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Effect of Eosinophils on Purinergic Receptor P2X₃ Expression in Mouse Sensory Neurons



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Introduction

- P2X₃ purinoceptors are activated by extracellular ATP released during cell stress.
- In asthma, sensitivity to inhaled ATP is increased.
- Airway eosinophilia, which is common in asthmatics, increases airway epithelial sensory nerve density.

Research Question

Do airway eosinophils increase P2X₃ expression in mouse sensory nerves?

Methods

Animals

1. WT: Normal level of eosinophils
 2. PHIL: Transgenic absence of eosinophils
 3. NJ16 (NJ1638 lineage): Transgenic chronic systemic eosinophilia
 4. NJ17 (NJ1726 lineage): Transgenic chronic pulmonary eosinophilia
- Airway sensory ganglia and dorsal root ganglia (thoracic vertebrae 1-5) were labeled for P2RX₃ mRNA using RNAscope in situ hybridization and imaged using an ApoTome confocal microscope (40X, 1.3 N.A.).
 - P2X₃ expression was determined by measuring the percentage of P2X₃ positive pixels within three, randomly assigned, non-overlapping 50x50 micron sections of neurons.
 - Data was analyzed using one-way ANOVA tests.

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Results

Figure 1: Nodose/Jugular Ganglia from WT and Chronic Airway Eosinophilic mice. RED = P2X₃ mRNA

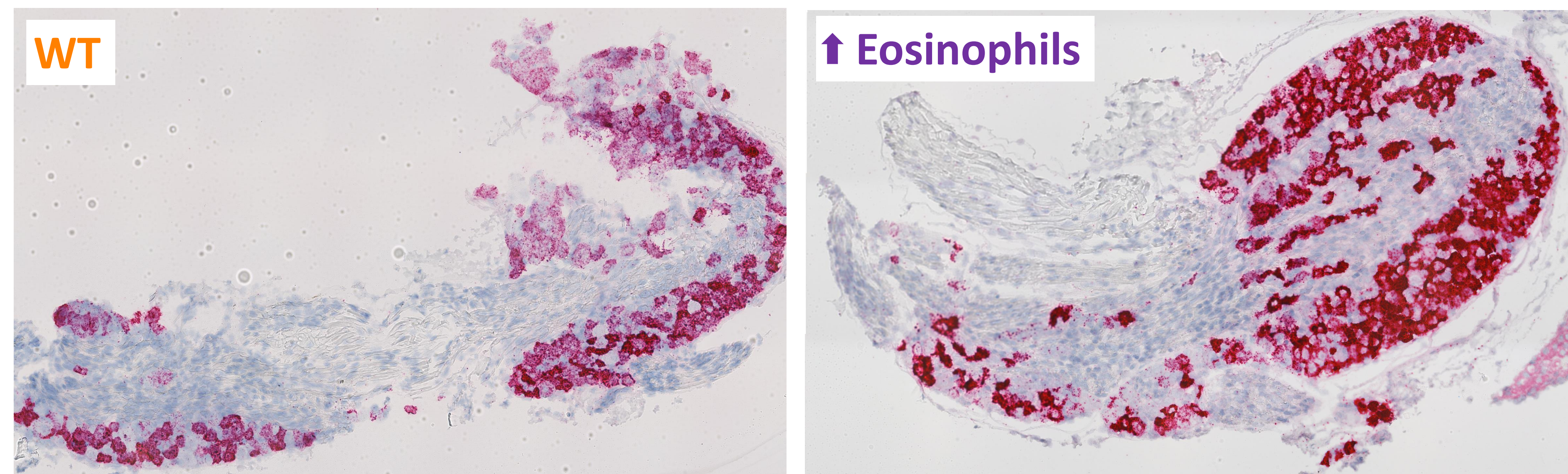
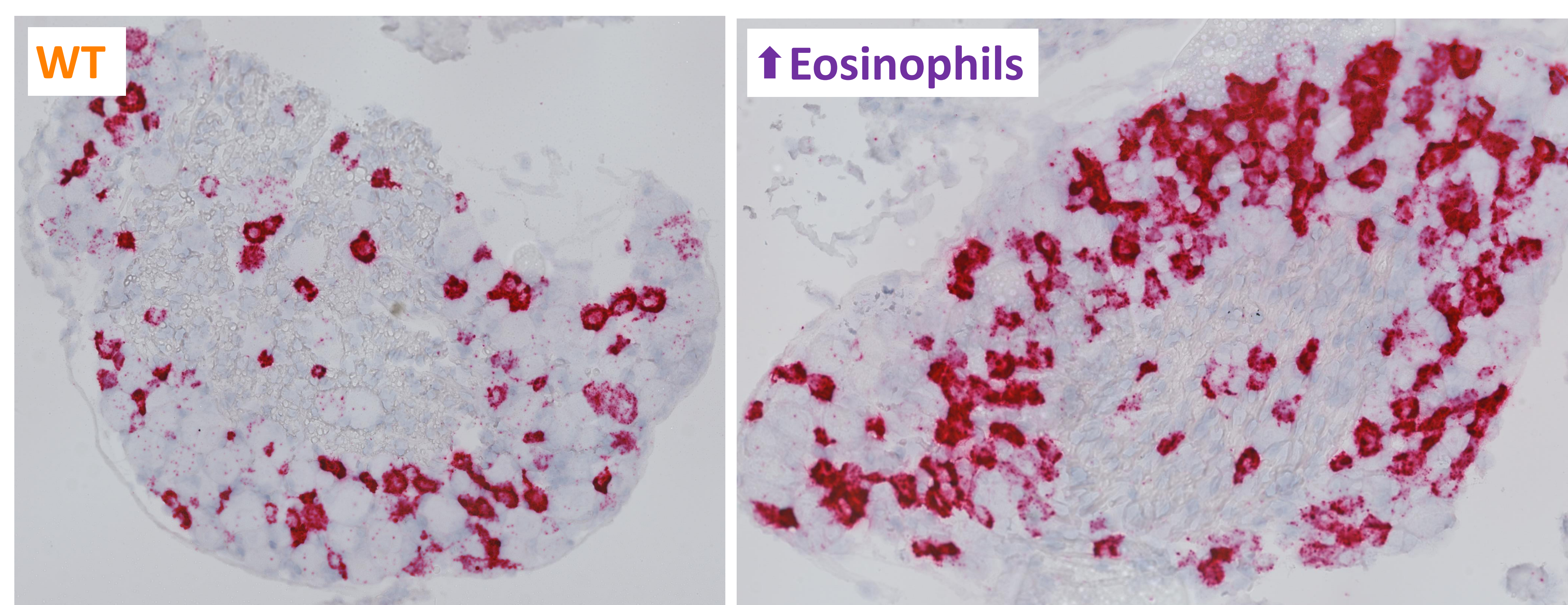


Figure 2: Dorsal Root Ganglia from WT and Chronic Airway Eosinophilic mice. RED = P2X₃ mRNA



Conclusions

- P2X₃ is highly expressed in airway sensory neurons.
- Eosinophils positively regulate sensory nerve P2X₃ expression.
- Eosinophils may increase neuronal ATP sensitivity in asthma.

Figure 3: P2X₃ expression is increased in the nodose/jugular ganglia of mice with chronic eosinophilia

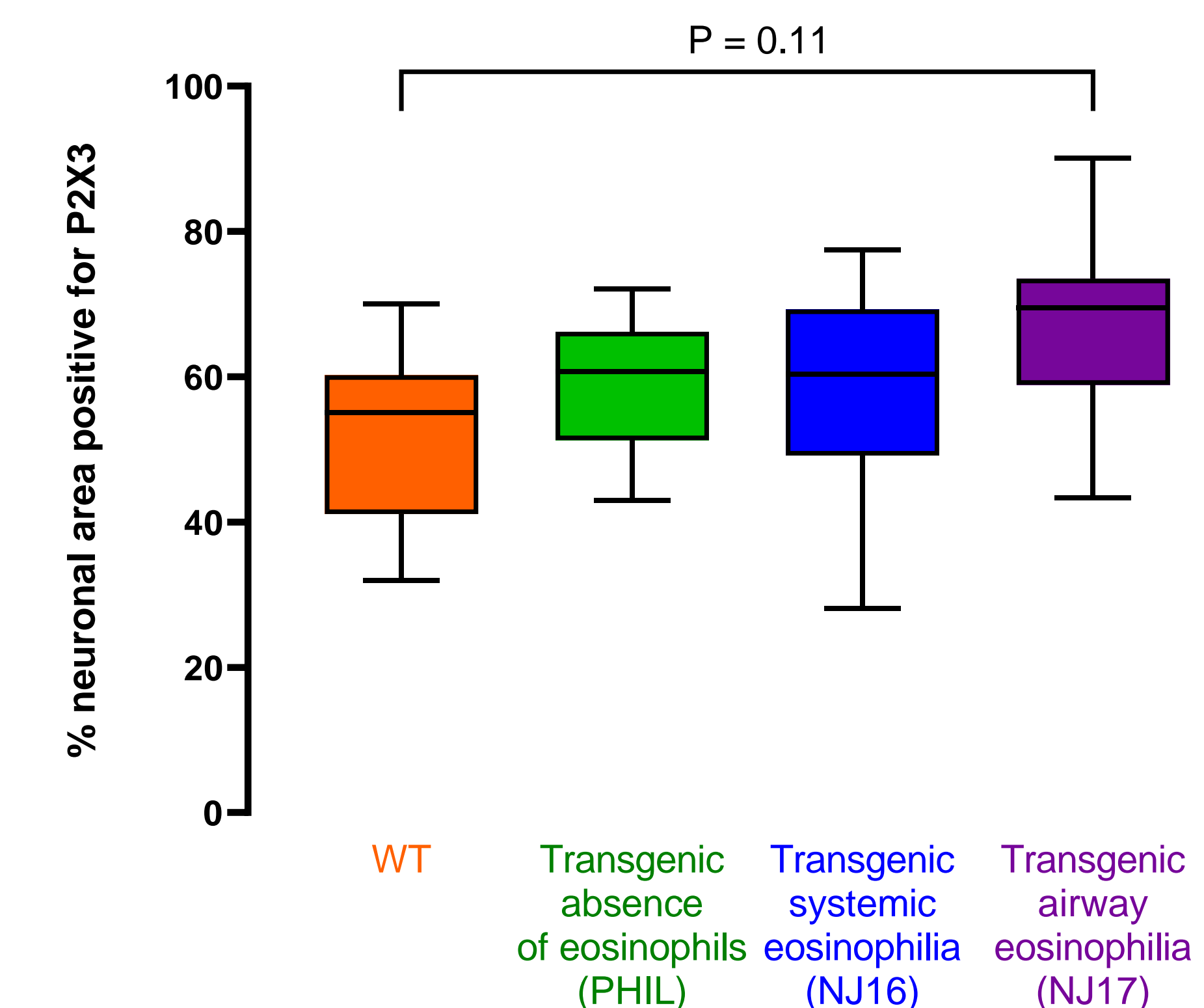
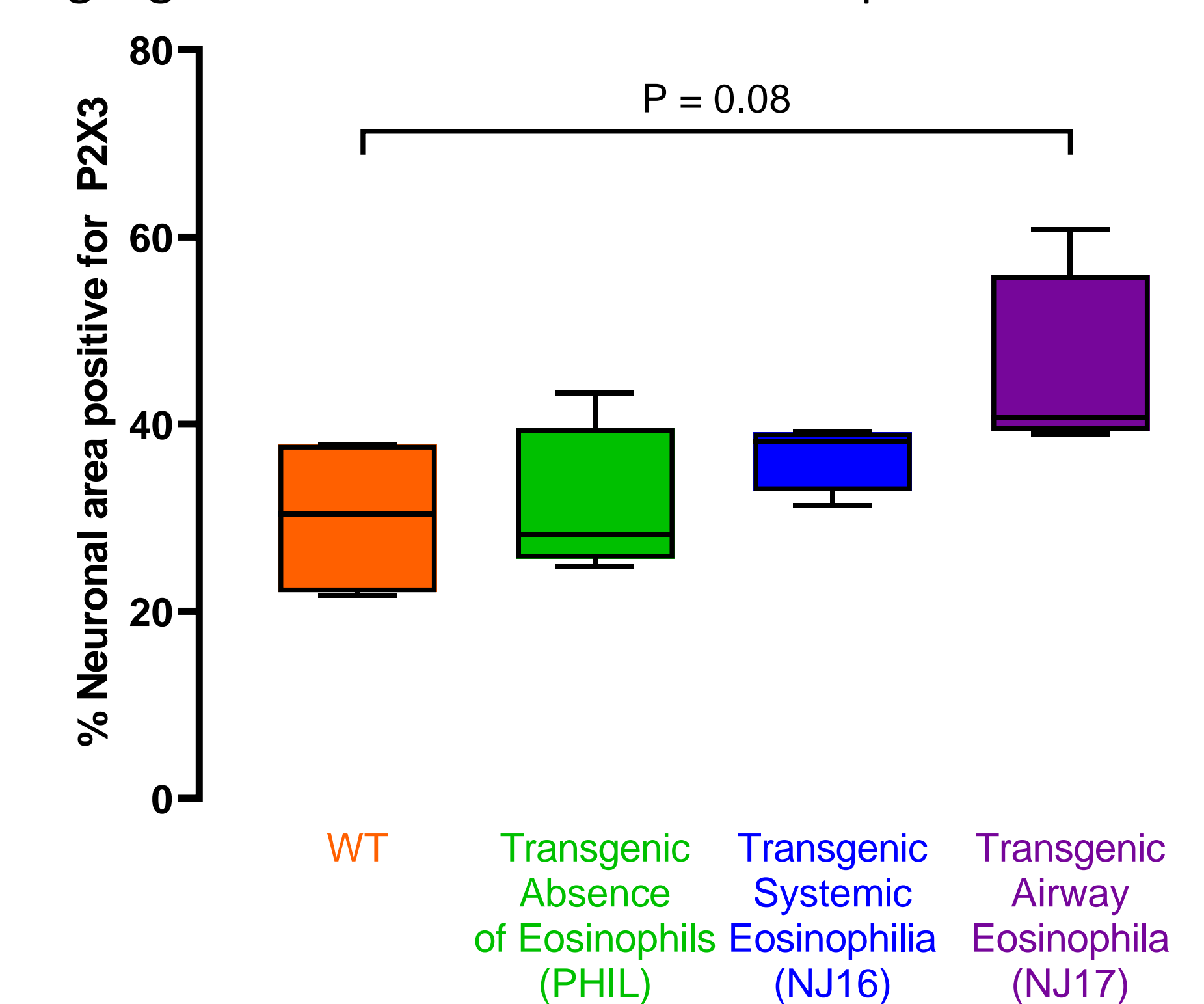


Figure 4: P2X₃ expression is increased in the dorsal root ganglia of mice with chronic eosinophilia



Asthma Lab

