

# Mind:

Where does HIV associated cognitive impairment fit in 2025?

## Chair:

Prof Jaime Vera

# Speakers:

Memory Sachikonye Prof Alan Winston ASK A QUESTION HERE





# Brain health: Are we still sane?

Memory Sachikonye
UK Community Advisory Board (UK-CAB)

memory.sachikonye@ukcab.net; X: memsachi

Ageing & HIV: A multidisciplinary meeting on HIV and Ageing 12 September 2025



# **Declarations**

None

Ageing & HIV: A multidisciplinary meeting on HIV and Ageing 12 September 2025





# Our brain health will cover

- 1. Our experiences
- 2. Our brain health fears
- 3. Impact on our daily lives
- 4. How we 'try' to cope
- 5. Interventions?

**RM1** Are these your notes? Make sure they are not visible on the slides - move them to the notes section

Rebecca Mbewe, 2025-04-21T11:32:29.083



# Hormonal changes and the brain

- Oestrogen supports memory, mood and cognitive functions
- During menopause: decline in oestrogen → can affect neurotransmitters (e.g., serotonin, dopamine)

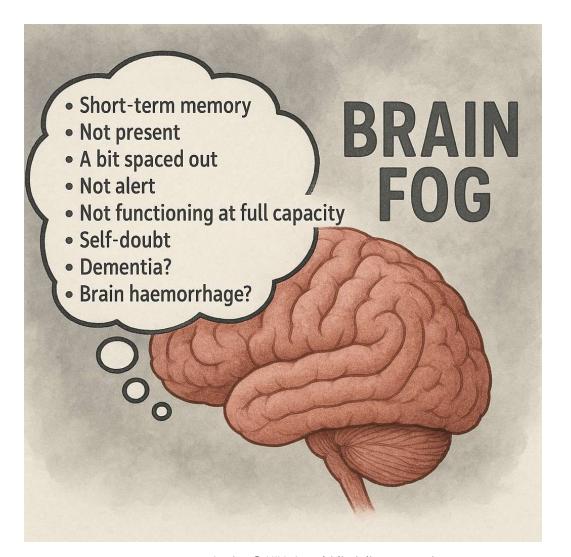
# Common changes:

- Brain fog
- Forgetfulness
- Mood swings
- Sleep disruption

Peri and menopausal women's experiences



Ageing & HIV: A multidisciplinary meeting on HIV and Ageing 12 September 2025











# Impact on our daily lives

# Memory lapses

- Where did I put that thing?
- I definitely know that face, but what's their name?

# Difficulty concentrating

- Can I still get my work done?
- Doing too many things that are left unfinished.





# **Anxiety or irritability**

- Am I losing myself?
- Double checking everything you do.
- Is it just stress, or something worse?
- What if it's something serious?

# Insomnia or poor-quality sleep

• affects how well we think, react, work, learn, and get along with others



# **Interventions**

- Brain scan that came back normal, but I am still worried 'H'
- Had a memory test that did show short-term memory issues 'J'
- Electronic and paper calendars
- Meditation
- "I have not had any interventions with doctors, but I try to double-check things when I can, utilize my calendar.



# **Coping strategies - selfcare**

### Mind and mood:

- . **Exercise:** Boosts endorphins, supports cognition
- . **Mindfulness & Meditation:** Helps reduce stress and improve focus
- . Sleep Hygiene: Regular schedule, limit screens, cool/dark room

### Diet:

. Omega-3s, leafy greens, berries, and hydration

### **Routine:**

- Use lists, alarms, and planners to stay organized
- Break tasks into small chunks



# **Coping strategies**

'I try to reassure myself,

meditate, use movement,

speaking to loved ones about my fears,

doing things that bring me joy,

so I'm not constantly worrying' – 'M'





- Menopause affects the brain—we are not imagining it
- . Daily habits can make a big difference



• Find support—you're not alone: my support network



# It's not just women, men too:

https://tinyurl.com/2hcx4uar



# to my peers at Positively UK who shared their own experiences

# IMPERIAL



# Where does HIV associated cognitive impairment fit in 2025?

Alan Winston August 2025

# Overview

1	Defining brain health in persons with HIV
2	Some HIV considerations
3	Some non-HIV considerations
4	Management

1

Defining brain health in persons with HIV

Defining cognitive disorders



Before ART AIDSdementia complex

ADC STAGE	CHARACTERISTICS	
Stage 0:(Normal)	Normal mental and motor function.	
Stage 0.5: (Equivocal/subclinic al)	Either minimal or equivocal symptoms of cognitive or motor dysfunction bolinic characteristic of ADC, or mild signs (snout response, slowed extremity movements), but without impairment of work or capacity to perform activities of daily living (ADL). Gait and strength are normal.	
Stage 1: (Mild)	Unequivocal evidence (symptoms, signs, neuropsychological test performance) of functional intellectual or motor impairment characteristic of ADC, but able to perform all but he more demanding aspects of work or ADL. Can walk without assistance.	
Stage 2: (Moderate)	Cannot work or maintain the more demanding aspects of daily life, but able to perform basic activities of self care. Ambulatory, but may require a single prop.	
Stage 3: (Severe)  Major intellectual incapacity (cannot follow news or personal e cannot sustain complex conversation, considerable slowing of or motor disability (cannot walk unassisted, requiring walker or support, usually with slowing and clumsiness of arms as well).		
Stage 4: (End stage)	Nearly vegetative. Intellectual and social comprehension and responses are at a rudimentary level. Nearly or absolutely mute. Paraparetic or paraplegic with double incontinence.	
	HIV InSite Knowledge Base Chapter Published June 1998	

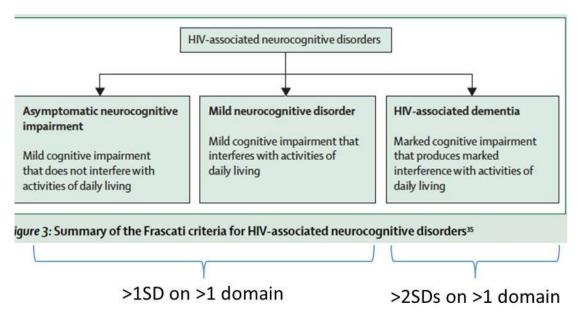
# HIV-associated neurocognitive disorders (HAND)

- 2007 criteria NIMH/NINDS working group
  - Defined at meeting in Frascati, Italy.
- Remains most used approach
  - "Currently people with HIV live longer with nearly undetectable viral loads, however neurocognitive impairment persists in a considerable proportion of the population. It is estimated that about 40-50% of people with HIV develop HAND".

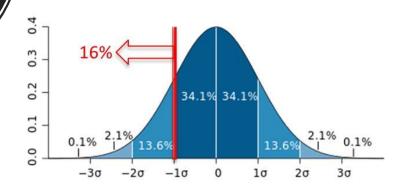


Antinori et al. Neurology. 2007

HAND has a high false classification rate

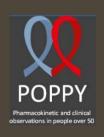


Compared with controls matched for age, sex and years of education Exclusion of confounders

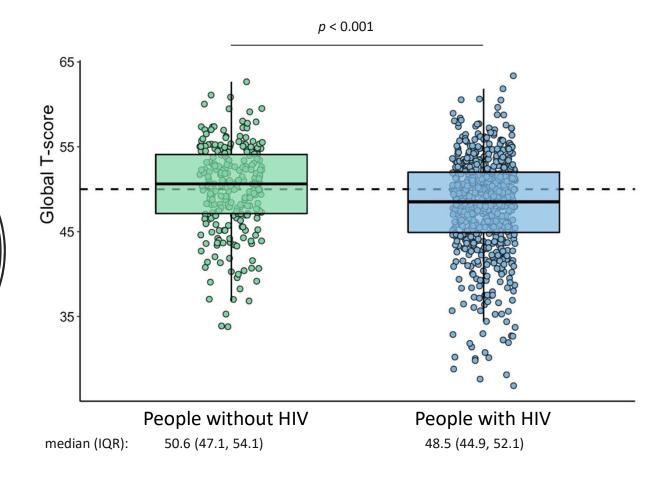


• False positive rate ~26%

Gisslen M et al. BMC Inf Dis. 2011



Cognitive function in the POPPY cohort



### Published: 09 July 2021

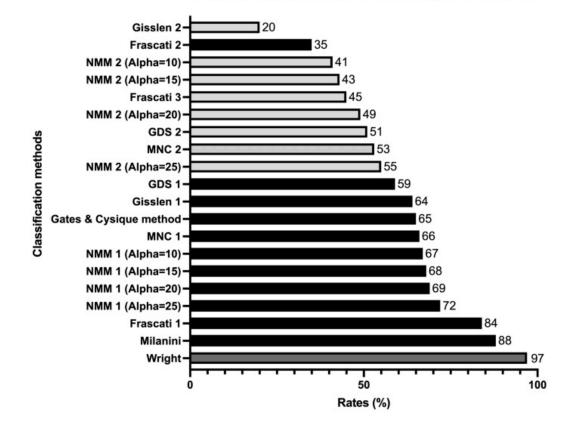
Rates of cognitive impairment in a South African cohort of people with HIV: variation by definitional criteria and lack of association with neuroimaging biomarkers

Anna J. Dreyer ☑, Sam Nightingale, Jodi M. Heaps-Woodruff, Michelle Henry, Hetta Gouse, Robert H. Paul, Kevin G. F. Thomas & John A. Joska

Journal of NeuroVirology 27, 579-594 (2021) Cite this article

Cognitive impairment in Cape Town

### Cognitive impairment rates according to 20 methods







# UK Civil Aviation Authority announces landmark changes for pilots and air traffic controllers living with HIV

20 June 2022 | Pilots, Medical



The UK Civil Aviation Authority, with the support of the Terrence Higgins Trust, the National AIDS Trust and the British HIV Association, has today issued a landmark package of measures for pilots living with HIV to support them flying and continuing with their careers.

The regulator has published important new guidance on how pilots living with HIV will be medically assessed to obtain and continue to hold their UK licences. The regulator's new guidance confirms that with a timely diagnosis of HIV and antiretroviral therapy, there is a much lower risk of a pilot suffering conditions that could impair their ability to safely fly aircraft.

This is a significant milestone in the global aeromedical assessment of HIV. It means there is a now a much clearer way for pilots living with HIV to obtain unrestricted (Class 1) medical certification in the UK. For some pilots living with HIV, it will also reduce the need for additional cognitive testing. The guidance also covers Class 3 medical certificate applicants who work, or wish to work, as air traffic control officers.



### CLINICAL SCIENCE

# Impact of HIV on Cognitive Performance in Professional Drivers

Gouse, Hetta PhD<sup>a</sup>; Masson, Catherine J. MA<sup>a</sup>; Henry, Michelle PhD<sup>b</sup>; Dreyer, Anna MA<sup>a</sup>; Robbins, Reuben N. PhD<sup>c</sup>; Kew, Greg MBChB, DOH, ACPHM (Occ Med)<sup>d</sup>; Joska, John A. MBChB, MMed (Psych), PhD, FCPsych (SA)<sup>a</sup>; London, Leslie MB ChB, MD, BSc Hons (Epid), DOH, M Med (PH)<sup>d</sup>; Marcotte, Thomas D. PhD<sup>e</sup>; Thomas, Kevin G. F. PhD<sup>f</sup>

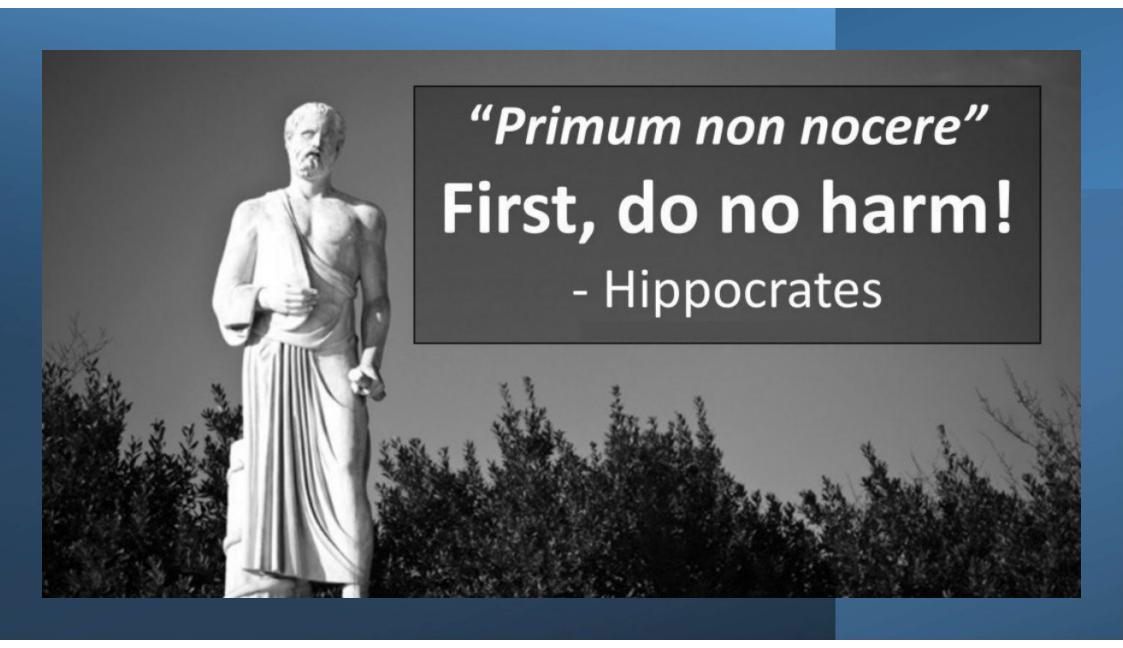
### Author Information ⊗

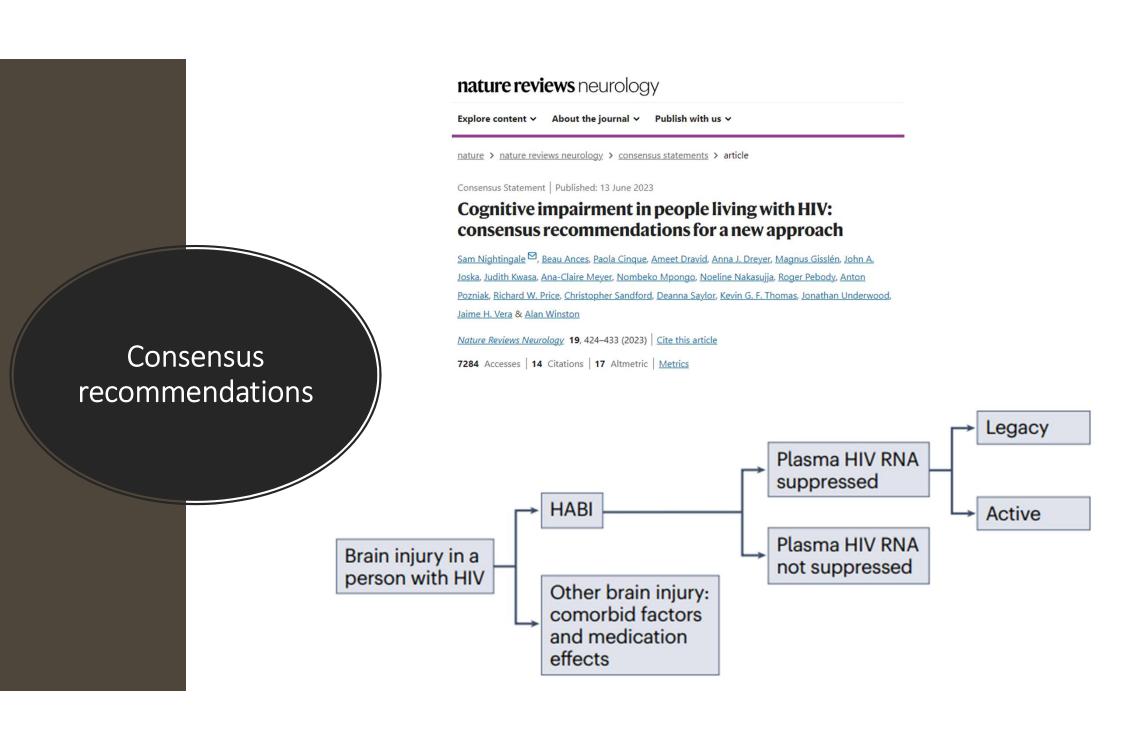
JAIDS Journal of Acquired Immune Deficiency Syndromes: April 15, 2022 - Volume 89 - Issue 5 - p 527-536

doi: 10.1097/QAI.0000000000002899

# **Conclusion:**

Drivers with HIV may be at risk of poorer long-term health and employment outcomes. Programs that monitor and support their longterm cognitive health are needed.





2

Some HIV considerations

CLINICAL SCIENCE

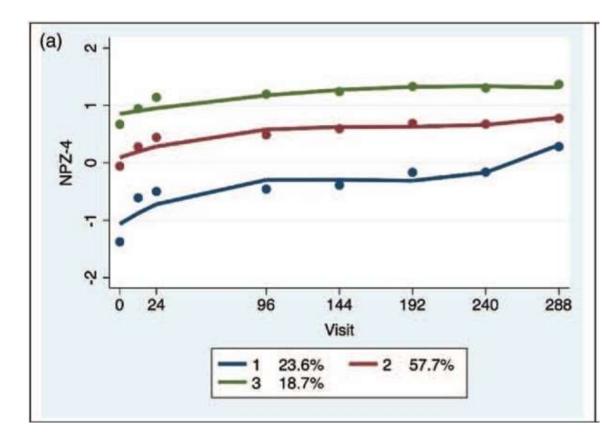
# Cognitive trajectories after treatment in acute HIV infection

Chan, Phillip<sup>a</sup>; Kerr, Stephen J.<sup>b,c</sup>; Kroon, Eugène<sup>a</sup>; Colby, Donn<sup>a,d</sup>; Sacdalan, Carlo<sup>a</sup>; Hellmuth, Joanna<sup>e</sup>; Reiss, Peter<sup>f</sup>; Vasan, Sandhya<sup>d,g</sup>; Ananworanich, Jintanat<sup>a,f</sup>; Valcour, Victor<sup>e</sup>; Spudich, Serena<sup>h</sup>; Paul, Robert<sup>i</sup>

Author Information⊙

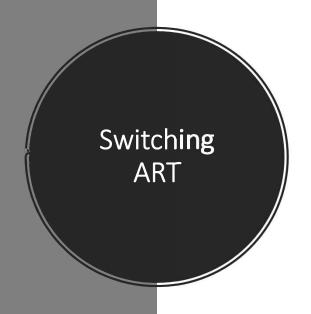
AIDS 35(6):p 883-888, May 1, 2021. | DOI: 10.1097/QAD.000000000002831

Implications of early ART



Cognitive performance, neuropsychiatric symptoms, and cerebrospinal fluid viral control following programmatic switch from efavirenz-based to dolutegravir-based antiretroviral therapy in South Africa (CONNECT): a prospective cohort study

Sam Nightingale, Anna J Dreyer, Kevin G F Thomas, Gert van Zyl, Eric Decloedt, Petrus J W Naude, Catherine Orrell, Phumla Sinxadi, Alan Winston, Saye Khoo, John A Joska



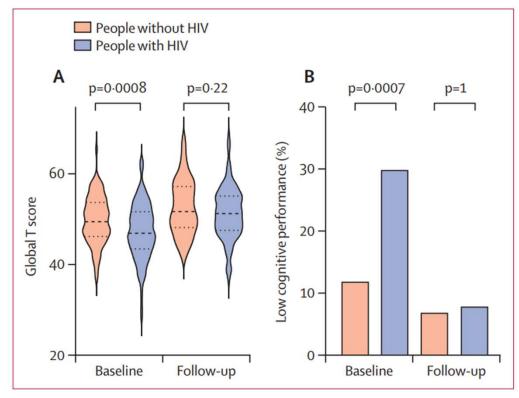


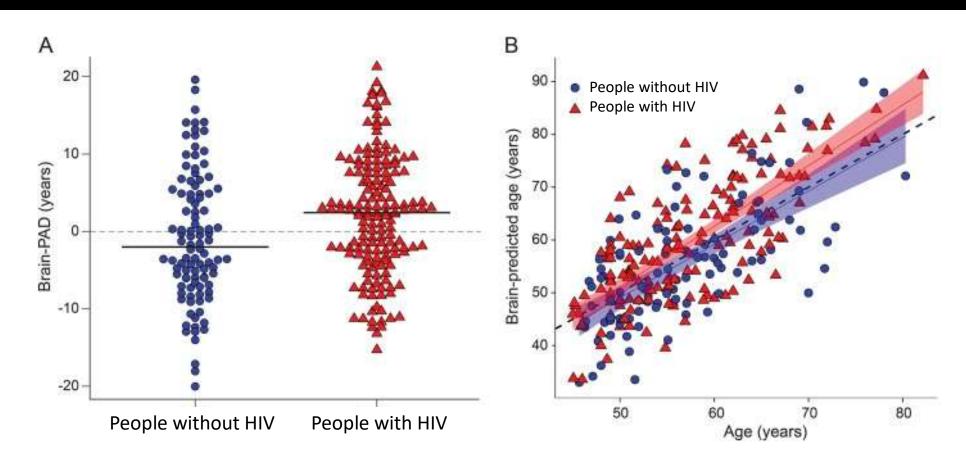
Figure 2: Cognitive performance at baseline (on efavirenz) and follow-up (on dolutegravir)

(A) Global T score. (B) Global deficit score.

Lancet HIV 2024; 11: e680-89



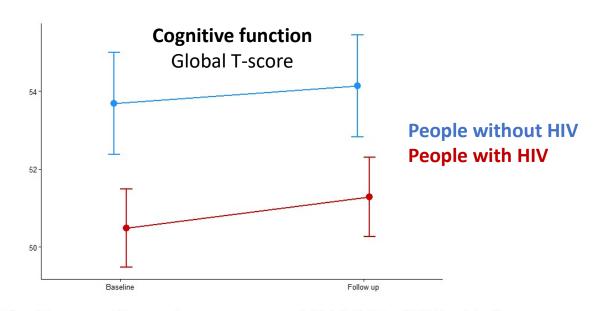
# Brain predicted age



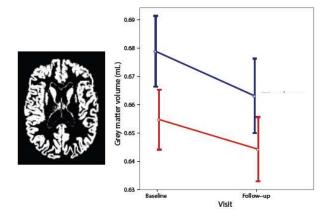
Neurology 2017 Apr 4;88(14):1349-1357



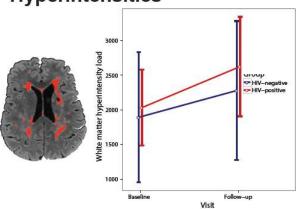
Longitudinal trajectories: COBRA



A) T1 - Grey matter volume



B) FLAIR - White Matter Hyperintensities



Clin Infect Dis. 2018 Jun 1;66(12):1899-1909



# 68-year-old woman

## **HIV** history

- HIV diagnosed 1997, commenced ART 2002
- Several ART combinations and drug-resistant virus
- Currently undetectable

### Other medical history, social, lifestyle

- BMI 34
- HbA1c in pre-diabetic range
- BP normal
- On a statin

## **Symptoms**

- 6-week history of headaches
- Finding it difficult to multitask

# Case 1 History

# 68-year-old woman

# **Neurological examination**

- Normal

# **Brain imaging**

- Some periventricular WM changes, no previous scan

### **CSF** examination

	Result
CSF Protein g/L	0.85
CSF WBC cells/μL	40
CSF Viral PCRs	negative
CSF HIV RNA copies/mL	1,200

#### Is this CSF HIV RNA escape?

#### Virological definition of CSF HIV RNA escape:

- either CSF HIV RNA detectable and plasma HIV RNA undetectable
- or both CSF HIV RNA and plasma HIV RNA detectable, with CSF HIV RNA higher than plasma HIV RNA





AIDS 2019: PMID 31790376

## Prevalence of CSF HIV RNA escape



SHORT COMMUNICATION 🙃 Open Access 🤄 📵 🕞 😒



#### Cerebrospinal fluid virology in people with HIV

Merle Henderson 🔀 Nuala Pepper, Manraj Bawa, David Muir, Alex Everitt, Nicola E. Mackie, Alan Winston

First published: 09 March 2023 | https://doi.org/10.1111/hiv.13471

Characteristic	Total cohort, n = 114	CSF escape, n = 19	No CSF escape, n = 95
Demographics			
Age (years), mean (SD)	48.8 (13.0)	51.6 (13.1)	48.3 (13.0)
Male sex, n (%)	81 (71)	10 (52)	71 (75)
Ethnicity, n (%)			
White	49 (43)	7 (37)	42 (44)
Black	30 (27)	7 (37)	23 (24)
Other	35 (31)	5 (26)	30 (32)

#### Symptoms of CSF HIV RNA escape

CLINICAL SCIENCE: CONCISE COMMUNICATION

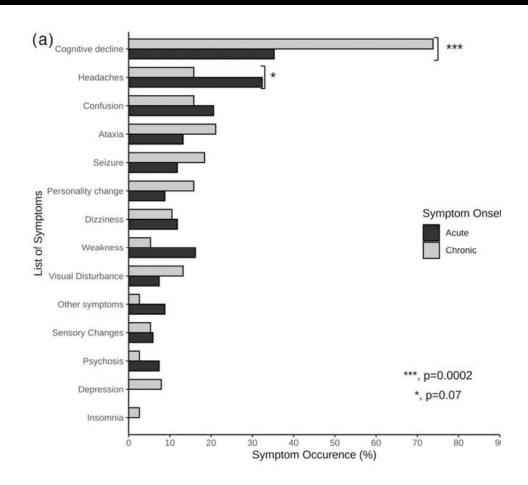
## The symptomatology of cerebrospinal fluid HIV RNA escape: a large case-series

Chan, Terrence Y.-H.a; De Zan, Valentinab; Gregg, Alistairc; Alagaratnam, Jasmini<sup>d,e</sup>; Gerevini, Simonettae; Antinori, Andreaf; D'Arminio Monforte, Antonellag; Saracino, Annalisah; Trunfio, Mattiai; Everitt, Alexi; Rackstraw, Simon<sup>a,k,l</sup>; Marta, Monica<sup>k,l</sup>; Calcagno, Andreaf; Cinque, Paolab; Winston, Alan<sup>c,d</sup>

#### Author Information (>)

AIDS 35(14):p 2341-2346, November 15, 2021. | DOI: 10.1097/QAD.000000000002992

Table 1 - Subject demographic and laboratory characterist	tics at the time of diagnosis ( $n = 106$ ).
Described the second of the	×
Baseline characteristics	
Age (years)	51 (47–56)
Male, n (%)	65 (61.3)
Time since first positive HIV test (months)	144 (75.5-212.5)



## Risk factors for CSF HIV RNA escape

- 1. Previous symptomatic CSF HIV RNA escape
- 2. Difficulty with ART compliance
- 3. Antiretroviral drug resistance
- 4. Use of boosted PIs (questionable)

## Clinical scenario

	Clinical Scenario	Biology	Presentation	CSF HIV RNA	CSF WCC
1	Asymptomatic	Equivalent to plasma blips	Incidental finding	<200 copies/mL	Normal
2	Neurologically symptomatic	Virological failure in CNS compartment	New or progressive neurological symptoms		Can be elevated
3	Secondary	Viral replication related to another infection	Presence of another infection		Often elevated

Curr HIV/AIDS Rep (2015) 12: 280

## Clinical scenario

	Clinical Scenario	Biology	Presentation	CSF HIV RNA	CSF WCC
1	Asymptomatic	Equivalent to plasma blips	Incidental finding	<200 copies/mL	Normal
2	Neurologically symptomatic	Virological failure in CNS compartment	New or progressive neurological symptoms		Can be elevated
3	Secondary	Viral replication related to another infection	Presence of another infection		Often elevated
4	Lag effect	CSF HIV RNA taking longer to suppress compared to plasma			Often normal

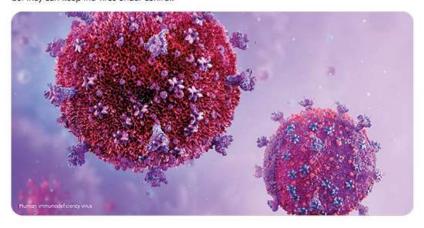
## Management CSF HIV RNA escape



#### WHAT YOU SHOULD KNOW ABOUT GENOTYPE TESTING FOR HIV DRUG RESISTANCE

Genotypic drug resistance testing can help make sure your HIV treatment is working effectively. If drug resistance is found, a new treatment can be used instead.

The human immunodeficiency virus (HIV) is the virus that causes HIV infection. The treatment for HIV is antiretroviral therapy (ART). ART medications stop the virus from multiplying. They cannot cure HIV but they can keep the virus under control.





3

Some non-HIV considerations



#### 82-year-old man

#### **HIV** and medical history

- Living with HIV for 30 years
- On ART with suppressed viral load
- Several AIDS defining illnesses including PCP in 1990s
- Hypertension and type II DM, both on optimal treatment

#### Social and lifestyle history

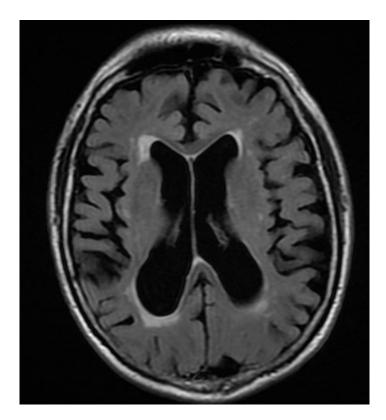
- Lives alone and independent
- Mental health is good

#### **Symptoms**

- Concentration difficulties over 3 years, progressively worsening
- Cognitive testing confirms impairment predominantly in short term memory

## 82-year-old man









Neurol Clin Pract. 2021 Oct; 11(5): e627-e633.

doi: 10.1212/CPJ.0000000000001060

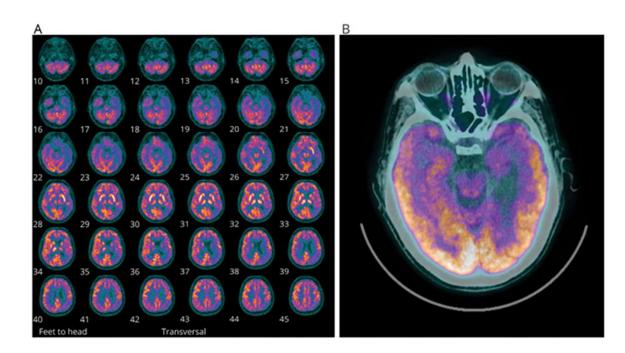
PMCID: PMC8610525

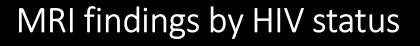
PMID: 34840876

#### Alzheimer Dementia in People Living With HIV

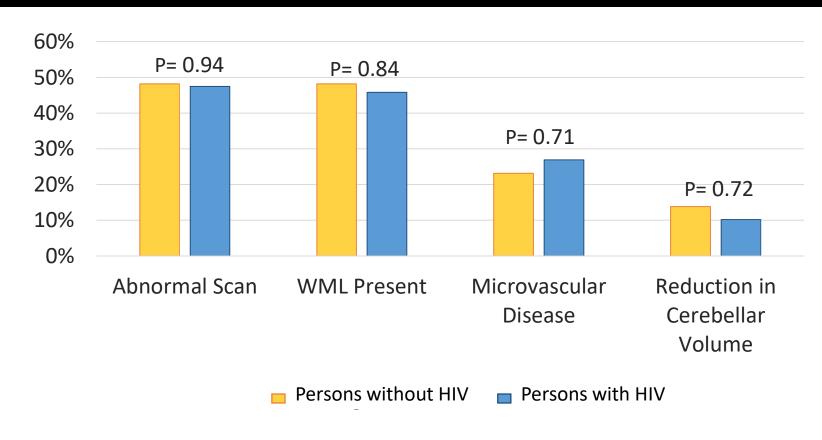
Andrea Calcagno, MD, MD, Luigi Celani, MD, Mattia Trunfio, MD, Giancarlo Orofino, MD, Daniele Imperiale, MD, Cristiana Atzori, BSc, Vincenzo Arena, MD, Gabriella d'Ettorre, MD, PhD, Giovanni Guaraldi, MD, Magnus Gisslen, MD, PhD, and Giovanni Di Perri, MD, PhD













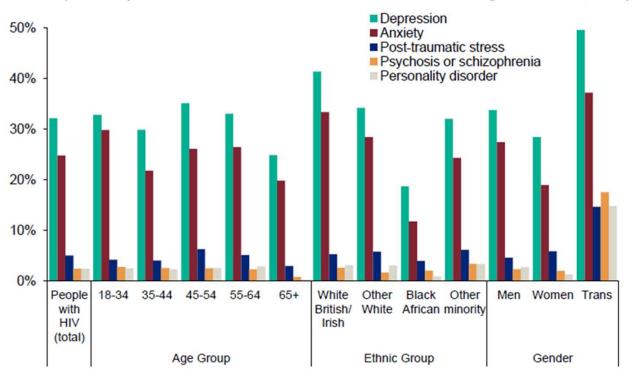
## Clusters of comorbidities

CVDs	STDs	Mental health	Cancers	Metabolic	Chest/other infections
Angina, CABG, Peripheral vascular disease, MI, Heart failure, Hypertension, End-stage renal disease	Gonorrhoea, Syphilis, LGV, Chlamydia, Hepatitis C	Depression, Anxiety, Panic attacks	Haematological cancer, Skin cancer, Solid organ cancer	Dyslipidaemia, Lipodystrophy, Hypertension	Pneumonia, Dizziness, Vertigo, CMV, Asthma, Bronchitis, COPD, Chest infection

AIDS. 2019 Oct 1;33(12):1871-1880.

#### Mental health

Self-reported prevalence of mental health conditions diagnosed in 4,422 people with HIV by age, ethnicity and gender





Positive Voices 2022: survey report - GOV.UK

Home > AIDS and Behavior > Article

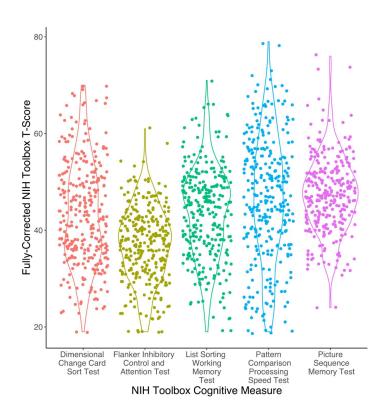
# HIV Stigma is Associated with Two-Year Decline in Cognitive Performance Among People with HIV

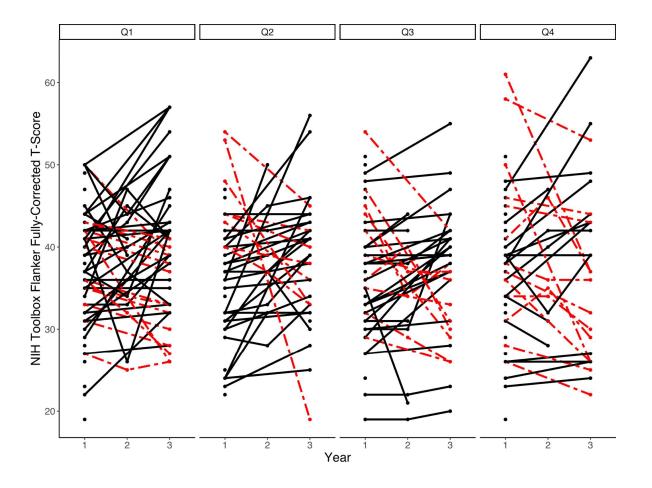
Original Paper | Published: 14 October 2024

Volume 29, pages 90–100, (2025) Cite this article

Download PDF ±

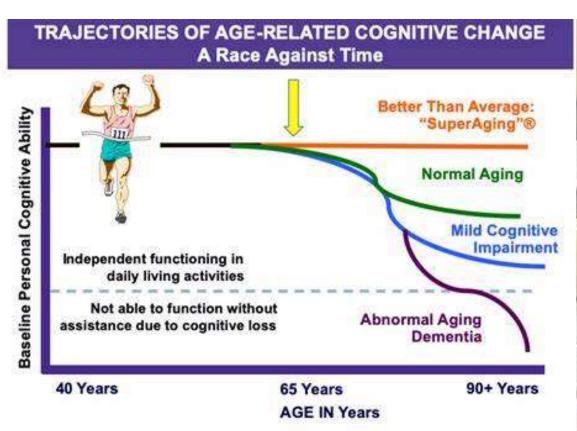
Access provided by JISC Springer Compact (publishing + reading)





AIDS Behav 2025 Jan;29(1):90-100

## Other considerations





# Anticholinergics

Anticholinergic medication use in elderly people living with HIV and self-reported neurocognitive impairment: a prospective cohort study

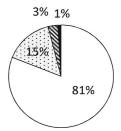
Bernadette Jakeman, Alexandra Scherrer, Manuel Battegay, Huldrych F. Gunthard, Anna Hachfeld, Alexandra Calmy, Patrick Schmid, Enos Bernasconi, Matthias Cavassini, Catia Marzolini 

on behalf of the Swiss HIV Cohort Study

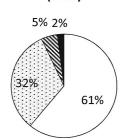
Author Notes

*Journal of Antimicrobial Chemotherapy*, Volume 77, Issue 2, February 2022, Pages 492–499, https://doi.org/10.1093/jac/dkab386

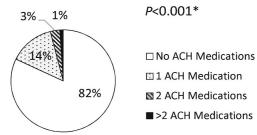
#### All Patients (n=1015)



#### Patients with SRNI (n=87)



#### Patients without SRNI (n=928)





ORIGINAL ARTICLE

#### Anticholinergic medications associated with falls and frailty in people with HIV

Jessica Doctor Alan Winston, Jaime H. Vera, Frank A. Post, Marta Boffito, Patrick W. G. Mallon, Jane Anderson, Christina Prechtl, Ian Williams, Margaret Johnson, Emmanouil Bagkeris, Memory Sachikonye, Caroline A. Sabin, for the POPPY Study ... See fewer authors

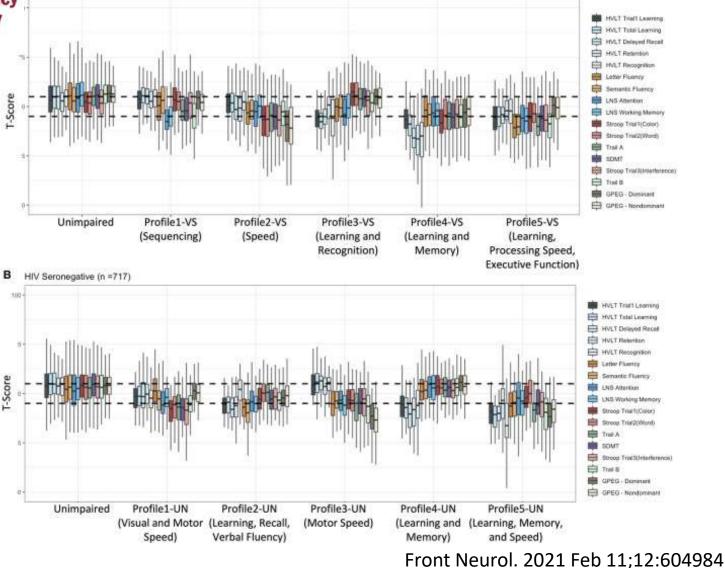
First published: 29 August 2023 | https://doi.org/10.1111/hiv.13532 | Citations: 2

Outcome	Number of ACMs	Adjustment					OR (95% CI)	p-value
	Any	Unadjusted		_ <u>_</u>	-	1	3.33 (1.88 - 5.9)	<0.001
		Level 1		-	<b>—</b>		2.47 (1.34 - 4.56)	0.004
		Level 2		-	<b>—</b>		1.93(0.93 - 3.97)	0.076
	1	Unadjusted		-			2.09 (1.03 - 4.23)	0.04
Falls		Level 1	-	-	<b>-</b>		1.66(0.79 - 3.49)	0.177
		Level 2	-	-	-		1.43(0.63 - 3.25)	0.389
	2 or more	Unadjusted			-		6.83 (3.31 – 14.11)	<0.001
		Level 1		<b>-</b>	-	—	4.53(2.06 - 9.98)	< 0.001
		Level 2		<u> </u>	-	<b>—</b>	3.58 (1.37 - 9.38)	0.009
	Any	Unadjusted		<b></b>	_		2.29 (1.48 - 3.55)	<0.001
		Level 1		-			1.8(1.12 - 2.92)	0.016
		Level 2		-	1		1.66 (0.94 - 2.95)	0.081
	1	Unadjusted		-	-1		1.93 (1.16 – 3.2)	0.011
Frailty		Level 1	1	-			1.62(0.93 - 2.82)	0.087
		Level 2	-	-			1.52(0.82 - 2.84)	0.186
	2 or more	Unadjusted		<b>—</b>	•	4	3.33(1.71 - 6.47)	< 0.001
		Level 1		-	_		2.26 (1.09 - 4.70)	0.028
		Level 2		-	—		2.12(0.89 - 5.02)	0.089
			0.50 1	1.0 2.0	10	90.160		
			0.50	1.0 2.0	4.0	8.0 16.0		

Odds ratio



Cognitive profiles in women with and without HIV



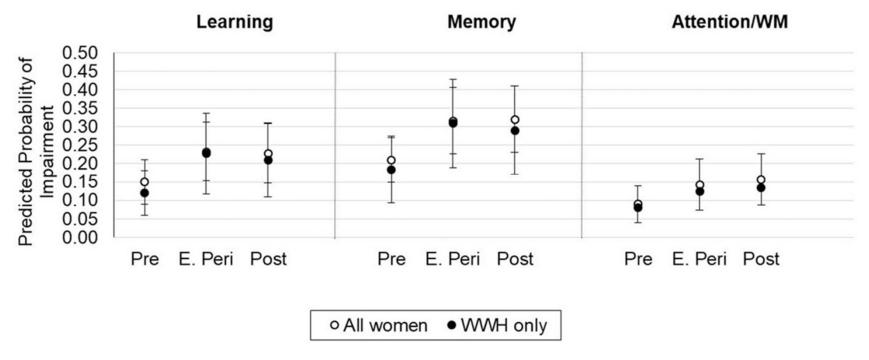
Women With HIV - Virally Suppressed (WWH-VS) (n =929)

#### **ORIGINAL STUDIES**

# Cognitive changes during the menopausal transition: a longitudinal study in women with and without HIV

Maki, Pauline M. PhD¹; Springer, Gayle MLA²; Anastos, Kathryn MD³; Gustafson, Deborah R. PhD⁴; Weber, Kathleen MS⁵; Vance, David PhD⁶; Dykxhoorn, Derek PhD⁷; Milam, Joel PhD՞8; Adimora, Adaora A. MD⁶; Kassaye, Seble G. MD¹0; Waldrop, Drenna PhD¹¹; Rubin, Leah. H. PhD²,1²

#### Menopause



Menopause 2021 Jan 11;28(4):360-368

4

Management considerations

# EACS guidelines



#### Algorithm

The person or their relatives complaining of, or care giver noting cognitive or CNS neurological symptoms - without or with optimally managed obvious confounding conditions(i)

→ Initial assessments(ii)

↓

Problems suspected

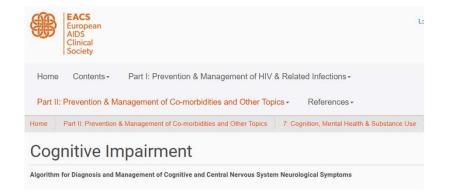
Evaluation for depression and anxiety and possible treatment(iii)

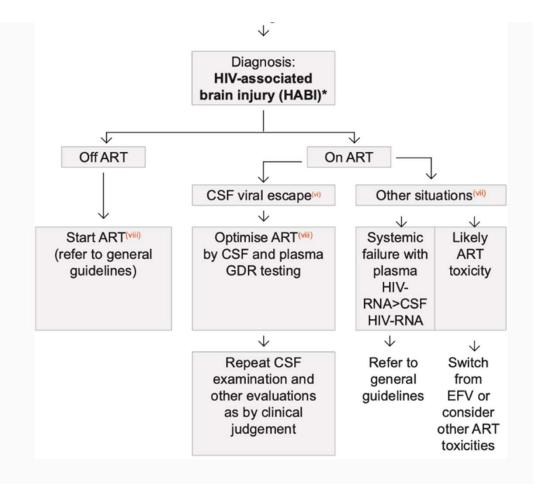
 $\downarrow$ 

Problems persisting but depression or anxiety excluded or optimally managed



# EACS guidelines





# Summary

1	<ul> <li>Defining brain health in persons with HIV</li> <li>Recently HABI definition has been proposed</li> </ul>
2	<ul> <li>Some HIV considerations</li> <li>Early initiation of ART may mitigate long term effects on cognitive health</li> <li>Don't forget to look for CSF HIV RNA escape</li> </ul>
3	<ul> <li>Some non-HIV considerations</li> <li>Impact on comorbidities and concomitant medication</li> <li>Social determinants of health, stigma, mental health have significant impact</li> </ul>
4	<ul> <li>Management</li> <li>Screening for contributing factors is paramount</li> <li>Recognising drivers and their management is paramount</li> </ul>





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