



HELCOM Stakeholder Conference 2025 A 'one Baltic' approach towards a sea unaffected by hazardous substances **31 March 2025** 



## PHARMACEUTICALS IN THE BALTIC SEA: Importance and risks Daniel Malnes

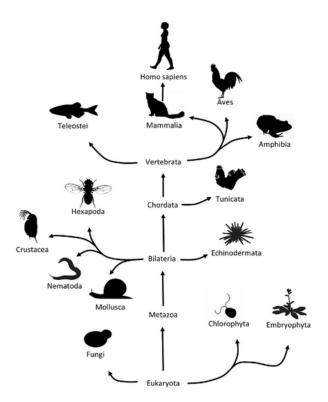




#### PHARMACEUTICALS: INTENTIONALLY BIOLOGICALLY RELEVANT

- Animal studies to prove efficiency in humans
  - Assumed that drug targets are conserved
- How far are drug targets conserved through evolution?
- Designed to affect biological processes at low concentrations
- Some pharmaceuticals designed to 'kill off' organisms
  - Antibacterials (affecting cyanobacteria?)
  - Antiparasitic products, insecticides and repellents
- Some pharmaceuticals designed to 'stop' biological processes
  - Contraception (affecting reproductive success)
  - Epilepsy (reducing unintentional movement)





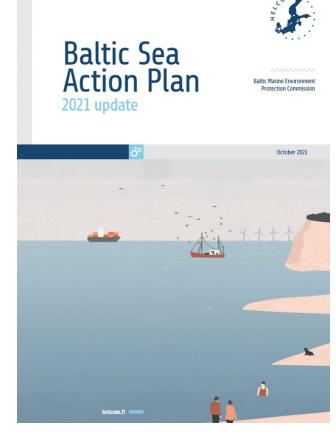
Evolutionary-conserved drug targets Verbruggen, B., et al. (2018). *Nucleic Acids Research*, 46(D1), D930-D936.



#### PHARMASEA PROJECT IN BRIEF (NOV 2023 - JAN 2025)

- Baltic Sea Action Plan 2021:
  - HL22: Improve the knowledge base on occurrence of pharmaceutical substances in the environment, their persistence and harmful effects and ensure availability of this information for a broad expert community by 2025.
  - HL23: Identify priority pharmaceuticals by 2024 utilising the best available knowledge on their releases into the aquatic environment, environmental effects and available data on their use in the region, for efficient risk reduction and for subsequent integration of these substances to HELCOM assessments, as indicators of the state of the Baltic Sea and environmental pressure





[3]







#### TWO DATA PIPELINES FOR SCIENCE-BASED POLICY DECISIONS

#### Occurrence data

#### Sales data

#### Data processing

#### Environmental hazard assessment

- 1. Priority substances
- 2. Substances of concern

- 1. Environmental matrix screening
- 2. Wastewater screening





• Proposed priority pharmaceuticals in the Baltic Sea

Pharmaceutical	Group	
17α-ethinylestradiol	Systemic hormonal preparations	
17β-estradiol	Systemic hormonal preparations	
Estrone	Systemic hormonal preparations	
Azithromycin	Antibacterials	
Clarithromycin	Antibacterials	
Erythromycin	Antibacterials	
Diclofenac	Non-steroidal anti-inflammatory drugs	
Ibuprofen	Non-steroidal anti-inflammatory drugs	
Carbamazepine	Antiepileptics	





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#### Systemic hormonal preparations

- Reduces likelihood of pregnancy ("toxic to reproduction")
- Some of natural origin (e.g., 17β-estradiol, estrone)
- Others synthetic (17 $\alpha$ ethinylestradiol)





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Carbamazepine		

#### **Antibacterials/antibiotics**

- Used against bacterial infections
- Affects cyanobacteria
- Can promote development of antimicrobial resistance





Proposed priority pharmaceuticals in the Baltic Sea

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### Non-steroidal anti-inflammatory drugs

- To treat pain, inflammation, and reduce high body temperature
- Sometimes available as creams/ointments





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#### **Antiepileptics**

- Reduces involuntary shaking
- EQS based on reproductive success in *Daphnia*



• Proposed pharmaceuticals of concern in the Baltic Sea

Pharmaceutical	Group
Metformin	Drugs used in diabetics
Clotrimazole	Antifungal
Norethisterone	Systemic hormonal preparations
Amoxicillin	Antibacterials
Ciprofloxacin	Antibacterials
Ofloxacin	Antibacterials
Sulfamethoxazole	Antibacterials
Tetracycline	Antibacterials
Trimethoprim	Antibacterials
Venlafaxine and O-desmethylvenlafaxine	Antidepressants
Cypermethrin	Antiparasitics





- 33% of Baltic Sea population included in assessment
- A total of 1,470 pharmaceuticals were assessed
- Precautionary approach showed 38 pharmaceuticals of interest
  - May be reduced after further 'desk-work'
  - Remaining pharmaceuticals subject to environmental/wastewater screening





	Pharmaceutical group	Number of pharmaceuticals of interest
	Antibiotics/Antibacterials	13
	Drugs used in erectile dysfunction	1
	Hormonal preparations	1
	Antineoplastic and immunomodulating agents	4
	Anti-acne preparations	2
	Antidepressants	2
	Calcium channel blockers	2
	X-ray contrast media	2
	Antidiarrheals, intestinal anti-inflammatory/antiinfective agents	1
	Antifungals	1
	Antihypertensives	1
	Antiinfectives and antiseptics	1
	Antipsychotic	1
	Antithrombotic agents	1 [12]
	Hypnotics and sedatives	1





- PharmaSea draft report results propose:
  - Priority substances (exceeding marine EQS)
  - Substances of concern (exceeding other EQS/PNECs)
  - Potential pharmaceuticals for sampling in
    - Environmental matrices
    - Wastewaters







# THANK YOU FOR THE ATTENTION PHARMACEUTICALS IN THE BALTIC SEA: Importance and risks

**Daniel Malnes**