

Taking the pulse of flies: Study reveals fly hearts respond to danger the same way human hearts do

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The fly's heart (left) - a minuscule structure made of two single rows of cells - is imaged non-invasively while the fly walks freely on a custom-made device (right). Credit: Charlotte Rosher, Moita lab, Champalimaud Foundation.

The sound of an accelerating heartbeat can instantly send chills down your spine. You know that sound means trouble. We are so accustomed



to the way our hearts seem to continuously mirror how we feel that we can easily imagine different hearts racing, aching or skipping a beat.

But do the hearts of other animals actually follow the same rules when in danger? When it comes to our fellow vertebrates—frogs, cats, antelope—the answer has been long-known to be "yes." But what about insects?

According to a study by scientists at the Champalimaud Centre for the Unknown in Lisbon, Portugal, which was published today (October 27th) in the journal *Current Biology*, the hearts of fruit flies respond to danger in very much the same way human hearts do.

"We were quite surprised by this result," recalls Marta Moita, the neuroscientist that led the project. "We know that when vertebrates face a threat, their autonomic nervous system kicks into action, generating the changes in cardiac activity that we are all familiar with. However, this system doesn't exist in insects, and so it was unclear whether they would exhibit similar cardiac alterations."

The team zoomed in on the fly's heart—a minuscule structure made up of two single rows of cells. The researchers followed the heart's activity through the fly's transparent exoskeleton while it was walking about by lighting up the heart's cells with fluorescent molecules. Occasionally, a dark expanding circle appeared on a large screen in front of the fly, mimicking an approaching threat.

"Amazingly, just like in humans, the fly's heart changed its activity depending on which defense response is assumed. If the fly decided to escape, the heart accelerated, but if the fly froze in place for a sustained period of time, its heart slowed down," Barrios recounts.

"This finding is very intriguing," Moita adds. "Since flies don't have an



autonomic nervous system, it means there's another mechanism at play here. The question is whether there's an <u>autonomic-nervous-system</u>-like structure that we are not aware of, or is there an entirely different mechanism yet to be discovered."

A change of heart

Remarkably, this unexpected result was only the first in a series of revelations. The next happened when the team examined the heart's activity in more detail.

"The structure of the fly's heart is very different from ours: it's just a single tube," Barrios explains. "And since its body is essentially divided into two sealed compartments, the heart alternates pumping in two directions."

The team tested whether the direction of pumping also changed depending on the fly's defensive response, and discovered that in both cases—escape and freezing—the heart was pumping more actively towards the front section of the fly.

"Pumping more nutrients into the front section while escaping makes sense. This is where the brain, legs and wings are located, so that's where the action is. But we didn't expect to see this while the fly was freezing," says Barrios.

According to the researchers, freezing is considered an energy-saving behavior. Indeed, the slowing down of the heart indicates just that. But then, what was the reason the heart was pumping more actively towards the front? The results of this next series of experiments ended up undermining yet another widely held scientific theory.



Freezing burns calories

"We suspected it meant that even though it was freezing, the fly was getting ready for action," Moita explains. "And that it was consuming energy to maintain this state of preparedness."

To test this hypothesis, the team compared the sugar levels of flies that froze with sugar levels of flies that were exposed to neutral images, and therefore didn't exhibit any defensive behaviors. The results were striking: flies that froze had significantly lower sugar levels.

"This finding refutes the generally held belief that freezing is a passive, energy-saving behavioral state," Moita argues. "Instead, it suggests that freezing is a state of active preparedness. Now, the question is—what is the fly preparing for? What is the range of actions that may follow freezing, and how is the choice between actions made by the brain?"

A new path

These questions join a string of new research avenues raised by this study. One of the most pressing is identifying the neural structure that controls cardiac responses to danger in flies, and deciphering how it works.

"Since flies and humans share many genes, the hearts of flies are commonly used to study various aspects of cardiology, particularly relating to disease," Barrios points out. "However, little attention has been directed towards how the fly's <u>heart</u> responds to danger."

"Now that we've demonstrated this new commonality, we can move forward to investigate how it happens. Eventually, we hope that insight gained in the fly will lead to an understanding of how the brain controls



behavior in other animals, including humans," Moita Concludes.

More information: Marta A. Moita, Threat induces cardiac and metabolic changes that negatively impact survival in flies, *Current Biology* (2021). DOI: 10.1016/j.cub.2021.10.013. www.cell.com/current-biology/f ... 0960-9822(21)01367-1

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