Neuroscience-based Nomenclature: improving clinical and scientific terminology in research and clinical psychopharmacology

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In 2008, the Neuroscience-based Nomenclature (NbN) was initiated under the leadership of the European College of Neuropsychopharmacology (ECNP), involving representatives from major bodies in the field of psychopharmacology and pharmacology: the American College of Neuropsychopharmacology (ACNP), the Asian College of Neuropsychopharmacology (AsCNP), the International College of Neuropsychopharmacology (CINP) and the International Union of Basic and Clinical Pharmacology (IUPHARM).

Psychiatrists typically prescribe ‘antidepressants’ and ‘neuroleptic’ or ‘antipsychotic’ drugs. The naming of the drugs is usually defined by the first intended use, and a substance usually keeps its name independently of any changing indications for use (or regulatory decisions). However, for over a decade now, things have changed: selective serotonin reuptake inhibitors (SSRIs) and other drugs initially developed for the treatment of depression are now generally also used as the main pharmacological treatments for anxiety disorders. Drugs such as quetiapine, initially developed for the treatment of psychotic symptoms, are now used in the treatment of depression, bipolar disorder, and, though off-label, for anxiety and other syndromes characterized by restlessness, anxiety, sleeping problems and agitation. Other terms such as typical v. atypical, first-, second- or third-generation antipsychotics or similar terms for certain groups of antidepressants were initially used to describe clinical or pharmacological features of the substances, but have since lost their precision due to the development of similar, but somewhat different drugs. For instance, atypicality which was initially seen as a categorical aspect of so-called antipsychotics [a drug is either typical which means that it acts via D2 dopaminergic receptors and produces extrapyramidal side-effects, or it is atypical, i.e. acts on serotonergic (5-HT2A) receptors and other neurotransmitter systems and has no extrapyramidal effects] is now seen more as a dimension, which is indeed influenced by pharmacological properties such as receptor binding profiles and dissociation rate, but also by dosage and other specific properties, e.g. in the case of partial agonists (Grunder et al. 2009). In addition, psychiatric diagnosis is moving from strict categories such as affective v. psychotic disorders towards a more dimensional characterization of symptoms (e.g. Sahakian et al. 2010; Robbins et al. 2012) and, also supported by genetic overlaps between, for instance, schizophrenia and bipolar disorder (e.g. Moskvina et al. 2009; Purcell et al. 2009). The research domain criteria (RDoC) approach promoted by the National Institute for Mental Health (NIMH) goes even further and aims at characterizing mental disorders purely based on a descriptive matrix of neuroscience-based symptoms (Insel et al. 2010). Thus, a terminology based on an initially intended use for one circumscribed disorder or application is not in line with recent developments in pharmacology and the neuroscience of mental illness and therefore is not timely any more. Furthermore, the old terminology can make it difficult for clinicians to explain to a patient suffering from anxiety why he or she should take an antidepressant drug (‘but I am not depressed, I have anxiety’), or why the depressed patient should take an antipsychotic (‘I am not schizophrenic’). Therefore, patients may be confused and also suffer additional stigma from the use of the old terminology.

To overcome these problems of the current terminology, the NbN aims to bring a neuroscience-based and pharmacologically driven approach: drugs are described by their pharmacological mechanism of action, the neurotransmitter modulated by the drug.
the mode of action (e.g. reuptake inhibition, enzyme inhibition, receptor action, or other modulatory effects).

There are multiple advantages of the NbN for many potential users: The communication between clinician and patient can be improved by not having to explain (and possibly excuse) the weaknesses of the current system and the changed use/application of medications. Research on effects of communication styles on medication adherence in mental disorders has shown that adherence can be enhanced when patients have a good understanding of their condition and the treatment options (e.g. Atreja et al. 2005). Also the communication between basic science and pre-clinical and clinical science will be improved. More and more journals such as the journal of Psychopharmacology and the journal Clinical Psychopharmacology and Neuroscience as well as Biological Psychiatry and its daughter journal Biological Psychiatry: Cognitive Neuroscience and Neuroimaging are all now requiring authors to use the NbN, although in a staged fashion to enable a transition and translation between the two systems (Krystal et al. 2016a, b; Nutt & Blier, 2016, Uchida et al. 2016), and other journals will certainly follow.

One main weakness of the NbN could be that it is still in development such that, for instance, drugs primarily used in child and adolescent psychiatry or in neurology are just being included. One example is the description of the mode of action of modafinil, which in the most recent version of the NbN is described as a dopamine (DA) reuptake inhibitor and wakefulness-promoting agent. However, the pharmacological action of modafinil is more complex, involving also a reuptake inhibiting effect for noradrenaline (Minzenberg & Carter, 2008) as well as modulating GABA, serotonin, histamine and orexin (Scoriels et al. 2013), and besides promoting wakefulness, it also has various effects on memory, emotional and motivational processes. However, the NbN is explicitly designed to be extended, updated and adapted; currently, it is automatically updated twice a year.

Due to its neuroscience base the NbN is not only flexible and adaptable to include novel pharmacological developments, but with increasing knowledge about the neurobiology of mental disorders the factor of matching a drug to a pathophysiological mechanism will become increasingly valuable for both clinicians and scientists. Progressing this vision will require the training of a new cadre of clinician scientists (Bullmore et al. 2009; Lehner & Insel, 2010; Insel et al. 2013), with novel programmes that focus on integrating basic and clinical neuroscience, translational medicine and novel methodologies. To achieve this, the NIH Neuroscience Blueprint has announced grants that target neuroscience education prior to entering university and emphasize neuroscience training at an early stage.

In conclusion, the NbN is a science-based approach to describing neuropsychopharmacological agents. It represents a major step that will improve scientific communication as well as the communication between clinicians and patients. We recommend that scientific publications should use the NbN where practical (in an intermediary period parallel to the ‘old’ terminology). In clinical situations, psychiatrists could begin using the mechanisms of action of drugs in addition to the ‘usual’ terms. This can also assist when explaining neurobiological models of disorders to patients, thereby enhancing understanding and adherence.


Declaration of Interest

B.J.S. is a member of the ACNP Nomenclature Workgroup.

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