

You've likely heard of the brain's gray matter. Here's why the white matter is important too

May 6 2022, by Christopher Filley



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Who has not contemplated how a memory is formed, a sentence



generated, a sunset appreciated, a creative act performed or a heinous crime committed?

The <u>human brain</u> is a three-pound organ that remains largely an enigma. But most people have heard of the brain's <u>gray matter</u>, which is needed for cognitive functions such as learning, remembering and reasoning.

More specifically, gray matter refers to regions throughout the brain where <u>nerve cells</u>—<u>known as neurons</u>—are concentrated. The region considered most important for cognition is <u>the cerebral cortex</u>, a thin layer of gray matter on the brain's surface.

But the other half of the brain—<u>the white matter</u>—is often overlooked. White matter lies below the cortex and also deeper in the brain. Wherever it is found, <u>white matter</u> connects neurons within the gray matter to each other.

I am a professor of neurology and psychiatry and the director of the behavioral neurology section at the University of Colorado Medical School. My work involves the evaluation, treatment and investigation of older adults with dementia and younger people with traumatic brain injury.

Finding out how these disorders affect the brain has motivated many years of my study. I believe that understanding white matter is perhaps a key to understanding these disorders. But so far, researchers have generally not given white matter the attention it deserves.

Figuring out the white matter

This lack of recognition largely stems from the difficulty in studying white matter. Because it's located below the surface of the brain, even the most high-tech imaging can't easily resolve its details. But recent



findings, made possible by advancements in brain imaging and autopsy examinations, are beginning to show researchers how critical white matter is.

White matter is comprised of many <u>billions of axons</u>, which are like long cables that carry electrical signals. Think of them as elongated tails that act as extensions of the neurons. The axons connect neurons to each other at junctions called synapses. That is where communication between neurons takes place.

Axons come together in bundles, or tracts, that course throughout the brain. Placed end to end, their combined length in a single human brain is approximately 85,000 miles. Many axons are <u>insulated with myelin</u>, a layer of mostly fat that speeds up electrical signaling, or communication, between neurons by up to 100 times.

This increased speed is crucial for <u>all brain functions</u> and is partly why Homo sapiens have unique mental capacities. While there's no doubt <u>our</u> <u>large brains</u> are due to evolution's addition of neurons over eons, there has been an even greater <u>increase in white matter</u> over evolutionary time.

This little-known fact has profound implications. The increased volume of white matter—mainly from the <u>myelin sheaths</u> that surround axons—enhances the efficiency of neurons in the gray matter to optimize <u>brain function</u>.

Imagine a nation of cities that are all functioning independently, but not linked to other cities by roads, wires, the internet or any other connections. This scenario would be analogous to the brain without white matter. Higher functions like language and memory are organized into networks in which gray matter regions are connected by white matter tracts. The more extensive and efficient those connections, the better the brain works.



White matter and Alzheimer's

Given its essential role in the connections between brain cells, <u>damaged</u> <u>white matter</u> can disturb any aspect of cognitive or emotional function. White matter pathology is present in many brain disorders and can be severe enough <u>to cause dementia</u>. Damage to myelin is common in these disorders, and when the disease or injury is more severe, axons can also be damaged.

More than 30 years ago, my colleagues and I described this syndrome as <u>white matter dementia</u>. In this condition, the dysfunctional white matter is no longer adequately performing as a connector, meaning that the gray matter cannot act together in a seamless and synchronous manner. The brain, in essence, has been disconnected from itself.

Equally important is the possibility that white matter dysfunction plays a role in many diseases currently thought to originate in gray matter. Some of these diseases stubbornly defy understanding. For example, I suspect white matter damage may be critical in the early phases of Alzheimer's disease and traumatic brain injury.

Alzheimer's is the most common type of dementia in older individuals. It can impair cognitive function and rob people of their very identity. No cure or <u>effective treatment</u> exists. Ever since <u>Alois Alzheimer's 1907</u> <u>observations</u> of gray matter proteins—called amyloid and tau—neuroscientists have believed the buildup of these proteins <u>is the</u> <u>central problem</u> behind Alzheimer's. Yet many drugs that remove these proteins <u>do not stop</u> the patients' cognitive decline.

<u>Recent findings increasingly suggest</u> that white matter damage—preceding the accumulation of those proteins—<u>may be the</u> <u>true culprit</u>. As brains age, they often experience gradual loss of blood flow from the narrowing of vessels that convey blood from the heart.



Lower blood flow heavily impacts white matter.

Remarkably, there is even evidence that inherited forms of Alzheimer's also feature <u>early white matter abnormalities</u>. That means therapies aimed at maintaining blood flow to white matter may prove more effective than attempting to dislodge proteins. One simple treatment likely to help is <u>controlling high blood pressure</u>, as this can reduce the severity of white matter abnormalities.

White matter and traumatic brain injury

Patients with <u>traumatic brain injury</u>, particularly those with moderate or severe injuries, can have lifelong disability. One of the most ominous outcomes of TBI is <u>chronic traumatic encephalopathy</u>, a brain disease believed to cause progressive and irreversible dementia. In TBI patients, the accumulation of tau protein in gray matter is evident.

Researchers have long recognized that white matter damage is common in people who have sustained a TBI. <u>Observations from the brains</u> of those with repetitive traumatic brain injuries—<u>football players</u> and military veterans have been frequently studied—have shown that white matter damage is prominent, and may precede the appearance of tangled proteins in the gray matter.

Among scientists, there is a burgeoning excitement over the <u>new interest</u> in white matter. Researchers are now beginning to acknowledge that the traditional focus on the study of <u>gray matter</u> has not produced the results they hoped. Learning more about the half of the <u>brain</u> known as white <u>matter</u> may help us in the years ahead to find the answers needed to alleviate the suffering of millions.

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Provided by The Conversation

Citation: You've likely heard of the brain's gray matter. Here's why the white matter is important too (2022, May 6) retrieved 6 May 2024 from <u>https://medicalxpress.com/news/2022-05-youve-heard-brain-gray-white.html</u>

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