

DEMYELINATING DISEASES — MS vs NMOSD vs MOGAD

Distinguishing the three main inflammatory demyelinating diseases. **Confirm AQP4-IgG and MOG-IgG serostatus (cell-based assay) before long-term treatment** — misdiagnosis leads to harmful or ineffective therapy.

Feature	Multiple Sclerosis	NMOSD (AQP4-IgG)	MOGAD (MOG-IgG)
Target antigen	None identified (T-cell-mediated CNS demyelination)	AQP4 water channel on astrocytes	MOG on the oligodendrocyte / myelin surface
Demographics	Young adults; F:M ~3:1	Median ~40 y; F:M up to ~9:1 ; more common in non-white populations	All ages incl. children; F ≈ M
Optic neuritis	Usually unilateral , mild–moderate	Severe , may be bilateral; posterior/chiasmal ; poor recovery	Often bilateral , painful, with disc edema; usually good recovery
Myelitis	Short-segment, partial	LETM ≥3 vertebral segments, central cord	LETM , often conus/cauda ; H-sign on axial
Other hallmark attacks	Brainstem/cerebellar, cognitive	Area postrema (intractable nausea/vomiting/hiccups), diencephalic, narcolepsy	ADEM (esp. children); cortical encephalitis with seizures (FLAMES)
Brain MRI	Periventricular ovoid (Dawson fingers), juxtacortical, infratentorial	Periependymal (around 3rd/4th ventricle & area postrema), diencephalic	Fluffy / ill-defined, ADEM-like, often resolves ; cortical FLAIR-hyperintensity
CSF oligoclonal bands	Positive ~85–95%	Usually negative (<20%)	Usually negative
Serology	No specific antibody	AQP4-IgG (cell-based assay)	MOG-IgG (cell-based assay); low/transient titers can be false-positive
Course	Relapsing → secondary progressive; progression <i>independent</i> of relapses	Relapsing; no progressive phase — disability is attack-driven	Monophasic or relapsing; disability attack-related
Attack recovery	Usually moderate	Often severe / incomplete	Often good
Acute treatment	IV methylprednisolone ± PLEX	IV steroids + early PLEX	IV steroids (steroid-responsive/dependent) ± IVIG/PLEX
Maintenance	DMTs (interferons → anti-CD20; see MS DMT sheet)	Eculizumab, ravulizumab, inebilizumab, satralizumab (FDA); rituximab	Off-label: rituximab (less reliable), IVIG , MMF, azathioprine — no FDA-approved therapy
⚠ Key pitfall	—	MS DMTs (IFN-β, natalizumab, fingolimod) WORSEN NMOSD	Don't over-call on a low/transient MOG titer; re-test if uncertain

Bottom line: MS — OCB-positive, short lesions, progressive course, treat with DMTs. NMOSD — AQP4-IgG, LETM & area postrema, severe attacks, treat with complement/IL-6/anti-CD20 (never MS DMTs). MOGAD — MOG-IgG, ADEM/bilateral ON, steroid-responsive, often monophasic; no approved therapy.