



RENCONTRE SCIENTIFIQUE

Perturbateurs endocriniens : les nouveaux défis de la recherche

Jeudi 13 juin 2024

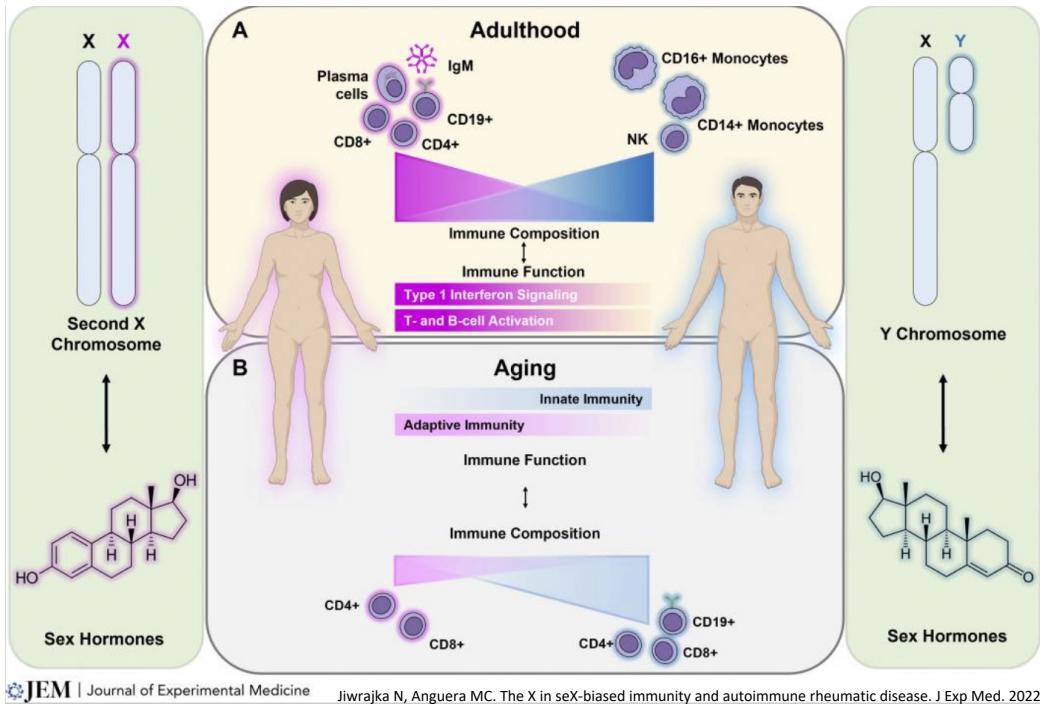
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Impact des perturbateurs endocriniens œstrogéniques sur le système immunitaire d'un poisson téléostéen, le bar européen

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Dimorphisme sexuel dans la réponse immunitaire :

- Estrogène endogène: 17 β -estradiol (E2)
 - Hormone femelle majeure
 - Importante pour reproduction sexuelle
- Système immunitaire féminin plus performant
- Maladies auto-immunes: majoritairement chez les femme (>80%)

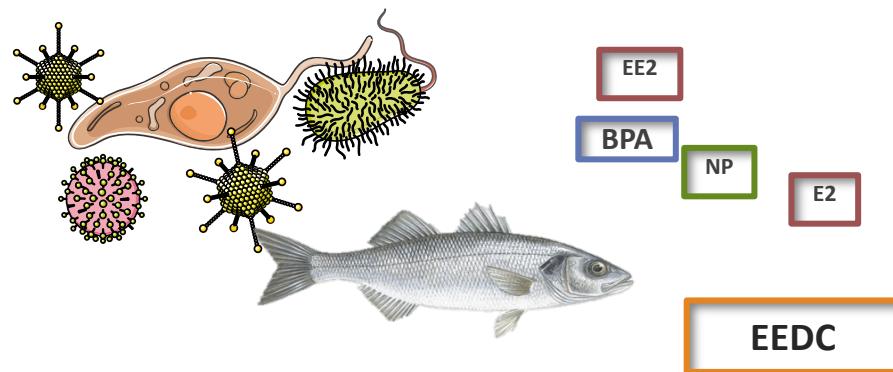


JEM | Journal of Experimental Medicine

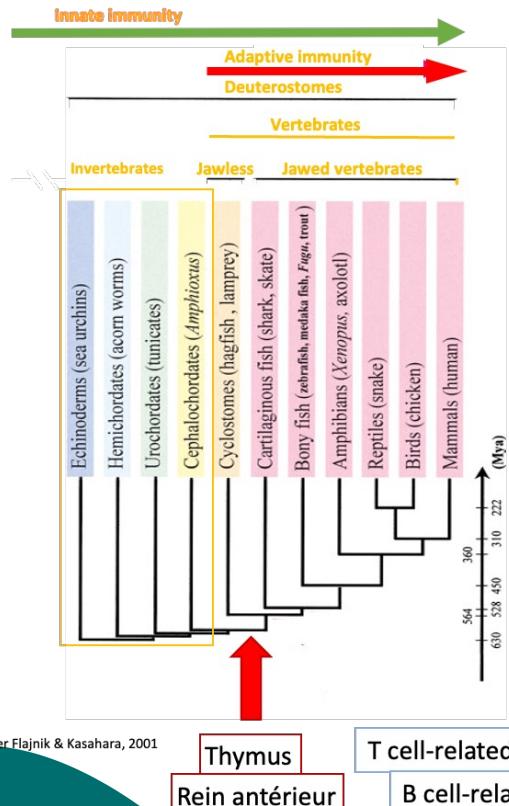
Jiwrajka N, Anguera MC. The X in seX-biased immunity and autoimmune rheumatic disease. J Exp Med. 2022

Relation entre la pollution anthropique (PE) et montée des maladies émergentes

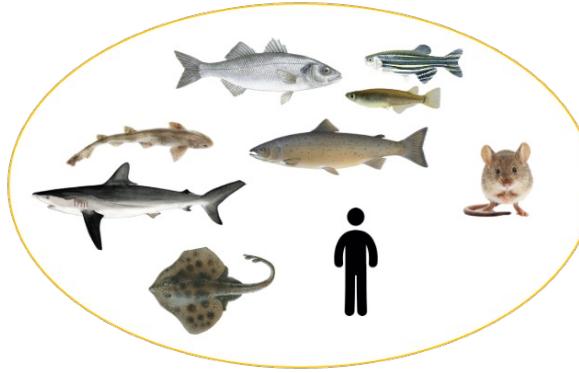
- Rôle de la modulation de l'immunité des organismes induite par l'environnement ?
- Quels traits immunitaires affectés par les divers facteurs de stress anthropiques ?



Immunité chez les vertébrés



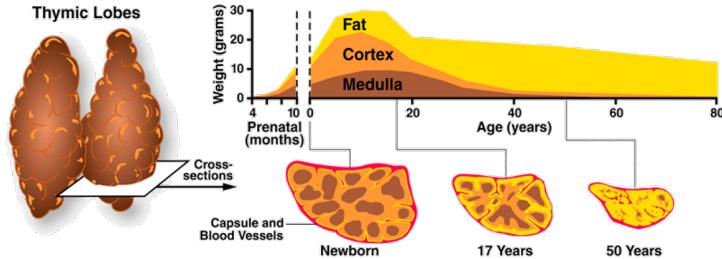
➤ Vertebrés ont développé un **système immunitaire adaptatif**



Importance du
thymus pour l'
immunité **adaptative**

Influence des estrogènes sur le SI : plasticité thymique

- Organe très **plastique**
- Involution thymique: tissu adipeux/ perte des espaces épithéliaux / réduction de la prolifération des T et de leur nombre: Emigration des cellules T en périphérie moins importante
- Variation associée aux fonctions **reproductives**
- Sous contrôle des hormones stéroïdes sexuelles

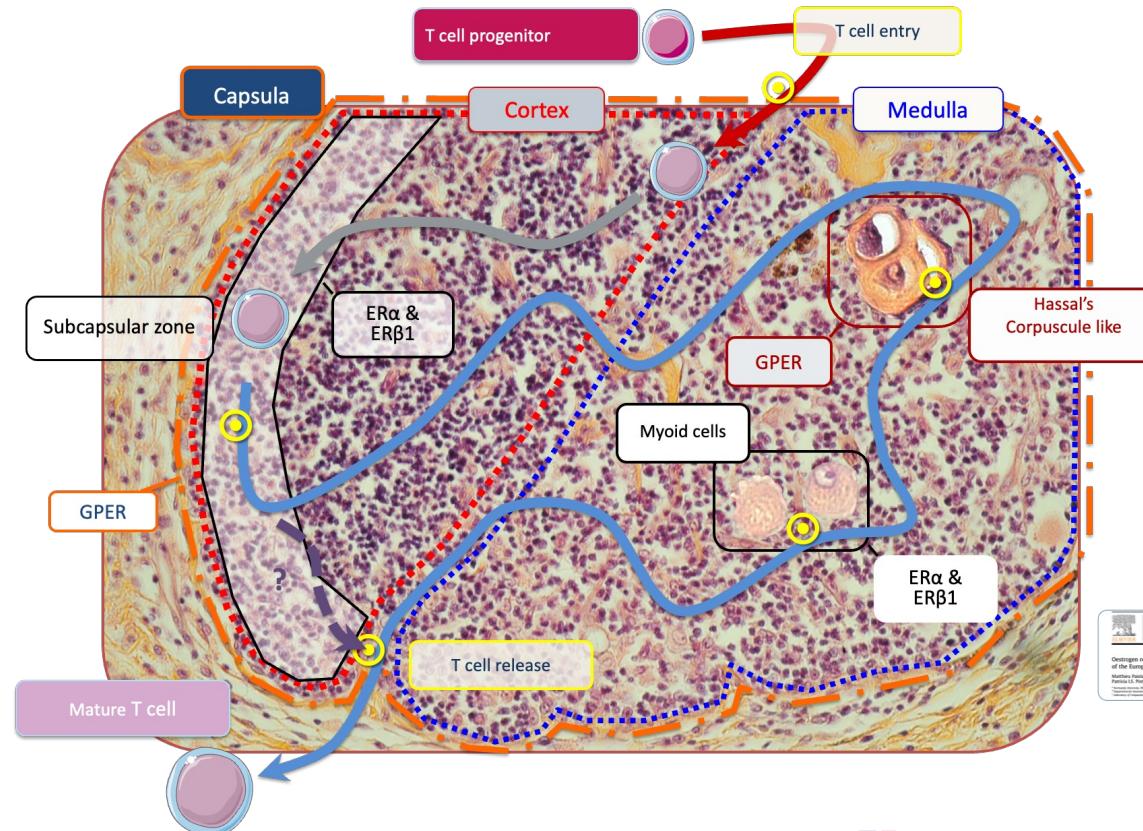


L'involution thymique réversible au cours de la grossesse met en lumière l'importance des œstrogènes dans la modulation de la plasticité thymique.

Distribution des différents récepteurs aux estrogènes dans le microenvironnement thymique

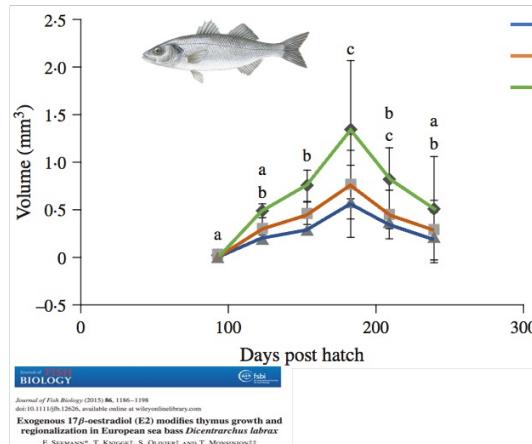


L'organisation morpho-fonctionnel du thymus est conservée du point de vue de l'évolution

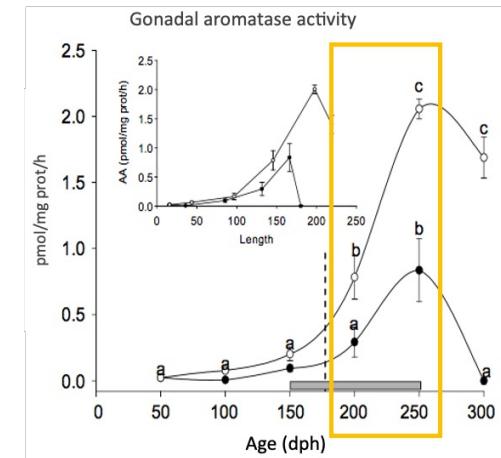
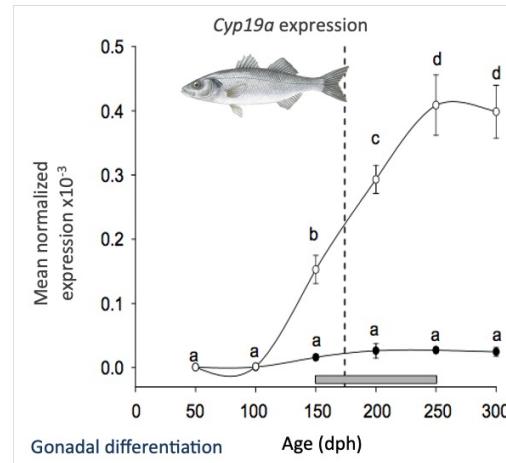


Open Access available at <https://doi.org/10.1007/s00162-024-02905-0>
Developmental and Comparative Immunology
Volume 100, June 2024, Article 103006
Original research article
Estrogen receptor distribution related to functional thymus anatomy of the European sea bass, *Dicentrarchus labrax*
Patricia S. Pinto • George Skopidakis • Spyros Mousoulis
Received: 10 January 2024 / Accepted: 10 May 2024 / Published online: 10 June 2024
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Mécanisme de régulation hormonale du SI identique et même plasticité thymique ?



Estradiol promotes Atrophie thymique

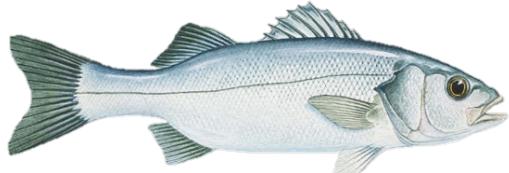
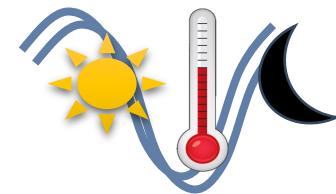
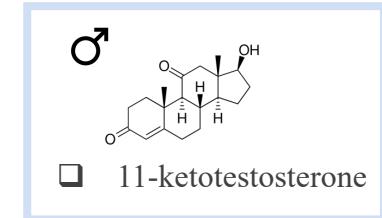
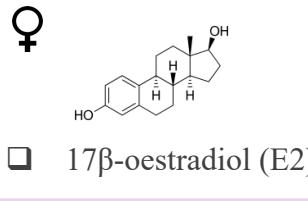


Blázquez et al 2008. Gen. Comp. Endocrinol. 158, 95–101.

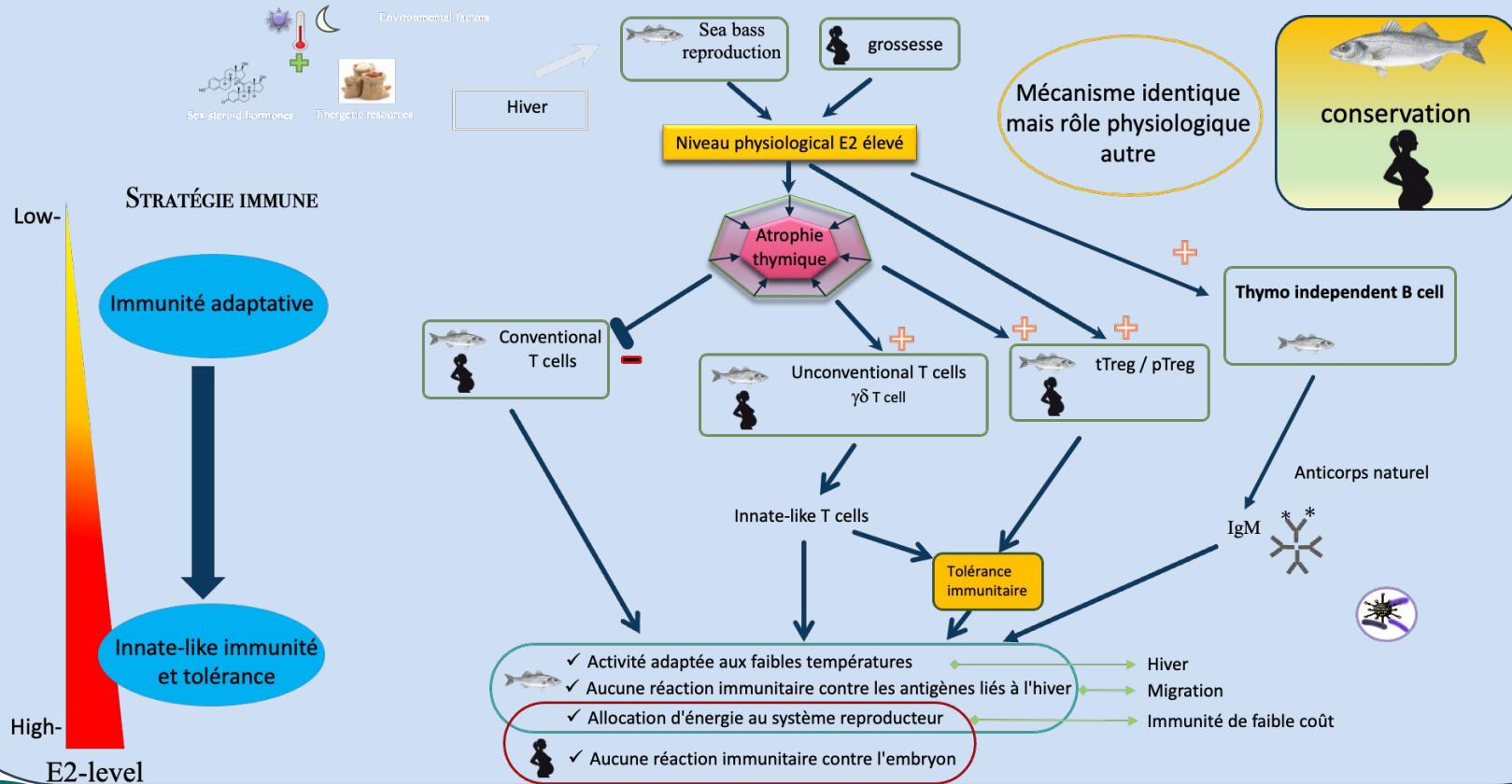
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Quelles sont les implications écologiques de la régulation estrogénique du SI en relation avec la reproduction ?

- Augmentation du taux plasmatique d'hormones sexuelles = reproduction
- Reproduction du bar = hiver
- hiver= changement important de la photopériode et température

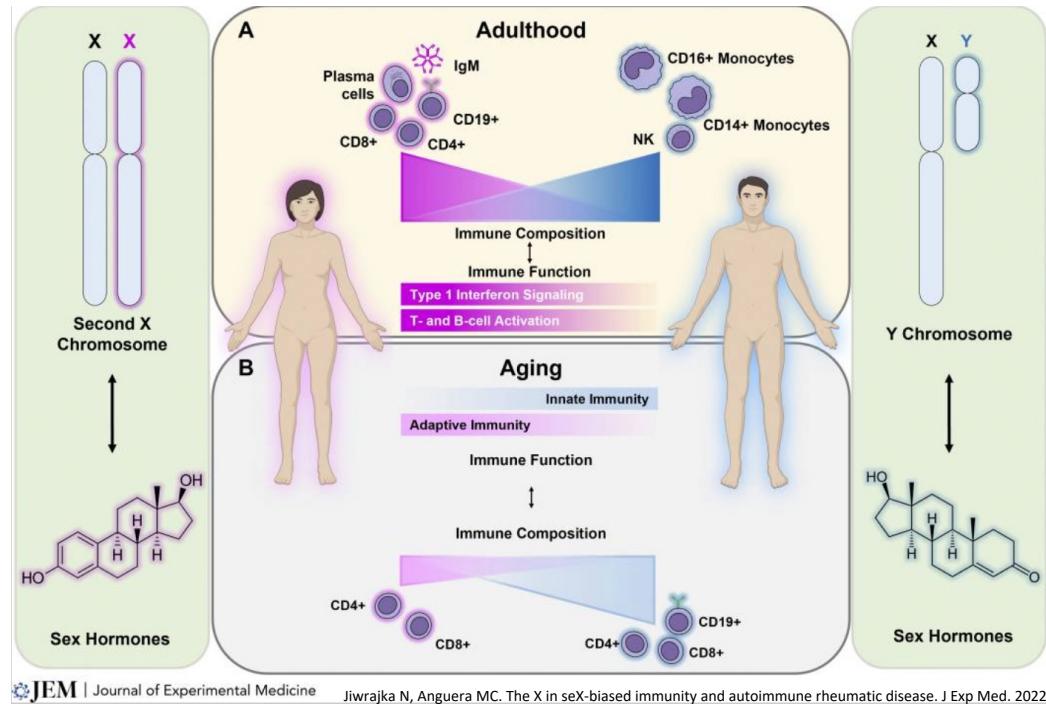


- Facteurs environnementaux
- Facteurs Endogènes



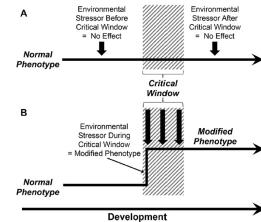
Plasticité du SI (thymus) avec âge

- Développement du SI
- Mise en place avant la **puberté**
- Diminution de la performance immunitaire avec l'âge



Détermination de fenêtres critiques durant l'ontogénèse du thymus

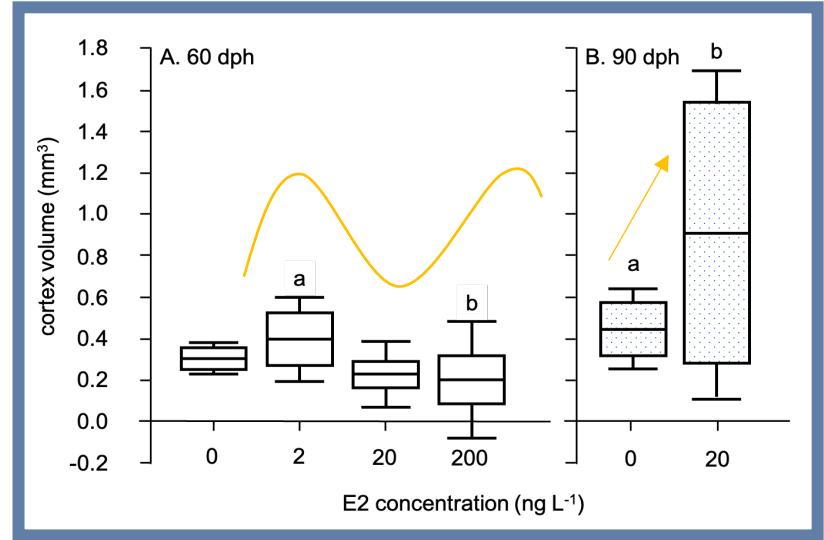
Experiment	E2 concentration [ng/L]	Initial fish age [dph]
1	0	67 dph
	20	
2	200	
	0	37
2		97
	20	127



- 60 dph *D. labrax* exposés à 2, 20 et 200 ng L⁻¹ E2 pendant 56 jours
- Alevins de différents âges (30, 90 et 120 dph) exposés à 20 ng L⁻¹ E2
- Evaluation des volumes du thymus total, cortex et médulla

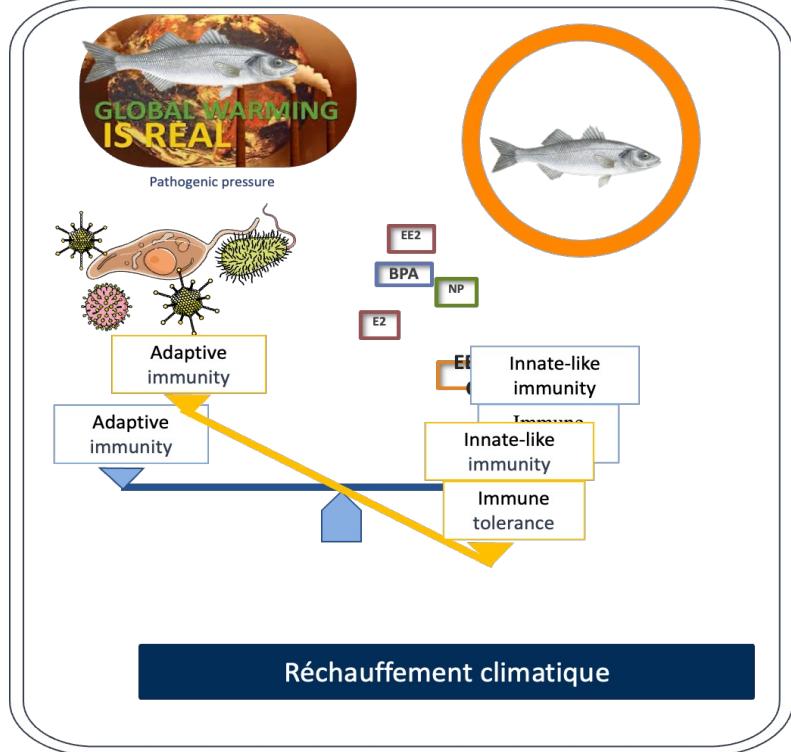
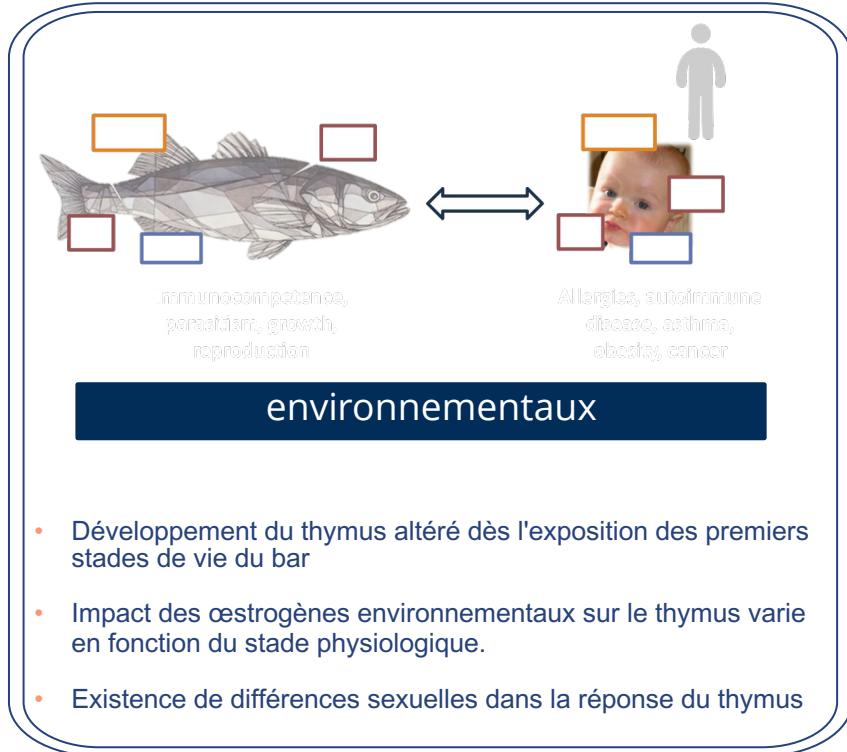
Thymus:

- Poissons de 60 dph
 - Courbe dose réponse non monotone
 - Atrophie du cortex
- Poissons de 90 dph
 - Hypertrophie thymique



E2 affecte la régionalisation du thymus et la maturation des cellules T

CONCLUSION



EFFECT OF OESTROGEN ON THE IMMUNE SYSTEM OF SEA BASS

Stock / Acclimation
E2 bath exposure (20 ng/l)
Group I: 47-54 g/fish
Group II: 60-90 g/fish
BCD1: 2.2×10^6 CFU/fish: 211-217 g/fish
BCD2: 1.0×10^6 CFU/fish: 223-226 g/fish
Critical windows based on thymus maturation

Bacterial Challenge with *Vibrio harveyi*

SUMMARY
This study evaluated the effects of oestrogen (E₂) on the immune system of sea bass (*Dicentrarchus labrax*). It was found that oestrogens affect thymus plasticity and T cell maturation, particularly during immune system ontogeny, therefore affecting the animals' immunocompetence and resistance to pathogens. Increasing our understanding of the impact of environmental contaminants on aquatic life lays the foundation for the enactment of stricter legislation to ensure safer aquatic environments for fish and humans.

KNOWLEDGE NEED
Endocrine Disrupting Chemicals (EDCs) can interfere with hormonal regulation in vertebrates and may disrupt their endocrine systems. A growing body of evidence indicates that many EDCs affect the immune systems of marine organisms, increasing their susceptibility to disease. Natural and synthetic oestrogens are among the most common EDCs. They are frequently found in surface waters, at disrupting concentrations, to which aquatic organisms are directly exposed. There is a clear need to gain further insights into the potential health effects on aquatic animals, such as sea bass. Specifically, little is known about effects of exposure to exogenous oestrogens during immune system ontogenesis and how this affects later immunocompetence.

POTENTIAL IMPACT

- Results provide evidence for the need for improved water quality for aquaculture species, which could lead to strengthened legislation, better sewage treatment, and reduction of farm runoff.
- A focus by the aquaculture industry on testing for and maintaining water quality, which will lead to improved overall stock health, in turn improving the economic viability of fish farming for certain species or in certain areas.

EATIP - Strategic Research and Innovation Agenda (SRIA) Thematic Area 2: Knowledge and Systems, Goal 1. To see the full list and descriptions of the thematic areas and goals, please visit: eatip.eu/?page_id=46

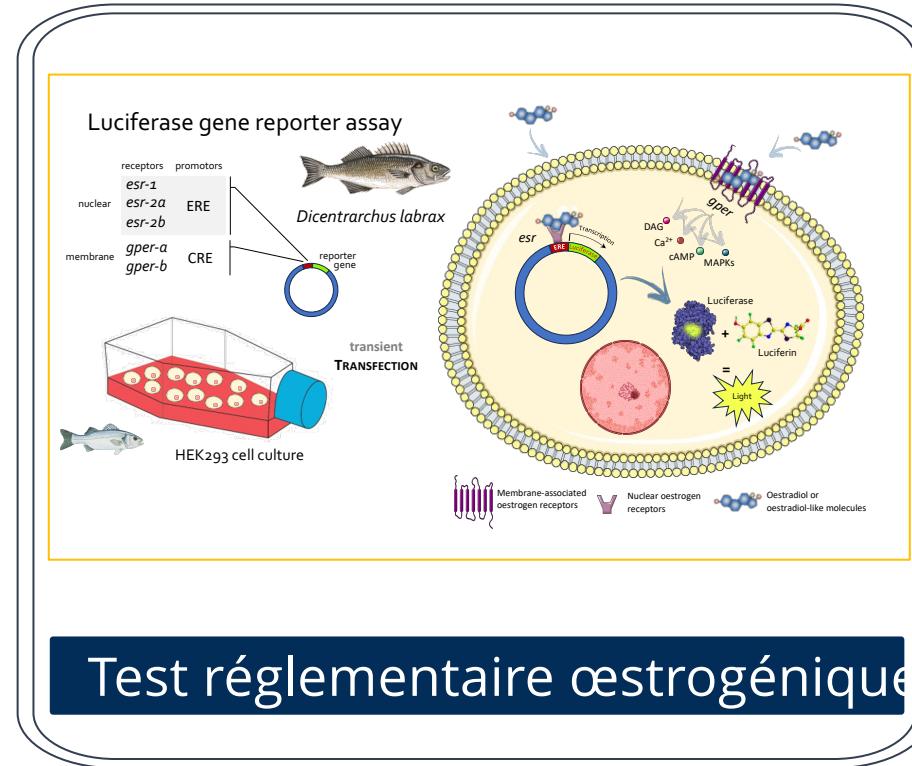
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Outlook



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Merci de votre attention!