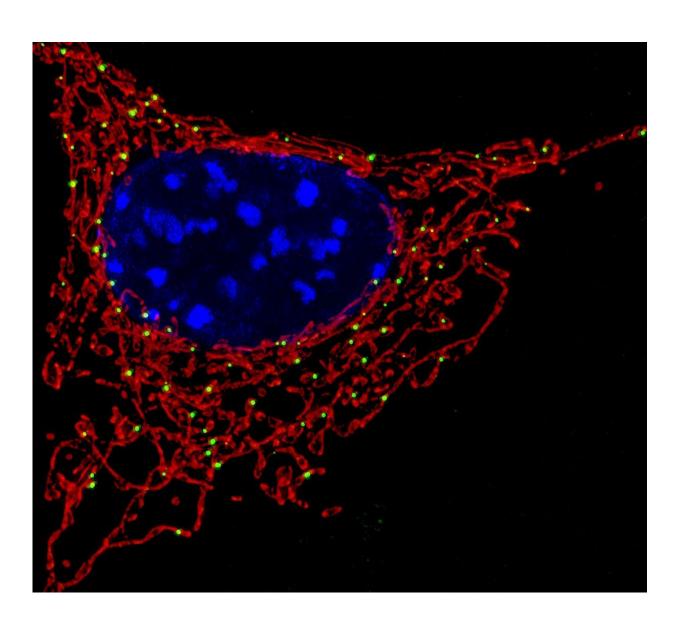


## Mitochondria work much like Tesla battery packs, study finds

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Mitochondria (in red) are organelles found in most cells. They generate a cell's chemical energy. Credit: Flickr/NICHD-U. Manor



For years, scientists assumed that mitochondria—the energy-generating centers of living cells—worked much like household batteries, generating energy from a chemical reaction inside a single chamber or cell. Now, UCLA researchers have shown that mitochondria are instead made up of many individual bioelectric units that generate energy in an array, similar to a Tesla electric car battery that packs thousands of battery cells to manage energy safely and provide fast access to very high current.

"Nobody had looked at this before because we were so locked into this way of thinking; the assumption was that one mitochondrion meant one battery," said Dr. Orian Shirihai, a professor of medicine in endocrinology and pharmacology at the David Geffen School of Medicine at UCLA and senior author of the study published in *EMBO Journal*. It is also not a coincidence that this has taken place in California, where an electric vehicle revolution has made its impact everywhere on campus.

Mitochondria are one type of organelle—tiny structures that perform specific functions within a cell. All <u>cells</u> in the human body, except for <u>red blood cells</u>, contain one or more—sometimes several thousand—<u>mitochondria</u>. These organelles have a smooth outer membrane and a wrinkled inner membrane that has folds, called cristae, extending toward the mitochondrion's center. Until now, researchers thought that the purpose of the inner membrane's wrinkly texture was simply to increase the surface area for energy production.

"Electric vehicle engineers told me about advantages of having many small battery cells instead of one large one; if something happens to one cell, the system can keep working, and multiple small batteries can provide a very high current when you need it," Shirihai said.



Tesla vehicles, for instance, are powered by 5,000 to 7,000 small <u>battery</u> <u>cells</u>, depending on the car model. The small batteries, arranged in a large grid, let the vehicles rapidly charge, efficiently cool and quickly use a large amount of power to accelerate.

Shirihai, using conventional microscopy, had been seeing that cells function well with a small number of very long mitochondria, which did not match the idea of many small batteries. He started to wonder whether each mitochondria really was a single large battery.

Shirihai and colleagues faced a difficult challenge to develop an approach to map the voltage on the membrane of mitochondria in living cells at a resolution never seen before. Two students, Dane Wolf and Mayuko Segawa, optimized a form of high-resolution microscopy to visualize the interior of mitochondria and watch energy production and voltage distribution inside the organelle.

"What the images told us was that each of these cristae is electrically independent, functioning as an autonomous battery," Shirihai said. "One cristae can get damaged and stop functioning while the others maintain their membrane potential."

Between each cristae, where the <u>inner membrane</u> of the mitochondria loops back outward, clusters of proteins form the boundaries of each cristae. Researchers knew that without these proteins, mitochondria were more sensitive to damage. Now, Shirihai has explained why: The proteins normally divide each cristae from their neighbors, acting as electrical insulators. In cells lacking those proteins, Shirihai found that each mitochondria became one giant battery.

"The battery experts I had originally talked to were very excited to hear that they were right," Shirihai said. "It turns out that mitochondria and Teslas, with their many small batteries, are a case of convergent



evolution."

The discovery may help lead to a new understanding of mitochondria's roles in aging, disease and medical complications, the researchers said. The authors note that many medical conditions are associated with severe disturbances in cristae structure.

**More information:** Dane M Wolf et al. Individual cristae within the same mitochondrion display different membrane potentials and are functionally independent, *The EMBO Journal* (2019). DOI: 10.15252/embj.2018101056

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