

Brain disease research reveals differences between sexes

March 16 2021



Human neurodegenerative diseases affect men and women differently, yet sex is rarely included in in vitro bioengineered models of neurodegenerative disease. Sex-related differences include a wide range of biochemical factors, gene expression, and biomechanical cues. These sex differences must be included in blood-brain barrier models to improve the understanding of sex differences in neurodegenerative disease and eventually realize personalized medicine. Credit: Callie Weber



Men and women are impacted differently by brain diseases, like Alzheimer's disease and Parkinson's disease. Researchers are urging their colleagues to remember those differences when researching treatments and cures.

In *APL Bioengineering*, University of Maryland scientists highlight a growing body of research suggesting sex differences play roles in how patients respond to <u>brain diseases</u>, as well as multiple sclerosis, motor neuron <u>disease</u>, and other brain ailments.

That is progress from just a few years ago, said Alisa Morss Clyne, director of the university's Vascular Kinetics Laboratory.

"I have worked with <u>vascular cells</u> for 20 years and, up until maybe about five years ago, if you asked if the sex of my cells mattered at all, I would have said no," Clyne said. Then, she worked on a difficult study in which data appeared "all over the place."

"We separated the cell data by sex, and it all made sense," Clyne said. "It was an awakening for me that we should be studying this."

As of 2020, an estimated 5.8 million Americans were diagnosed with Alzheimer's disease, another 1 million with Parkinson's disease, 914,000 with multiple sclerosis, and 63,000 with motor neuron disease. These diseases happen when <u>nerve cells</u> in the brain and nervous system quit working and, ultimately, die.

The changes are associated with the breakdown of what is called the <u>blood-brain barrier</u>—a border of cells that keeps the wrong kind of molecules in the bloodstream from entering the brain and damaging it.

Published research has shown differences in the blood-<u>brain</u> barriers of men and women. Some of the research suggests the barrier can be



stronger in women than men, and the barriers in men and women are built and behave differently.

That could factor into known differences in the sexes, such as Alzheimer's disease being more prevalent in older <u>women</u> than men, while Parkinson's impacts men more frequently and tends to do so more severely.

The authors said they hope their article will serve as a reminder to researchers not just in their own field, but across the sciences, that accounting for sex differences leads to better results.

"I think there is an awakening in the past 10 years or so that you cannot ignore sex differences," Clyne said. "My goal is to inspire people to include sex differences in their research, no matter what research they are doing."

More information: "Sex differences in the blood-brain barrier and neurodegenerative diseases" *APL Bioengineering*, <u>aip.scitation.org/doi/10.1063/5.0035610</u>

Provided by American Institute of Physics

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