

Recombinant Human Perlecan Domain-V Laminin-like Globulin Domain 3 (rhPDVLG3): A Novel Basement Membrane-Derived Matrikine Targeting Neurovascular Injury and Repair

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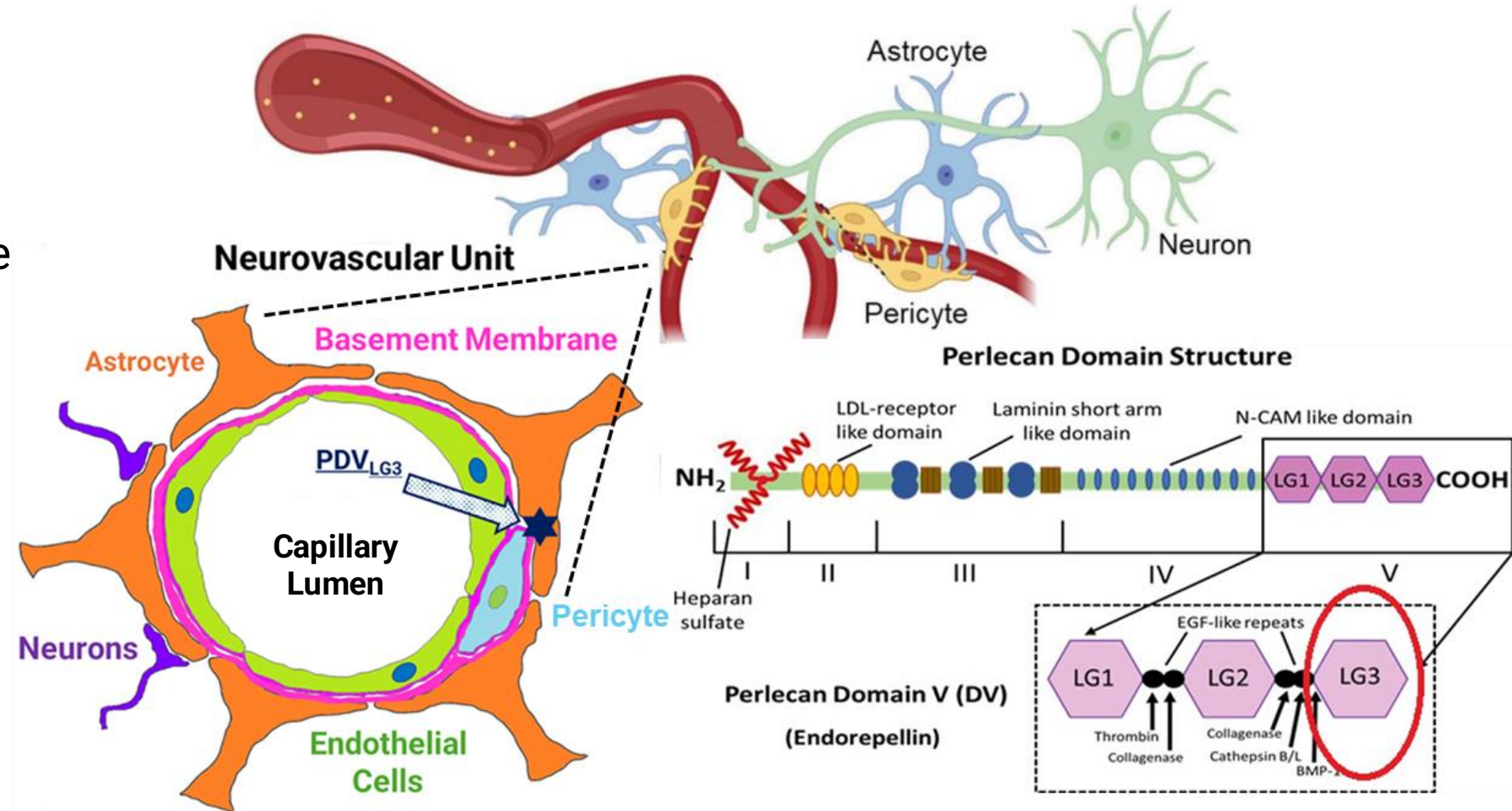
March 13th, 2025
ASENT, Bethesda MD

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Perlecan Domain V – LG3

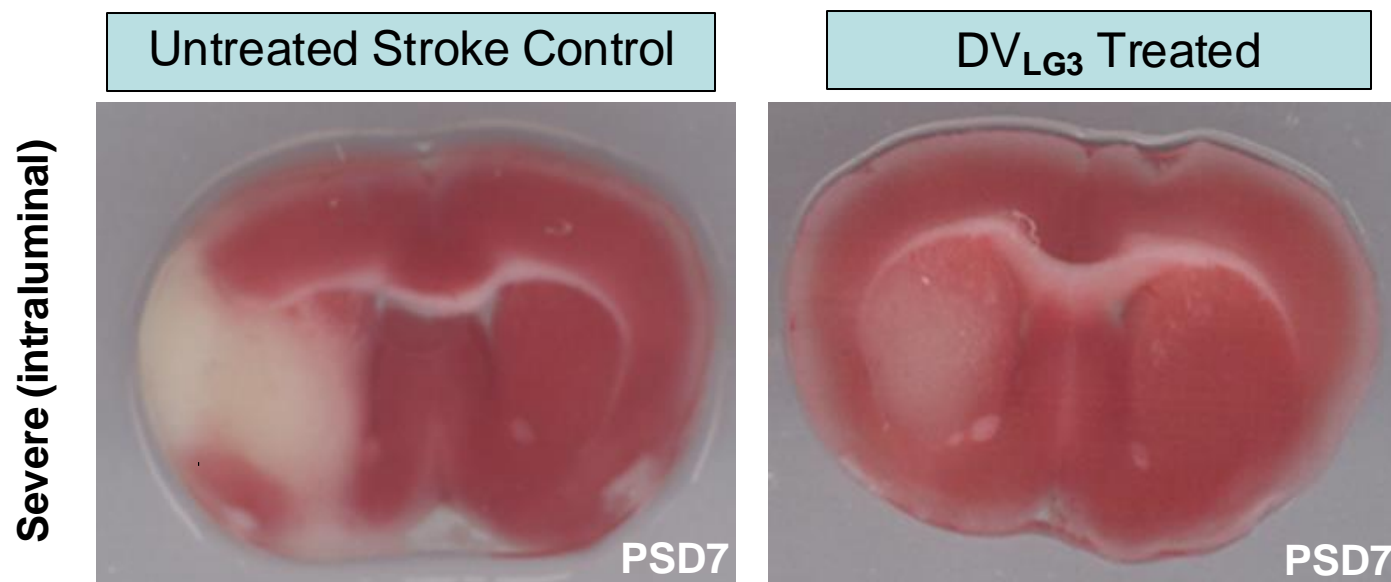
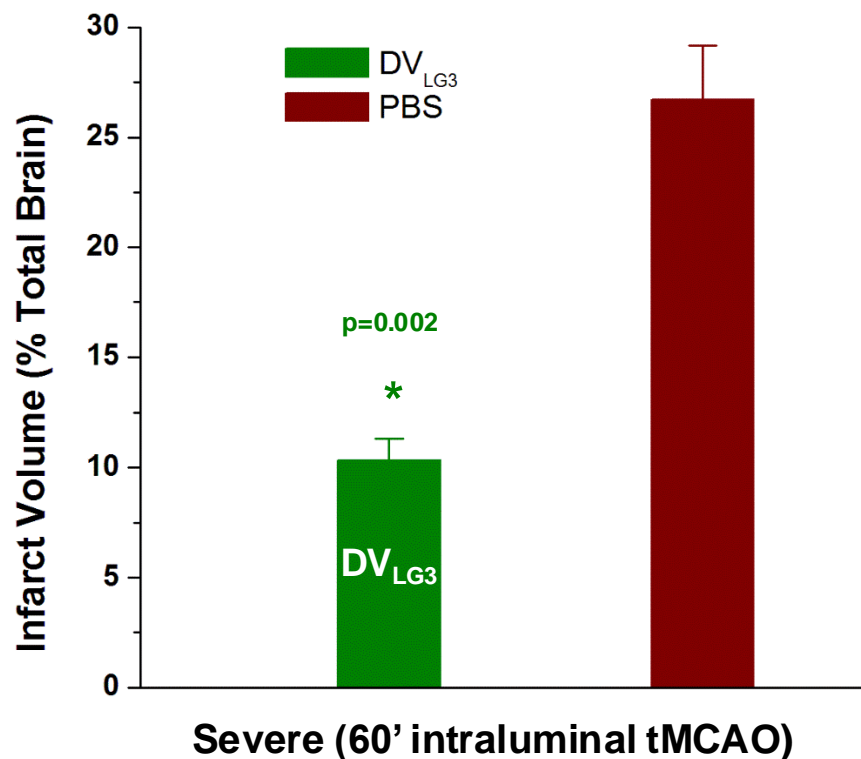
A First-in-Class Matrikine Therapeutic: Designed by Evolution, Developed by Stream

- 550M year-old protein
- 21 kDa c-terminus of Perlecan Domain V
- Elevated in brain of stroke and TBI patients
- Paracrine signal: 40nm distance
- Pleiotropic and anti-apoptotic: Interacts with all cell types of the NVU (through integrin R's)



LG3 is Acutely Neuroprotective

Quantification & Imaging of infarct volume following severe stroke (MCAo) in mice.



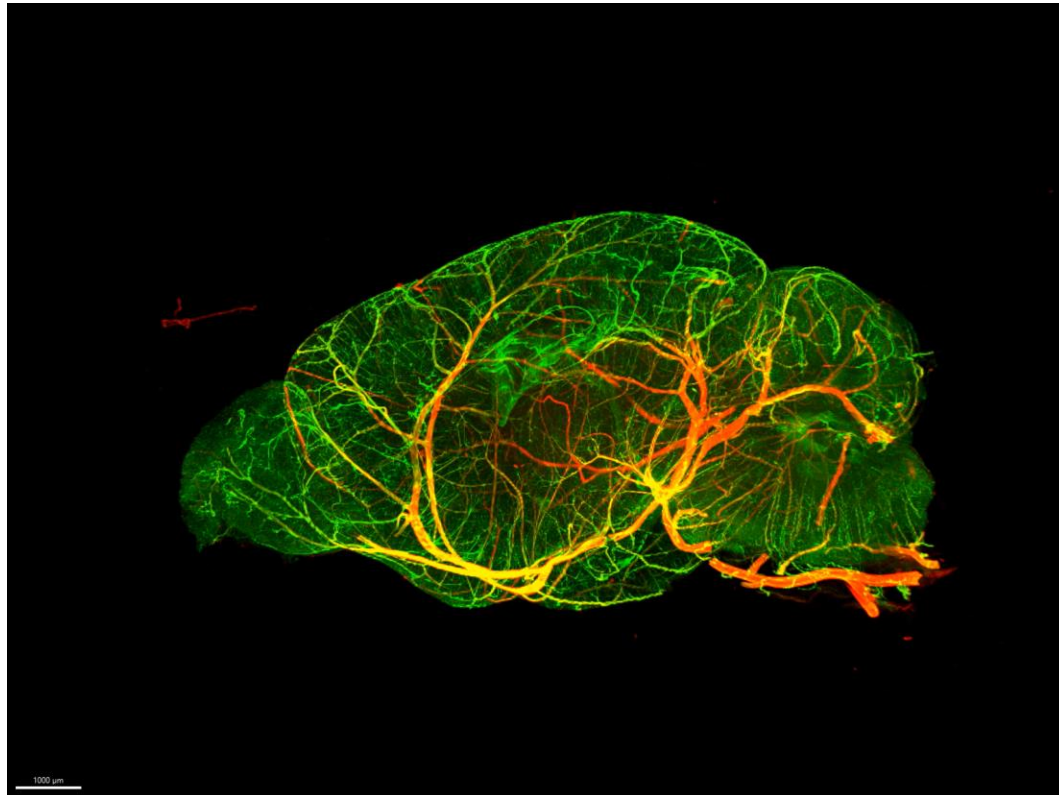
Drug Selection Study: LG3 only shown. 7 days after 60 min transient MCAO filament stroke in mice. Single 6 mg/kg IP dose of LG3 at reperfusion

Stroke:
Mouse MCAO

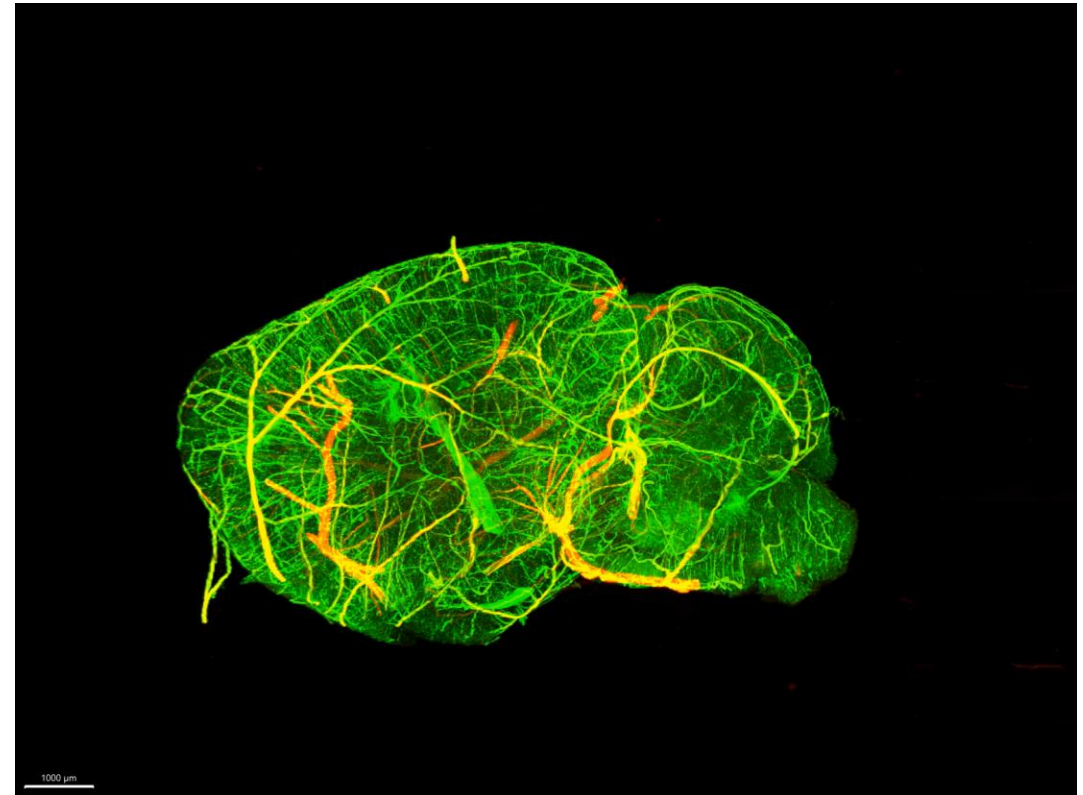
LG3 Preserves Peri-infarct Vasculature

Tissue Clearing + Light Sheet Microscopy 72hr Post CCA-MCA Stroke

Stroke - PBS

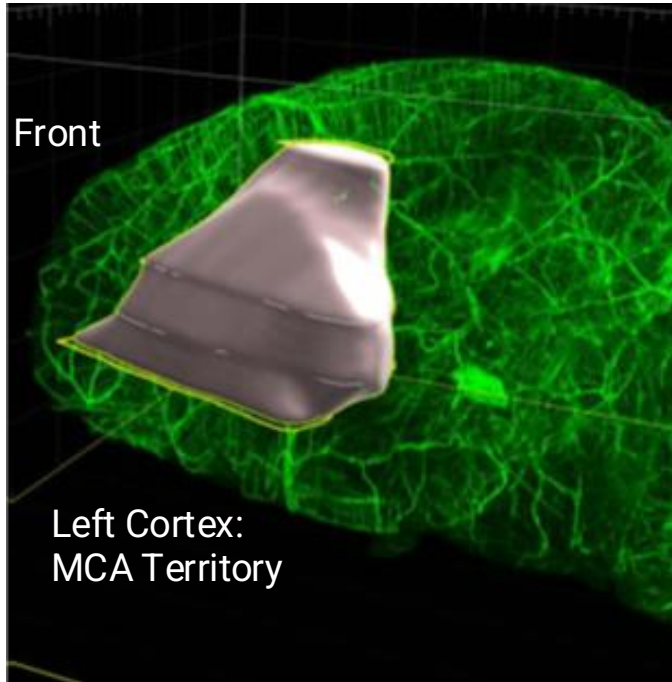


Stroke - LG3

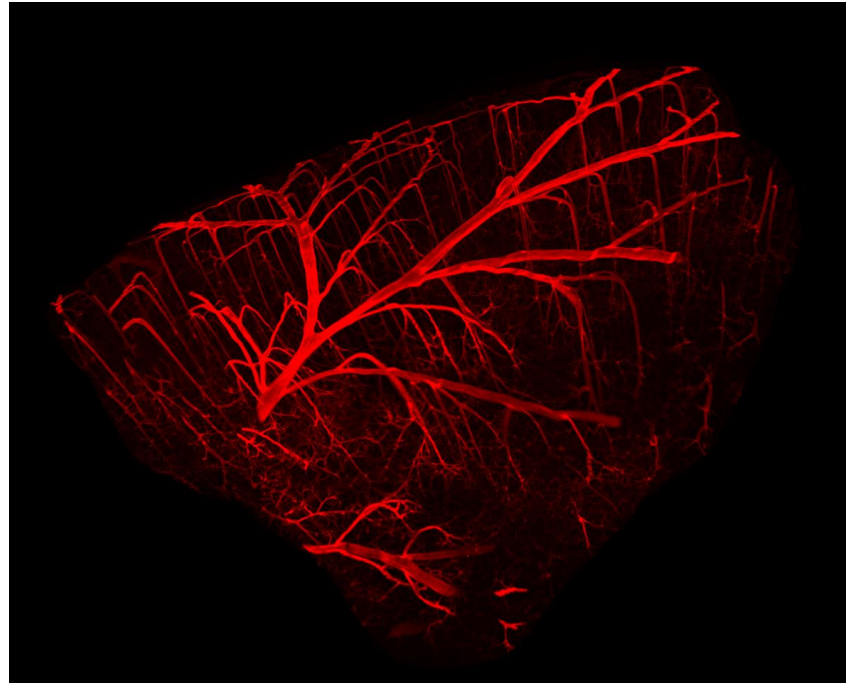


Focus: Penumbra Core 3D Vasculature Tracing

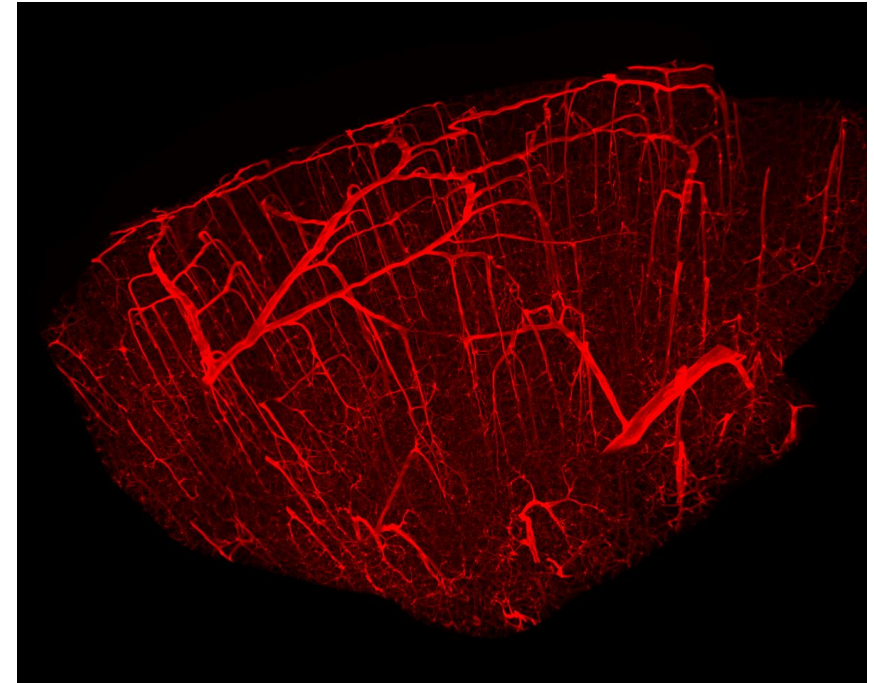
Stroke Region of Interest



Stroke - PBS

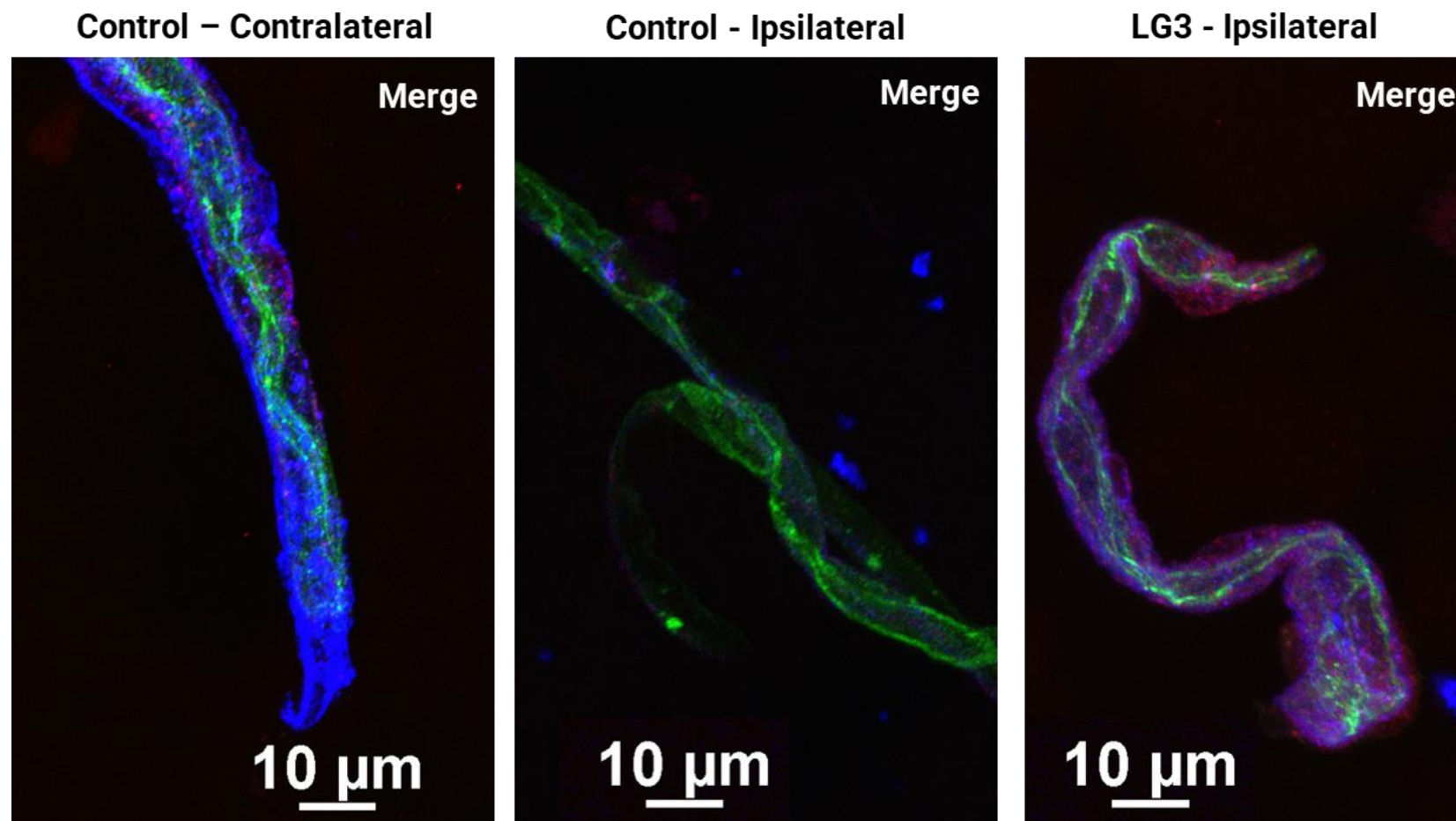
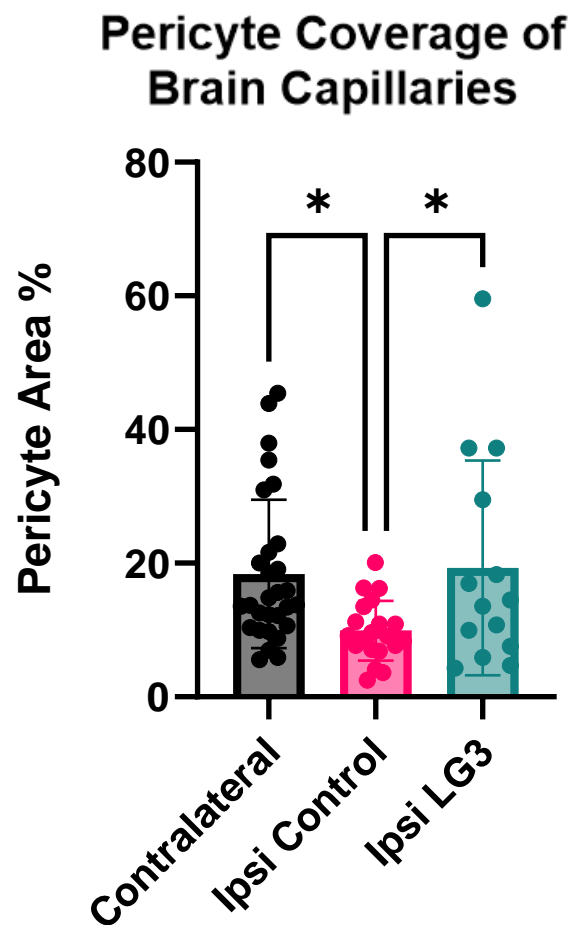


Stroke - LG3



LG3 Improves CNS Pericyte Coverage

Isolated Capillaries + Confocal Microscopy 72hr Post CCA-MCA Stroke



Labeling: PDGFR- β Claudin-5 Lectin

Summary: LG3 – Novel Neurotherapeutic

- Novel, first in class (matrikine) investigational protein
 - *Structurally derived from Perlecan within the extracellular matrix of NVU*
- Endogenously released in response to traumatic or ischemic injury
 - *21kDa matrikine evolutionarily primed for NVU signaling*
- Homes to and persists at sites of neurovascular injury
 - *Drug tracing demonstrates ~48hr parenchymal residence*
- Restores NVU homeostasis: Pleiotropic action → all cell types
 - *Reduces endothelial apoptosis*
 - *Modulates pericyte-microvessel dynamics*
 - *Downregulates glial inflammatory signaling*
 - *Directly neuroprotective and anti-apoptotic*
- Is acutely neuroprotective and functionally restorative
 - *In multiple species and CNS injury models (Stroke, TBI, Alzheimer's)*
- Clinical trials to be initiated soon
 - *Ultimately across a wide range of acute and chronic neurodegenerative diseases*

Acknowledgements

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