



NeuroRecovery Lab

ADVANCING NEUROREHABILITATION THROUGH NEUROMODULATION



CHU
de Liège

CLINIQUE DE LA CONSCIENCE ET DE
NEUROREVALIDATION

Exploring Neurophysiological & Neuroimaging Alterations in Concussion

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NeuroRehab & Consciousness Clinic

University Hospital of Liege





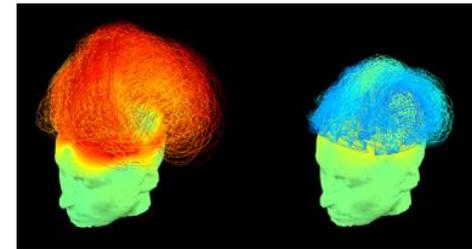
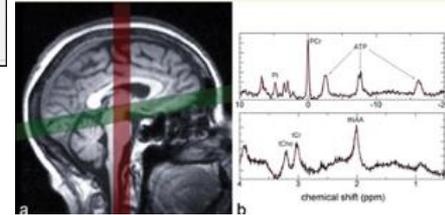
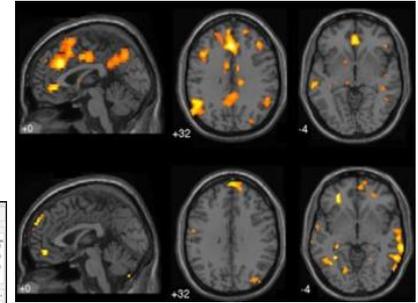
Concussions & persistent symptoms

- Concussion or mild TBI = 90% total TBI
 - 30-50% develop persistent post concussion symptoms
 - Somatic, cognitive, emotional, sleep disturbances
 - 10-27% unemployed at 12 month (TRACK-TBI)
 - Standard exams usually normal (CT-MRI)
-
- ➔ Lack of uniformity for the diagnostic & management
 - ➔ Underdiagnosed (& reported) – silent epidemic

Advanced imaging techniques



- Functional Magnetic Resonance Imaging - fMRI
- Diffusion Tensor Imaging – DTI
- Susceptibility Weighted Imaging – SWI
- Magnetic Resonance Spectroscopy – MRS
- Positron Emission Tomography – PET
- Electrophysiology – EEG
- Transcranial Magnetic Stimulation – TMS
- ...



New (exploratory) imaging techniques



Amsterdam Consensus Conference 2022
205 studies – 81 (f)MRI & 29 EEG

Systematic review

Role of biomarkers and emerging technologies in defining and assessing neurobiological recovery after sport-related concussion: a systematic review

Jason B Tabor ,¹ Benjamin L Brett ,² Lindsay Nelson,² Timothy Meier,²
Linden C Penner,¹ Andrew R Mayer,³ Ruben J Echemendia ,^{4,5}
Thomas McAllister ,⁶ William P Meehan, III,^{7,8} Jon Patricios ,⁹
Michael Makdissi,^{10,11} Silvia Bressan,¹² Gavin A Davis ,¹³ Zahra Premji ,¹⁴
Kathryn J Schneider ,¹ Henrik Zetterberg,¹⁵ Michael McCrea ,²



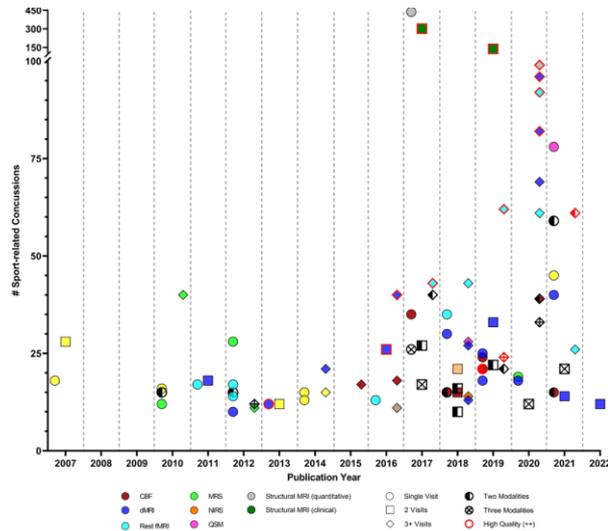
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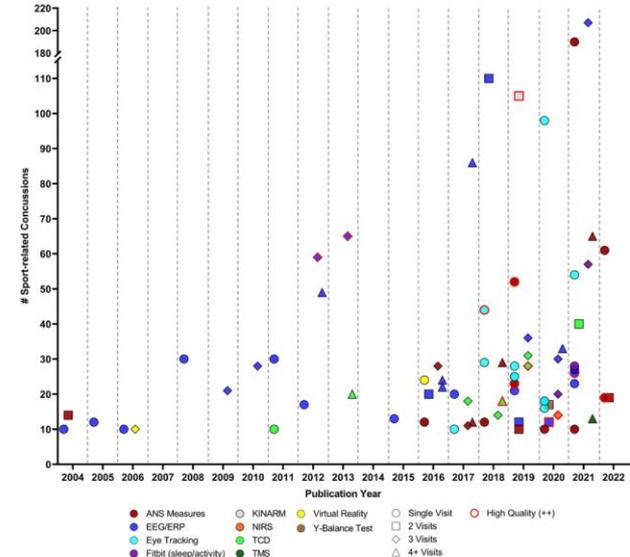
Systematic review

Role of biomarkers and emerging technologies in defining and assessing neurobiological recovery after sport-related concussion: a systematic review

Neuroimaging



Electrophysiology





New (exploratory) imaging techniques

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Role of biomarkers and emerging technologies in defining and assessing neurobiological recovery after sport-related concussion: a systematic review

Systematic review

Key messages

- Use in research for diagnosis & prognosis
- Promising sensitivity for assessing acute neurobiological effects
- Microstructural & functional alterations
- Alterations persists beyond clinical clearance

*BUT heterogeneity & mixed findings
& lack of specificity*

KEY RECOMMENDATIONS

- ⇒ Advances in neuroimaging, fluid biomarkers, genetic testing and emerging technologies continue to provide valuable research tools for the investigation of sport-related concussion neurobiology in the areas of diagnosis, prognosis and recovery.
- ⇒ While these technologies show potential for eventual clinical use, their translation is limited by methodological inconsistencies and a lack of generalisability, based on the existing evidence.
- ⇒ Progression to further validate their clinical utility will require large, multisite, prospective longitudinal studies implementing standardised operating procedures, common data elements, consistent data collection time points and more sophisticated biostatistical approaches to data analysis.



Functional network alterations

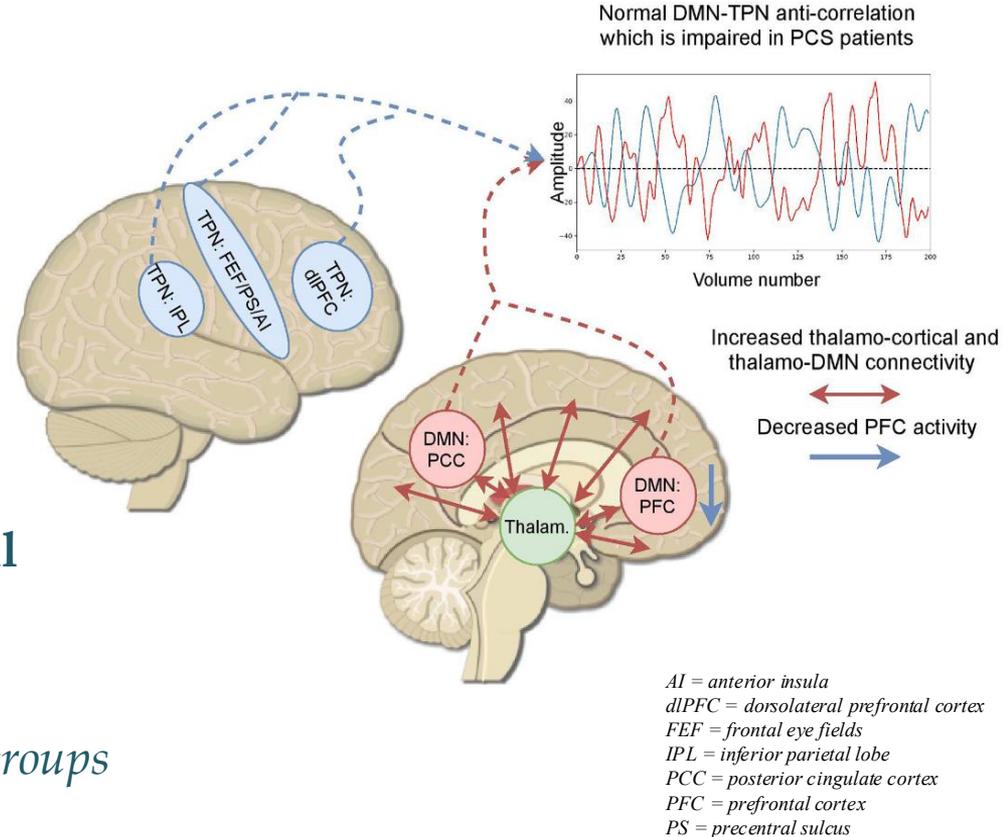
Scoping review - 80 studies

- 10 conventional MRI
- 24 DWI
- 23 fMRI
- 9 EEG/MEG
- 14 others (eg, fNRIS)

fMRI: DMN/TPN & thalamo-cortical

*BUT heterogeneity & mixed findings
& lack of longitudinal studies & control groups*

Mortaheb et al. Frontiers Neurol. 2021



Long term alterations



Observational longitudinal study

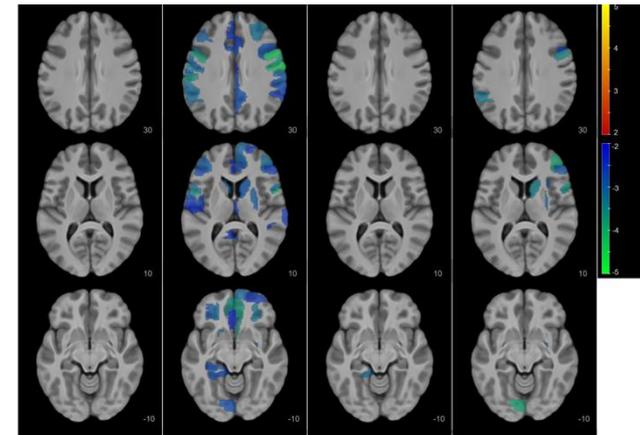
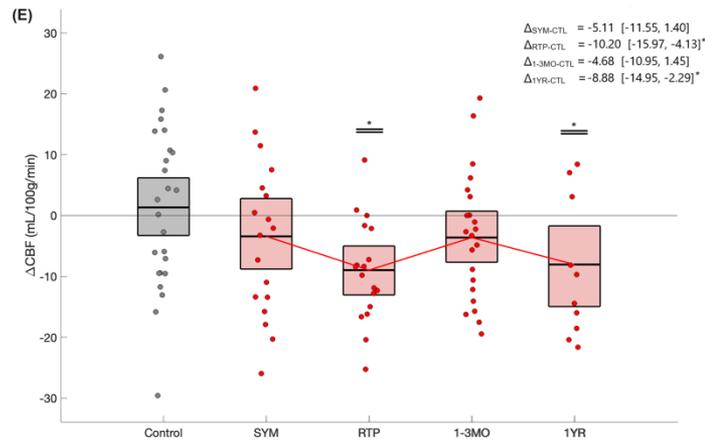
- N= 187 - 25 concussed & 27 injury-free athletes (male & female)
- fMRI (functional connectivity, CBF, DTI)
- Baseline -> symptomatic -> RTP – medical clearance -> 1-3 month -> 1-year



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- N= 187 - 25 concussed & 27 injury-free athletes (male & female)
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Medial temporal regions

**CBF : decreased at symptomatic
persisted beyond medical clearance & 1-year**

Short term & subclinical alterations

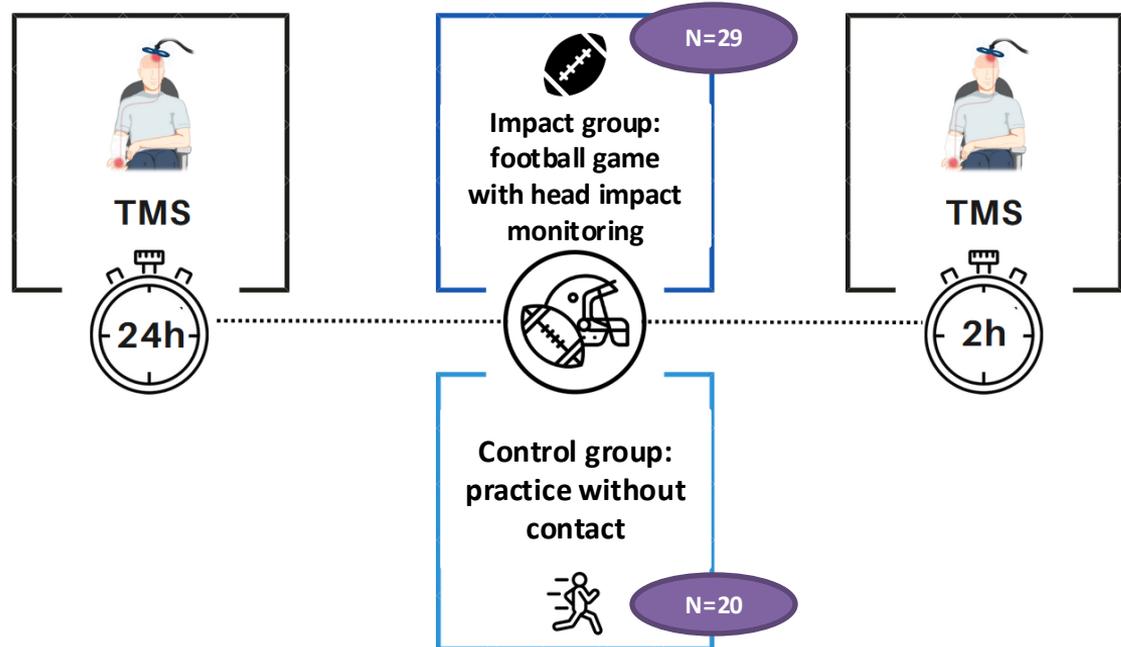


Head impacts

→ Assess intracortical inhibition (**TMS**) post game (impacts) versus post training (without impacts)

Male athletes (n=29)
Varsity football
McGill & UdeM
2021-2022 seasons
Random assignment

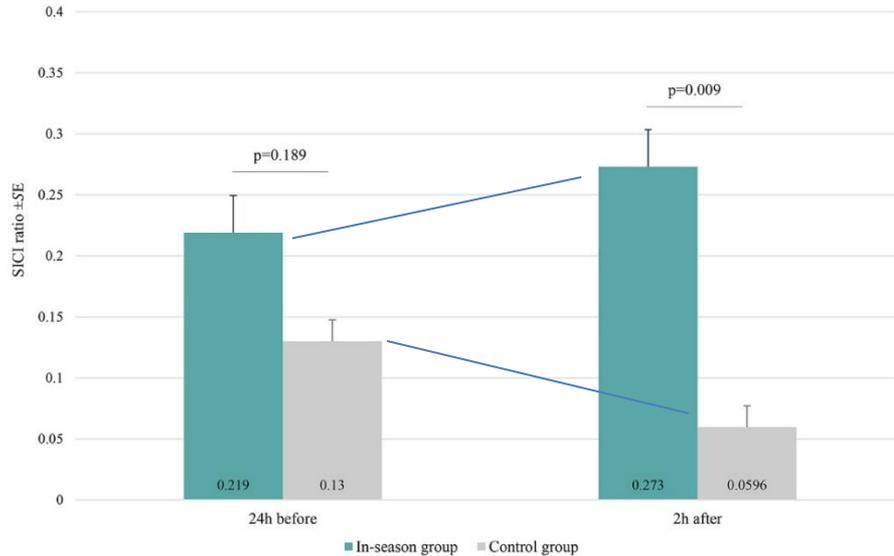
23 yo
185 cm
95 kg



Short term & subclinical alterations



Intracortical inhibition – TMS

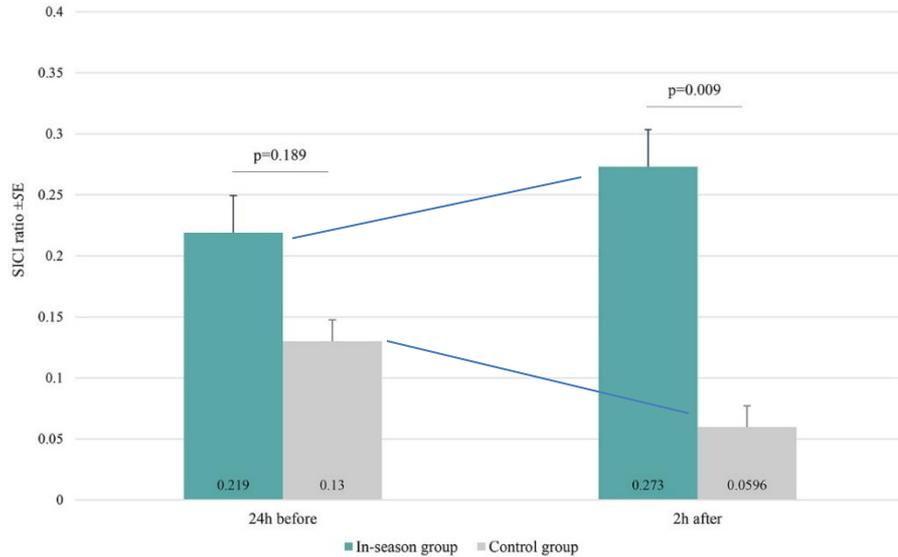


➔ Short term changes in cortical excitability following head impacts during games

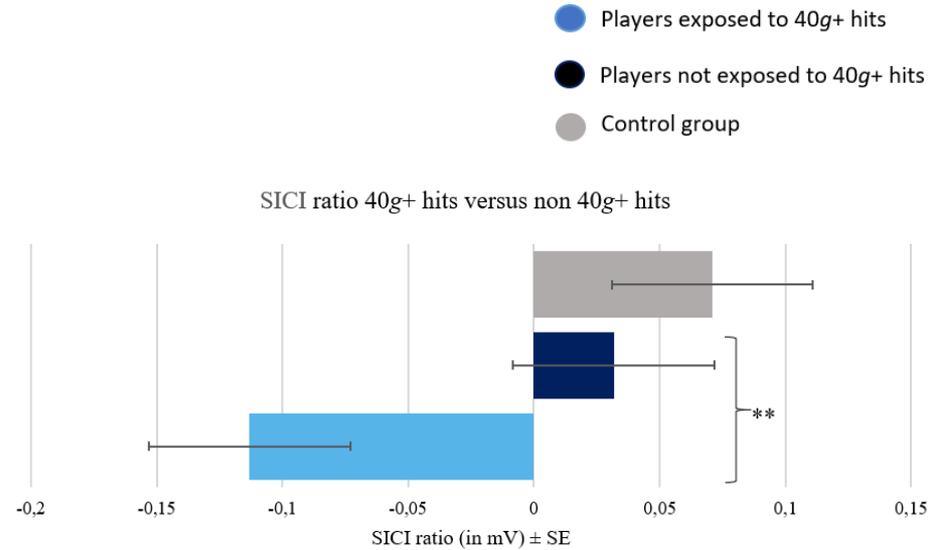
Short term & subclinical alterations



Intracortical inhibition – TMS



TMS & impacts



➔ Short term changes in cortical excitability following head impacts during games

➔ Deleterious intracortical disinhibition with 40+g impacts (neurometabolic cascade)



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In clinical practice?



NeuroRecovery Lab

Concussion care pathway CHU Liège



Neuropsychology & physiotherapy



PM&R & neurology



Clinical & high-density EEG

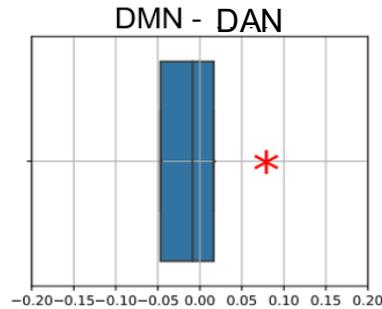
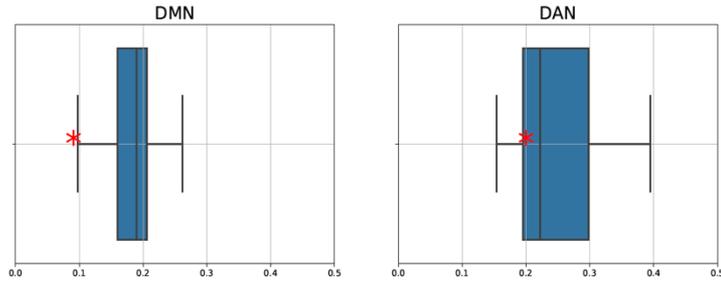


Structural & functional MRI

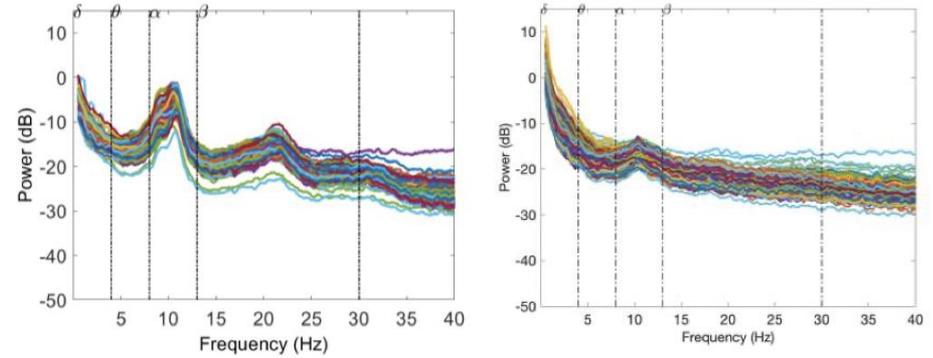
Concussion care pathway



fMRI

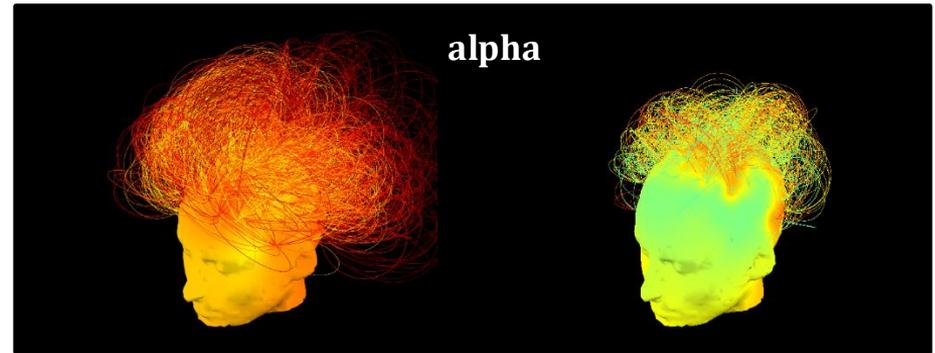


HD-EEG



Control

Patient



Conclusions



Added value to understand 1. pathophysiological mechanisms of concussion
2. course of (subclinical) recovery

Objectification of microstructural & functional alterations

- ➔ Functional connectivity // persistent symptoms
- ➔ Subclinical & long-term alterations beyond clinical clearance

Use in clinical practice?

Large scale longitudinal studies

Need for clinical translation (fMRI ➔ EEG)

Enhanced specificity for patient's level interpretation





Thank you !



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www.giganeurorecovery.uliege.be