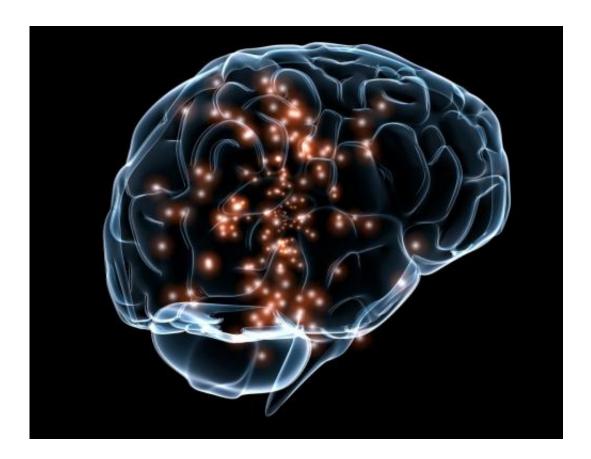


## New maps reveal the individual brain changes linked to different mental illness

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A breakthrough project mapping brain changes in nearly 1,300 people diagnosed with six different types of mental illness has revealed the extraordinary diversity of brain changes found in people with conditions like major depression and schizophrenia.



The study, published in *Nature Neuroscience* and led by researchers at Monash University's Turner Institute for Brain and Mental Health and School of Psychological Sciences, used brain imaging to measure the size, or volume, of over 1,000 different brain regions.

"Over the past few decades, researchers have mapped <u>brain areas</u> showing reduced volume in people diagnosed with a wide variety of mental illness, but this work has largely focused on group averages, which makes it difficult to understand what is happening in the brains of individual people," said Ph.D. student Ms Ashlea Segal, who led the research. "For example, knowing that the average height of the Australian population is about 1.7 m tells me very little about the height of my next-door neighbor," she added.

The team used new statistical techniques developed Prof Andre Marquand at the Donders Institute, Netherlands, who co-led the project, to map regions in the brain showing unusually small or large volumes in people diagnosed with either schizophrenia, depression, <u>bipolar disorder</u>, <u>obsessive-compulsive disorder</u>, <u>attention-deficit hyperactivity disorder</u>, or autism spectrum disorder.

"We used a statistical model to establish expectations about <u>brain size</u> given someone's age and sex. We can then quantify how much an individual person's brain volume deviates from these expectations, much like the growth charts commonly used for height and weight in pediatrics," said Professor Alex Fornito, who led the research team.

"We confirmed earlier findings that the specific brain regions showing large deviations in brain volume vary a lot across individuals, with no more than 7% of people with the same diagnosis showing a major deviation in the same brain area," said Professor Fornito. "This result means that it is difficult to pinpoint treatment targets or causal mechanisms by focusing on group averages alone. It may also explain



why people with the same diagnosis show wide variability in their symptom profiles and treatment outcomes," he added.

The team then investigated the connectivity of the areas showing large volume deviations. "Because the brain is a network, dysfunction in one area can spread to affect other, connected sites. We found that, while deviations occurred in distinct brain regions across different people, they were often connected to common upstream or downstream areas, meaning they aggregated within the same brain circuits," said Ms Segal.

"It's possible that this circuit-level overlap explains commonalities between people with the same diagnosis, such as, for example, why two people with schizophrenia generally have more symptoms in common than a person with schizophrenia and one with depression."

The team leveraged their new approach to identify potential treatment targets for different disorders. "We found that certain specific brain circuits were preferentially involved in some disorders, suggesting that they are potential treatment targets," explained Ms Segal.

"However, our findings suggest that these targets will only be appropriate for a subset of people. For instance, we found evidence that brain circuits linked to frontal areas were preferentially involved in depression. These circuits are commonly used as targets for non-invasive brain stimulation therapies, but our data suggest that they may only effective targets for around 1/3 of people."

The approach developed by the team opens new opportunities for mapping brain changes in mental illness. "The framework we have developed allows us to understand the diversity of <u>brain changes</u> in people with <u>mental illness</u> at different levels, from individual regions through to more widespread <u>brain circuits</u> and networks, offering a deeper insight into how the <u>brain</u> is affected in individual people," said



Professor Fornito.

**More information:** Regional, circuit and network heterogeneity of brain abnormalities in psychiatric disorders, *Nature Neuroscience* (2023). DOI: 10.1038/s41593-023-01404-6

Provided by Monash University

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