



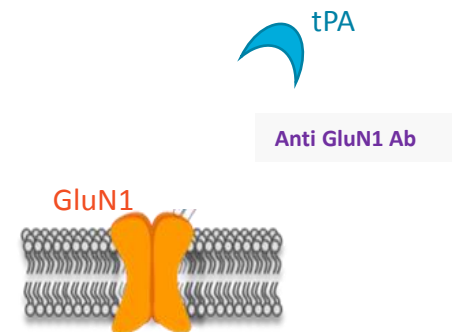
SELECTED OPPORTUNITIES IN NEUROSCIENCE

Antibody Protecting from BSCB leakage induced CNS damages
(BIO13193)

ANTIBODY PROTECTING FROM BSCB LEAKAGE INDUCED CNS DAMAGES (BIO13193)

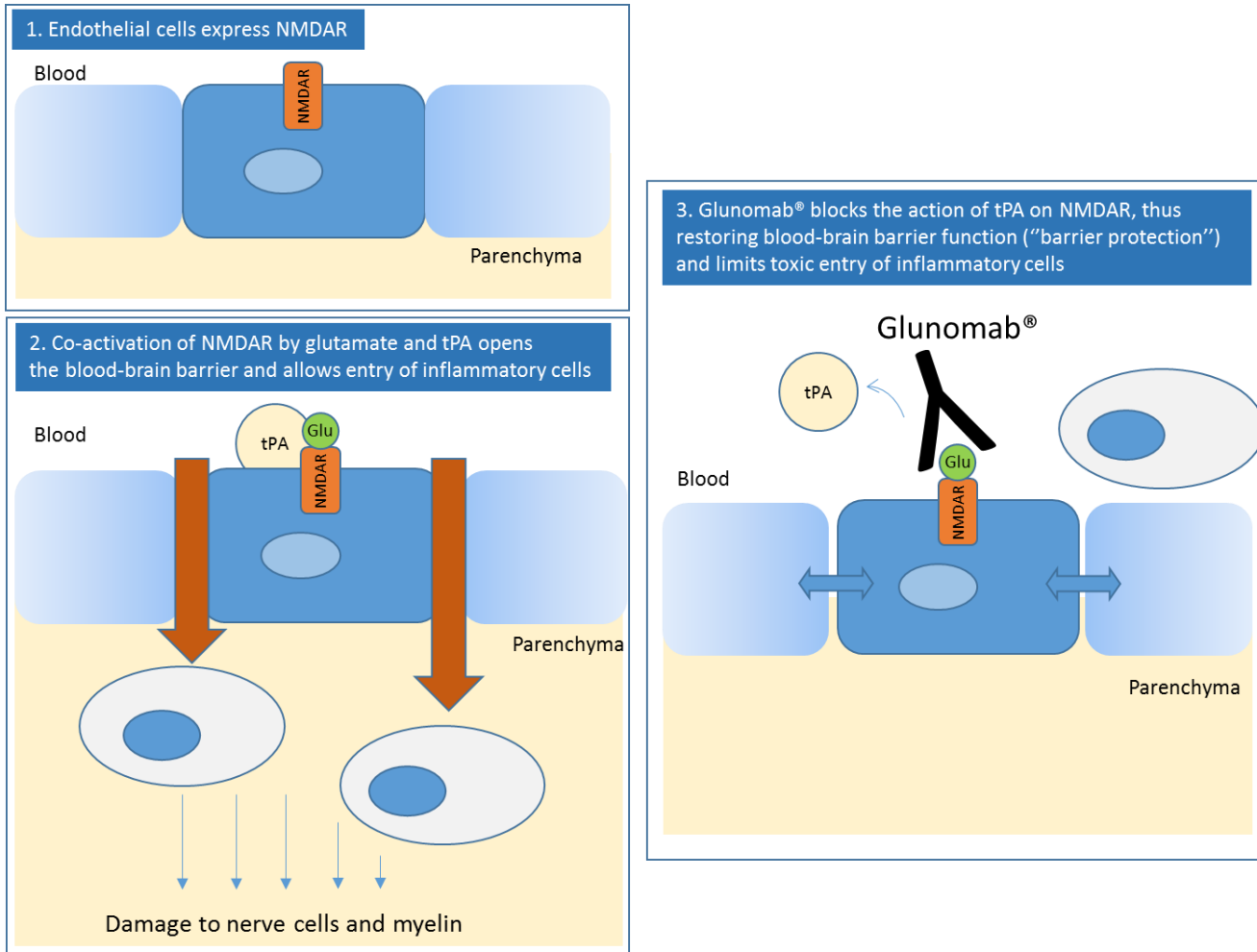
Product factsheet

- ▶ **Product:** Anti-GluN1 monoclonal antibody (GluNomab)
- ▶ **Mechanism:**
 - ◆ tPA positively regulates NMDAR through its interaction with GluN1 subunit in the extracellular compartment.
 - ◆ anti-GluN1 antibody prevents tPA binding on GluN1 subunit of NMDAR
 - ◆ BSCB opening in Multiple sclerosis and EAE allows CNS entry of neurotoxic elements (fibrinogen, ...)
- ▶ **Phase of development:** in vivo PoC
 - ◆ The anti-GluN1 monoclonal antibody (**GluNomab**) prevents tPA/NMDAR induced BBB/BSCB opening
 - ◆ GluNomab is protective in a multiple sclerosis mouse model
- ▶ **Potential applications:** multiple sclerosis, neurodegenerative diseases
- ▶ **Patent:** WO2014187879 published on Jan 22, 2015
- ▶ **Ref:**
 - ◆ Macrez et al. Brain 2016
 - ◆ Reijerkerk et al. J Neurochem. 2010
 - ◆ Gaberel et al. Neuropharmacology 2013
 - ◆ Macrez et al. Stroke 2011



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Rationale in Multiple Sclerosis

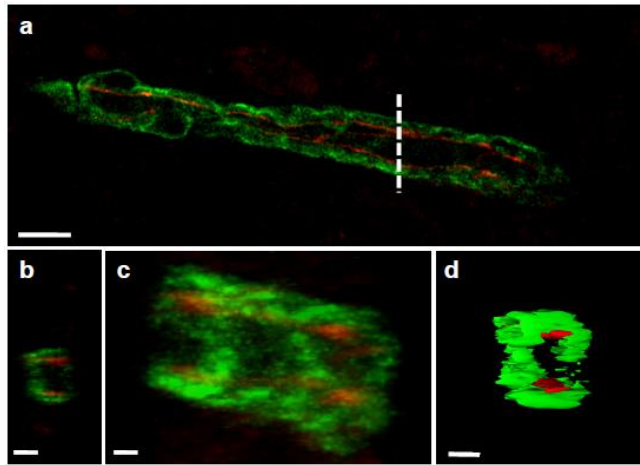


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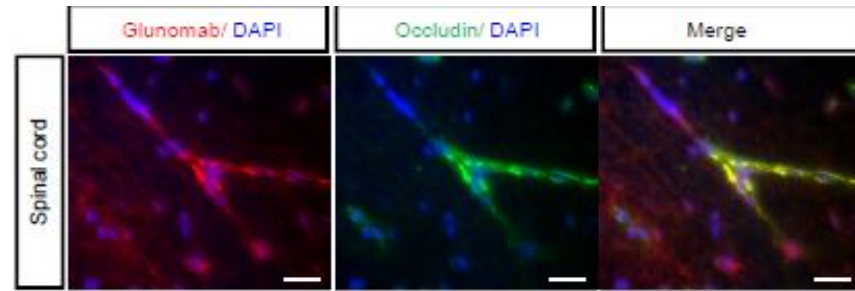
Rationale in Multiple Sclerosis

NMDARs localized at the luminal surface of endothelial cells can be blocked by Glunomab to prevent the blood-spinal cord barrier leakage

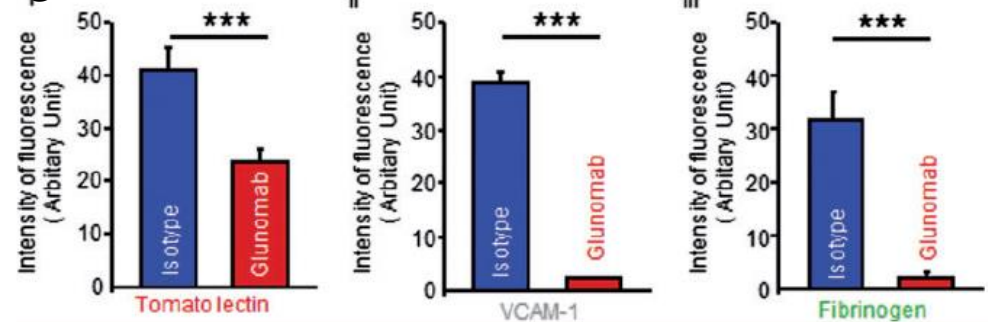
A



B



C



A: Confocal analysis of Glunomab staining (red) location in a collagen labeled (green) white matter capillary

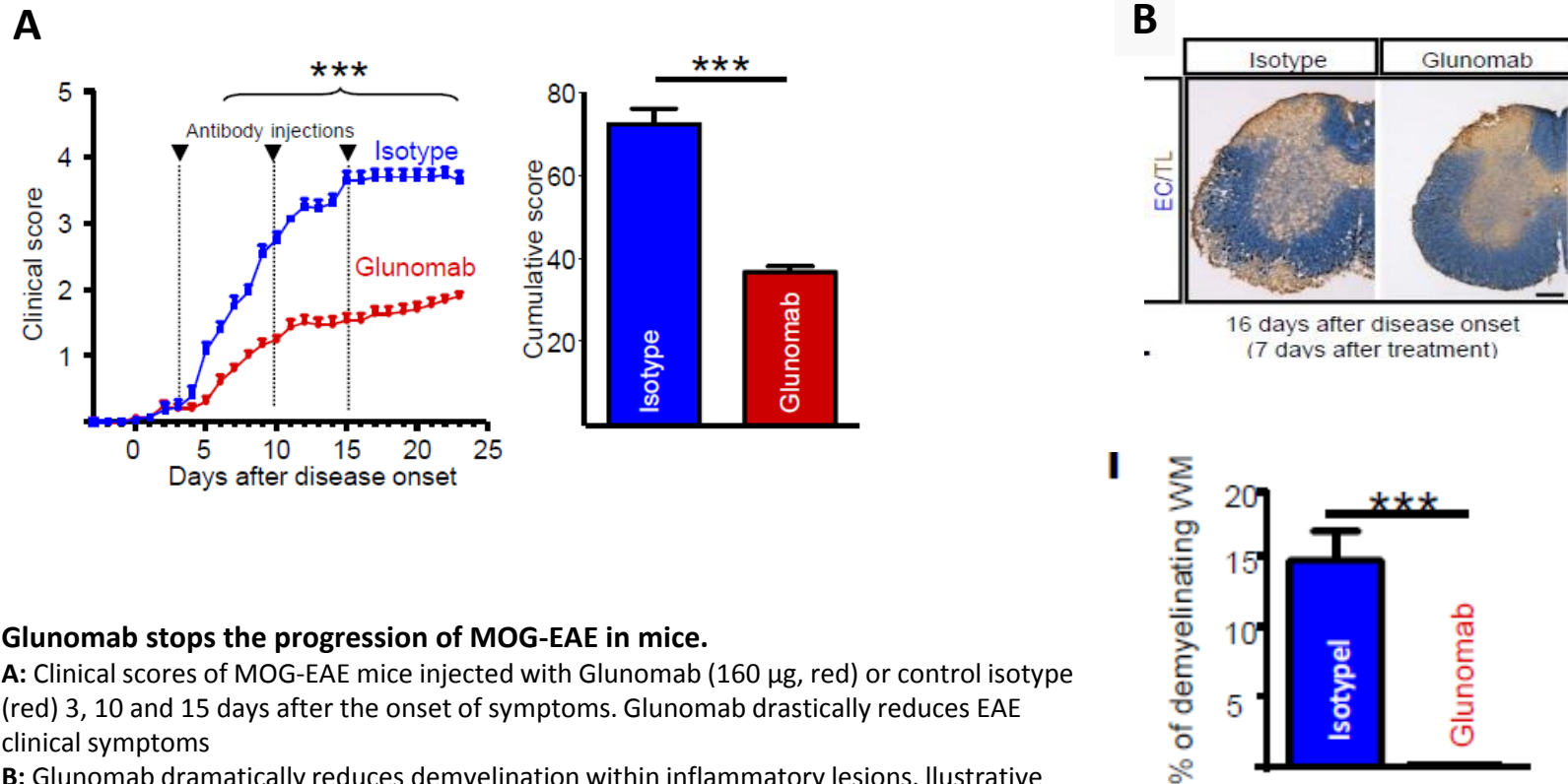
B: Immunostaining for Glunomab (in red) and occludin (green) in the spinal cord of mice showing colocalization at tight junctions.

C: Glunomab limits BSCB permeabilisation: Quantification of immunofluorescence intensity in spinal cord tissue

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Rationale in Multiple Sclerosis

Disease progression and demyelination is prevented by Glunomab



Glunomab stops the progression of MOG-EAE in mice.

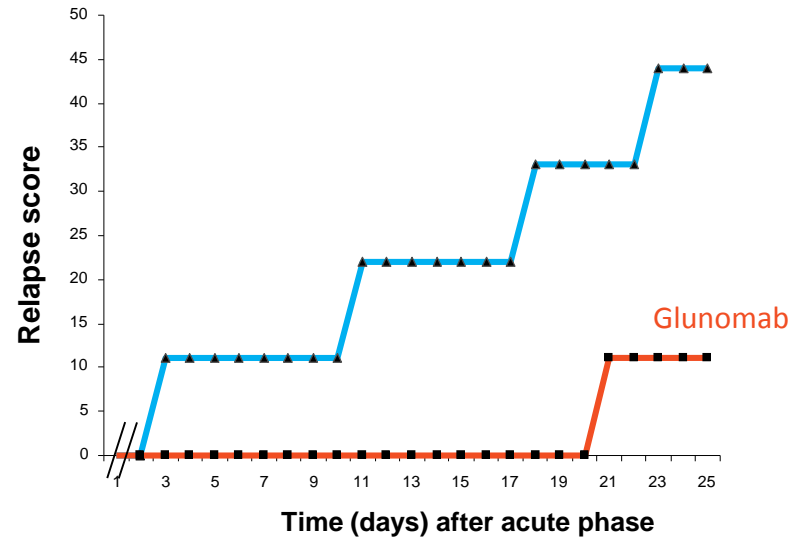
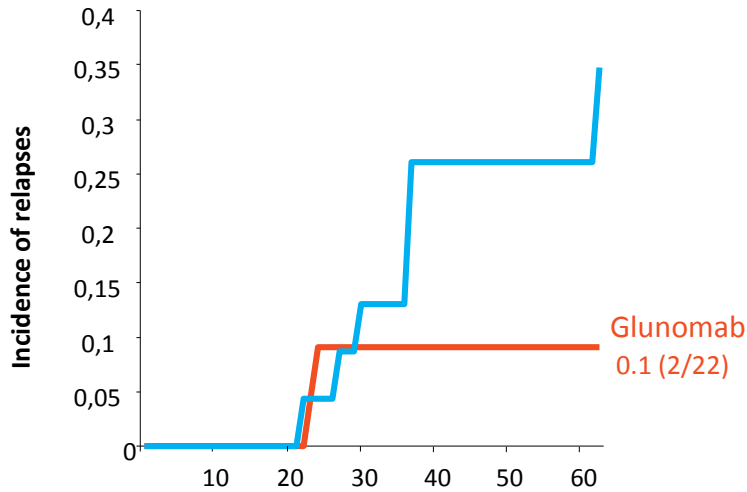
A: Clinical scores of MOG-EAE mice injected with Glunomab (160 μ g, red) or control isotype (red) 3, 10 and 15 days after the onset of symptoms. Glunomab drastically reduces EAE clinical symptoms

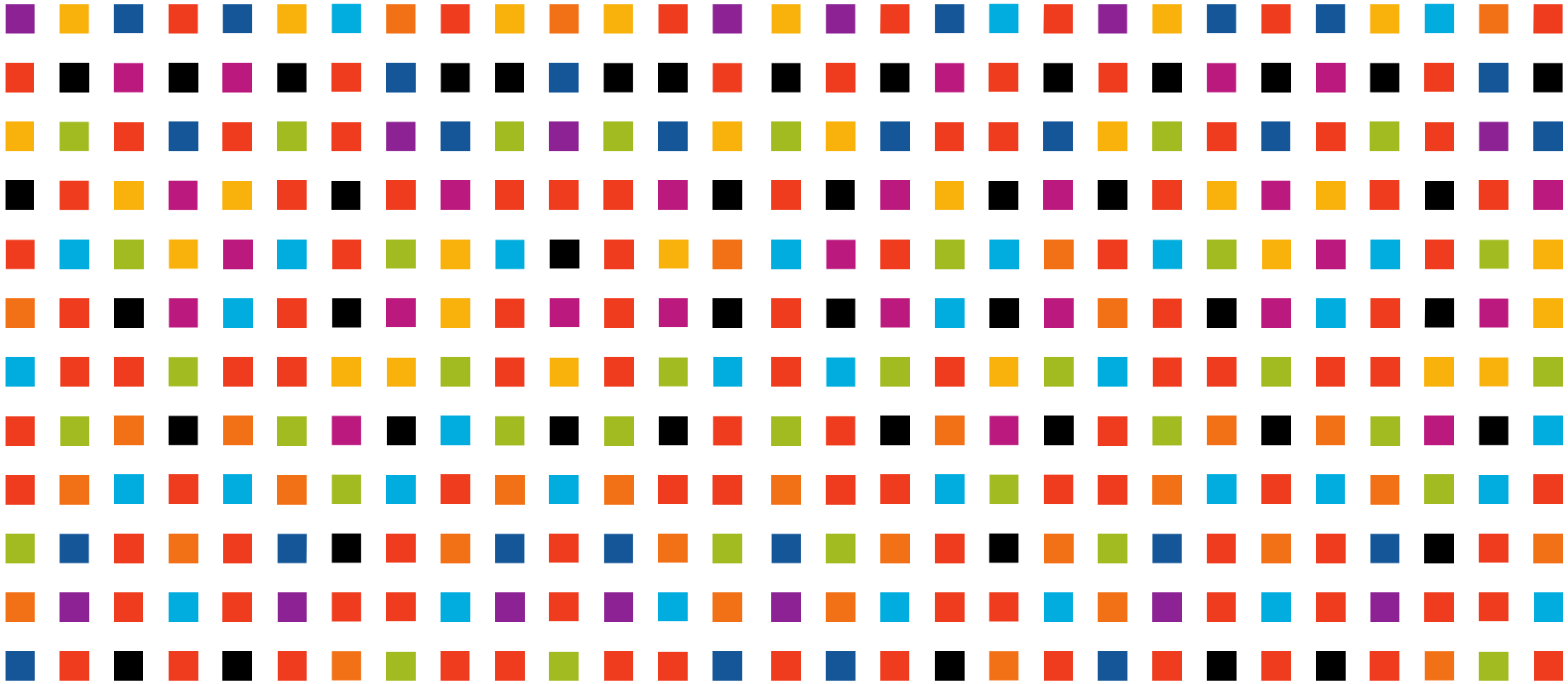
B: Glunomab dramatically reduces demyelination within inflammatory lesions. Illustrative picture (upper panel) of spinal cord of isotype and glunomab injected mice and quantification (lower panel).

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Rationale in Multiple Sclerosis

Glunomab has a therapeutic effect on relapses of plp-EAE mice





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