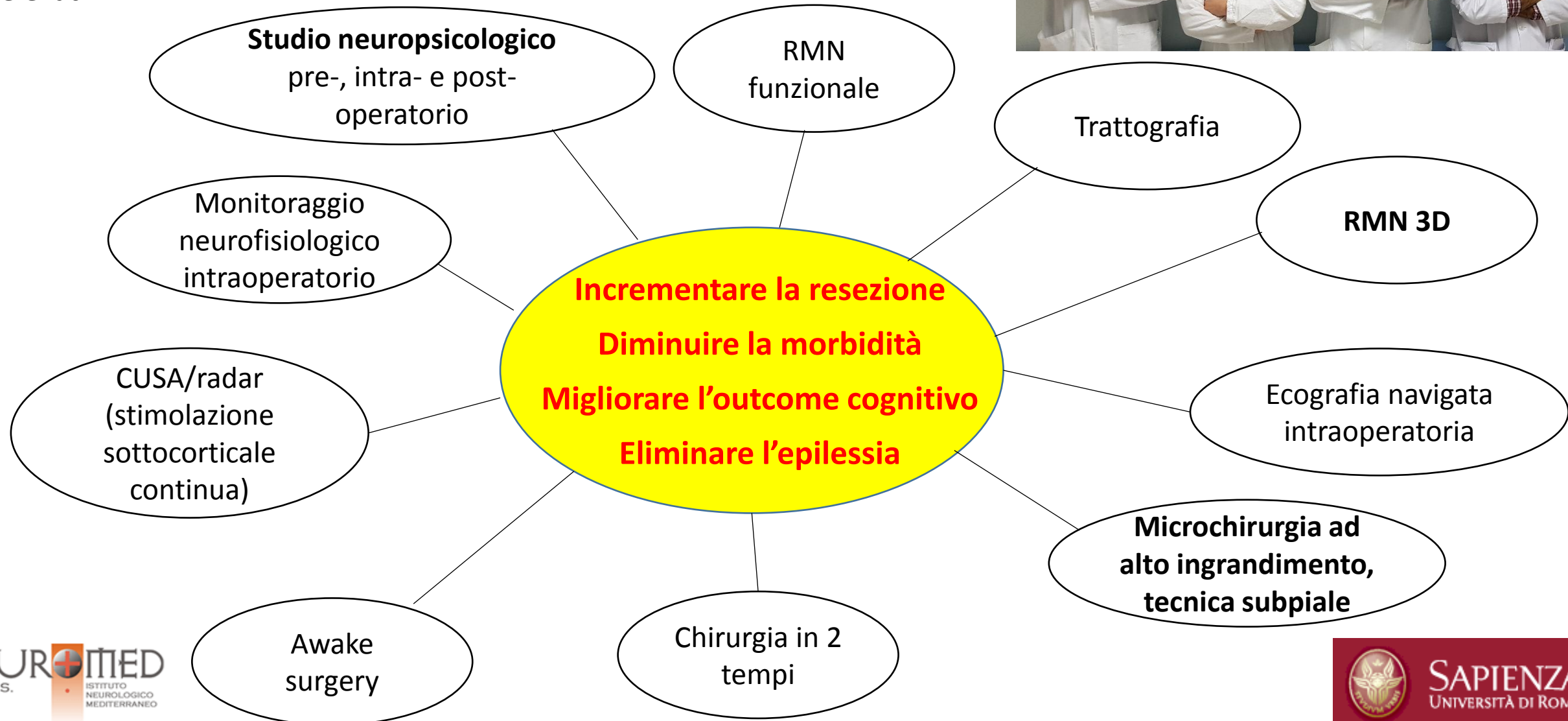
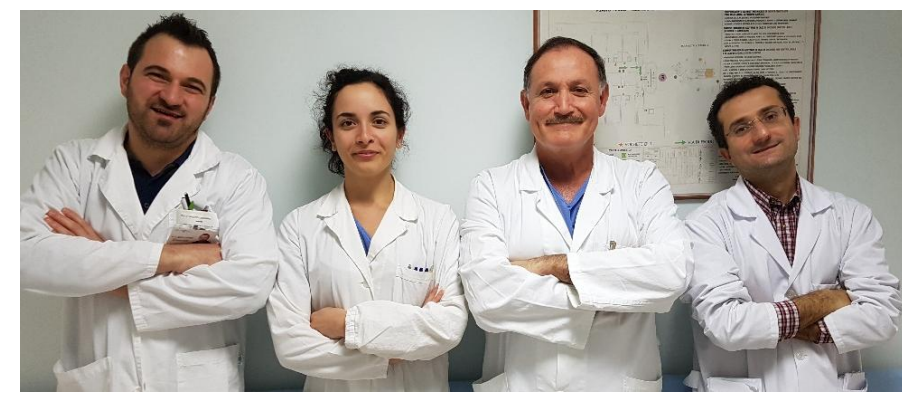


After
transylvian
microsurgery



Conclusioni - 1

I gliomi, specie se di basso grado, vanno operati precocemente, per migliorare la qualità della vita (crisi epilettiche) e ritardare l'inevitabile progressione a gradi più elevati



CONCLUSIONI- 2



Nei tumori glioneuronalni, i fattori critici sul risultato epilettologico e cognitivo sono:

- intervento precoce
- resezione totale

I pazienti con epilessia farmacoresistente necessitano di una valutazione prechirurgica epilettologica, in particolare nell'epilessia temporale (nel 40% dei casi è stata eseguita una resezione estesa al di là della asportazione della lesione, tutti con storia di epilessia superiore ad un anno)

Outcome epilettologico buono (85% di Engel I). Miglioramento cognitivo molto frequente, legato alla guarigione dell'epilessia e alla sospensione dei farmaci antiepilettici