Supplementary Material

Supplementary Table 1: Time-related principles derived from the Brain health: time matters in multiple sclerosis³ policy report recommendations

Brain health: time matters in multiple sclerosis recommendation	Time-related principles
Educate family and primary care physicians about the importance of promptly referring people with suspected MS to a neurologist, and ultimately to a specialist clinic, to speed up diagnosis and treatment initiation	 Reporting of symptoms to a healthcare professional without delay Prompt or urgent referral to a neurologist after first report of MS-related symptoms to a healthcare professional
Improve access to specialist care for MS: make diagnostic and monitoring procedures more widely accessible, increase the numbers of healthcare professionals who specialise in the management of MS, and ensure that these specialists provide prompt diagnostic and support services for people with suspected MS and those who have been newly diagnosed with the disease	 Prompt MRI scan after referral to a neurologist Rapid diagnosis after referral to a neurologist Timely offer of cognitive testing after MS diagnosis
Adopt the latest accepted diagnostic criteria, in order to diagnose MS as early as possible	No principle for Round 1 was derived from this recommendation, it was developed into a consensus statement for Round 4.
Ensure that MS healthcare professionals can take the time to educate people with MS about strategies to manage their disease. Emphasise the importance of a 'brain-healthy' lifestyle, the benefits of early treatment with therapies that can modify the disease course, the likely consequences of inadequate or suboptimal treatment and the goal of minimizing disease activity while optimizing safety	 Extended appointment with neurologist following diagnosis to discuss implications of MS diagnosis Early discussion about the importance of living a brainhealthy lifestyle Regular reminders about living a brainhealthy lifestyle Regular check-ups (with a primary care physician or at a neurology clinic) to manage comorbidities
Implement a shared decision-making process that embodies dialogue between people with MS and healthcare professionals. A well-informed and proactive collaboration between people with MS and their healthcare team is vital to successful management of the disease	 Early discussion with patient about the benefits of early treatment with therapies that can modify the disease course Early inclusion of patients in informed, shared decision making Early discussion with patient about the aims of treatment
Make the full range of disease-modifying therapies available to people with active relapsing forms of MS, regardless of their treatment history, to speed up adoption of the most appropriate treatment strategy that optimizes effectiveness and safety for each individual	 Evaluation of suitability/eligibility for treatment shortly after MS diagnosis Regular review of whether patient meets eligibility criteria for a DMT Prompt offer of treatment with a DMT after patient becomes eligible (based on national guidelines)
Include evidence from monitoring via regular clinical evaluation and scheduled/unscheduled MRI brain scans in any definitions of disease activity or suboptimal response, in order to assist in the rapid identification of treatment failure and the decision to switch treatment	Regular MRI scans
Ensure that MS healthcare professionals can take the time to monitor disease activity in people with MS	Regular clinical evaluations
Agree and implement standardized data collection techniques, protocols and data sets, nationally and internationally, to track clinical and subclinical events in routine practice. Incorporate these into a clinical management tool to facilitate individualized practice	• Regular inclusion of patient data in MS database
Maintain treatment with a disease-modifying therapy for as long as a person with MS would be at risk of inflammatory disease activity if they were not receiving treatment; in the case of a suboptimal response, make a prompt decision about whether or not to switch therapy	 Regular review of current treatment/treatment options Rapid offer of an alternative DMT when response to current DMT is suboptimal Regular review of the aims of treatment

MRI, magnetic resonance imaging; MS, multiple sclerosis.

Supplementary Table 2: Variables that reflect the timing of events in the MS care pathway, derived from the principles of brain health-focused care agreed in round 1

Principle	Va	riable
Reporting of symptoms to a healthcare professional without delay	1.	Time <u>from</u> person experiencing first symptoms possibly related to MS pre-diagnosis to reporting the symptoms to any healthcare professional
Prompt or urgent referral to a neurologist after first report of MS-related symptoms to a healthcare professional	2.	Time <u>from</u> person first reporting symptoms possibly related to MS to a healthcare professional pre-diagnosis <u>to</u> referral to a neurologist
Prompt MRI scan after referral to a neurologist	3.	Time <u>from</u> referral to a neurologist by a healthcare professional to first MRI scan (if MRI scan was not performed before referral)
	4.	Time <u>from</u> referral to a neurologist by a healthcare professional <u>to</u> completion of diagnostic workup
Rapid diagnosis after referral to a neurologist	5. 6.	Time <u>from</u> completion of diagnostic workup <u>to</u> appointment to discuss results Time <u>from</u> referral to a neurologist by a healthcare professional <u>to</u>
Extended appointment with neurologist following	7.	accurate MS diagnosis for a typical patient Length of appointment with neurologist following diagnosis to discuss
diagnosis to discuss implications of MS diagnosis Timely offer of cognitive testing after MS diagnosis	8.	implications of MS diagnosis Time <u>from</u> MS diagnosis to offer of cognitive screening
Early discussion about the importance of living a brain-healthy lifestyle	8. 9.	Time <u>from</u> MS diagnosis to discussion about the importance of living a brain-healthy lifestyle
	10.	Time <u>from</u> MS diagnosis to referral to additional appropriate services to support 'brain-healthy lifestyle' modifications (may not be applicable for all people with MS)
Regular reminders about living a brain-healthy lifestyle		<u>Frequency</u> of active, documented discussion about living a brain-healthy lifestyle
Regular check-ups (with a primary care physician		<u>Frequency</u> of routine reminders about living a brain-healthy lifestyle <u>Frequency</u> of check-ups (with specialist, primary care physician, or at a
or at a neurology clinic) to manage comorbidities Early discussion with patient about the benefits of		neurology clinic) to screen for and/or manage comorbidities Time <u>from</u> MS diagnosis to discussion with patients about the pros and
early treatment with therapies that can modify the disease course	14.	cons of early treatment with a DMT
Evaluation of suitability/eligibility for treatment shortly after MS diagnosis	15.	Time <u>from</u> MS diagnosis <u>to</u> evaluation of suitability/eligibility for a DMT
Regular review of whether patient meets eligibility criteria for a DMT	16.	<u>Frequency</u> of review of whether a patient currently not taking a DMT is eligible for a DMT, based on applicable guidelines
Prompt offer of treatment with a DMT after patient becomes eligible (based on national guidelines)		Time <u>from</u> patient becoming eligible for a DMT (based on applicable guidelines) to patient being offered a DMT
		Time <u>from</u> patient deciding to commence treatment with a DMT <u>to</u> starting a DMT
Early discussion with patient about the aims of treatment	19.	Time <u>from</u> MS diagnosis to discussion with patient about aims of treatment
Early inclusion of patients in informed, shared decision making		No variable for Round 2 was derived from this principle, it was developed into a consensus statement for Round 4.
Regular clinical evaluations	20.	Frequency of follow-up clinical evaluations by MS team
Regular MRI scans	21.	Frequency of MRI scans
Regular review of current treatment/treatment options	22.	<u>Frequency</u> of review of whether current DMT is effective/appropriate and consideration of alternatives as appropriate
Rapid offer of an alternative DMT when response to current DMT is suboptimal	23.	Time <u>from</u> recognition that response to current DMT is suboptimal <u>to</u> offer of an appropriate alternative DMT
Regular inclusion of patient data in MS database		No variable for Round 2 was derived from this principle, it was developed into a consensus statement for Round 4.
Regular review of the aims of treatment	24.	<u>Frequency</u> of review of treatment aims with the patient
n/a*	25.	Time <u>from</u> patient experiencing new or worsened symptoms to reporting those symptoms to an MS healthcare professional

*Derived from Delphi Panel free-text comments in Round 1.

DMT, disease-modifying therapy; MRI, magnetic resonance imaging; MS, multiple sclerosis.

Supplementary Table 3: Statements for which consensus (≥75% agreement) was not reached by the Delphi Consensus Panel in round 5 and a summary of the reasons given

Statement	Summary of reasons for Delphi Panel non-agreement
 Achievable standard: Cognitive screening should be offered to all patients with MS within 3 months of diagnosis Aspirational standard: Cognitive screening should be offered to all patients with MS within 4 weeks of diagnosis Aspirational standard: The MS team should engage patients with MS in an active, documented discussion about living a brain-healthy lifestyle at least once every 5 months 	 Cognitive performance may be influenced by emotional or treatment-related factors soon after diagnosis Not all centres are able to offer cognitive screening Results are unlikely to influence treatment decisions at an early stage in the care pathway Every 6 months is sufficient Frequency should be aligned with that of other routine clinic visits
 Core standard: The MS team should send a routine written reminder to every patient with MS about living a brain-healthy lifestyle at least once every year Achievable standard: The MS team should send a routine written reminder to every patient with MS about living a brain-healthy lifestyle at least once every 6 months Aspirational standard: The MS team should send a routine written reminder to every patient with MS about living a brain-healthy lifestyle at least once every 6 months Aspirational standard: The MS team should send a routine written reminder to every patient with MS about living a brain-healthy lifestyle at least once every 3 months 	 Administrative costs Lack of evidence of effectiveness Disruption to patients Preference for face-to-face reminders
 Aspirational standard: The MS team should review at least once every 3 months whether each patient with MS who is not receiving a DMT is eligible for one, based on applicable guidelines Aspirational standard: The MS team should perform a follow-up clinical evaluation of each patient at least once every 3 months 	 Every 6 months is sufficient Inconvenience to patients Increased healthcare costs Lack of proven added benefit Every 6 months is sufficient Inconvenience to patients Increased healthcare costs Lack of proven added benefit
• Core standard: If a patient's response to their current DMT is judged to be suboptimal, an appropriate, alternative DMT should be offered within 4 months	 4 months is too long a delay

DMT, disease-modifying therapy; MS, multiple sclerosis.