

## Exploring [<sup>11</sup>C]CPPC as a CSF1R-targeted PET Imaging Marker for Early Parkinson's Disease Severity.

### Supplementary Data

Supplementary table 1.

Group	Sex	Age	PMI	Lewy Path	ADNC	Other	Inferior parietal cortex		Caudate		Midbrain		Basal ganglia	
							GM	WM	GM	WM	GM	WM	GM	WM
Control	F	90	22	None	None; A0B0C0		5.5	ND	0.7	ND	4.4	ND	0.9	0.5
Control	M	72	20	None	None; A0B0C0		8.8	8.0	ND	ND	ND	ND	ND	ND
Control	F	65	23	None	None; A0B0C0		80.7	4.2	ND	ND	ND	ND	ND	ND
Control	M	82	15	None	None; A0B0C0	CVD	13.6	6.7	22.0	5.3	12.6	ND	31.5	2.8
Control	F	68	12	None	None; A0B0C0		8.3	1.9	3.4	1.0	1.7	0.2	4.9	0.7
Control	M	97	16	None	None; A0B0C0		6.4	4.3	0.0	0.1	4.2	0.8	ND	1.66
PD	M	77	16	Neocortical	Low; A1B1C1		119.1	63.7	1.9	0.5	1.9	0.6	0.7	0.0
PD	M	91	19	Neocortical	Low; A1B2C1	CVD	12.6	4.4	18.0	2.1	14.6	4.2	17.6	6.7
PD	M	72	15	Brainstem	None; A0B0C0	Hypoxic- ischemic	7.3	0.7	ND	ND	8.1	ND	3.9	0.9
PD	M	65	6	Limbic	Very Low; A1B0C0	CVD + AA	94.4	44.3	19.5	1.0147	10.0	3.3	10.4	3.7
PD	F	67	16	Neocortical	Low; A1B1C2		198.8	101.9	ND	ND	ND	ND	ND	ND
PD	M	90	16	Brainstem	Low; A1B1C1	CVD	101.2	55.6	23.5	3.5	9.6	ND	47.1	18.2

$B_{\max}$  (fmol/mg) sites of  $^3\text{H}$ -JHU11761 (CSF1R) in frozen sections. GM, grey matter; WM, white matter. ND=not detected. CVD = microvascular ischemic disease. AA = amyloid angiopathy. Lewy pathology = brainstem, limbic, or neocortical. ADNC = Alzheimer's disease related change (very low, low, moderate, high). PMI = Post Mortem Interval.

**Supplementary table 2.**

Region	F	p		t	p		
Anterior cingulate	1.65	0.217	HC vs. Mild PD	-2.56	0.338		
			HC vs. Mod PD	1.13	0.866		
			Mild PD vs. Mod PD	3.69	0.263		
Brainstem	5.25	0.015	*	HC vs. Mild PD	-1.4	0.522	
			HC vs. Mod PD	3.88	0.060		
			Mild PD vs. Mod PD	5.28	0.012	*	
Cerebellar Cortex	5.59	0.012	*	HC vs. Mild PD	-1.02	0.738	
			HC vs. Mod PD	4.74	0.031	*	
			Mild PD vs. Mod PD	5.77	0.011	*	
Striatum	9.68	0.001	**	HC vs. Mild PD	-0.68	0.902	
			HC vs. Mod PD	7.68	0.002	**	
			Mild PD vs. Mod PD	8.35	0.002	**	
Frontal cortex	5.36	0.014	*	HC vs. Mild PD	-1.13	0.646	
			HC vs. Mod PD	4.11	0.043	*	
			Mild PD vs. Mod PD	5.24	0.012	*	
Hippocampus	6.39	0.008	*	HC vs. Mild PD	-1.47	0.519	
			HC vs. Mod PD	4.59	0.031	*	
			Mild PD vs. Mod PD	6.05	0.006	*	
Occipital cortex	3.24	0.062		HC vs. Mild PD	-1.22	0.677	
				HC vs. Mod PD	3.46	0.156	
				Mild PD vs. Mod PD	4.68	0.051	
Parietal cortex	5.75	0.011	*	HC vs. Mild PD	-0.99	0.748	
				HC vs. Mod PD	4.81	0.027	*
				Mild PD vs. Mod PD	5.80	0.010	*
Posterior cingulate	2.69	0.094		HC vs. Mild PD	-2.08	0.477	
				HC vs. Mod PD	3.15	0.341	
				Mild PD vs. Mod PD	5.23	0.079	
Pallidum	4.39	0.027	*	HC vs. Mild PD	1.48	0.716	
				HC vs. Mod PD	5.60	0.068	
				Mild PD vs. Mod PD	7.08	0.023	*
Temporal cortex	5.58	0.012	*	HC vs. Mild PD	-1.86	0.457	
				HC vs. Mod PD	4.69	0.057	
				Mild PD vs. Mod PD	6.55	0.009	*
Thalamus	6.48	0.007	*	HC vs. Mild PD	-1.13	0.713	
				HC vs. Mod PD	5.36	0.019	*
				Mild PD vs. Mod PD	6.49	0.006	*

F values for ANOVA comparing regional [<sup>11</sup>C]CPPC V<sub>T</sub> across healthy controls (HC), PD patients with mean

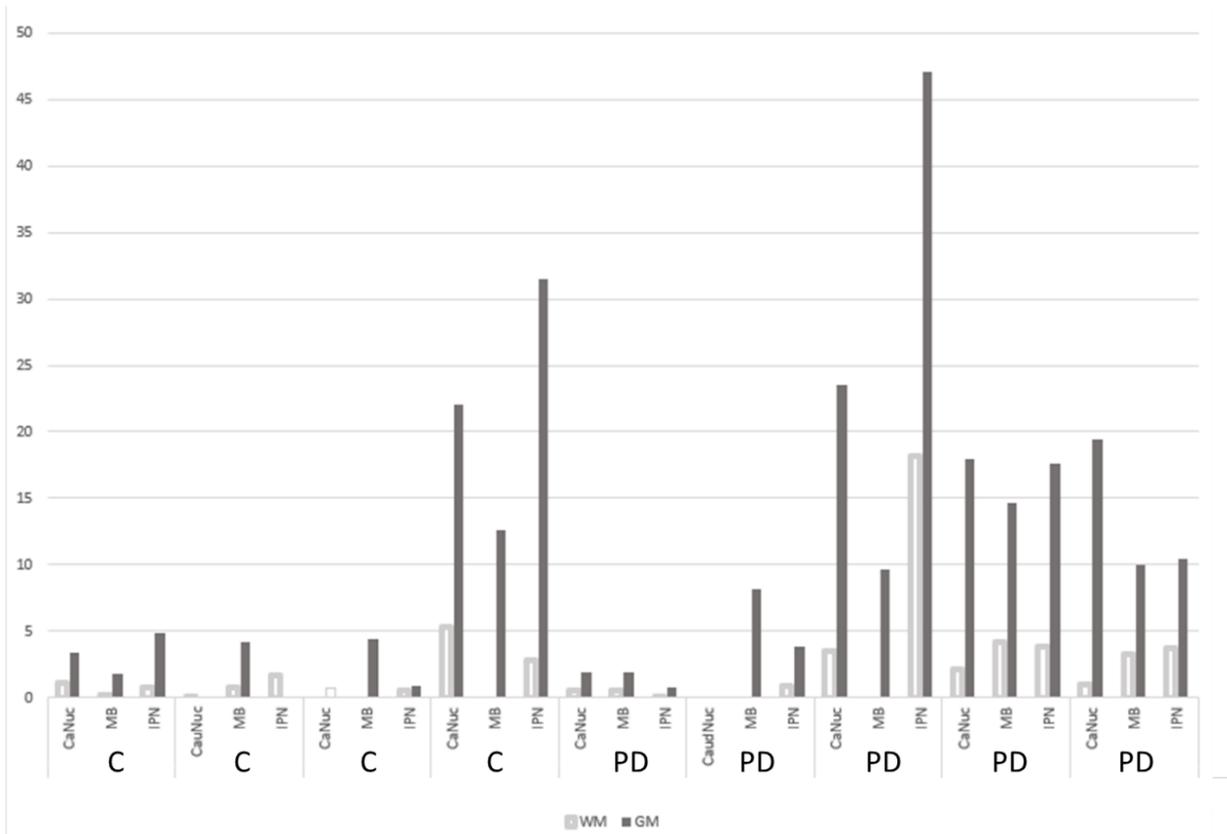
MDS-UPDRS part II below median (Mild PD), and PD patients with mean MDS-UPDRS part II at median or

above (Moderate PD), all with Hoehn & Yahr stage  $\leq 2$ . T values for pair-wise post hoc tests between groups with Tukey's test. \* $p < 0.05$  (uncorrected). \*\* $p < 0.005$  (corrected for multiple comparisons).

**Supplementary table 3.** Antibody identification for IHC and western blot experiments.

<b>Target</b>	<b>Antibody</b>	<b>Source Company</b>	<b>Catalog Number</b>
<i>IHC</i>			
CSF1R	M-CSFR/CD115 (6B9B9)	NOVUS Biologicals	NBP2-37292
IBA1	Anti-Iba1	Wako	019-19741
anti-Rabbit IgG	Goat anti-Rabbit IgG (H+L) Cross-Adsorbed Secondary Antibody	Alexa Fluor	488
anti-Mouse IgG	Goat anti-Mouse IgG (H+L) Cross-Adsorbed Secondary Antibody	Alexa Fluor	594
<i>Western Blot</i>			
CSF1R	CD115 (c-fms) Monoclonal Antibody	Invitrogen	14-1152-82
Actin	beta Actin Antibody (C4)	Santa Cruz	sc-47778
anti-Mouse IgG	Peroxidase AffiniPure® Goat Anti-Mouse IgG (H+L)	Jackson ImmunoResearch	115-035-146

**Supplementary Figure 1.**



5 nM binding survey of <sup>3</sup>H-JHU11761 binding in 5 healthy controls (C) and 4 subjects with Parkinson's disease (PD) in inferior parietal cortex (IPC), midbrain (MB), and caudate nucleus (CaNuc).