

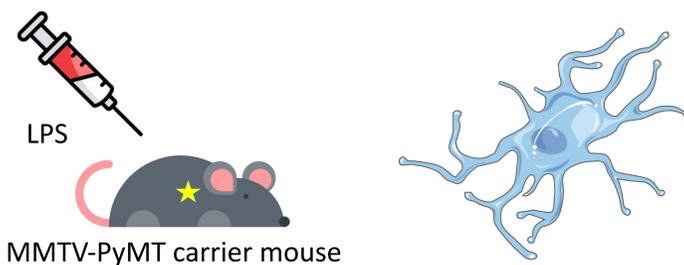
Effects of LPS stimulation on Microglial Activation in the Hippocampus and Hypothalamus of MMTV-PyMT carrier mice.



Sanya Shah, Macy Parmelee, Elizabeth Smithy, Gaushik Mouli

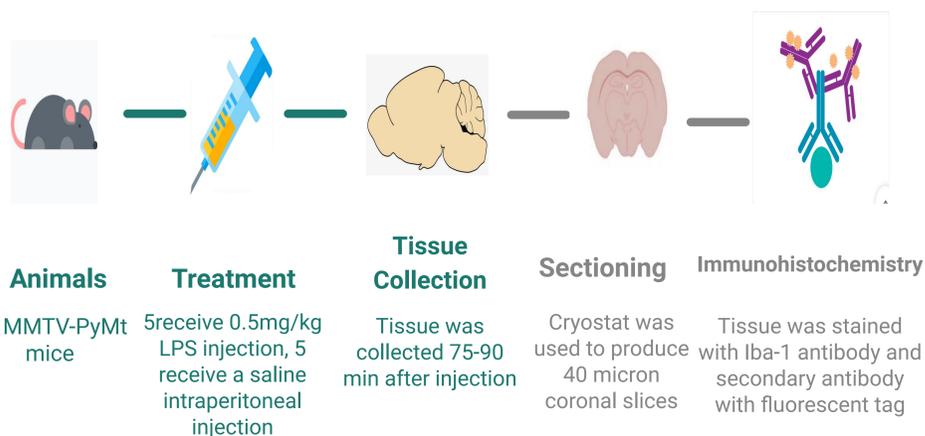
Intro

- Microglia are the primary innate immune system cells of the central nervous system (CNS) (Dong et al., 2019).
- Our goal is to research differentialized stress responses in distinct regions of the brain, particularly within mice prone to developing breast cancer.
- We analyzed microglia response profiles in the hippocampus and hypothalamus in MMTV-PyMT carrier mice that have experienced LPS peripheral immune challenge.



Microglia within the hypothalamus will be more vulnerable to LPS stimulation as compared to microglia within the prefrontal cortex because the hypothalamus maintains brain and body homeostasis. Hypothalamic microglia will have a greater increase in activation number and exhibit the M1 activation state (greater soma size and smaller process length).

Methods



Results

Microglial response to LPS in the Hypothalamus

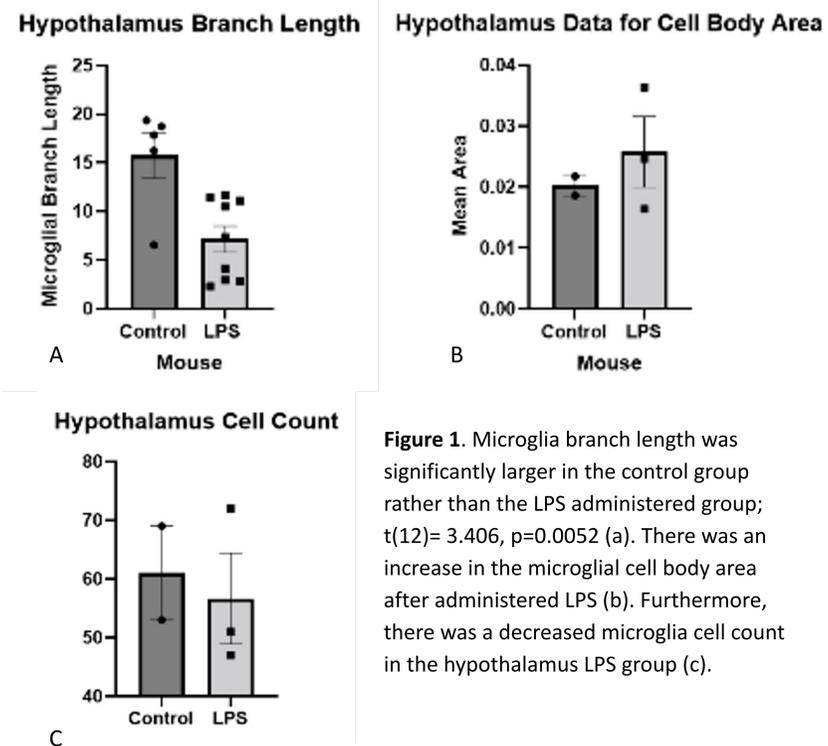


Figure 1. Microglia branch length was significantly larger in the control group rather than the LPS administered group; $t(12) = 3.406, p = 0.0052$ (a). There was an increase in the microglial cell body area after administered LPS (b). Furthermore, there was a decreased microglia cell count in the hypothalamus LPS group (c).

Microglial response to LPS in the Hippocampus

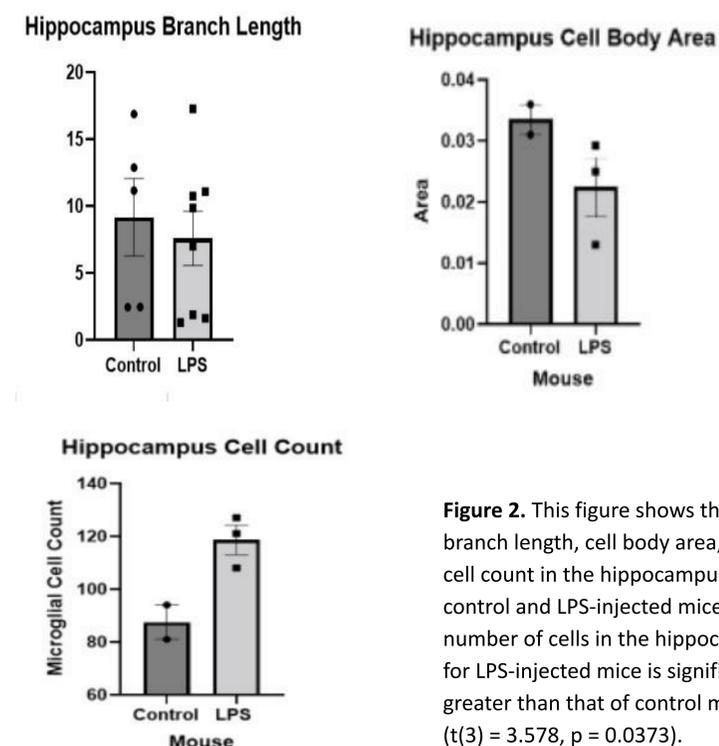


Figure 2. This figure shows the branch length, cell body area, and cell count in the hippocampus for control and LPS-injected mice. The number of cells in the hippocampus for LPS-injected mice is significantly greater than that of control mice ($t(3) = 3.578, p = 0.0373$).

Conclusions

- LPS mice demonstrated significantly higher microglia cell counts than control mice in the hippocampus
- LPS mice demonstrated significantly shorter branch lengths than control mice in the hypothalamus
- LPS exposure induces activated morphology in microglia, but rarely to a significant extent in the hypothalamus and hippocampus

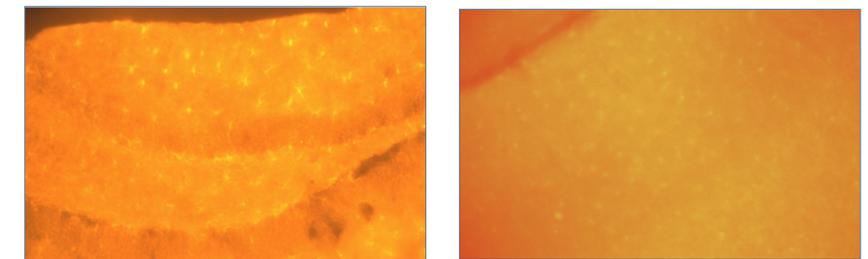


Figure 3. Fluorescent images of hippocampus of mouse 3 (left) and 8 (right) taken at 20x magnification. Mouse 3 was the LPS experimental mouse and demonstrates a greater degree of IBA-1 staining.

Acknowledgments

We would like to thank Dr. Jeremy Borniger at Cold Spring Harbor Laboratory for donating the mice tissue. We would also like to thank the Graduate Research Consultant Program funded by OUR, Research and Discovery Course Development Grant funded by OUR, and the Psychology and Neuroscience Undergraduate Research Grant funded by Lindquist Undergraduate Research Award for funding. Additionally, we would like to thank the Center for Faculty of Excellence, the College of Arts and Sciences & the department of Psychology and Neuroscience for additional funding and support.

References

- Bollinger, J. L., Burns, C. M. B., & Wellman, C. L. (2016). Differential effects of stress on microglial cell activation in male and female medial prefrontal cortex. *Brain, behavior, and immunity*, 52, 88-97.
- Calcia, M. A., Bonsall, D. R., Bloomfield, P. S., Selvaraj, S., Barichello, T., & Howes, O. D. (2016). Stress and neuroinflammation: a systematic review of the effects of stress on microglia and the implications for mental illness. *Psychopharmacology*, 233(9), 1637-1650.
- Dong, H., Wang, Y., Zhang, X., Zhang, X., Qian, Y., Ding, H., & Zhang, S. (2019). Stabilization of brain mast cells alleviates LPS-induced neuroinflammation by inhibiting microglia activation. *Frontiers in cellular neuroscience*, 13, 191.
- Folick, A., Koliwad, S., Valdearcos, M. (2021). Microglial Lipid Biology in the Hypothalamic Regulation of Metabolic Homeostasis. *Frontiers*, 27. <https://doi.org/10.3389/fendo.2021.668396>