| Cell | Molecule | Potential impact on remyelination |
|---|--|--|
| type | Marilian and distributions of the | Township of the Con- |
| Oligodendroglial Precursor Cells (OPCs) | Myelin-associated neurite | Impairing remyelination |
| | outgrowth inhibitor NogoA | Impeding neurite outgrowth |
| | N. 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 | Enhanced remyelination by anti-NogoA-Ab ozanezumab |
| | Membrane-bound repulsive | Impairing remyelination |
| | guidance molecule A (RGMa) | Modulates T cell response |
| | | Impeding neurite outgrowth |
| | NT | Enhanced remyelination by anti-RGMa-Ab elezanumab |
| | Nogo-receptor interacting | Impairing remyelination |
| | protein LINGO-1 | Impeding OPC differentiation |
| | Encolor and the (Encolor dis- | Enhanced remyelination by anti-LINGO-1-Ab opicinumab |
| | Envelope protein (Env) of the | Impairing remyelination |
| | pathogenic human endogenous | Impeding OPC differentiation |
| | retrovirus type W (pHERV-W) | Enhanced remyelination and reduced neurodegeneration by |
| | Simvastatin | anti-pHERV-W Env-Ab temelimab |
| | | (Potentially) stimulating OPC differentiation and survival |
| | Quetiapine | Stimulating OPC differentiation and myelin protein production |
| | Clemastine fumarate | Stimulating OPC differentiation and remyelination |
| | Chitinase 3-like-3 (Chi3l3) | Activating epidermal growth factor receptor (EGFR) |
| Neural Stem Cells (NSCs) | | Inducing pro-oligodendrogenic transcription factor signature |
| | Clia | Enhancing oligodendrogenesis |
| | Gli1 | Impairing remyelination |
| | | Immobilizing and impeding the differentiation of resident NSCs |
| | C:L1 | Enhanced remyelination by GANT61 (SMI of Gli1) |
| | Sirt1 | Impairing remyelination |
| | | Suppressing oligodendrogenesis from NSCs Activating downstream Akt and p38 MAPK |
| | | Limiting the expansion of SVZ NSCs and OPCs |
| | | Enhanced oligodendrogenesis by EX-527 (SMI of Sirt1) |
| | nuclear factor I X (NFIX) | Impairing remyelination |
| | nuclear factor 174 (141174) | Suppressing oligodendrogenesis from NSCs |
| | B-cell leukemia homeodomain | Early regulator of SVZ neuro- und oligodendrogenesis |
| | 1 (Pbx1) | Priming factor to activate neuron-specific genes Dcx and Th |
| | prospero-related homeobox 1 | Enhancing SGZ neurogenesis and neuronal differentiation |
| | gene (Prox1) | Downstream target of β-catenin-TCF/LEF signaling |
| | drosha and nuclear factor IB | Enhancing oligodendrogenesis |
| | (NFIB) | Drosha enhances SGZ neurogenesis and suppresses |
| | (IVI ID) | oligodendrogenesis via suppression of NFIB |
| | nuclear factor-erythroid 2- | Enhancing SGZ neurogenesis |
| | related factor 2 (NRF2) | Enhancing neuronal differentiation |
| | | Regulating the expression of several antioxidant enzymes |
| | fibroblast growth factor | Promoting remyelination |
| | receptor-3 (FGFR3) | Enhancing SVZ oligodendrogenesis |
| | . , , , | Promoting migration of SVZ-derived OPCs to more distal areas |
| Microglia and peripherally-derived | Fractalkine receptor (CX3CR1) | Promoting remyelination |
| | - | Increasing microglial phagocytic capacity |
| | | Improving extracellular myelin debris clearance |
| | Triggering receptor expressed | Promoting remyelination |
| | on myeloid cells (TREM2) | Increasing microglial phagocytic capacity |
| | | Improving extracellular myelin debris clearance |
| 1 | | 1 0 |

| | TAM family receptors MerTK | Promoting remyelination |
|--|-------------------------------|---|
| | and Axl | Activating microglia and increasing microglial phagocytic |
| | and Axi | |
| | | capacity |
| | | Improving extracellular myelin debris clearance |
| | Envelope protein (Env) of the | Impairing remyelination |
| | pathogenic human endogenous | Stimulating microglia-associated inflammation and |
| | retrovirus type W (pHERV-W) | neurodegeneration |
| | | Decreasing microglial phagocytic capacity |
| | Matrix metalloproteinase 7 | Promoting remyelination |
| | (MMP7) | Splitting fibronectin aggregates that disrupt OPC differentiation |
| | | Secreted by microglia |
| | Long noncoding RNA | Impairing remyelination |
| | (lncRNA) GAS5 | Averting microglial M2 polarization by suppressing TRF4 |
| | CXCL12 | Promoting OPC differentiation |
| | | Secreted by microglia |
| | Semaphorin 3F | Promoting OPC differentiation |
| | _ | Improving OPC recruitment |
| | | Secreted by microglia |
| | Activin-A | Promoting OPC differentiation |
| | | Downstream activation of Rac-Cdc42 GTPases, Akt and mTOR |
| | | Secreted by microglia |
| | Galectin-3 | Promoting OPC differentiation |
| | | Improving myelin integrity and function |
| | | Secreted by microglia |

Table S1. Molecules and their potential impact on remyelination in the central nervous system (CNS). Ab=antibody; SMI=small molecule inhibitor; SVZ=subventricular zone; SGZ=subgranular zone; MAPK=Mitogen-activated protein kinases; TAM= TYRO3, Axl and MerTK family of receptor tyrosine kinases (RTKs).