	Large lesion volume subgroup	Small lesion volume subgroup	Test statistic	Significance (p-value)
Ν	32	70		
age (median, range) (years)	32.8 (20.7 - 49.7)	33.2 (20.1 - 56.2)	t = -0.544	0.588
sex = M (%)	11 (34%)	23 (32%)	$\chi^2 = 0$	1
Disease duration (median, range) (months)	4 (0 - 13)	4 (1 - 8)	W = 1085.5	1
Diagnosis at baseline = CIS (%)	0 (0%)	28 (40%)	χ ² = 15.692	< 0.0001
Diagnosis at final visit = CIS (%)	0 (0%)	16 (22.9%)	χ ² = 7.033	0.008
Follow-up duration (median, range) (months)	33.57 (3.44 - 70.5)	21.99 (5.69 - 68.8)	W = 1332	0.097
Baseline EDSS (median, range)	1.5 (0 - 3)	1.5 (0 - 3.5)	χ ² = 7.527	0.275
Baseline T2 lesion volume (ml) (mean)	5.83 ± 4.22	0.67 ± 0.10	W = 2240	< 0.0001
Baseline Cortical thickness (mm) (mean)	2.61 ± 0.12	2.65 ± 0.10	t = -1.497	0.140
Baseline NAWM sT1w/T2w (mean)	0.35 ± 0.04	0.37 ± 0.03	t = -3.140	0.003
Fulfilled NEDA-3 criteria during follow-up (%)	5 (15.6%)	18 (25.7%)	χ ² = 2.61	0.11

Supplementary Table 1. Demographic, imaging and clinical characteristics of the large lesion and small lesion subgroups.

	Model 3
age at baseline	-0.001 ± 0.0004
coefficient (95% CI)	
follow-up time coefficient (95% CI)	0.0003 ± 0.001
lesion volume coefficient (95% CI)	-0.008 ± 0.003**
cortical thickness coefficient (95% CI)	0.021 ± 0.015*
Interaction between lesion volume and follow-up time coefficient	0.0001 ±
(95% CI)	0.00005**
Interaction between cortical thickness and follow-up time	-0.0002 ± 0.0004
coefficient (95% CI)	
intercept (95% CI)	0.389 ± 0.045***
Number of patients	102
Number observations	455

marginal R ²	0.054
conditional R ²	0.834

Supplementary Table 2. Association between eroded sT1w/T2w and other imaging

parameters. *p<0.1**p<0.05 ***p<0.01

	Model 3
age at baseline	-0.0005 ±
coefficient (95% CI)	0.0004
follow-up time coefficient (95% CI)	-0.0004 ± 0.001
lesion volume coefficient (95% CI)	-0.010 ± 0.003***
cortical thickness coefficient (95% CI)	0.028 ± 0.014*
Interaction between lesion volume and follow-up time coefficient	0.0001 ±
(95% CI)	0.00005

Interaction between cortical thickness and follow-up time	-0.0001 ±
coefficient (95% CI)	0.0003
intercept (95% CI)	0.371 ±
	0.043***
Number of patients	102
Number observations	356
marginal R ²	0.082
conditional R ²	0.896

Supplementary Table 3. Association between eroded sT1w/T2w, other imaging

parameters and NEDA-3. *p<0.1**p<0.05 ***p<0.01