The New Science of Mind



By ERIC R. KANDEL



THESE days it is easy to get irritated with the exaggerated interpretations of brain imaging — for example, that a single fMRI scan can reveal our innermost feelings — and with inflated claims about our understanding of the biological basis of our higher mental processes.

Such irritation has led a number of thoughtful people to declare that we can never achieve a truly sophisticated understanding of the biological foundation of complex mental activity.

In fact, recent newspaper articles have argued that psychiatry is a "semi-science" whose practitioners cannot base their treatment of mental disorders on the same empirical evidence as physicians who treat disorders of the body can. The problem for many people is that we cannot point to the underlying biological bases of most psychiatric disorders. In fact, we are nowhere near understanding them as well as we understand disorders of the liver or the heart.

But this is starting to change.

Consider the biology of depression. We are beginning to discern the outlines of a complex neural circuit that

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becomes disordered in depressive illnesses. Helen Mayberg, at Emory University, and other scientists used brain-scanning techniques to identify several components of this circuit, two of which are particularly important.

One is Area 25 (the subcallosal cingulate region), which mediates our unconscious and motor responses to emotional stress; the other is the right anterior insula, a region where self-awareness and interpersonal experience come together.

These two regions connect to the hypothalamus, which plays a role in basic functions like sleep, appetite and libido, and to three other important regions of the brain: the amygdala, which evaluates emotional salience; the hippocampus, which is concerned with memory; and the prefrontal cortex, which is the seat of executive function and self-esteem. All of these regions can be disturbed in depressive illnesses.

In a recent study of people with depression, Professor Mayberg gave each person one of two types of treatment: cognitive behavioral therapy, a form of psychotherapy that trains people to view their feelings in more positive terms, or an antidepressant medication. She found that people who started with below-average baseline activity in the right anterior insula responded well to cognitive behavioral therapy, but not to the antidepressant. People with above-average activity responded to the antidepressant, but not to cognitive behavioral therapy. Thus, Professor Mayberg found that she could predict a depressed person's response to specific treatments from the baseline activity in the right anterior insula.

These results show us four very important things about the biology of mental disorders. First, the neural circuits disturbed by psychiatric disorders are likely to be very complex.

Second, we can identify specific, measurable markers of a mental disorder, and those biomarkers can predict the outcome of two different treatments: psychotherapy and medication.

Third, psychotherapy is a biological treatment, a brain therapy. It produces lasting, detectable physical changes in our brain, much as learning does.

And fourth, the effects of psychotherapy can be studied empirically. Aaron Beck, who pioneered the use of cognitive behavioral therapy, long insisted that psychotherapy has an empirical basis, that it is a science. Other forms of psychotherapy have been slower to move in this direction, in part because a number of psychotherapists believed that human behavior is too difficult to study in scientific terms.

ANY discussion of the biological basis of psychiatric disorders must include genetics. And, indeed, we are beginning to fit new pieces into the puzzle of how genetic mutations influence brain development.

Most mutations produce small differences in our genes, but scientists have recently discovered that some mutations give rise to structural differences in our chromosomes. Such differences are known as copy number variations.

People with copy number variations may be missing a small piece of DNA from a chromosome, or they may have an extra piece of that DNA.

Matthew State, at the University of California, San Francisco, has discovered a remarkable copy number variation involving chromosome 7. An extra copy of a particular segment of this chromosome greatly increases the risk of autism, which is characterized by social isolation. Yet the loss of that same segment results in Williams syndrome, a disorder characterized by intense sociability.

This single segment of chromosome 7 contains about 25 of the 21,000 or so genes in our genome, yet an extra copy or a missing copy has profound, and radically different, effects on social behavior.

The second finding is de novo point mutations, which arise spontaneously in the sperm of adult men. Sperm divide every 15 days. This continuous division and copying of DNA leads to errors, and the rate of error increases significantly with age: a 20-year-old will have an average of 25 de novo point mutations in his sperm, whereas a 40-year-old will have 65. These mutations are one reason older fathers are more likely to have children with autism and schizophrenia.

Our understanding of the biology of mental disorders has been slow in coming, but recent advances like these have shown us that mental disorders are biological in nature, that people are not responsible for having schizophrenia or depression, and that individual biology and genetics make significant contributions.

The result of such work is a new, unified science of mind that uses the combined power of cognitive psychology and neuroscience to examine the great remaining mysteries of mind: how we think, feel and experience ourselves as conscious human beings.

This new science of mind is based on the principle that our mind and our brain are inseparable. The brain is a complex biological organ possessing immense computational capability: it constructs our sensory experience, regulates our thoughts and emotions, and controls our actions. It is responsible not only for relatively simple motor behaviors like running and eating, but also for complex acts that we consider quintessentially human, like thinking, speaking and creating works of art. Looked at from this perspective, our mind is a set of operations carried out by our brain. The same principle of unity applies to mental disorders.

In years to come, this increased understanding of the physical workings of our brain will provide us with important insight into brain disorders, whether psychiatric or neurological. But if we persevere, it will do even more: it will give us new insights into who we are as human beings.

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