The impact of socioeconomic status on subsequent neurological outcomes in multiple sclerosis

Supplementary Material

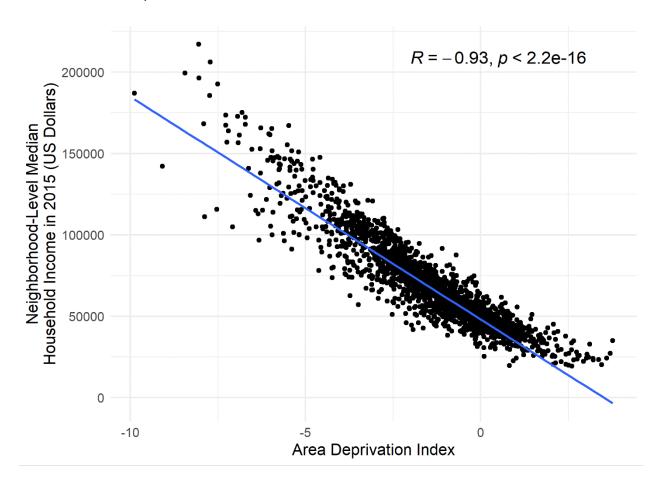
Zip Code Distribution of Research Participants

In this study, the mean (and standard deviation) number of participants per zip code was 2.5 (3.5). The maximum number of participants in a single zip code was 29 participants. Six (out of 723) zip codes had 20 or more participants. To preserve confidentiality given the relatively small number of study participants per zip code, we did not report zip code specific distribution of participants.

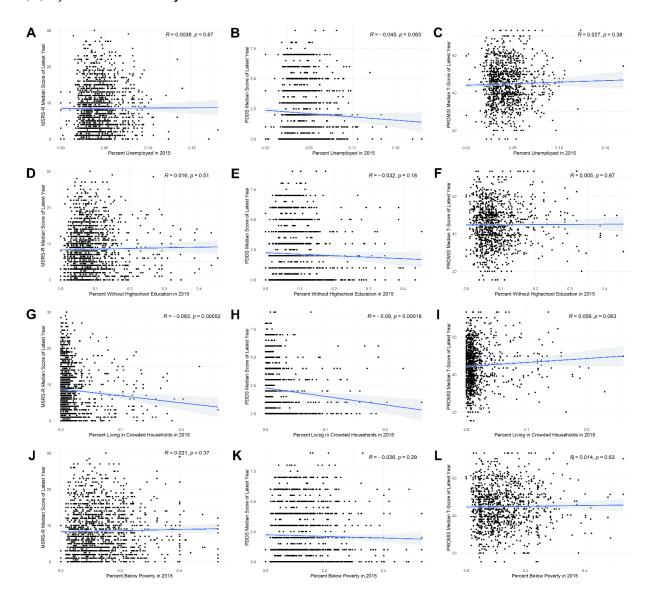
Additional Neighborhood Indicator of Socioeconomic Status (SES)

Using the zip codes of the home residence of each participant at enrollment to query the American Community Survey SES datasets from 2014 to 2018 (abbreviated as 2018), we derived components of the neighborhood SES indicator, and calculated the area deprivation index (ADI) in 2018 as described in the Methods of the main text (section 2.3). The covariate-adjusted regression models using ADI in 2018 were shown in eTable 3.

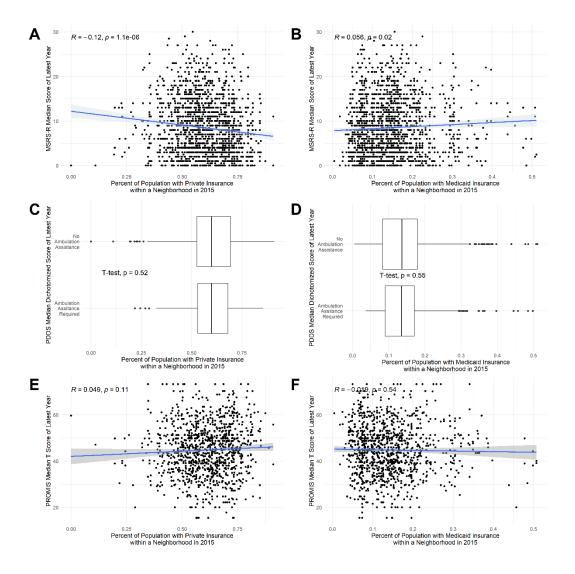
eFigure 1. Pearson correlation between *neighborhood median household income* in 2015 and *area deprivation index* in 2015 in the combined data.



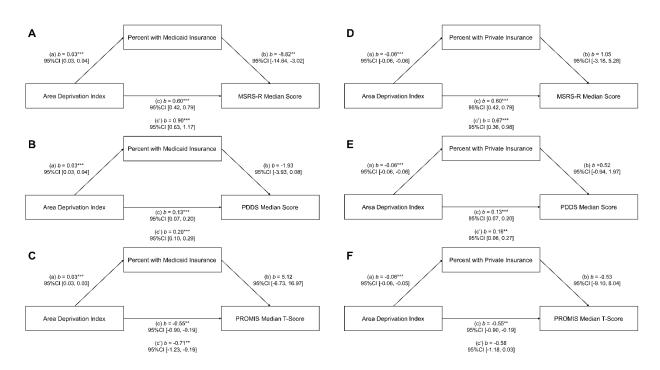
eFigure 2. Scatterplots showing the *non-significant* Pearson correlations between the components of the area deprivation index (ADI) in 2015 and subsequent patient-reported outcomes (PROs) of neurological and physical function. The ADI components in 2015 included percent unemployed (**A-C**), percent without high school education (**D-F**), percent living in crowded households (**G-I**), percent below poverty line (**J-L**) within the zip code of a participant's residential address. The PROs included the median scores of MSRS-R (**A**, **D**, **G**, **J**), PDDS (**B**, **E**, **H**, **K**), and PROMIS-physical function (**C**, **F**, **I**, **L**) from the latest year between 2018 and 2020.



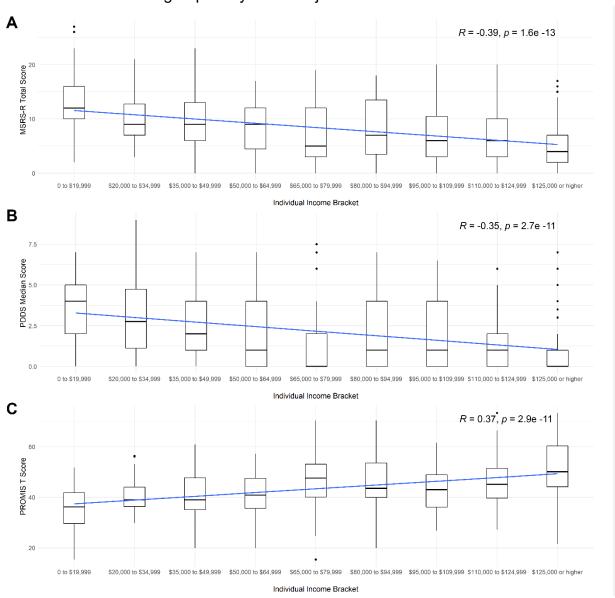
eFigure 3. Pearson correlations between neighborhood insurance coverage in 2015 and subsequent patient-reported outcomes (PROs) of neurological and physical function. Insurance coverage included percent with either *private* (**A**, **C**, **D**) or *Medicaid* (**B**, **D**, **F**) insurance within a neighborhood of the participant's residential zip code in 2015. The PROs included median scores of MSRS-R (**A**, **B**), PDDS (**C**, **D**), and PROMIS-physical function (**E**, **F**) from the latest year between 2018 and 2020.



eFigure 4. Covariate-adjusted mediation analyses to assess the mediating effect of healthcare insurance coverage. The covariates included age of latest patientreported outcome (PRO), sex, race and ethnicity, age of first symptom onset, disease duration, and type of disease-modifying therapy. Path model assessed whether the association between area deprivation index (ADI) in 2015 and PROs (i.e., MSRS-R, PDDS, and PROMIS T-Score) from the latest year between 2018 and 2020 was mediated by neighborhood percent of population with Medicaid insurance or by neighborhood percent of population with private insurance within the same zip code as the participant's residence. Path c represents the covariate-adjusted effect of ADI on PROs without mediators in the model. Path c' (indirect effect) represents the covariateadjusted effect of ADI on PROs with mediators in the model. Path a represents the effect of ADI on neighborhood percent of population with private medical insurance or Medicaid insurance. Path b represents the effect of neighborhood percent of population with private or Medicaid insurance on PROs. Non-standardized β regression coefficients, 95% confidence intervals, and significance thresholds (*p<0.05, **p<0.01, ***p<0.001) as reported by PROCESS Macro in SPSS are shown. Please also refer to Table 3 in the main text for the significance level for the mediating effect of healthcare insurance coverage.



eFigure 5. Association between *individual-level household income brackets* in 2020 and the median scores of the patient-reported outcomes of neurological and physical function in 2020 in a subgroup analysis of the joint cohorts.



eTable 1. Eligibility criteria of the original cohort studies. The current study dataset that drew from the original cohort studies only included adult participants (age 18 years or older) with a neurologist confirmed diagnosis of multiple sclerosis.

Cohort Study	Inclusion	Exclusion
PROMOTE ¹	Age 7 years or older	None
	Neurologist-confirmed diagnosis of multiple sclerosis (MS) or related disorders, including a first central nervous system demyelinating episode with a positive MRI scan or abnormal MRI scans characteristic of MS but no clinical symptoms of the disease OR controls	
Pittsburgh COVID-19	Age 18 years or older	None
and MS Study	Neurologist-confirmed diagnosis of multiple sclerosis (MS) or related disorders OR controls	
	Live in the United States	
CUIMC ² MS Genetics Study	Neurologist-confirmed diagnosis of multiple sclerosis (MS) or related disorders	None
CUIMC ² COVID-19	Age 18 years or older	None
and MS Study ⁴	Neurologist-confirmed diagnosis of multiple sclerosis (MS) or related disorders OR controls	
	Live in the United States	
GEMS ³	Age 5 years or older	None
	First-degree family members of patients with multiple sclerosis (MS), or patients who have received anti-TNF alpha therapy as treatment for inflammatory diseases other than MS (e.g., Crohn's disease, psoriasis and rheumatoid arthritis) with OR without a neurologist-confirmed diagnosis of MS	
	OR	
	Patients that have been referred for an evaluation of first presentation of neurological symptoms but do not have a diagnosis of MS	
	Live in the United States	

- Prospective Investigation of Multiple Sclerosis in the Three Rivers Region (PROMOTE), NCT02994121
- 2. Columbia University Irving Medical Center (CUIMC)
- 3. Genes and Environment in Multiple Sclerosis (GEMS), NCT01353547
- 4. Also referred to as Multiple Sclerosis (MS) Patient Response to COVID-19 Outbreak Study

eTable 2. Covariate-adjusted regression analysis using *area deprivation index* in **2015** as the exposure and the median scores of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020.

			MSRS-R			PDDS		PROMIS			
	Predictors	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	
	ADI	0.62	0.36, 0.89	<0.001	0.11	0.02, 0.20	0.02	-0.49	-1.08, 0.10	0.106	
	Age of latest PRO	0.16	0.11, 0.21	<0.001	0.09	0.07, 0.10	<0.001	-0.5	-0.63, -0.37	<0.001	
	Sex, Male	-0.01	-0.78, 0.76	0.98	0.55	0.29, 0.81	<0.001	-0.24	-1.98, 1.49	0.785	
horts	Sex, Other	3.71	-8.09, 15.50	0.538	0.71	-3.27, 4.70	0.726	-3.53	-22.73, 15.67	0.718	
ry Co	Race and Ethnicity, Other	1.23	0.10, 2.36	0.033	0.11	-0.28, 0.49	0.586	0.09	-3.04, 3.21	0.956	
Discovery Cohorts	Age of First Symptom Onset	-0.08	-0.13, -0.02	0.005	-0.04	-0.06, -0.02	<0.001	0.19	0.07, 0.32	0.003	
Ö	Disease Duration	0	-0.06, 0.06	0.933	0	-0.02, 0.02	0.633	0.03	-0.12, 0.17	0.716	
	DMT Group, High Efficacy	0.94	0.02, 1.86	0.046	0.55	0.24, 0.87	0.001	-2.31	-4.37, -0.24	0.029	
	DMT Group, Standard Efficacy	-1.44	-2.41, -0.46	0.004	-0.37	-0.70, -0.04	0.03	2.05	-0.10, 4.20	0.061	
	Observations	1248			1234			645			
	R ² / R ² adjusted	0.104 / 0.09	97		0.208 / 0.20	02		0.244 / 0.234			
			0.21,			0.03,	0.000	0.0	-1.12,	0.005	
	ADI	0.46	0.72	<0.001	0.12	0.21	0.009	-0.6	-0.08	0.025	
	ADI Age of latest PRO	0.46	0.72 0.10, 0.25	<0.001	0.12		<0.009	-0.6	-	<0.001	
S			0.72 0.10,			0.21 0.04,			-0.08 -0.56,		
ohorts	Age of latest PRO	0.17	0.72 0.10, 0.25 -0.87,	<0.001	0.07	0.21 0.04, 0.10 -0.28,	<0.001	-0.4	-0.08 -0.56, -0.24 -0.96,	<0.001	
ition Cohorts	Age of latest PRO Sex, Male Race and	0.17	0.72 0.10, 0.25 -0.87, 1.53 -1.60,	<0.001 0.588	0.07 0.15	0.21 0.04, 0.10 -0.28, 0.57 -0.46,	<0.001 0.496	-0.4 1.58	-0.08 -0.56, -0.24 -0.96, 4.11 -3.79,	<0.001 0.221	
eplication Cohorts	Age of latest PRO Sex, Male Race and Ethnicity, Other Age of First Symptom Onset Disease Duration	0.17 0.33 -0.24	0.72 0.10, 0.25 -0.87, 1.53 -1.60, 1.12 -0.15, -0.01 -0.15, 0.05	<0.001 0.588 0.725	0.07 0.15 0.02	0.21 0.04, 0.10 -0.28, 0.57 -0.46, 0.50 -0.04,	<0.001 0.496 0.939	-0.4 1.58 -0.92	-0.08 -0.56, -0.24 -0.96, 4.11 -3.79, 1.95 -0.06,	<0.001 0.221 0.53	
Replication Cohorts	Age of latest PRO Sex, Male Race and Ethnicity, Other Age of First Symptom Onset Disease Duration DMT Group, High Efficacy	0.17 0.33 -0.24 -0.08	0.72 0.10, 0.25 -0.87, 1.53 -1.60, 1.12 -0.15, -0.01 -0.15, 0.05 -1.67, 2.32	<0.001 0.588 0.725 0.036	0.07 0.15 0.02 -0.01	0.21 0.04, 0.10 -0.28, 0.57 -0.46, 0.50 -0.04, 0.01 -0.00, 0.06 -0.61, 0.80	<0.001 0.496 0.939 0.335	-0.4 1.58 -0.92 0.09	-0.08 -0.56, -0.24 -0.96, 4.11 -3.79, 1.95 -0.06, 0.24 -0.26, 0.12 -6.55, 1.29	<0.001 0.221 0.53 0.248	
Replication Cohorts	Age of latest PRO Sex, Male Race and Ethnicity, Other Age of First Symptom Onset Disease Duration DMT Group, High Efficacy DMT Group, Standard Efficacy	0.17 0.33 -0.24 -0.08 -0.05 0.32 -2.43	0.72 0.10, 0.25 -0.87, 1.53 -1.60, 1.12 -0.15, -0.01 -0.15, 0.05 -1.67,	<0.001 0.588 0.725 0.036 0.313	0.07 0.15 0.02 -0.01 0.03 0.1 -1.02	0.21 0.04, 0.10 -0.28, 0.57 -0.46, 0.50 -0.04, 0.01 -0.00, 0.06 -0.61,	<0.001 0.496 0.939 0.335 0.084	-0.4 1.58 -0.92 0.09 -0.07 -2.63 2.68	-0.08 -0.56, -0.24 -0.96, 4.11 -3.79, 1.95 -0.06, 0.24 -0.26, 0.12 -6.55,	<0.001 0.221 0.53 0.248 0.467	
Replication Cohorts	Age of latest PRO Sex, Male Race and Ethnicity, Other Age of First Symptom Onset Disease Duration DMT Group, High Efficacy DMT Group,	0.17 0.33 -0.24 -0.08 -0.05 0.32	0.72 0.10, 0.25 -0.87, 1.53 -1.60, 1.12 -0.15, -0.01 -0.15, 0.05 -1.67, 2.32 -4.50, -0.36	<0.001 0.588 0.725 0.036 0.313 0.751	0.07 0.15 0.02 -0.01 0.03 0.1	0.21 0.04, 0.10 -0.28, 0.57 -0.46, 0.50 -0.04, 0.01 -0.00, 0.06 -0.61, 0.80 -1.75, -0.29	<0.001 0.496 0.939 0.335 0.084 0.789	-0.4 1.58 -0.92 0.09 -0.07 -2.63	-0.08 -0.56, -0.24 -0.96, 4.11 -3.79, 1.95 -0.06, 0.24 -0.26, 0.12 -6.55, 1.29 -1.41, 6.77	<0.001 0.221 0.53 0.248 0.467 0.188	

^{1.} Beta-coefficient from regression analysis adjusting for covariates as described in the Methods.

^{2. 95%} confidence intervals

eTable 3. Covariate-adjusted regression analysis using *area deprivation index* in **2018** as the exposure and the median scores of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020.

		MSRS-R				PDDS		PROMIS			
	Predictors	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	
	ADI	0.63	-0.90, -0.36	<0.001	0.12	-0.21, -0.03	0.011	-0.6	0.00, 1.19	0.048	
	Age of latest PRO	0.16	0.11, 0.22	<0.001	0.09	0.07, 0.10	<0.001	-0.5	-0.63, -0.38	<0.001	
	Sex, Male	-0.01	-0.78, 0.75	0.972	0.55	0.29, 0.81	<0.001	-0.31	-2.05, 1.42	0.722	
orts	Sex, Other	3.62	-8.18, 15.41	0.547	0.72	-3.27, 4.70	0.724	-3.64	-22.81, 15.53	0.709	
, Coho	Race and Ethnicity, Other	1.29	0.17, 2.42	0.025	0.11	-0.27, 0.50	0.566	0.03	-3.09, 3.14	0.987	
Discovery Cohorts	Age of First Symptom Onset	-0.08	-0.13, -0.02	0.005	-0.04	-0.06, -0.02	<0.001	0.19	0.07, 0.32	0.003	
Disc	Disease Duration	0	-0.06, 0.06	0.92	0	-0.02, 0.02	0.634	0.03	-0.11, 0.17	0.714	
	DMT Group, High Efficacy	0.94	0.02, 1.86	0.046	0.55	0.24, 0.87	0.001	-2.36	-4.43, -0.30	0.025	
	DMT Group, Standard Efficacy	-1.44	-2.42, -0.47	0.004	-0.37	-0.70, -0.04	0.03	2.02	-0.12, 4.16	0.064	
	Observations	1248			1234			645			
	R2 / R2 adjusted	0.104 / 0.09			0.208 / 0.20)2		0.246 / 0.235			
	ADI	0.53	-0.79, - 0.26	<0.001	0.14	-0.23, -0.04	0.004	-0.71	0.17, 1.26	0.011	
	Age of latest PRO	0.17	0.09, 0.25	<0.001	0.07	0.04, 0.10	<0.001	-0.39	-0.55, -0.24	<0.001	
	Sex, Male	0.39	-0.80, 1.59	0.517	0.16	-0.26, 0.59	0.446	1.47	-1.07, 4.00	0.256	
horts	Race and Ethnicity, Other	-0.26	-1.61, 1.09	0.707	0.01	-0.46, 0.49	0.963	-0.83	-3.69, 2.03	0.57	
Replication Cohorts	Age of First Symptom Onset	-0.08	-0.15, -0.00	0.042	-0.01	-0.04, 0.01	0.358	0.08	-0.06, 0.23	0.264	
plicati	Disease Duration	-0.04	-0.14, 0.05	0.36	0.03	-0.00, 0.06	0.071	-0.08	-0.27, 0.11	0.424	
Re	DMT Group, High Efficacy	0.33	-1.66, 2.32	0.748	0.1	-0.60, 0.80	0.787	-2.64	-6.55, 1.28	0.186	
	DMT Group, Standard Efficacy	-2.43	-4.50, -0.37	0.021	-1.02	-1.75, -0.29	0.006	2.68	-1.40, 6.76	0.198	
	Observations 421 R2 / R2 adjusted 0.157 / 0.140				416			378			
	1 	0.157 / 0.14	10	·	0.258 / 0.24		·	0.217 / 0.200			

^{1.} Beta-coefficient from regression analysis adjusting for covariates as described in the Methods.

^{2. 95%} confidence intervals

eTable 4. Covariate-adjusted regression analysis of *neighborhood median household income* in 2015 as the exposure and the median score of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020 from the joint cohorts.

		MSRS-R			PDDS		PROMIS			
Predictors	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	
Median Household Income	-3.80e-05	-5.05e-05, -2.56e-05	<0.001	-8.58e-06	-1.28e-05, -4.32e-06	0.045	2.55e-05	5.96e-07, 5.05e-05	<0.001	
Age of Latest PRO	0.16	0.12, 0.21	<0.001	-0.46	-0.56, -0.36	<0.001	0.08	0.07, 0.10	<0.001	
Sex, Male	0.09	-0.57, 0.74	0.791	0.28	-1.14, 1.71	0.698	0.47	0.25, 0.69	<0.001	
Sex, Other	3.46	-8.07, 14.99	0.556	-4.04	-23.56, 15.47	0.684	0.84	-3.08, 4.75	0.674	
Race and Ethnicity, Other	0.45	-0.43, 1.33	0.316	-0.4	-2.42, 1.63	0.7	0.01	-0.29, 0.31	0.958	
Age of First Symptom Onset	-0.08	-0.12, -0.03	0.001	0.15	0.05, 0.24	0.003	-0.03	-0.05, -0.01	<0.001	
Disease Duration	-0.01	-0.06, 0.04	0.575	-0.01	-0.12, 0.10	0.844	0.01	-0.01, 0.03	0.245	
DMT Group, High Efficacy	0.63	-0.19, 1.46	0.131	-1.99	-3.78, -0.20	0.03	0.44	0.16, 0.72	0.002	
DMT Group, Standard Efficacy	-1.81	-2.68, -0.94	<0.001	2.65	0.76, 4.53	0.006	-0.52	-0.82, -0.22	0.001	
Observations	1665			1646			1020			
R ² / R ² adjusted	usted 0.118 / 0.113				19		0.227 / 0.220			

- 1. Beta-coefficient from regression analysis adjusting for covariates as described in the Methods.
- 2. 95% confidence intervals

eTable 5. Covariate-adjusted regression analysis of *neighborhood median home value* in 2015 as the exposure and the median score of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020 from the joint cohorts.

		MSRS-R			PDDS		PROMIS			
Predictors	Estimate ¹	95% CI ²	P-	Estimate ¹	95% CI ²	P-	Estimate ¹	95% CI ²	P-	
			value			value			value	
Median	-6.50e-06	-8.16e-06,	<0.001	-1.54e-06	-2.11e-06,	0.002	4.98e-06	1.81e-06,	<0.001	
Home Value		-4.84e-06			-9.65e-07			8.14e-06		
Age of Latest PRO	0.16	0.11, 0.20	<0.001	-0.45	-0.55, -0.35	<0.001	0.08	0.07, 0.10	<0.001	
Sex, Male	0.14	-0.51, 0.79	0.674	0.22	-1.20, 1.64	0.761	0.48	0.26, 0.70	<0.001	
Sex, Other	5.84	-5.62, 17.31	0.318	-5.9	-25.40, 13.60	0.553	1.4	-2.50, 5.31	0.481	
Race and Ethnicity, Other	1.2	0.33, 2.06	0.007	-1.11	-3.14, 0.91	0.281	0.18	-0.12, 0.47	0.246	
Age of First Symptom Onset	-0.07	-0.12, -0.03	0.001	0.14	0.05, 0.24	0.004	-0.03	-0.04, -0.01	<0.001	
Disease Duration	-0.01	-0.06, 0.04	0.77	-0.02	-0.13, 0.09	0.704	0.01	-0.01, 0.03	0.175	
DMT Group, High Efficacy	0.76	-0.06, 1.58	0.068	-2.19	-3.98, -0.39	0.017	0.47	0.19, 0.75	0.001	
DMT Group, Standard Efficacy	-1.72	-2.58, -0.86	<0.001	2.51	0.64, 4.39	0.009	-0.49	-0.79, -0.20	0.001	
Observations	1666			1647			1021			
R ² / R ² adjusted	0.130 / 0.12	25		0.230 / 0.22	25	-	0.232 / 0.225			

- 1. Beta-coefficient from regression analysis adjusting for covariates as described in the Methods.
- 2. 95% confidence intervals

eTable 6 Covariate-adjusted regression analysis of *neighborhood percentage of population with college education* in 2015 as the exposure and the median score of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020 from the joint cohorts.

		MSRS-R			PDDS		PROMIS			
Predictors	Estimate ¹	95% CI ²	P-	Estimate ¹	95% CI ²	P-	Estimate ¹	95% CI ²	P-	
			value			value			value	
Percent with College Education	-7.31	-8.99, -5.64	<0.001	-1.62	-2.20, -1.05	<0.001	9.31	5.73, 12.89	<0.001	
Age of Latest PRO	0.17	0.12, 0.21	<0.001	0.08	0.07, 0.10	<0.001	-0.45	-0.55, -0.35	<0.001	
Sex, Male	0.16	-0.48, 0.81	0.62	0.48	0.26, 0.71	<0.001	0.03	-1.38, 1.44	0.966	
Sex, Other	5.27	-6.14, 16.68	0.365	1.24	-2.66, 5.13	0.534	-6.52	-25.85, 12.80	0.508	
Race and Ethnicity, Other	0.76	-0.10, 1.62	0.083	0.08	-0.22, 0.37	0.618	-0.45	-2.44, 1.54	0.656	
Age of First Symptom Onset	-0.07	-0.12, -0.03	0.001	-0.03	-0.04, -0.01	<0.001	0.13	0.04, 0.23	0.007	
Disease Duration	-0.01	-0.06, 0.04	0.768	0.01	-0.01, 0.03	0.181	-0.03	-0.14, 0.08	0.596	
DMT Group, High Efficacy	0.74	-0.08, 1.55	0.077	0.46	0.18, 0.74	0.001	-2.41	-4.19, -0.63	0.008	
DMT Group, Standard Efficacy	-1.74	-2.60, -0.88	<0.001	-0.5	-0.80, -0.21	0.001	2.19	0.32, 4.06	0.022	
Observations	1668			1649			1022			
R ² / R ² adjusted	0.138 / 0.13	33		0.231 / 0.2	26		0.245 / 0.238			

^{1.} Beta-coefficient from regression analysis adjusting for covariates as described in the Methods.

^{2. 95%} confidence intervals

eTable 7. Covariate-adjusted regression analysis of *neighborhood percentage of population with more than one occupant per room* (indicating a crowded household) in 2015 as the exposure and the median score of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020 from the joint cohorts.

		MSRS-R			PDDS		PROMIS			
Predictors	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	
Percent Living in Crowded Households	-16.83	-28.78, -4.89	0.006	-3.86	-7.91, 0.18	0.061	-3.94	-27.30, 19.41	0.741	
Age of Latest PRO	0.17	0.12, 0.21	<0.001	0.08	0.07, 0.10	<0.001	-0.47	-0.56, - 0.37	<0.001	
Sex, Male	-0.05	-0.71, 0.60	0.875	0.44	0.21, 0.66	<0.001	0.47	-0.95, 1.89	0.518	
Sex, Other	3.7	-7.93, 15.33	0.533	0.89	-3.04, 4.82	0.656	-3.67	-23.24, 15.89	0.713	
Race and Ethnicity, Other	1.32	0.41, 2.24	0.005	0.2	-0.11, 0.52	0.199	-0.51	-2.68, 1.66	0.645	
Age of First Symptom Onset	-0.08	-0.12, -0.03	0.001	-0.03	-0.05, -0.02	<0.001	0.15	0.05, 0.25	0.002	
Disease Duration	-0.02	-0.07, 0.03	0.539	0.01	-0.01, 0.03	0.263	-0.01	-0.12, 0.11	0.895	
DMT Group, High Efficacy	0.48	-0.35, 1.31	0.254	0.41	0.12, 0.69	0.005	-1.76	-3.54, 0.03	0.053	
DMT Group, Standard Efficacy	-2.03	-2.91, -1.16	<0.001	-0.57	-0.87, -0.27	<0.001	2.86	0.99, 4.74	0.003	
Observations	1668			1649			1022			
R ² / R ² adjusted	0.104 / 0.099			0.218 / 0.214			0.226 / 0.219			

- 1. Beta-coefficient from regression analysis adjusting for covariates as described in the Methods.
- 2. 95% confidence intervals

eTable 8. Covariate-adjusted regression analysis of *neighborhood percentage of population without high school education* in 2015 as the exposure and the median score of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020 from the joint cohorts.

		MSRS-R			PDDS		PROMIS			
Predictors	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	
Percent without High School Education	4.14	-1.26, 9.54	0.133	0.82	-1.01, 2.64	0.381	-11.39	-22.25, -0.52	0.04	
Age of Latest PRO	0.17	0.13, 0.22	<0.001	0.08	0.07, 0.10	<0.001	-0.47	-0.57, -0.37	<0.001	
Sex, Male	-0.04	-0.70, 0.62	0.901	0.44	0.22, 0.66	<0.001	0.39	-1.03, 1.81	0.59	
Sex, Other	3.22	-8.42, 14.86	0.587	0.78	-3.15, 4.71	0.697	-4.22	-23.73, 15.29	0.672	
Race and Ethnicity, Other	0.72	-0.20, 1.63	0.126	0.07	-0.24, 0.38	0.654	0.1	-2.03, 2.24	0.923	
Age of First Symptom Onset	-0.08	-0.13, -0.04	<0.001	-0.03	-0.05, - 0.02	<0.001	0.15	0.05, 0.25	0.002	
Disease Duration	-0.02	-0.07, 0.03	0.514	0.01	-0.01, 0.03	0.273	-0.01	-0.12, 0.10	0.861	
DMT Group, High Efficacy	0.44	-0.39, 1.27	0.3	0.4	0.11, 0.68	0.006	-1.79	-3.57, -0.01	0.049	
DMT Group, Standard Efficacy	-2.02	-2.90, -1.15	<0.001	-0.57	-0.87, - 0.27	<0.001	2.79	0.92, 4.66	0.003	
Observations	1668			1649			1022			
R ² / R ² adjusted	0.101 / 0.096			0.217 / 0.212			0.228 / 0.222			

¹ Beta-coefficient from regression analysis adjusting for covariates as described in the Methods.

^{2 95%} confidence intervals

eTable 9. Covariate-adjusted regression analysis of *neighborhood percentage of population unemployed* in 2015 as the exposure and the median score of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020 from the joint cohorts.

		MSRS-R			PDDS		PROMIS			
Predictors	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	
Percent Unemployed	6.36	-10.09, 22.81	0.448	-0.22	-5.81, 5.36	0.937	-2.46	-37.74, 32.81	0.891	
Age of Latest PRO	0.17	0.13, 0.22	<0.001	0.08	0.07, 0.10	<0.001	-0.46	-0.56, -0.37	<0.001	
Sex, Male	-0.05	-0.71, 0.61	0.88	0.43	0.21, 0.66	<0.001	0.46	-0.96, 1.88	0.524	
Sex, Other	3.03	-8.62, 14.67	0.61	0.75	-3.17, 4.68	0.706	-3.79	-23.34, 15.76	0.704	
Race and Ethnicity, Other	0.82	-0.09, 1.74	0.077	0.12	-0.19, 0.43	0.462	-0.61	-2.69, 1.47	0.564	
Age of First Symptom Onset	-0.08	-0.12, -0.03	0.001	-0.03	-0.05, -0.02	<0.001	0.15	0.05, 0.25	0.002	
Disease Duration	-0.02	-0.07, 0.03	0.527	0.01	-0.01, 0.03	0.277	-0.01	-0.12, 0.10	0.884	
DMT Group, High Efficacy	0.45	-0.38, 1.28	0.291	0.4	0.11, 0.68	0.006	-1.77	-3.56, 0.01	0.052	
DMT Group, Standard Efficacy	-2.03	-2.91, -1.16	<0.001	-0.57	-0.87, -0.27	<0.001	2.86	0.99, 4.73	0.003	
Observations	1668			1649			1022			
R ² / R ² adjusted	0.100 / 0.09	95		0.216 / 0.212			0.226 / 0.219			

- 1. Beta-coefficient from regression analysis adjusting for covariates as described in the Methods.
- 2. 95% confidence intervals

eTable 10. Covariate-adjusted regression analysis of *neighborhood percentage of population living below the poverty line* in 2015 as the exposure and the median score of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020 from the joint cohorts.

		MSRS-R			PDDS		PROMIS			
Predictors	Estimate ¹	95% Cl ²	P-value	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	
Percent Below Poverty Line	1.54	-2.00, 5.09	0.393	0.23	-0.97, 1.43	0.706	1.14	-6.63, 8.92	0.773	
Age of Latest PRO	0.17	0.13, 0.22	<0.001	0.08	0.07, 0.10	<0.001	-0.46	-0.56, -0.37	<0.001	
Sex, Male	-0.05	-0.71, 0.61	0.876	0.44	0.21, 0.66	<0.001	0.48	-0.94, 1.90	0.509	
Sex, Other	2.87	-8.78, 14.53	0.629	0.72	-3.21, 4.65	0.719	-3.97	-23.54, 15.61	0.691	
Race and Ethnicity, Other	0.8	-0.12, 1.72	0.088	0.09	-0.22, 0.41	0.554	-0.73	-2.81, 1.36	0.495	
Age of First Symptom Onset	-0.08	-0.12, -0.03	0.001	-0.03	-0.05, -0.02	<0.001	0.15	0.05, 0.25	0.002	
Disease Duration	-0.02	-0.07, 0.03	0.516	0.01	-0.01, 0.03	0.273	-0.01	-0.12, 0.10	0.892	
DMT Group, High Efficacy	0.45	-0.38, 1.28	0.283	0.4	0.12, 0.68	0.006	-1.75	-3.54, 0.04	0.055	
DMT Group, Standard Efficacy	-2.02	-2.89, -1.14	<0.001	-0.57	-0.87, -0.27	<0.001	2.89	1.01, 4.77	0.003	
Observations	1668			1649			1022			
R ² / R ² adjusted	0.100 / 0.09	95		0.216 / 0.2	12		0.226 / 0.219			

- 1. Beta-coefficient from regression analysis adjusting for covariates as described in the Methods.
- 2. 95% confidence intervals

eTable 11. Covariate-adjusted regression analysis using *neighborhood percent of population with private insurance* in 2015 as the exposure and the median scores of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020 from the joint cohorts.

		MSRS-R			PDDS		PROMIS			
Predictors	Estimate ¹	95% CI ²	P-Value	Estimate ¹	95% CI ²	P-Value	Estimate ¹	95% CI ²	P-Value	
Percent with Private Insurance	1.2	-3.03, 5.42	0.578	0.68	-0.77, 2.12	0.356	-0.34	-9.23, 8.54	0.94	
ADI	0.71	0.40, 1.02	<0.001	0.18	0.08, 0.29	0.001	-0.65	-1.28, -0.02	0.043	
Age of Latest PRO	0.17	0.12, 0.21	<0.001	0.08	0.07, 0.10	<0.001	-0.46	-0.56, -0.36	<0.001	
Sex, Male	0.12	-0.53, 0.78	0.709	0.48	0.25, 0.70	<0.001	0.2	-1.22, 1.63	0.779	
Sex, Other	4.09	-7.40, 15.59	0.485	0.97	-2.94, 4.88	0.626	-4.77	-24.24, 14.69	0.63	
Race and Ethnicity, Other	0.41	-0.48, 1.30	0.366	0.01	-0.29, 0.32	0.931	-0.21	-2.29, 1.87	0.841	
Age of First Symptom Onset	-0.07	-0.12, -0.03	0.001	-0.03	-0.04, -0.01	<0.001	0.14	0.05, 0.24	0.004	
Disease Duration	-0.01	-0.06, 0.04	0.674	0.01	-0.01, 0.03	0.204	-0.02	-0.13, 0.09	0.734	
DMT Group, High Efficacy	0.68	-0.14, 1.50	0.106	0.45	0.17, 0.73	0.002	-2.17	-3.96, -0.38	0.018	
DMT Group, Standard Efficacy	-1.76	-2.63, -0.89	<0.001	-0.51	-0.81, -0.21	0.001	2.43	0.55, 4.31	0.011	
Observations	1668			1649			1022			
R ² / R ² adjusted	0.124 / 0.1	19		0.226 / 0.22	21		0.233 / 0.226			

^{1.} Beta-coefficient from regression analysis adjusting for covariates as described in the Methods. These analyses *additionally* adjust for the area deprivation index (**ADI**) in 2015.

^{2. 95%} confidence intervals

eTable 12. Covariate-adjusted regression analysis using *neighborhood percent of population with Medicaid insurance* in 2015 as the exposure and the median scores of patient-reported outcomes of neurological and physical function from the latest year between 2018 and 2020 from the joint cohorts.

		MSRS-R			PDDS		PROMIS			
Predictors	Estimate ¹	95% CI ²	P-Value	Estimate ¹	95% CI ²	P-Value	Estimate ¹	95% CI ²	P-Value	
Percent with Medicaid Insurance	-9.53	-15.39, -3.67	0.001	-2.37	-4.38, -0.36	0.021	5.38	-7.06, 17.82	0.396	
ADI	0.96	0.69, 1.23	<0.001	0.22	0.13, 0.32	<0.001	-0.81	-1.36, -0.25	0.004	
Age of Latest PRO	0.16	0.12, 0.21	<0.001	0.08	0.07, 0.10	<0.001	-0.46	-0.55, -0.36	<0.001	
Sex, Male	0.15	-0.50, 0.80	0.649	0.48	0.26, 0.70	<0.001	0.19	-1.23, 1.61	0.793	
Sex, Other	4.18	-7.28, 15.64	0.475	1	-2.91, 4.90	0.616	-4.8	-24.25, 14.66	0.629	
Race and Ethnicity, Other	0.8	-0.11, 1.72	0.086	0.1	-0.22, 0.41	0.545	-0.52	-2.67, 1.64	0.638	
Age of First Symptom Onset	-0.07	-0.12, -0.03	0.001	-0.03	-0.04, -0.01	<0.001	0.14	0.04, 0.24	0.004	
Disease Duration	-0.01	-0.06, 0.04	0.714	0.01	-0.01, 0.03	0.195	-0.02	-0.13, 0.09	0.72	
DMT Group, High Efficacy	0.74	-0.08, 1.56	0.077	0.46	0.18, 0.75	0.001	-2.21	-4.00, -0.42	0.016	
DMT Group, Standard Efficacy	-1.73	-2.59, -0.86	<0.001	-0.5	-0.80, -0.20	0.001	2.42	0.54, 4.30	0.012	
Observations	1668			1649			1022			
R ² / R ² adjusted	0.129 / 0.12	24		0.228 / 0.223			0.234 / 0.227			

^{1.} Beta-coefficient from regression analysis adjusting for covariates as described in the Methods. These analyses *additionally* adjust for the area deprivation index (**ADI**) in 2015.

^{2. 95%} confidence intervals

eTable 13. Covariate-adjusted regression analysis using nine *individual household income brackets* ranging from "\$0 to \$19,999" to "\$125,000 or higher" as exposure and the patient-reported outcomes of neurological and physical function for each participant in subgroups of the joint cohorts that accounts for the area deprivation index.

	MSRS-R			PDDS			PROMIS		
Predictors	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value	Estimate ¹	95% CI ²	P-value
Individual Household Income Bracket	-0.71	-0.92, -0.50	<0.001	-0.25	-0.33, -0.17	<0.001	1.29	0.87, 1.71	<0.001
Age of latest PRO	0.25	0.16, 0.34	<0.001	0.1	0.07, 0.13	<0.001	-0.51	-0.69, -0.34	<0.001
Sex, Male	-0.09	-1.53, 1.36	0.905	0.55	0.01, 1.09	0.047	-0.41	-3.25, 2.42	0.776
Sex, Other	0.98	-8.95, 10.91	0.846	1.06	-2.67, 4.79	0.576	-8.4	-26.90, 10.10	0.372
Race and Ethnicity, Other	1.19	-1.15, 3.52	0.317	-0.31	-1.19, 0.56	0.484	0.74	-3.89, 5.38	0.752
Age of First Symptom Onset	-0.19	-0.27, -0.10	<0.001	-0.06	-0.09, -0.02	0.001	0.24	0.08, 0.41	0.004
Disease Duration	-0.13	-0.23, -0.02	0.018	-0.03	-0.07, 0.01	0.172	0.09	-0.11, 0.29	0.365
DMT Group, High Efficacy	1.27	-0.33, 2.86	0.119	0.45	-0.15, 1.05	0.141	-1.91	-4.96, 1.15	0.22
DMT Group, Standard Efficacy	0.39	-1.31, 2.08	0.653	-0.18	-0.82, 0.45	0.574	2.72	-0.57, 6.01	0.105
ADI	0.19	-0.22, 0.60	0.365	-0.02	-0.18, 0.13	0.756	-0.19	-1.00, 0.62	0.648
Observations	320			320			284		
R ² / R ² adjusted	0.264 / 0.240			0.306 / 0.284			0.342 / 0.318		

^{1.} Beta-coefficient from regression analysis adjusting for covariates as described in the Methods. These analyses *additionally* adjust for the area deprivation index (**ADI**) in 2015.

^{2. 95%} confidence intervals