

Treatment Algorithm Multiple Sclerosis Disease-modifying Therapies Notes – See full policy for details	
Treatments	with [*] suffix should be agreed at MDT (see policy section 2)
[Note 1]	Where clinical or radiological markers indicate a poor prognosis for rapidly developing permanent disability, ofatumumab or ocrelizumab may be considered after a single clinical episode with MRI activity. Physicians and patients should weigh up the risks against the potential benefit. This must be agreed by the MDT.
[Note 2]	Alemtuzumab , ocrelizumab , ofatumumab and cladribine may be a safer option than natalizumab when John Cunningham virus (JCV) serology is high-index positive.
[Note 3]	Intolerance to treatment; If a patient satisfies the eligibility criteria for a first line therapy, and then is relapse-free on a drug to which they become intolerant, they may be switched to another DMT even though their relapses may now fall outside the eligibility window.
[Note 4]	NHS England 2014 clinical commissioning policy states that fingolimod can be used as an alternative to natalizumab for those patients receiving natalizumab who are at high risk of developing PML as defined by the following: (i) JCV exposure indicated by anti-JCV antibody positive status, (ii) Receiving an immunosuppressant prior to receiving natalizumab , or (iii) Natalizumab treatment duration of >2 years. If patients develop a severe adverse effect to natalizumab (e.g. anaphylaxis), and they have not previously received fingolimod , then it may be appropriate to use fingolimod .
[Note 5]	For cladribine to be given, NICE specifically defined treatment failure as "1 relapse in the previous year and MRI evidence of disease activity."
[Note 6]	For fingolimod : under previous guidance, fingolimod may be given if patients have an unchanged or increased relapse rate or ongoing severe relapses compared with the previous year despite treatment with beta interferon or glatiramer acetate . This is now extended to include disease activity on dimethyl fumarate , diroximel fumarate , teriflunomide and ponesimod .
[Note 7]	Autologous haematopoietic stem cell treatment (AHSCT) for autoimmunity is commissioned at specialised centres and should be discussed at a specialist MDT.

After considering all these options, it may be appropriate to continue the second line therapy, despite evidence of disease activity. None of the drugs promise 100% efficacy and some patients and physicians may choose to tolerate some disease activity without changing drugs.

[Note 8]