

Eating Disorders Update 2019

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Objectives

1. Review diagnosis, comorbidity, risk, and treatment of ED.
2. Discuss the complexity of severe and enduring EDs.
3. Possibly discuss ICAT???

Part 1: Overview: Diagnosis, Comorbidity, and Risk

Eating Disorders: The Big Issue (Schmidt et al; Lancet)

- Peak age of onset: 15-25 years.
- Average duration of disorder: 6 years.
- AN and BN more common in young women, but men/boys affected; BED equally split between genders.
- EDs diagnosed in people at increasingly younger ages.
- AN is one of the most common chronic illnesses of adolescence- similar to incidence rates of Type I Diabetes in rates.

Eating Disorders: The Big Issue (Schmidt et al; Lancet)

- Mortality rates for EDs are 2x general population and AN is 6x greater.
- 1 of 3 people with BN or BED will become obese.
- 1 in 4 ED patients is unemployed.
- Women with EDs are more likely to stay childless and need fertility treatment to conceive.
- Women with ED have significant problems feeding and interacting with their children.

Eating Disorders: The Big Issue (Schmidt et al; Lancet)

- Caregivers of ED patients spend nearly twice as much time care-giving (24 h/week vs. 14 h/week) than caregivers of patients with other disorders (e.g. cancer, psychosis, dementia).
- Recent estimates: 20 million people in Europe have an ED with a cost of 1 trillion Euros per year (financial cost-249 billion and burden of disease cost-763 billion)

DSM-5 Criteria for Anorexia Nervosa

- A.** Restriction of energy intake relative to requirements, leading to a **significantly low body weight in the context of age, sex, developmental** trajectory, and physical health. Significantly low weight is defined as a weight that is less than minimally normal or, for children and adolescents, less than that minimally expected.
- B.** Intense fear of gaining weight or becoming fat, or persistent behavior that interferes with weight gain, even though at a significantly low weight.
- C.** Disturbance in the way in which one's body weight or shape is experienced, undue influence of body weight or shape on self-evaluation, or persistent lack of recognition of the seriousness of the current low body weight.

DSM-5 Criteria for Bulimia Nervosa

- A.** Recurrent episodes of binge eating. An episode of binge eating is characterized by **both** of the following:
 - 1.** Eating, in a discrete period of time (e.g., within any 2-hour period), an amount of food that is definitely larger than what most individuals would eat in a similar period of time under similar circumstances.
 - 2.** A sense of lack of control over eating during the episode (e.g., a feeling that one cannot stop eating or control what or how much one is eating)
- B.** Recurrent inappropriate compensatory behavior in order to prevent weight gain, such as self-induced vomiting; misuse of laxatives, diuretics, and other medications; fasting; or excessive exercise.
- C.** The binge eating and inappropriate compensatory behaviors both occur, on average, at least once a week for 3 months.
- D.** Self-evaluation is unduly influenced by body shape and weight.
- E.** The disturbance does not occur exclusively during episodes of anorexia nervosa.

DSM-5 Criteria for Binge Eating Disorder

- A. Recurrent episodes of binge eating.
- B. The binge-eating episodes are associated with three (or more) of the following:
 - 1. eating much more rapidly than normal
 - 2. eating until feeling uncomfortably full
 - 3. eating large amounts of food when not feeling physically hungry
 - 4. eating alone because of being embarrassed by how much one is eating
 - 5. feeling disgusted with oneself, depressed, or very guilty after overeating
- C. Marked distress regarding binge eating is present.
- D. The binge eating occurs, on average, at least once a week for 3 months.
- E. The binge eating is not associated with the recurrent use of inappropriate compensatory behavior and does not occur exclusively during the course of bulimia nervosa or anorexia nervosa.

Other Feeding and Eating Disorders

- 1. Rumination disorder
- 2. PICA
- 3. Avoidant restrictive food intake disorder (ARFID)

Comorbidity/Associated Problems

Lifetime Psychiatric Diagnoses in ED Patients

	ANR	ANBP	BN
Depression [median %(N)]	40 (5)	73 (5)	48.5 (8)
OCD [median %(N)]	20 (7)	20 (6)	15 (8)
Social Phobia [median %(N)]	26 (4)	28 (4)	18.5 (4)
Panic [median %(N)]	9 (8)	14 (5)	10 (7)
Substance Use [median %(N)]	5 (5)	15 (4)	22.5 (6)
Phobia [median %(N)]	15 (3)	18 (3)	12 (3)

Zonneville-Bender et al., 2004; Kaye et al., 2004; Bulik et al., 2004; Binford & leGrange, 2005; Godart et al., 2004; Ricca et al., 2001; Halmi et al., 1991; Herzog et al., 1992; Brewerton et al., 1995; Fornari et al., 1991; Braun et al., 1994

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Twelve-year course and outcome of bulimia nervosa

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Prien, Germany*

Twelve Year Course and Outcome of BN

- 196 females met DSM IV criteria for BN
- Admitted to ED hospital in Bavaria between 1985 and 1988
- Interview based assessment of ED and psychiatric comorbidity
- Assessed at admission, d/c, 2yr, 6yr, and 12yr follow-up

Twelve Year Course and Outcome of BN

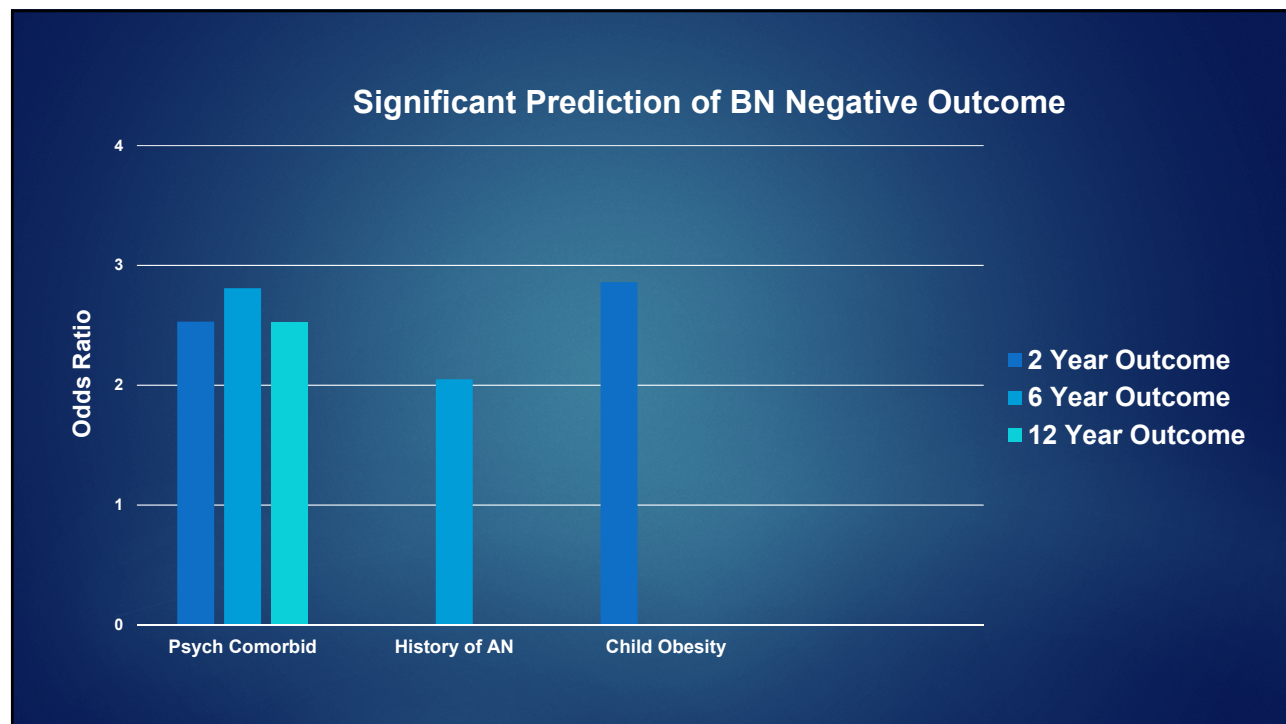
12 year follow up

- 167 patients assessed
- 117 (70.1%) recovered
- Standardized mortality: 2.36

Twelve Year Course and Outcome of BN

Predictor Analysis (Baseline predictors → 12 year outcome)

1. Presence of lifetime psychiatric disorder (+ BN)
2. History of AN
3. Childhood obesity
4. Age of onset
5. Duration of illness
6. Frequency of binge episodes
7. Previous treatment



Predisposing Factors

Genetics and epigenetics of eating disorders

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Video abstract



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Abstract: Eating disorders (EDs) are serious psychiatric conditions influenced by biological, psychological, and sociocultural factors. A better understanding of the genetics of these complex traits and the development of more sophisticated molecular biology tools have advanced our understanding of the etiology of EDs. The aim of this review is to critically evaluate the literature on the genetic research conducted on three major EDs: anorexia nervosa, bulimia nervosa, and binge eating disorder. We will first review the diagnostic criteria, clinical features, prevalence, and prognosis of anorexia nervosa, bulimia nervosa, and binge eating disorder, followed by a review of family, twin, and adoption studies. We then review the history of genetic studies of EDs, covering linkage analysis, candidate-gene association studies, genome-wide association studies, and the study of rare variants in EDs. Our review also incorporates a translational perspective by covering animal models of ED-related phenotypes. Finally, we review the nascent field of epigenetics of EDs and a look forward to future directions for ED genetic research.

Keywords: anorexia nervosa, binge eating disorder, bulimia nervosa, animal models, genome-wide association studies, high-throughput sequencing

Overview of eating disorders

Eating disorders (EDs) are serious psychiatric conditions with significant morbidity and mortality. That EDs have a genetic component may come as a surprise to many due to widespread misperception of them being disorders of volition. Research over the past decade has confirmed that genes do indeed play a role, and animal models of core related phenotypes are assisting with defining the underlying biology of these pernicious illnesses. In this review, we focus on three major EDs: anorexia nervosa (AN), bulimia nervosa (BN), and binge eating disorder (BED). Most of the genetic research has focused on AN and BN; less information is available for BED due to its status as a newly recognized ED diagnosis.

Anorexia nervosa

AN (International Classification of Diseases, 10th revision [ICD-10]: F50.00) is a serious ED with substantial morbidity and the highest lifetime mortality among psychiatric disorders.¹⁻³ Low weight or body mass index (BMI) is the sine qua non of AN and the primary target of initial treatment.^{4,5} Symptoms of AN include persistent restriction of food intake, an intense fear of gaining weight or persistent behavior that interferes with weight gain, and a distorted body image.¹ There are two subtypes of AN: restricting subtype (ICD-10: F50.01) and binge/purge subtype (ICD-10: F50.02).⁶

REGULAR ARTICLE

5-HT_{1A} Receptor Binding Is Increased After Recovery from Bulimia Nervosa Compared to Control Women and Is Associated with Behavioral Inhibition in Both Groups

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ABSTRACT

Objective: Because altered serotonin (5-HT) function appears to persist after recovery from bulimia nervosa (RBN), we investigated the 5-HT_{1A} receptor, which could contribute to regulation of appetite, mood, impulse control, or the response to antidepressants.

Method: Thirteen RBN individuals were compared to 21 healthy control women (CW) using positron emission tomography and [carbonyl-¹¹C]WAY100635 ([¹¹C]WAY).

Results: RBN had a 23–34% elevation of [¹¹C]WAY binding potential (BP)_B in subgenual cingulate, mesial temporal, and parietal regions after adjustments for multiple comparisons. For CW, [¹¹C]WAY BP_B was related negatively to novelty seeking, whereas for RBN, [¹¹C]WAY

BP_B was related positively to harm avoidance and negatively related to sensation seeking.

Discussion: Alterations of 5-HT_{1A} receptor function may provide new insight into efficacy of 5-HT medication in BN, as well as symptoms such as the ability to inhibit or self-control the expression of behaviors related to stimulus seeking, aggression, and impulsivity. © 2010 by Wiley Periodicals, Inc.

Keywords: bulimia nervosa; 5-HT_{1A} receptor; positron emission tomography; behavioral inhibition; subgenual cingulate; mesial temporal cortex

(Int J Eat Disord 2010; 00:000–000)

Biological Factors

Personality Factors

Personality subtyping and bulimia nervosa: psychopathological and genetic correlates

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ABSTRACT

Background. There is empirical evidence suggesting that individuals with bulimia nervosa vary considerably in terms of psychiatric co-morbidity and personality functioning. In this study, latent profile analysis was used to attempt to identify clusters of bulimic subjects based on psychiatric co-morbidity and personality.

Method. A total of 178 women with bulimia nervosa or a subclinical variant of bulimia nervosa completed a series of self-report inventories of co-morbid psychopathology and personality, and also provided a buccal smear sample for genetic analyses.

Results. Three clusters of bulimic women were identified: an affective-perfectionistic cluster, an impulsive cluster, and a low co-morbid psychopathology cluster. The clusters showed expected differences on external validation tests with both personality and eating-disorder measures. The impulsive cluster showed the highest elevations on dissocial behavior and the lowest scores on compulsivity, while the affective-perfectionistic cluster showed the highest levels of eating-disorder symptoms. The clusters did not differ on genetic variations of the serotonin transporter gene.

Conclusions. This study corroborates previous findings suggesting that the bulimia nervosa diagnostic category is comprised of three classes of individuals based on co-morbid psychopathology and personality. These differences may have significant etiological and treatment implications.

Childhood Stress

Effects of multiple forms of childhood abuse and adult sexual assault on current eating disorder symptoms

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Sexual Assault

ABSTRACT

The objective of this study is to examine the effect of recent adult sexual assault on current eating disorder symptoms when controlling for the effects of multiple forms of childhood abuse. A total of 489 undergraduate women completed the Eating Disorder Examination-Questionnaire, and surveys regarding childhood abuse and sexual assault that had occurred in the previous three months. Approximately 30% of the sample indicated recent unwanted sexual experiences. Childhood emotional abuse contributed unique variance to the prediction of current ED symptoms, but sexual and physical abuse did not. Recent sexual assault contributed additional unique variance to current ED symptoms when controlling for childhood abuse, thus both emotional abuse in childhood and sexual assault in adulthood contributed unique variance to ED symptoms.

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Weight & Performance-Related Activities

Aspects of disordered eating continuum in elite high-intensity sports

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Dieting is an important risk factor for disordered eating and eating disorders. Disordered eating occurs on a continuum from dieting and restrictive eating, abnormal eating behavior, and finally clinical eating disorders. The prevalence of eating disorders is increased in elite athletes and for this group the cause of starting to diet is related to (a) perception of the paradigm of appearance in the specific sport, (b) perceived performance improvements, and (c) sociocultural pressures for thinness or an "ideal" body. Athletes most at risk for disordered eating are those involved in sports emphasizing a thin body size/shape, a high power-to-weight ratio, and/or sports utilizing weight categories,

such as in some high-intensity sports. In addition to dieting, personality factors, pressure to lose weight, frequent weight cycling, early start of sport-specific training, overtraining, injuries, and unfortunate coaching behavior, are important risk factors. To prevent disordered eating and eating disorders, the athletes have to practice healthy eating, and the medical staff of teams and parents must be able to recognize symptoms indicating risk for eating disorders. Coaches and leaders must accept that disordered eating can be a problem in the athletic community and that openness regarding this challenge is important.

Socio-Cultural Factors

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Sociocultural pressures and adolescent eating in the absence of hunger

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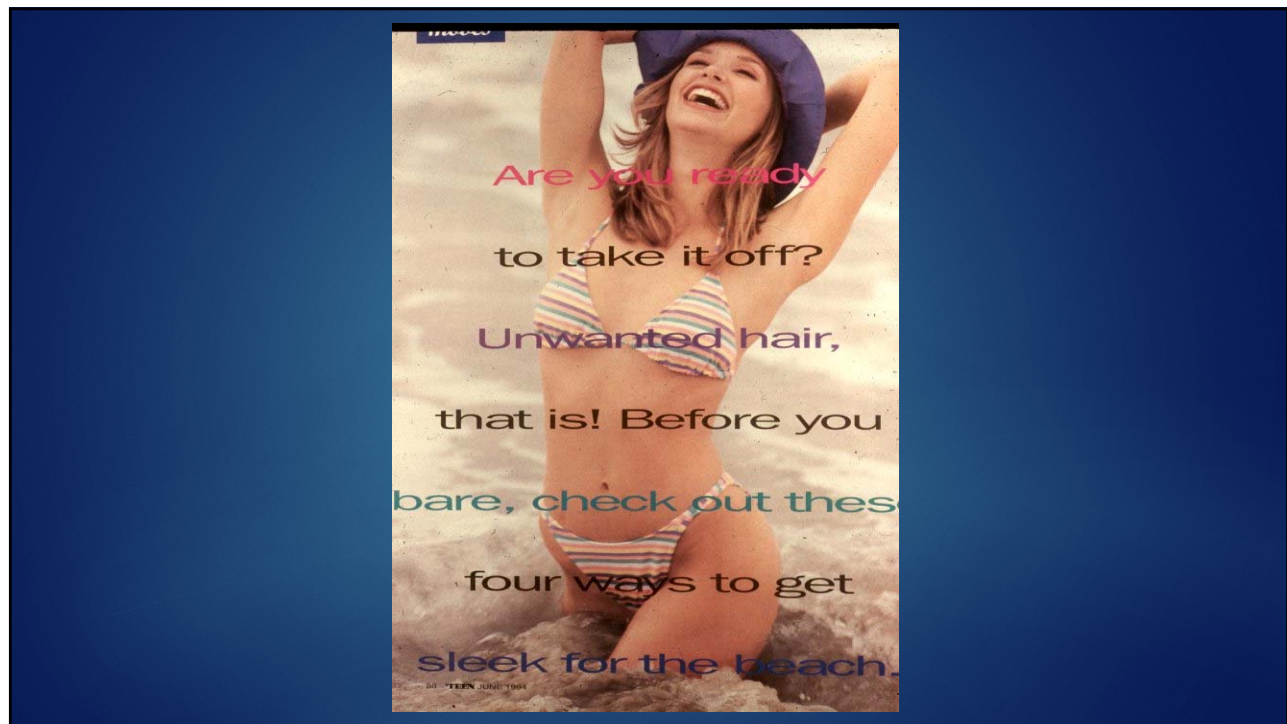
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The latest lingo on the short scene:

1. **Hip-huggers** are cut low on the hips.
2. **Biker-length shorts** fall just above the knee and are cut close to the leg.
3. **Short shorts** have a two-and-one-half-inch inseam.
4. **Boxer-cut shorts** are similar to traditional boxer underwear.
5. **Paper-bag waist shorts** have a belt that cinches at the waist, creating gothors.
6. **Micro shorts** have one-and-one-half or two-inch inseams.

PHOTOGRAPH BY KEVIN BAUM

SHORT LEGS?

Best bet: Micro-shorts

The short story: The more leg that shows, the longer your legs will look. Choose a fabric with a small pattern to help in proportion with your body frame.

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want your legs to look longer?

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Choose the one just right for you!

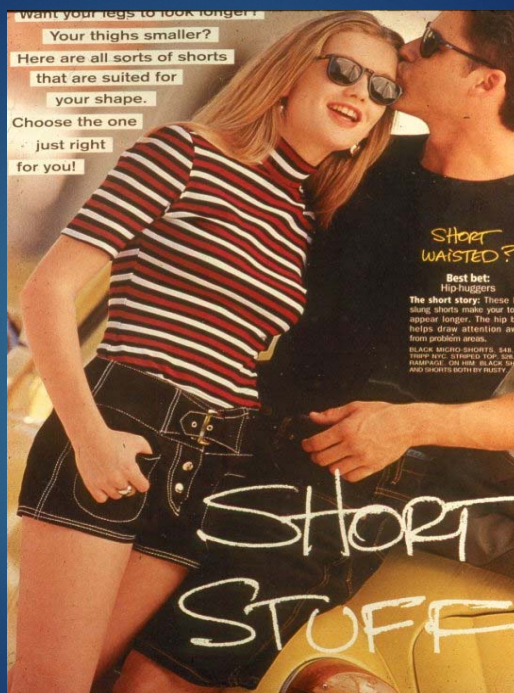
SHORT WAISTED?

Best bet: Hip-huggers

The short story: These lo-sling shorts make your legs appear longer. The hip belt helps draw attention away from problem areas.

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SHORT STUFF



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This summer can be the best summer of your life! Make sure you don't miss out on all the fun at the beach, parties, and at the pool!

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Send me _____.

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These are the parts of my body I want to change:

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☐ Neck ☐ Waist ☐ Thighs
☐ Chest ☐ But ☐ Legs

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PRINT

Name _____

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City _____ State _____ Zip _____

PERSONAL BODY GOALS FORM

(Answer only if you want to)

I weigh _____ pounds; I WANT to weigh _____

These are the parts of my body I want to change:

Check as many as you want:

- | | | |
|--------------------------------|--------------------------------|---------------------------------|
| <input type="checkbox"/> Face | <input type="checkbox"/> Arms | <input type="checkbox"/> Hips |
| <input type="checkbox"/> Neck | <input type="checkbox"/> Waist | <input type="checkbox"/> Thighs |
| <input type="checkbox"/> Chest | <input type="checkbox"/> Butt | <input type="checkbox"/> Legs |

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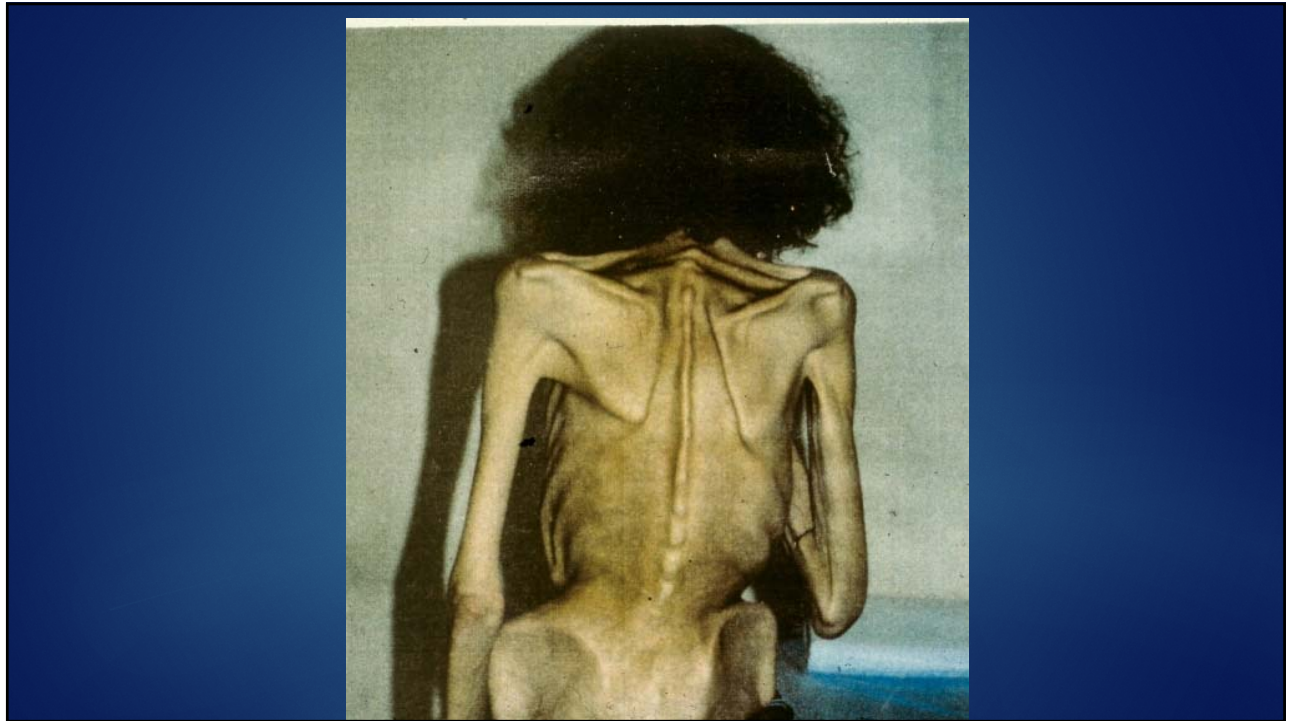
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Part II: Treatment

The Prevalence and Correlates of Eating Disorders in the National Comorbidity Survey Replication

James I. Hudson, Eva Hiripi, Harrison G. Pope, Jr., and Ronald C. Kessler

Background: Little population-based data exist on the prevalence or correlates of eating disorders.

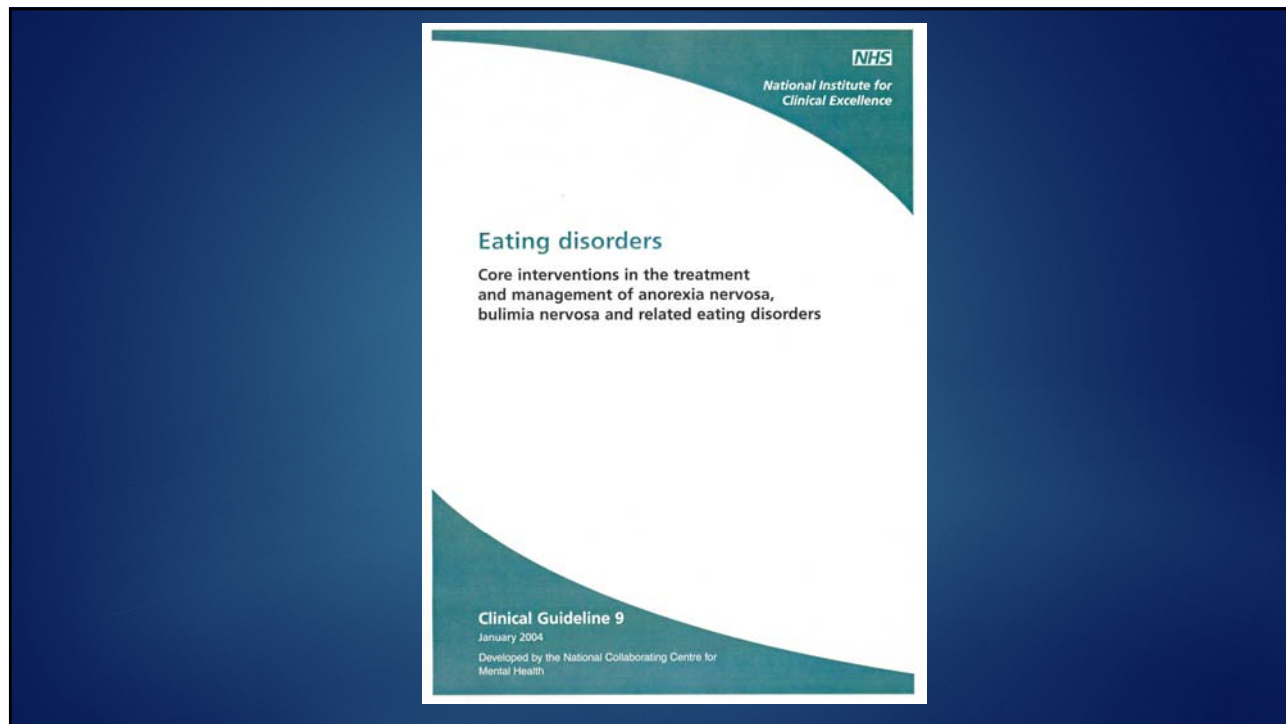
Methods: Prevalence and correlates of eating disorders from the National Comorbidity Replication, a nationally representative face-to-face household survey ($n = 9282$), conducted in 2001–2003, were assessed using the WHO Composite International Diagnostic Interview.

Results: Lifetime prevalence estimates of DSM-IV anorexia nervosa, bulimia nervosa, and binge eating disorder are .9%, 1.5%, and 3.5% among women, and .3%, .5%, and 2.0% among men. Survival analysis based on retrospective age-of-onset reports suggests that risk of bulimia nervosa and binge eating disorder increased with successive birth cohorts. All 3 disorders are significantly comorbid with many other DSM-IV disorders. Lifetime anorexia nervosa is significantly associated with low current weight (body-mass index < 18.5), whereas lifetime binge eating disorder is associated with current severe obesity (body-mass index ≥ 40). Although most respondents with 12-month bulimia nervosa and binge eating disorder report some role impairment (data unavailable for anorexia nervosa since no respondents met criteria for 12-month prevalence), only a minority of cases ever sought treatment.

Conclusions: Eating disorders, although relatively uncommon, represent a public health concern because they are frequently associated with other psychopathology and role impairment, and are frequently under-treated.

Lifetime and 12 Month Treatment of DSM IV EDs-NCS-R (n=2,980)

	AN	BN	BED
ED Tx			
Lifetime (%)	33.8	43.2	43.6
Any Psychiatric Tx			
Lifetime (%)	50	63.2	51.2



How to Find the NICE Guideline

www.NICE.org.uk

Anorexia Nervosa (NICE Guidelines)

- Most people with anorexia nervosa should be managed on an outpatient basis with psychological treatment by a service that is competent in giving that treatment and assessing the physical risk of people with eating disorders.

ED Treatment Team



Anorexia Nervosa (NICE Guidelines Cont.)

- People with anorexia nervosa requiring inpatient treatment should be admitted to a setting that can provide the skilled implementation of refeeding with careful physical monitoring (particularly in the first few days of refeeding) in combination with psychosocial interventions.

Anorexia Nervosa (NICE Guidelines Cont.)

- Family interventions that directly address the eating disorder should be offered to children and adolescents with anorexia nervosa.

What is the Maudsley Family Based Approach?

- Outpatient weight restoration treatment
- ~ Twenty sessions over 6-12 months
- Puts PARENTS in charge of weight restoration (appropriate control, ultimately relinquished)
- Contrary to traditional separation of parents and child
- No assumption about etiology of AN

Treatment of AN

- Maudsley or Family Based Treatment has not been shown to be effective for patients over 18
- No specific treatment has been shown to be effective for AN adults in a scientific study
 - RCTs of psychological treatment are plagued by high dropout and small effects regardless of treatment type

Pharmacotherapy and AN

Antidepressants (fluoxetine, venlafaxine, citalopram, tricyclics)

- Small trials
- High dropout rates
- 4 placebo controlled trials did not find evidence of weight gain or improvement in ED psychopathology

Atypical Antipsychotics (olanzapine, risperidone)

- Small trials
- Not well controlled
- Limited effect on weight or psychopathology-(obsessional thinking?)

Kaplan & Howlett, 2010; Dynamed, 2015

Fluoxetine After Weight Restoration in Anorexia Nervosa A Randomized Controlled Trial

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ANOREXIA NERVOSA IS A SERIOUS psychiatric illness with substantial morbidity and a lifetime mortality arguably as high as that associated with any psychiatric illness.¹ A major contributor to the poor prognosis of this illness is the high rate of relapse following initial treatment. Despite successful weight restoration, 30% to 50% of patients require rehospitalization within 1 year of discharge.²⁻⁵ This discouraging experience has prompted interest in interventions aimed at preventing deterioration following weight restoration.

Patients with anorexia nervosa often exhibit symptoms of other psychiatric disorders,⁶ such as depression and obsessive-compulsive disorder, which are responsive to antidepressant medication, suggesting that pharmacological interventions might be of use. Surprisingly,

See also p 2659 and Patient Page.

Context Antidepressant medication is frequently prescribed for patients with anorexia nervosa.

Objective To determine whether fluoxetine can promote recovery and prolong time-to-relapse among patients with anorexia nervosa following weight restoration.

Design, Setting, and Participants Randomized, double-blind, placebo-controlled trial. From January 2000 until May 2005, 93 patients with anorexia nervosa received intensive inpatient or day-program treatment at the New York State Psychiatric Institute or Toronto General Hospital. Participants regained weight to a minimum body mass index (calculated as weight in kilograms divided by the square of height in meters) of 19.0 and were then eligible to participate in the randomized phase of the trial.

Interventions Participants were randomly assigned to receive fluoxetine or placebo and were treated for up to 1 year as outpatients in double-blind fashion. All patients also received individual cognitive behavioral therapy.

Main Outcome Measures The primary outcome measures were time-to-relapse and the proportion of patients successfully completing 1 year of treatment.

Results Forty-nine patients were assigned to fluoxetine and 44 to placebo. Similar percentages of patients assigned to fluoxetine and to placebo maintained a body mass index of at least 18.5 and remained in the study for 52 weeks (fluoxetine, 26.5%; placebo, 31.5%; $P = .57$). In a Cox proportional hazards analysis, with prerandomization body mass index, site, and diagnostic subtype as covariates, there was no significant difference between fluoxetine and placebo in time-to-relapse (hazard ratio, 1.12; 95% CI, 0.65-2.01; $P = .64$).

Conclusions This study failed to demonstrate any benefit from fluoxetine in the treatment of patients with anorexia nervosa following weight restoration. Future efforts should focus on developing new models to understand the persistence of this illness and on exploring new psychological and pharmacological treatment approaches.

Trial Registration clinicaltrials.gov Identifier: NCT00288574

JAMA. 2006;295:2605-2612

www.jama.com

ingly, virtually all of the controlled trials of medication (most of which have been conducted during the initial phase of treatment when patients are underweight) have shown no benefit of medication compared with placebo.⁷⁻⁹ Despite this lack of evidence of effectiveness, a substantial number of pa-

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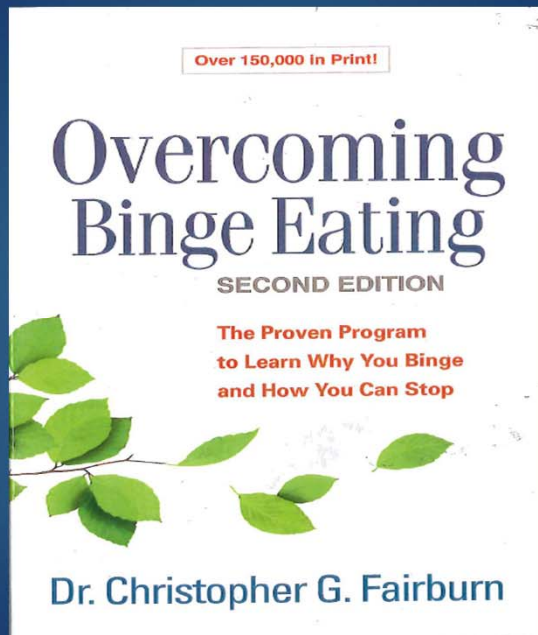
(Reprinted) JAMA, June 14, 2006—Vol 295, No. 22 2605

Pharmacotherapy and AN (Walsh et al., 2006)

- RCT with 93 weight restored AN patients (post discharge)
- Fluoxetine vs. placebo for 1 year
- No difference in relapse or number successfully completing 1 year of treatment (Relapse: Plac=55%; Fluox = 57%)

Bulimia Nervosa (NICE Guidelines)

- As a possible first step, patients with bulimia nervosa should be encouraged to follow an evidence-based self-help programme.



Bulimia Nervosa (NICE Guidelines cont.)

- As an alternative or additional first step to using an evidence-based self-help program, adults with bulimia nervosa may be offered a trial of an antidepressant drug.

Bulimia Nervosa (NICE Guidelines cont.)

- As an alternative or additional first step to using an evidence-based self-help program, adults with bulimia nervosa may be offered a trial of an antidepressant drug.

Pharmacotherapy and BN

- Antidepressants are the medications recommended in NICE guideline
- SSRIs, particularly fluoxetine, are drugs of choice
- High dropout rate (30-40%)
- Add antidepressant to structured psychotherapy if no reduction in symptoms after 10 sessions

Dynamed, 2015

Bulimia Nervosa (NICE Guidelines cont.)

- Cognitive behaviour therapy for bulimia nervosa (CBT-BN), a specifically adapted form of CBT, should be offered to adults with bulimia nervosa. The course of treatment should be for 16 to 20 sessions over 4 to 5 months.

Bulimia Nervosa (NICE Guidelines cont.)

- Adolescents with bulimia nervosa may be treated with CBT-BN, adapted as needed to suit their age, circumstances and level of development, and including the family as appropriate.

Royal Australian and New Zealand College of Psychiatrists Practice Guidelines for the Treatment of Eating Disorders (2014)

Binge Eating Disorder

1. First line treatment is psychological treatment-CBT
2. Consider topiramate or orlistat for binge control and obesity
3. Consider high dose fluoxetine-alone if psychological treatment is not available
4. Combined treatment (drug and psychological) may offer added benefit

Pharmacotherapy of BED

	Binge Eating	Wt. Loss
Tricyclics (Imipramine, Desipramine)	++	±
SSRIs/NRIs (Citalopram, Fluvoxamine, Sertraline)	++	+
Orlistat	±	++
Anti-epileptics (Topiramate , Zonisamide)	++	+++

± = Minimal effect
+ = Small effect
++ = Moderate effect
+++ = Significant effect

Mitchell et al., 2013

Other Treatments Being Tested or Recently Distributed

- Affect regulation based treatments for BN and BED
 - Integrative Cognitive Affective Therapy (ICAT)
 - Dialectical Behavior Therapy (DBT) for BN and BED
- Lisdexamphetamine (LDX, Vyvanse) for BED (FDA approved)
- Intranasal naloxone for binge eating

Hospital Based Treatment

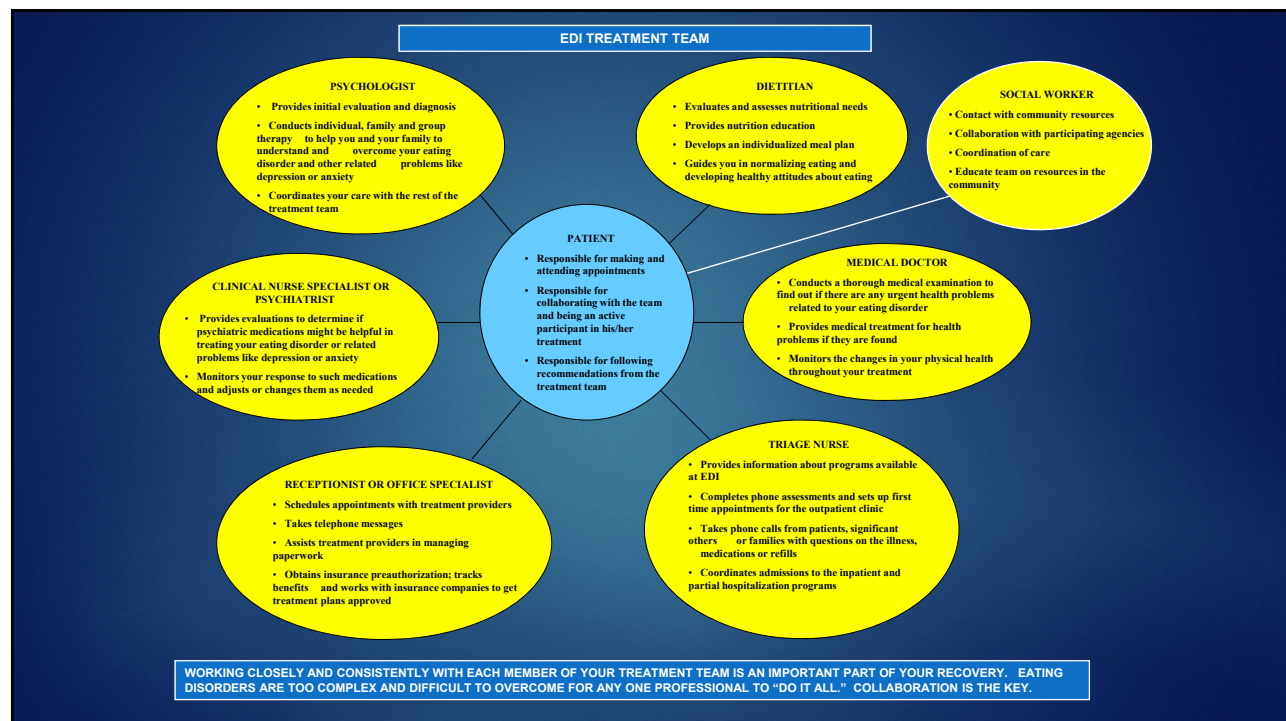
Continuum of Care

- Self help
- Outpatient
- Intensive outpatient
- Partial hospital
- Residential
- Inpatient

Factors to Consider in need for Hospitalization

- Medical complications
- Suicidality
- Body weight
- Low motivation to recover
- Comorbid disorders
- Impairment in ability to care for self
- Extreme environmental stress
- Treatment availability

Team Approach to Eating Disorders



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Questions?
(Particularly about ED treatment in rural areas)

Thank You!

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Hospital and Clinic Teams