IMAGING: What procedures and how they can help?

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Disclaimer

I am an HIV physician

Overview:

1 MR imaging of brain

2 MR spectroscopy

Functional MRI / other modalities / PET



MR Imaging of the brain

Contraindications





MR Imaging of the brain

Contraindications

- Implanted devices
- Metallic foreign bodies
- Unstable patients
- Claustrophobia
- Pregnancy?



MR Imaging of the brain





Open MRI scanner



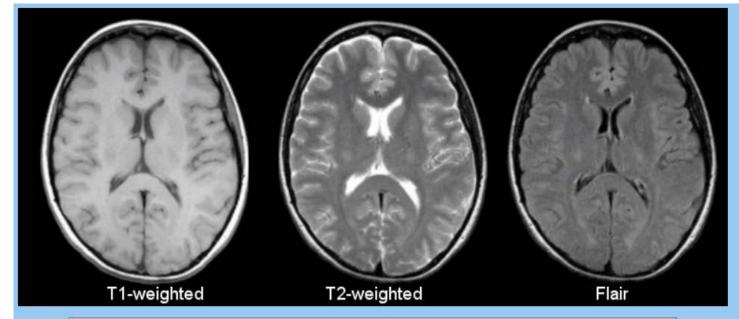
Basic physics

- A powerful, external magnetic field is employed to align the protons that are normally randomly oriented within the water nuclei of the tissue being examined.
- This alignment (or magnetisation) is then disrupted by introduction of an external Radio Frequency (RF) energy.
- The nuclei return to their resting position through various relaxation processes and in so doing emit RF energy.
- After a certain period the emitted signals are measured.
- By varying the sequence of RF pulses applied & collected, different types of images are created.
 - **Repetition Time (TR)** is the amount of time between successive pulse sequences applied to the same slice.
 - **Time to Echo (TE)** is the time between the delivery of the RF pulse and the receipt of the echo signal.

Common sequences

	TR (msec)	TE (msec)	
T1-Weighted (short TR and TE)	500	14	
T2-Weighted (long TR and TE)	4000	90	
Flair (very long TR and TE	9000	114	

Common sequences



Tissue	T1-Weighted	T2-Weighted	Flair
CSF	Dark	Bright	Dark
White Matter	Light	Dark Gray	Dark Gray
Cortex	Gray	Light Gray	Light Gray
Fat (within bone marrow)	Bright	Light	Light
Inflammation (infection, demyelination)	Dark	Bright	Bright



What is it good for?

Differential diagnoses

Very common differential diagnoses in PWH with cognitive disorders:

- Vascular
- Neurodegenerative

<u>Less common differential diagnoses in PWH with cognitive disorders:</u>

- Inflammatory and demyelinating lesions
- AIDS defining OI and tumours
- Other CNS infections and tumours
- Trauma

Some cases

82 year old British man

- HIV positive for 30 years
- On ART with suppressed viral load
- Concentration difficulties over 3 years, progressively worsening
- Cognitive testing confirms impairment predominantly in short term memory
- Several AIDS defining illnesses including PCP in 1990s
- Hypertension and type II DM

Some cases

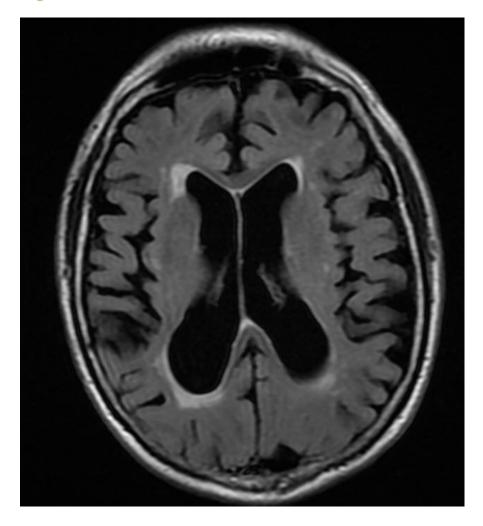
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Differential diagnoses



80 year old man





Case 2

52 year old woman

- HIV positive for 15 years
- Born in India
- On ART with suppressed viral load
- Concentration difficulties over 2 years
- Not had cognitive testing yet
- No ADI
- Hypertension (poorly controlled), type II DM, hyperlipidaemia, raised BMI (over 30)

Differential diagnoses?

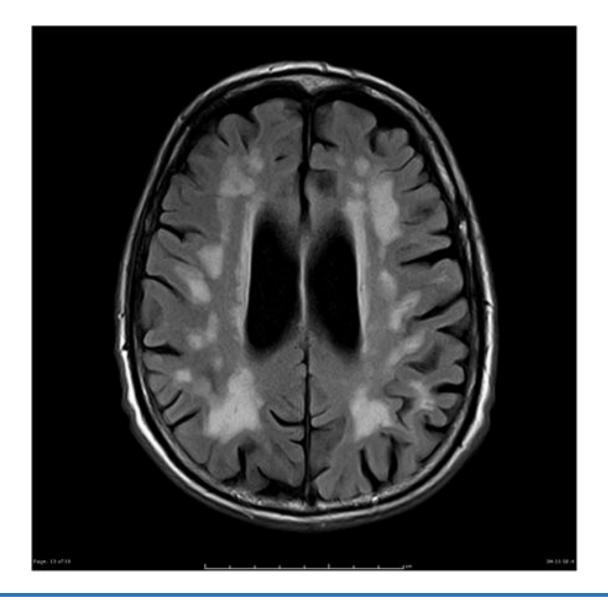


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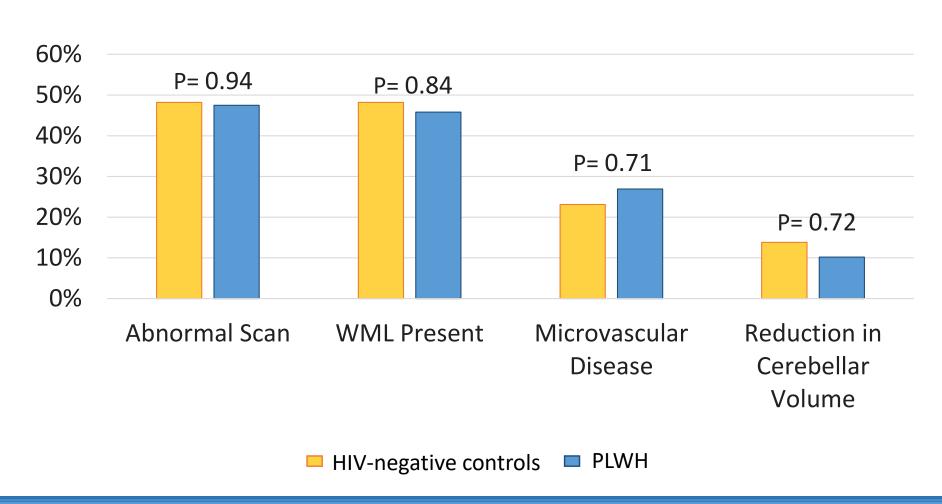
Differential diagnoses?





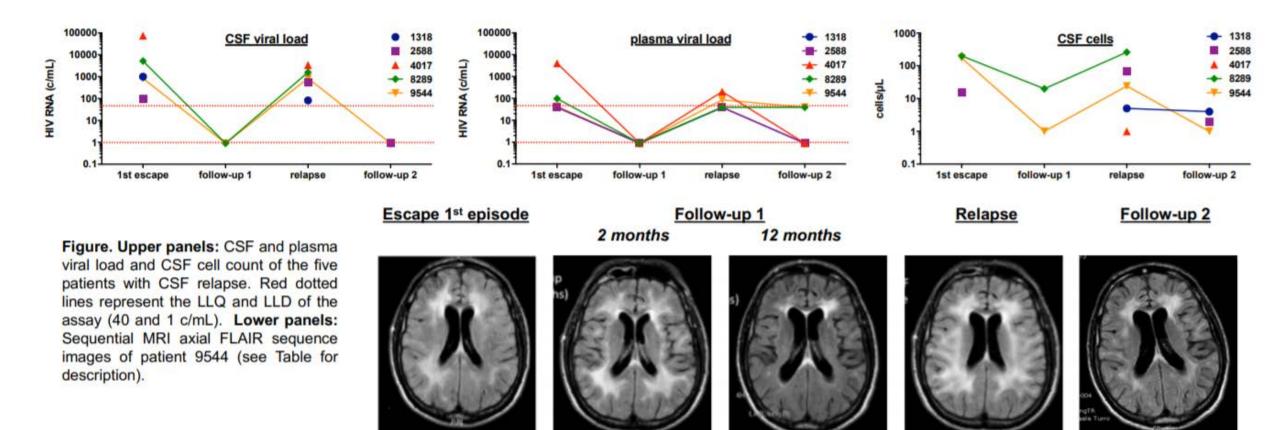
Case 2

MRI Findings by HIV Status





CSF escape





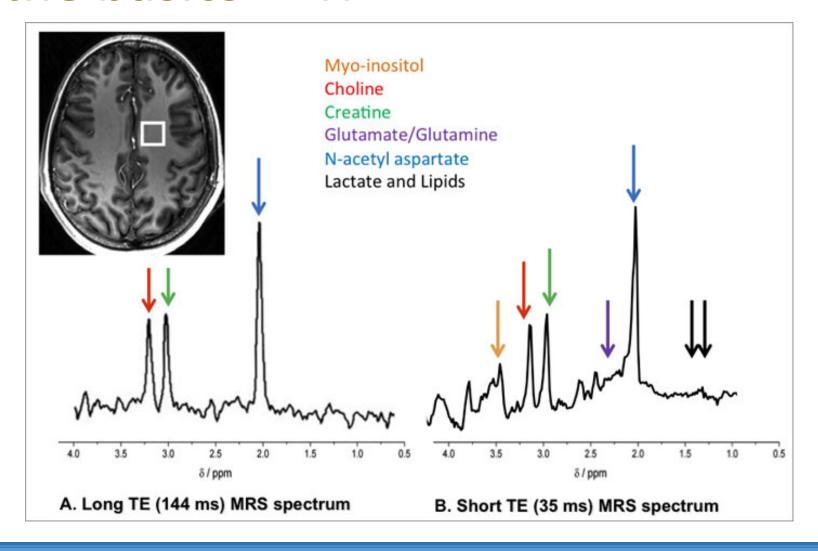
Overview:

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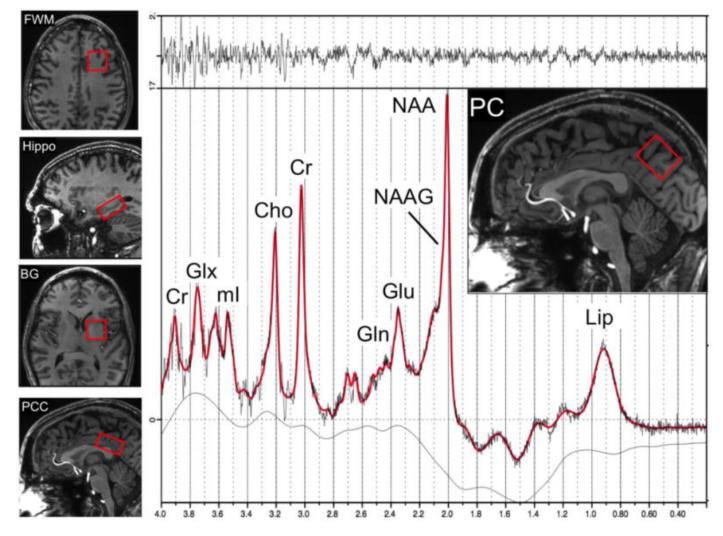
2 MR spectroscopy

Functional MRI / other modalities / PET

MRS the basics -1H

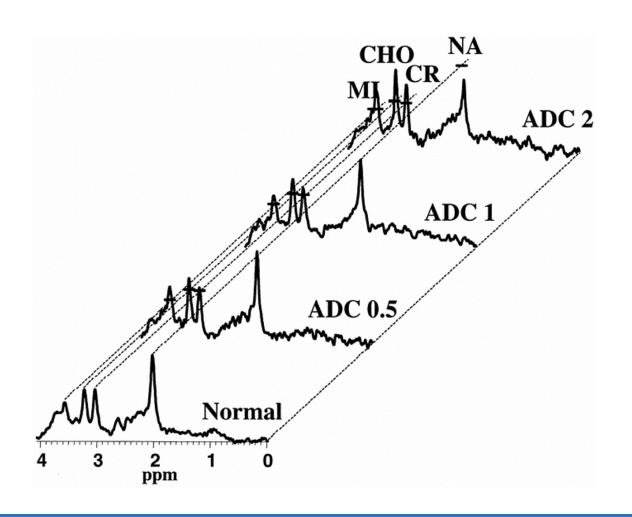


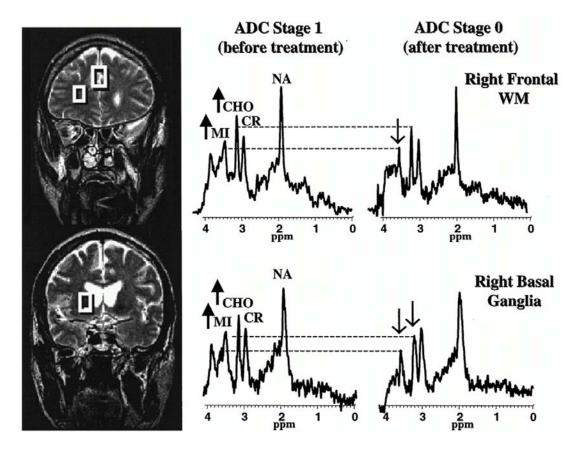
MRS the basics - 7-Tesla





Clinical utilities?





Case 3

42 year old man 2007

- Originally from Eritrea
- Diagnosed with HIV 2007
- Presented with symptomatic HIV
 - Concentration difficulties, weight loss
- MRI scan brain normal
- Cognitive testing severe impairment
- CSF exam unremarkable (CSF RNA 1log lower plasma)
- Commenced TDF/FTC/efavirenz



Case 3

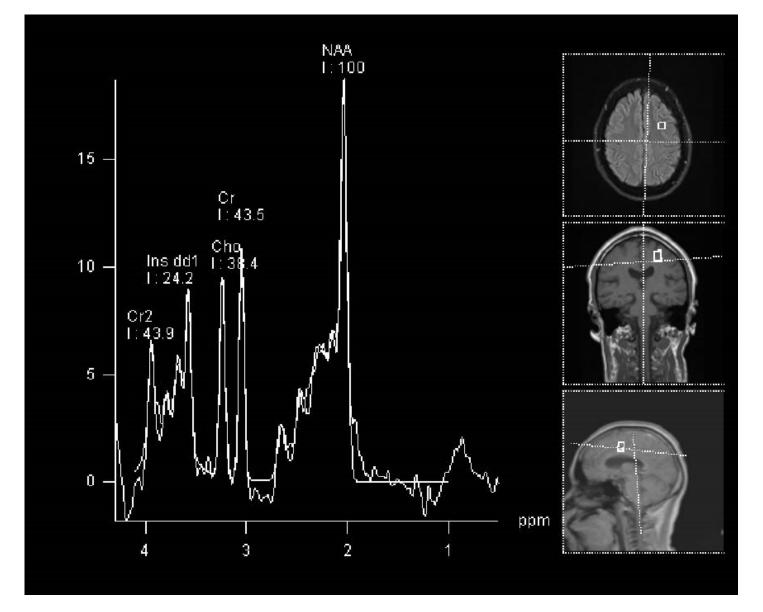
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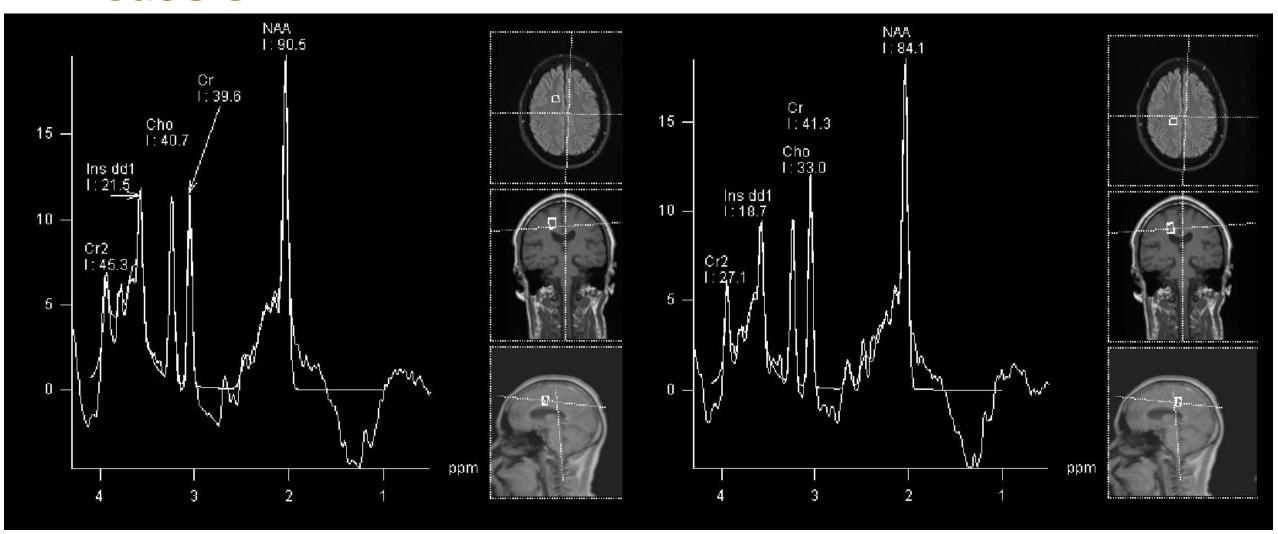
2007 to 2019

- Ongoing concentration complaints
- Attended neurology, psychiatry, psychology
 - Premorbid personality disorder, PTSD, depression
- ALWAYS suppressed in plasma
- Repeat CSF exams undetectable HIV RNA
- Repeat MRI brain normal
- Switched off efavirenz no change
- Cognitive testing deteriorating

Case 3



Case 3



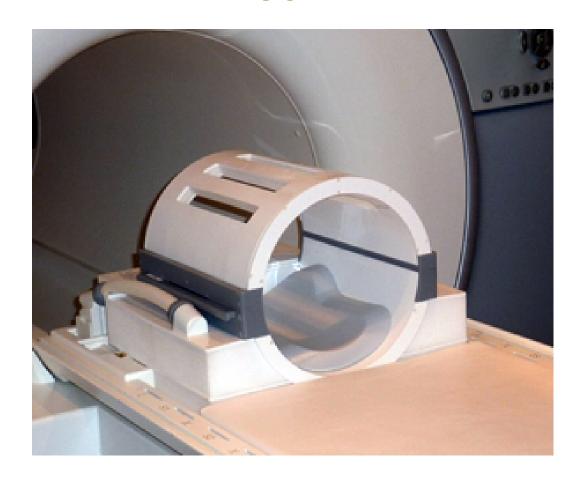


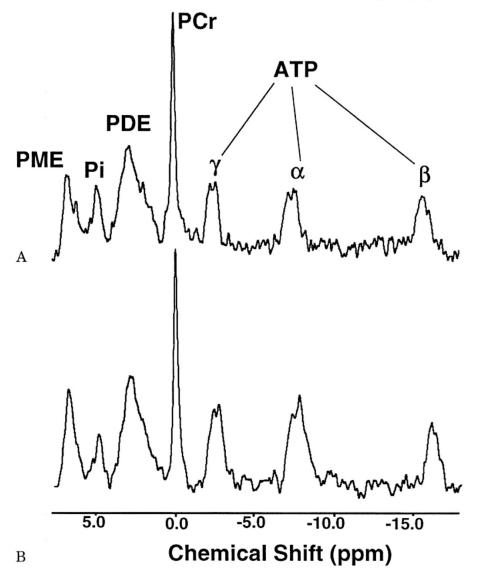
Clinical use of brain ¹H-MRS

- Is available at ICHNT
- In cases where we suspect HIV may still be contributing to cognitive disorders and other biomarkers have not proven useful
- We infrequently use 'intensification' ART strategies such as maraviroc –
 if we do take this approach longitudinal MRS can be helpful in
 monitoring progress



Other types of MRS - ^{31}P





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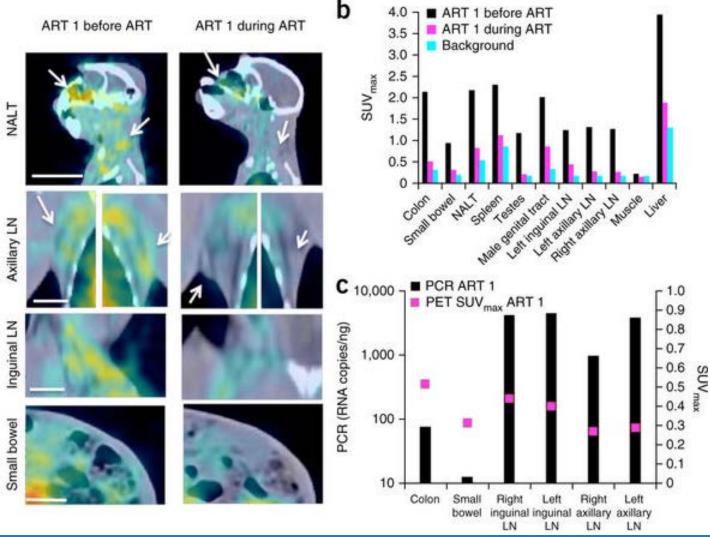
Functional MRI / other modalities / PET

Quantification of HIV - Gp120 Ab



Whole-body immunoPET reveals active SIV dynamics in viremic and antiretroviral therapy—treated macaques

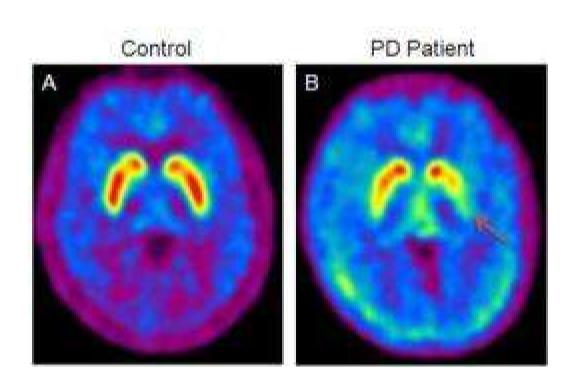
Philip J Santangelo, Kenneth A Rogers, Chiara Zurla, Emmeline L Blanchard, Sanjeev Gumber, Karen Strait, Fawn Connor-Stroud, David M Schuster, Praveen K Amancha, Jung Joo Hong, Siddappa N Byrareddy, James A Hoxie, Brani Vidakovic, Aftab A Ansari, Eric Hunter & Francois Villinger



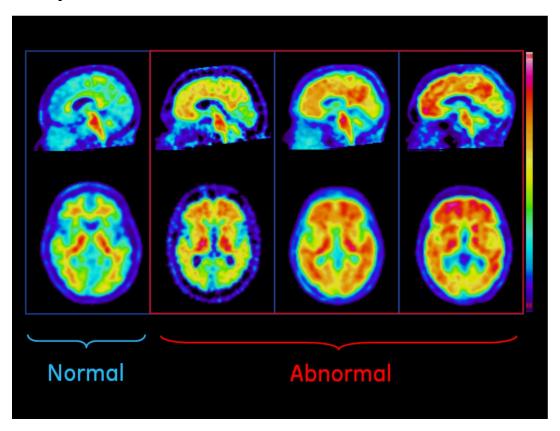
PET in clinical use

Presynaptic dopaminergic imaging:

DaTscan

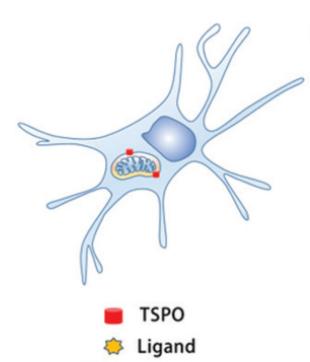


Amyloid PET



TSPO PET

RESTING MICROGLIA



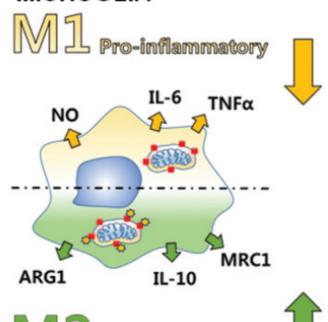
Mitochondrion

NEUROINFLAMMATION

TSPO Up-regulation/ Ligand binding

NEUROPROTECTION

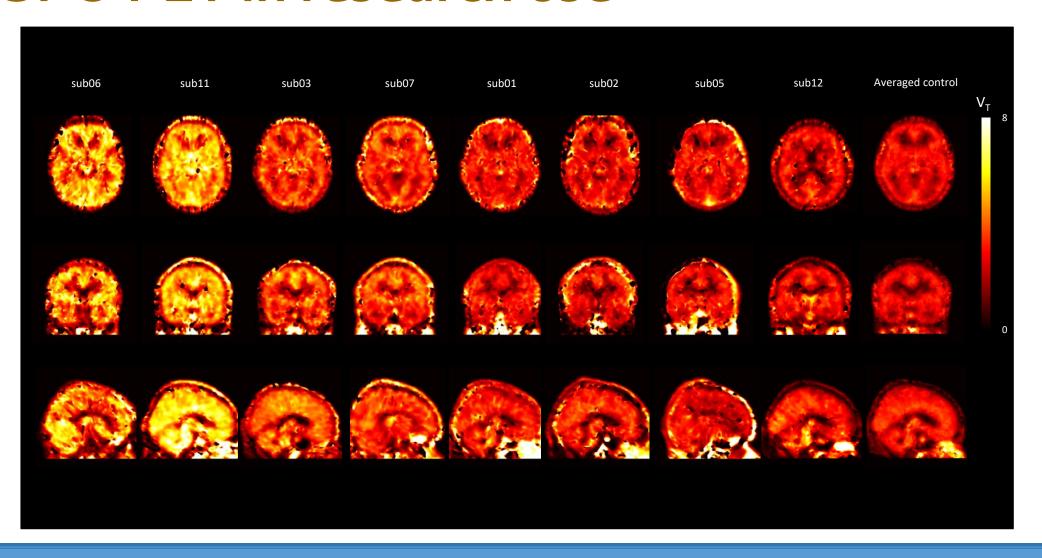
REACTIVE MICROGLIA



M2 Anti-inflammatory



TSPO PET in research use





Summary: MRI imaging in PWH on ART

MR imaging of brain

Important in defining differential diagnoses

Common DDx:

- Vascular changes
- Neurodegenerative diseases
- HIV related changes

Important routine investigation in PWH with cognitive disorders

Often abnormal in PWH but not necessarily HIV related

MR spectroscopy of brain

May be useful modality to detect ongoing inflammation

May be a useful modality to detect treatment responses

Reserve for challenging management cases