

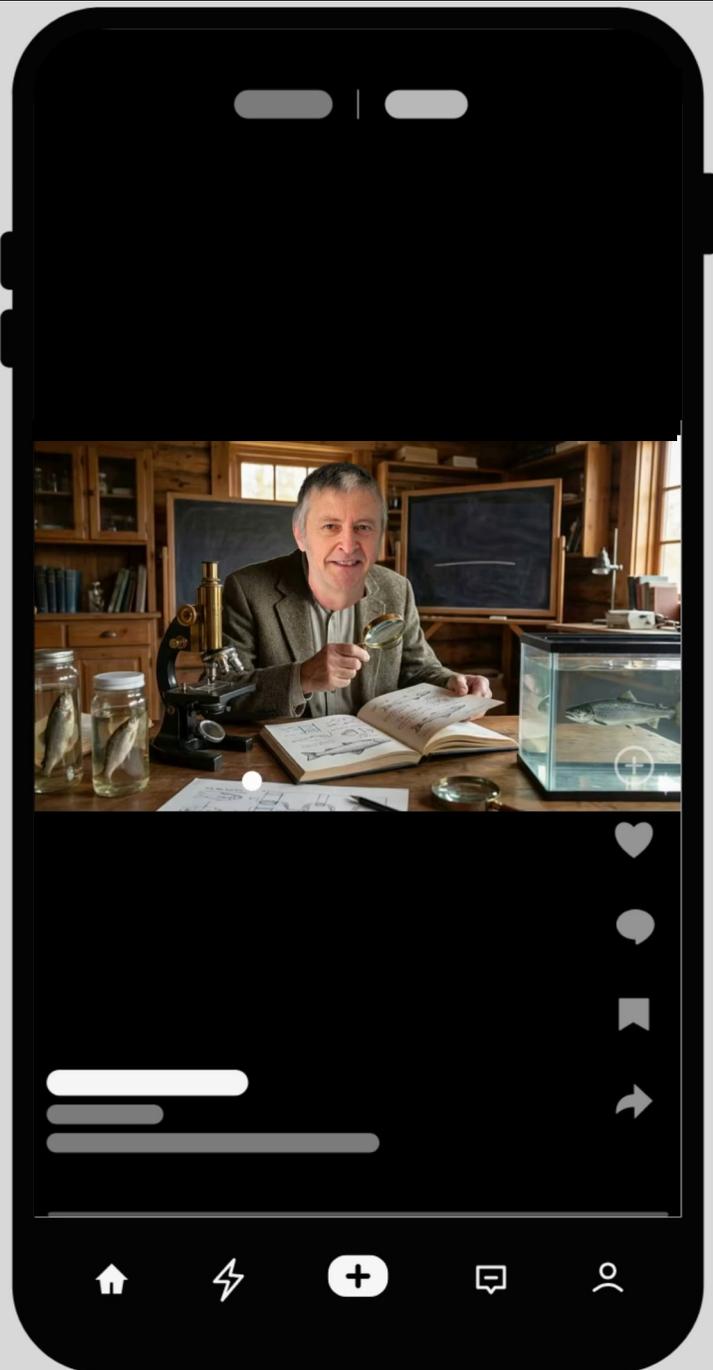
Stillehavslaksen har løsningen, hvordan kan vi gi den videre til Atlantisk laks?

Crisp ResisT et FHF finansiert prosjekt

Lene Sveen, Mark D. Fast, Torstein Tengs, Rachel A. Kline, Judit Aguilar Marti, Dominic Kurian, Gerrit Timmerhaus, Marianne Vaadal, Ross D. Houston, James E. Bron, Sean J. Monaghan, Haitham H. Mohammed, Rose Ruiz Daniels, Sarah Salisbury, Diego Robledo, Mark Braceland, Miroslava Hansen, Anne Wargelius, Teshome Tilahun, Ajna Ødegård, Tone Kari Knudsdatter, Matt Watsmuth, Nicholas Robinson



Alle prosjekter starter med en idè....



Nick Robinson

Nofima

Jeg har en drøm

- 1.Smitte naturlig luseresistente laksearter med lus
2. Identifisere genene som er involvert
3. Validere genene ved hjelp av genredigering
4. Bruke denne informasjonen til å gjøre Atlantisk laks mer lik stillehavslaks

Med målrettet avl, vaksinerings eller genredigering

CRISP RESIST



CRISP RESIST





BAKGRUNN

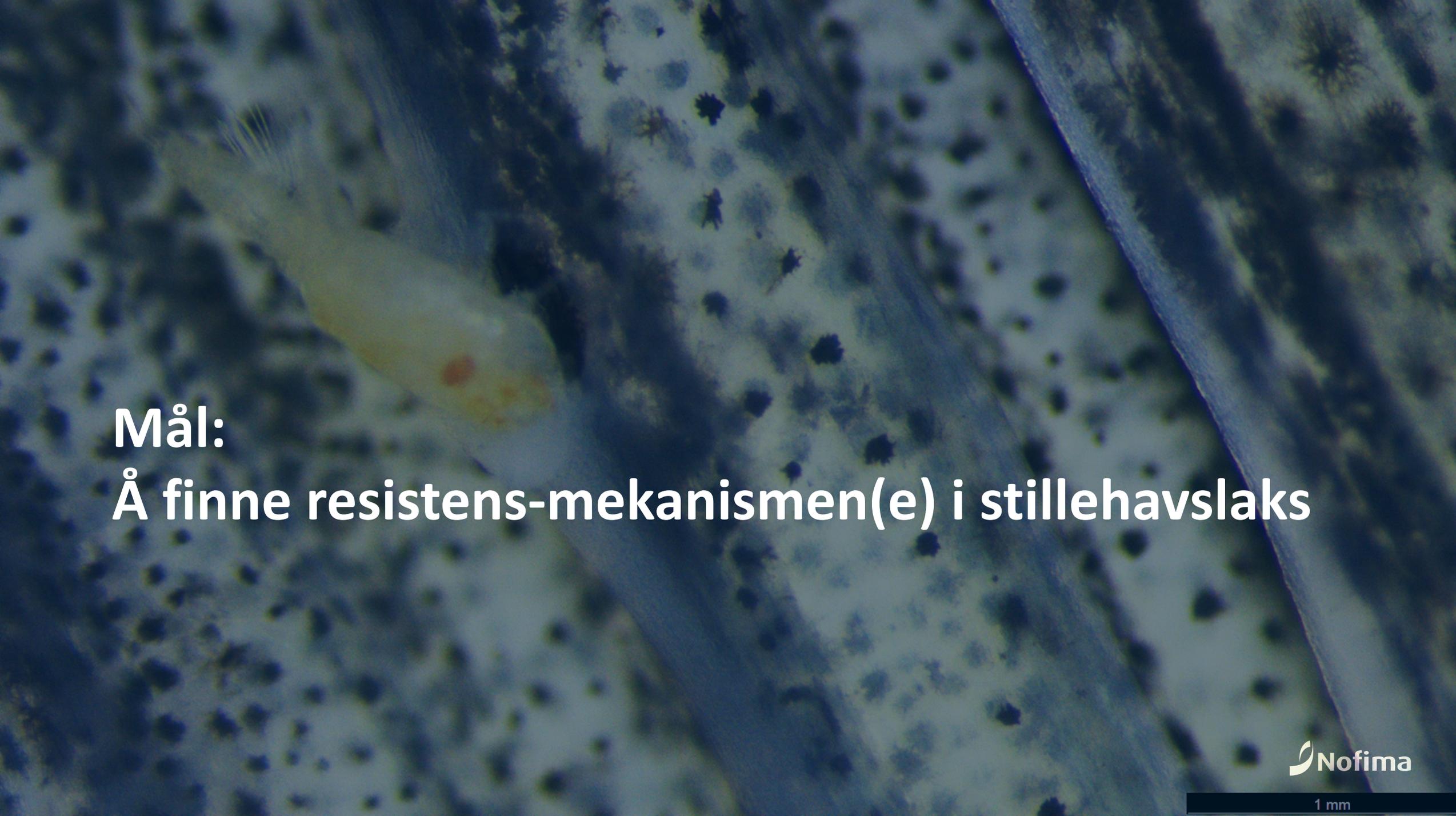
FORSØKSDESIGN

RESULTATER

GENREDIGERING

VEIEN VIDERE





Mål:
Å finne resistens-mekanismen(e) i stillehavslaks

Stillehavslaks

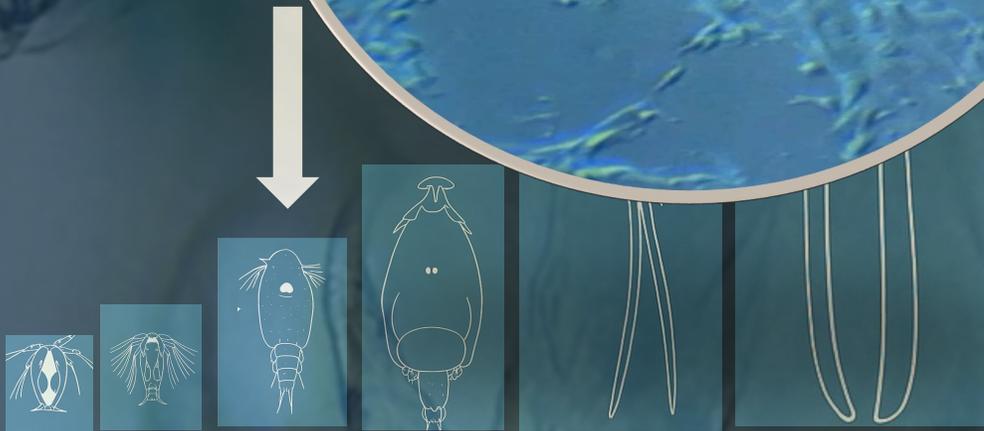
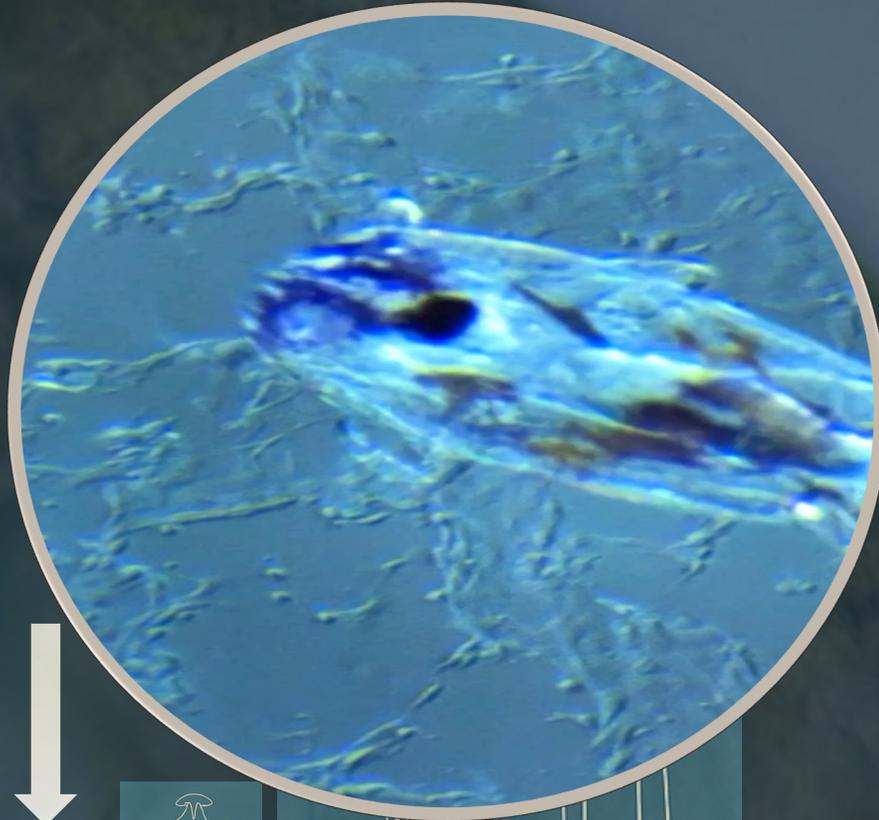
Pink salmon (Pukkel laks)

Chum salmon (ketalaks)

Coho salmon (Sølv laks)

Atlantic salmon (Atlantisk laks)

Forsøksdesign



Mark FAST

UPEI, Canada

Ingen lus på Chum eller Pink (Forsøk 1)

24 timer



12 timer

Ingen lus på Chum eller Pink (Forsøk 2)

24 timer



12 timer

All fisk bedøvet og smittet med lus (forsøk 3)



12 timer

Forsøksdesign



Forsøksdesign



7 dager

BAKGRUNN

FORSØKSDESGIN

RESULTATER

GENREDIGERING

VEIEN VIDERE



BAKGRUNN

FORSØKSDESGIN

RESULTATER

GENREDIGERING

VEIEN VIDERE

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Keratinocytes drive the epithelial hyperplasia key to sea lice resistance in coho salmon

Research article | [Open access](#) | Published: 29 July 2024

Volume 22, article number 160, (2024) | [Cite this article](#)

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> [Cell Tissue Res.](#) 2025 Aug;401(2):181-211. doi: 10.1007/s00441-025-03976-0. Epub 2025 Jun 4.

Local inflammation at the salmon louse (*Lepeophtheirus salmonis*) attachment site contributes to copepodid rejection in coho salmon (*Oncorhynchus kisutch*)

Lene Sveen ¹, Mark D Fast ², Torstein Tengs ³, Rachel A Kline ⁴, Judit Aguilar Marti ⁴, Dominic Kurian ⁴, Gerrit Timmerhaus ³, Marianne Vaadal ³, Ross D Houston ⁵, James E Bron ⁶, Sean J Monaghan ⁶, Haitham H Mohammed ^{7 8}, Rose Ruiz Daniels ^{6 4}, Sarah Salisbury ⁴, Diego Robledo ⁴, Mark Braceland ^{7 9}, Miroslava Hansen ^{3 10}, Nicholas Robinson ^{3 11}

Affiliations + expand

PMID: 40464919 | PMCID: [PMC12325551](#) | DOI: [10.1007/s00441-025-03976-0](#)

Abstract

The study investigates the susceptibility of Atlantic salmon (*Salmo salar*) and Pacific salmon species (pink salmon, *Oncorhynchus gorbuscha*; coho salmon, *Oncorhynchus kisutch*; and chum salmon, *Oncorhynchus keta*) to the parasitic salmon lice (*Lepeophtheirus salmonis*). The research had two main objectives: to characterize the morphology of the scaly skin in four salmonid species and to compare the cellular response at the louse attachment site in coho salmon and Atlantic salmon. Three consecutive challenge trials were conducted, with successful louse infestation only achieved across all four species in the third trial using mild anesthesia with tricaine methanesulfonate. Skin and fin samples were collected at 12, 24, 36, 48, 60, and 168 h post-

[S. J. Salisbury](#)

[Sveen, R. D. H](#)

 4280 Acc

Abstract

Background

Tre forsøk på å smitte fisken



Chum og pink
merker at lusa biter,
må bedøves

Differential rejection of salmon lice by pink and chum salmon: disease consequences and expression of proinflammatory genes

[Simon R M Jones](#)¹, [Mark D Fast](#), [Stewart C Johnson](#), [David B Groman](#)

Affiliations + expand

PMID: 17629118 DOI: [10.3354/dao075229](#)

[Free article](#)

Abstract

Comparative transcriptomics of Atlantic *Salmo salar*, chum *Oncorhynchus keta* and pink salmon *O. gorbuscha* during infections with salmon lice *Lepeophtheirus salmonis*

[Ben J G Sutherland](#), [Kim W Koczka](#), [Motoshige Yasuike](#), [Stuart G Jantzen](#), [Ryosuke Yazawa](#), [Ben F Koop](#)¹, [Simon R M Jones](#)

Affiliations + expand

PMID: 24628956 PMCID: [PMC4004277](#) DOI: [10.1186/1471-2164-15-200](#)

Abstract

Background: Salmon species vary in susceptibility to infections with the salmon louse (*Lepeophtheirus salmonis*). Comparing mechanisms underlying responses in susceptible and resistant species is important for estimating impacts of infections on wild salmon, selective breeding of farmed salmon, and expanding our knowledge of fish immune responses to ectoparasites. Herein we report three *L. salmonis* experimental infection trials of co-habited Atlantic *Salmo salar*, chum *Oncorhynchus keta* and pink salmon *O. gorbuscha*, profiling hematocrit, blood cortisol concentrations, and transcriptomic responses of the anterior kidney and skin to the infection.

Results: In all trials, infection densities (lice per host weight (g)) were consistently highest on chum salmon, followed by Atlantic salmon, and lowest in pink salmon. At 43 days post-exposure, all lice had developed to motile stages, and infection density was uniformly low among species. Hematocrit was reduced in infected Atlantic and chum salmon, and cortisol was elevated in infected chum salmon. Systemic transcriptomic responses were profiled in all species and large differences in response functions were identified between Atlantic and Pacific (chum and pink) salmon. Pink and chum salmon up-regulated acute phase response genes, including complement and coagulation components, and down-regulated antiviral immune genes. The pink salmon response involved the largest and most diverse iron sequestration and homeostasis mechanisms.

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theirus salmonis copepodids
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of unexposed chum. Neither
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ties following either
21 d after low exposure. An
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1beta) in fin and head kidney
n this species. Together,
L. salmonis in the juvenile

Resultater

Chum og pink
merker at **lusa biter**,
må bedøves

Kjempespennende

> [Dis Aquat Organ. 2007 May 9;75\(3\):229-38. doi: 10.3354/dao075229.](#)

Differential rejection of salmon lice by pink and chum salmon: disease consequences and expression of proinflammatory genes

Simon R M Jones ¹, Mark D Fast, Stewart C Johnson, David B Groman

Affiliations + expand

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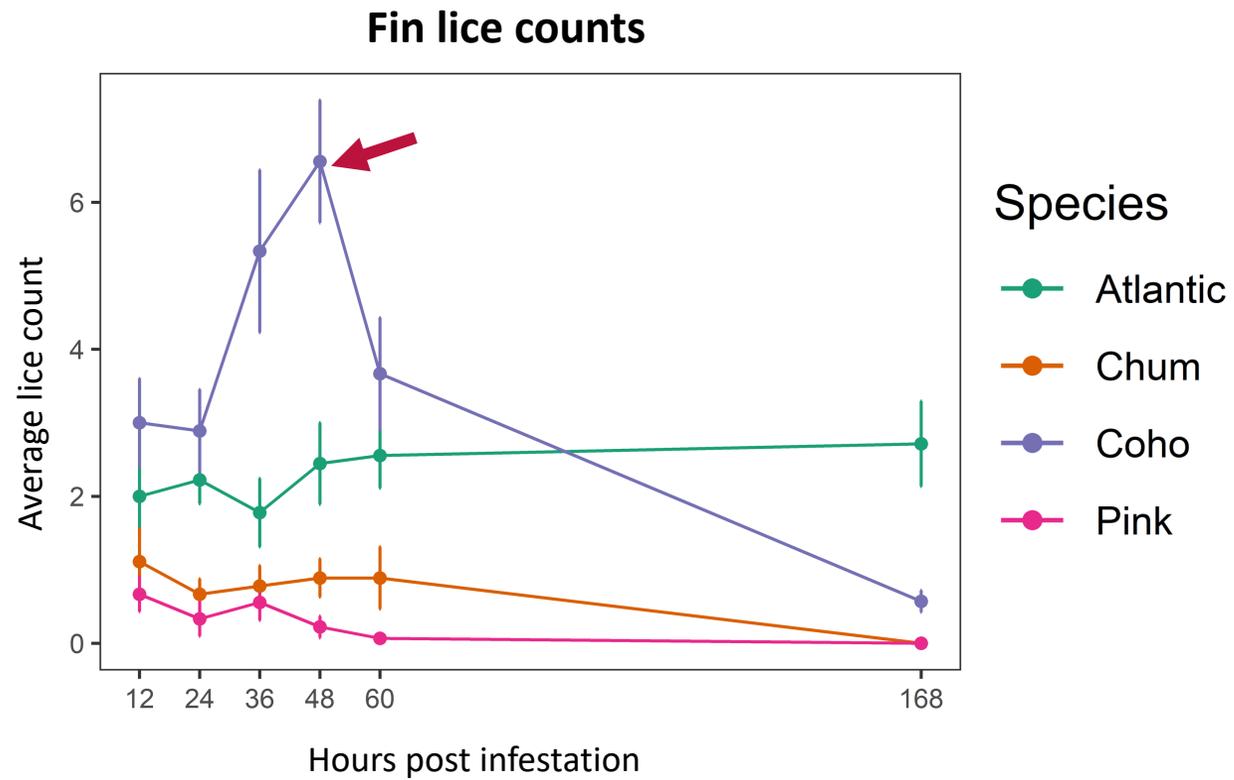
They shake it off!
(How?)

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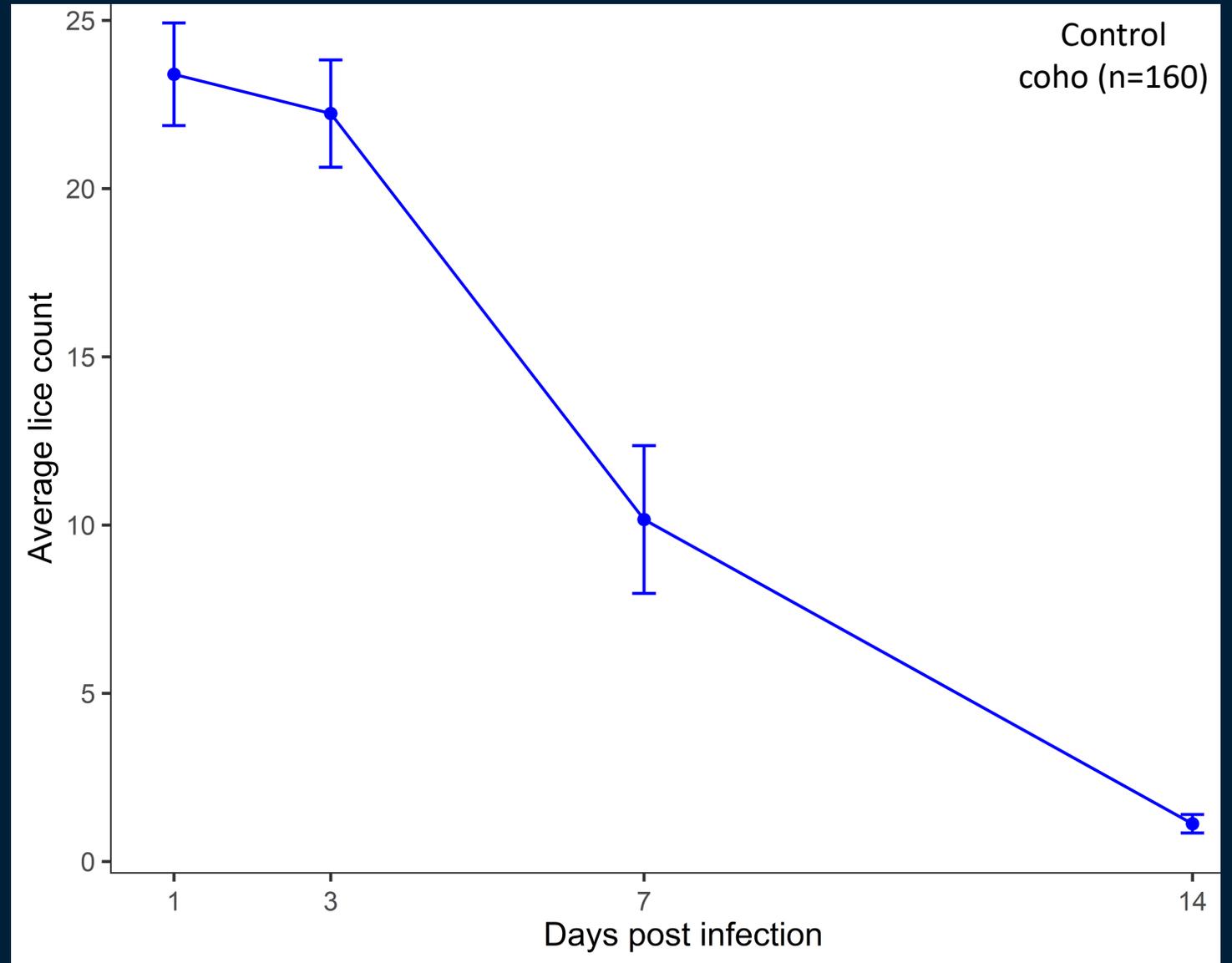
Resultater



Flest lus på Coho

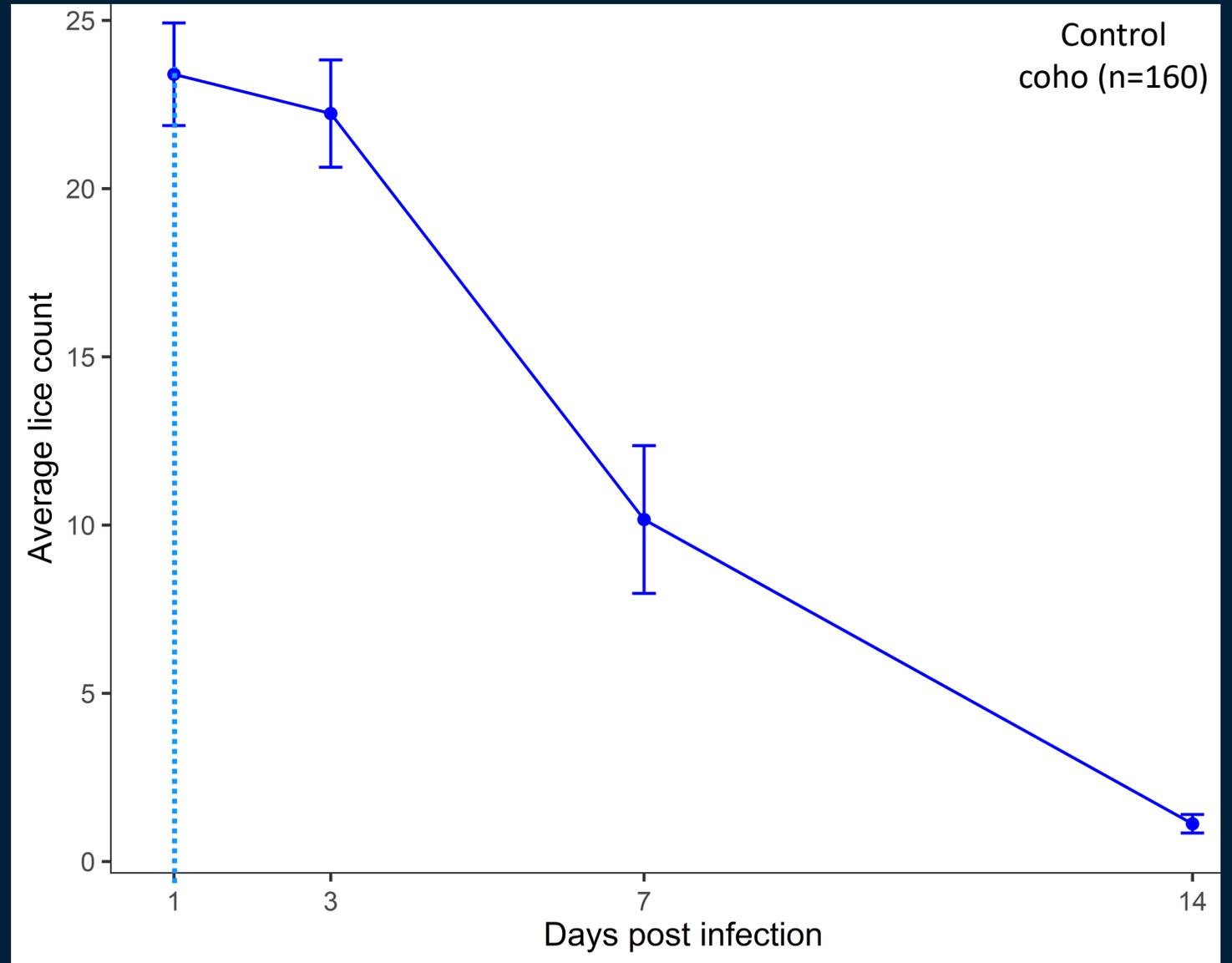
Resultater

1. Lusa setter seg på coho laks



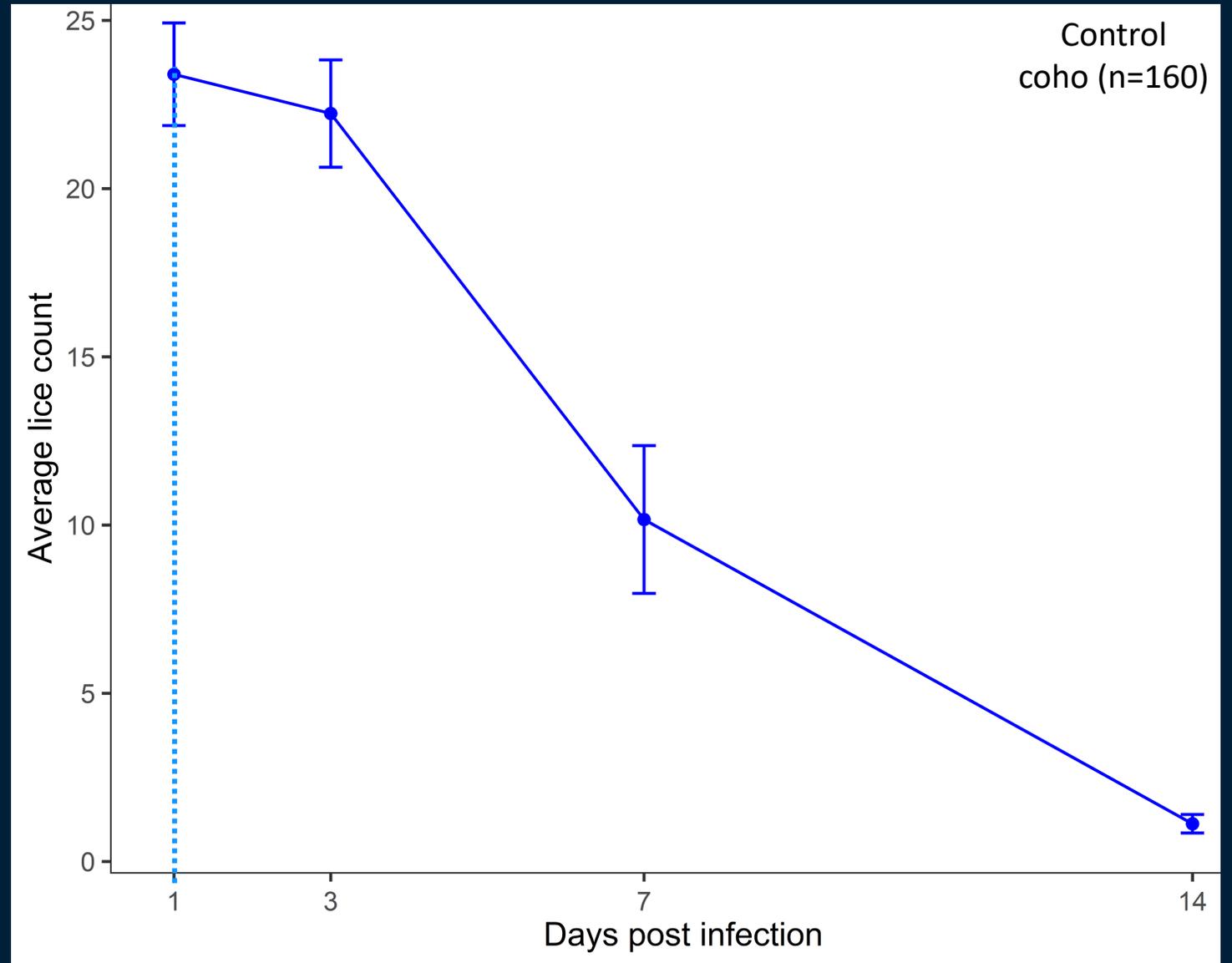
Resultater

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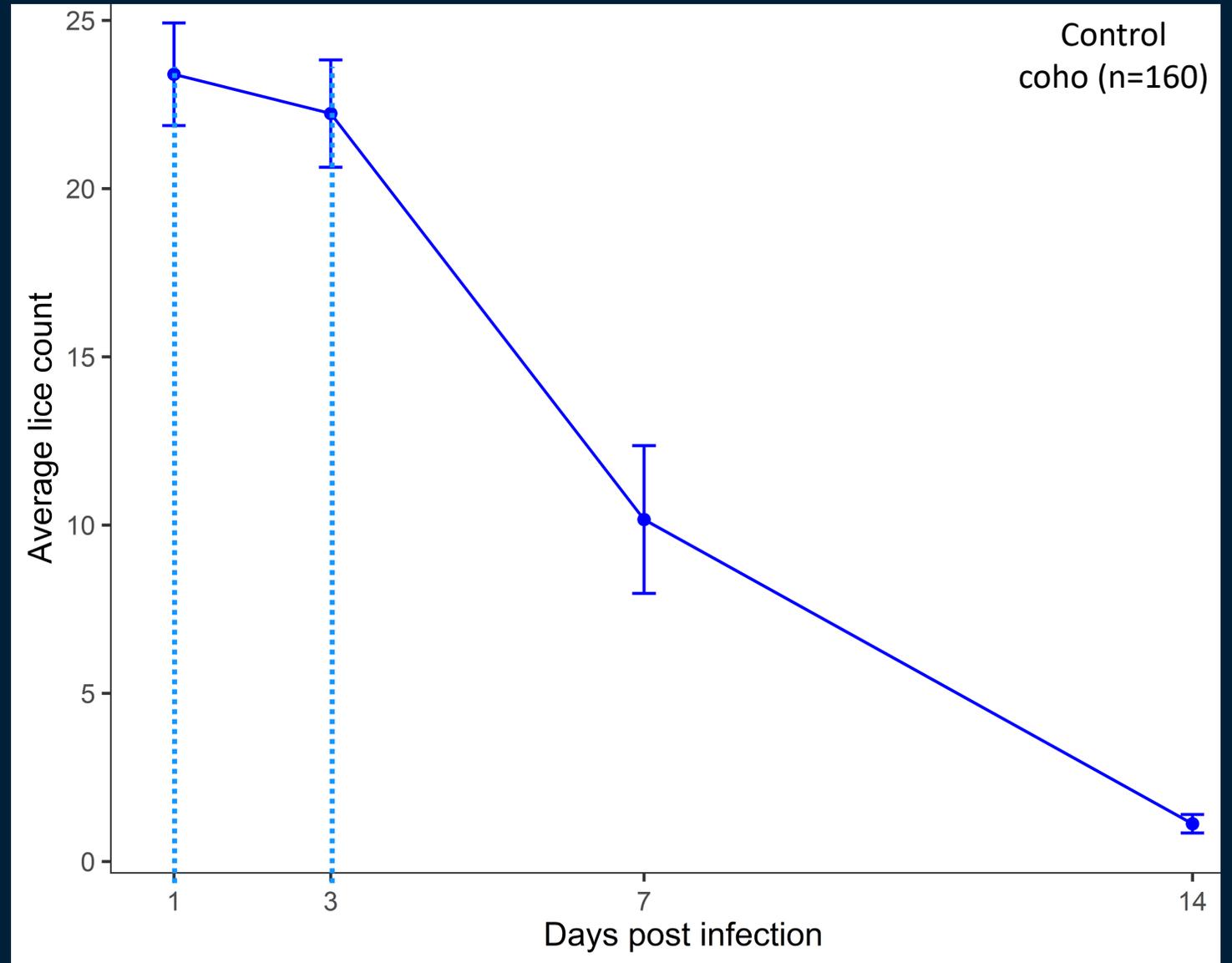
Resultater

1. Responsen starter umiddelbart



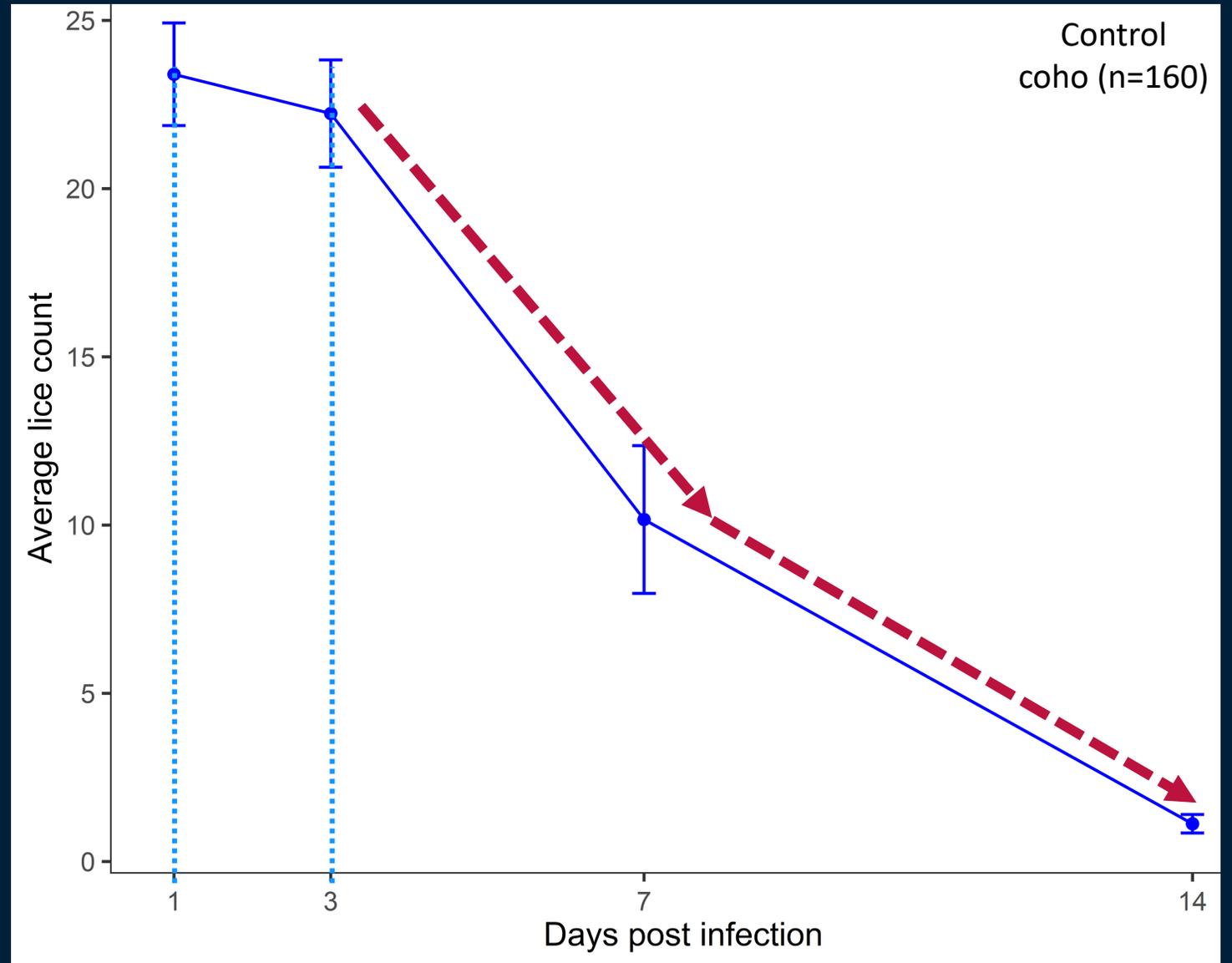
Resultater

2. Gradvis redusksjon i lusetall



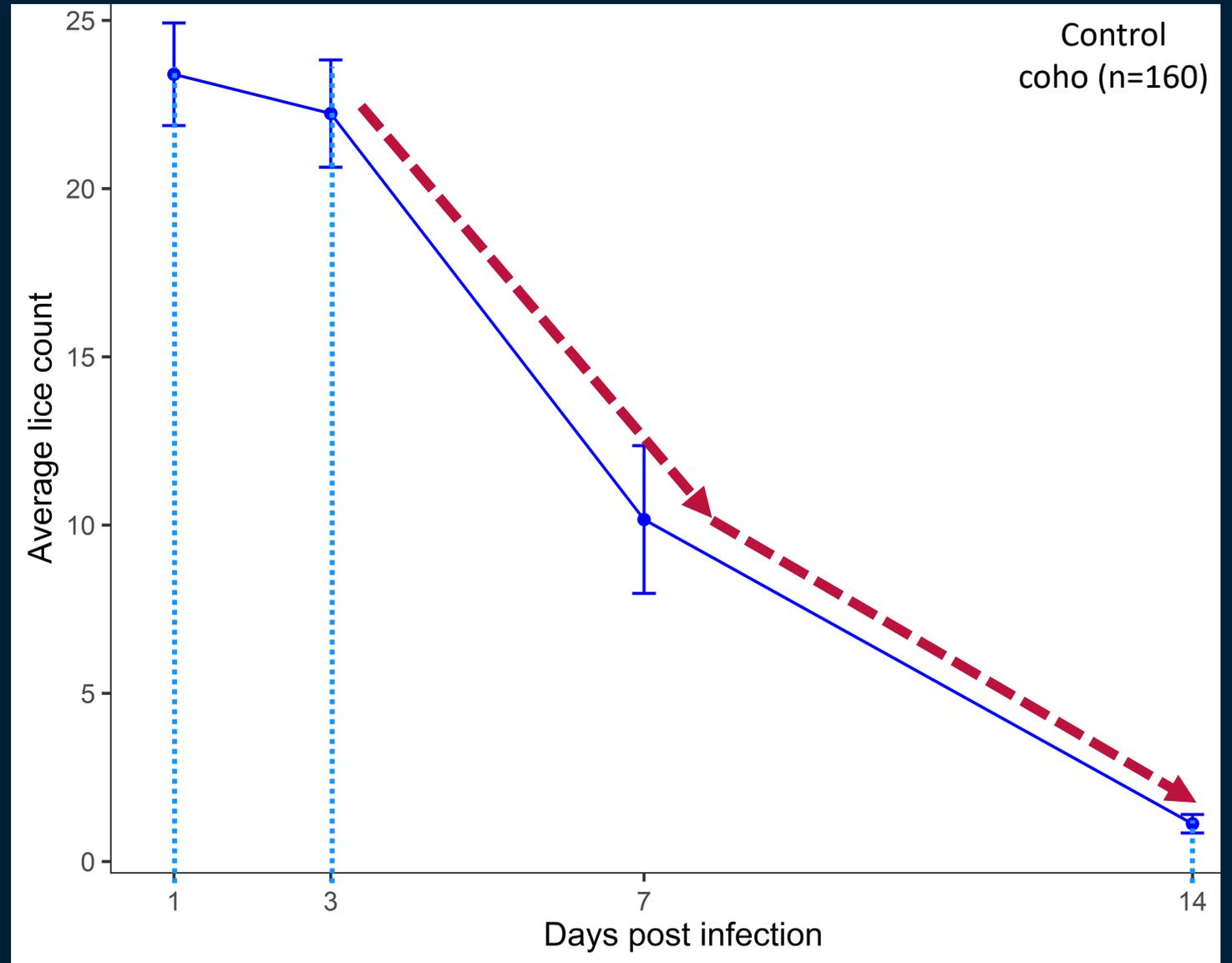
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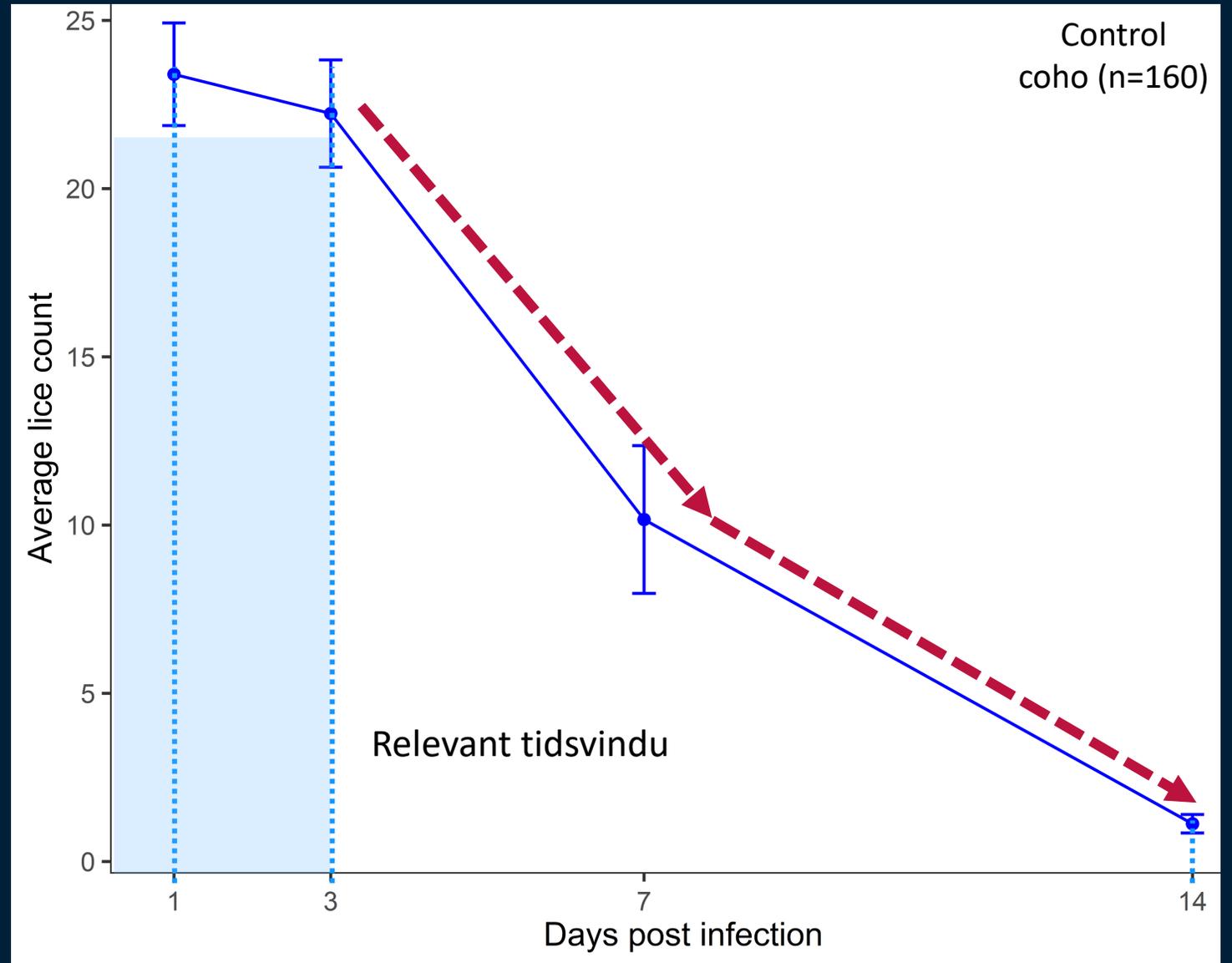
Resultater

3. Nesten ingen lus etter 14 dager



Resultater

3. Nesten ingen lus etter 14 dager



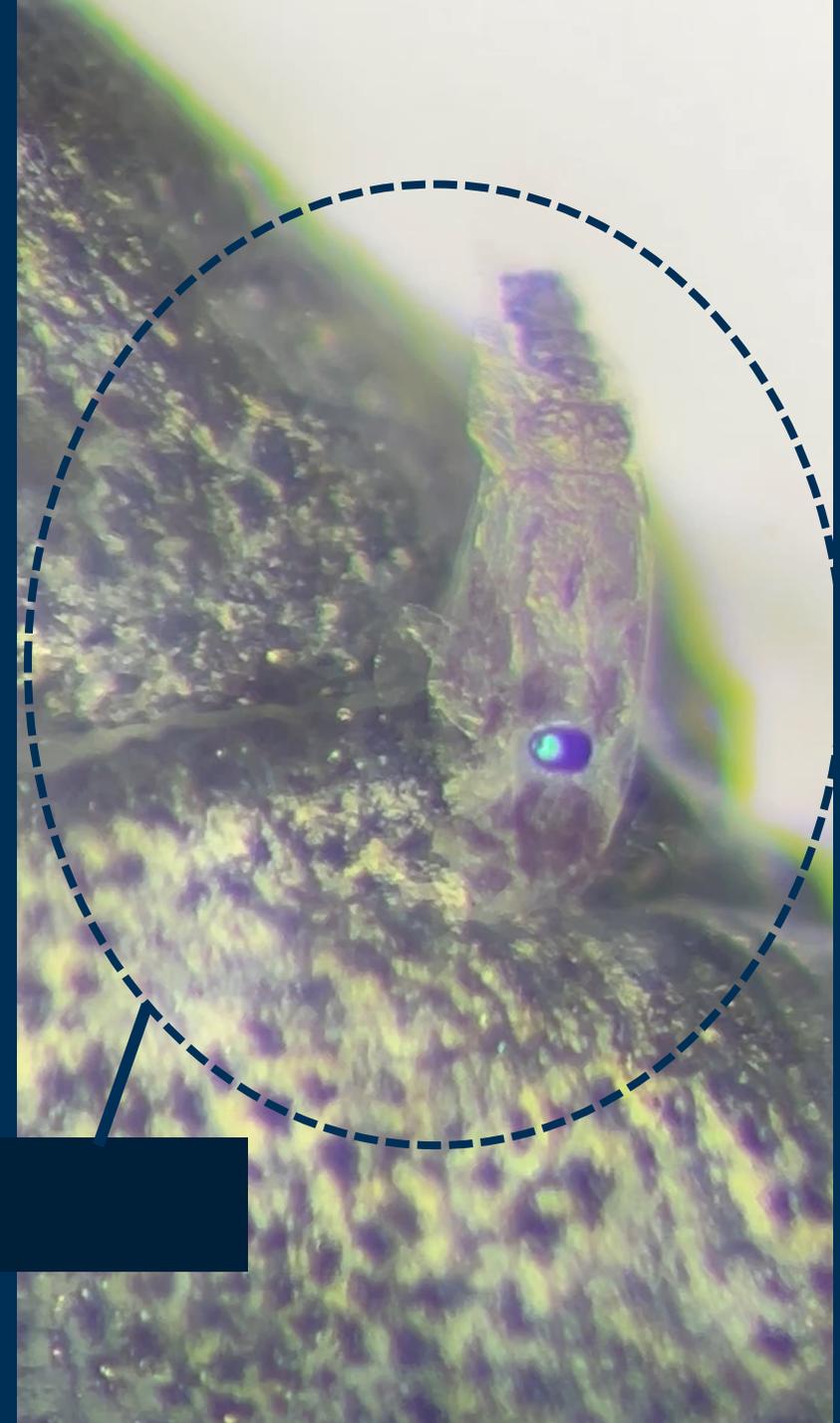
Resultater



Sarah Salisbury
Roslin institute
29



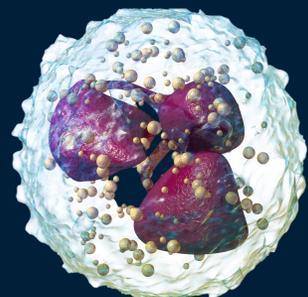
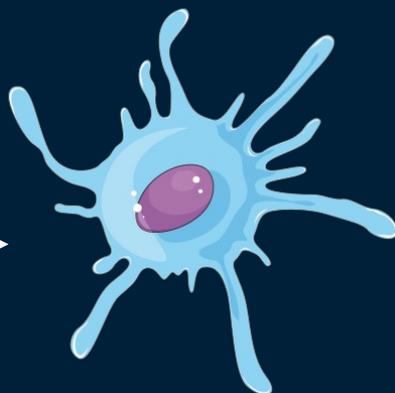
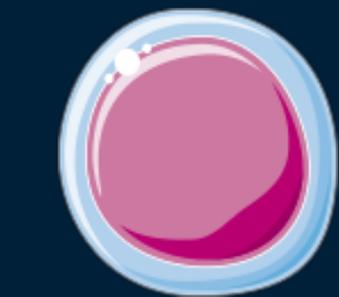
Rose R. Daniels
Roslin institute



Encellekjerne sekvensering

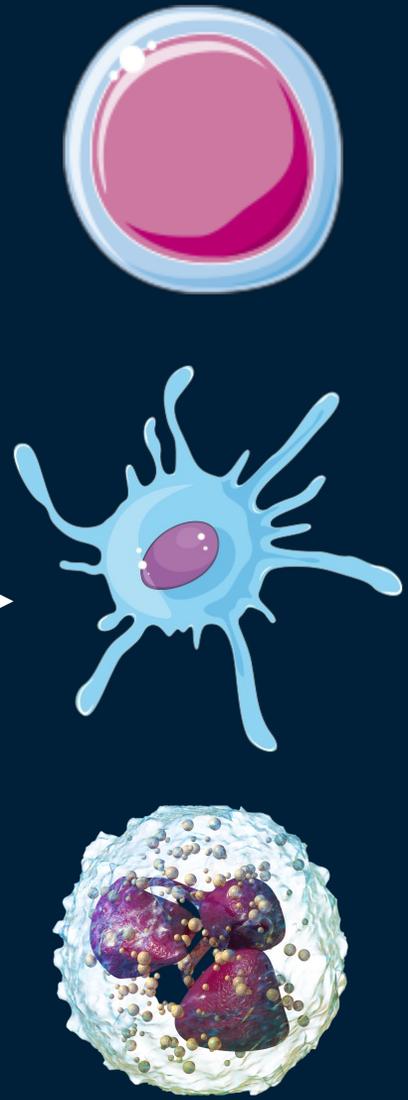
Resultater

Fiskeskinn /
finne



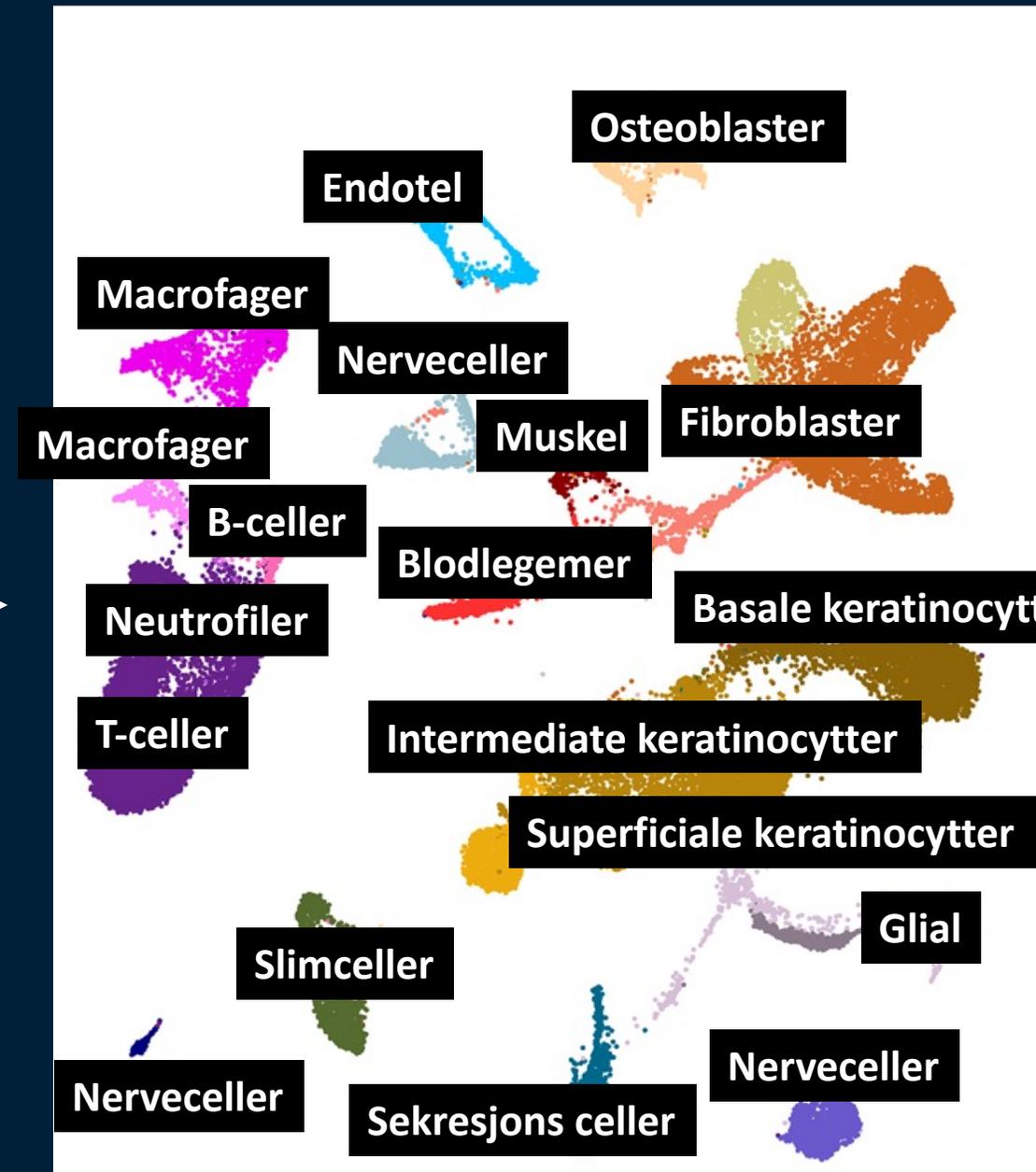
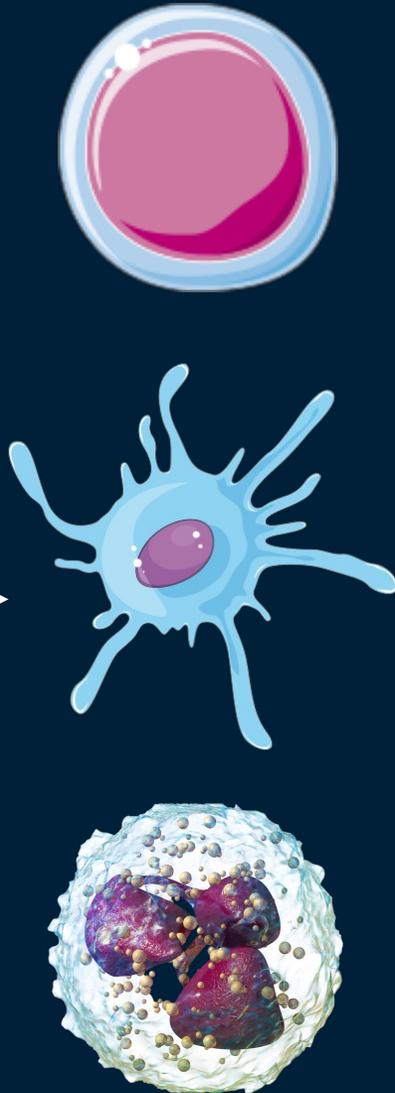
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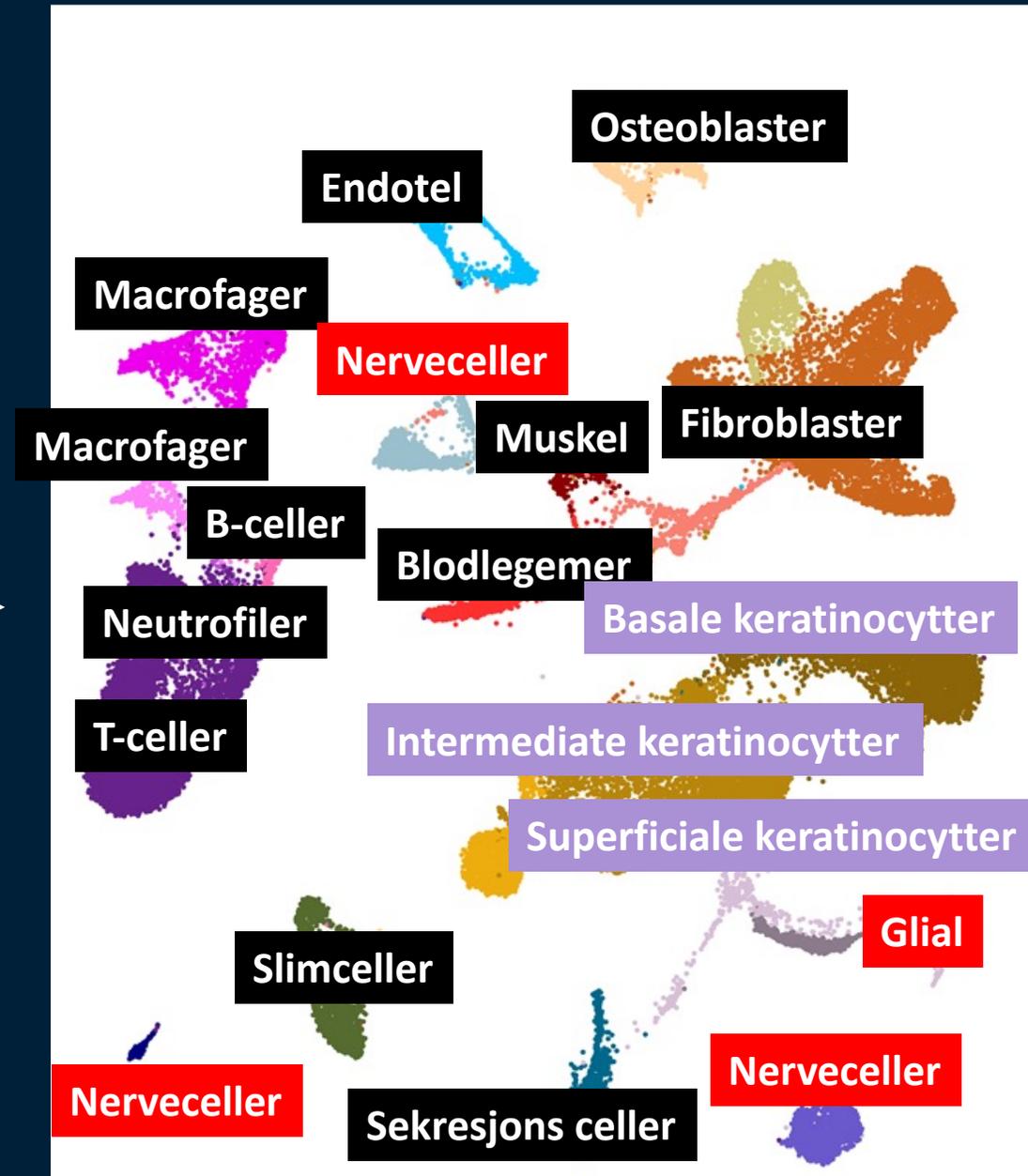
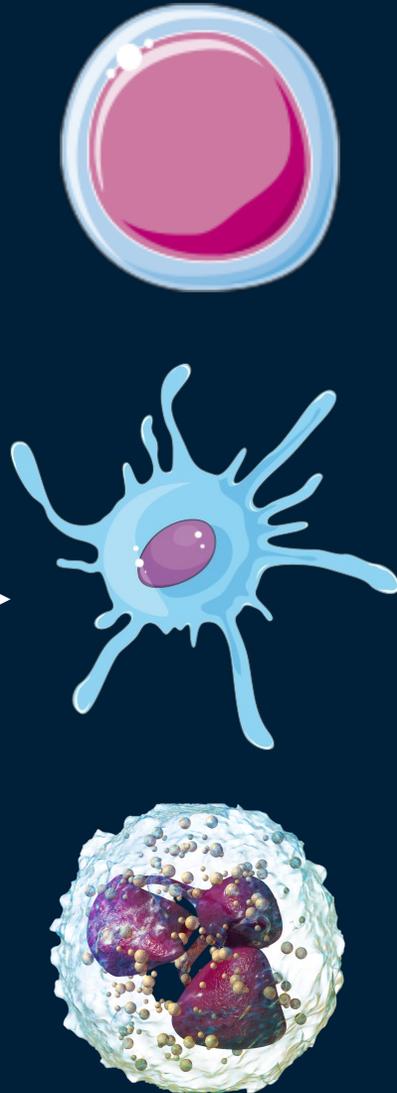
Resultater

Fiskeskinn /
finne



Resultater

Fiskeskinn /
finne

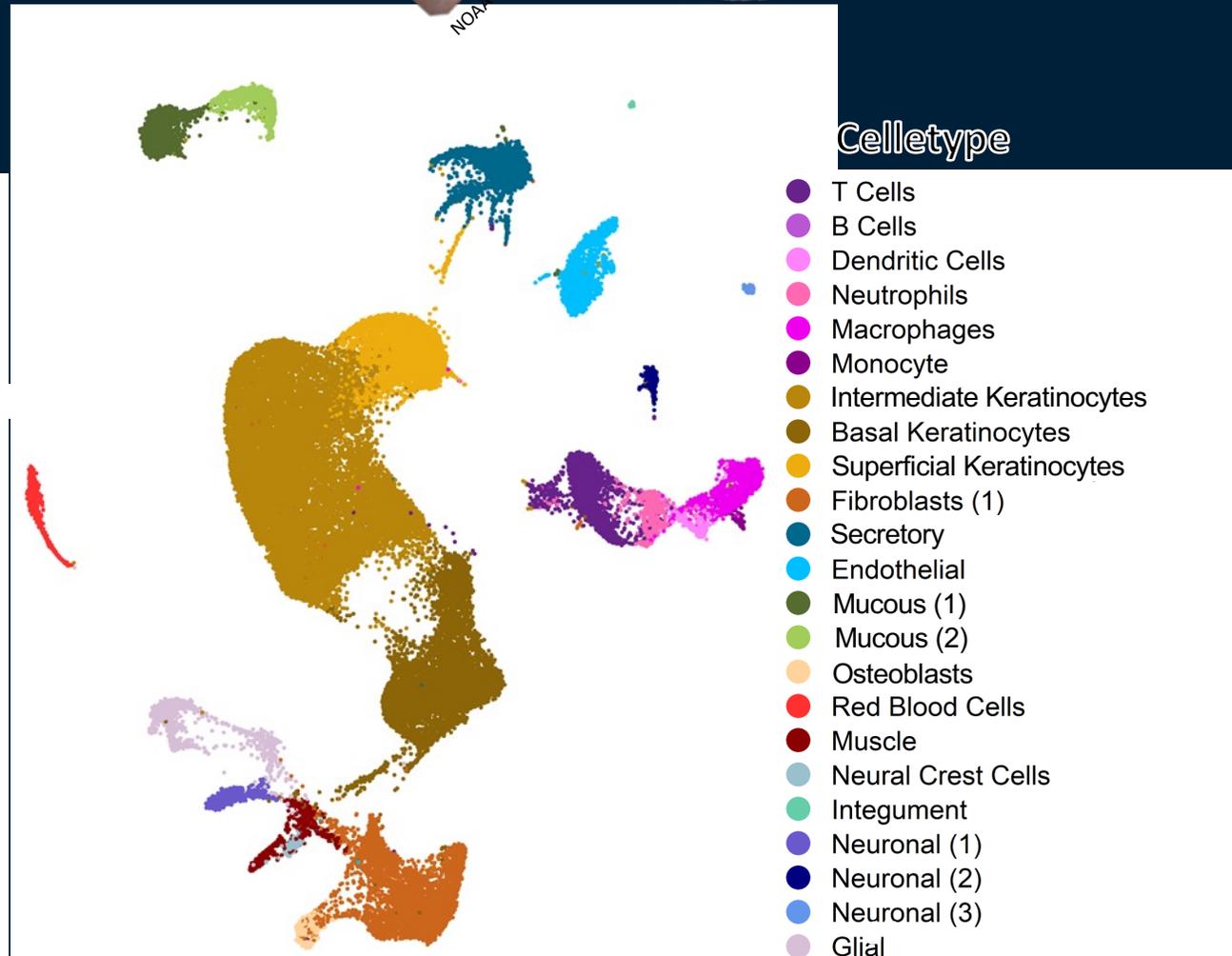
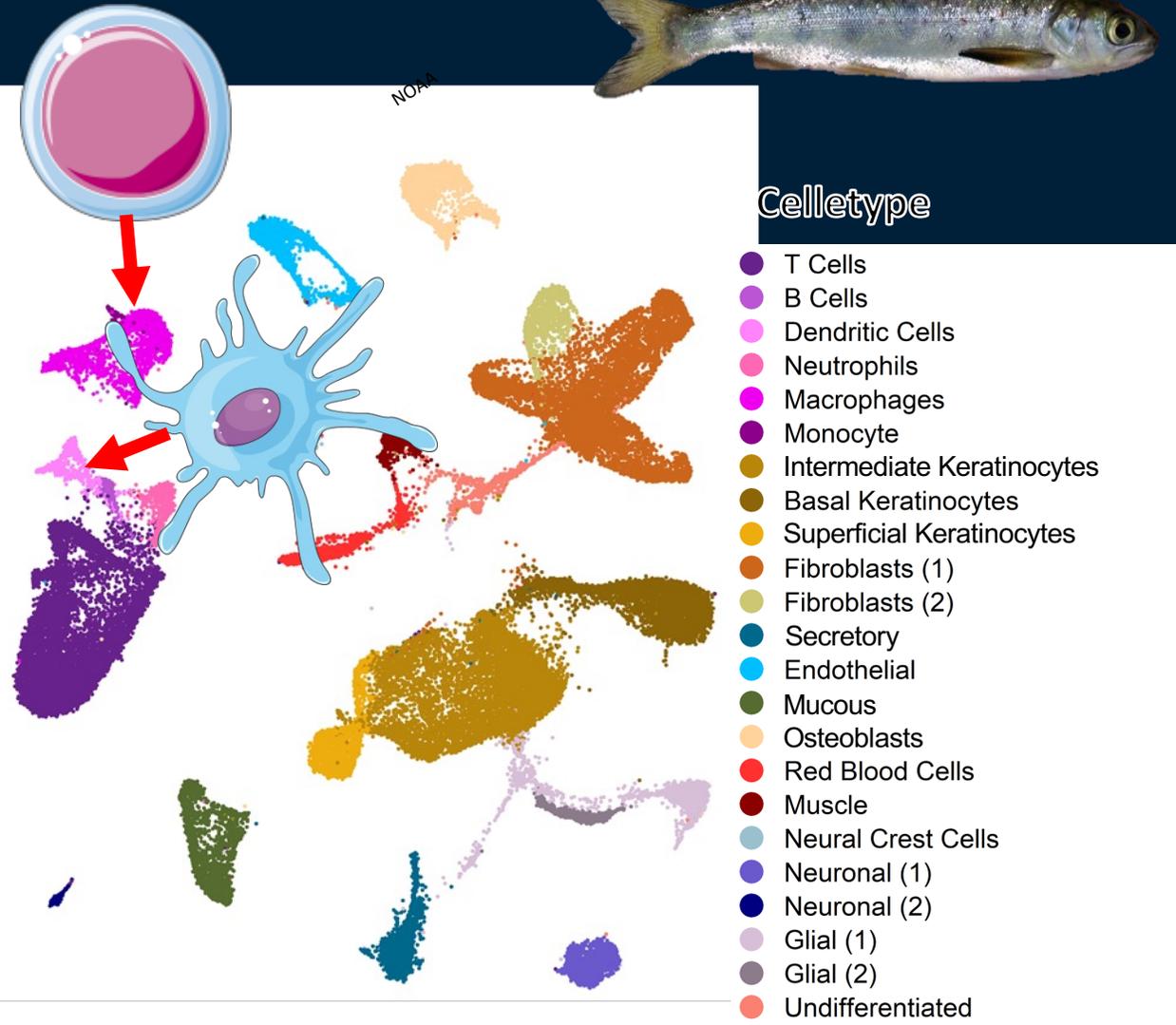


Resultater

ATLANTIC



COHO

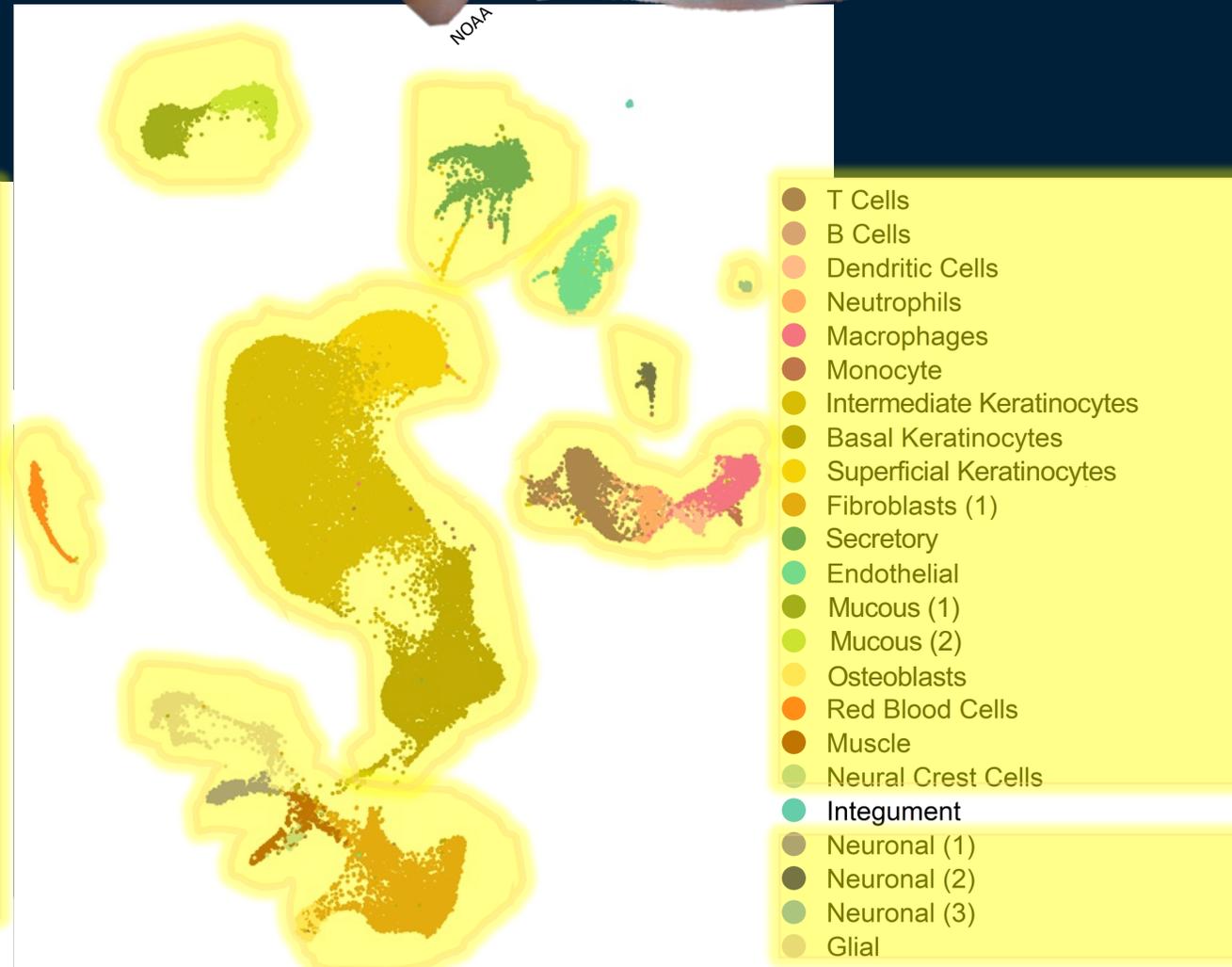
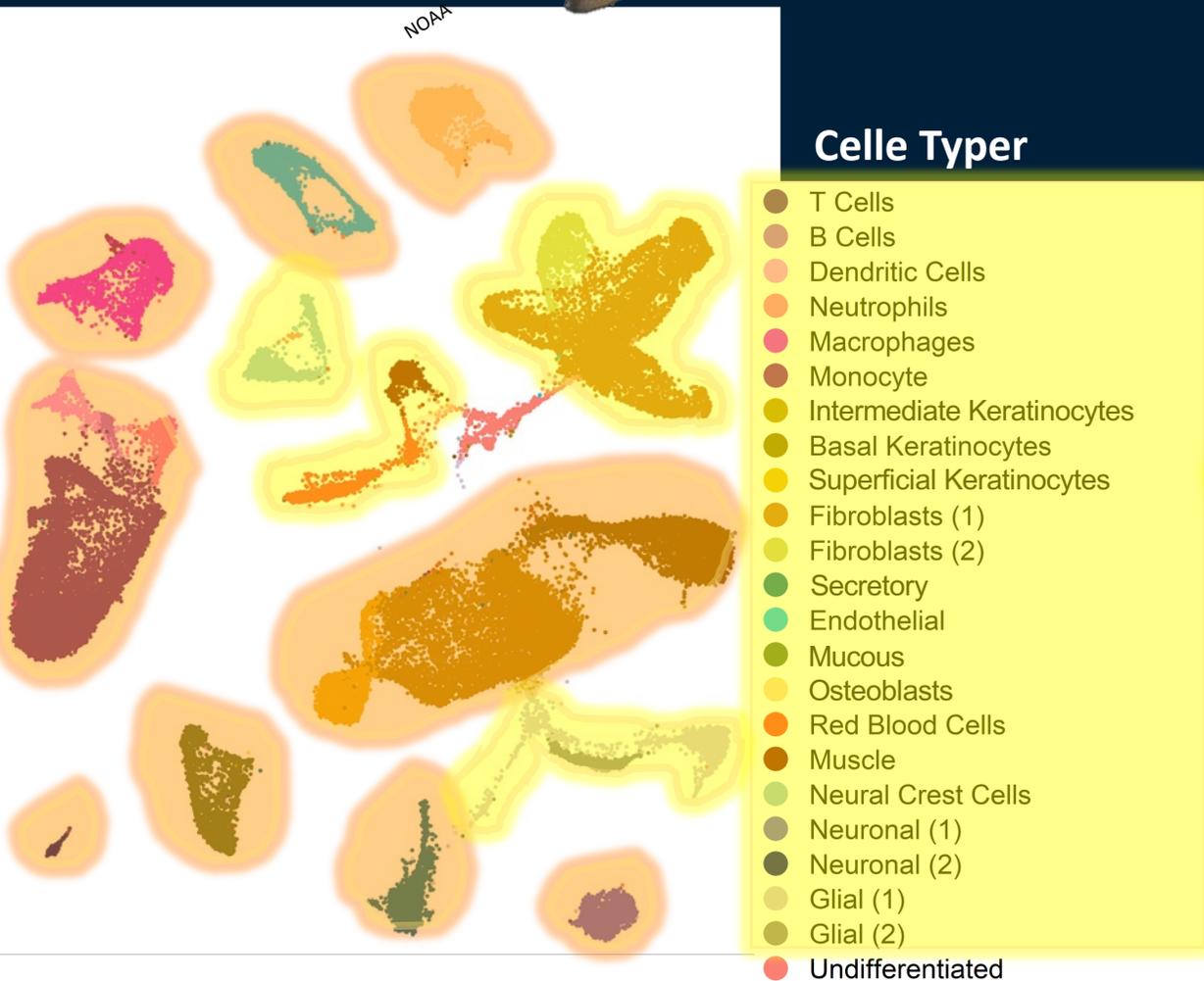


Resultater

ATLANTIC



COHO



CHUM



PINK



Cell Types



- T Cells
- B Cells
- Dendritic Cells
- Neutrophils
- Macrophages
- Monocytes
- Superficial Keratinocytes
- Intermediate Keratinocytes
- Basal Keratinocytes
- Fibroblasts (1)
- Fibroblasts (2)
- Endothelial
- Secretory
- Mucous
- Red Blood Cells
- Muscle (1)
- Muscle (2)
- Neural Crest Cells
- Neuronal (3)
- Collagen Enriched Cells

Cell Types



- T Cells
- B Cells
- Dendritic Cells
- Neutrophils
- Macrophages
- Superficial Keratinocytes
- Intermediate to Superficial Keratinocytes
- Intermediate Keratinocytes
- Basal Keratinocytes
- Fibroblasts
- Endothelial
- Secretory
- Mucous
- Red Blood Cells
- Muscle
- Muscle Satellite Cells
- Neural Crest Cells
- Neuronal (2)
- Glial

CHUM



PINK



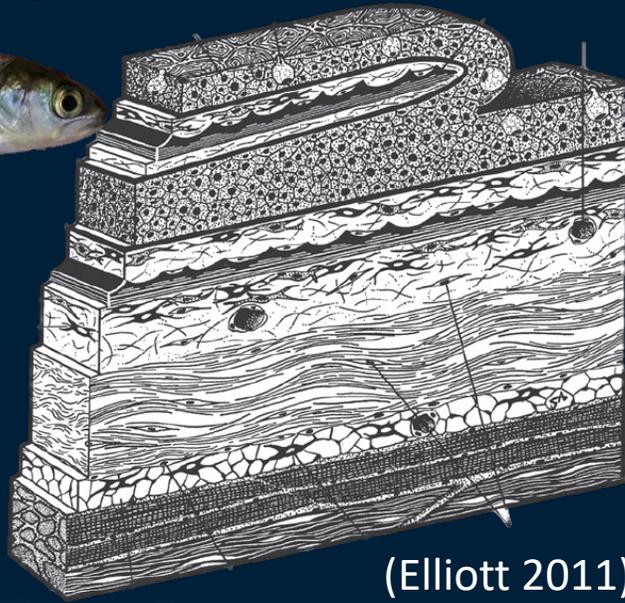
Cell Types

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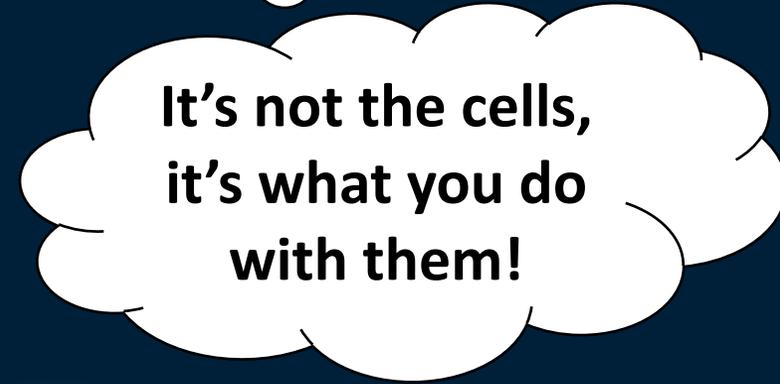
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- Muscle
- Muscle Satellite Cells
- Neural Crest Cells
- Neuronal (2)
- Glial

ATLANTIC



(Elliott 2011)

COHO



It's not the cells, it's what you do with them!

Alle fire arter har samme celletyper

CHUM



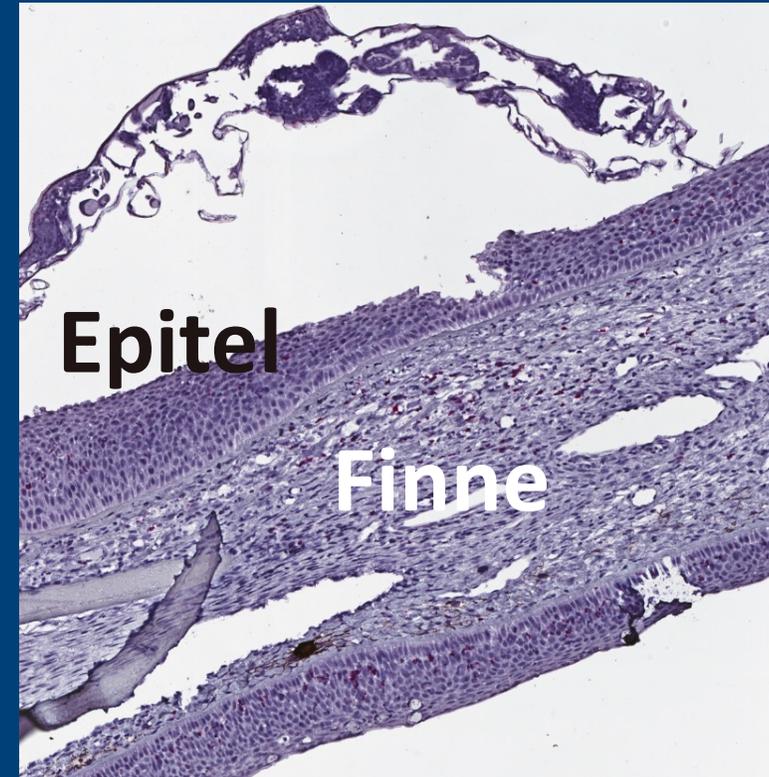
PINK



Isn't that a bit boring?

Resultater

Epitelcellene (Keratinocytter)



Resultater

Epitelcellene (Keratinocytter)



Resultater



Basal keratinocytes



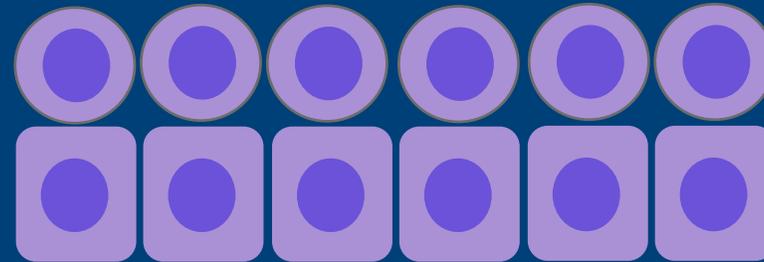
Finne

Resultater



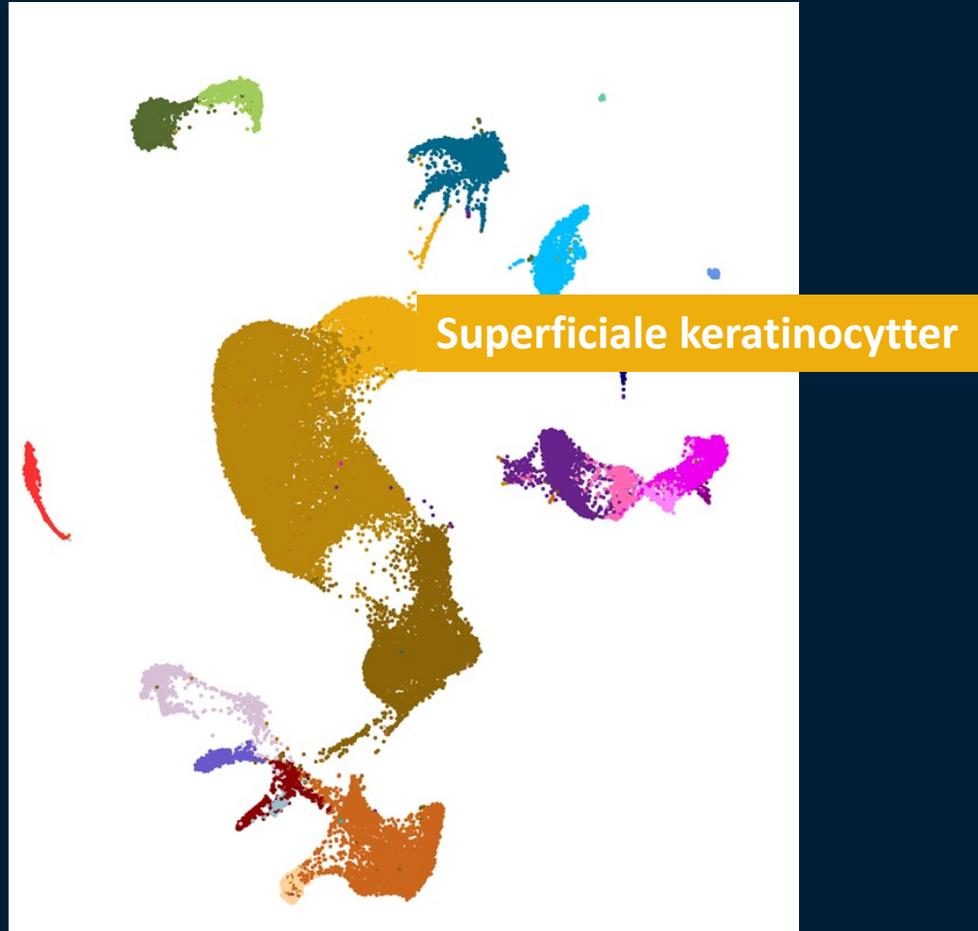
Intermediate keratinocytes

Intermediate keratinocytes

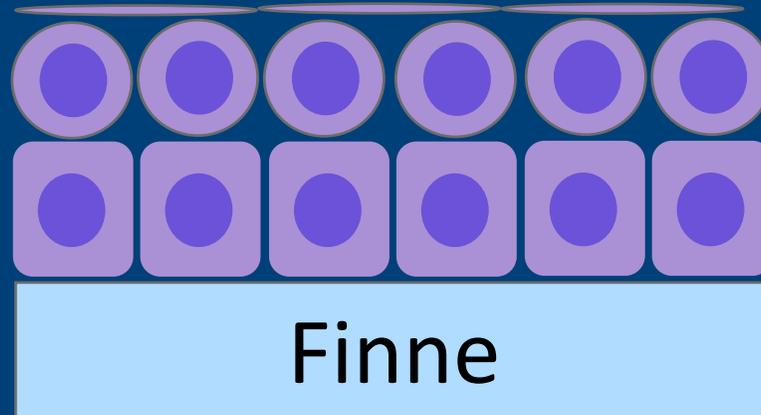


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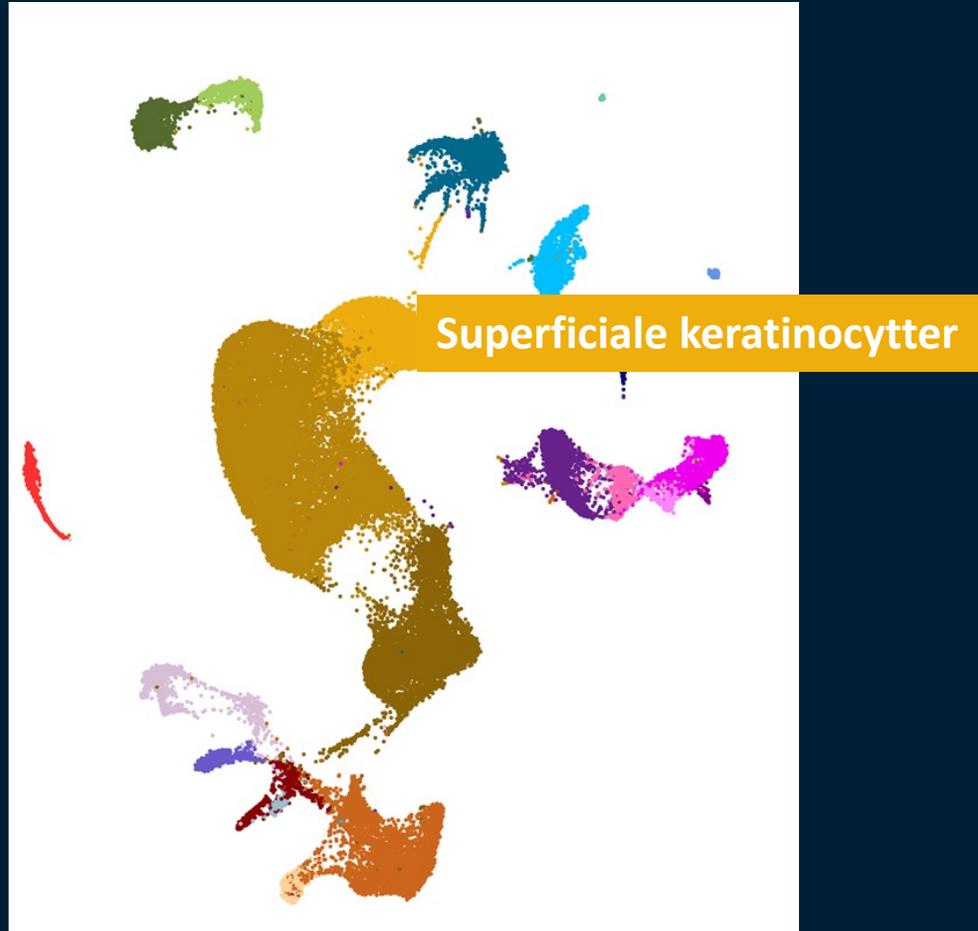
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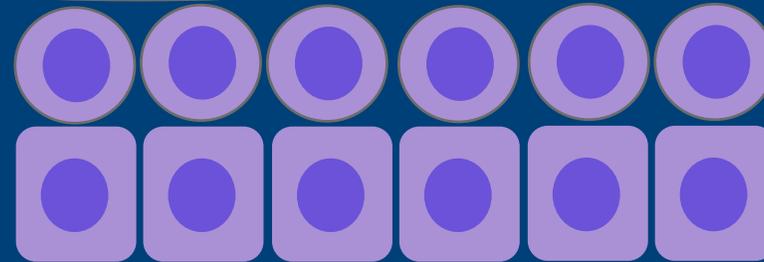
Superficial keratinocytes



Resultater

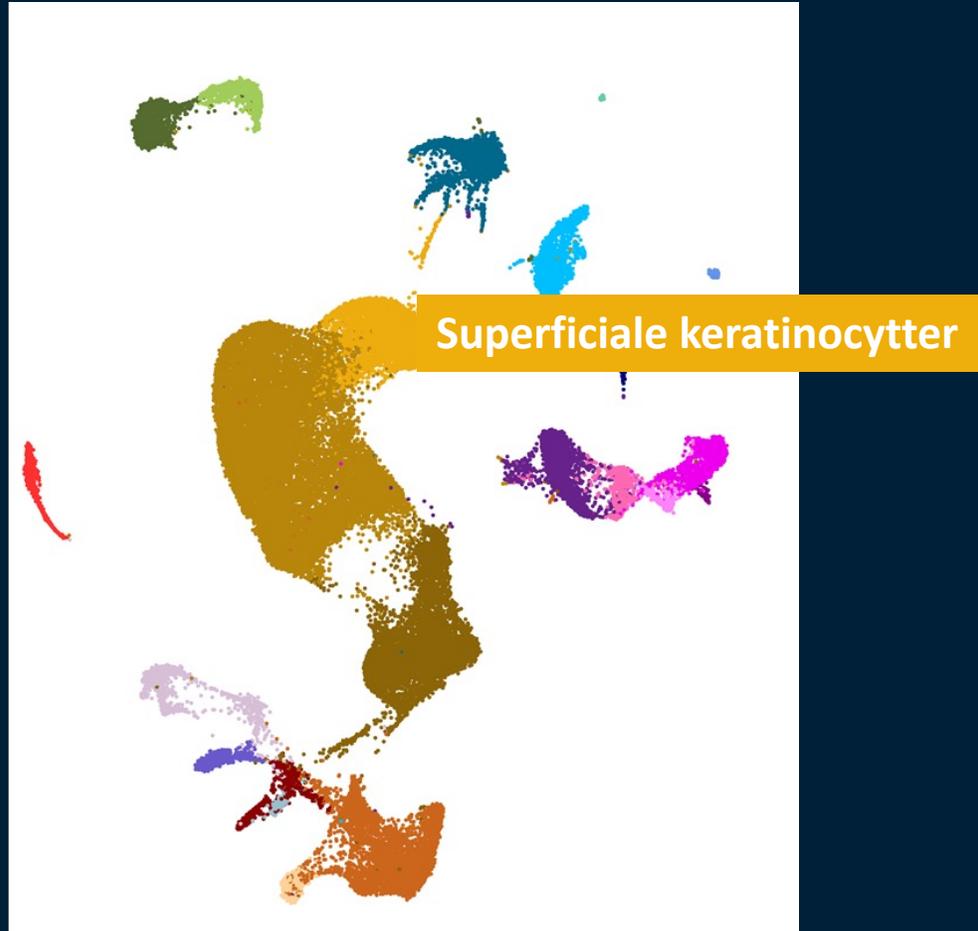


Lus

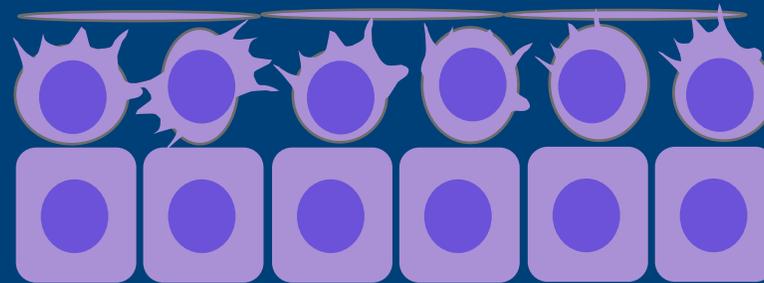


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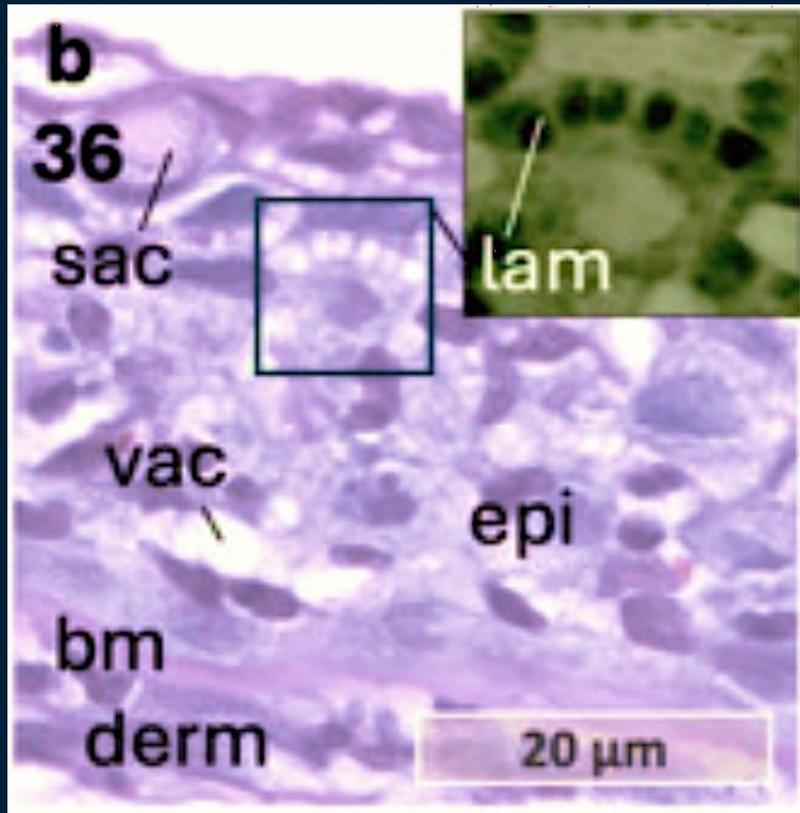


Lus

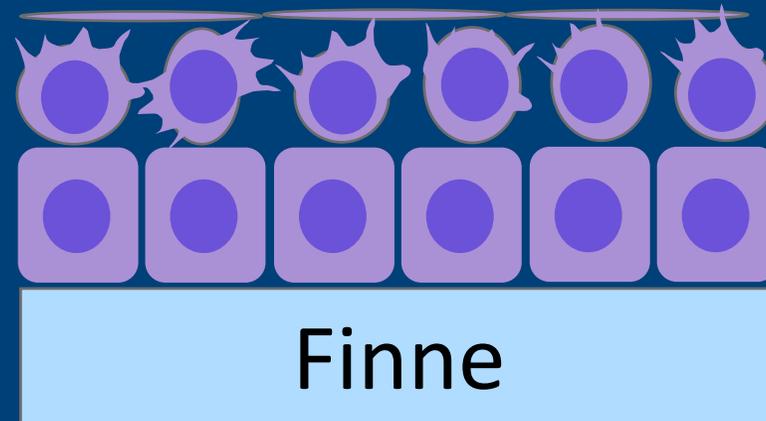


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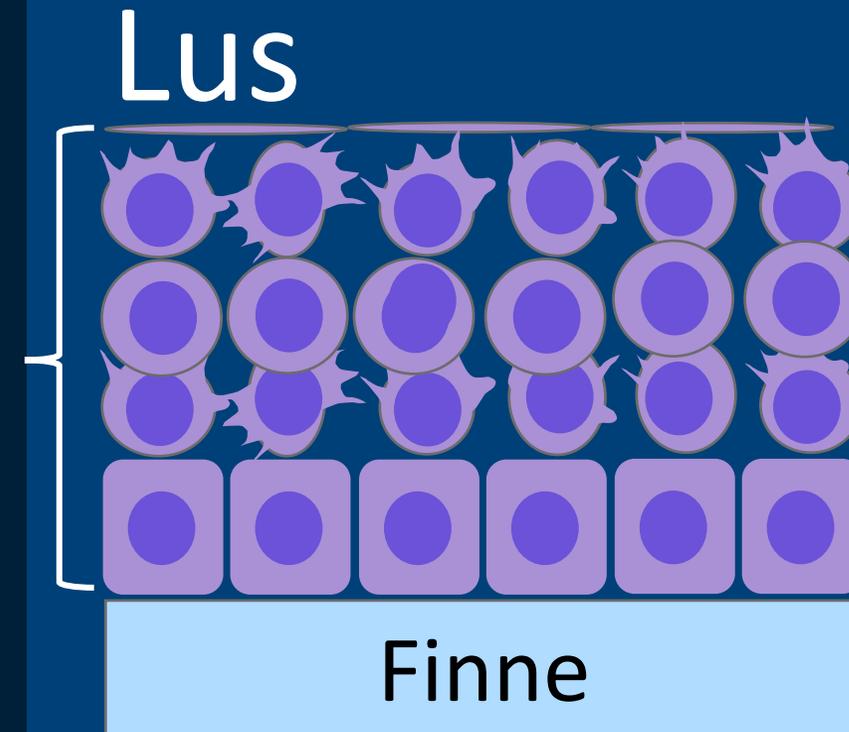


Lus

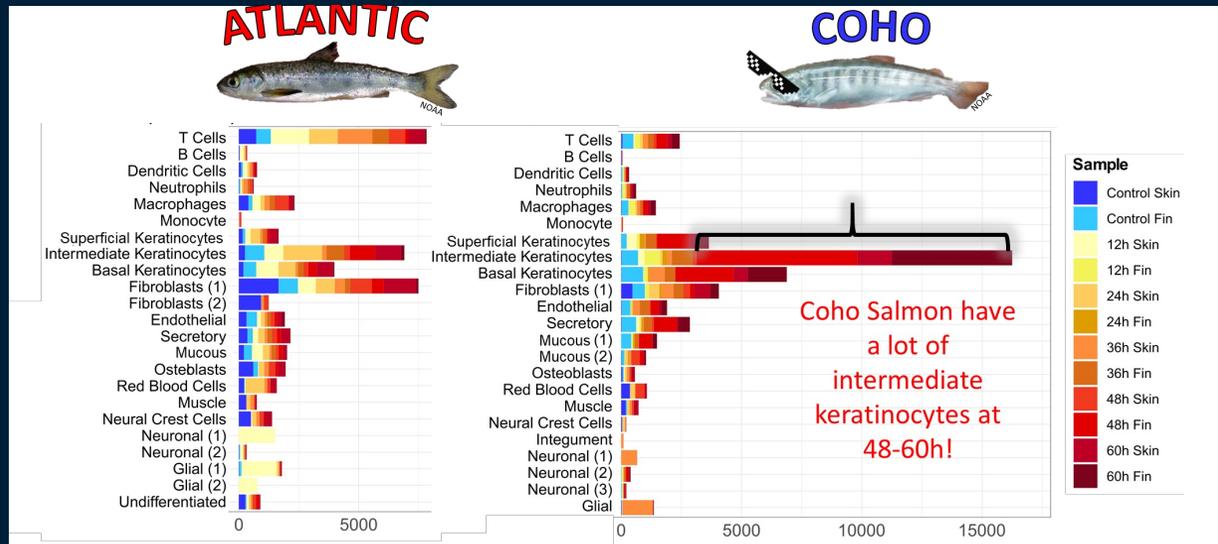


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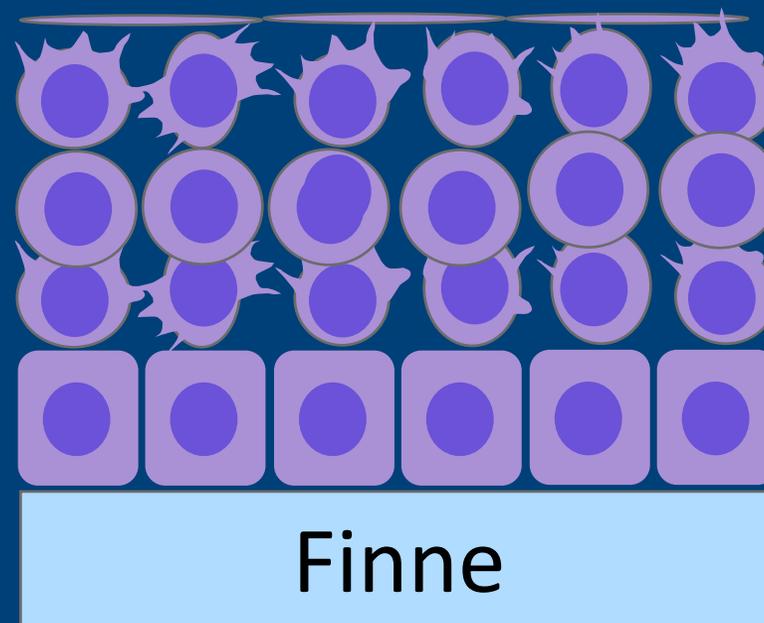
Økning i antall
intermediate
epitelceller 40 – 60
timer etter smitte



Resultater

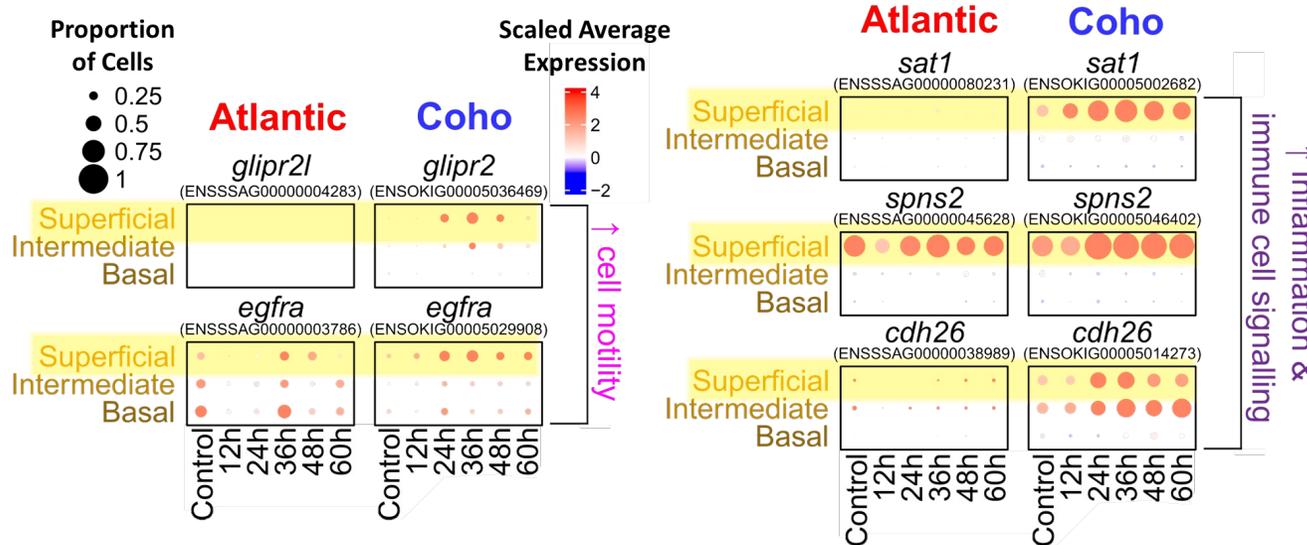


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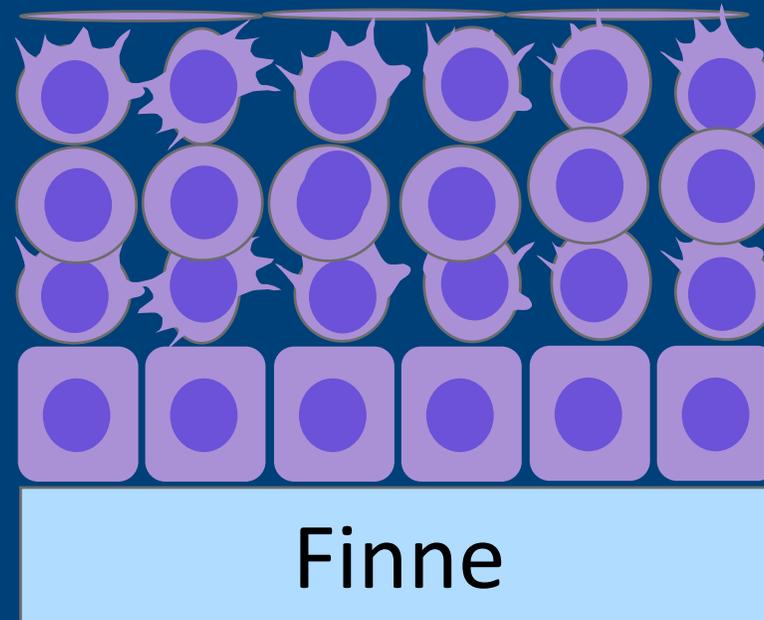


Resultater

Epitelcellene uttrykker forskjellige gener

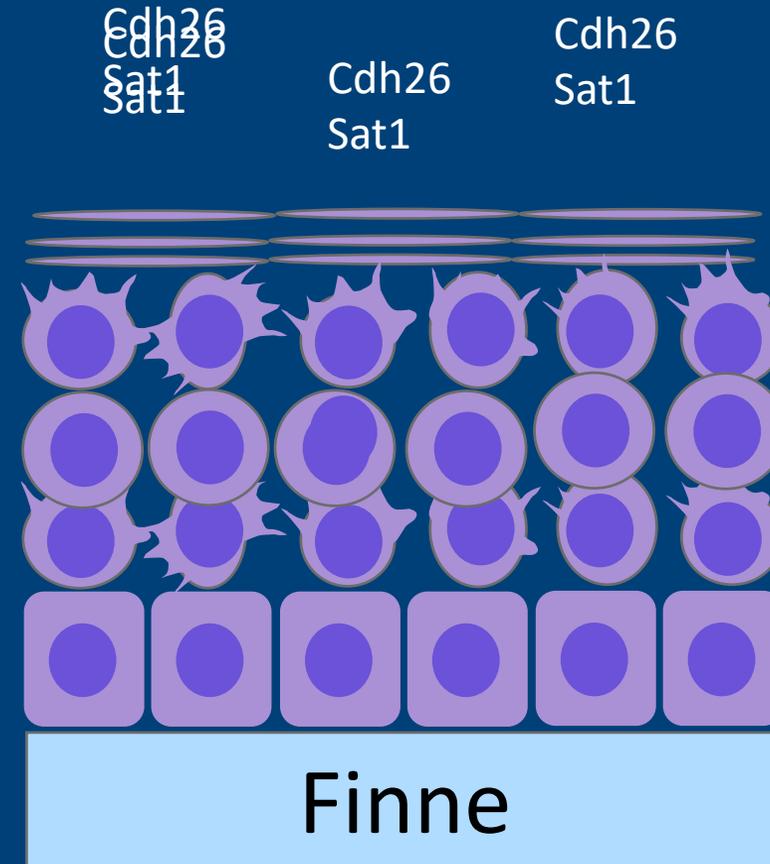
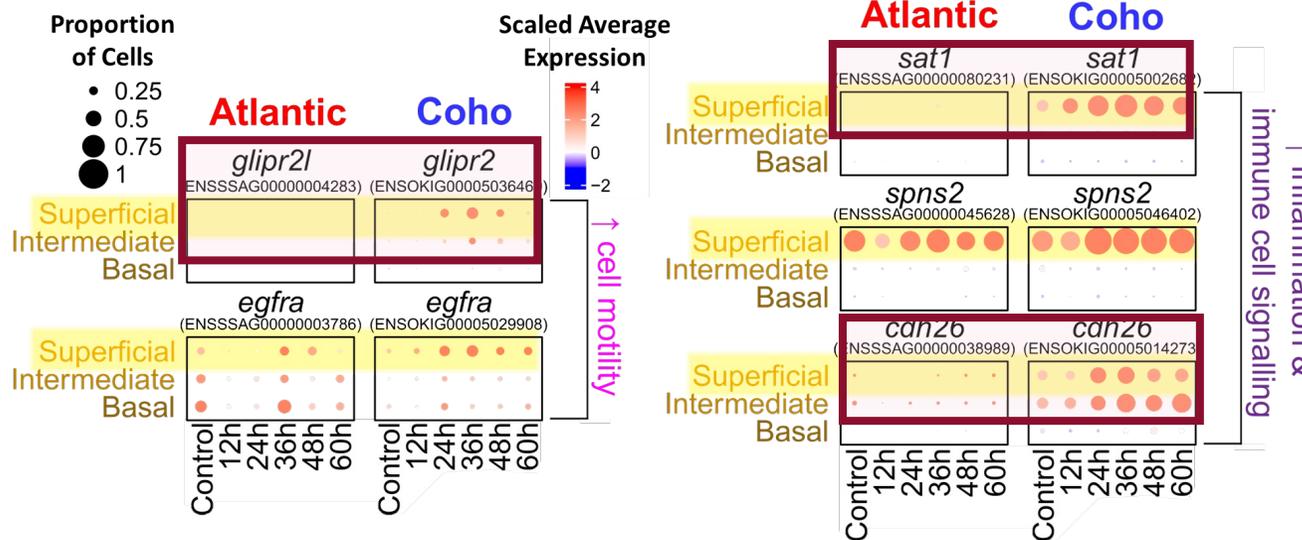


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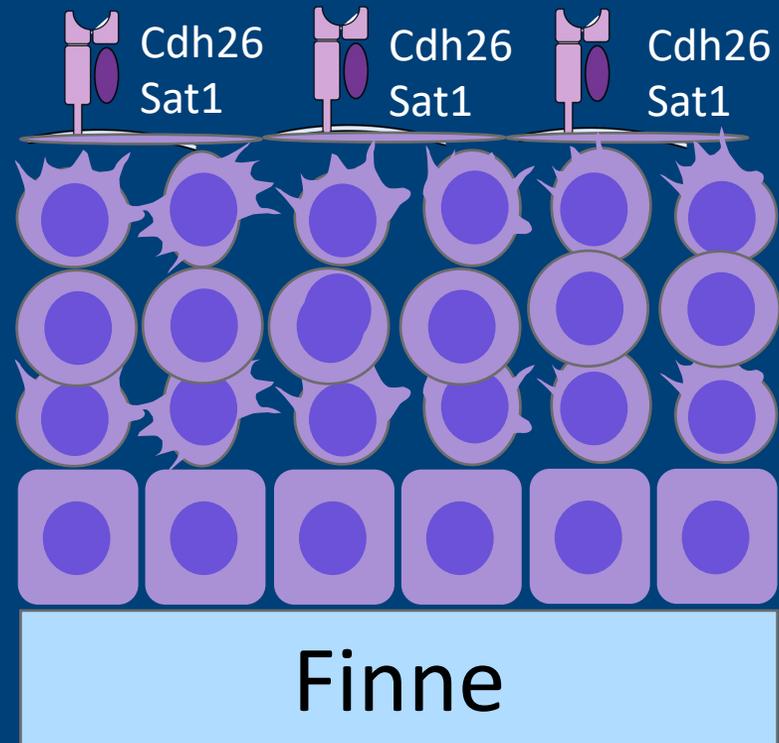
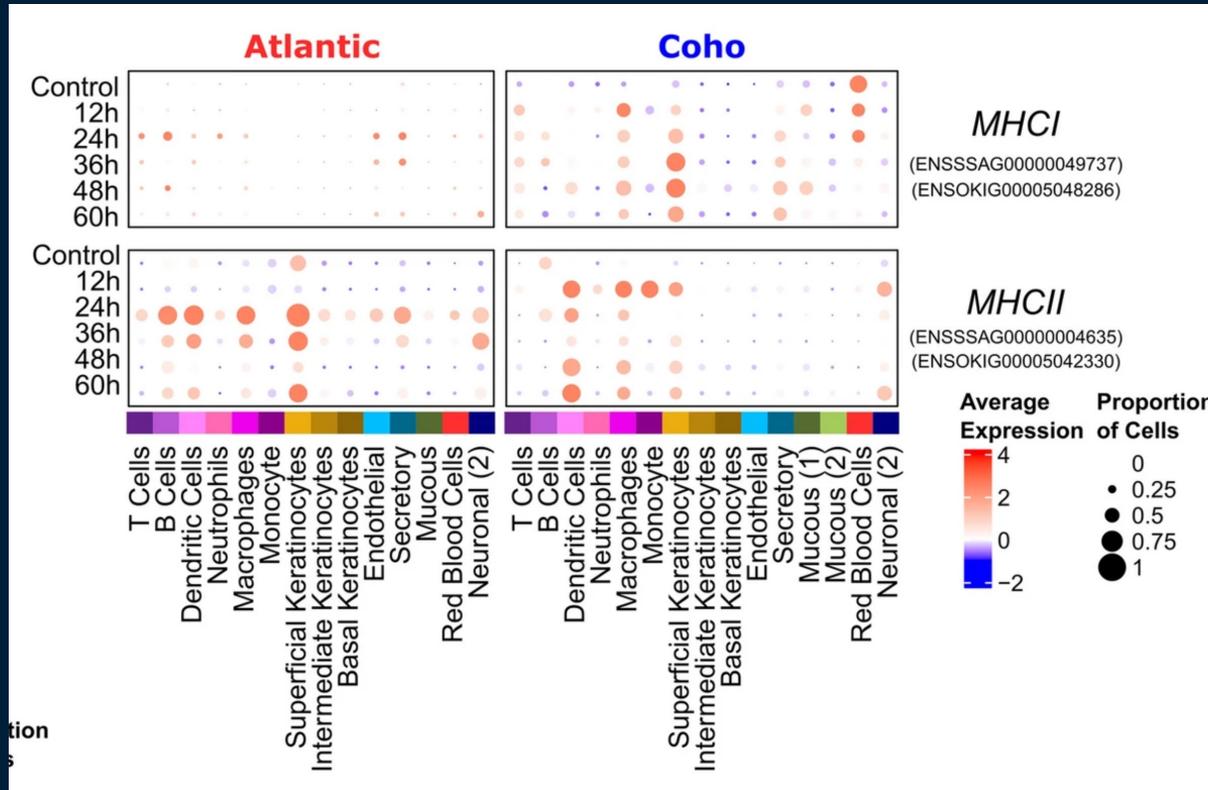
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Epitelcellene uttrykker forskjellige gener

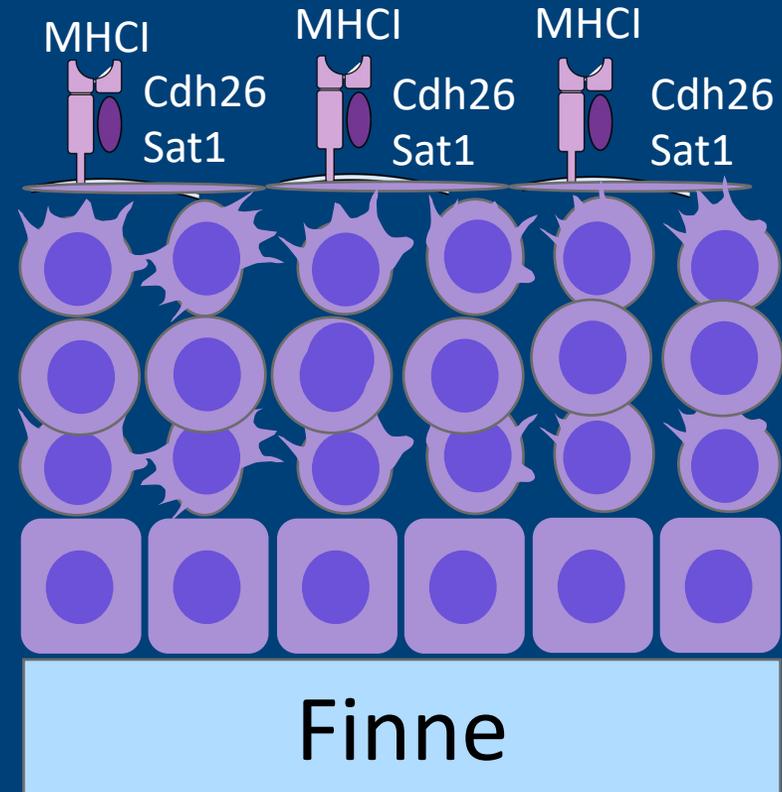
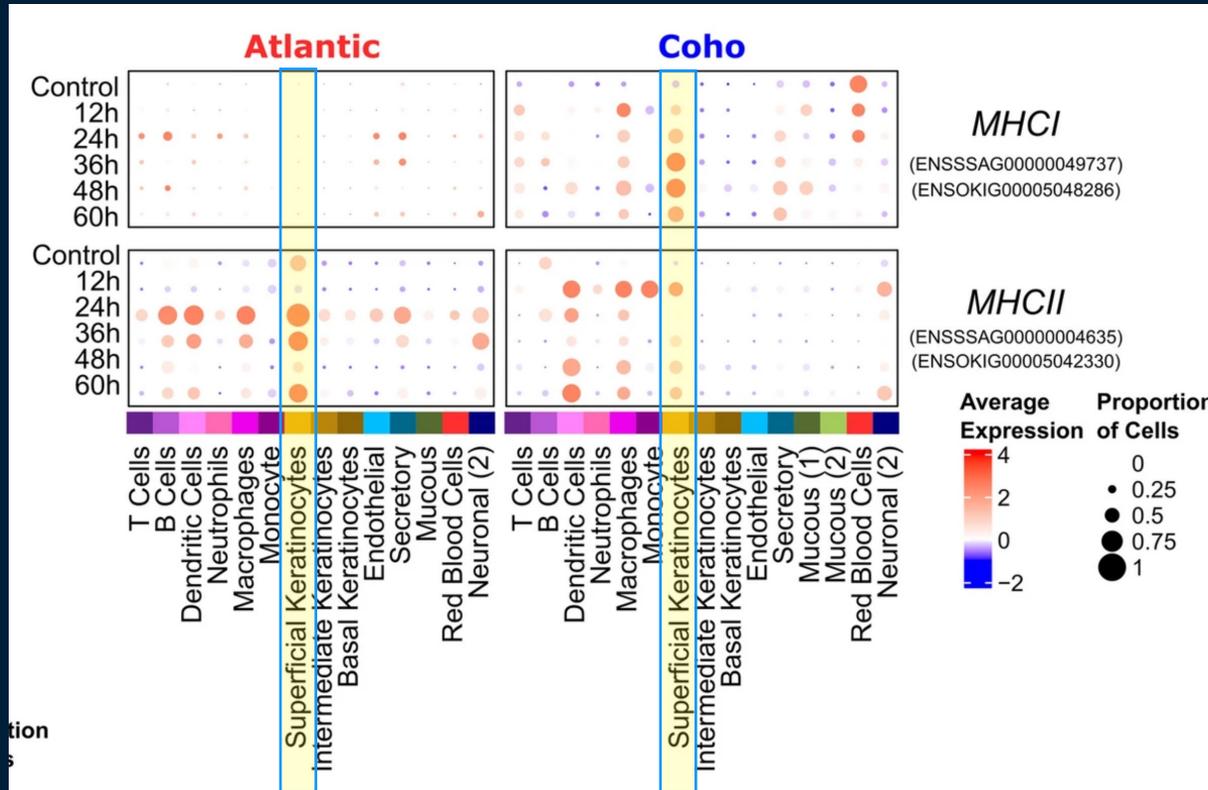


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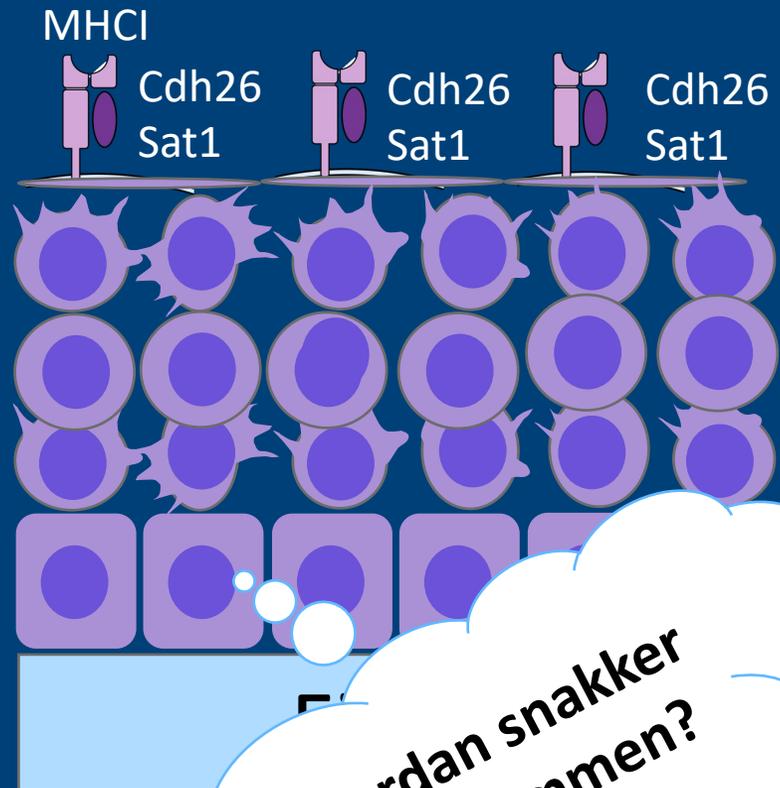
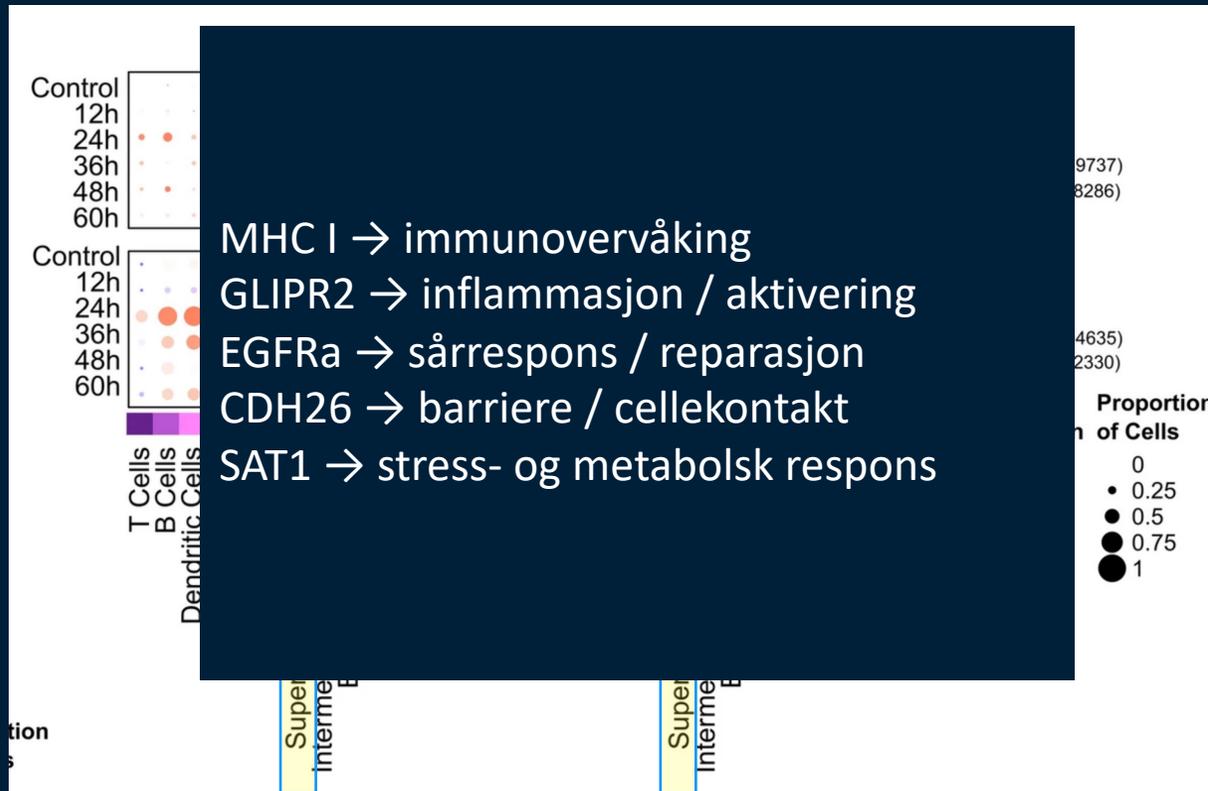
Epitelcellene uttrykker forskjellige gener



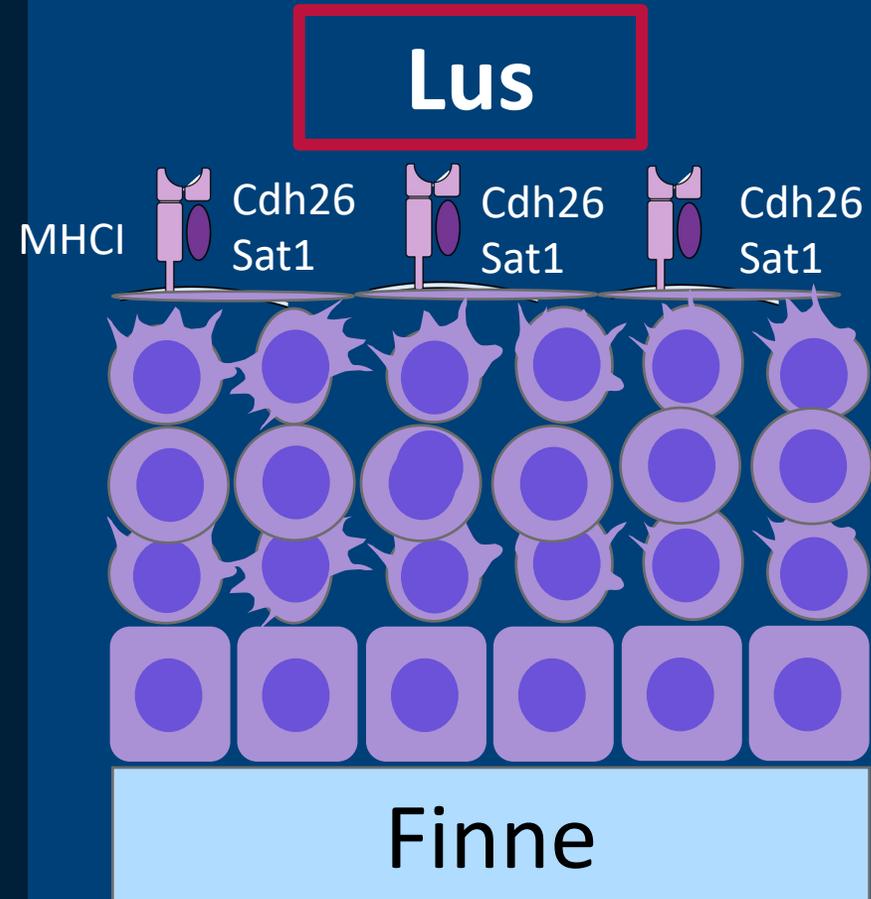
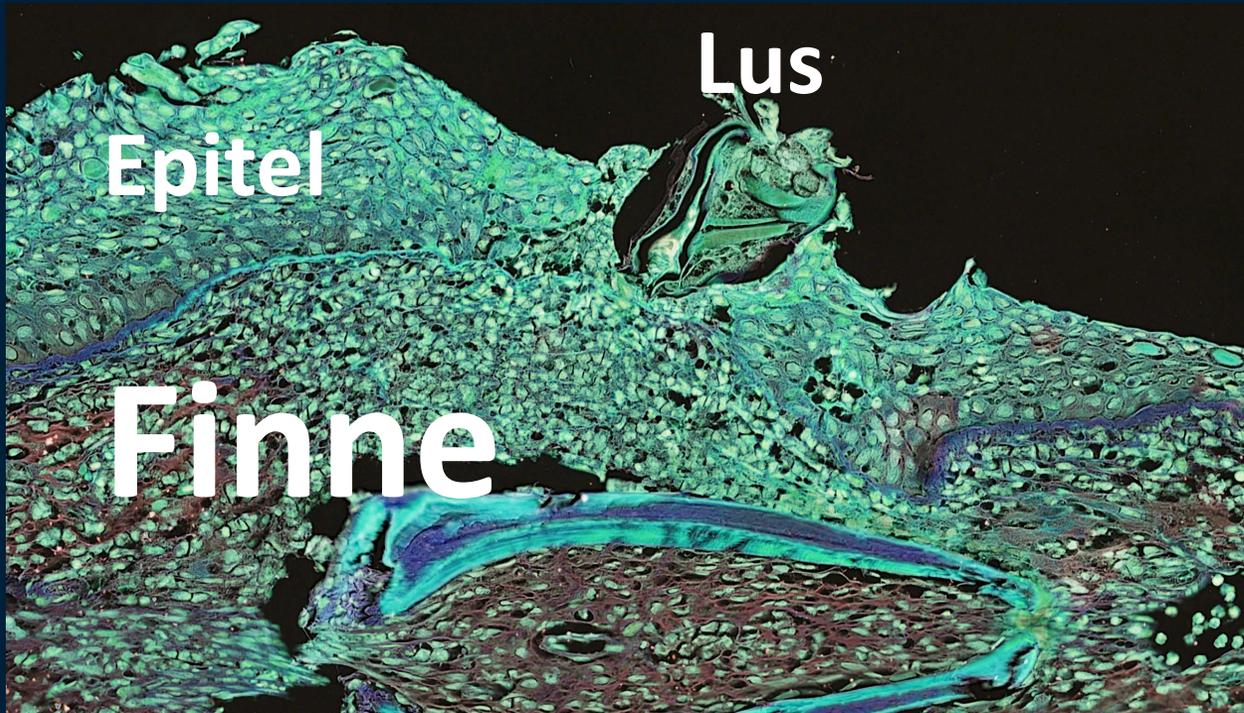
Resultater



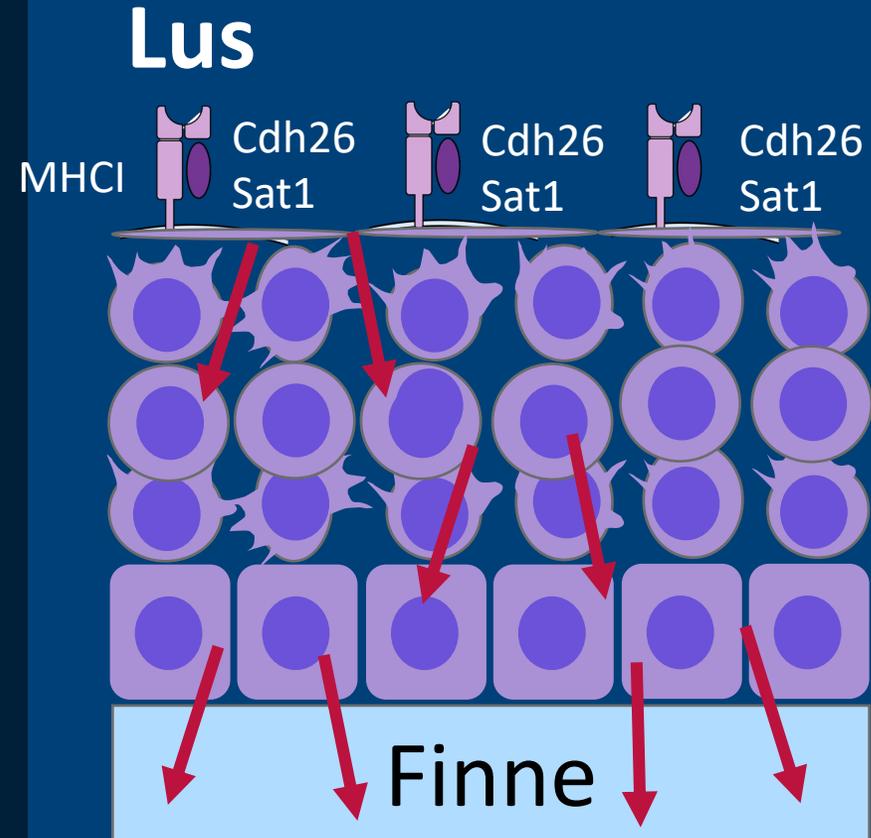
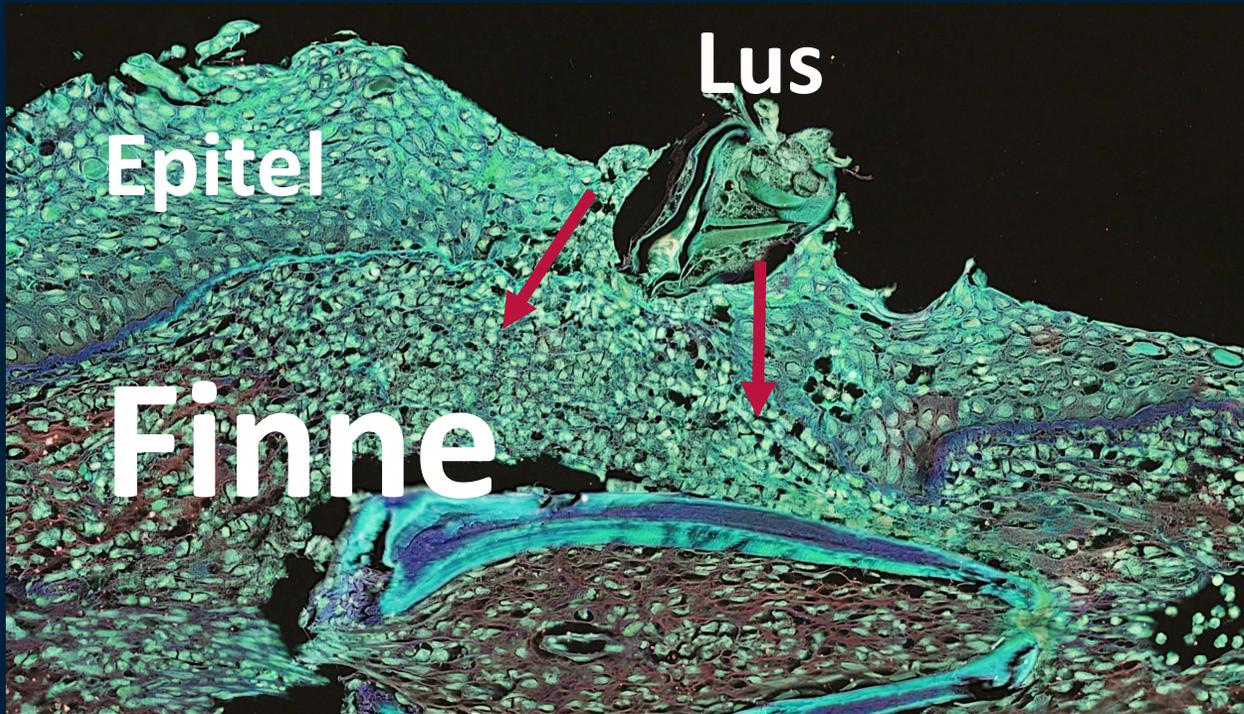
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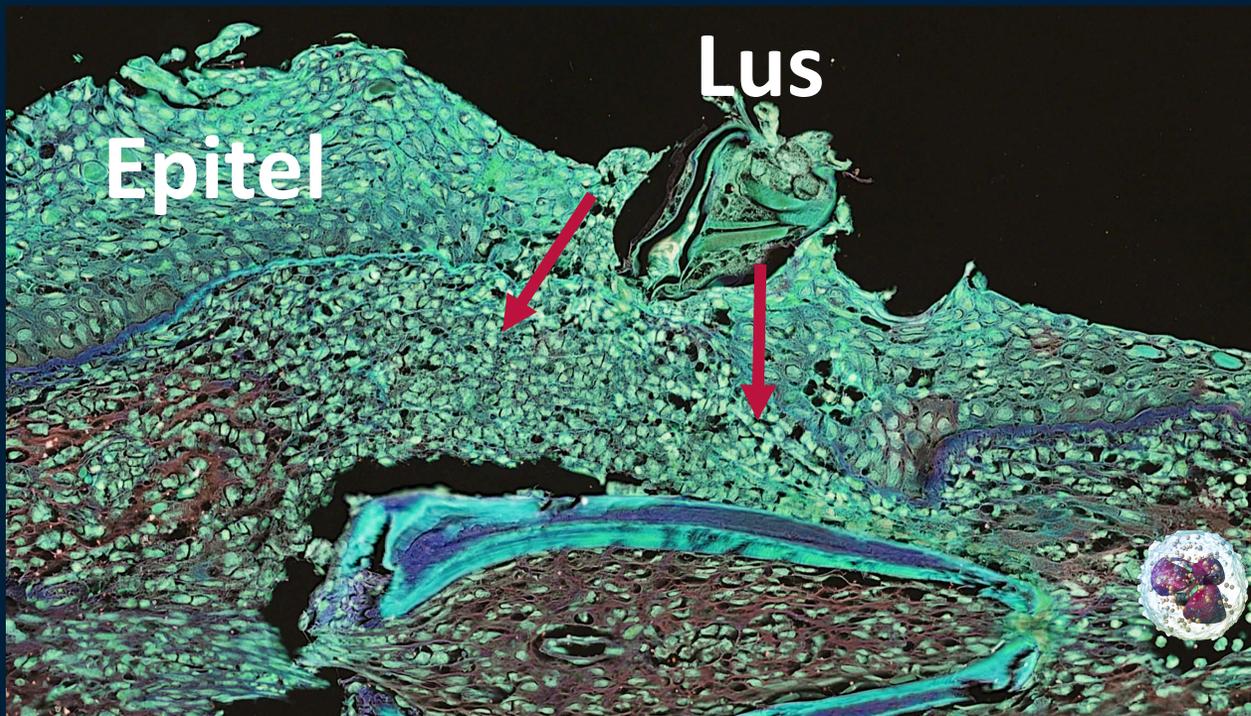
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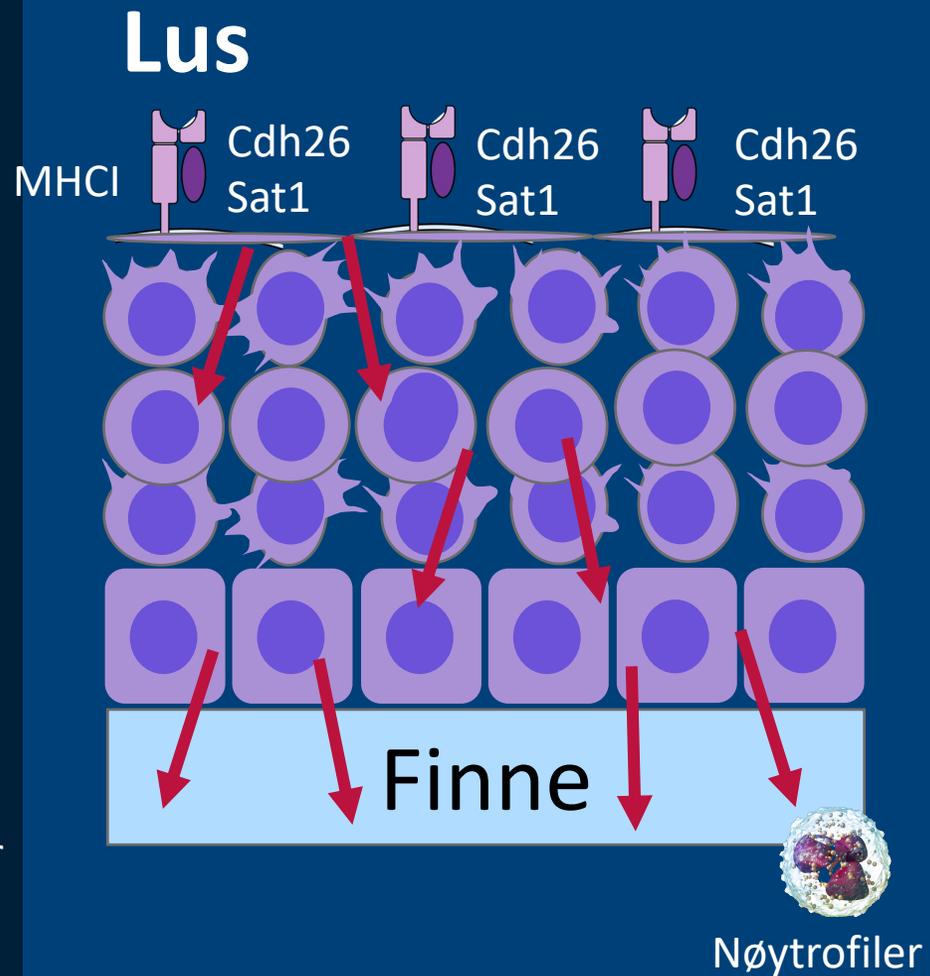
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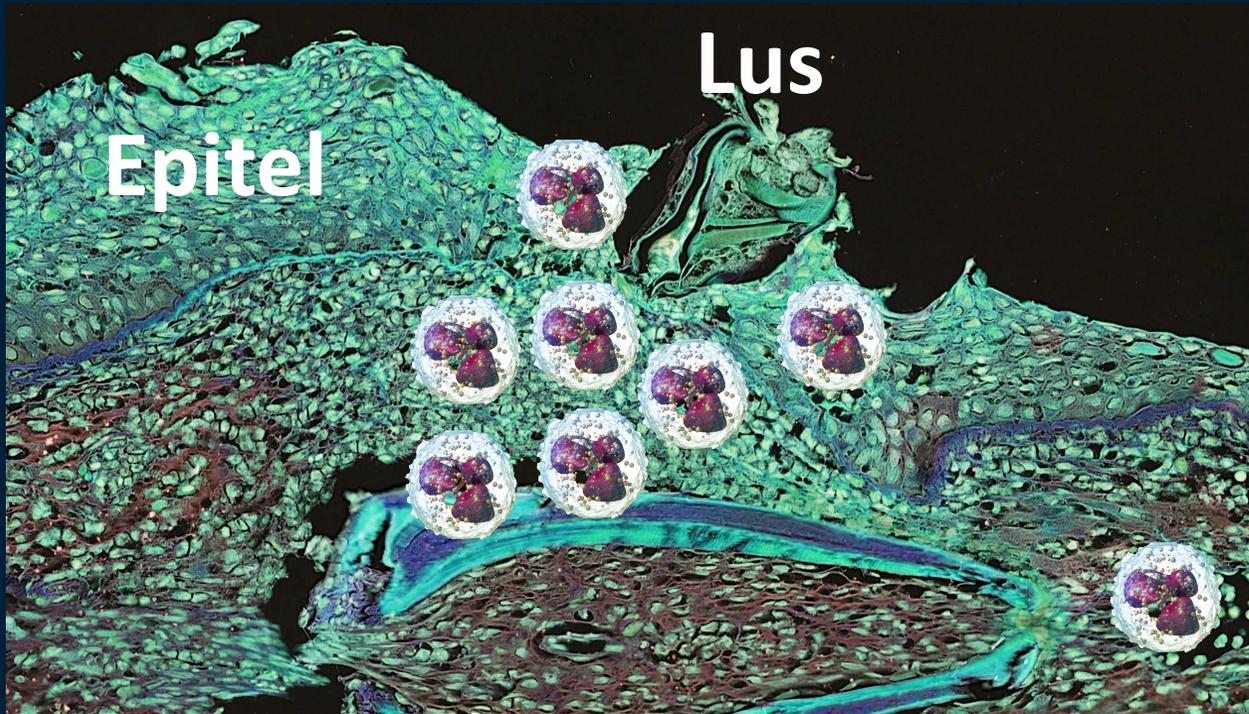
Resultater



Nøytrofiler

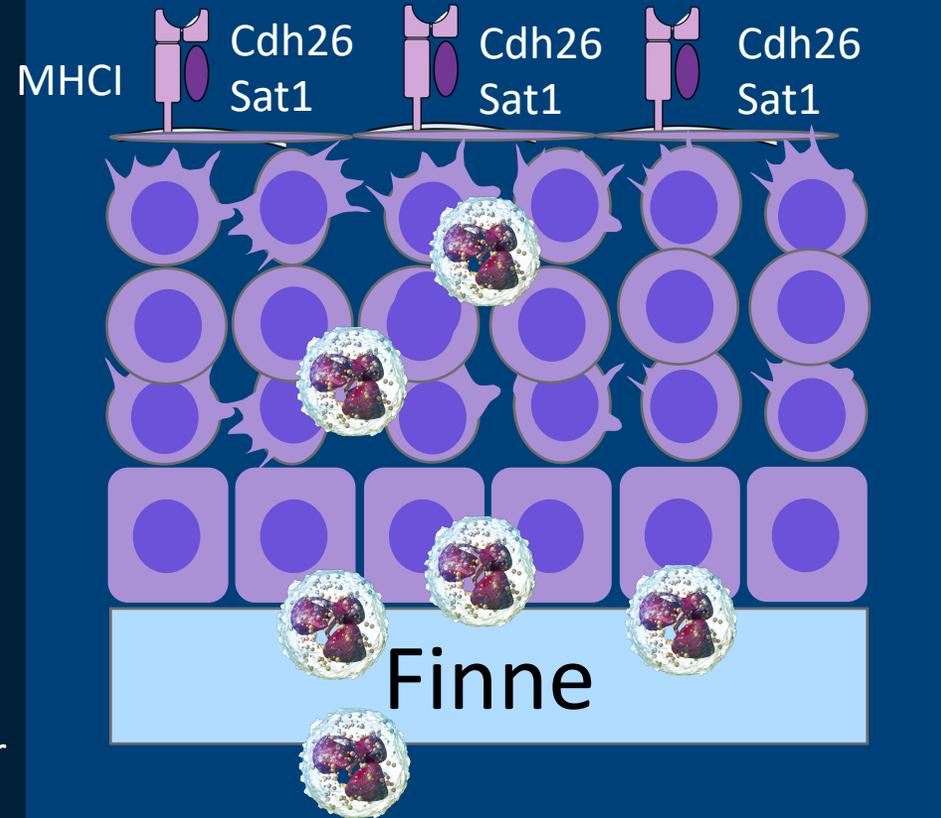


Resultater



Nøytrofiler

Lus



Nøytrofiler

Resultater

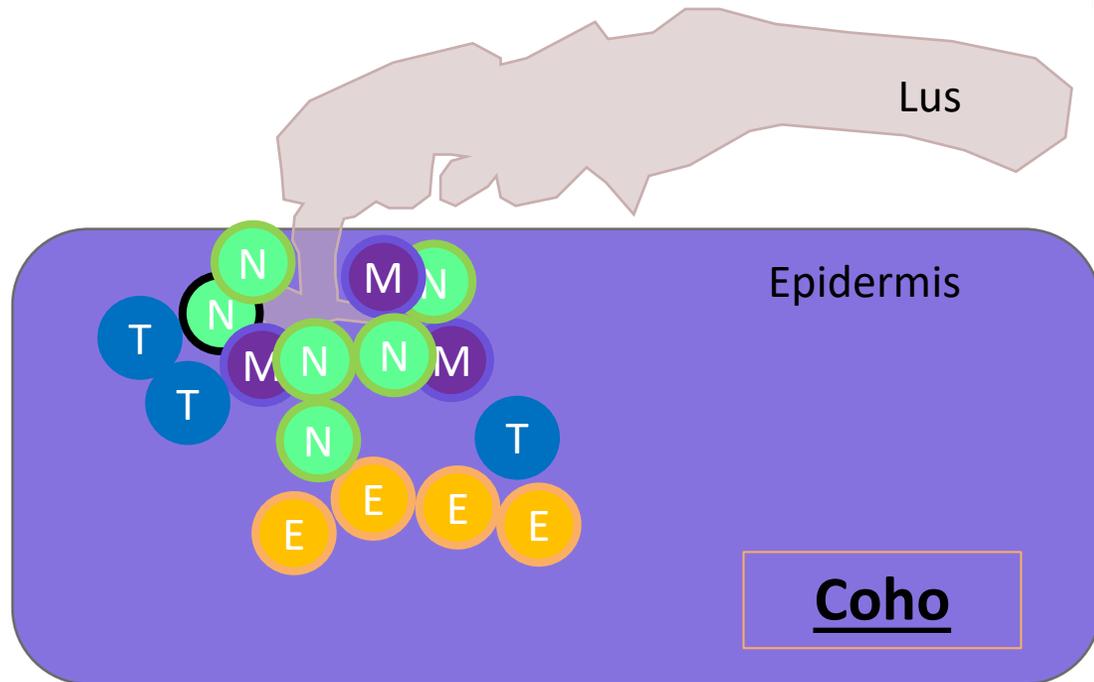


Oppsummering resultater

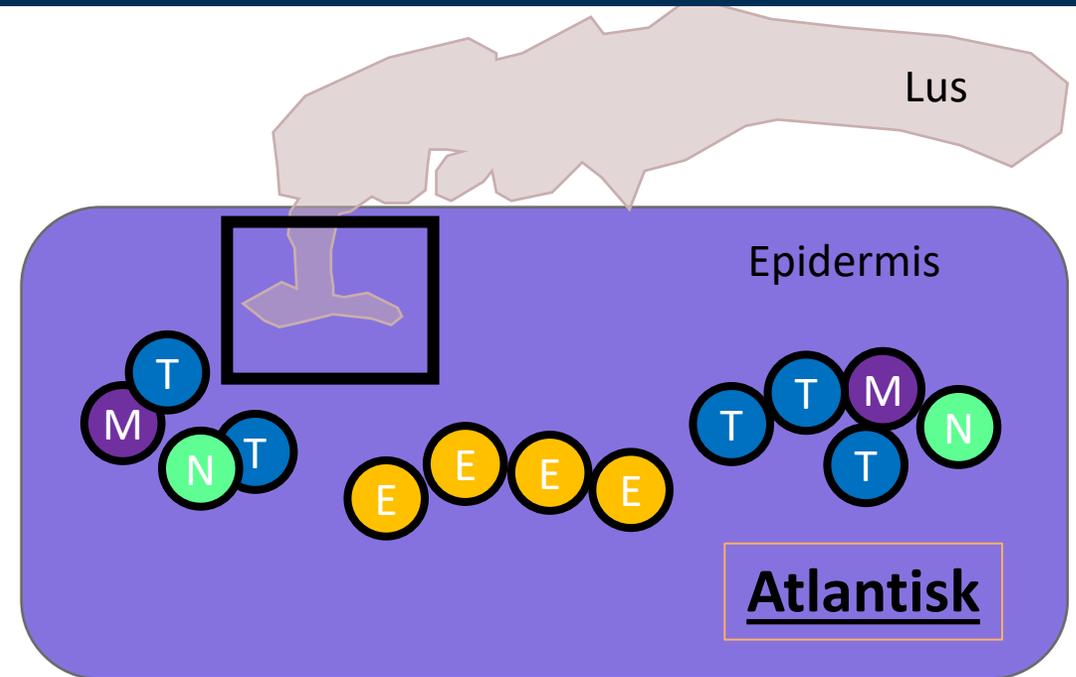
Oppsummering

Chum og Pink

Første strategi er "shake it off"
Immunceller...



Immunceller målrettet mot
festepunktet



Immunceller mangler spesifikk
retning, reagerer på erosjon av
epitel

BAKGRUNN

FORSØKSDESGIN

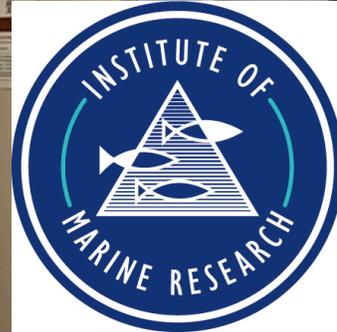
RESULTATER

GENREDIGERING

VEIEN VIDERE

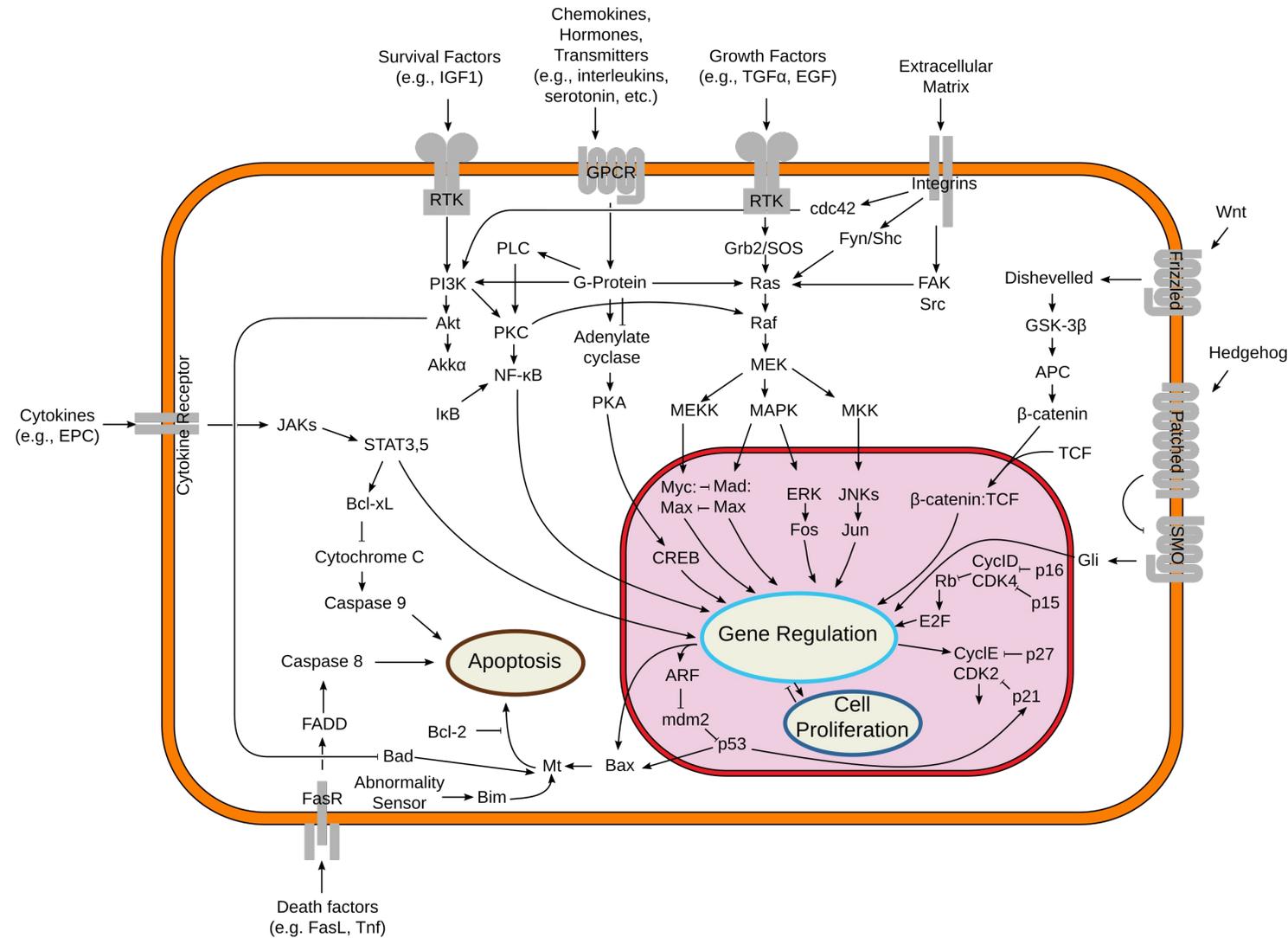


 Nofima



Genredigering

Hvilken gener skal vi velge?



Genredigering

Yamanaka-faktorer er en gruppe på fire proteiner **Oct4, Sox2, Klf4 og c-Myc**,

som kan omprogrammere **voksne, spesialiserte celler** tilbake til **ung tilstand**

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Article | Published: 07 March 2022

In vivo partial reprogramming alters age-associated molecular changes during physiological aging in mice

[Kristen C. Browder](#), [Pradeep Reddy](#), [Mako Yamamoto](#), [Amin Haghani](#), [Isabel Guillen Guillen](#), [Sanjeeb Sahu](#), [Chao Wang](#), [Yosu Luque](#), [Javier Prieto](#), [Lei Shi](#), [Kensaku Shojima](#), [Tomoaki Hishida](#), [Zijuan Lai](#), [Qingling Li](#), [Feroza K. Choudhury](#), [Weng R. Wong](#), [Yuxin Liang](#), [Dewakar Sangaraju](#), [Wendy Sandoval](#), [Concepcion Rodriguez Esteban](#), [Estrella Nuñez Delicado](#), [Pedro Guillen Garcia](#), [Michal Pawlak](#), [Jason A. Vander Heiden](#), ... [Juan Carlos Izpisua Belmonte](#)  [+ Show authors](#)

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Abstract

Partial reprogramming by expression of reprogramming factors (Oct4, Sox2, Klf4 and c-Myc)



16 forskjellige typer genredigert laks er under uttesting

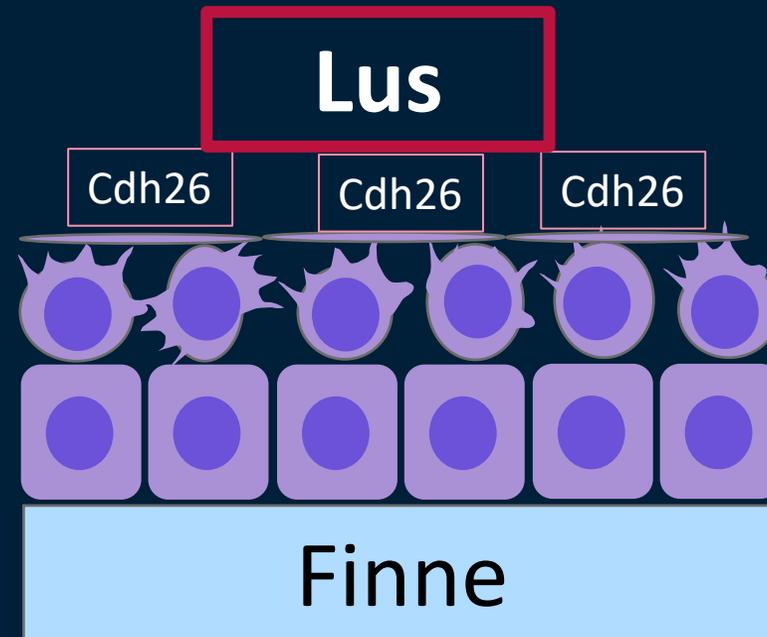
Genredigering

Cdh26 → immune modulering, tiltrekke seg hvite blodlegmer (neutrophiler)

Pbrm_1 → Proliferering

Mrc1a → Reseptor for gjenkjenning av parasitter og aktivering av medfødt immunrespons

Socs3 → Regulerer inflammasjon og immunrespons



Genredigering

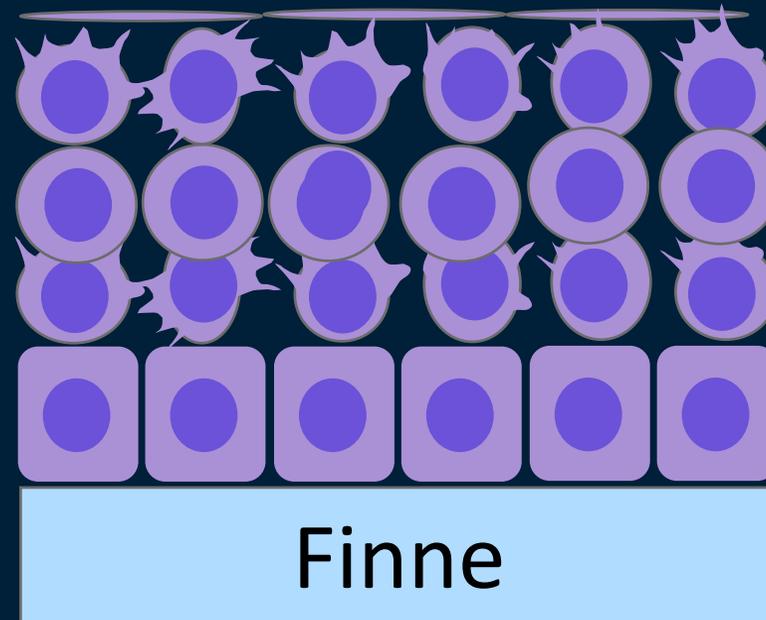
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Lus



Genredigering

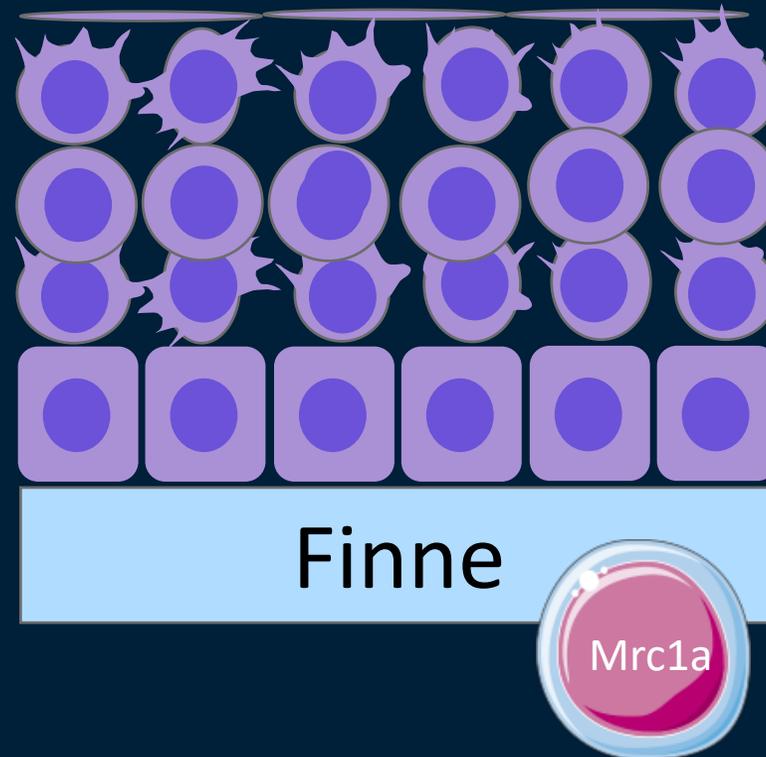
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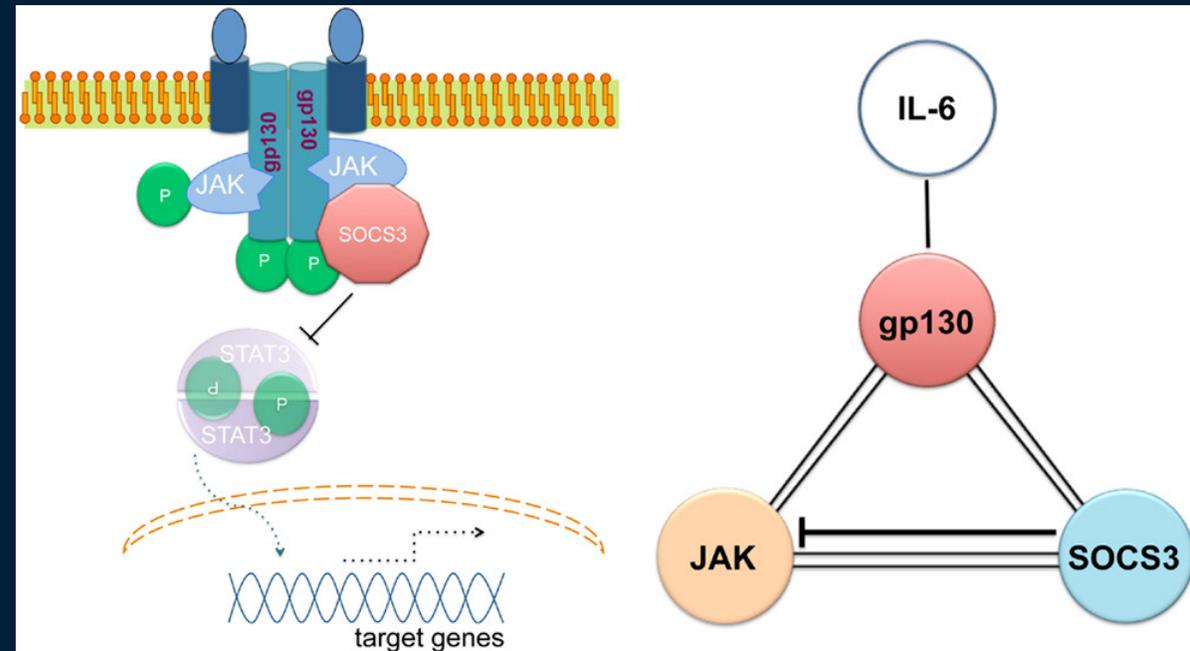
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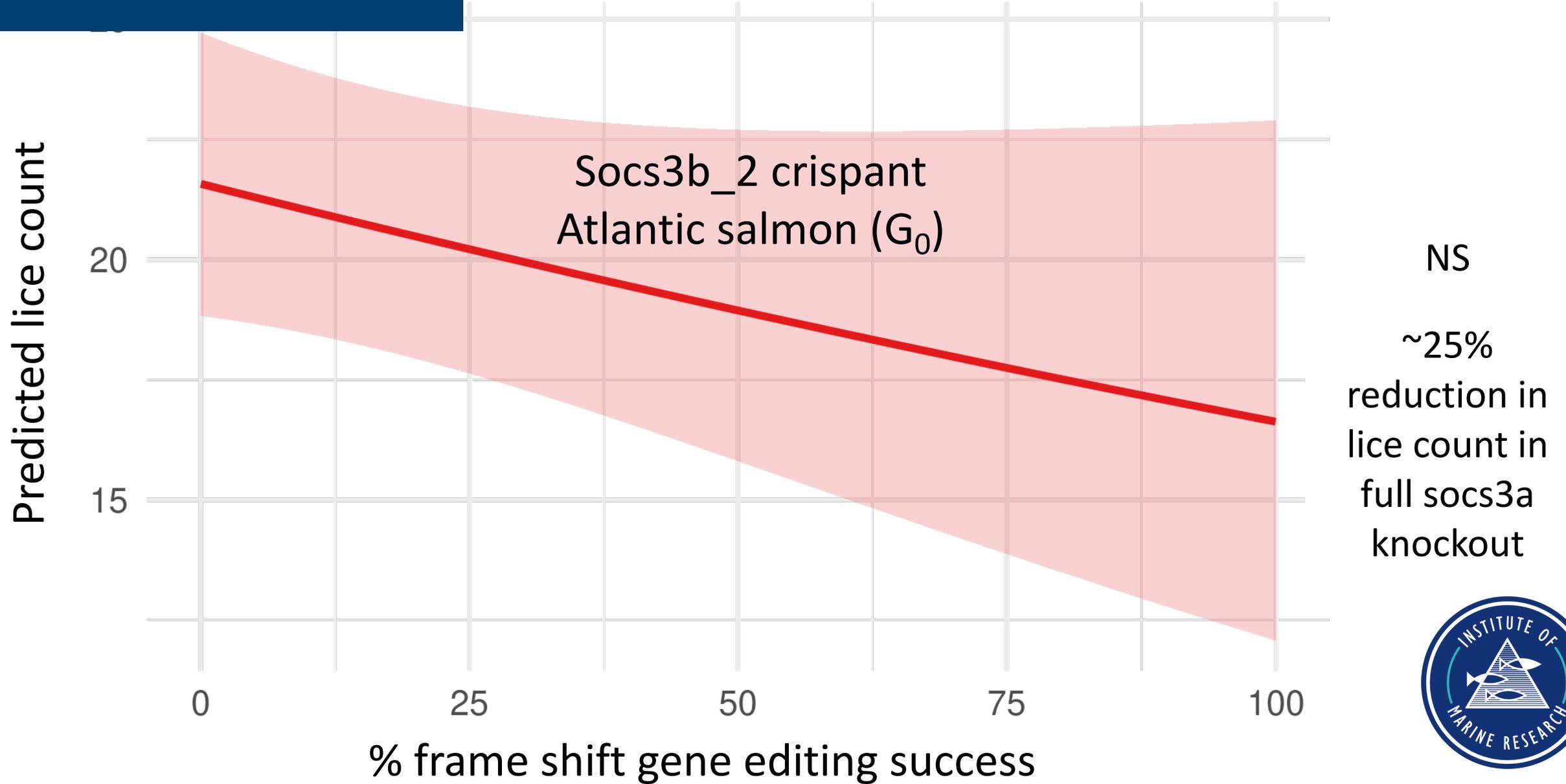
Socs3 → Regulerer inflammasjon og immunrespons

SOCS3 hovedregulator



Carrow og Rottenberg 2014

Genredigering



BAKGRUNN

FORSØKSDESGIN

RESULTATER

GENREDIGERING

VEIEN VIDERE



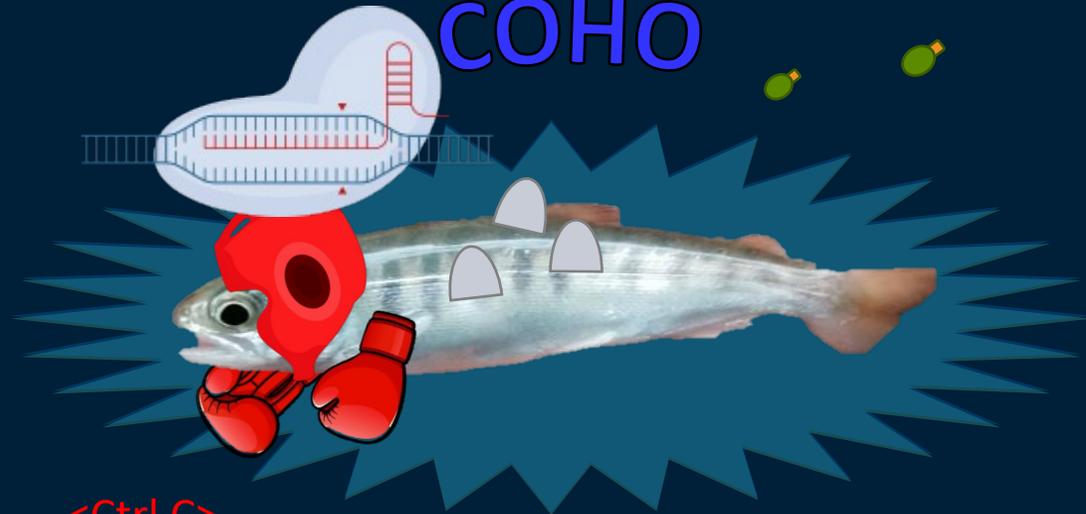
Veien videre

ATLANTIC



<Ctrl V>

COHO



<Ctrl C>

Slide by: Sarah Salisbury

Resultatene identifiserer flere signalveier og gener som bidrar til å gjøre coho laks resistent mot lus.

Genredigering eller selektiv avl kan brukes for å forbedre Atlantisk laks sin medfødte immunitet mot lakselus

Slide av: Sarah Salisbury



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