Table e-1 Mean scores clinical measures when excluding patients with late onset depressive disorder (LOD) and healthy controls (HC) on medications potentially affecting these scores*

Clinical assessment	LOD	НС	P
	n=33	n=24	
UPDRS Motor Examination mean (SD)	0.7 (1.5)	1.9 (2.5)	.004
	n= 19	n=30	
RBD1Q "yes" n (%)	5 (13.8)	2 (6.6)	.08
RBDSQ mean (SD)	4.1 (3.1)	2.1 (2.1)	.02
	n=26	n=24	
SCOPA-AUT mean (SD)	14.3 (7.7)	7.6 (5.1)	.001

Mann Whitney U test, Fisher's Exact test

Abbreviations UPDRS= Unified Parkinson's Disease Rating scale, RBD1Q= REM Sleep Behavior Disorder Single-Question Screen, RBDSQ= The REM sleep Behaviour Disorder screening questionnaire, SCOPA-AUT= Scales for Outcomes in Parkinson's disease- Autonomic.

*Potentially affecting RBDSQ, RBD1Q; Citalopram, Sertraline, Escitalopram, Venlafaxine, Fluoxetine, Nortriptyline, Amytriptyline, Paroxetine, Olanzapine and Risperidone, SCOPA AUT: Nortriptyline, Risperidone, Olanzapine, Pregabalin, Lisinopril, Atenolol, Losartan, Amlodipine, Ramipril, Bisoprolol and Bendroflumethiazide, MDS-UPDRS motor examination: Olanzapine, Risperidone and Amlodipine.

Table e-2 Baseline characteristics of patients with late onset depressive disorder (LOD) and healthy controls (HC) who had a ¹²³l-ioflupane SPECT

	LOD (n=29)	HC (n=25)	р
Age in years mean (SD)	67.7 (7.5)	69.7 (9.0)	.37
Male/ Female n (%)	13 (45)	9 (36)	.35
Caucasian n (%)	25 (86.2)	24 (96)	.23
Family history of PD/ dementia n (%)	9 (31)	6 (24)	.40
<12 years education n (%)	8 (27.6)	5 (20)	.37
Q risk score ^ mean (SD)	18.8 (11.6)	19.2 (12.1)	.96
Anti-hypertensive treatment n (%)	10 (34.5)	4 (16.4)	.11
Diabetes n (%)	5 (17)	3 (12)	.44
Current/ Previous smoker n (%)	15 (51.7)	10 (40)	.28*

Independent samples t test, Mann Whitney U test and Fisher's exact

Abbreviations; PD =Parkinson's disease

[^]Q risk score=Cardiovascular risk test calculator (https://qrisk.org/2017/, 2017), using the age, sex, ethnicity, smoking status, diabetes status and evidence of; chronic kidney disease, atrial fibrillation, blood pressure treatment and Rheumatoid arthritis, 2 HC did not complete the q risk

^{*}Current and ex-smoker vs Non smoker

Table e-3 Comparison of MRI findings between patients with late onset depressive disorder (LOD) and healthy controls (HC)

Region	LOD	HC	р	
	Mean m	Mean mm³ (SD)		
	Median [1st			
Third Ventricle	3.11 (0.15)	3.15 (0.12)	.20	
	3.11 [3.01;3.22]	3.18 [3.04; 3.25]		
Fourth Ventricle	3.27 (0.11)	3.27 (0.10)	.48	
	3.27 [2.20;3.33]	3.26 [3.19;3.36]		
Right Caudate	3.49 (0.05)	3.49 (0.05)	.47	
	3.48 [3.46;3.53]	3.48 [3.43;3.51]		
Left Caudate	3.46 (0.05)	3.47 (0.05)	.89	
	3.45 [3.41;3.49]	3.46 [3.43;3.51]		
Right Putamen	3.59 (0.04)	3.59 (0.02)	.19	
	3.59 [3.55;3.63]	3.59 [3.57;3.62]		
Left Putamen	3.60 (0.05)	3.60 (0.03)	.26	
	3.60 [3.56;3.63]	3.60 [3.58;3.63]		
Right Hippocampus	3.55 (0.04)	3.58 (0.03)	.08	
	3.56 [3.52;3.58]	3.57 [3.54;3.61]		
Left Hippocampus	3.54 (0.05)	3.55 (0.03)	.45	
	3.54 [3.50;3.59]	3.55 [3.53;3.59]		
Right Amygdala	3.18 (0.05)	3.21 (0.03)	.07	
	3.20 [316;3.21]	3.21 [3.2;3.24]		
Left Amygdala	3.19 (0.05)	3.21 (0.03)	.24	
	3.19 [3.17;3.24]	3.21 [3.18;3.24)		
Total white matter	5.61 (0.05)	5.61 (0.05)	.30	
	5.61 [5.58;5.65]	5.61 [5.57;5.65]		
Total grey matter	5.64 (0.03)	5.65 (0.03)	.83	
	5.64 [5.62;5.67]	5.65 [5.63;5.67]		
Total white matter lesion	3.04 (0.60)	3.16 (0.63)	.96	
	3.04 [2.54;3.43]	3.25 [2.62;3.56]		

Analysis of variance (ANCOVA) with total intracranial volume (TIV), type of scanner, age and gender as a covariate.

Abbreviations: mm³=millimeters cube, SD= standard deviation.

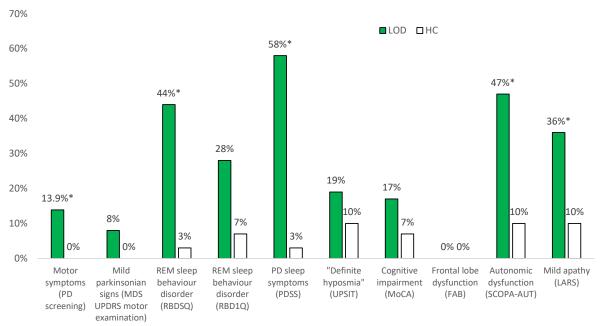


Figure e-1 Patients with LOD (n=36) and healthy controls (HC) (n=30) that had abnormal scores on the clinical scales when using cut-offs

LOD= late onset depressive disorder, PD screening= Parkinson's disease screening questionnaire, MPS= Mild Parkinsonian signs, RBD1Q= REM Sleep Behavior Disorder Single-Question Screen, RBDSQ= The REM sleep Behaviour Disorder screening questionnaire, PDSS=Parkinson's disease sleep scale, UPSIT= University of Pennsylvania Smell Identification Test, MoCA= Montreal Cognitive Assessment, FAB=Frontal Assessment Battery, SCOPA-AUT= Scales for Outcomes in Parkinson's disease- Autonomic, , LARS= Lille Apathy Rating Scale

^{*}p<0.05 using a Fisher's Exact between participant groups

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