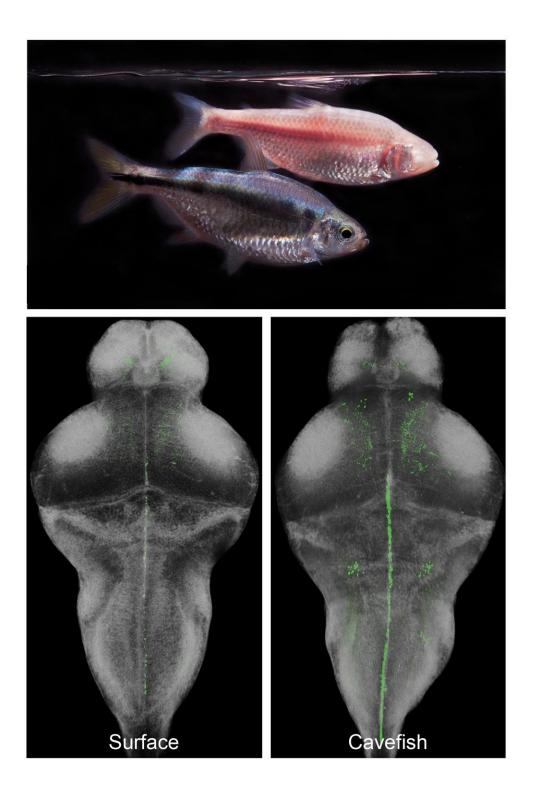


Sleepless in Latin America: Blind cavefish, extreme environments and insomnia

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Hypocretin, previously implicated in human narcolepsy, is expressed at higher levels in the cavefish brain (A. mexicanus). Image of Mexican cavefish and surface fish counterparts (top). Brains of surface fish (left) and cavefish (right) with wake-promoting Hypocretin protein depicted in green (bottom). Credit: Alex Keene, Florida Atlantic University

Nearly all animals studied need to sleep, but little is known about why some animals sleep most of the day, while others sleep very little. As a result of living in total and permanent darkness in a small location in northeast Mexico, the eyeless, tiny blind Mexican cavefish (*Astyanax mexicanus*) has evolved sleeplessness, snoozing far less than their riverdwelling relatives. These fish have been studied for nearly 100 years for their fascinating traits, yet little is known about how their behavior evolves. Because of their lack of sleep, they provide a great model to study human sleep disorders such as insomnia.

A new study published in *eLife* by neuroscientists in the Charles E. Schmidt College of Science at Florida Atlantic University is providing new insight into the evolution of sleep.

For the study, the researchers found that the neuropeptide Hypocretin, previously implicated in human narcolepsy, is expressed at higher levels in the cavefish brain. While loss of Hypocretin has been shown to cause narcolepsy in human and animal models, cavefish evolved to have higher levels of Hypocretin resulting in sleeplessness.

"These findings suggest that differences in Hypocretin production may explain variation in sleep between animal species, or even between individual people," said Alex Keene, Ph.D., senior author and an associate professor in the Department of Biological Sciences and a member of the FAU Brain Institute. "It may also provide important



insight into how we might build a brain that does not need to sleep."

The team of scientists, led by Keene, set out to uncover the mechanics behind sleep loss in A. mexicanus to find out more about how deficiencies in Hypocretin signaling are associated with altered sleep and narcolepsy, a chronic sleep disorder that causes overwhelming daytime drowsiness, in diverse vertebrate organisms.

To accomplish that, Keene and his team conducted a comparative study among A. mexicanus surface and cavefish <u>brain</u> cells. Their results show that the number of Hypocretin cells in cavefish is significantly higher than that of their surface-dwelling relatives.

Next, they showed that inhibiting Hypocretin signaling, genetically or pharmacologically, increased sleep in cavefish, while it had very little effect on surface fish.

"A central question moving forward is why Hypocretin functions differently in cavefish," said Keene. "We want to understand what advantage sleep loss provides these fish in the cave environment."

More information: James B Jaggard et al. Hypocretin underlies the evolution of sleep loss in the Mexican cavefish, *eLife* (2018). <u>DOI:</u> 10.7554/eLife.32637

Provided by Florida Atlantic University

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