# **Comments** Controversies

## Unifying neurology and psychiatry—a 'prescient but premature' notion?

We read with interest Dr. Henry A. Nasrallah's call to unify psychiatry and neurology into 1 specialty (Current PSYCHIATRY, From the Editor, August 2013, p. 8-9; http://bit.ly/16wImL3). Trends in this direction already are evident. Medical schools, including New York University School of Medicine, the University of Massachusetts Medical School, and the Medical University of South Carolina, have 6-year combined psychiatry and neurology residency programs that prepare students for board exams in either specialty or both specialties. We offer considerations that we hope will energize and inform the discussion, in turn moving us toward discovery of an optimal framework for all stakeholders.

In support of unification, Dr. Nasrallah's editorial points to research advances in pharmacology, neuroimaging, and genetics. He writes that a "neuropharmacological revolution" is occurring, and it includes the discovery of medications that control symptoms of mood and anxiety disorders. The use of psychotropic medications certainly is escalating; >1 in every 10 Americans currently takes an

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antidepressant,12 reflecting a >400% increase over the past 2 decades.<sup>2</sup>

However, increasing use does not prove efficacy or establish causal mechanisms. Controversy persists regarding antidepressant efficacy, particularly for mild-to-moderate depression. The Sequenced Treatment Alternatives to Relieve Depression trial<sup>3</sup> was the largest study to evaluate antidepressant effectiveness.3 Summarizing the findings, Thomas Insel, MD, Director of the National Institute of Mental Health stated, "Most important, this study demonstrates that for at least 70% of patients, appropriate treatment with an SSRI [selective serotonin reuptake inhibitor] is not enough."4 What is clear is that the benefits of antidepressants are smaller than originally thought, and may be limited to patients with particular types of (severe) depression.

We agree that developments in neuroimaging and genetics of mental illness are exciting and some day may change the relationship between neurology and psychiatry. However, advanced neuroimaging technologies such as functional magnetic resonance imaging (fMRI) and diffusion tensor imaging (or diffusion MRI) have shown abnormalities involving brain-wide networks in psychiatric disorders5 not distinct regional abnormalities. In other words, psychiatry deals with mental phenomena that are modular in nature and not always reducible to their molecular origins, whereas in neurologic disorders, research linear cause-effect relationships and regional abnormalities are more common.

Genome-wide association studies have shown that few polymorphisms can be associated with numerous mental disorders6 (ie, multifinality), and that several genes may be associated



with a particular disorder (ie, equifinality). Multifinality and equfinality are characteristics of complex disorders, with nonlinear gene × gene and gene × environment interactions. In contrast, most heritable neurologic disorders follow a Mendelian pattern of inheritance and predictable cause-effect neuropathology. Furthermore, recent advances in epigenetics—in which environmental and social factors modify gene expression and affect patterns of inheritance<sup>7</sup>—have further complicated our understanding of genetic contributions to psychiatric illness, and have revealed limitations in reductionist models.

Although neurologic and psychiatric disorders ultimately reflect cerebral pathology, the former commonly result from circumscribed lesions, and the latter are more complexly determined, resulting from disruptions in brain circuitry influenced by genetic, epigenetic, environmental, and social factors.7 As much as we might like it, research in psychiatry has not progressed to the point of fully explaining the brain-based processes underlying complex psychiatric disorders, decipher complex gene × gene and gene × environment interactions, or associate certain genes and biological pathways with specific disorders. A move to formally combine neurology and

psychiatry may be prescient but premature because different sets of skills and methodologies might be required of clinicians and researchers in either specialty. More research is needed to shape and refine our disease and training models, and inform evidencebased practice.

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## Flying high with Bromo-DragonFLY

I enjoyed reading "New 'legal' highs: Kratom and methoxetamine" (CURRENT PSYCHIATRY, Pearls, August 2013, p.54-55; http://bit.ly/1gmPVFB), which discussed legal substances of abuse with adverse effects. I appreciate Dr. Julianna Troy's recommendation to be familiar with "legal" drugs given their prevalence and potential for adverse effects when taken alone or with psychotropics.

Another legal substance, Bromo-DragonFLY—so named for the resemblance of its chemical structure to the insect—is illegal in many European countries and is being abused in the United States, notably in New Orleans, Louisiana.¹ Available as a white or off-white powder, or as paper "blotters," the drug exerts serotonergic and noradrenergic receptor agonism.¹ It is characterized by slow onset but long duration of action.

Intoxication with Bromo-Dragon-FLY beyond the usual dose of 800 to 1,300 µg can result in dystonia, tachycardia, hypertension, psychosis, tachypnea, vasoconstriction with necrosis, seizures, hepatic or renal dysfunction, and death. Treatment comprises hydration, respiratory support, and benzodiazepines, muscle relaxants, or antipsychotics. 1-4

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## Clarifying a statement about efficacy of FGA, SGA

We appreciate Dr. Nasrallah referencing our work to assert that "most SGAs are similar to FGAs." (CURRENT PSYCHIATRY, Comments & Controversies, October 2013, p. 39-40; http://bit.ly/177QOy6). What we actually found in our 2003¹ and 2011² meta-analyses was that "some antipsychotics are more efficacious than others," and the first-generation antipsychotic vs second-generation antipsychotic distinction is not very useful clinically.³ These findings have repeatedly been replicated.³

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## CORRECTION

In Table 2 of "Expanding medication options for pediatric ADHD" (CURRENT PSYCHIATRY, December 2013, pp. 20-29), "Metadate CD" should have been "Metadate CR."