

Researchers map structure of mitochondria at different life stages

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This partial figure from the paper illustrates the steps taken to compare the mitochondria of young and old mice. First, thin tissue slices were imaged with an advanced imaging method called serial block face-scanning electron microscopy. Then, the mitochondria were identified and 3D reconstructions were overlaid on the tissue images. Finally, the 3D reconstructions were analyzed for volume, surface area, and other metrics. Credit: *Advanced Biology* (2023). DOI: 10.1002/adbi.202300186



Mitochondria, organelles with an inner and outer membrane, are responsible for creating the energy that cells use to survive, and their morphology is key to accomplishing this task. The inner mitochondrial membrane contains folds, called cristae, that maximize the surface area available for mitochondrial energetic processes.

Although <u>oxidative stress</u> is a normal part of the reactions that take place inside <u>mitochondria</u>, excess stress and dysfunction are associated with disease and aging. Yet, as the number and function of mitochondria are highly dependent on what tissue they're in, scientists are interested in exploring the changes that occur in mitochondria across tissues as we age.

The lab of Antentor O. Hinton Jr., assistant professor of molecular physiology and biophysics, came together with collaborators from across Vanderbilt University, Vanderbilt University Medical Center, the U.S., and Brazil to determine the differences in the 3D <u>morphology</u> of mitochondria and their cristae in <u>brown adipose tissue</u>.

BAT uses the cristae within mitochondria for thermogenesis—the dissipation of heat that helps humans classify as warm-blooded animals. Scientists have observed a decrease in BAT thermogenic function in both older rodents and humans, and the decline has been associated with the development of metabolic disorders, including obesity and diabetes.

Changes in mitochondrial morphology in the context of BAT is not well understood, so researchers, led by Hinton lab Ph.D. student Amber Crabtree, developed a quantitative 3D electron microscopy approach to map cristae network organization in the BAT of adult and aged mice.

The researchers' findings, <u>published</u> in *Advanced Biology* under the title "Defining Mitochondrial Cristae Morphology Changes Induced by Aging in Brown Adipose Tissue" and featured on the cover in January, build



upon prior work that suggests that the surface area, volume, and complexity of mitochondria grow as mice age. However, Crabtree and Hinton also showed that the surface area, volume, and complexity of the cristae decrease during aging.

"These findings highlight the importance of considering changes in overall cristae density and not relying solely on changes in mitochondrial mass to quantify mitochondrial morphology changes," the authors said in the paper.

The researchers also reported that the overall shape of mitochondria influences the thermogenic capacity of BAT, as high-thermogenic BAT mitochondria were round or spherical, while low-thermogenic mitochondria were more elongated. Reduced thermogenic capacity was also associated with aging and obesity, which suggests that mitochondrial structure and function may have implications for overall metabolic health.

Hinton hopes that future studies will determine whether there are sexdependent differences in BAT mitochondrial structure across aging. Specifically, probing mitochondrial structure and function within BAT and across other tissues and other species may broaden our understanding of aging mechanisms and identify potential therapies to treat ailments associated with aging and mitochondrial dysfunction.

More information: Amber Crabtree et al, Defining Mitochondrial Cristae Morphology Changes Induced by Aging in Brown Adipose Tissue, *Advanced Biology* (2023). <u>DOI: 10.1002/adbi.202300186</u>



Provided by Vanderbilt University

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