

# SSMG. Bruxelles, février 2014.

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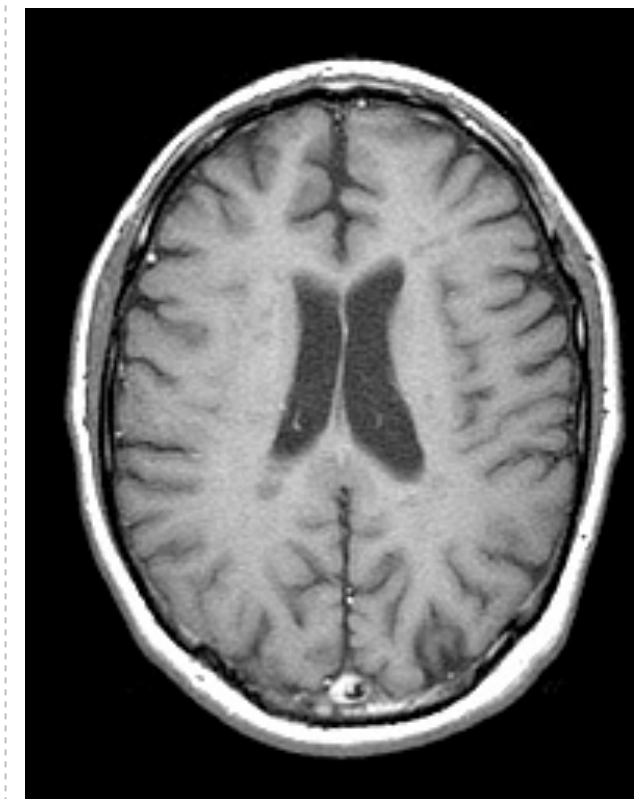
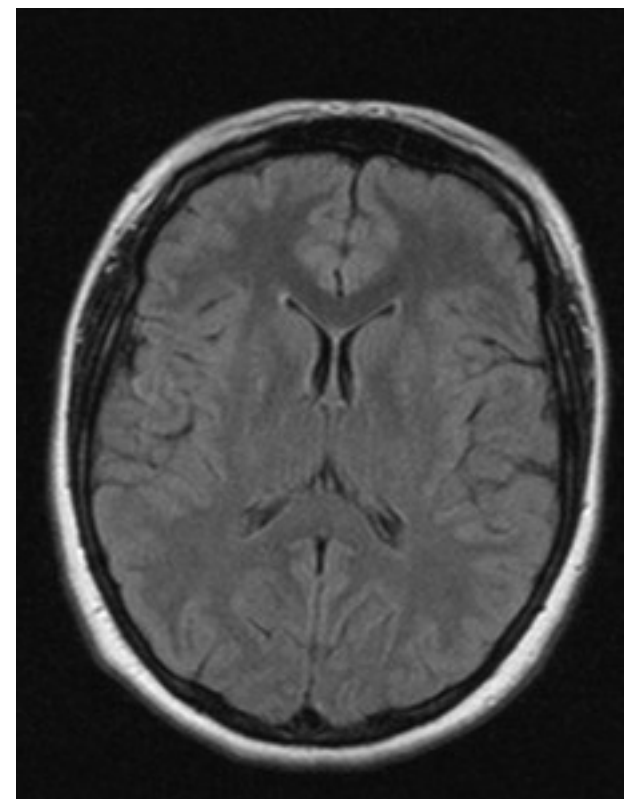
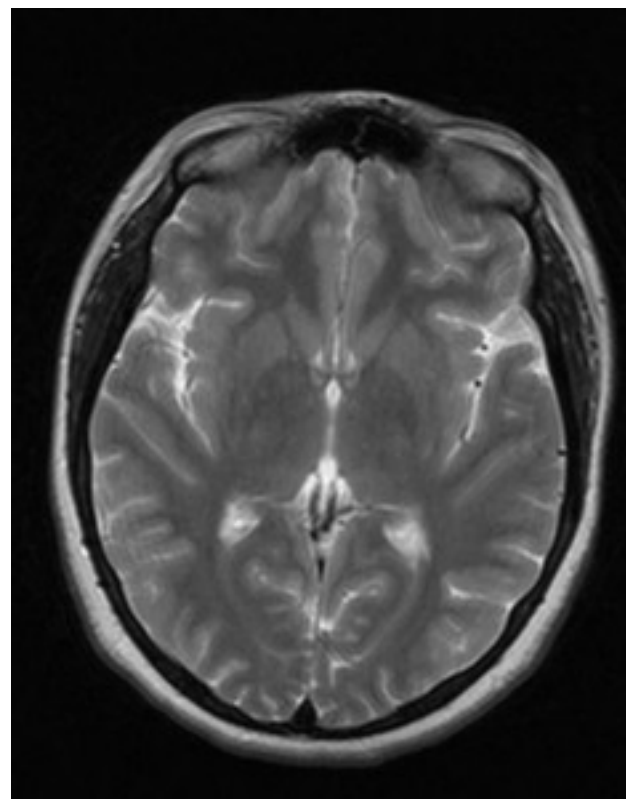
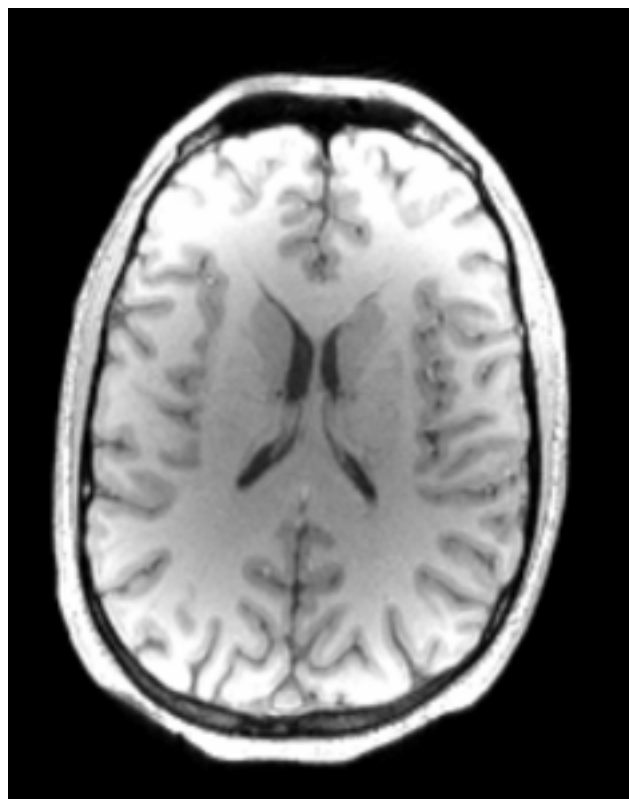


# Sommaire

- Introduction
- SEP
- Traumatismes cérébraux
- Conclusion



# I. Introduction



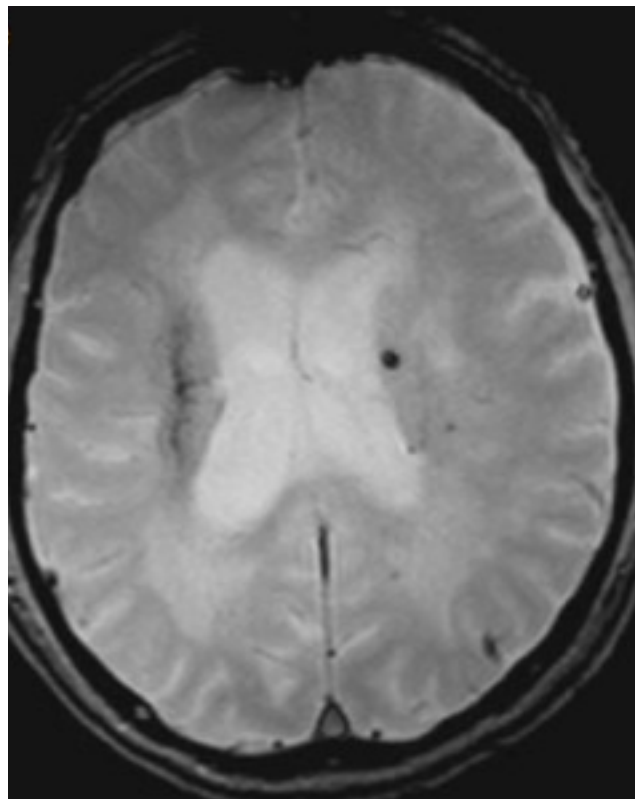
T1

T2

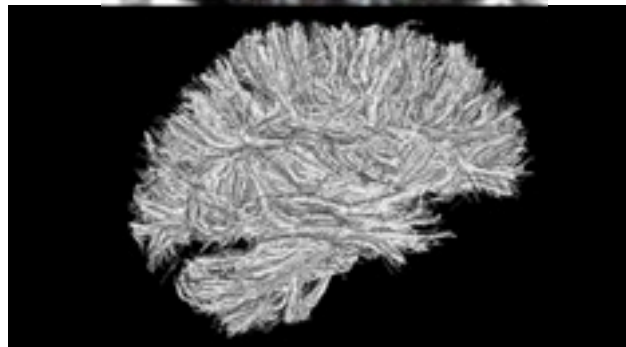
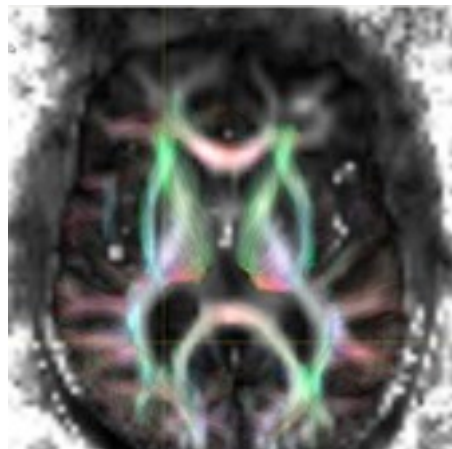
FLAIR

T1+G

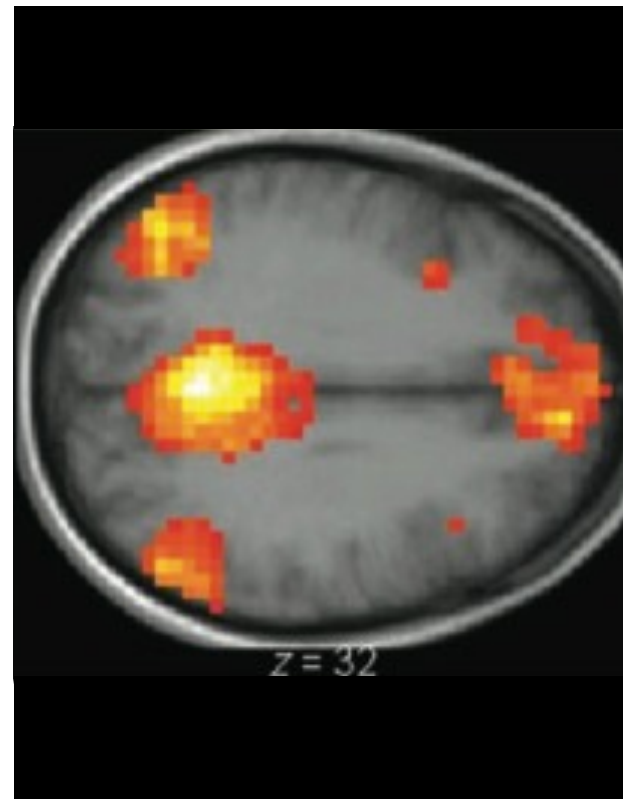
# I. Introduction



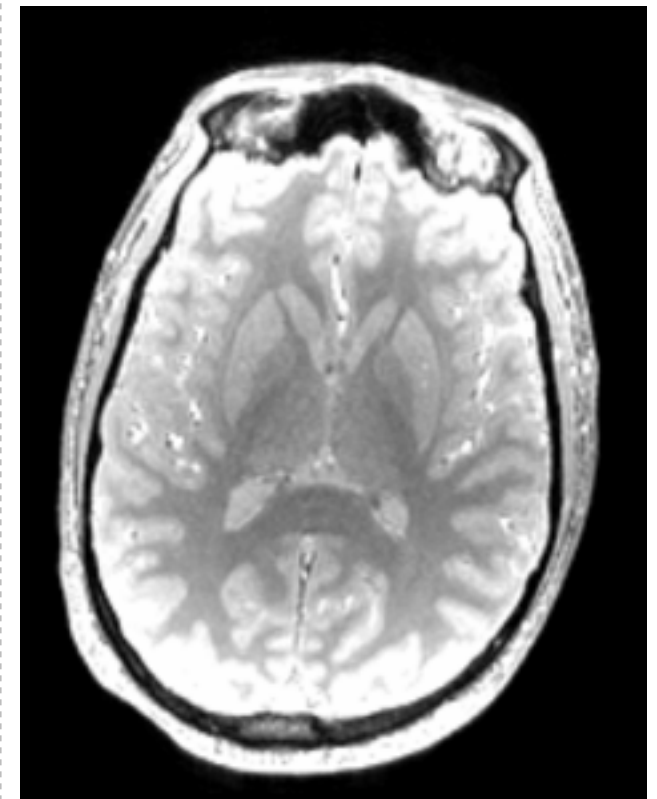
T2\*



Diff



fMRI



MTR

# III. SEP

- ❏ L'IRM cérébrale avec des séquences classique permet de faire le diagnostic sur base d'un seul examen (+ clinique et anamnèse bien sûr).
- ❏ NB: CT-Scan = inutile, ne permet pas de visualiser les lésions.

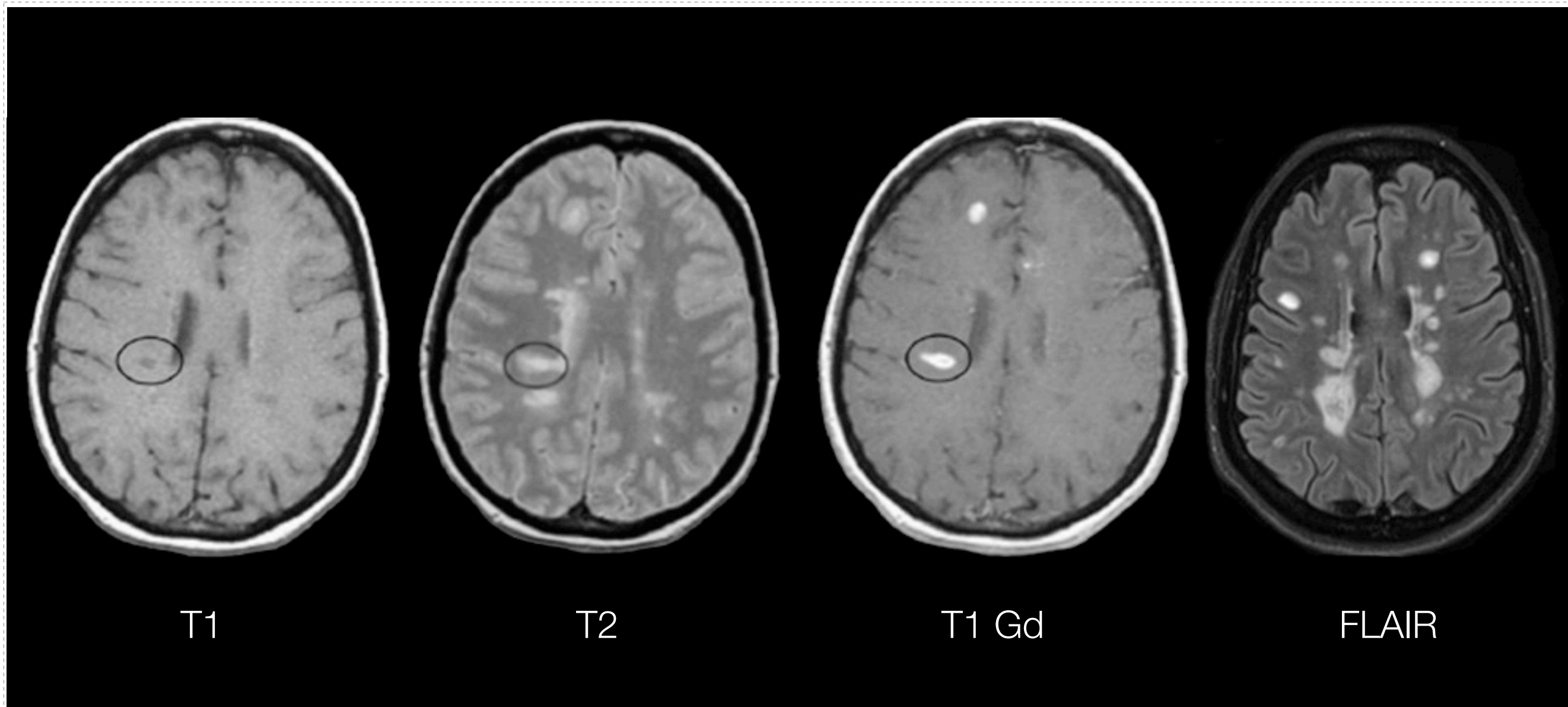
**TABLE 4: The 2010 McDonald Criteria for Diagnosis of MS**

Clinical Presentation	Additional Data Needed for MS Diagnosis
≥2 attacks <sup>a</sup> ; objective clinical evidence of ≥2 lesions or objective clinical evidence of 1 lesion with reasonable historical evidence of a prior attack <sup>b</sup>	None <sup>c</sup>
≥2 attacks <sup>a</sup> ; objective clinical evidence of 1 lesion	Dissemination in space, demonstrated by: ≥1 T2 lesion in at least 2 of 4 MS-typical regions of the CNS (periventricular, juxtacortical, infratentorial, or spinal cord) <sup>d</sup> ; or Await a further clinical attack <sup>a</sup> implicating a different CNS site
1 attack <sup>a</sup> ; objective clinical evidence of ≥2 lesions	Dissemination in time, demonstrated by: Simultaneous presence of asymptomatic gadolinium-enhancing and nonenhancing lesions at any time; or A new T2 and/or gadolinium-enhancing lesion(s) on follow-up MRI, irrespective of its timing with reference to a baseline scan; or Await a second clinical attack <sup>a</sup>
1 attack <sup>a</sup> ; objective clinical evidence of 1 lesion (clinically isolated syndrome)	Dissemination in space and time, demonstrated by: For DIS: ≥1 T2 lesion in at least 2 of 4 MS-typical regions of the CNS (periventricular, juxtacortical, infratentorial, or spinal cord) <sup>d</sup> ; or Await a second clinical attack <sup>a</sup> implicating a different CNS site; and For DIT: Simultaneous presence of asymptomatic gadolinium-enhancing and nonenhancing lesions at any time; or A new T2 and/or gadolinium-enhancing lesion(s) on follow-up MRI, irrespective of its timing with reference to a baseline scan; or Await a second clinical attack <sup>a</sup>
Insidious neurological progression suggestive of MS (PPMS)	1 year of disease progression (retrospectively or prospectively determined) plus 2 of 3 of the following criteria <sup>d</sup> : 1. Evidence for DIS in the brain based on ≥1 T2 lesions in the MS-characteristic (periventricular, juxtacortical, or infratentorial) regions 2. Evidence for DIS in the spinal cord based on ≥2 T2 lesions in the cord 3. Positive CSF (isoelectric focusing evidence of oligoclonal bands and/or elevated IgG index)

Polman et al, *Ann Neurol*, 2011

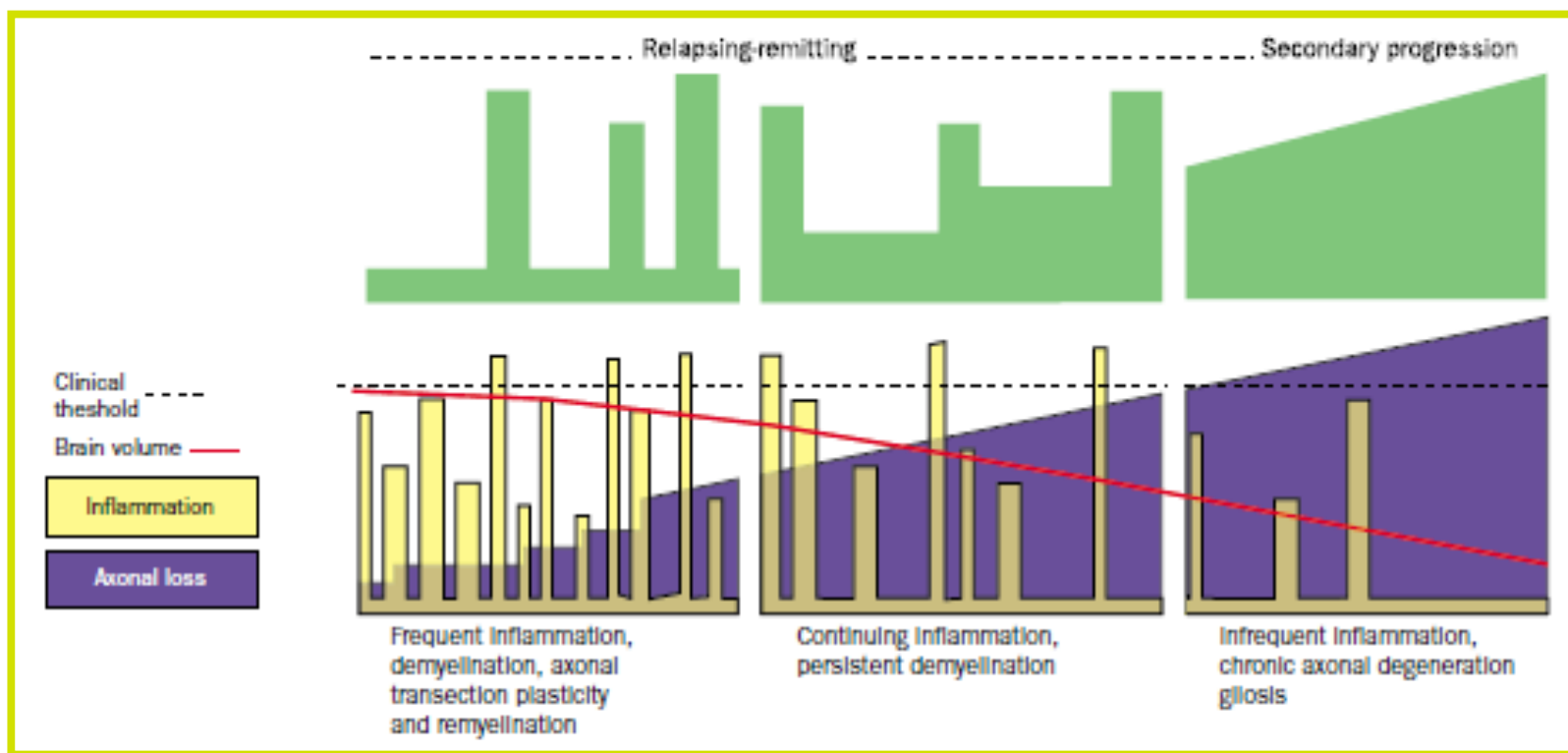






# III. SEP

# III. SEP



Compston & Coles, *Lancet*, 2002

Actuellement: paradoxe clinico-radiologique

## The clinico-radiological paradox in multiple sclerosis revisited Frederik Barkhof

The use of magnetic resonance imaging as a surrogate outcome measure in clinical trials, or even as a prognosticator in the assessment of the natural evolution, assumes a close relationship between extent and rate of development of magnetic resonance imaging abnormalities with the clinical status and rate of development of disability. While it may seem obvious that patients who develop new lesions are worse off than those without new lesions, the association between clinical findings and radiological extent of involvement is generally poor. In this review, various confounders are discussed, including inappropriate clinical rating, lack of histopathological specificity (especially for axonal loss), neglect of spinal cord involvement, underestimation of damage to the normal appearing brain tissue (both white and gray matter), and masking effects of cortical adaptation. It is concluded that much progression has been made in magnetic resonance techniques so that the clinico-radiological dissociation has indeed proved to be a paradox. Thus, the relevance of normal appearing brain tissue damage, residual brain volume, spinal cord damage and cerebral plasticity had to be reiterated. The increased awareness of the subtle interplay between these dimensions should be kept in mind when magnetic resonance is used as a surrogate outcome measure. This corroborates with conventional wisdom that one should not rely on a single magnetic resonance measure, but take full advantage of the fact that magnetic resonance is able to provide multidimensional information. *Curr Opin Neurol* 15:239-245. © 2002 Lippincott Williams & Wilkins.

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*Current Opinion in Neurology* 2002, 15:239-245

### Abbreviations

EDSS Expanded Disability Status Scale  
MRI magnetic resonance imaging  
MSFC Multiple Sclerosis Functional Composite

© 2002 Lippincott Williams & Wilkins  
1360-7540

### Introduction

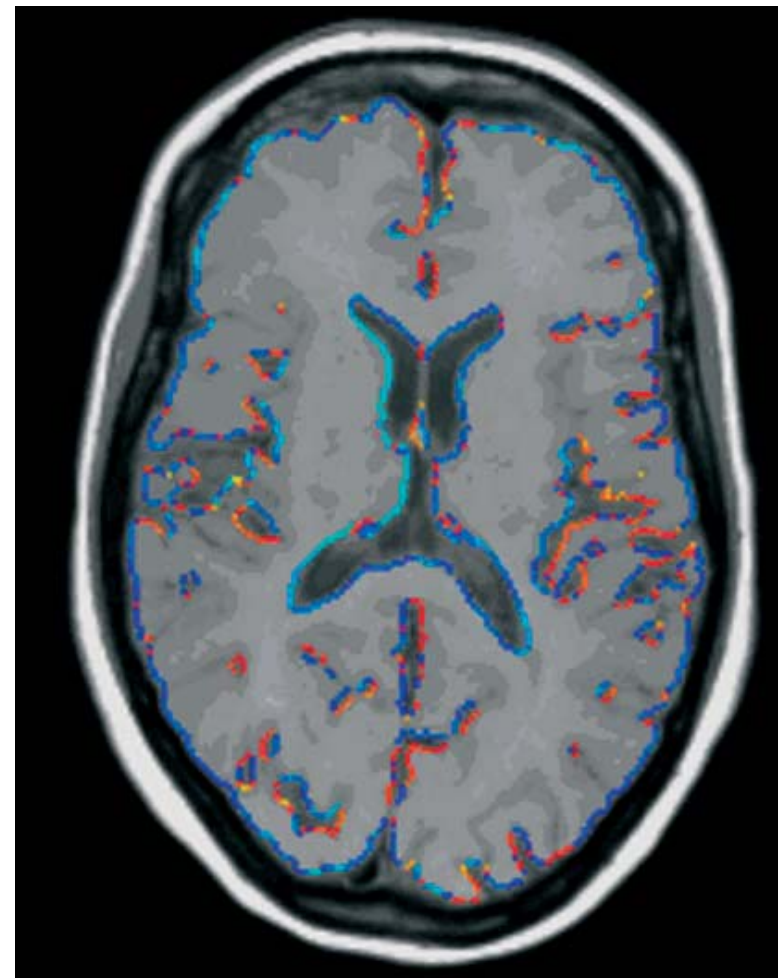
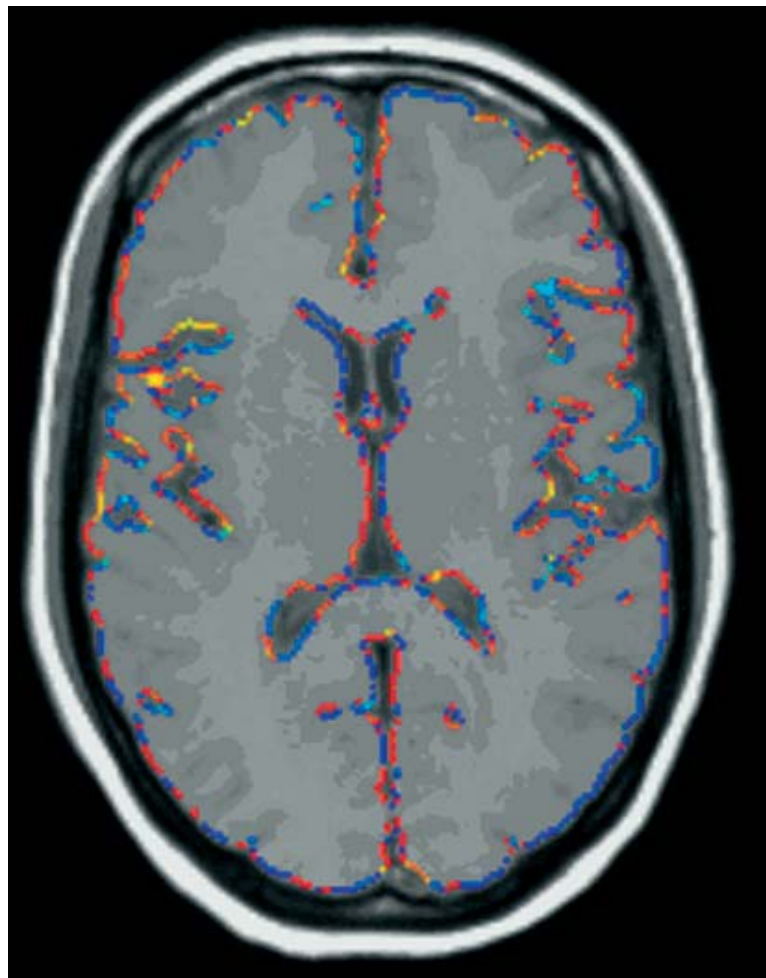
In the field of multiple sclerosis, magnetic resonance imaging (MRI) is widely applied to ascertain the diagnosis. In the recently published diagnostic criteria [1\*\*], MRI is given a much more prominent role, allowing one to demonstrate not only dissemination in space, but also dissemination in time [2\*]. Its diagnostic sensitivity reflects the ability to identify clinically silent lesions, which, when studied sequentially, display a dynamic pattern of waxing and waning that is governed by a variety of immunological and other factors [3-10,11\*,12,13]. Unlike the clinical picture, there are frequent new silent lesions, which can be used to study the natural course or attempts to modify the natural course. In fact, gadolinium-enhanced and T2-weighted MRI is widely used to monitor treatment efficacy, serving as an important secondary outcome measure in most phase III clinical trials [14\*,15,16], and often being used as the primary outcome in exploratory ('proof-of-concept') phase II studies [17-20].

The use of MRI as a surrogate outcome measure in clinical trials, or even as a prognosticator in the assessment of the natural evolution, assumes a close relationship between extent and rate of development of MRI abnormalities with the clinical status and rate of development of disability. While it may seem obvious that patients who develop new lesions are worse off than those without new lesions, the association between clinical findings and radiological extent of involvement generally is poor. In this review, various confounders will be discussed to what has optimistically been labeled 'the clinico-radiological paradox', assuming that the relation is indeed tighter when controlled for these confounders. This review discusses progress that has been made in overcoming some of these limitations, including inappropriate clinical rating, lack of histopathological specificity (especially for axonal loss), neglect of spinal cord involvement, underestimation of damage to the normal appearing brain tissue (both white and gray matter), and compensation by cortical adaptation.

### New rating scales: including cognitive assessment

The limitations of the Expanded Disability Status Scale (EDSS) are well known. They include incomplete coverage of central nervous system domains, non-linearity, and observer bias. In an attempt to develop a more comprehensive scale, the Multiple Sclerosis Functional Composite (MSFC) has been developed. The MSFC combines information from three independent

# III. SEP



🔸 Développement: mesure atrophie/VBM

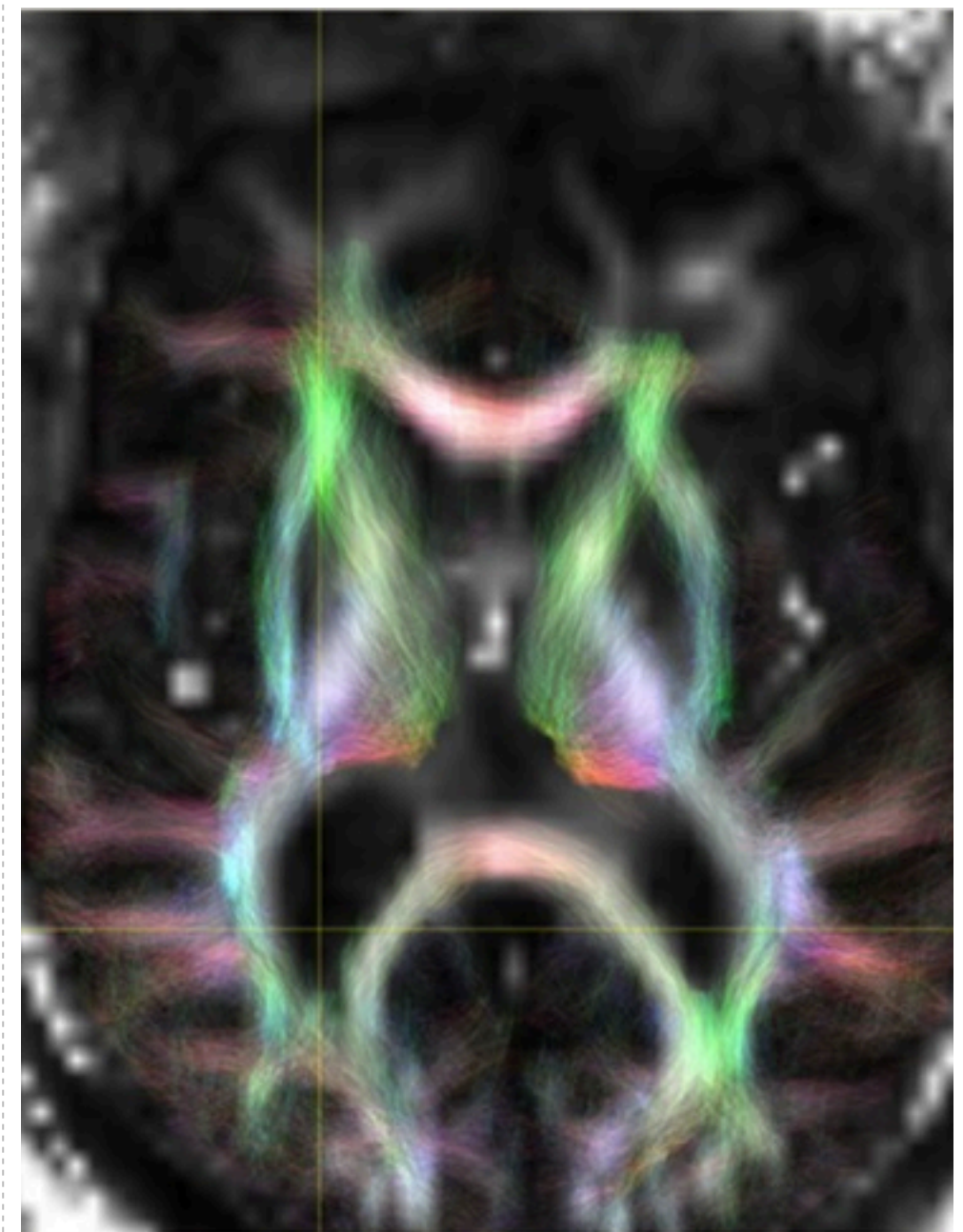
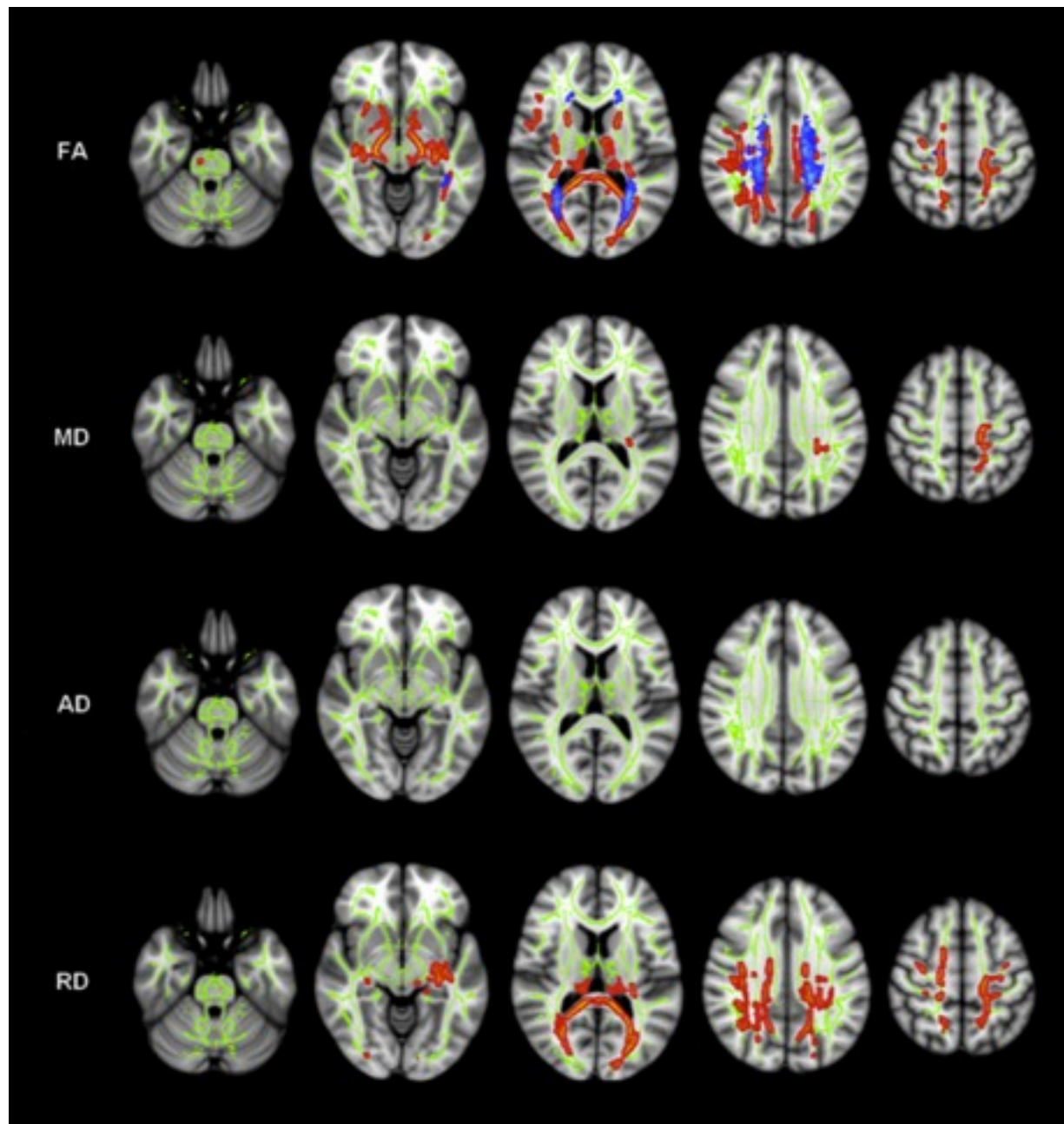
Bakshi et al, *Lancet Neurol*, 2008





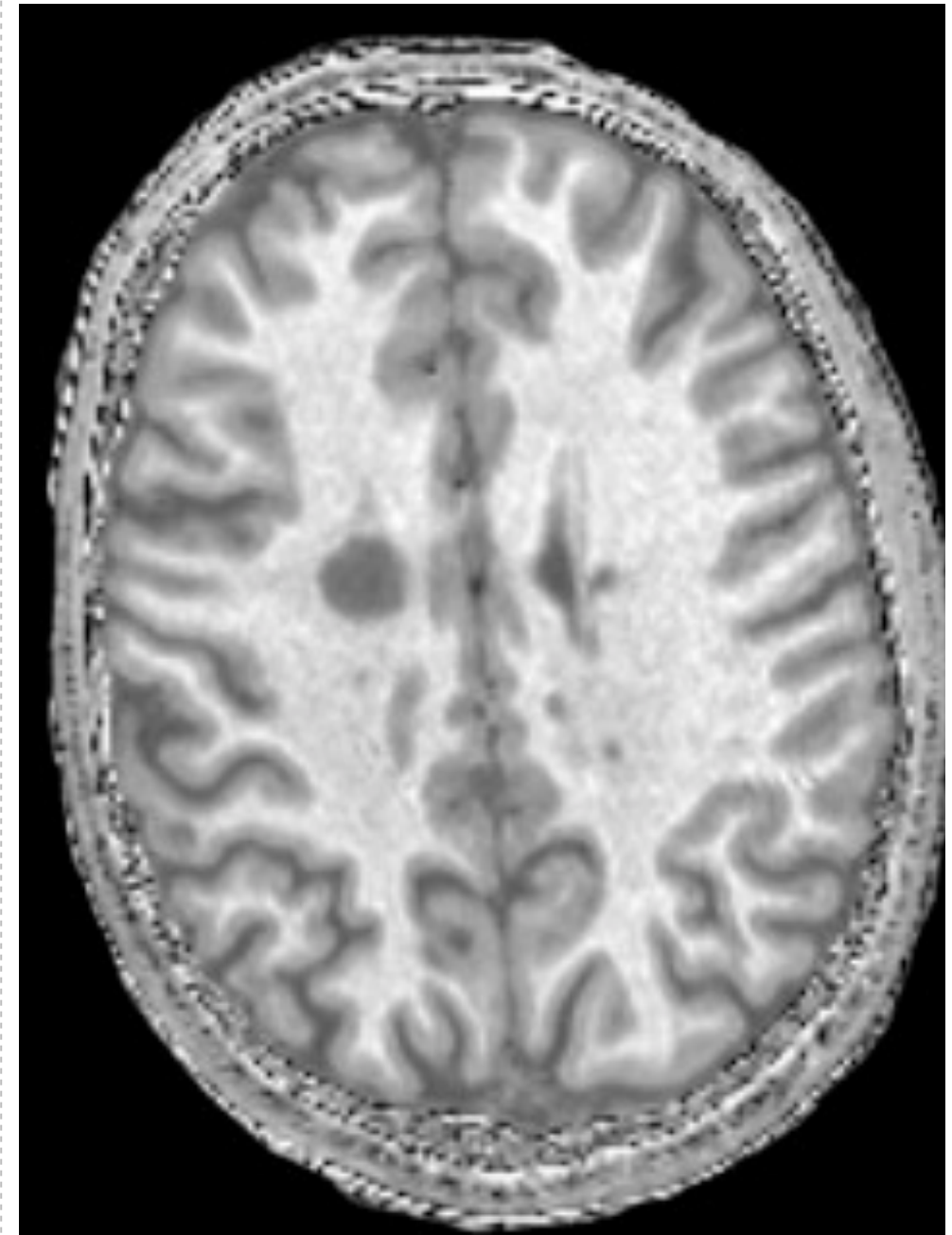
# III. SEP

 Dével.:  
 Diffusion  
 (MD/FA/  
 DTI)



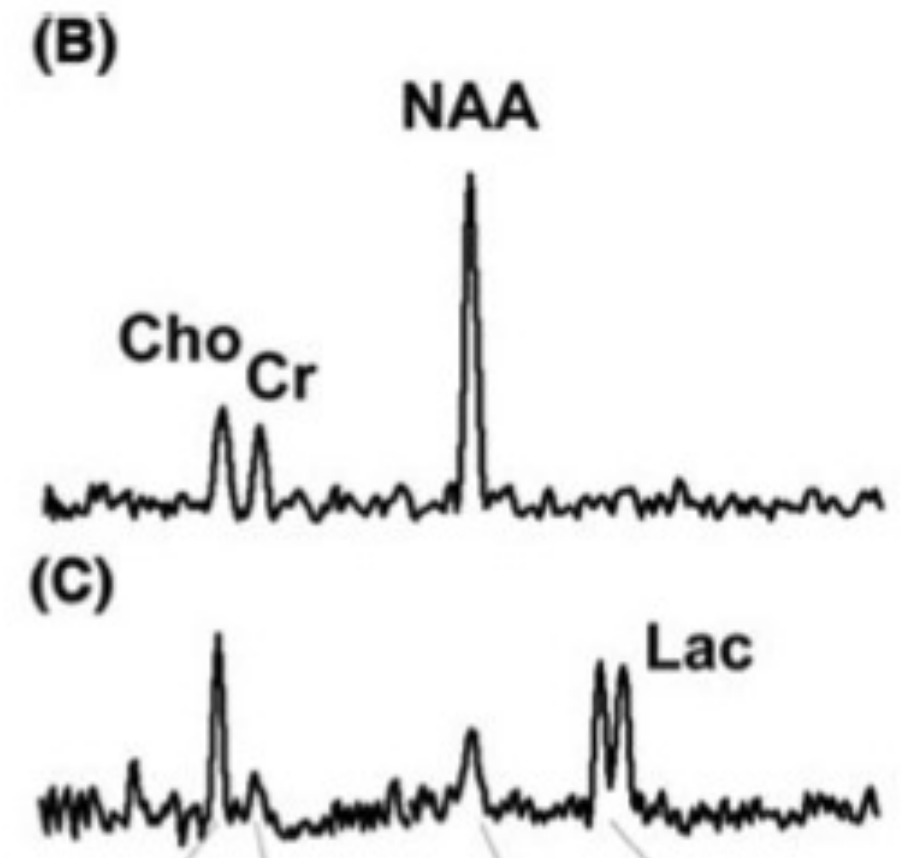
# III. SEP

🔸 Développement: Transfert de Magnétisation



# III. SEP

- ❖ H-MRS: mesure de certains métabolites reflétant l'intégrité structurelle et fonctionnelle du tissu cérébral.

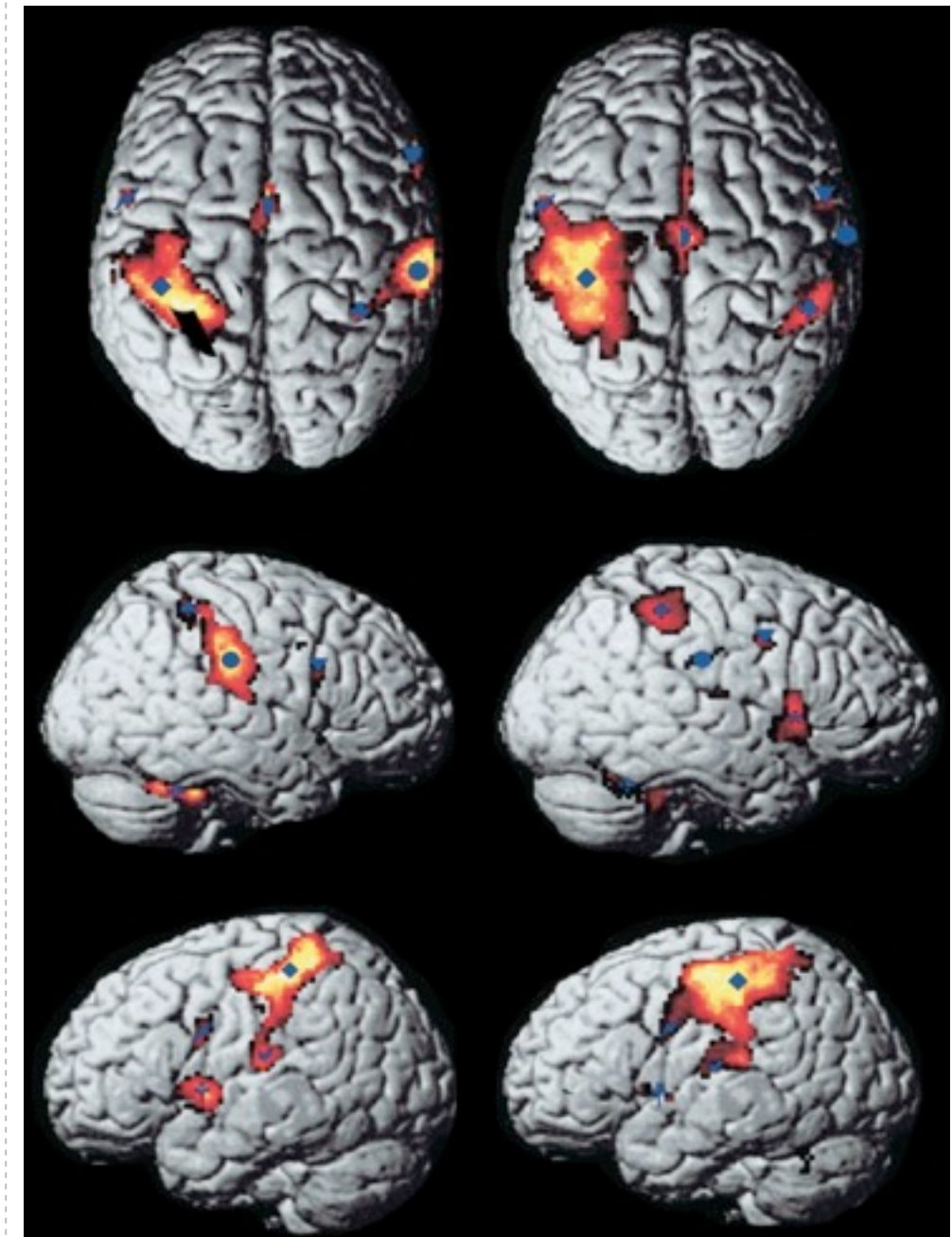


Filippi et al, in: Handbook of multiple sclerosis, Cook (ed) 2006



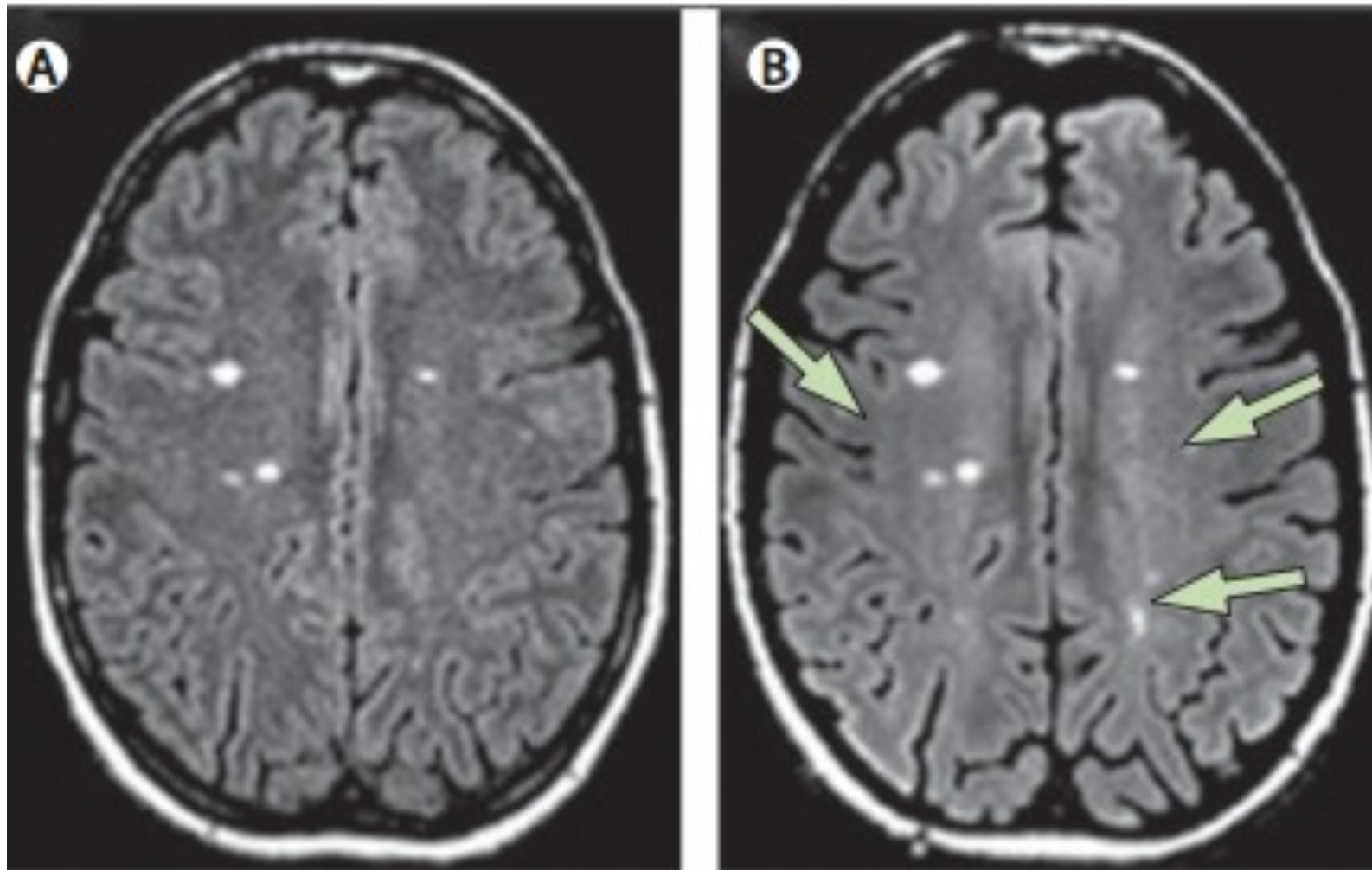
# III. SEP

- ❏ Développement: fMRI
  - ❏ modification des RSNs y compris DMN chez patients SEP
  - ❏ Modifications corrélées aux épreuves cognitives et au phénotype SEP
  - ❏ Pourrait représenter l'effet de la neuroplasticité initialement





# III. SEP



Bakshi et al, *Lancet Neurol* 2008

- ❖ Développement: UHF MRI
- ❖ Permet de démontrer lésions plus fines, expliquant certaines anomalies des séquences non-conventionnelles dans NAWM
- ❖ Permet de réaliser IRM au sodium (étude gradient Na + intra/extra => reflet intégrité membranaire)

# III. SEP

🍯 Futur/autres:

🍯 MWF

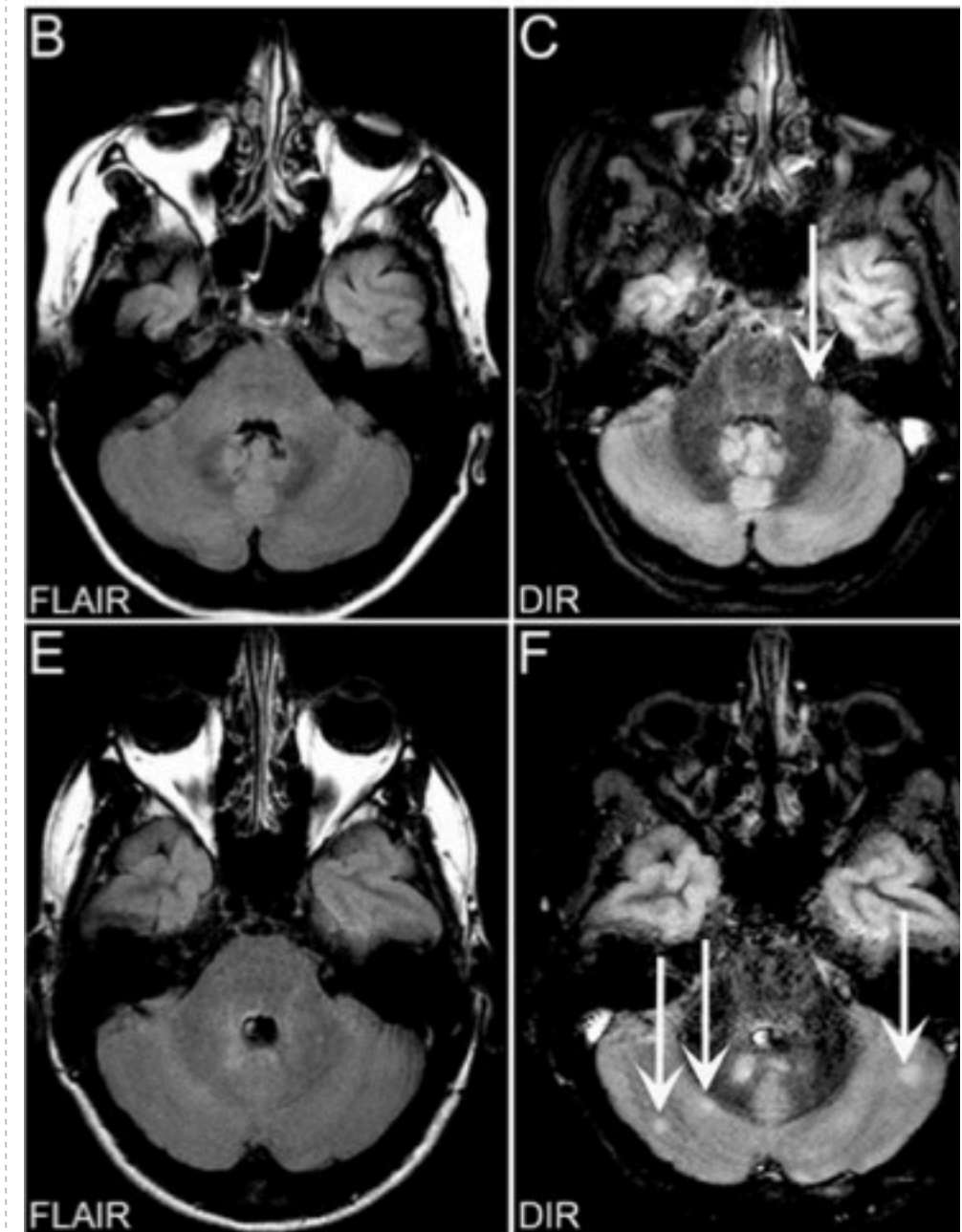
🍯 Perfusion MRI

🍯 SWI

🍯 DIR

🍯 Iron deposit quantification

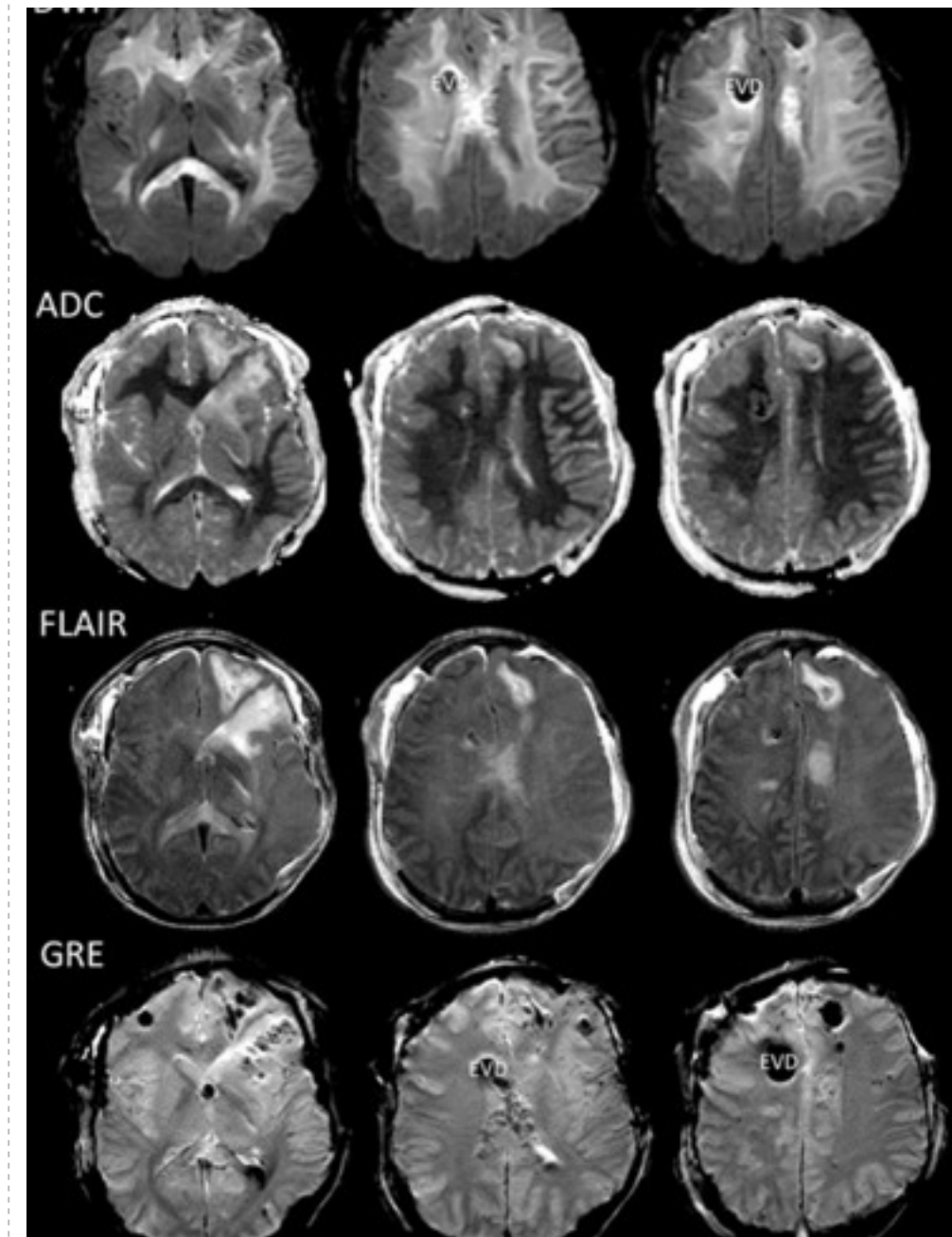
🍯 Connectome



# V. Trauma

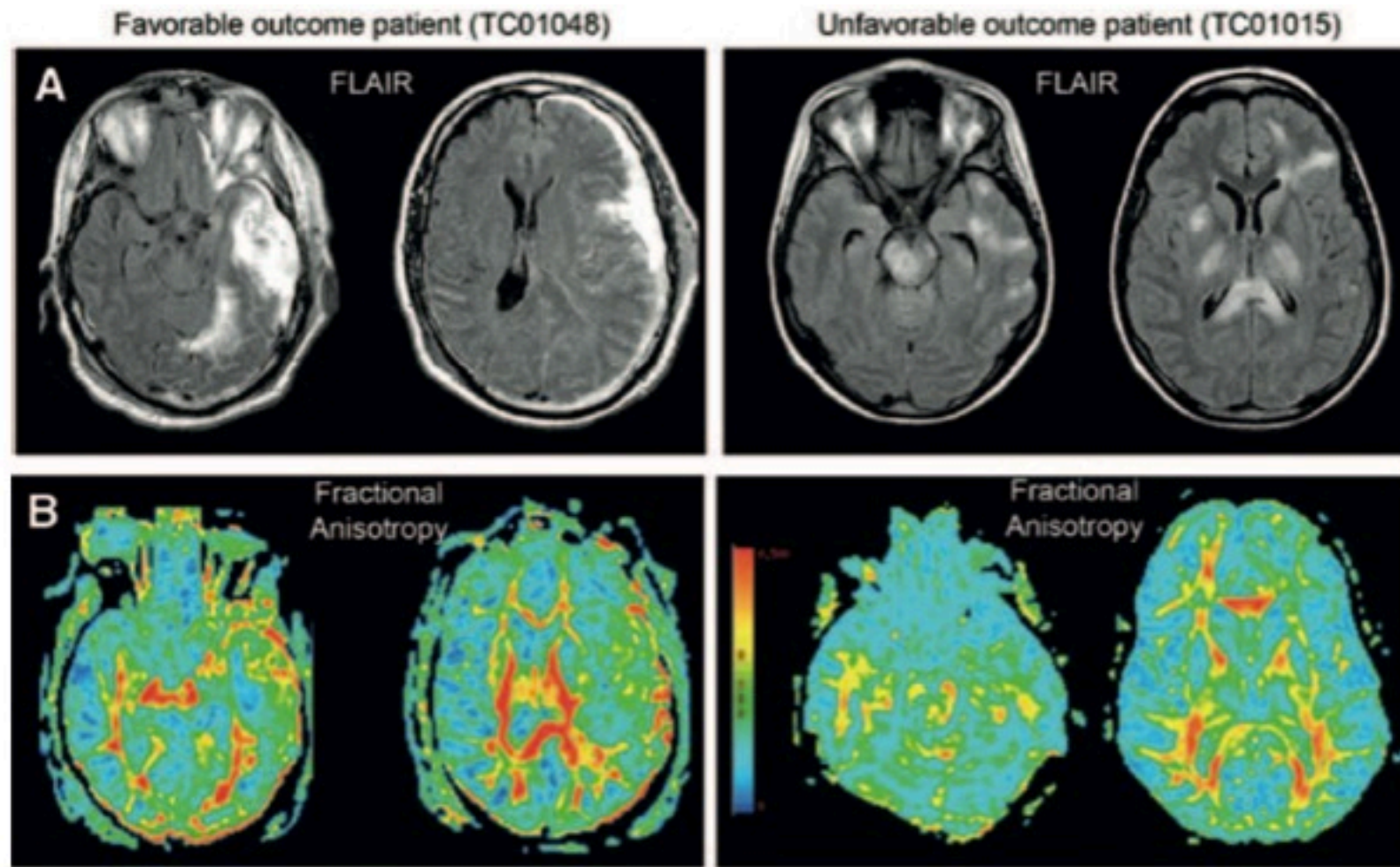
- ❏ Traumatisme crânien sévère: CT-Scan Cérébral !
- ❏ Trauma léger/modéré: Canadian head-CT rule détermine nécessité de scanner.
- ❏ Place de la MRI: montre DAI, meilleure résolution pour tronc et fosse post. Peu/pas d'intérêt pour prise en charge aiguë.

Edlow et al, *Neurocrit Care*, 2013



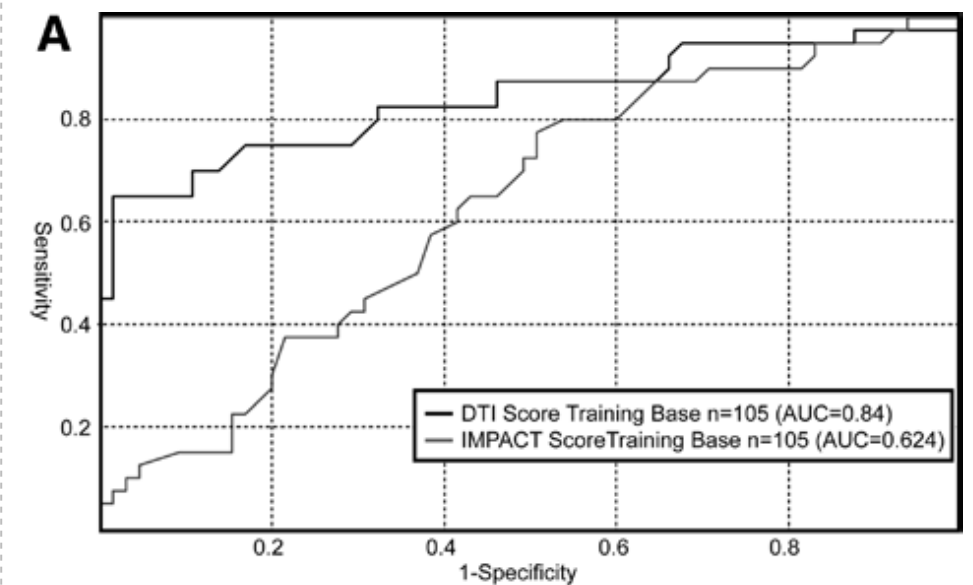


# V. Trauma



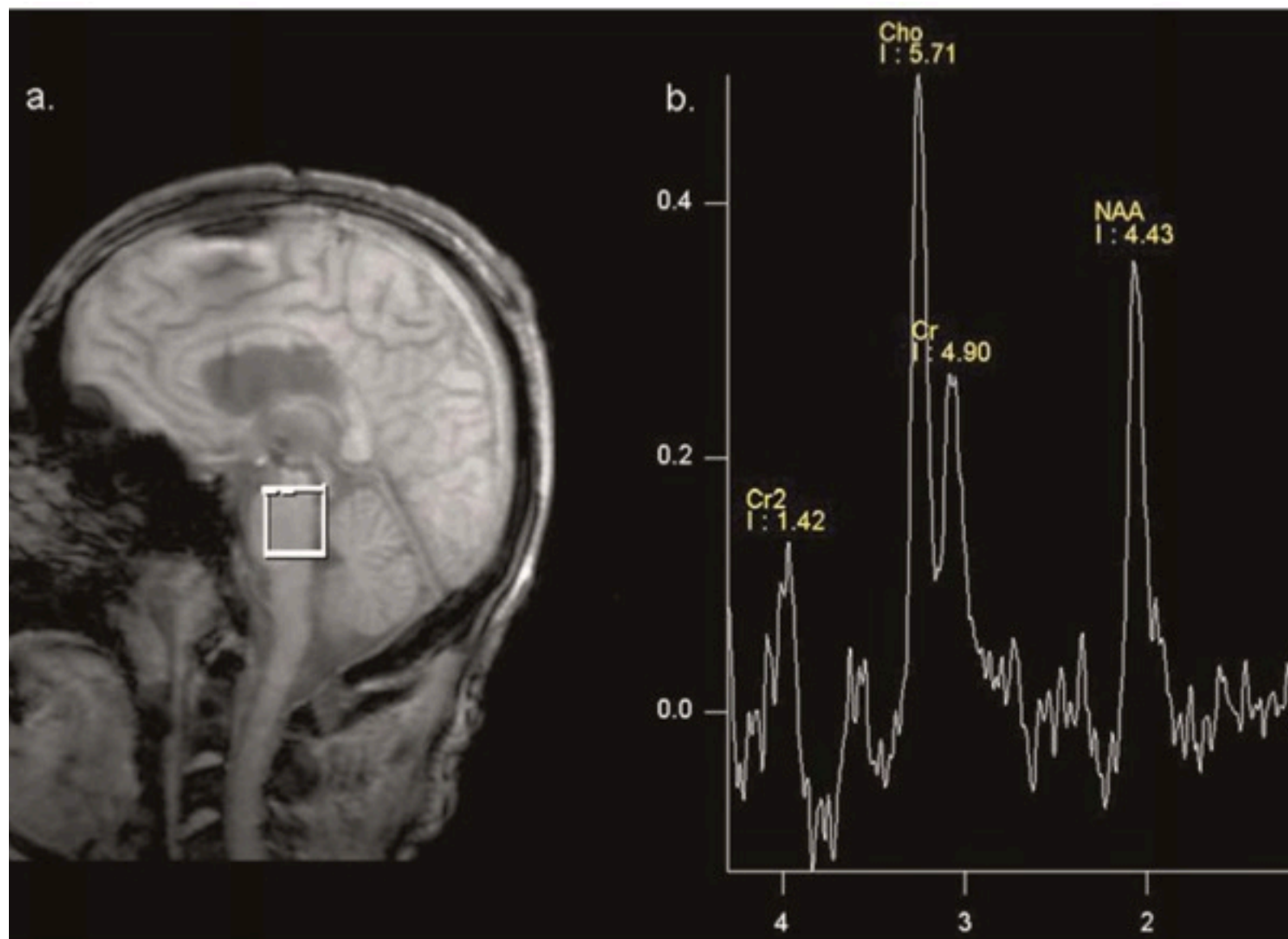
Galanaud *et al*, *Anesthesiology*, 2012

En développement:  
utilisation IRM (DTI) en aigu  
comme outil pronostic





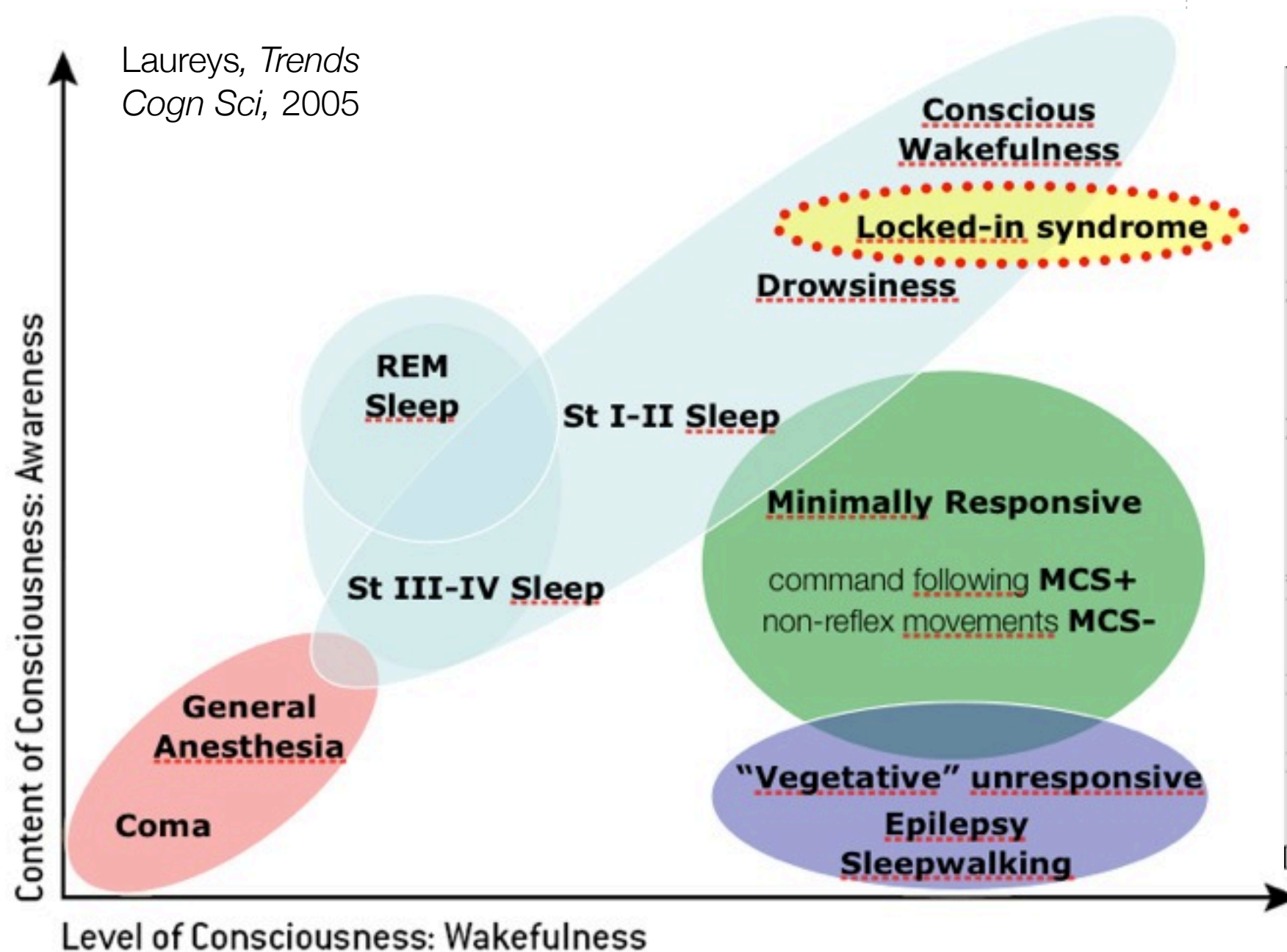
# V. Trauma



- ❖ Développement: H-MRS
- ❖ NAA/Cr plus faible chez patients avec mauvais outcome (GOS 1-3)
- ❖ GOS anti-corrélé avec NAA et corrélé avec La

Tshibanda et al, *Prog Brain Res*, 2009

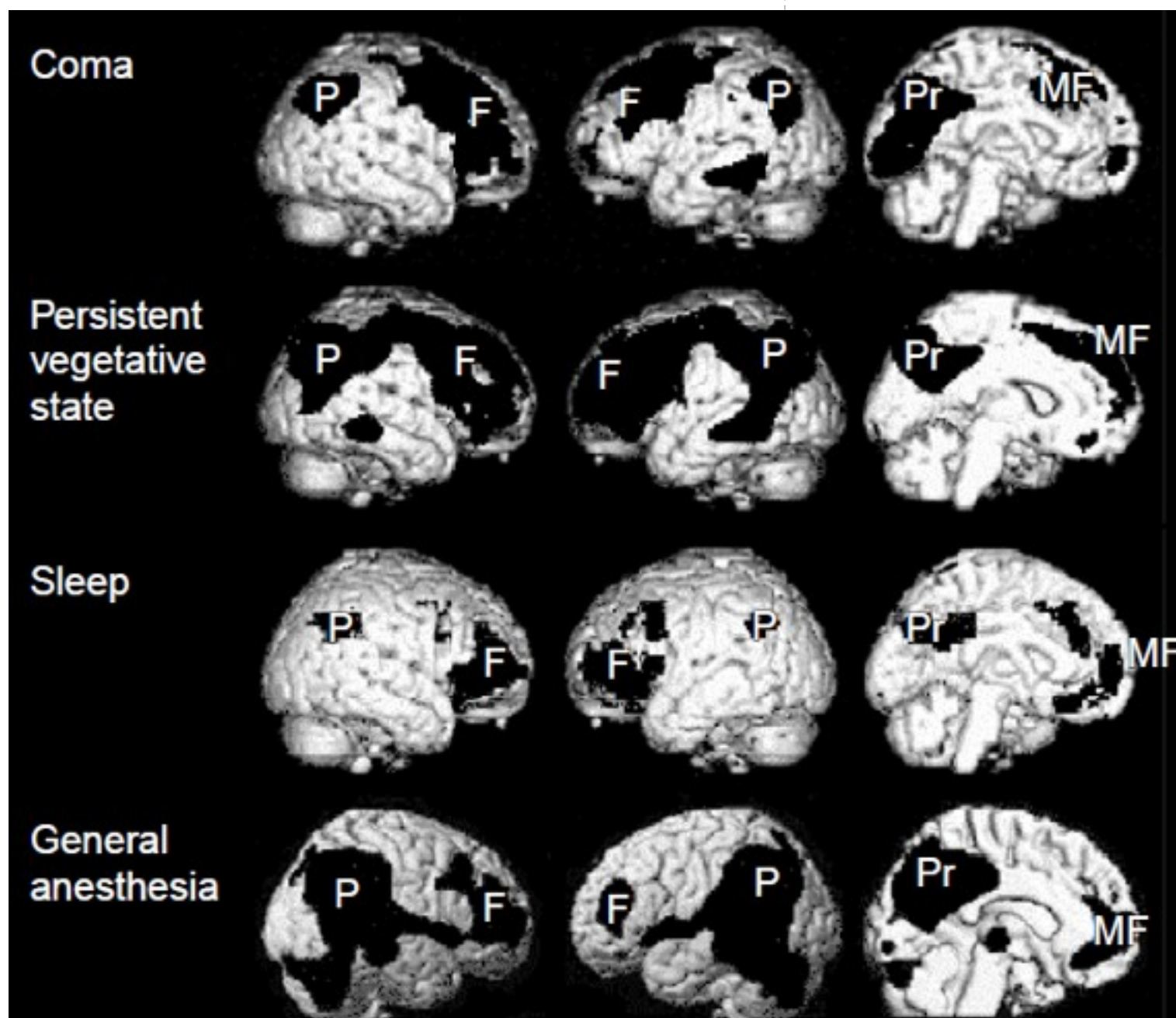
# V. Trauma



JFK COMA RECOVERY SCALE - REVISED ©2004	
Record Form	
Patient:	Date:
<b>AUDITORY FUNCTION SCALE</b>	
4 - Consistent Movement to Command *	
3 - Reproducible Movement to Command *	
2 - Localization to Sound	
1 - Auditory Startle	
0 - None	
<b>VISUAL FUNCTION SCALE</b>	
5 - Object Recognition *	
4 - Object Localization: Reaching *	
3 - Visual Pursuit *	
2 - Fixation *	
1 - Visual Startle	
0 - None	
<b>MOTOR FUNCTION SCALE</b>	
6 - Functional Object Use †	
5 - Automatic Motor Response *	
4 - Object Manipulation *	
3 - Localization to Noxious Stimulation *	
2 - Flexion Withdrawal	
1 - Abnormal Posturing	
0 - None/Flaccid	
<b>OROMOTOR/VERBAL FUNCTION SCALE</b>	
3 - Intelligible Verbalization *	
2 - Vocalization/Oral Movement	
1 - Oral Reflexive Movement	
0 - None	
<b>COMMUNICATION SCALE</b>	
2 - Functional: Accurate †	
1 - Non-Functional: Intentional *	
0 - None	
<b>AROUSAL SCALE</b>	
3 - Attention	
2 - Eye Opening w/o Stimulation	
1 - Eye Opening with Stimulation	
0 - Unarousable	
<b>TOTAL SCORE</b>	

Giacino, Arch Phys Med Rehab, 2004

# V. Trauma



 PET-CT

Vigilance conservée  
Pas de «conscience»

Pas de vigilance  
Pas de «conscience»

TRENDS in  
Cognitive  
Sciences

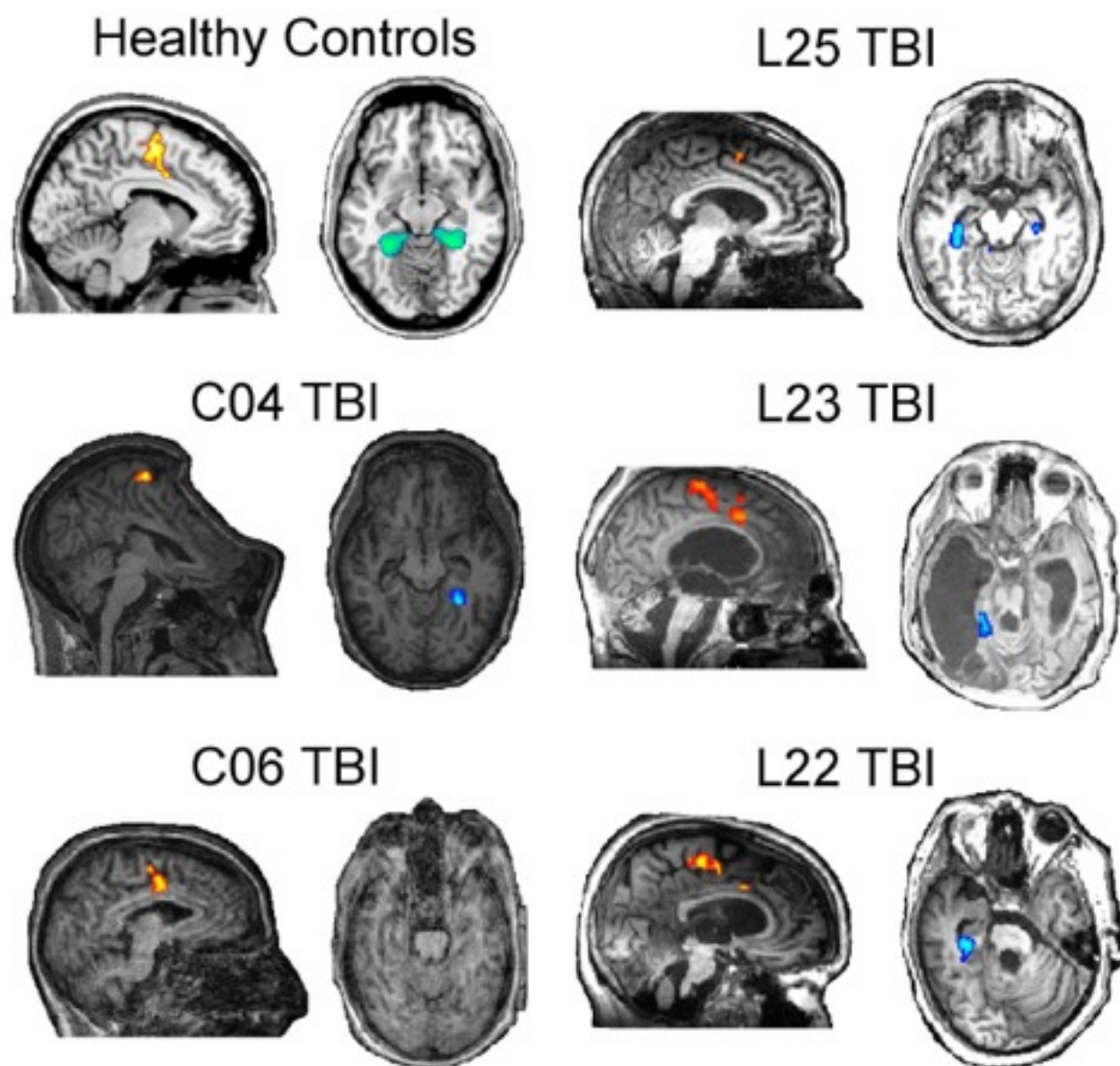
Baars, Ramsoy, Laureys,  
*Trends in Neurosciences*  
2003



# V. Trauma

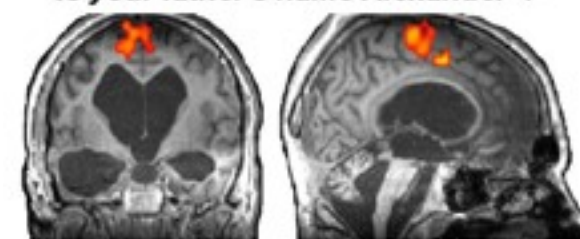
❖ fMRI: paradigme actif

Monti & Vanhaudenhuyse, et al  
*New England J Med* 2010

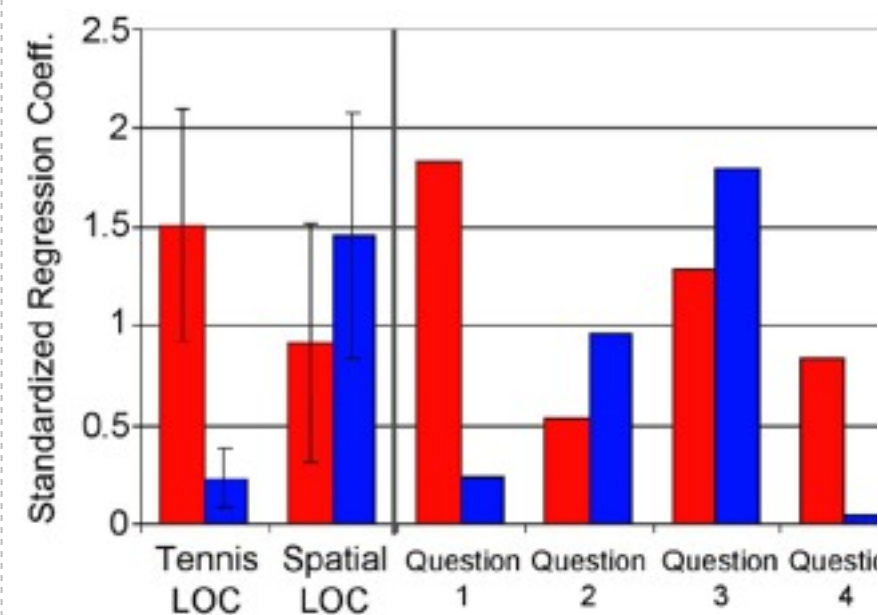
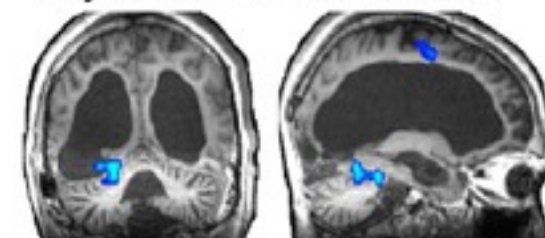


Imagine **Tennis** to answer 'YES'  
Imagine **Navigating** to answer 'NO'

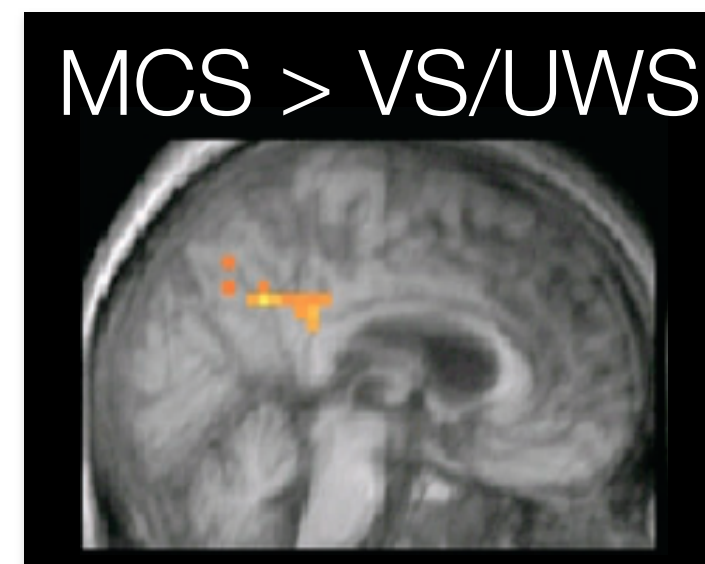
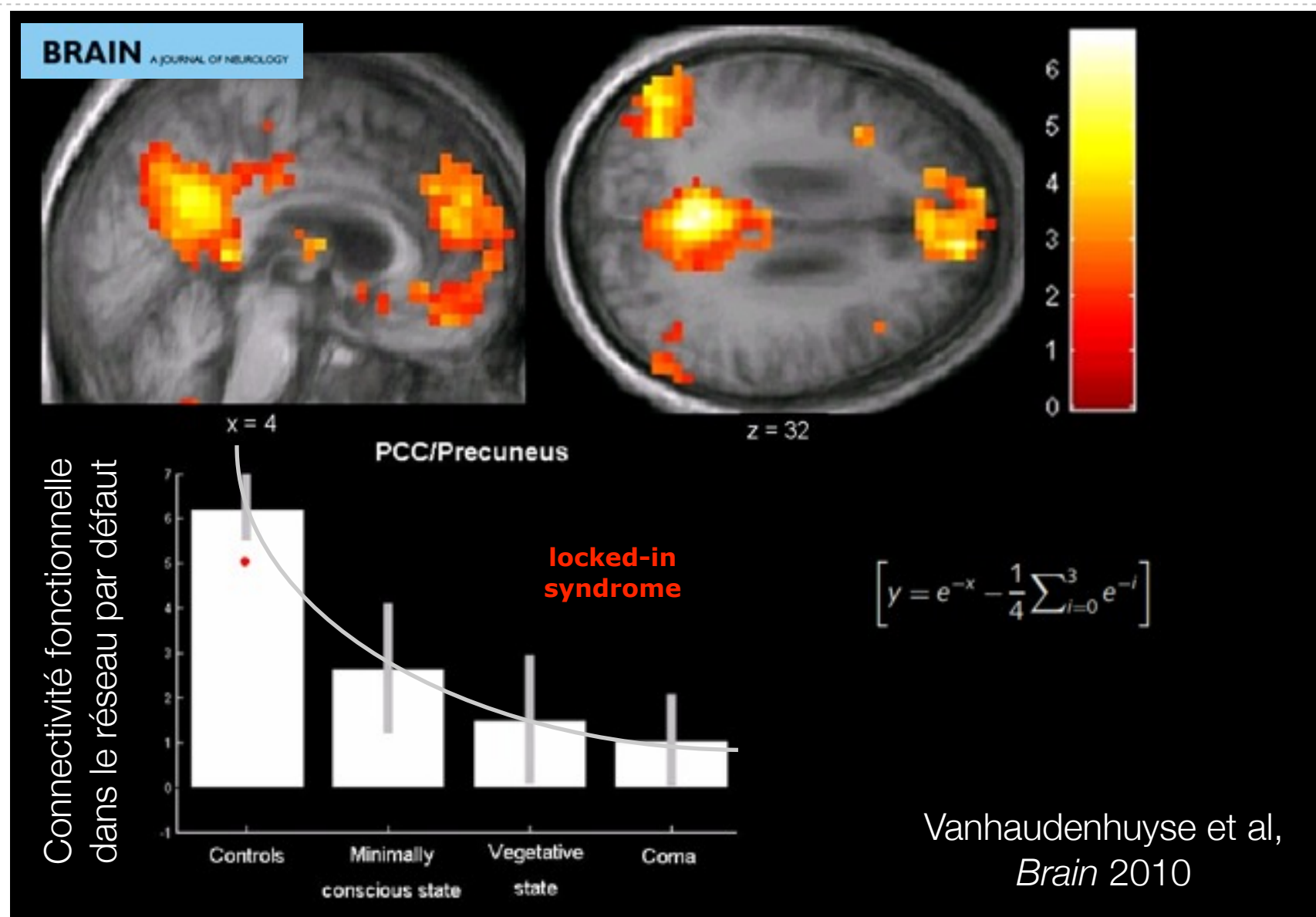
Is your father's name Alexander ?



Is your father's name Thomas ?





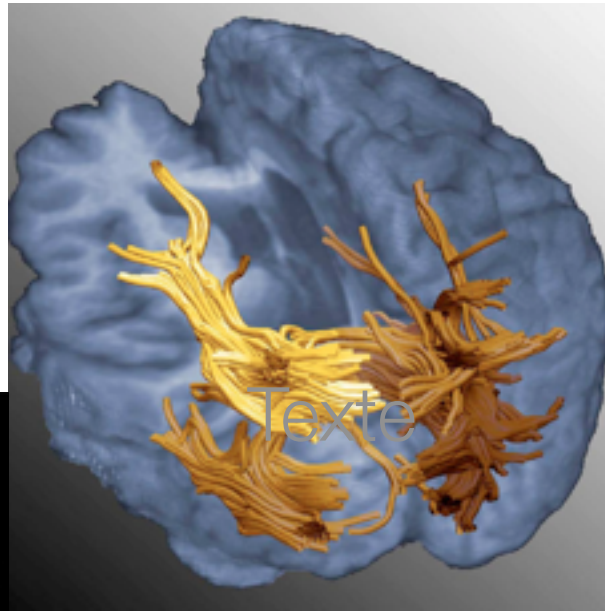
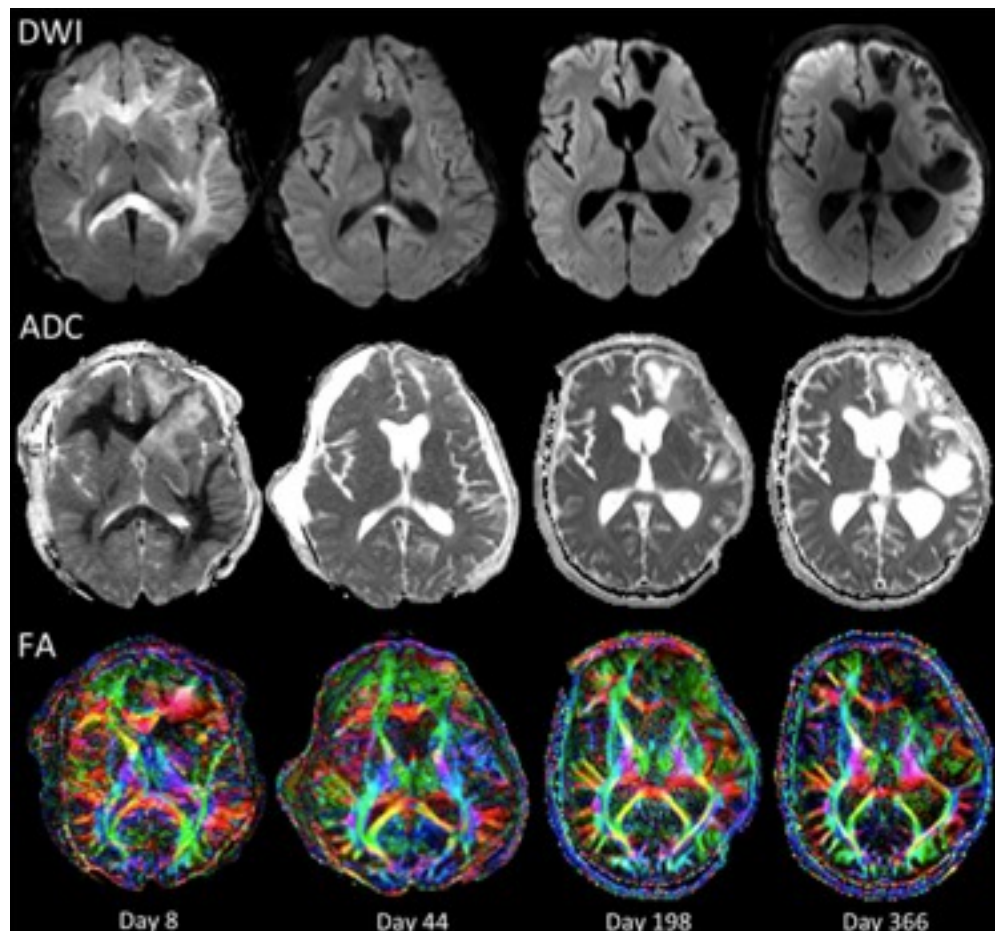


# V. Trauma

🔲 fMRI: resting state

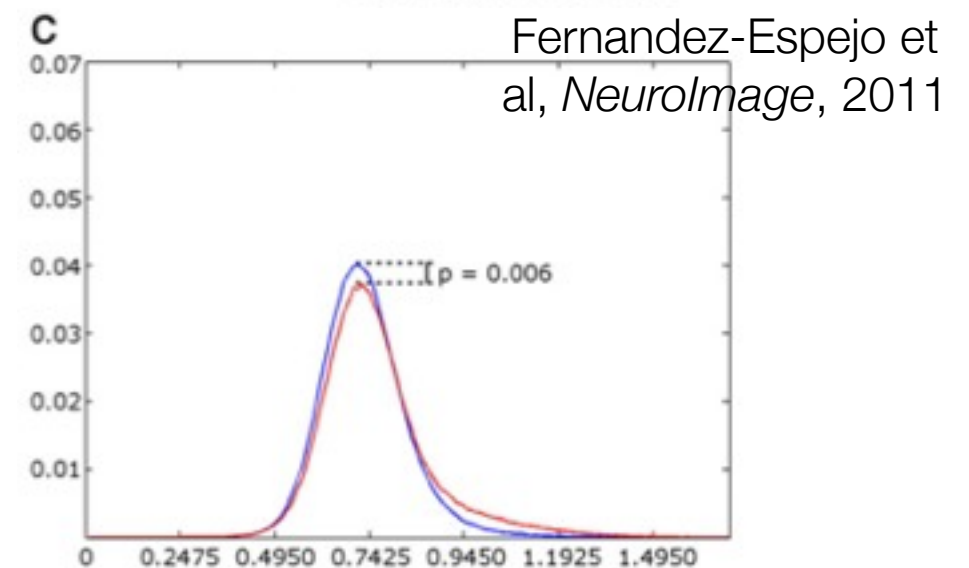
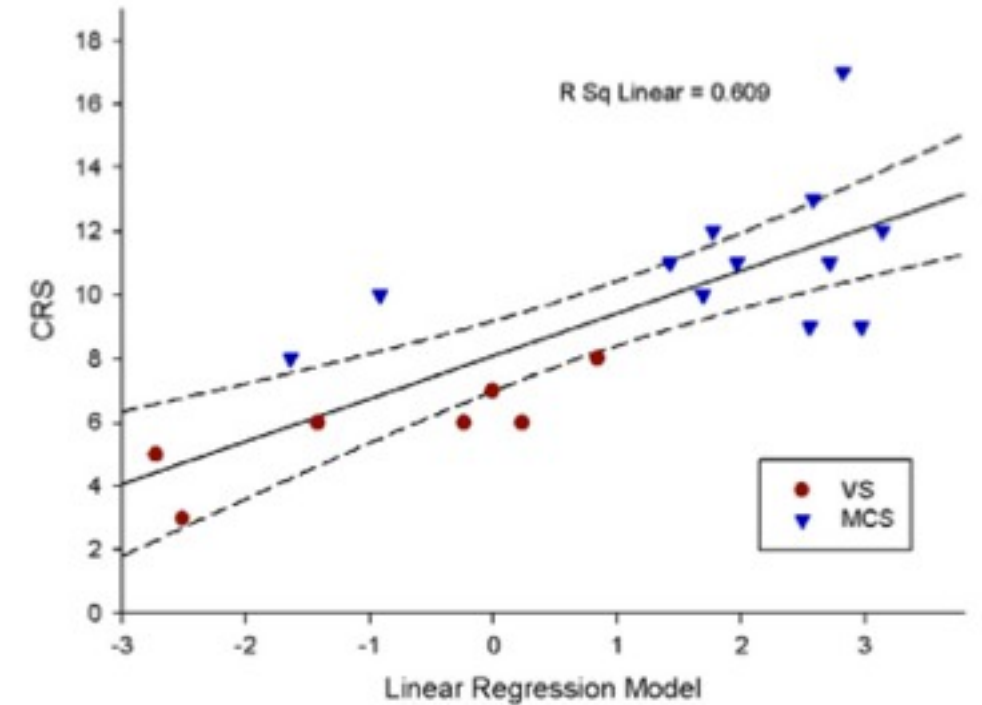
# V. Trauma

## Diffusion et DTI



Voss et al, *J Clin Invest*, 2006

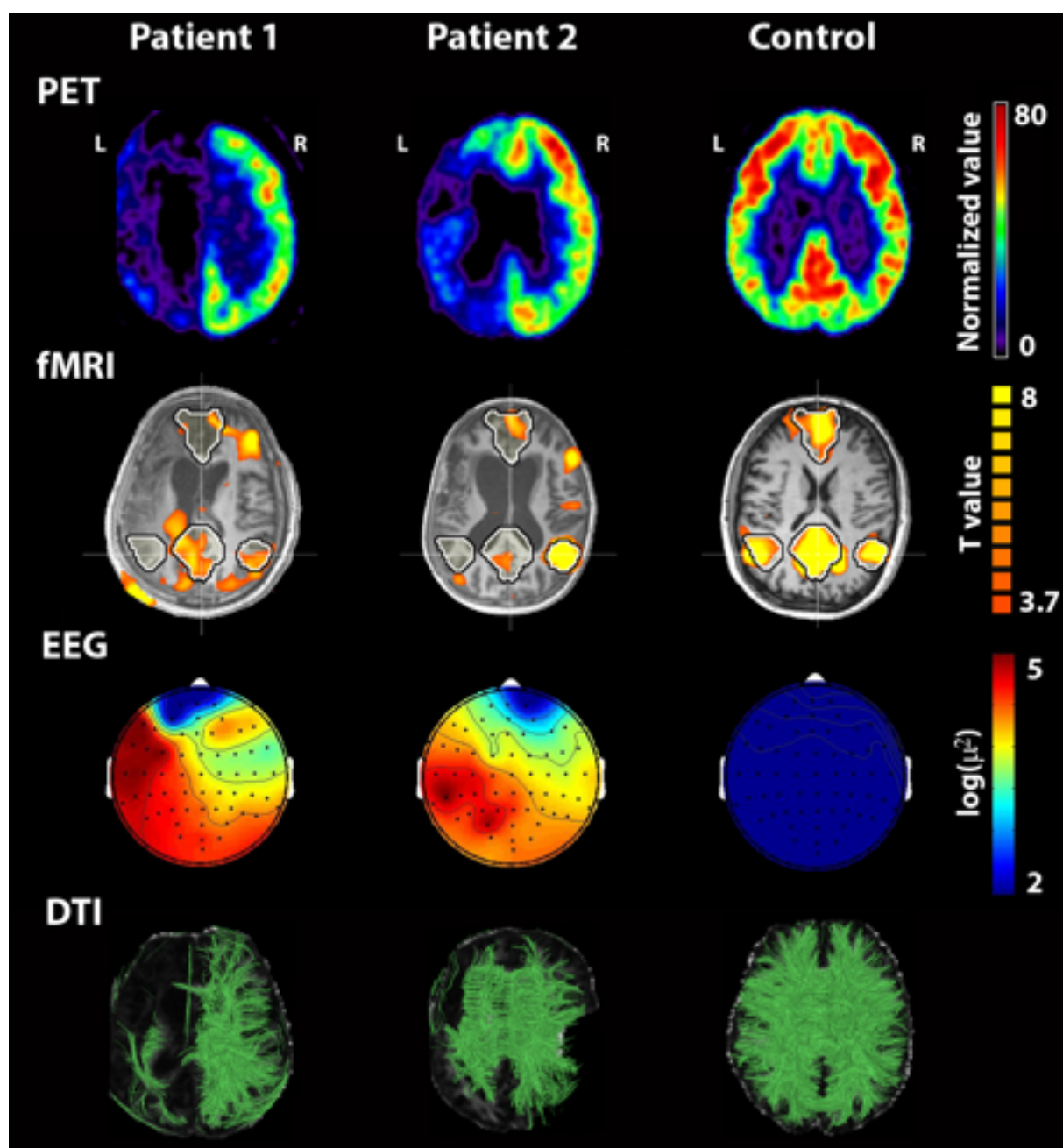
Edlow et al, *Neurocrit Care*, 2013



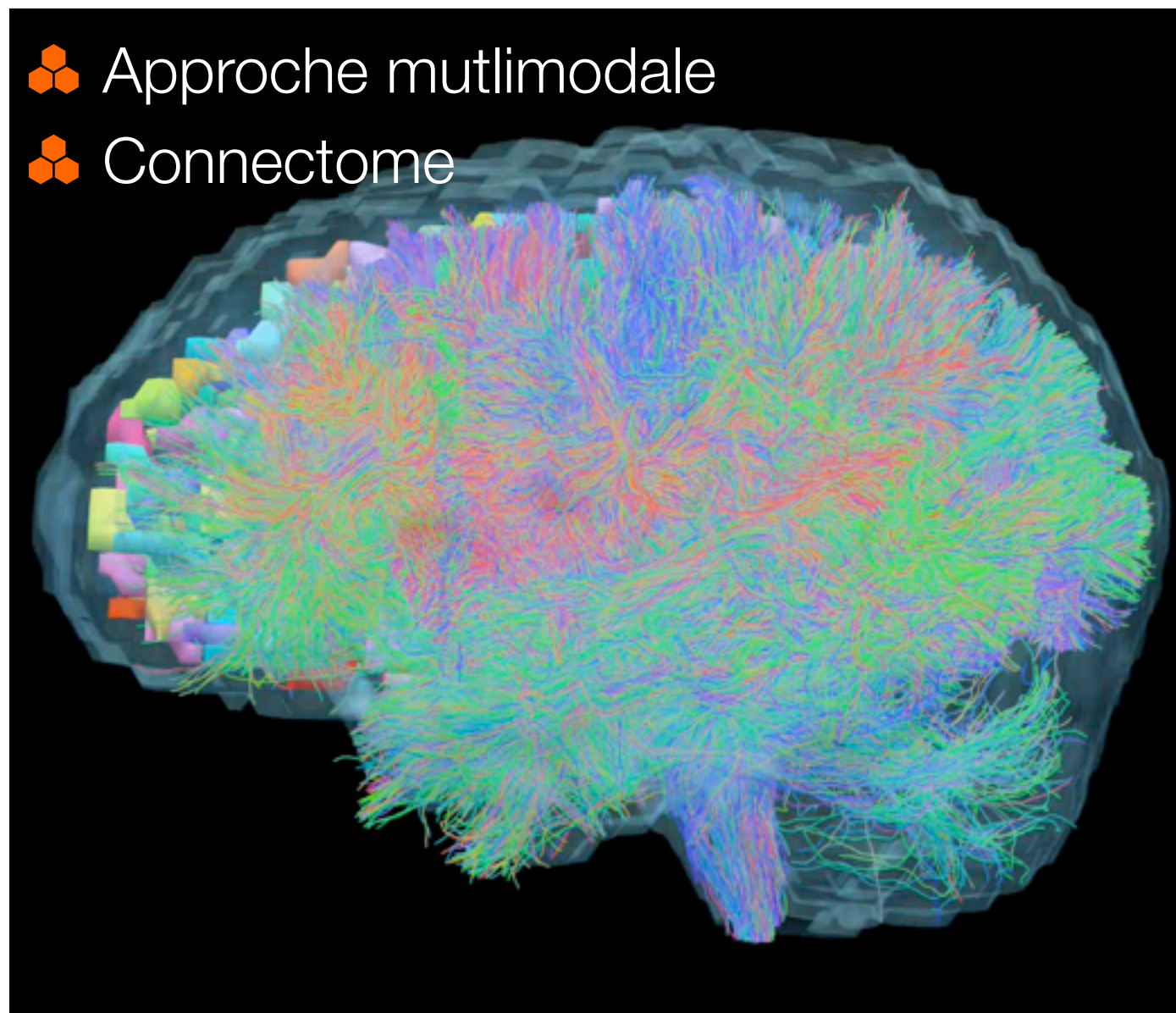
Fernandez-Espejo et al, *NeuroImage*, 2011



# V. Trauma



 Approche multimodale  
 Connectome



Erik Ziegler, Cyclotron Art Committee

# VI. Conclusion

- ❏ Merci.
- ❏ L'ensemble des illustrations non référencées dans la partie SEP m'ont été gentiment fournies par le Dr E. Lommers, que je remercie chaleureusement.

