

Stability in Cognition and Employment in People with Relapsing Multiple Sclerosis Treated with Cladribine Tablets: Two-year Phase IV CLARIFY-MS Study

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Supplementary Appendix

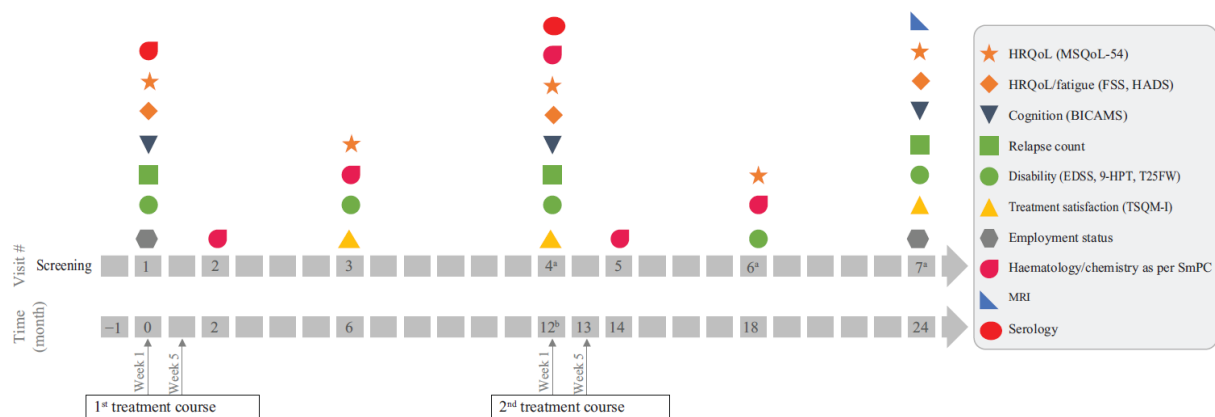


Figure S1: CLARIFY-MS study design

9-HPT, Nine-Hole Peg Test; BICAMS, Brief International Cognitive Assessment for Multiple Sclerosis; EDSS, Expanded Disability Status Scale; FSS, Fatigue Severity Scale; HADS, Hospital Anxiety and Depression Scale; HRQoL, health-related quality of life; MRI, magnetic resonance imaging; MSQoL-54, Multiple Sclerosis Quality of Life-54 instrument; RMS, relapsing multiple sclerosis; SPC, summary of product characteristics; T25FW, Timed 25-Foot Walk; TSQM, Treatment Satisfaction Questionnaire for Medication.

^aThere was a 0–3-month window for these trial visits. ^bThe second treatment course may have been delayed for some participants.

Figure adapted from: Brochet B, et al. *Mult Scler Relat Disord* 2022. (CC BY)

Table S1: Participant baseline demographics and disease characteristics

	Tx-naïve n = 134	Tx-experienced n = 348	Total N = 482
Mean age ± SD, years	35.2 ± 11.3	38.3 ± 9.9	37.4 ± 10.4
Female, n (%)	89 (66.4)	249 (71.6)	338 (70.1)
Mean time since onset of MS ± SD, months	42.6 ± 58.7	120.9 ± 90.3	99.1 ± 89.8
Mean time since diagnosis of MS ± SD, months	16.40 ± 32.97	95.62 ± 76.47	73.59 ± 76.04
No. of relapses within 12 months before baseline visit, n (%)			
0	0 (0.0)	4 (1.1)	4 (0.8)
1	35 (26.1)	225 (64.7)	260 (53.9)
≥2	99 (73.9)	119 (34.2)	218 (45.2)
EDSS, median (range)	2.0 (0.0–5.0)	2.5 (0.0–5.0)	2.5 (0.0–5.0)
DMT use in prior 6 months to start of study, n (%)	0 (0.0)	287 (82.5)	287 (59.5)
BICAMS (BICAMS set)	n = 108	n = 291	N = 399
SDMT, median (Q1; Q3)	52.5 (45.0; 60.0)	50.0 (41.5; 58.0)	51.0 (43.0; 59.0)
CVLT-II, median (Q1; Q3)	56.0 (46.0; 62.0)	55.5 (45.0; 64.0)	56.0 (45.0; 63.0)
BVMT-R, median (Q1; Q3)	24.0 (18.0; 28.0)	23.5 (17.0; 29.0)	24.0 (17.0; 29.0)

BICAMS, Brief International Cognitive Assessment for Multiple Sclerosis;

BVMT-R, Brief Visuospatial Memory Test–Revised; CVLT-II, California Verbal Learning Test

Second Edition; DMT, disease-modifying therapy; EDSS, Expanded Disability Status Scale;

MS, multiple sclerosis; Q, quartile; SD, standard deviation; SDMT, Symbol Digit Modalities Test;

Tx, treatment.

Table S2: SDMT scores in the TS

Visit	Statistics	Absolute	Absolute change from baseline
Total			
N = 482			
Baseline	n (%)	472 (97.9)	-
	Mean (SD)	50.7 (12.9)	-
	Median	51.0	-
	Q1; Q3	43.0; 59.0	-
M12	n (%)	459 (96.8)	451 (95.1)
	Mean (SD)	52.0 (14.0)	1.4 (9.1)
	Median	52.0	2.0
	Q1; Q3	44.0; 61.0	-2.0; 6.0
M24	n (%)	427 (92.6)	420 (91.1)
	Mean (SD)	53.0 (13.9)	2.0 (8.4)
	Median	54.0	2.0
	Q1; Q3	45.0; 62.0	-2.0; 6.0

M, month; Q, quartile; SD, standard deviation; SDMT, Symbol Digit Modalities Test; TS, treated set.

Table S3: Shift of change categories in cognitive function from M12 to M24 versus CLARIFY-MS baseline to M12 – 4-point individual-level difference in SDMT scores

Baseline to M12	Change from M12 to M24 in SDMT scores				
	Total (N = 482)				
	Increase n (%)	Stable n (%)	Decrease n (%)	Missing n (%)	Total n (%)
Increase	24 (5.0)	68 (14.1)	55 (11.4)	17 (3.5)	164 (34.0)
Stable	55 (11.4)	92 (19.1)	32 (6.6)	15 (3.1)	194 (40.2)
Decrease	49 (10.2)	26 (5.4)	13 (2.7)	5 (1.0)	93 (19.3)
Missing	3 (0.6)	3 (0.6)	0	25 (5.2)	31 (6.4)
Total	131 (27.2)	189 (39.2)	100 (20.7)	62 (12.9)	482 (100.0) ^a

^aDue to rounding to one decimal digit in each category, percentages may not sum up to 100.0% in total.

Increase, change in SDMT ≥ 4 ; Stable, $-4 < \text{change in SDMT} < 4$; Decrease, change in SDMT ≤ -4 .
M, month; SDMT, Symbol Digit Modalities Test.

Table S4: Shift of change categories in cognitive function from M12 to M24 versus CLARIFY-MS baseline to M12 – 8-point individual-level difference in SDMT scores

Baseline to M12	Change from M12 to M24 in SDMT scores				
	Total (N = 482)				
	Increase n (%)	Stable n (%)	Decrease n (%)	Missing n (%)	Total n (%)
Increase	4 (0.8)	55 (11.4)	16 (3.3)	11 (2.3)	86 (17.8)
Stable	30 (6.2)	246 (51.0)	25 (5.2)	24 (5.0)	325 (67.4)
Decrease	14 (2.9)	22 (4.6)	2 (0.4)	2 (0.4)	40 (8.3)
Missing	1 (0.2)	5 (1.0)	0	25 (5.2)	31 (6.4)
Total	49 (10.2)	328 (68.0)	43 (8.9)	62 (12.9)	482 (100.0) ^a

^aDue to rounding to one decimal digit in each category, percentages may not sum up to 100.0% in total.

Increase, change in SDMT ≥ 4 ; Stable, $-4 < \text{change in SDMT} < 4$; Decrease, change in SDMT ≤ -4 .
M, month; SDMT, Symbol Digit Modalities Test.

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