

Obsessive compulsive and related disorders: comparing DSM-5 and ICD-11

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Obsessive-compulsive disorder (OCD) has been recognized as mainly characterized by compulsivity rather than anxiety and, therefore, was removed from the anxiety disorders chapter and given its own in both the American Psychiatric Association (APA) *Diagnostic and Statistical Manual of Mental Disorders* (DSM-5) and the Beta Draft Version of the 11th revision of the World Health Organization (WHO) *International Classification of Diseases* (ICD-11). This revised clustering is based on increasing evidence of common affected neurocircuits between disorders, differently from previous classification systems based on interrater agreement. In this article, we focus on the classification of obsessive-compulsive and related disorders (OCRDs), examining the differences in approach adopted by these 2 nosological systems, with particular attention to the proposed changes in the forthcoming ICD-11. At this stage, notable differences in the ICD classification are emerging from the previous revision, apparently converging toward a reformulation of OCRDs that is closer to the DSM-5.

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Introduction

Obsessive-compulsive disorder (OCD) is a chronic and disabling neuropsychiatric disorder, with a lifetime prevalence of 2%.¹ Despite its prevalence, OCD is poorly recognized, underdiagnosed, and undertreated, resulting in considerable cost and burden to the individual and to the health economy.² Better diagnosis of OCD is a recognized public health priority.³ OCD is currently classified by the World Health Organization (WHO) in the *International Classification of Diseases and Related Health Problems, Tenth Revision* (ICD-10)⁴ and the American Psychiatric Association (APA) in the *Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition* (DSM-5).⁵ However, these 2 major diagnostic systems show marked differences, either in the description of the disorder or in the diagnostic

criteria, specifiers, and differential diagnosis. It is noteworthy that whereas the current version of the ICD-10 was approved in 1990, the DSM-5 was released in 2013, and therefore has benefitted from recent advances in the understanding of the neurobiology of OCD and its nosological relationship with several other disorders characterized by obsessive-compulsive symptomatology.

The removal of OCD from the Anxiety Disorders chapter and the establishment of the Obsessive-Compulsive and Related Disorders (OCRDs) chapter reflects the hypothesis of common external validators other than anxiety within OCRDs,⁶ which represent a group of disorders characterized by compulsivity. The idea of grouping disorders on the basis of compulsive features dates back to Kraepelin's description of "compulsive insanity" (1899),⁷ in which "compulsive ideas and compulsive apprehensions dominate the clinical picture." Although extremely relevant, Kraepelin's conceptualization was based on his clinical intuition, whereas the OCRDs chapter in both the DSM-5 and the ICD-11 Beta Draft is now based on increasing

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evidence of common affected neurocircuits between disorders and reflects the shift of paradigm from the previous clustering rationale, based on interrater agreement. Although the affected neurocircuits and the specific biomarkers for OCRDs are not explicitly mentioned in the DSM-5, this nosological shift is consistent with the increasing evidence of distinct neurobiological profiles separating OCD and anxiety disorders. The neurobiology of OCD has shifted from the anxiety-avoidance paradigm—involving amygdala and prefrontal cortex network dysfunctions as key components of anxiety disorders⁸—to a dysfunction of the orbitofronto-striato-pallido-thalamic circuitry and reward circuitry.^{9,10} This also encompasses the identification of distinct compensatory mechanisms, which may represent candidate endophenotypes (see “Future Directions” section).¹¹

ICD-10 and DSM-5: Differences in the Approach to OCD and Related Disorders

Phenomenology of obsessions and compulsions

Although obsessions and compulsions, similarly defined, represent the core features of OCD in both ICD-10 and DSM-5, there are several differences between the definitions provided in the two systems (see Table 1).

In ICD-10, OCD is characterized by “recurrent obsessional thoughts or compulsive acts.” Obsessions are defined as “ideas, images or impulses,” whereas compulsions are defined as “stereotyped behaviors that are repeated again and again.” These definitions imply a conceptualization of obsessions being uniquely cognitive events, with no mention of the increasingly recognized non-cognitive events called “sensory phenomena,” which may precede compulsions. “Sensory phenomena” is a term that encompasses a variety of subjective experiences, also referred to as “premonitory urges,” “sensory tics,” “just-right perceptions,” “sensory experiences,” “feelings of incompleteness,” and “not just-right phenomena.”¹² Some authors have hypothesized that such phenomena represent specific pathological pathways and therefore contribute to the identification of OCD subgroups that are more specific.¹³ The presence of these experiences may also have therapeutic consequences, since they may represent a treatment response predictor to pharmacological and behavioral therapy.^{14–17} On the other hand, compulsions are conceptualized as being uniquely motor behaviors, with no consideration of mental rituals. Moreover there is no mention of a functional relationship between the 2 (ie, that compulsions may arise as an attempt to reduce anxiety or distress caused by obsessions). Conversely, the DSM-5 refers to obsessions as “recurrent and persistent thoughts, urges, or images” and to compulsions as “repetitive behaviors or mental acts,” underlining in

both definitions a mutual relationship between the 2, namely that that the individual may attempt to neutralize obsessions with some other thought or actions (ie, compulsions) or feels driven to perform compulsions in response to an obsession.

Additional differences concern the definition of “distress” associated with obsessions and compulsions. The ICD-10 states that obsessions and compulsions generate distress and are not pleasurable experiences, without clarifying the associated specific affects, whereas the DSM-5 states that they can be accompanied by a broad range of affective responses, some of which represent substrates for new research, such as panic attacks, strong feelings of disgust, or a sense of “incompleteness.” For instance, the feeling of disgust is a prominent negative affect in OCD, and a growing body of research suggests that abnormal disgust responses may be implicated in the OCD symptomatology, with self-reported disgust positively correlating with the severity of other OCD-spectrum symptoms.¹⁸ The feeling of disgust was also proposed as a prime candidate for a “new” domain of temperament, and it was also proposed to play a role in the pathogenesis and maintenance of OCD.¹⁹ In addition, whereas the ICD-10 requires a minimum duration for obsessions or compulsions, which must be present almost every day for at least 2 weeks, the DSM-5 does not focus on a specific duration requirement but rather requires symptoms to be time-consuming. This broader conceptualization of OCD can be expected to improve the diagnostic sensitivity of the instrument and capture more cases that might otherwise have been missed.

Specifiers and differential diagnosis

There are also notable differences between ICD-10 and DSM-5 in the use of specifiers and differential diagnosis. The ICD-10 specifiers pertain to the diagnostic picture of OCD, with 3 main diagnostic presentations: predominantly obsessional thoughts, predominantly compulsive acts, mixed obsessional thoughts and acts. No specifiers are provided in regard to the degree of insight, which is implicitly considered as fair/good in the definition of symptoms (“they must be recognized as the individual’s own thoughts or impulses”). The validity and utility of dividing OCD according to these specifiers has been questioned, and it is thought they will change under the ICD-11: according to the online ICD-11 Beta Draft, a new “degree of insight” specifier may be added (see the following paragraphs). Insight can be notably poor or absent in approximately 20% of OCD cases, and in the presence of poor insight, the diagnosis may be missed or confused with delusional disorders. Interestingly, poor insight cases of OCD do not require preferential treatment with antipsychotics and appear to respond

TABLE 1. Summary of DSM-5,⁵ ICD-10,⁴ and ICD-11 (draft)⁴⁰ approach to OCD

	DSM-5	ICD-10	ICD-11 (draft)
Phenomenology (Obsessions (O); Compulsions (C))	O and C defined separately; functional relationship between O and C; C can be mental acts and not only behaviors.	Shared definition of O and C; O are only thoughts and C are only behaviors; no functional relationship between O and C.	O and C defined separately; functional relationship between O and C; C can also be mental acts.
Distress	Time-consuming or clinically significant distress or functional impairment.	Distress or interference with activities.	Time-consuming or significant distress or significant functional impairment.
Duration of symptoms	No criteria.	Most days for equal or more than 2 weeks.	O and C must be time consuming (eg, taking more than 1 hour per day).
Insight	Good/fair or absent/delusional.	They must be recognized as the individual's own thoughts or impulses.	Fair to good; poor; no insight.
Differential diagnosis	OCD can be diagnosed in presence of depressive disorders, Tourette syndrome, schizophrenia, and in presence of delusional OCD beliefs.	OCD can be diagnosed with depressive disorders according to timing rules; OCD cannot be diagnosed in presence of schizophrenia or Tourette syndrome.	N.a.
Specifiers	Degree of insight; presence/absence of tics.	Predominantly O; predominantly C; mixed O and C.	Degree of insight.

just as well as insightful cases to first line treatment with selective serotonin reuptake inhibitors (SSRIs).²⁰ On the other hand, patients with poor insight may be more difficult to engage in cognitive behavioral therapies involving exposure and response prevention.²¹ Therefore, including a specifier that validates poor insight as a recognized “subgroup” could be expected to improve the detection and treatment of OCD and draws attention to poor insight as a cardinal feature of the disorder.

Conversely, DSM-5 has 2 specifiers: the degree of insight (ranging from absent or delusional to good or fair) and the presence or absence of tics. This latter specifier also reflects the distinct “differential diagnosis” approach of DSM-5 compared to ICD-10. The ICD-10 takes a “hierarchical” approach to taxonomy and rules out the diagnosis of OCD in the presence of Tourette syndrome, as a “higher order” disorder, though high rates of comorbidity between these disorders are increasingly recognized.²² Moreover, the ICD-10 does not permit an OCD diagnosis in schizophrenia, whereas it suggests careful examination in the presence of depressive disorders: in these latter disorders, OCD can be diagnosed according to onset and persistence. The diagnosis of OCD is only allowed in acute depressive disorders if the OCD occurred first, and in chronic depressive disorders if the OCD symptoms persist for long periods in the absence of depression. Nevertheless, occasional panic attacks or mild phobic symptoms are no bar to the diagnosis. This approach may be considered unduly restrictive, as it hampers the diagnosis and treatment of OCD in the presence of major psychiatric disorders with which OCD is known to share considerable comorbidity, such as affective disorders (approximately 60%)²³ schizophrenia (approximately 25%).²⁴ Indeed, the relationship between obsessive-compulsive symptomatology and psychosis is long-known, and the

prevalence of OC symptoms in schizophrenia and ultra-high risk (UHR) populations is higher than in the general population,²⁵ so that OCD has also been hypothesized as a prodrome for schizophrenia.²⁶

In contrast, DSM-5 permits an OCD diagnosis in the presence of Tourette syndrome, schizophrenia, depressive disorder, and delusional beliefs, suggesting a differential diagnosis with a broader range of disorders such as anxiety and depressive disorders, other OCDs, OC personality disorder, and eating disorders. This extended and non-hierarchical approach to OCD differential diagnosis in the DSM-5 is not without consequences at a clinical level. Though the DSM-5 states OCD can be distinguished from the ruminations of major depressive disorder (which are usually mood-congruent and not necessarily experienced as intrusive or distressing), sometimes in clinical practice this differentiation may not be so obvious. Thus, there may be a bias toward over-diagnosis of comorbid OCD in ambiguous cases of depression. From a clinical perspective, obsessions that present uniquely in depressive phases may be suggestive of a bipolar disorder, rather than OCD, as well as an episodic course of OCD;^{27,28} nevertheless the differentiation between ruminative thoughts and obsessions is not always clear, since they both are repetitive cognitive intrusions accompanied by negative emotions, are difficult to dismiss, and are subjectively experienced as loss of mental control.²⁹ These issues may also apply to the depressive phase of bipolar disorder with important clinical implications, since some treatments for OCD (eg, ultra-high dose SSRI) can potentially worsen bipolar symptomatology.³⁰ Moreover, the clear identification of comorbidities between OCD and bipolar disorder may impact the treatment algorithm, favoring the administration of antipsychotics or mood stabilizers.^{30,31}

On the other hand, allowing the diagnosis of OCD in schizophrenia, given the recognition of the high comorbidity rates, has opened the debate on the still controversial “schizo-obsessive” subtype^{32–34} and on tentative therapeutic approaches for this highly disabled subgroup of patients.

Obsessive compulsive and related disorders

Arguably, designation by the DSM-5 of a new family of OCRDs, on the basis of age of onset, comorbidities, neurobiological factors, and treatment response, constitutes one of the major modern day advances in taxonomy and will generate considerable and much-needed new interest in the evaluation and treatment of disorders characterized by compulsivity.

A recent review³⁵ underlines the clinical significance of grouping OCD with disorders such as body dysmorphic disorder (BDD), trichotillomania (TTM; hair-pulling disorder), excoriation (skin-picking) disorder, and hoarding disorder, and highlights their similarities across a range of validators. Other authors have gone further and have suggested dividing the OCRDs into 2 subgroups, one comprising “cognitive OCRDs” (ie, OCD, BDD, hoarding disorder) and the second comprising “body-focused repetitive behavioral disorders” (ie, TTM and skin-picking disorder).^{36,37} This proposal gains some support from the results of a large twin study examining the structure of genetic and environmental risk factors of OCRDs.³⁸ Two latent liability factors for OCRDs were found that were largely under genetic control. The first latent factor was common across all OCRDs and was therefore conceptualized as a nonspecific genetic vulnerability, possibly explaining the phenomenologic similarities, patterns of comorbidity, and familiarity described in the OCD literature. The second factor, which was also strongly genetically influenced, loaded exclusively on TTM and skin-picking disorder, whereas environmental factors were shown to be largely disorder-specific.³⁸ In contrast, other contributions³⁹ have questioned the empirical validity and practical utility of the new OCRDs DSM-5 chapter. The authors critically review the grouping of these disorders on the basis of relatively limited data and suggest that the OCD concepts are based on the superficial form of symptoms and lacks scientific merit.³⁹ Notwithstanding, the ICD-11 seems to be moving toward the approach of an OCRDs classification, tracing the DSM-5 model (see the next paragraph).

OCD and OCRDs in the Future ICD-11: Diagnosis and Classification

Recently, the Beta Draft of the ICD-11 (not final, not approved by the WHO and updated daily) has been

published online.^{40,41} Even though the diagnostic criteria are not yet available, the general description of the disorder seems to reflect the acceptance of already published recommendations for the revised classification of OCD.⁴²

By definition, OCD in the draft ICD-11 is

... characterized by the presence of obsessions or compulsions, or both. Obsessions are repetitive and persistent thoughts, images, or urges (impulses) that are intrusive, unwanted, and commonly associated with anxiety. The individual attempts to ignore or suppress obsessions or neutralize them by performing compulsions. Compulsions are repetitive behaviors or mental acts that the individual feels driven to perform in response to an obsession, according to rigid rules, or to achieve a sense of “completeness.” In order for obsessive-compulsive disorder to be diagnosed, obsessions and compulsions must be time consuming (eg, taking more than 1 hour per day), and result in significant distress or significant impairment in personal, family, social, educational, occupational or other important areas of functioning.

This reformulated definition, although not final, seems to accept many of the recommendations recently proposed⁴² for the revision of the diagnostic guidelines, differential diagnosis, and specifiers of OCD. First, the draft ICD-11 has adopted the word “urge” in the definition of obsessions, which is consistent with the DSM-5 definition and is intended to reduce confusion with the impulse control disorders. However, in recognition of the growing neuropsychological evidence suggesting that impulsivity contributes to the generation of obsessive-compulsive behavior, the term “impulse” has not been completely replaced, maintaining an option for clinicians to use both words, and probably facilitating the translation to languages other than English. In addition, whereas the ICD definition still mentions the common presence of anxiety, the statement from the previous ICD-10, in which “anxiety is almost invariably present,” has been reformulated. This is consistent with the removal of OCD from the anxiety disorder section in the DSM, and seems to reflect the recognition of anxiety symptoms as variable and heterogeneous in OCD and, therefore, as a less stable and reliable key indicator of OCD.

Also, in accordance with the DSM, in the draft ICD-11 a functional relationship between obsessions and compulsions has been made explicit. Regarding obsessions, the recommendation that they may be either behaviors or mental acts has been accepted, as well as the removal of the term “stereotyped”, which might be potentially confusing with stereotypies or stereotypic movement

disorder.⁴² Addition of the “sense of completeness” is also noteworthy, due to the growing evidence that supports the key role and prevalence of the so-called “sensory phenomena” or “not-just-right-experiences” in OCD patients.^{12,43} Last, 3 other changes are to be underlined: the modification of specifiers, which have been partly aligned to those in DSM-5, namely including the degree of insight, ranging from absent insight or delusional beliefs, poor insight, to good or fair insight (the presence of tics has not so far been included); further clarification of the functional consequences of the disorder; and the duration requirement (namely that obsessions and compulsions must be time consuming (eg, take more than 1 hour per day). Up to now, the rules for differential diagnosis are still to be elucidated.

Another key change from the ICD-10 to the ICD-11 Beta Draft involves the move to classify OCD within a broad cluster of Obsessive Compulsive and Related Disorders that is similar to, but extends beyond, that of the DSM-5. OCRDs in ICD-11 Beta Draft are defined as “a group of disorders characterized by repetitive thoughts and behaviours that are believed to share similarities in etiology, genetic determinants, and affected neurocircuits or are commonly co-occurring.” This section currently includes obsessive-compulsive disorder, body dysmorphic disorder (BDD), olfactory reference disorder, hypochondriasis, hoarding disorder, and body-focused repetitive behavior disorders (excoriation disorder and trichotillomania). Up to now, it also comprises other specified/unspecified obsessive-compulsive and related disorders, with no detailed description or definition.

The draft ICD-11 classification also proposes that this group of OCRDs may be divided into subsets, based largely on phenomenology and existing (limited) evidence of shared treatment-response profiles.⁴⁴ For example, one subset may include disorders for which cognitive phenomena, such as obsessions, intrusive thoughts, and preoccupations, are central and in response individuals engage in excessive related repetitive behaviors (ie, obsessive-compulsive disorder, body dysmorphic disorder, hypochondriasis, and olfactory reference disorder). Another may include OCRDs that are primarily characterized by body-focused repetitive behaviors and involve recurrent and habitual behaviors directed at the integument and lack a prominent cognitive aspect (eg, hair-pulling, skin-picking).

New ICD-11 Disorders

In the ICD-11, the OCRDs will also encompass 2 new ICD disorders: hoarding disorder^{40,45} and olfactory reference disorder.^{40,46} Hoarding disorder is characterized by excessive accumulation of and attachment to possessions regardless of their actual value, resulting in

cluttered living spaces, the use and safety of which are compromised. Excessive acquisition, characterized by repetitive urges or behaviors related to buying, stealing, or amassing items, including those that are free, is considered integral to the diagnosis, as is difficulty discarding, due to a perceived need to save items and distress associated with discarding them. In recognition of the poor levels of insight commonly found in cases of hoarding disorder, the degree of insight is included as a specifier. The proposed specifier of “severe domestic squalor”⁴⁵ has not so far been included. Thus, the draft ICD-11 definition is similar to that in the DSM-5, although in the DSM-5 “excessive acquisition” is not essential and is included at the level of specifier.

Olfactory reference disorder is a chronically disabling disorder characterized by persistent preoccupation with emitting a perceived foul or offensive body odor that is either unnoticeable or only slightly noticeable to others and is more common in certain cultures, eg, Asia, Africa. Though the exact prevalence rates are still unknown, owing to a lack of epidemiological studies, a total of 84 case reports worldwide was estimated,⁴⁷ with most reports consisting of case reports or small case series. The largest series are from Japan (N = 38), Canada (N = 36), Nigeria (N = 32), Saudi Arabia (N = 15), and Brazil (n = 14).⁴⁸ In contrast, in the DSM-5, this condition (named “olfactory reference syndrome,” ORS) is not listed as an independent diagnosis, but rather falls in the “Other Specified OCRDs” section and in the “Glossary of Cultural Concepts of Distress” as a restricted variant of the Japanese *Taijin kyofusho* (“interpersonal fear disorder”).

Clustering OCRDs: From DSM-5 to ICD-11

The ICD-11 has refined and extended the DSM-5 approach to OCRDs clustering, with an arguably stronger focus on the growing evidence supporting a coherent underpinning neurobiology. Both systems recognize compulsivity, rather than anxiety, as the main core dimension of the OCRDs. However, whereas the “etiological similarities” are not defined in the DSM-5, which refers to “a range of diagnostic validators,” the ICD-11 explicitly underlines genetic and neurobiological similarities as a basis for linking the disorders.

Both the DSM-5 and ICD-11 include five disorders in the OCRD cluster (OCD, body dysmorphic disorder, hoarding disorder, excoriation disorder, and trichotillomania), which may represent the core disorders of the section. However, the ICD-11 additionally includes olfactory reference syndrome and hypochondriasis as OCRDs, whereas the DSM-5 includes instead “substance/medication induced OCRDs” and “OCRDs due to another medical condition.” The inclusion of hypochondriasis in the ICD-11 OCRDs is probably due to

an emphasis on mental “preoccupation” in its description and to the presence of excessive, repetitive checking and healthcare-seeking behaviors. Hypochondriasis has historically been included in the “anxiety and fear-related disorders” cluster, based on the phenomenological overlap with anxiety disorders including symptoms of hypervigilance toward bodily symptoms and fear-related avoidance. At this preliminary juncture (October 2015), the draft ICD-11 lists hypochondriasis in the OCRDs and the disorder has been removed from the anxiety disorders grouping. This is different from the DSM-5, which still includes “illness anxiety disorder (hypochondriasis)” in the anxiety disorders section. It is to be expected that by moving hypochondriasis into the OCRDs section, the ICD-11 would generate potentially fruitful new research perspectives that may advance understanding of the underpinning neuropsychological mechanisms and encourage new treatment development. On the other hand, the DSM-5 has broadened the OCRDs cluster to include “substance/medication induced OCRDs” and “OCRDs due to another medical condition.” These diagnoses are not considered specific mental disorders in a narrow sense, but are recognized as generic conditions that may be encountered by a mental health clinician. Their inclusion as diagnoses in the DSM-5 OCRDs is intended to raise awareness of the “non-psychiatric” origins of some obsessive-compulsive syndromes, and improve differential diagnosis and treatment.

Obsessive-Compulsive Personality Traits

A major revision to the personality disorder section of the ICD classification has been proposed,⁴⁹ based on the growing evidence that supports a dimensional approach toward personality pathology⁵⁰ as opposed to personality disorder categories.⁵¹ These dimensions of personality pathology would include “anankastic features,” such as perfectionism, extreme need to control their own and others’ behavior, and rigid adherence to rules consistent with the ICD-10 diagnosis of anankastic (obsessive-compulsive) personality disorder.⁴⁹

The ICD-11 plans to include a system of “multiple parenting,” which allows certain diagnostic categories that could legitimately be placed in more than 1 section of the classification to be cross-referenced. Multiple parenting is expected to enhance clinical utility by ensuring increased recognition of cross-referenced disorders in differential diagnosis or as comorbidities to improve treatment-planning.⁵² Anankastic personality disorder is highly comorbid with OCD and shares a significant familial link,^{53,54} as well as a similar neuropsychological profile, reflecting cognitive inflexibility and perseveration that corresponds to that of individuals with OCD.⁵⁵ The ICD-11 workgroup on OCD

proposes that the anankastic variant of personality disorders is cross-referenced in the OCRD grouping,⁵² based on significant similarities in phenomenology (although important differences also exist such as a lack of obsessions and compulsions), as well as major comorbidity with a range of obsessive-compulsive disorders.^{56,57}

Future Directions

It is clear that while significant progress is being made in classifying OCD and related disorders, with clear evidence of a convergence in the conceptualization of these disorders emerging from the 2 major global classificatory systems, a great deal of uncertainty remains. While the aspiration of “carving nature at its joints” may not be readily achievable, there are grounds to believe a better understanding of the neurobiological underpinnings will inform a more rational classification and further refine the clinical definition of individual disorders and their taxonomy.

In order to overcome the limitations of the current diagnostic systems and to address the need for a new approach to classifying mental disorders, based on dimensions of observable behavior and neurobiological measures,⁵⁸ the National Institute of Mental Health (NIMH) has recently launched the Research Domain Criteria (RDoC) project. This aims to create a framework that integrates the most recent contributions in neuroscience and genomics, with the ultimate goal of “precision medicine,” namely a diagnostic refinement based on a deeper understanding of the circuitries and networks of psychiatric disorders considered to be brain diseases.⁵⁹ It is to be expected that future iterations of the DSM and ICD systems will align with the RDoC approach, with the aim of bringing neuroscience and clinical practice closer together (see Table 2).

Endophenotypes represent a still debated concept, yet they are a hot topic in neuropsychiatric research. They can be conceptualized as a special kind of biomarker, encompassing heritable neurobiological and neurobehavioral characteristics that play an important role for bridging the gap between the microscopic level (eg, molecular genetics) and the macroscopic level (eg, clinical symptoms) in neuropsychiatric disorders.^{60,61} Neither of the existing nosological systems have so far included an adequate integration of the current knowledge about neural circuits, neurotransmitters, and behavior, as proposed by the NIMH’s RDoC project.^{58,62} However, in the DSM-5, there has been an attempt to facilitate the identification and fine-tuning of psychiatric endophenotypes, through the introduction of a “Cross-Cutting Symptom Measures” chapter. This chapter integrates a “dimensional assessment” with a “categorical diagnosis” approach. It aims at addressing

TABLE 2. Summary of relevant topics to be considered

Topic	Relevance	DSM-5	ICD-11 (draft)
Endophenotypes	Deeper understanding of pathophysiologic processes; differential treatment response.	No endophenotypes evaluation or description.	No endophenotypes evaluation or description.
Genetic and immunologic characteristics	Integrative view of OCD pathophysiology; differentiation with autoimmune syndromes (eg, PANS, PANDAS).	Brief description of genetic characteristics in the "Risk and Prognostic Factors" paragraph for some OCRDs; no differential diagnosis with PANS or PANDAS.	No description of genetic characteristics for OCRDs; no differential diagnosis with PANS or PANDAS.
Neuroprogression and clinical staging	Inform therapeutics of OCD; develop sequential treatments; identify disorder trajectories.	Brief description of OCRDs course with no mention to neuroprogression or clinical staging models.	No mention to disorders' course or neuroprogression or clinical staging models.
Comorbidities subgroups	Develop specific treatments based on comorbidities for complex clinical pictures.	Description and prevalence rates of comorbidities; no subgrouping or clinical implications based on comorbidities.	No description of comorbidities.
Dimensional measures	Assessment of co-occurring symptoms; identification and fine-tuning of psychiatric endophenotypes.	"Cross-cutting Symptom Measures" section to identify additional areas of inquiry significant for prognosis or treatment.	No additional dimensional measures.
Gender differences	Understanding differential clinical expression phenomena in males/females; facilitate evaluation of postpartum OCD.	Brief description of gender differences in the "Gender-Related Diagnostic Issues" paragraph for some OCRDs; brief mention to "peripartum OCD."	No specific mention to gender differences.

the issue of co-occurring symptoms across mental disorders⁶³ as an adjunct tool "to give clinicians quantitative ratings that characterize patients in a way that is simple, useful, and clinically meaningful."⁶⁴ From this perspective, the identification of specific endophenotypes, which may cross traditional categorical diagnostic boundaries, would be expected to help in both the diagnostic and therapeutic process, by providing a deeper understanding of the pathophysiological processes specific to a disorder and the differential response to specific treatments.⁶⁵ For example, a better understanding of the cognitive dimension of OCRDs, in terms of overlapping attentional and planning deficits, may be particularly helpful for clarifying the strength of relationship between different OCRDs and devising new treatment strategies that target cognitive deficits.⁶⁶

Genetic studies in OCD are relatively scarce in comparison to those in schizophrenia and affective disorders.⁶⁷ In addition, the immunologic characteristics of OCRDs do not currently receive adequate attention. Such findings may contribute to the development of an integrative view of OCD pathophysiology, as for example the "glutamate-based genetic immune hypothesis."⁶⁸

The classification of issues pertaining to the pediatric acute-onset neuropsychiatric syndrome (PANS) highlights some of these deficiencies. PANS refers to a syndrome characterized by an abrupt, dramatic onset of OCD (meeting DSM-IV criteria) or severely restricted food intake, with the concurrent presence of additional neuropsychiatric symptoms, with similarly severe and acute onset, triggered by any infectious agent (not only streptococcal infection) in addition to non-infectious triggers, which are yet to be fully determined.⁶⁹ This syndrome has been proposed as an expanded clinical

entity compared to the previous pediatric autoimmune neuropsychiatric disorders associated with streptococcal infections (PANDAS).⁶⁹ Though recognized by the National Institute of Health (NIH), PANS and PANDAS are not yet validated nosological constructs⁷⁰ and are not listed as a diagnosis by the ICD-10 or the DSM-5. From this perspective, a useful future specifier for OCD would be the mode of onset (acute versus progressive), which may help differentiate autoimmune clinical pictures.

A relevant issue that pertains to both diagnostic systems is the concept of staging and "neuroprogression." Neither the DSM-5 nor the draft ICD-11 have incorporated the long-debated concept of clinical staging, which may differentiate early, milder clinical phenomena from those that accompany illness progression and chronicity. This is already a reality in several fields of medicine, and is based on the assumption that early intervention can produce better clinical and functional outcomes. Nevertheless, it is a currently neglected dimension in most psychiatric disorders;⁷¹ the concept of clinical staging is emerging in the research on schizophrenia,⁷² together with the aim of an identification of the trajectory of the disorder and the investigation of the after effects of the duration of untreated disorder, though this still seems to be a backward approach in the OCD research.

In sum, what is currently missing in both diagnostic systems is the integration of a "clinimetric" perspective,^{73,74} a concept referring to a number of clinical issues that do not find room in the current taxonomy, comprising clinical staging and severity. Such integration would also inform therapeutics of OCD and may help developing sequential treatment strategies, which are, up to now, scarcely investigated.^{75,76} Data from the

most recent meta-analyses and practice guidelines⁷⁷ indicate that serotonin reuptake inhibitors (SRIs) still represent the first-line pharmacological treatment (after a trial with cognitive behavioral therapy [CBT]), yet a large percentage of patients still does not achieve a symptom remission, particularly when the clinical picture includes the presence of comorbidities. Therefore, a diagnostic system allowing the identification of the clinimetric properties would facilitate research and the development of specific treatments strategies in complex clinical pictures and comorbid conditions.⁷⁸

Last, gender evaluation is a relevant factor that should be taken in account in OCD, since gender differences have been shown in the clinical expression of OCD phenomenon.⁷⁹ This is also relevant in regard to postpartum psychiatric disorders; whereas a good deal of attention has been paid to postpartum depression and psychosis, less effort has been devoted to studying OCD with a perinatal onset.

Conclusions

The classification of OCD underwent a noticeable change in the process of revision from DSM-IV to DSM-5. Although the diagnostic criteria for the disorder remained almost the same, the clustering in a new, dedicated chapter of OCRDs reflects the results of 2 decades of intensive study in the field of OCD and the recognition of close relationships and overlaps between this group of disorders.

The DSM-5 has paved the way for the forthcoming revision of the existing ICD-10, which has become outdated. The definitive ICD-11 is expected to be released 2018. A draft version in the latest stage of revision has been made available online and demonstrates prominent changes. Indeed, despite differences in the origins, purpose, and scope of the 2 classification systems,⁸⁰ the process of harmonization between them seems to be crystallizing in the field of OCRDs. These modifications are consistent with the aim of a “scientifically valid” classification system, since the proposed changes are grounded upon recent findings and a thorough review of the latest contribution in the literature. While more remains to be achieved, this represents an important step forward in the development of a “globally applicable” classification system, as it bridges the gap between the 2 nosological systems and facilitates the process of diagnosis worldwide and in both clinical and research contexts.

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REFERENCES:

- Ruscio AM, Stein DJ, Chiu WT, Kessler RC. The epidemiology of obsessive-compulsive disorder in the National Comorbidity Survey Replication. *Mol Psychiatry*. 2010; **15**(1): 53–63.
- Hollander E, Stein D, Fineberg NA, Legault M. Quality of life outcomes in patients with obsessive-compulsive disorder: relationship to treatment response and symptom relapse. *J Clin Psychiatry*. 2010; **71**(6): 784–792.
- Fineberg NA, Baldwin DS, Menchon JM, et al. The Obsessive Compulsive and Related Disorders Research Network Manifesto for a European research network into obsessive-compulsive and related disorders. *Eur Neuropsychopharmacol*. 2013; **23**(7): 561–568.
- World Health Organization. *International Statistical Classification of Diseases and Related Health Problems*. 10th rev. Geneva: WHO; 1992.
- American Psychiatric Association. *Diagnostic and Statistical Manual of Mental Disorders*. 5th ed. Arlington, VA: American Psychiatric Publishing; 2013.
- Stein DJ, Fineberg NA, Bienvenu OJ, et al. Should OCD be classified as an anxiety disorder in DSM-V? *Depress Anxiety*. 2010; **27**(6): 495–506.
- Kraepelin E. *Psychiatry: A Textbook for Students and Physicians*. Canton, MA: Science History Publications [Original work published 1899].
- Shin LM, Liberzon I. The neurocircuitry of fear, stress, and anxiety disorders. *Neuropsychopharmacology*. 2010; **35**(1): 169–191.
- Menzies L, Chamberlain SR, Laird AR, Thelen SM, Sahakian BJ, Bullmore ET. Integrating evidence from neuroimaging and neuropsychological studies of obsessive-compulsive disorder: the orbitofronto-striatal model revisited. *Neurosci Biobehav Rev*. 2008; **32**(3): 525–549.
- Figeo M, Vink M, de Geus F, et al. Dysfunctional reward circuitry in obsessive-compulsive disorder. *Biol Psychiatry*. 2011; **69**(9): 867–874.
- de Vries FE, de Wit SJ, Cath DC, et al. Compensatory frontoparietal activity during working memory: an endophenotype of obsessive-compulsive disorder. *Biol Psychiatry*. 2014; **76**(11): 878–887.
- Ferrão YA, Shavitt RG, Prado H, et al. Sensory phenomena associated with repetitive behaviors in obsessive-compulsive disorder: an exploratory study of 1001 patients. *Psychiatry Res*. 2012; **197**(3): 253–258.
- Miguel EC, Baer L, Coffey BJ, et al. Phenomenological differences appearing with repetitive behaviors in obsessive-compulsive disorder and Gilles de la Tourette’s syndrome. *Br J Psychiatry*. 1997; **170**(2): 140–145.
- Miguel EC, do Rosário-Campos MC, Prado HDS, et al. Sensory phenomena in obsessive-compulsive disorder and Tourette’s disorder. *J Clin Psychiatry*. 2000; **61**(2): 150–156.
- Summerfeldt LJ. Understanding and treating incompleteness in obsessive-compulsive disorder. *J Clin Psychology*. 2004; **60**(11): 1155–1168.
- Shavitt RG, Belotto C, Curi M, et al. Clinical features associated with treatment response in obsessive-compulsive disorder. *Compr Psychiatry*. 2006; **47**(4): 276–281.
- Katerberg H, Cath DC, Denys DA, et al. The role of the COMT Val (158)Met polymorphism in the phenotypic expression of obsessive-compulsive disorder. *Am J Med Genet B Neuropsychiatr Gen*. 2010; **153B**(1): 167–176.

18. Whitton AE, Henry JD, Grisham JR. Cognitive and psychophysiological correlates of disgust in obsessive-compulsive disorder. *Br J Clin Psychol*. 2015; **54**(1): 16–33.
19. Olatunji BO, Tart CD, Ciesielski BG, McGrath PB, Smits JA. Specificity of disgust vulnerability in the distinction and treatment of OCD. *J Psychiatr Res*. 2011; **45**(9): 1236–1242.
20. Eisen JL, Rasmussen SA, Phillips KA, et al. Insight and treatment outcome in obsessive-compulsive disorder. *Compr Psychiatry*. 2001; **42**(6): 494–497.
21. Visser HA, van Megen H, van Oppen P, et al. Inference-based approach versus cognitive behavioral therapy in the treatment of obsessive-compulsive disorder with poor insight: a 24-session randomized controlled trial. *Psychother Psychosom*. 2015; **84**(5): 284–293.
22. Nordstrom EJ, Burton FH. A transgenic model of comorbid Tourette's syndrome and obsessive-compulsive disorder circuitry. *Mol Psychiatry*. 2002; **7**(6): 617–625; 524.
23. Fineberg NA, Hengartner MP, Bergbaum C, Gale T, Rössler W, Angst J. Lifetime comorbidity of obsessive-compulsive disorder and sub-threshold obsessive-compulsive symptomatology in the community: impact, prevalence, socio-demographic and clinical characteristics. *Int J Psychiatry Clin Pract*. 2013; **17**(3): 188–196.
24. Mukhopadhaya K, Krishnaiah R, Taye T, et al. Obsessive-compulsive disorder in UK clozapine-treated schizophrenia and schizoaffective disorder: a cause for clinical concern. *J Psychopharmacol*. 2009; **23**(1): 6–13.
25. Byerly M, Goodman W, Acholonu W, Bugno R, Rush AJ. Obsessive compulsive symptoms in schizophrenia: frequency and clinical features. *Schizophr Res*. 2005; **76**(2–3): 309–316.
26. Bottas A, Cooke RG, Richter MA. Comorbidity and pathophysiology of obsessive-compulsive disorder in schizophrenia: is there evidence for a schizo-obsessive subtype of schizophrenia? *J Psychiatry Neurosci*. 2005; **30**(3): 187–193.
27. Swartz CM, Shen WW. Is episodic obsessive compulsive disorder bipolar? A report of four cases. *J Affect Disord*. 1999; **56**(1): 61–66.
28. Zutshi A, Kamath P, Reddy YC. Bipolar and nonbipolar obsessive-compulsive disorder: a clinical exploration. *Compr Psychiatry*. 2007; **48**(3): 245–251.
29. Wahl K, Schönfeld S, Hissbach J, et al. Differences and similarities between obsessive and ruminative thoughts in obsessive-compulsive and depressed patients: a comparative study. *J Behav Ther Exp Psychiatry*. 2011; **42**(4): 454–461.
30. Amerio A, Odone A, Liapis CC, Ghaemi SN. Diagnostic validity of comorbid bipolar disorder and obsessive-compulsive disorder: a systematic review. *Acta Psychiatr Scand*. 2014; **129**(5): 343–358.
31. Perugi G, Toni C, Frare F, Traverso MC, Hantouche E, Akiskal HS. Obsessive-compulsive-bipolar comorbidity: a systematic exploration of clinical features and treatment outcome. *J Clin Psychiatry*. 2002; **63**(12): 1129–1134.
32. Poyurovsky M, Weizman A, Weizman R. Obsessive-compulsive disorder in schizophrenia: clinical characteristics and treatment. *CNS Drugs*. 2004; **18**(14): 989–1010.
33. Rajkumar RP, Reddy YC, Kandavel T. Clinical profile of “schizo-obsessive” disorder: a comparative study. *Compr Psychiatry*. 2008; **49**(3): 262–268.
34. Varlakova Y, Patel D, Mukhopadhaya K, et al. The neurocognitive and behavioural impact of comorbid obsessive-compulsive syndrome in schizophrenia. In De Haan L, Schirmbeck F, Zink M, eds. *Obsessive-Compulsive Symptoms in Schizophrenia*. Cham, Switzerland: Springer International Publishing; 2015: 91–125.
35. Ameringen M, Patterson B, Simpson W. DSM-5 obsessive-compulsive and related disorders: clinical implication of new criteria. *Depress Anxiety*. 2014; **31**(6): 487–493.
36. Phillips KA, Stein DJ, Rauch SL, et al. Should an obsessive-compulsive spectrum grouping of disorders be included in DSM-V? *Depress Anxiety*. 2010; **27**(6): 528–555.
37. Stein DJ, Grant JE, Franklin ME, et al. Trichotillomania (hair pulling disorder), skin picking disorder, and stereotypic movement disorder: toward DSM-V. *Depress Anxiety*. 2010; **27**(6): 611–626.
38. Monzani B, Rijdsdijk F, Harris J, Mataix-Cols D. The structure of genetic and environmental risk factors for dimensional representations of DSM-5 obsessive-compulsive spectrum disorders. *JAMA Psychiatry*. 2014; **71**(2): 182–189.
39. Abramowitz JS, Jacoby RJ. Obsessive-compulsive and related disorders: a critical review of the new diagnostic class. *Ann Rev Clin Psychol*. 2015; **11**: 165–186.
40. World Health Organization. ICD-11 Beta Draft. <http://apps.who.int/classifications/icd11/browse/l-m/en>. Accessed October 30, 2015.
41. Luciano M. The ICD-11 beta draft is available online. *World Psychiatry*. 2015; **14**(3): 375–376.
42. Simpson HB, Reddy YC. Obsessive-compulsive disorder for ICD-11: proposed changes to the diagnostic guidelines and specifiers. *Rev Bras Psiquiatr*. 2014; **36**(Suppl 1): 3–13.
43. Hellriegel J, Barber C, Wikramanayake M, Fineberg N, Mandy WPL. Is ‘not just right experience’ (njre) in obsessive-compulsive disorder part of an autistic phenotype? Submitted.
44. Grant J, Chamberlain S, Odlaug B. Evidence-based treatment for obsessive-compulsive and related disorders (OCDs). Clinical Guide to OCDs. Oxford, 2014. Please clarify type of entry. Is this a book chapter? If so, please provide name(s) of editors, name/location of publisher, and page numbers for chapter. If a journal article, please clarify journal title and provide volume/page numbers.
45. Fontenelle LF, Grant JE. Hoarding disorder: a new diagnostic category in ICD-11? *Rev Bras Psiquiatr*. 2014; **36**(Suppl 1): 28–39.
46. Veale D, Matsunaga H. Body dysmorphic disorder and olfactory reference disorder: proposals for ICD-11. *Rev Bras Psiquiatr*. 2014; **36**(Suppl 1): 14–20.
47. Begum M, McKenna PJ. Olfactory reference syndrome: a systematic review of the world literature. *Psychol Med*. 2011; **41**(3): 453–461.
48. Phillips KA, Menard W. Olfactory reference syndrome: demographic and clinical features of imagined body odor. *Gen Hosp Psychiatry*. 2011; **33**(4): 398–406.
49. Tyrer P, Reed GM, Crawford MJ. Classification, assessment, prevalence, and effect of personality disorder. *Lancet*. 2015; **385**(9969): 717–726.
50. Bernstein DP, Iscan C, Maser J, Boards of Directors of the Association for Research in Personality Disorders; International Society for the Study of Personality Disorders. Opinions of personality disorder experts regarding the DSM-IV personality disorders classification system. *J Pers Disord*. 2007; **21**(5): 536–551.
51. Skodol AE, Gunderson JG, Shea MT, et al. The Collaborative Longitudinal Personality Disorders Study (CLPS): overview and implications. *J Pers Disord*. 2005; **19**(5): 487–504.
52. Stein DJ, Kogan CS, Atmaca M, et al. The classification of Obsessive-Compulsive and Related Disorders in the ICD-11. *J Affect Disord*. 2016; **190**: 663–674.
53. Bienvenu OJ, Samuels JF, Wuyek LA, et al. Is obsessive-compulsive disorder an anxiety disorder, and what, if any, are spectrum conditions? A family study perspective. *Psychol Med*. 2012; **42**(1): 1–13.
54. Samuels J, Nestadt G, Bienvenu OJ, et al. Personality disorders and normal personality dimensions in obsessive-compulsive disorder. *Br J Psychiatry*. 2000; **177**(5): 457–462.
55. Fineberg NA, Day GA, de Koenigswarter N, et al. The neuropsychology of obsessive-compulsive personality disorder: a new analysis. *CNS Spectr*. 2015; **20**(5): 490–499.
56. Coles ME, Pinto A, Mancebo MC, Rasmussen SA, Eisen JL. OCD with comorbid OCPD: a subtype of OCD? *J Psychiatr Res*. 2008; **42**(4): 289–296.

57. Pinto A, Mancebo MC, Eisen JL, Pagano ME, Rasmussen SA. The Brown Longitudinal Obsessive Compulsive Study: clinical features and symptoms of the sample at intake. *J Clin Psychiatry*. 2006; **67**(5): 703-711.
58. Insel T, Cuthbert B, Garvey M, *et al*. Research domain criteria (RDoC): toward a new classification framework for research on mental disorders. *Am J Psychiatry*. 2010; **167**(7): 748-751.
59. Insel TR. The NIMH research domain criteria (RDoC) project: precision medicine for psychiatry. *Am J Psychiatry*. 2014; **171**(4): 395-397.
60. Gottesman II, Gould TD. The endophenotype concept in psychiatry: etymology and strategic intentions. *Am J Psychiatry*. 2003; **160**(4): 636-645.
61. Insel TR, Cuthbert BN. Endophenotypes: bridging genomic complexity and disorder heterogeneity. *Biol Psychiatry*. 2009; **66**(11): 988-989.
62. Cuthbert BN. The RDoC framework: facilitating transition from ICD/DSM to dimensional approaches that integrate neuroscience and psychopathology. *World Psychiatry*. 2014; **13**(1): 28-35.
63. Clarke DE, Kuhl EA. DSM-5 cross-cutting symptom measures: a step towards the future of psychiatric care? *World Psychiatry*. 2014; **13**(3): 314-316.
64. Berry EA, Heaton PT, Kelton CM. National estimates of the inpatient burden of pediatric bipolar disorder in the United States. *J Ment Health Policy Econ*. 2011; **14**(3): 115-123.
65. Phillips ML. The emerging role of neuroimaging in psychiatry: characterizing treatment-relevant endophenotypes. *Am J Psychiatry*. 2007; **164**(5): 697-699.
66. Fineberg NA, Chamberlain SR, Goudriaan AE, *et al*. New developments in human neurocognition: clinical, genetic, and brain imaging correlates of impulsivity and compulsivity. *CNS Spectr*. 2014; **19**(1): 69-89.
67. Murphy DL, Moya PR, Fox MA, Rubenstein LM, Wendland JR, Timpano KR. Anxiety and affective disorder comorbidity related to serotonin and other neurotransmitter systems: obsessive-compulsive disorder as an example of overlapping clinical and genetic heterogeneity. *Philos Trans R Soc Lond B Biol Sci*. 2013; **368**(1615): 20120435.
68. Rotge JY, Aouizerate B, Tignol J, *et al*. The glutamate-based genetic immune hypothesis in obsessive-compulsive disorder: an integrative approach from genes to symptoms. *Neuroscience*. 2010; **165**(2): 408-417.
69. Swedo SE, Leckman JF, Rose NR. From research subgroup to clinical syndrome: modifying the PANDAS criteria to describe PANS (pediatric acute-onset neuropsychiatric syndrome). *Pediatrics & Therapeutics*. 2012; **2**: 113.
70. Pichichero ME. The PANDAS syndrome. *Adv Exp Med Biol*. 2009; **634**: 205-216.
71. Fava GA, Kellner R. Staging: a neglected dimension in psychiatric classification. *Acta Psychiatr Scand*. 1993; **87**(4): 225-230.
72. McGorry PD, Hickie IB, Yung AR, Pantelis C, Jackson HJ. Clinical staging of psychiatric disorders: a heuristic framework for choosing earlier, safer and more effective interventions. *Aust NZ J Psychiatry*. 2006; **40**(8): 616-622.
73. Feinstein AR. T. Duckett Jones Memorial Lecture: the Jones criteria and the challenges of clinimetrics. *Circulation*. 1982; **66**(1): 1-5.
74. Fava GA, Rafanelli C, Tomba E. The clinical process in psychiatry: a clinimetric approach. *J Clin Psychiatry*. 2012; **73**(2): 177-184.
75. Albert U, Brunatto C. Obsessive-compulsive disorder in adults: Efficacy of combined and sequential treatments. *Clinical Neuropsychiatry*. 2009; **6**(2): 83-93.
76. Albert U, Barbaro F, Aguglia A, Maina G, Bogetto F. [Combined treatments in obsessive-compulsive disorder: current knowledge and future prospects.] *Riv Psichiatr*. 2012; **47**(4): 255-268 [In Italian.]
77. Koran ML. *Practice Guideline for the Treatment of Patients with Obsessive-Compulsive Disorder*. Arlington, VA: American Psychiatric Association; 2007. http://www.psych.org/psych_pract/treatg/pg/prac_guide.cfm.
78. Pallanti S, Grassi G. Pharmacologic treatment of obsessive-compulsive disorder comorbidity. *Exp Opin Pharmacother*. 2014; **15**(17): 2543-2552.
79. Mathis MAD, Alvarenga PD, Funaro G, *et al*. Gender differences in obsessive-compulsive disorder: a literature review. *Rev Bras Psiquiatr*. 2011; **33**(4): 390-399.
80. First MB. Harmonisation of ICD-11 and DSM-V: opportunities and challenges. *Br J Psychiatry*. 2009; **195**(5): 382-390.