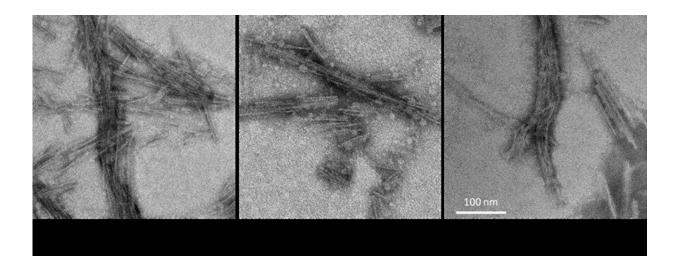


Scientists develop test for uncommon brain diseases

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Representative negative-stained transmission electron microscopy images of 4R RT-QuIC products seeded with brain homogenates from individuals with the designated diseases - frontotemporal dementia and Parkinsonism linked to chromosome 17; corticobasal degeneration; and progressive supranuclear palsy. Credit: NIAID

National Institutes of Health (NIH) scientists have developed an ultrasensitive new test to detect abnormal forms of the protein tau associated with uncommon types of neurodegenerative diseases called tauopathies. As they describe in *Acta Neuropathologica*, this advance gives them hope of using cerebrospinal fluid, or CSF—an accessible patient sample—to diagnose these and perhaps other, more common



neurological diseases, such as Alzheimer's disease.

Scientists have linked the abnormal deposition of tau in the brain to at least 25 different neurodegenerative diseases. However, to accurately diagnose these diseases, brain tissue often must be analyzed after the patient has died.

For their study, the researchers used the same test concept they developed when using post-mortem <u>brain tissue</u> samples to detect the abnormal tau types associated with Pick disease, Alzheimer's disease and chronic traumatic encephalopathy (CTE). They adapted the test to use CSF for the detection of abnormal tau of progressive supranuclear palsy (PSP), corticobasal degeneration (CBD), and other less common tauopathies.

They detected abnormal tau in CSF from both living and deceased patients. In one case, the test led to a corrected diagnosis in a patient who had died from CBD, but who was initially diagnosed with PSP. The new test is called 4R RT-QuIC—which stands for 4-repeat tau protein amplified in a <u>real-time</u>, quaking-induced <u>conversion process</u>.

The test was developed at NIH's Rocky Mountain Laboratories, part of the National Institute of Allergy and Infectious Diseases. Study collaborators, who provided patient specimens and clinical guidance, included investigators from the Mayo Clinic in Jacksonville, Florida; the University of Bologna, Italy; the University of California, San Diego; the University of California, San Francisco; Indiana University; and the University of Verona in Italy.

The researchers plan to continue evaluating the clinical performance of 4R RT-QuIC by analyzing larger sets of CSF samples. One focus will be to compare <u>test</u> results from tauopathy patients who agree to provide CSF samples both before and after death. The scientists hope this type



of evaluation will help them better understand how abnormal tau in CSF evolves during brain <u>disease</u>.

More information: Eri Saijo et al, 4-Repeat tau seeds and templating subtypes as brain and CSF biomarkers of frontotemporal lobar degeneration, *Acta Neuropathologica* (2019). <u>DOI:</u> 10.1007/s00401-019-02080-2

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