Olfactory Identification is Associated with Neuroimaging Biomarkers

Shannon L. Risacher, PhD

Co-Leader, Neuroimaging Core, Indiana Alzheimer Disease Center Indiana University School of Medicine



Data Blitz – ADC Meeting, 4/21/2018



Olfaction in AD

- Multiple studies have shown that patients with MCI and AD show impaired olfactory identification (Doty 1987, Djordjevic 2008, Devanand 2010).
- Studies have also shown an association between olfactory identification impairment and medial temporal lobe neurodegeneration (Growdon 2015, Murphy 2003).
- However, studies have suggested a minimal association with amyloid deposition, which was mostly driven by differences in diagnostic group (Growdon 2015, Bahar-Fuchs 2010).

Goal

With the advent of PET imaging techniques to measure tau deposition in vivo, we sought to evaluate the relationship between olfactory identification and cerebral tau deposition on [¹⁸F]Flortaucipir PET, as well as medial temporal (MTL) neurodegeneration, in participants atrisk for Alzheimer's disease



Methods

 33 individuals were included in the analysis, including 19 cognitively normal older adults (CN), 7 with subjective cognitive decline (SCD), and 5 with MCI

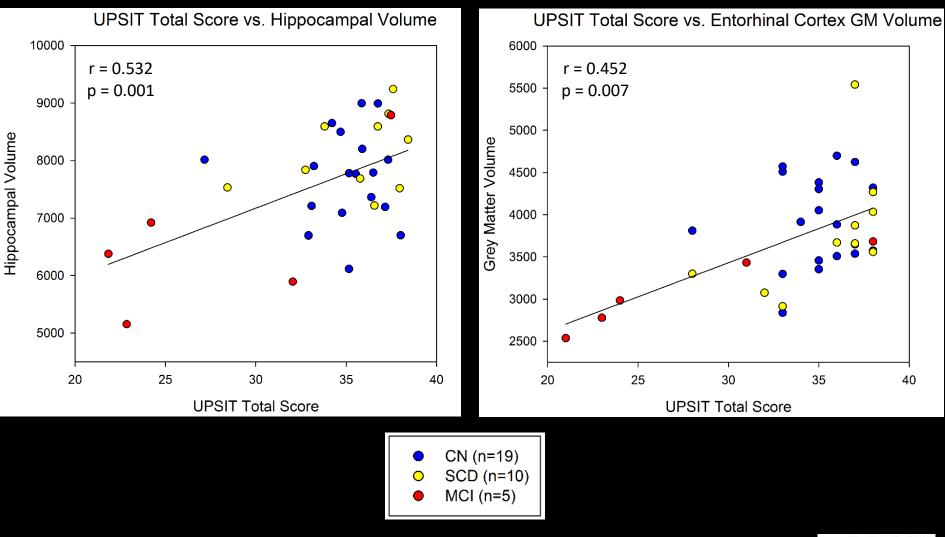
	CN	SCD	MCI	p-value
	(n=19)	(n=10)	(n=5)	
Age (years)	68.5 (6.9)	72.2 (6.4)	75.7 (10.6)	ns
Education (years)	17.6 (2.0)	17.5 (2.0)	16.0 (4.0)	ns
Sex (M, F)	4, 15	4, 6	4, 1	0.046
CDR – Sum of Boxes [‡]	0.2 (0.3)	0.1 (0.2)	1.2 (1.6)	0.003
MoCA Total Score [‡]	26.8 (2.5)	26.1 (2.3)	22.5 (3.8)	0.024
Craft Story Immediate Recall ^{‡,§}	16.4 (3.5)	16.7 (2.1)	12.0 (2.7)	0.016
Craft Story Delayed Recall ^{‡,§}	15.8 (3.5)	17.5 (2.7)	10.1 (4.9)	0.001
CCI – Self (12 items) ^{‡,¶}	16.2 (4.1)	24.0 (4.4)	33.5 (12.7)	<0.001
CCI – Informant (12 items) ^{‡,#}	17.4 (6.6)	15.2 (3.8)	26.1 (2.8)	ns
UPSIT Total Score**	34.5 (2.3)	35.7 (3.3)	28.7 (7.0)	0.006

Methods



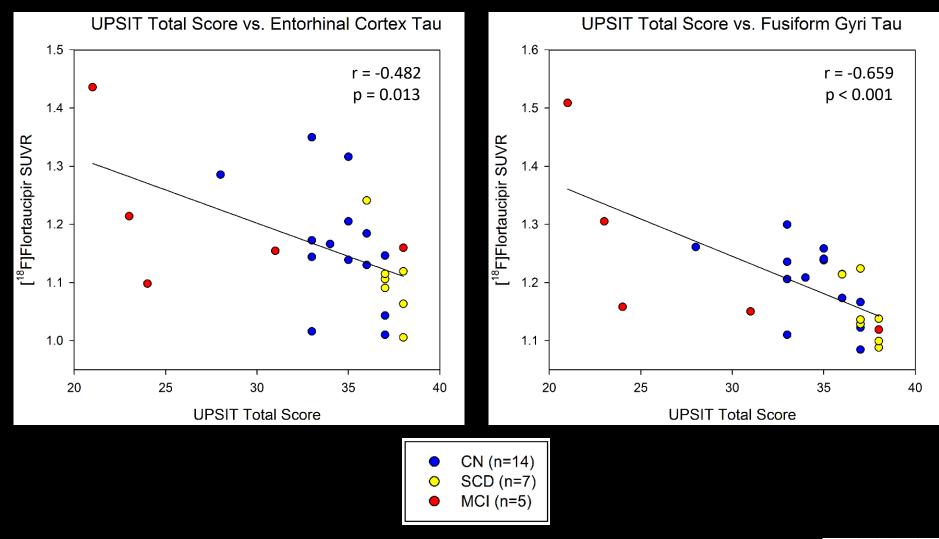
 The relationship between a neurodegeneration and tag a linear regression model, intracranial volume (MRI o

https://www.sciencedirect.com/science/article/pii/S2095881115300202 http://www.hnoinfo.com/fb/fb0405arnold/tes_top.html



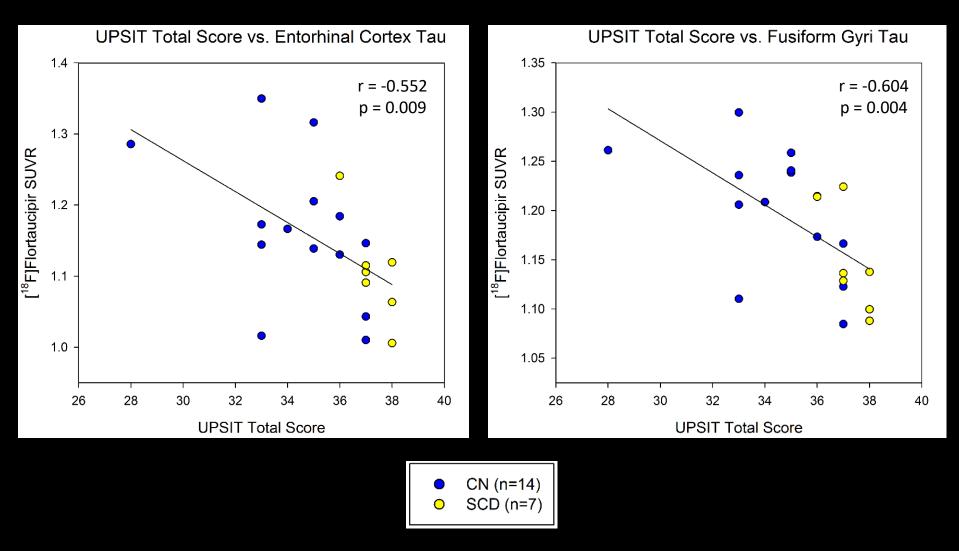
<u>iad</u>c

Risacher et al. (2017) Alz & Dementia: DACM





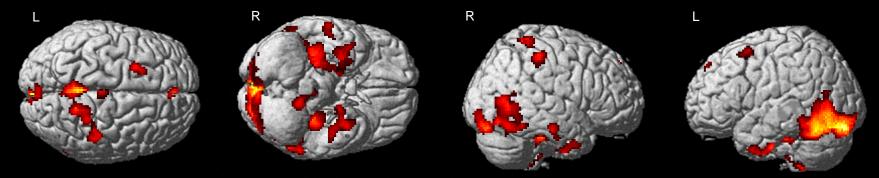
Risacher et al. (2017) Alz & Dementia: DACM



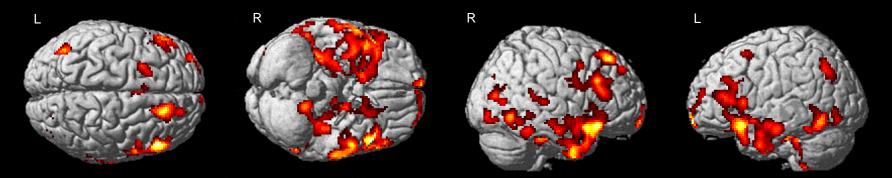


Risacher et al. (2017) Alz & Dementia: DACM

A) Association between Tau Deposition and UPSIT Total Score in all participants



B) Association between Tau Deposition and UPSIT Total Score in CN and SCD only





Cluster p<0.05 (FWE) Risacher et al. (2017) *Alz & Dementia: DACM*

Conclusions



- Olfactory identification was not associated with amyloid (*data not shown*) but was associated with MTL neurodegeneration.
- Olfactory identification was associated with MTL tau deposition both on regional and voxel-wise analysis.
- Similar results were observed in cognitively normal individuals only (CN/SCD).
- This test may have utility to detect AD neuropathology in early disease stages.



Acknowledgments



Dept of Radiology & Imaging Sciences

Andrew SaykinHeEileen TallmanWeRachael DeardorffJohGary HutchinsKarBrenna McDonaldKar

Heather Polson Wendy Territo John West Karmen Yoder

Eli Lilly/Avid Pharmaceuticals for assistance with the AV-1451/ Flortaucipir tracer Liana Apostolova Trina Bird Jared Brosch Steve Brown Martin Farlow Evan Finley Tatiana Foroud Yolanda Graham-Dotson Su Gao Bradley Glazier Kala Hall TJ Olivares Fred Unverzagt

Indiana Alzheimer Disease Center

<u>Funding</u>: National Institute on Aging (K01 AG049050, R01 AG19771, P30 AG10133), Alzheimer's Association, Indiana Clinical and Translational Science Institute, Indiana University-IU Health Strategic Research Initiative