



# Decoding equine IBH: Single-cell insights into allergen-specific immune dysregulation

14th International Veterinary Immunology Symposium

#### Johanna Geisler

Vetsuisse Faculty, University of Berne Department of Clinical Research and Veterinary Public Health



## $u^{^{b}}$

## Equine insect bite hypersensitivity (IBH)



Seasonal allergic IgE mediated dermatitis with

#### Th2 dominance

#### Cause:

· Bites from Culicoides spp.

#### Clinical signs:

Pruritic papules → self-trauma

#### Diagnosis:

- Clinical signs, exclusion of other diseases
- IgE testing (intradermal or serological)

#### **Treatment**

Strict insect control



https://infravec2.eu/wpcontent/uploads/2017/08/Culicoides -nubeculosus.jpg



Video taken by Fenja Johanna Von der Hören









Genetics of Skin Disease in Horses - Scientific Figure on ResearchGate. Available from: https://www.researchgate.net/figure/Clinical-signs-of-IBH-A-B-Alopecia-broken-hairs-and-focal-ulcerations-in-the-mane\_fig1\_342092138 [accessed 12 Jun, 2023]

 $oldsymbol{u}^{\scriptscriptstyle b}$ 

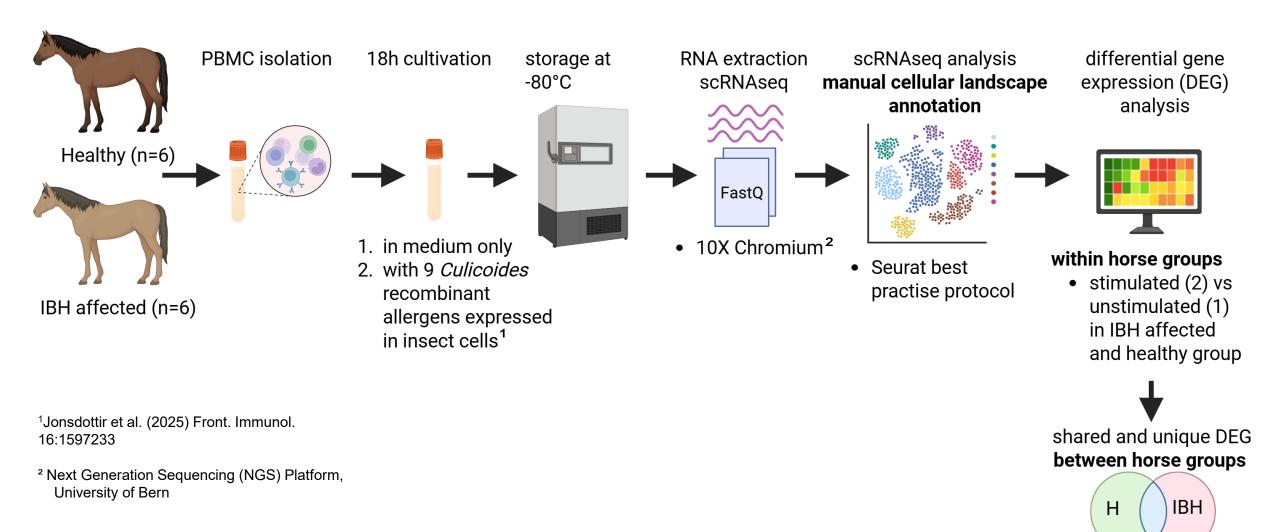
### Research aim



How do PBMC from healthy and IBH horses differ in their allergen response on a transcriptional level?

## $u^{\scriptscriptstyle b}$ Methods

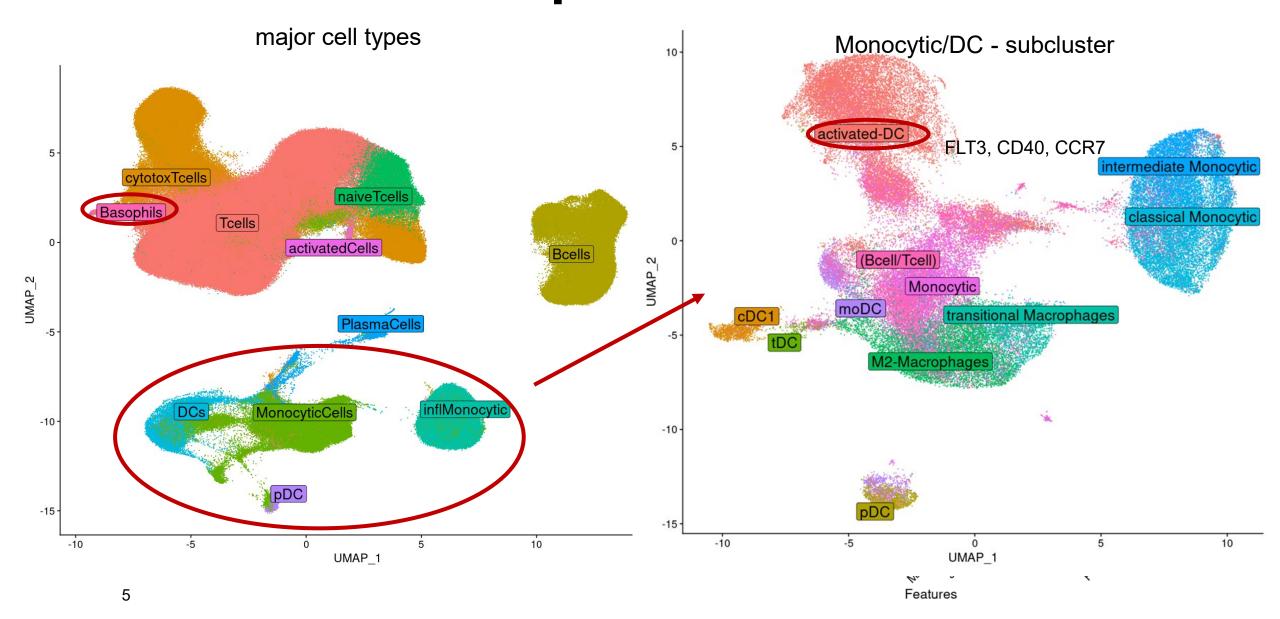




## $u^{^{\scriptscriptstyle b}}$

## **Annotation of equine PBMC**

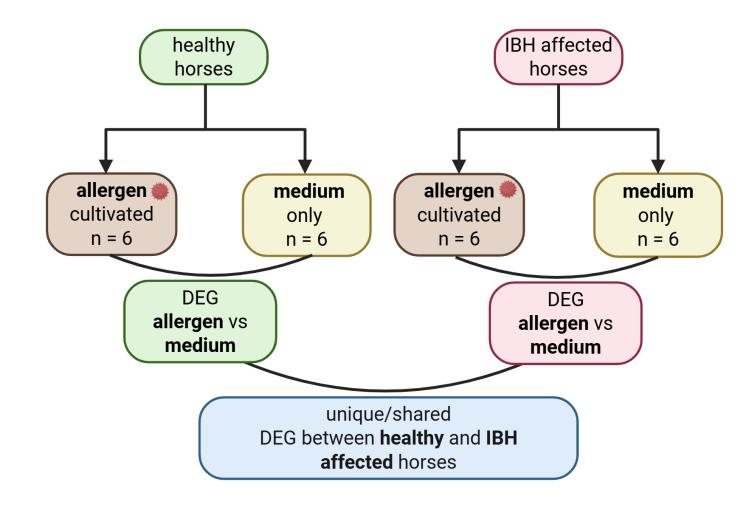




### $u^{^{\scriptscriptstyle b}}$

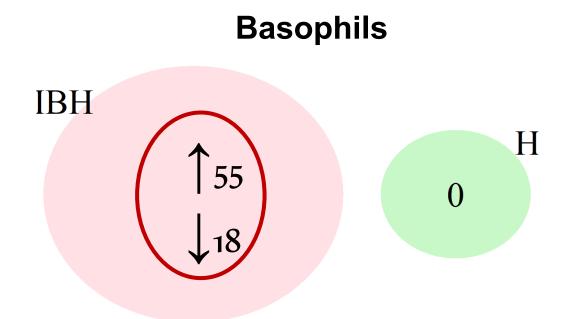
## Shared and unique DEG within horse groups stimulated vs unstimulated

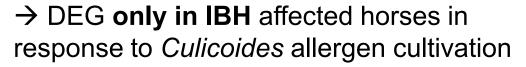


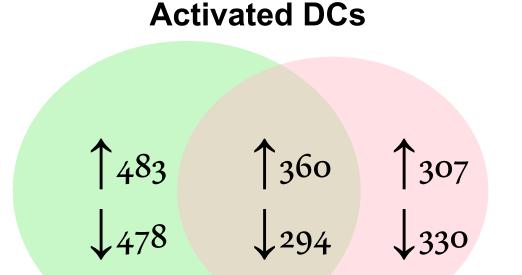


## Shared and unique DEG within horse groups stimulated vs unstimulated





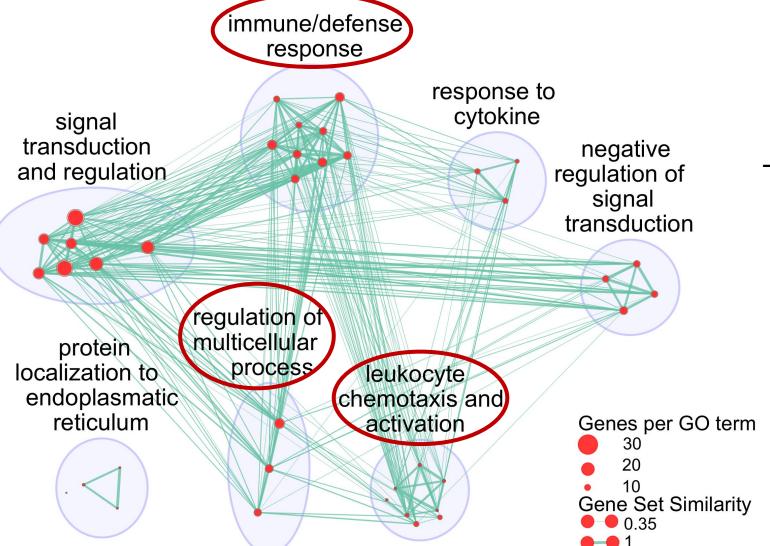


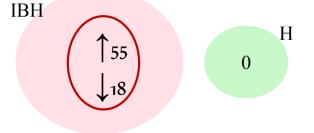


- → enrichment analysis (EA) with g:profiler¹
- → visualization with EnrichmentMap and AutoAnnotate in cytoscape<sup>2</sup>

## Immune priming and initiation of inflammation by IBH basophils







→ No biological process associations with downregulated genes

#### Cytokines:

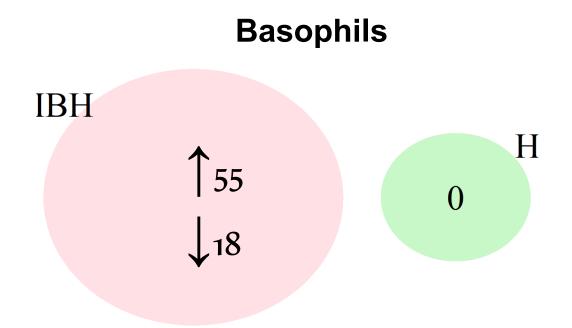
CSF2, IL6, IL31

#### Chemokines:

- CXCL8, CCL4, CCL1
- → Promotion of Th17 and Th2 immune response, pruritus and inflammation

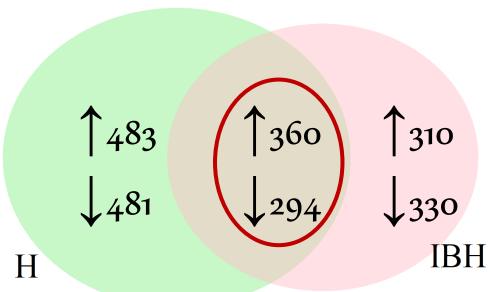
## Shared and unique DEG within horse groups stimulated vs unstimulated





→ DEG **only in IBH** affected horses in response to *Culicoides* allergen cultivation





- → enrichment analysis (EA) with g:profiler¹
- → visualization with EnrichmentMap and AutoAnnotate in cytoscape<sup>2</sup>

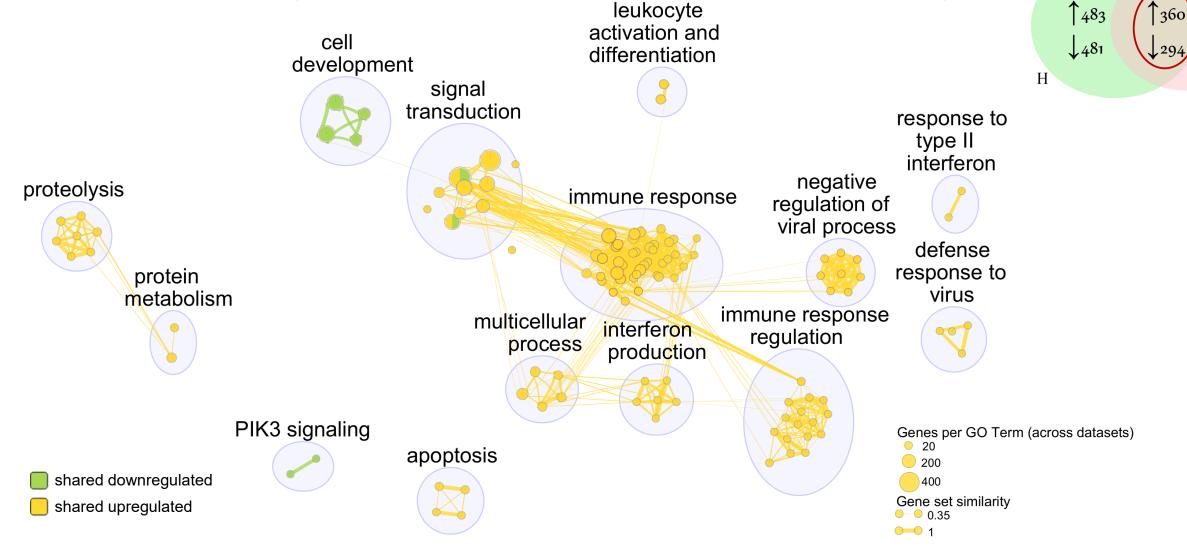
activated DC - broad immune activation and regulation in IBH and healthy

Graduate School for Cellular and Biomedical Sciences

310

,330

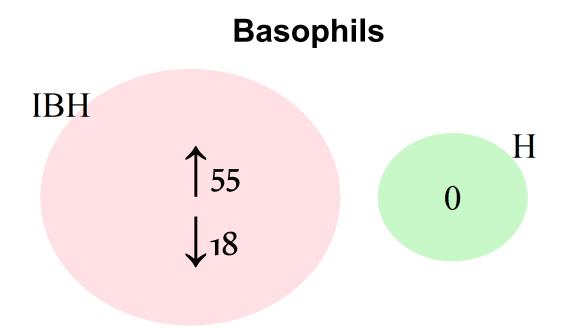
**IBH** 

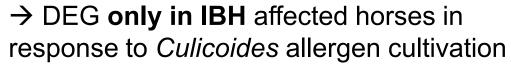


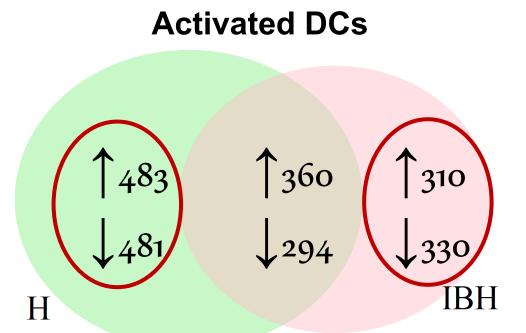
## $u^{^{b}}$

## Shared and unique DEG within horse groups stimulated vs unstimulated









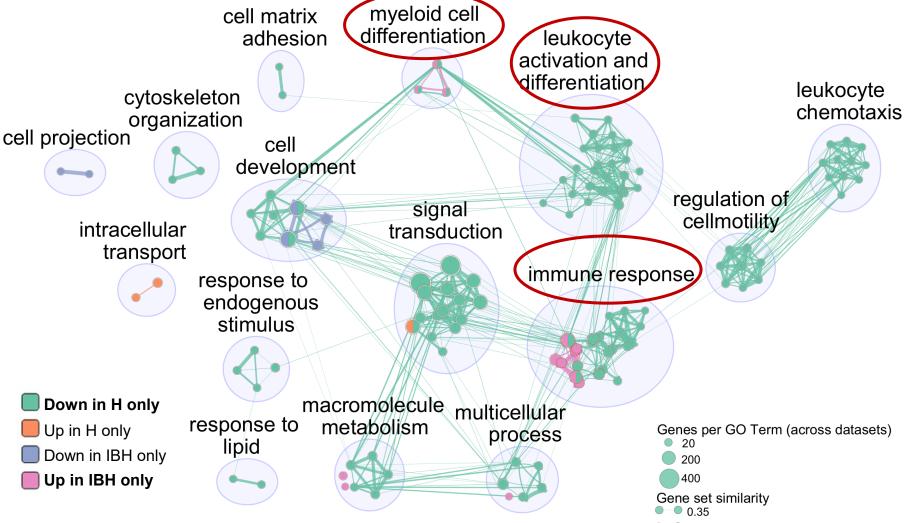
- → enrichment analysis (EA) with g:profiler¹
- → visualization with EnrichmentMap and AutoAnnotate in cytoscape<sup>2</sup>

 $u^{^{\mathsf{b}}}$ 

Distinct transcriptional programs in IBH affected and H activated DC



360



#### Down in H only:

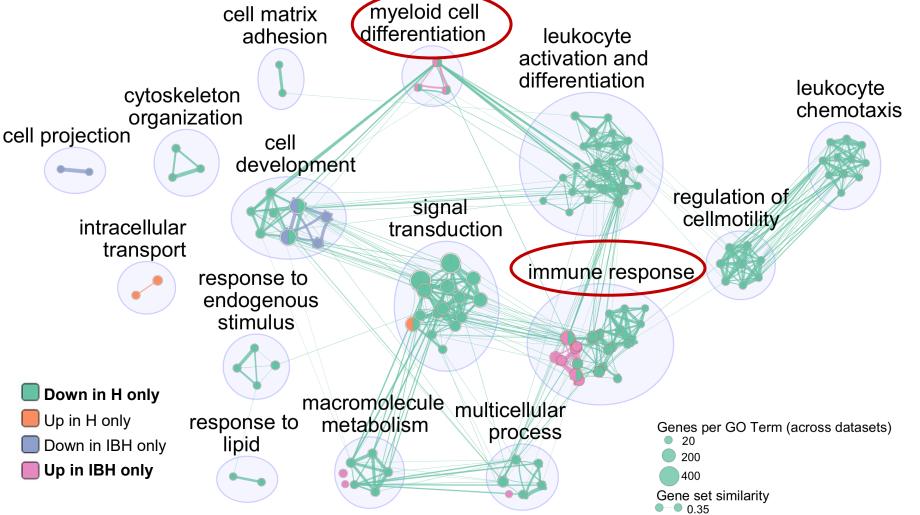
- CSFR1, SPI1
- → DC development and maturation
- IL16, IL2
- → monocyte, eosinophil and CD4+ recruitment
- CD63, CTSS
- → Ag processing and presentation
- TYROBP
- → Signal transduction for DC activation

 $u^{^{\mathsf{b}}}$ 

## Distinct transcriptional programs in IBH affected and H activated DC



360



#### Up in IBH only:

- MYD88, CASP8, UBD, MAFB,
   CEBPB
- → maturation and differentiation
- PTAFR, ITGB8
- → regulatory properties
- CLEC5A, TRIM21
- → pattern recognition and Ag presentation

## $u^{\scriptscriptstyle b}$ Summary and conclusion



#### Stimulation of PBMC with *Culicoides* allergen:

#### **Basophils**

→ activation only in IBH affected horses

#### **Activated DC**

- → broad immune activation in both horse groups
- → Healthy horse specific reduced maturation, T cell recruitment and MHC II antigen presentation
- → IBH specific enhanced antigen presentation, maturation, pattern recognition, and regulatory properties

Activated dendritic cells in IBH affected horses bear a transcriptional program associated with enhanced specialization and functionality possibly shaped by antigen specific IgE mediated basophil activation

### $u^{^{b}}$

### Thank you for your attention! - Questions?





Image generated by ChatGPT (OpenAI)

#### Acknowledgements

- Prof Dr Eliane Marti (Supervisor)
- Stephanie Talker, PhD (Coadvisor)
- Vidhya Jagannathan, PhD (bioinformatic supervision)
- Experimental Pathology, University of Iceland, Keldur (collaboration)
- NGS Platform University of Bern
- Dorontina Mahmuti and Delia Catherine Bogenstätter (Lab. technicians)
- Swiss Assocciation for Animal Science (financial support)

Organisationseinheit