Estimating the evolution of disease in the Parkinson's Progression Markers Initiative

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Supplementary Appendix

[FIGUREA1.JPEG HERE]

Figure A.1: Distribution of estimate of time shift, $\hat{\delta}_{\iota}$ by disease status. **Abbreviations**: PD, Parkinson Disease; HC, Healthy Control.

[FIGUREA2.JPEG HERE]

Figure A.2: Correlation between random slopes for pair of outcomes.

Abbreviations: PIGD, Posture Instability and Gait Difficulty; RBD, REM sleep behavior disorder; UPSIT, University of Pennsylvania Smell Identification Test; MOCA, Montreal Cognitive Assessment; HVLT, Hopkins Verbal Learning Test; GDS, Geriatric Depression Scale; SFT, Semantic Fluency Test; SCOPA, Scale for Outcomes in Parkinson's; LINEORT, Line Orientation Test; UPDRS, Unified Parkinson Disease Rating Scale; STAI, State Trait Anxiety Index; PD, Parkinson Disease; HC, Healthy Control.

[FIGUREA3A.JPEG and FIGUREA3B.JPEG HERE]

(a) For 17 outcomes (b) For 9 outcomes with UPDRS separated **Figure A.3**: Population level predicted severity for female with mean age and median latent time of a healthy control. The legend is ordered by the age at which the predicted severity level for each outcome is 0.5 (sliced horizontally). **Abbreviations**: PIGD, Posture Instability and Gait Difficulty; RBD, REM sleep behavior disorder; UPSIT, University of Pennsylvania Smell Identification Test; MOCA, Montreal Cognitive Assessment; HVLT, Hopkins Verbal Learning Test; GDS, Geriatric Depression Scale; SFT, Semantic Fluency Test; SCOPA, Scale for Outcomes in Parkinson's; LINEORT, Line Orientation Test; UPDRS, Unified Parkinson Disease Rating Scale; STAI, State Trait Anxiety Index; PD, Parkinson Disease; HC, Healthy Control.

[FIGUREA4ALJPEG AND FIGUREA4B.JPEG HERE]

(a) Female.

(b) Male.

Figure A.4: Positional variance diagram of the central ordering. The x-axis is the event position.

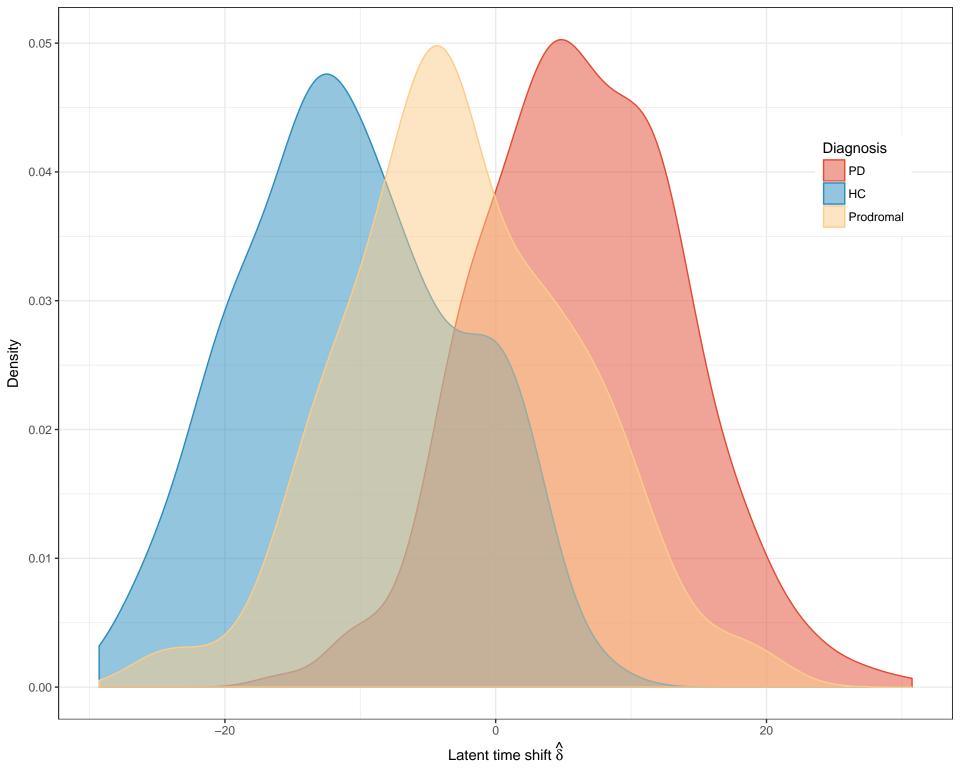
Abbreviations: PIGD, Posture Instability and Gait Difficulty; RBD, REM sleep behavior disorder;

UPSIT, University of Pennsylvania Smell Identification Test; MOCA, Montreal Cognitive Assessment; HVLT, Hopkins Verbal Learning Test; GDS, Geriatric Depression Scale; SFT, Semantic Fluency Test; SCOPA, Scale for Outcomes in Parkinson's; LINEORT, Line Orientation Test; UPDRS, Unified Parkinson Disease Rating Scale; STAI, State Trait Anxiety Index; PD, Parkinson Disease; HC, Healthy Control.

[FIGUREA5.JPEG HERE]

Figure A.5: Positional variance diagram of the central ordering

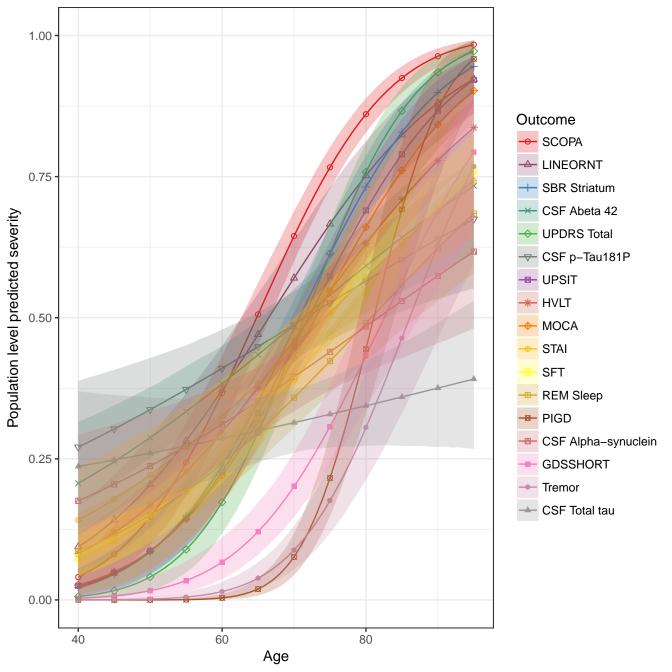
Abbreviations: PIGD, Posture Instability and Gait Difficulty; RBD, REM sleep behavior disorder; UPSIT, University of Pennsylvania Smell Identification Test; MOCA, Montreal Cognitive Assessment; HVLT, Hopkins Verbal Learning Test; GDS, Geriatric Depression Scale; SFT, Semantic Fluency Test; SCOPA, Scale for Outcomes in Parkinson's; LINEORT, Line Orientation Test; UPDRS, Unified Parkinson Disease Rating Scale; STAI, State Trait Anxiety Index; PD, Parkinson Disease; HC, Healthy Control.

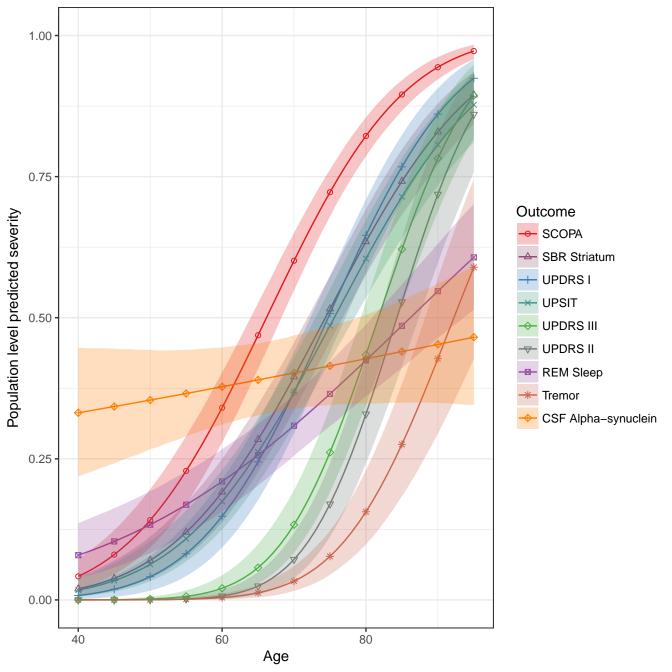


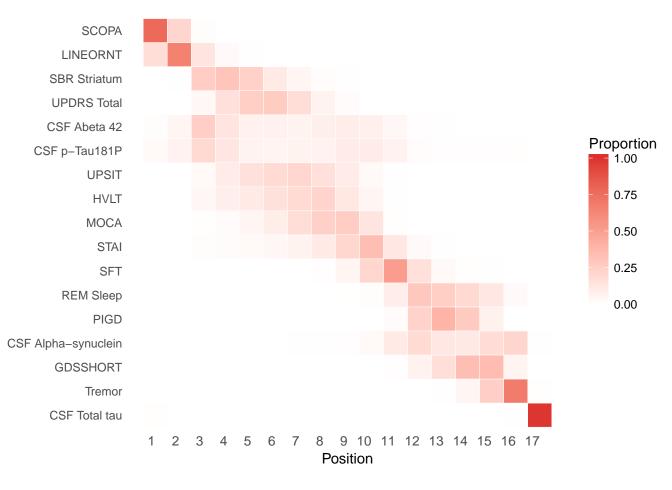
UPSIT																	1
UPDRS Total																1	0.0647
Tremor															1	0.5447	0.0626
STAI														1	0.2784	0.4523	-0.0585
SFT													1	0.2345	0.0348	0.1317	-0.0257
SCOPA												1	0.1315	0.4366	0.0937	0.4062	-0.025
SBR Striatum											1	0.3717	0.0844	0.1113	-0.3266	0.2989	0.0971
REM Sleep										1	0.2183	0.2655	0.1294	0.2381	0.0267	0.1187	-0.0343
PIGD									1	0.1459	0.3091	0.3663	0.1748	0.1855	0.2096	0.606	0.035
MOCA								1	0.2326	0.1748	0.0357	0.124	0.3907	0.3017	0.0638	0.2298	-0.0467
LINEORNT							1	0.2388	0.0119	0.0046	0.0232	0.0113	0.1152	0.1915	0.1164	0.0929	0.0142
HVLT						1	0.3473	0.4223	0.0902	0.0422	0.0908	-0.0441	0.397	0.1958	-0.013	0.0732	0.0493
GDSSHORT					1	0.285	0.1604	0.2156	0.2108	0.1644	0.1085	0.4096	0.2789	0.6638	0.1003	0.2976	-0.0413
CSF Total tau				1	-0.0531	-0.0598	-0.0097	-0.1062	-0.1174	-0.0562	0.0562	-0.0525	-0.1121	-0.0347	0.001	0.0122	0.0444
CSF p-Tau181P			1	0.0561	0.0023	-0.0401	-0.018	0.1038	0.0135	0.0055	-0.2517	-0.0065	-0.1006	0.0796	0.0918	0.0027	-0.0446
CSF Alpha-synuclein		1	0.1115	0.1449	-0.0521	0.0952	0.0067	0.0409	-0.0798	-0.0078	0.0226	0.0325	-0.0727	-0.0494	-0.0151	-0.0515	0.0203
CSF Abeta 42	1	0.1052	0.5427	0.2236	0.0625	0.0647	0.0447	0.1798	0.0273	0.0658	-0.2042	-0.0739	0.0349	0.1413	0.111	0.0437	-0.0408
CSF Abeta 42 رچحی رچحی	peta A2	nuclein re	UIR CSFT	GDS GDS	HORT	HULLING	ORNI	MOCA	PIGD REP	SPR S	itiatum e	SCOPA	est.	STAL	Iremot UPDP	5 TOtal	JPSIT
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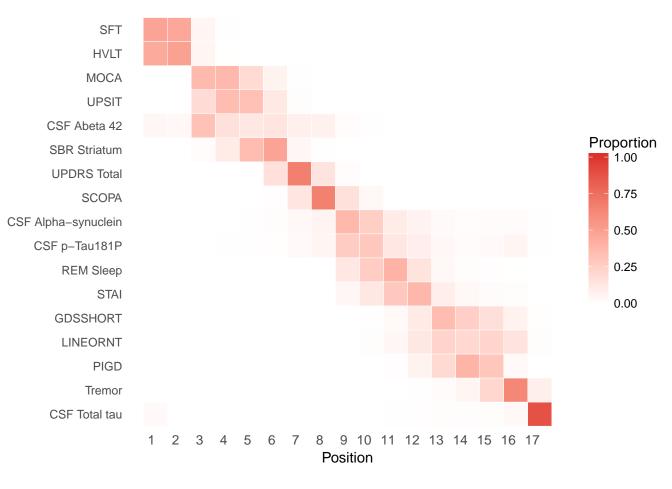
0.5 0.0 -0.5 -1.0

ρ 1.0









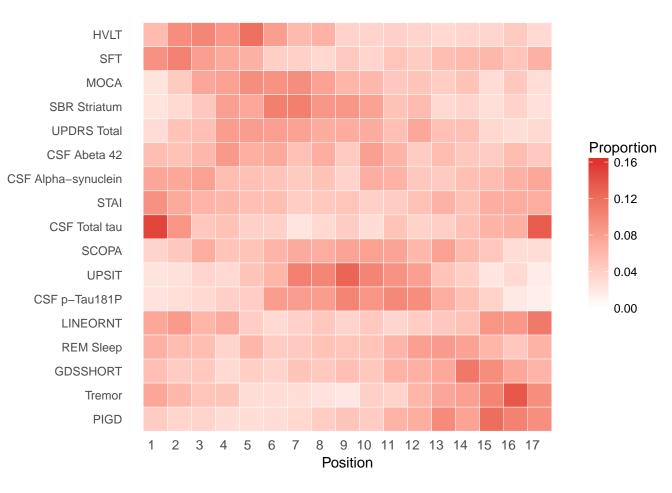


Table A. 1: Results of posterior estimates of parameters and corresponding 95%

credible interval for all 17 outcomes from the Parkinson's Progression Markers

Initiative (PPMI).

Pos	t. 95%		Post.	95%		Post.	95%		
	an Credible	Parameter		Credible	Parameter	Mean	Credible		
CSF Abe		M	10CA		Tremor				
Intercept -0.0		Intercept	-1.871	(-2.304,-1.431)	Intercept	-0.203	(-0.975,0.572)		
Age 0.0		Age	0.030	(0.023,0.036)	Age	-0.009	(-0.021,0.002)		
Female -0.0	(, , ,	Female	-0.284	(-0.427,-0.148)	Female	-0.176	(-0.419,0.065)		
Latent time, γ_1 0.0		Latent time, γ_8	0.945	(0.917,0.974)	Latent time, γ_{15}	0.094	(0.071,0.114)		
Error Variance, σ_1 0.4		Error Variance, σ_8	0.945	(0.917,0.974)	Error Variance, σ_{15}	0.811	(0.796,0.827)		
CSF Alpha-s			PIGD		UPDRS Total				
Intercept 0.8		Intercept	-2.250	(-3.054,-1.4)	Intercept	-0.524	(-0.985,-0.06)		
Age -0.0		Age	0.016	(0.003,0.028)	Age	0.010	(0.003,0.017)		
Female -0.1	(, , ,	Female	0.076	(-0.164,0.328)	Female	-0.047	(-0.188,0.098)		
Latent time, γ_2 0.0		Latent time, γ_9	0.114	(0.085,0.139)	Latent time, γ_{16}	0.073	(0.054,0.088)		
Error Variance, σ_2 0.4	Error Variance, σ_9	1.156	(1.134,1.178)	Error Variance, σ_{16} 0.349 (0.343,0.356)					
CSF p-Ta			A Sleep		UPSIT				
Intercept 0.3		Intercept	0.317	(-0.121,0.745)	Intercept	-1.103	(-1.567,-0.642)		
Age -0.0		Age	-0.006	(-0.013,0.001)	Age	0.019	(0.012,0.026)		
Female 0.0	(, , ,	Female	-0.115	(-0.256,0.027)	Female	-0.210	(-0.352,-0.067)		
Latent time, γ_3 0.0		Latent time, γ_{10}	0.040	(0.03,0.05)	Latent time, γ_{17}	0.043	(0.032,0.055)		
Error Variance, σ_3 0.7	Error Variance, σ_{10}		(0.568,0.6)	Error Variance, σ_{17}	0.218	(0.059,0.446)			
CSF Tota			Striatu		Heterogeneous parameters				
Intercept 1.7		Intercept	-1.079	(-1.476,-0.681)	Intercept, τ_0	5.002	(4.19,5.831)		
Age -0.0		Age	0.018	(0.012, 0.025)	Age, τ_1	-0.001	(-0.012,0.009)		
Female -0.2 Latent time. γ_4 0.0		Female	-0.084	(-0.211, 0.043)	Female, τ_2	-0.063	(-0.287,0.159)		
Latent time, γ_4 0.0 Error Variance, σ_4 0.2		Latent time, γ_{11} Error Variance, σ_{11}	0.049	(0.035, 0.061)					
GDSSH			COPA	(0.2,0.224)	Model comparison criteria				
Intercept -0.3	Intercept -2.319 (-2.685,-1.933)			WAIC 117860.0					
Age -0.0		Age	0.033	(0.027,0.039)		20166.7	0		
Female -0.0		Female	0.504	(0.388,0.618)	10010	20100.7			
Latent time, γ_5 0.0	()	Latent time, γ_{12}	0.038	(0.029, 0.047)					
Error Variance, σ_5 1.1		Error Variance, σ_{12}		(0.456,0.482)					
HVL		· · · · ·	SFT	(0.100,0.102)					
Intercept -1.1		Intercept	-0.905	(-1.286,-0.536)					
Age 0.0		Age	0.017	(0.011,0.023)					
Female -0.3		0	-0.492	(-0.616,-0.368)					
Latent time, γ_6 0.0		Latent time, γ_{13}	0.022	(0.016,0.029)					
Error Variance, σ_6 0.5	51 (0.534,0.568)	Error Variance, σ_{13}	0.429	(0.417,0.443)					
LINEO	RNT		STAI						
Intercept -2.0	41 (-2.609,-1.493)	Intercept	0.701	(0.26,1.139)					
Age 0.0		Age	-0.012	(-0.018,-0.005)					
Female 0.7	63 (0.592,0.932)	Female	0.136	(0.002,0.274)					
Latent time, γ_7 0.0	28 (0.019,0.038)	Latent time, γ_{14}	0.044	(0.033,0.054)					
Error Variance, σ_7 1.1	35 (1.152,1.22)	Error Variance, σ_{14}	0.594	(0.578,0.61)					