Obsessive-Compulsive Disorder: Diagnosis and Management

JILL N. FENSKE, MD, and THOMAS L. SCHWENK, MD, University of Michigan Medical School, Ann Arbor, Michigan

Obsessive-compulsive disorder is an illness that can cause marked distress and disability. It often goes unrecognized and is undertreated. Primary care physicians should be familiar with the various ways obsessive-compulsive disorder can present and should be able to recognize clues to the presence of obsessions or compulsions. Proper diagnosis and

education about the nature of the disorder are important first steps in recovery. Treatment is rarely curative, but patients can have significant improvement in symptoms. Recommended first-line therapy is cognitive behavior therapy with exposure and response prevention or a selective serotonin reuptake inhibitor. The medication doses required for treatment of obsessive-compulsive disorder are often higher than those for other indications, and the length of time to response is typically longer. There are a variety of options for treatment-resistant obsessive-compulsive disorder, including augmentation of a selective serotonin reuptake inhibitor with an atypical antipsychotic. Obsessive-compulsive disorder is a chronic condition with a high rate of relapse. Discontinuation of treatment should be undertaken with caution. Patients should be closely monitored for comorbid depression and suicidal ideation. (Am Fam Physician. 2009;80(3):239-245. Copyright © 2009 American Academy of Family Physicians.)

▶ Patient information: A handout on obsessivecompulsive disorder, written by the authors of this article, is available at http://www.aafp. org/afp/20090801/239s1.html.

This article exempifies the AAFP 2009 Annual Clinical Focus on management of chronic illness.

ЕВ СМЕ

This clinical content conforms to AAFP criteria for evidence-based continuing medical education (EB CME). See CME Quiz on page 225.



The online version of this article includes supple-

mental content at http:// www.aafp.org/afp.

bsessive-compulsive disorder (OCD) is a neuropsychiatric disorder characterized by recurrent distressing thoughts and repetitive behaviors or mental rituals performed to reduce anxiety. Symptoms are often accompanied by feelings of shame and secrecy because patients realize the thoughts and behaviors are excessive or unreasonable. This secrecy, along with a lack of recognition of OCD symptoms by health care professionals, often leads to a long delay in diagnosis and treatment. OCD has a reputation of being difficult to treat, but there are many effective treatments available.

Epidemiology

The lifetime prevalence of OCD is 1.6 percent.1 Symptoms usually begin during adolescence, and more than 50 percent of affected persons have symptom onset before their mid-20s.1 OCD has substantial adverse effects on well-being; more than one half of patients report moderate to severe distress



from obsessions and compulsions.² OCD interferes with work performance, social interactions, and family relationships. It is a chronic disorder and is likely to persist if not treated effectively. Nearly 70 percent of patients report a continuous course of symptoms, and 23 percent experience a waxing and waning course.³ The average time to treatment after meeting diagnostic criteria for OCD is 11 years.³ This delay is attributed to many factors, including reluctance of patients to report symptoms and underrecognition of OCD by physicians.

Pathogenesis

The current model for the pathogenesis of OCD is complex. Neuroimaging studies show involvement of the dorsolateral prefrontal cortex, basal ganglia, and thalamus.⁴ Because of the response to selective serotonin reuptake inhibitors (SSRIs), it is hypothesized that the serotonin system is heavily involved in the neurochemistry of OCD. Family studies have shown that

www.aafp.org/afp

American Family Physician 239

Downloaded from the American Family Physician Web site at www.aafp.org/afp. Copyright © 2009 American Academy of Family Physicians. For the private, noncommercial use of one individual user of the Web site. All other rights reserved. Contact copyrights@aafp.org for copyright questions and/or permission requests.

Clinical recommendation	Evidence rating	References
Patients with OCD should be monitored for psychiatric comorbidities and suicide risk.	С	3, 19, 21
Cognitive behavior therapy with exposure and response prevention is an effective treatment for OCD.	А	21-23
SSRIs are an effective treatment for OCD and are recommended as first-line pharmacologic therapy.	А	21, 24, 25
A trial of SSRI therapy should continue for eight to 12 weeks, with at least four to six weeks at the maximal tolerable dosage.	С	21
SSRIs should be taken for at least one to two years before attempting to discontinue. Exposure and response prevention "booster" sessions should be considered to prevent relapse.	С	21
Augmentation of SSRI therapy with atypical antipsychotic agents is effective in some patients with treatment-resistant OCD.	В	16, 21, 30

OCD = obsessive-compulsive disorder; *SSRI* = selective serotonin reuptake inhibitor.

A = consistent, good-quality patient-oriented evidence; B = inconsistent or limited-quality patient-oriented evidence; C = consensus, diseaseoriented evidence, usual practice, expert opinion, or case series. For information about the SORT evidence rating system, go to http://www.aafp. org/afpsort.xml.

genetics have a role in the etiology of OCD, particularly in the early-onset form of the disorder.⁵ An immunologic component has also been proposed, based on the association of OCD with pediatric autoimmune neuropsychiatric disorder associated with streptococcal infections (PANDAS), in which children develop an abrupt onset of OCD symptoms or tics after infection with group A *Streptococcus*.

Table 1. Common	Obsessions a	and Compulsions
-----------------	--------------	-----------------

Туре	Examples		
Obsessions			
Aggressive impulses	Images of hurting a child or parent		
Contamination	Becoming contaminated by shaking hands with another person		
Need for order	Intense distress when objects are disordered or asymmetric		
Religious	Blasphemous thoughts, concerns about unknowingly sinning		
Repeated doubts	Wondering if a door was left unlocked		
Sexual imagery	Recurrent pornographic images		
Compulsions			
Checking	Repeatedly checking locks, alarms, appliances		
Cleaning	Handwashing		
Hoarding	Saving trash or unnecessary items		
Mental acts	Praying, counting, repeating words silently		
Ordering	Reordering objects to achieve symmetry		
Reassurance- seeking	Asking others for reassurance		
Repetitive actions	Walking in and out of a doorway multiple times		

Diagnosis

Obsessions are recurrent intrusive thoughts or images that cause marked distress. Patients usually recognize that the thoughts are self-generated and inappropriate. Some common obsessions involve contamination, doubts about whether an important task has been performed, or worries that an action will harm another person (*Table 1*). Compulsions are repetitive activities or mental rituals designed to counteract the anxiety caused by obsessions. Common compulsions include handwashing, checking, ordering, praying, counting, and seeking reassurance (*Table 1*). The *Diagnostic and Statistical Manual of Mental Disorders*, 4th ed., states that to meet the criteria for OCD, the obsessions cannot be excessive worries about everyday problems, and they must cause marked distress (*Table 2*).⁶

OCD is a heterogeneous disorder with several subtypes and a multitude of manifestations (*Table 3*).⁷⁻¹⁵ There are several associated disorders (often referred to as OCD

Table 2. Diagnostic Criteria for Obsessive-Compulsive Disorder

Recurrent obsessions or compulsions

- Obsessions and compulsions are severe enough to be time consuming (more than one hour daily) or to cause marked distress or significant impairment
- At some point during the course of the disorder, the person has recognized that the obsessions or compulsions are excessive or unreasonable
- If another axis I disorder is present, the content of the obsessions or compulsions is not restricted to it
- The disturbance is not a result of physiologic effects of a substance or medical condition

Information from reference 6.

spectrum disorders), such as body dysmorphic disorder, trichotillomania, hypochondriasis, and eating disorders. These disorders have similar features and respond to the same therapies used to treat OCD.

Patients are often reluctant to report symptoms of OCD, which they may find embarrassing. Physicians should maintain a high awareness for the possibility of OCD in patients with general complaints of anxiety or depression. Patients may offer clues by alluding to intrusive thoughts or repetitive behaviors. Avoidance of particular locations or objects, excessive concerns about illness or injury, and repetitive reassurance-seeking are common. Chapped hands may signal excessive handwashing. If OCD is suspected, the use of a few simple screening questions can be helpful (Table 4).16 Standardized diagnostic tools are available, but most are not practical for use in primary care. There are some brief patient self-report inventories that may be useful; two commonly used tools are the Obsessive-Compulsive Inventory-Revised¹⁷ and the Florida Obsessive-Compulsive Inventory¹⁸ (Online Figure A). Psychiatric referral is indicated if there is diagnostic uncertainty.

Comorbidities

The rate of psychiatric comorbidity in patients with OCD is high, particularly in those with severe OCD. In one longitudinal study, more than 90 percent of patients with OCD met the criteria for at least one other axis I diagnosis in their lifetime.³ The most common comorbid diagnosis is major depressive disorder, which affects two thirds of persons with OCD at some point in life.³ Panic disorder, social phobia, specific phobias, and substance abuse are also com-

mon. The risk of suicide in persons with OCD is high; more than 50 percent experience suicidal ideation, and 15 percent have attempted suicide.³ Depression and hopelessness are major correlates of suicidal behavior in persons with OCD.¹⁹ Patients with OCD should be carefully monitored for suicide risk and symptoms of depression.

Treatment

Once a diagnosis of OCD has been established, it is important to provide education and support. Although

Table 3. Subtypes of Obsessive-Compulsive Disorder

Subtype	Features
Early-onset ^{7,8}	Symptom onset before puberty
	Higher frequency of tics and other psychiatric comorbiditie
	Onset of compulsions often predates obsessions
	Compulsions often severe and frequent
	Less responsive to first-line treatments
	Strong familial link (17 percent among first-degree relative
Hoarding ⁹	Less insight than in other OCD subtypes
	Symptoms often more severe
	Higher rates of psychiatric comorbidities, especially social phobia
	Greater degree of global impairment
	May be less responsive to psychological treatment
"Just right" ^{10,11}	Patients wish to have things "perfect," "certain," or "unde control"
	Results in a need to repeat certain actions until the uncomfortable feeling subsides
	"Not-just-right" experiences are common in all forms of OCD, but for some patients it is the primary manifestatic
Primary	25 percent of patients with OCD lack overt compulsions
obsessional ¹²	Patients are not free from rituals, which may be mental (e.g., praying, counting, reciting "good words")
	Common themes of obsessions are sex, violence, religion
	Historically thought to be less responsive to treatment, but does respond to medication and exposure and response prevention
Scrupulosity ¹³	Religious or moral obsessions
	Devastating form of OCD for patients to whom faith or religious affiliation is important
	Obsessions focus on whether one has committed a sin, or involve blasphemous thoughts
	Compulsions include prayer, reassurance-seeking from clergy, excessive confession
Fic-related ^{14,15}	Significant overlap with early-onset OCD
	Many patients meet criteria for Tourette syndrome
	High rate of comorbid conditions (e.g., attention-deficit/ hyperactivity disorder, body dysmorphic disorder, trichotillomania, social anxiety, mood disorders)
	Hoarding and somatic obsessions are common
	Often requires combination treatment with a selective serotonin reuptake inhibitor and an atypical antipsychoti

full remission is rare in patients with OCD, significant improvement is common. Evidence-based medical and behavioral therapies are available to reduce the severity and frequency of obsessions and compulsions. However, it may take weeks to months for these therapies to become effective. Physicians should inform patients about this delay in treatment response, provide support, and encourage adherence during the early phase of treatment.

It is helpful to quantify the severity of symptoms and impairment before and during treatment for OCD. This

Table 4. Screening Questions for Obsessive-Compulsive Disorder

- Do you have thoughts or images that keep coming back to you and are difficult to put out of your head? For example, being contaminated by something, having something terrible happen to you or someone you care about, or doing something terrible?
- Do you ever feel the need to perform certain actions that don't make sense or that you don't want to do, such as washing, cleaning, counting, or checking things over and over?

Adapted with permission from Canadian Psychiatric Association. Clinical practice guidelines. Management of anxiety disorders [published correction appears in Can J Psychiatry. 2006;51(10):623]. Can J Psychiatry. 2006;51(8 suppl 2):445.

may be done with standardized rating scales or by a patient estimate of the time spent each day engaging in obsessive-compulsive thoughts or behaviors. The Yale-Brown Obsessive Compulsive Scale (Y-BOCS) is a reliable tool for measuring OCD symptom severity.²⁰ It is also important to monitor the effect of OCD symptoms on relationships, work, self-care, and recreational time.

Treatment is indicated when OCD symptoms cause impairment in functioning or significant distress for the patient. Reasonable treatment goals would be spending less than one hour per day on obsessivecompulsive behaviors, with minimal interference with daily tasks.²¹ A treatment strategy algorithm is provided in *Figure 1.*^{16,21}

Psychiatric consultation is recommended for patients with severe OCD, as measured by the Y-BOCS. For patients with mild or subclinical symptoms, education and support may be sufficient. High-quality self-help materials are available that explain the nature of the disorder, its manifestations, and available treatments (see online patient handout).

PSYCHOLOGICAL TREATMENTS

Psychological treatments are effective for OCD.²² These treatments should be administered by a properly trained health care professional, most commonly a psychologist

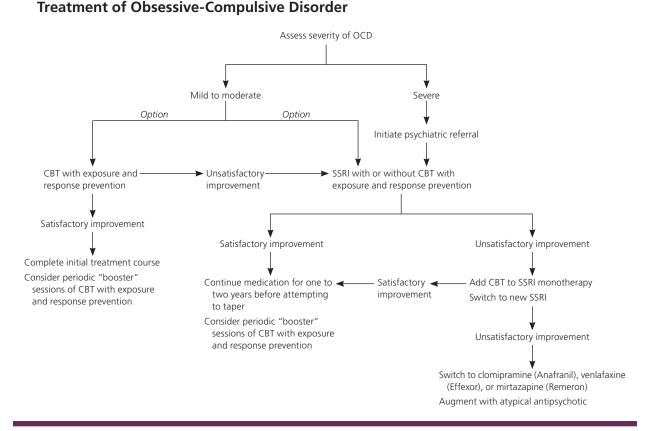


Figure 1. Treatment algorithm for patients with obsessive-compulsive disorder. (CBT = cognitive behavior therapy; OCD = obsessive-compulsive disorder; SSRI = selective serotonin reuptake inhibitor.)

Information from references 16 and 21.

or social worker. Cognitive behavior therapy (CBT) is the method of psychotherapy most often used; there is no evidence for the use of psychodynamic psychotherapy or "talk therapy" for treatment of OCD. Exposure and response prevention is a key element of CBT that has been proven effective in the treatment of OCD.²³ Patients are taught to confront situations that create fear related to their obsessions, and to avoid performing compulsive behaviors in response. The feared situations may be confronted directly (e.g., touching objects in a public restroom), or through imagined encounters (e.g., imagining shaking hands with another person). Patients refrain from performing rituals until the level of anxiety dissipates. Exposure and response prevention is usually performed in 13 to 20 weekly sessions, with each session lasting one to two hours.²¹

PHARMACOTHERAPY

OCD exhibits a highly selective response to serotonergic medications. Clomipramine (Anafranil), a tricyclic antidepressant with a strong serotonergic effect, was historically the first-line pharmacologic treatment for OCD. However, because of concerns about the safety and adverse effects of tricyclic agents, SSRIs have become first-line pharmacologic treatments for OCD. Fluoxetine (Prozac), fluvoxamine, paroxetine (Paxil), and sertraline (Zoloft) have been approved by the U.S. Food and Drug Administration for the treatment of OCD. Citalopram (Celexa) and escitalopram (Lexapro) are also commonly used. Approximately 60 to 70 percent of patients experience some degree of improvement in OCD symptoms

with SSRI treatment.²⁴ A recent Cochrane review confirmed the effectiveness of SSRIs for the treatment of OCD (absolute risk reduction = 8 to 17 percent; number needed to treat = 6 to 12).²⁵

The dosage of SSRI required to achieve treatment effect for OCD is often higher than the recommended dosages for other indications (*Table 5*).²¹ The dosage should be increased over four to six weeks until the maximal dosage is achieved, or until further increase is limited by adverse effects.²¹ Higher-than-maximal dosages are sometimes used, with careful monitoring for serotonin syndrome. Early signs of serotonin syndrome include anxiety, tremor, tachycardia, and sweating.²⁶ The patient should continue taking the SSRI for eight to 12 weeks, with at least four to six weeks at the maximal tolerable dosage.²¹ It usually takes at

least four to six weeks for patients to note any significant improvement in symptoms; for some, it may take 10 to 12 weeks or longer.

If medical therapy is successful, it should be continued for at least one to two years.²¹ If the patient chooses to discontinue pharmacotherapy, the dosage should be gradually tapered over several months. If symptoms worsen during this time, the original dosage should be resumed, and further attempts at discontinuing medication should be approached with reservation. Some patients require lifelong medical therapy.

Initial data suggest that the response to psychological treatments may be more durable than medication.²⁷ Periodic exposure and response prevention "booster" sessions are recommended to lower the risk of relapse when psychological therapy is discontinued.²¹ Initiating psychological treatments before a trial of medication discontinuation may also be an effective strategy to lower the risk of relapse.

If an adequate trial of an SSRI or psychological therapy does not result in a satisfactory response, one option is to initiate combined treatment. If the patient prefers to continue with medical therapy alone, a trial of a different SSRI is indicated.²¹ If there is no response to trials of at least two SSRIs, clomipramine may be considered.²¹ Clomipramine can cause anticholinergic adverse effects and, rarely, arrhythmia or seizures. It should be started at a low dose (25 mg) with gradual titration to minimize adverse reactions. Venlafaxine (Effexor) is another option for second-line treatment; the extended-release form was shown in a randomized controlled trial to be

Table 5. Typical SSRI Dosages in Patients with Obsessive-Compulsive Disorder

SSRI	Starting dosage (mg per day)	Target dosage (mg per day)	Maximal dosage (mg per day)
Citalopram (Celexa)	20	40 to 60	80
Escitalopram (Lexapro)	10	20	40
Fluoxetine (Prozac)*	20	40 to 60	80
Fluvoxamine*	50	200	300
Paroxetine (Paxil)*	20	40 to 60	60
Sertraline (Zoloft)*	50	200	200

SSRI = selective serotonin reuptake inhibitor.

*—Approved by the U.S. Food and Drug Administration for treatment of obsessivecompulsive disorder.

Adapted with permission from Koran LM, Hanna GL, Hollander E, Nestadt G, Simpson HB, for the American Psychiatric Association. Practice guideline for the treatment of patients with obsessive-compulsive disorder. Am J Psychiatry. 2007;164(7 suppl):22.

equivalent to paroxetine.²⁸ A small preliminary study suggested that mirtazapine (Remeron) may also be an effective treatment for OCD.²⁹

Another option for patients with OCD who have partially responded to SSRI therapy is the addition of an atypical antipsychotic. Antipsychotic augmentation is indicated only after a three-month trial of an SSRI at the maximal tolerated dosage.³⁰ Risperidone (Risperdal) has the strongest evidence base for use as an adjunctive agent; however, quetiapine (Seroquel) and olanzapine (Zyprexa) are also used.¹⁶ Antipsychotic augmentation is particularly beneficial in patients with comorbid tics. An SSRI in combination with risperidone or haloperidol (formerly Haldol) is the preferred treatment in these patients.³⁰

Patients with treatment-resistant OCD should be referred to a subspecialist. There are a variety of treatment options for these patients, but the evidence for most therapies is based on small preliminary studies or expert opinion.^{16,21} Partial hospitalization and residential treatment facilities are options for patients with severe, treatment-resistant OCD.

COMPLEMENTARY AND ALTERNATIVE MEDICINE

There are limited trials of complementary and alternative medicine approaches for the treatment of OCD. Initial studies have suggested beneficial effects for moderate-intensity aerobic exercise and mindfulness interventions (e.g., meditative breathing).^{31,32} There has long been interest in the use of St. John's wort for treatment of OCD. A recent double-blind study did not support the effectiveness of this treatment, although further study has been recommended.³³

Special Considerations for Childhood OCD

Although OCD in childhood can occur in isolation, there is a high rate of comorbidity with mood disorders, tic disorders, attention-deficit/hyperactivity disorder, and developmental abnormalities. Children with abrupt onset of obsessive-compulsive symptoms or tics should be evaluated for group A *Streptococcus* infection, with possible PANDAS. Children with OCD should be referred to a subspecialist. CBT with exposure and response prevention is the preferred initial treatment modality. SSRI treatment may be indicated in patients with severe symptoms, or when there is lack of improvement with CBT alone.²¹

The authors thank Joseph Himle, PhD, of the University of Michigan Medical School Department of Psychiatry, for assistance with the preparation of the manuscript.

The Authors

JILL N. FENSKE, MD, is a clinical lecturer in the Department of Family Medicine at the University of Michigan Medical School, Ann Arbor.

THOMAS L. SCHWENK, MD, is the George A. Dean, MD, Chair and Professor of Family Medicine at the University of Michigan Medical School, an associate editor of *Journal Watch*, and a section editor of UpToDate.

Address correspondence to Jill N. Fenske, MD, Chelsea Health Center, 14700 E. Old U.S. Highway 12, Chelsea, MI 48118 (e-mail: jnfenske@ med.umich.edu). Reprints are not available from the authors.

Author disclosure: Nothing to disclose.

REFERENCES

- Kessler RC, Chiu WT, Demler O, Merikangas KR, Walters EE. Prevalence, severity, and comorbidity of 12-month DSM-IV disorders in the National Comorbidity Survey Replication [published correction appears in Arch Gen Psychiatry. 2005;62(7):709]. Arch Gen Psychiatry. 2005;62(6):617-627.
- 2. Koran LM. Quality of life in obsessive-compulsive disorder. *Psychiatr Clin North Am.* 2000;23(3):509-517.
- 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.
- Friedlander L, Desrocher M. Neuroimaging studies of obsessive-compulsive disorder in adults and children. *Clin Psychol Rev.* 2006;26(1):32-49.
- Pauls DL. The genetics of obsessive compulsive disorder: a review of the evidence. Am J Med Genet C Semin Med Genet. 2008;148(2):133-139.
- American Psychiatric Association. *Diagnostic and Statistical Manual of* Mental Disorders. 4th ed., text revision. Washington, DC: American Psychiatric Association; 2000.
- Chabane N, Delorme R, Millet B, Mouren MC, Leboyer M, Pauls D. Earlyonset obsessive-compulsive disorder: a subgroup with a specific clinical and familial pattern? J Child Psychol Psychiatry. 2005;46(8):881-887.
- Rosario-Campos MC, Leckman JF, Mercadante MT, et al. Adults with early-onset obsessive-compulsive disorder. *Am J Psychiatry*. 2001;158(11):1899-1903.
- 9. Wheaton M, Timpano KR, Lasalle-Ricci VH, Murphy D. Characterizing the hoarding phenotype in individuals with OCD: associations with comorbidity, severity and gender. *J Anxiety Disord*. 2008;22(2):243-252.
- Coles ME, Frost RO, Heimberg RG, Rhéaume J. "Not just right experiences": perfectionism, obsessive-compulsive features and general psychopathology. *Behav Res Ther.* 2003;41(6):681-700.
- Rasmussen SA, Eisen JL. The epidemiology and clinical features of obsessive compulsive disorder. *Psychiatr Clin North Am.* 1992;15(4):743-758.
- McKay D, Abramowitz JS, Calamari JE, et al. A critical evaluation of obsessive-compulsive disorder subtypes: symptoms versus mechanisms. *Clin Psychol Rev.* 2004;24(3):283-313.
- 13. Nelson EA, Abramowitz JS, Whiteside SP, Deacon BJ. Scrupulosity in patients with obsessive-compulsive disorder: relationship to clinical and cognitive phenomena. J Anxiety Disord. 2006;20(8):1071-1086.
- 14. Diniz JB, Rosario-Campos MC, Hounie AG, et al. Chronic tics and Tourette syndrome in patients with obsessive-compulsive disorder. *J Psychiatr Res.* 2006;40(6):487-493.
- McDougle CJ, Goodman WK, Leckman JF, Lee NC, Heninger GR, Price LH. Haloperidol addition in fluvoxamine-refractory obsessive-compulsive disorder. A double-blind, placebo-controlled study in patients with and without tics. Arch Gen Psychiatry. 1994;51(4):302-308.
- Canadian Psychiatric Association. Clinical practice guidelines. Management of anxiety disorders [published correction appears in *Can J Psychiatry*. 2006;51(10):623]. *Can J Psychiatry*. 2006;51(8 suppl 2):95-915.

- Foa EB, Huppert JD, Leiberg S, et al. The Obsessive-Compulsive Inventory: development and validation of a short version. *Psychol Assess*. 2002;14(4):485-496.
- Storch EA, Kaufman DA, Bagner D, et al. Florida Obsessive-Compulsive Inventory: development, reliability, and validity [published correction appears in *J Clin Psychol.* 2007;63(12):1265]. *J Clin Psychol.* 2007;63(9):851-859.
- 19. KamathP,ReddyYC,KandavelT.Suicidalbehaviorinobsessive-compulsive disorder. J Clin Psychiatry. 2007;68(11):1741-1750.
- Rush JA. Handbook of Psychiatric Measures. Washington, DC: American Psychiatric Association; 2000:572-574.
- Koran LM, Hanna GL, Hollander E, Nestadt G, Simpson HB, for the American Psychiatric Association. Practice guideline for the treatment of patients with obsessive-compulsive disorder. *Am J Psychiatry.* 2007; 164(7 suppl):5-53.
- Gava I, Barbui C, Aguglia E, et al. Psychological treatments versus treatment as usual for obsessive compulsive disorder (OCD). *Cochrane Database Syst Rev.* 2007;(2):CD005333.
- Fisher PL, Wells A. How effective are cognitive and behavioral treatments for obsessive-compulsive disorder? A clinical significance analysis. *Behav Res Ther.* 2005;43(12):1543-1558.
- Eddy KT, Dutra L, Bradley R, Westen D. A multidimensional meta-analysis of psychotherapy and pharmacotherapy for obsessive-compulsive disorder. *Clin Psychol Rev.* 2004;24(8):1011-1030.
- Soomro GM, Altman D, Rajagopal S, Oakley-Browne M. Selective serotonin re-uptake inhibitors (SSRIs) versus placebo for obsessive compulsive disorder (OCD). Cochrane Database Syst Rev. 2008;(1):CD001765.

- Adams SM, Miller KE, Zylstra RG. Pharmacologic management of adult depression. Am Fam Physician. 2008;77(6):785-792.
- Simpson HB, Liebowitz MR, Foa EB, et al. Post-treatment effects of exposure therapy and clomipramine in obsessive-compulsive disorder. *Depress Anxiety*. 2004;19(4):225-233.
- Denys D, van der Wee N, van Megen HJ, Westenberg HG. A double blind comparison of venlafaxine and paroxetine in obsessive-compulsive disorder. J Clin Psychopharmacol. 2003;23(6):568-575.
- Koran LM, Gamel NN, Choung HW, Smith EH, Aboujaoude EN. Mirtazapine for obsessive-compulsive disorder: an open trial followed by double-blind discontinuation. J Clin Psychiatry. 2005;66(4):515-520.
- Bloch MH, Landeros-Weisenberger A, Kelmendi B, Coric V, Bracken MB, Leckman JF. A systematic review: antipsychotic augmentation with treatment refractory obsessive-compulsive disorder [published correction appears in *Mol Psychiatry*. 2006;11(8):795]. *Mol Psychiatry*. 2006;11(7):622-632.
- Brown RA, Abrantes AM, Strong DR, et al. A pilot study of moderateintensity aerobic exercise for obsessive compulsive disorder. *J Nerv Ment Dis.* 2007;195(6):514-520.
- Hanstede M, Gidron Y, Nyklícek I. The effects of a mindfulness intervention on obsessive-compulsive symptoms in a non-clinical student population. J Nerv Ment Dis. 2008;196(10):776-779.
- Kobak KA, Taylor LV, Bystritsky A, et al. St John's wort versus placebo in obsessive-compulsive disorder: results from a double-blind study. *Int Clin Psychopharmacol.* 2005;20(6):299-304.