

## **NeuroMITilus Unraveling the role of mitochondria in coping with hypoxia/reoxygenation stress in *Mytilus* nervous system**

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### **Abstract**

Mussels are ecologically dominant in the intertidal zones and have successfully adapted to fluctuations in oxygen (O<sub>2</sub>) availability, due to both the intrinsic characteristics of the environment and the deoxygenation of aquatic habitats caused by eutrophication and warming. Facing **hypoxia-reoxygenation** (H/R) stress requires exquisite homeostatic responses, especially in tissue with a high O<sub>2</sub> and metabolic demand, such as the **nervous system** (NS). In hypoxia-sensitive organisms such as mammals, **mitochondria** are the hub of H/R-induced neural damage, since O<sub>2</sub> deficiency impairs ATP production, whereas a surge of O<sub>2</sub> causes an excessive production of toxic O<sub>2</sub> reactive species (ROS).

The main objective of *NeuroMITilus* is to expand the current concept of tolerance to H/R stress by elucidating the resilience mechanisms specific of the NS in the Mediterranean mussel, *Mytilus galloprovincialis* (*Mg*), with a particular focus on **mitochondrial homeostasis** (dynamics, function, bioenergetics), autophagy, cell death and inflammatory responses. Molecular dissection of the pathways underlying these resilience strategies will allow us to identify specific mechanisms needed to face the H/R stress.

The cutting-edge experimental approaches planned in the project - including molecular and cellular biology techniques, "Omics", biochemical and computational analyses, imaging, and physiology techniques - will make possible to reach new knowledge on the **molecular mechanisms of environmental adaptation** in mussel NS. The latter has also a strong biomedical impact: hypoxia and hypoxia-reperfusion occur in medical conditions in humans, such as stroke. Therefore, the novel data on hypoxia-tolerant molluscs will bring to light the evolutionarily tested solutions to prevent or mitigate the damage associated with H/R.