

Guidelines for the Prevention & Treatment of Pediatric Obesity

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Title: Prevention and Treatment of Pediatric Obesity: An Endocrine Society Clinical Practice Guideline Based on Expert Opinion

Short Title: Guidelines for the Prevention & Treatment of Pediatric Obesity

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Abstract

Objective: To formulate practice guidelines for the treatment and prevention of pediatric obesity.

Participants: An Endocrine Society-appointed Task Force of experts, methodologist, and medical writer.

Evidence: This evidence-based guideline was developed using the Grading of Recommendations, Assessment, Development, and Evaluation system to describe the strength of recommendations and the quality of evidence, which was generally low or very low.

Consensus Process: Two group meetings, several conference calls, and e-mail communications enabled consensus. Committees and members of The Endocrine Society and Lawson Wilkins Pediatric Endocrine Society reviewed and commented on the preliminary drafts.

Conclusions: We recommend: defining overweight as BMI $\geq 85^{\text{th}}$ percentile but $< 95^{\text{th}}$ percentile, and obesity as BMI $\geq 95^{\text{th}}$ percentile; against routine endocrine studies unless the height velocity is attenuated or inappropriate for the family background or stage of puberty; referring to a geneticist if there is evidence of a genetic syndrome; evaluating for obesity-associated co-morbidities in children with BMI $\geq 85^{\text{th}}$ percentile; prescribing and supporting intensive lifestyle (dietary, physical activity, and behavioral) modification as the prerequisite for any treatment.

We suggest that pharmacotherapy (in combination with lifestyle modification) be considered in 1) obese children only after failure of a formal program of intensive lifestyle modification and in 2) overweight children only if severe co-morbidities persist despite intensive lifestyle modification, particularly in children with a strong family history of type 2 diabetes or premature cardiovascular disease. Pharmacotherapy should be provided only by clinicians who are experienced in the use of anti-obesity agents and aware of the potential for adverse reactions. We suggest bariatric surgery for adolescents with BMI > 50 , or > 40 with severe co-morbidities in whom lifestyle modifications and/or pharmacotherapy have failed. Candidates for surgery and their families must be psychologically stable and capable of adhering to lifestyle modifications.

Access to experienced surgeons and sophisticated multidisciplinary teams who assess the benefits and risks of surgery is obligatory,.

Clinicians should emphasize the prevention of obesity by recommending breast-feeding of infants for at least 6 months and advocating that schools provide for 60 minutes of moderate to vigorous daily exercise in all grades. We suggest that clinicians educate children and parents through anticipatory guidance about healthy dietary and activity habits; and advocate for restricting the availability of unhealthy food choices in schools, policies to ban advertising unhealthy food choices to children, and community redesign to maximize opportunities for safe walking and bike riding to school, athletic activities, and neighborhood shopping.

Method of development of evidence-based guidelines

The Clinical Guidelines Subcommittee of The Endocrine Society identified pediatric obesity as a priority area requiring practice guidelines and appointed a Task Force to formulate evidence-based recommendations.

Accordingly, the purpose of these guidelines is to summarize information concerning:

- The seriousness of pediatric obesity and overweight.
- The diagnostic criteria.
- The available treatments and when to apply them.
- The available measures to prevent overweight and obesity.

The Task Force elected to use the GRADE method proposed by an international group with expertise in development and implementation of evidence-based guidelines (1). The Task Force used systematic reviews of available evidence to inform its key recommendations, and consistent language and graphical descriptions of both the strength of recommendations and the quality of evidence. The strength of a recommendation is indicated by the number **1** (strong recommendation, associated with the phrase “we recommend”) or **2** (weak recommendation, associated with the phrase “we suggest”). The quality of the evidence is indicated by cross-filled circles, such that ⊕○○○ denotes very low quality, ⊕⊕○○ low quality, ⊕⊕⊕○ moderate quality, and ⊕⊕⊕⊕ high quality evidence. A detailed description of this grading scheme has been published elsewhere (2).

Each recommendation is followed by a description of the evidence, the values that panelists considered in making the recommendation, and in some instances remarks, a section in

which panelists offer either technical comments or caveats. These remarks usually reflect unsystematic observations and should be considered suggestions.

Making strong recommendations with low quality evidence

For many issues for which the evidence base is of low or very low quality, the Task Force, nonetheless, elected to make strong recommendations. As noted by Guyatt and colleagues, “The strength of any recommendation depends on the following two factors: the tradeoff between the benefits and risks and burdens; and the quality of the evidence regarding treatment effect...” (3). A category 1 (strong) recommendation can be made when “the tradeoff is clear enough that most patients, despite differences in values, would make the same choice....” A category 2 (weak) recommendation is made when “the tradeoff is less clear, and individual patient values will likely lead to different choices...”(3).

We concur with the statement of Snow and colleagues: “Clinical practice guidelines are guides only and may not apply to all patients and all clinical situations. Thus, they are not intended to override clinicians’ judgment” (4). Endocrine Society clinical guidelines are valid for 3 years, after which time they are revised.

1.0 The problem with obesity

The objective of interventions in overweight and obese patients is the prevention or amelioration of obesity-related co-morbidities, e.g., glucose intolerance and type 2 diabetes mellitus, metabolic syndrome, dyslipidemia, and hypertension.

The prevalence of obesity, body mass index (BMI) \geq 95th percentile, increased almost four-fold for 6- to 11-year-old children and three-fold for 12- to 19-year-olds between the

surveys of 1963-1970 and 1999-2000 (5). For infants (0-23 months) the increase was from 7.2% in 1976-1980 to 11.6%; for 2- to 5-year-olds, from 5% to 13.9% during this time (6). Although rates vary among different ethnic groups, the overall prevalence of childhood obesity is 17.1% (7). The standards for weight and BMI percentiles are derived from the data of NHANES II (1976-1980) and NHES I (1963-1965) and II (1966-1970), when children were leaner.

The prevalence of overweight (BMI $\geq 85^{\text{th}}$ but $\leq 95^{\text{th}}$ percentile) in the same period increased by 2.5-fold to 37.2% in boys and girls older than 6 years of age. Exact rates vary according to ethnicity(6).

The prevalence of pediatric obesity varies according to ethnicity, with especially high rates among Mexican Americans and African Americans. The role of poverty, although tending to be associated with a higher prevalence of obesity, is inconsistent across ethnicity and gender (8).

1.1 Association with adult disease

This increased prevalence is particularly important as childhood overweight and obesity are predictive of adult overweight and obesity. After adjusting for parental obesity, the odds ratio for a BMI >27.8 for men and >27.3 for women aged 21-29 years increased from 1.3 when children aged 1-2 years had a BMI $>85^{\text{th}}$ percentile to an odds ratio of 17.5 when children aged 15-17 years had a BMI $>85^{\text{th}}$ percentile (9). The results are similar when looking at overweight adults at age 35 years compared with childhood BMIs at the 75^{th} , 85^{th} , and 95^{th} percentiles (10, 11). A BMI $\geq 85^{\text{th}}$ percentile during the pre-school and elementary school years is associated with a significantly increased risk of overweight in adolescents (12).

A major concern regarding the increased prevalence of obesity is its association with cardiovascular risk factors. Autopsy studies show the presence of not only fatty streaks but also

fibrous plaques in the aorta and coronary arteries of obese teenagers (13, 14). The prevalence of cardiovascular risk factors (hypertriglyceridemia, high LDL cholesterol, low HDL cholesterol, hyperinsulinemia, and hypertension) increases with the rise in BMI. For children and adolescents with a BMI between the 85th and 94th percentile, 19% had two or more cardiovascular risk factors and 5% had three or more. When the BMI was $\geq 95^{\text{th}}$ percentile, 39% had two or more risk factors and 18% had three or more. When the BMI was $\geq 99^{\text{th}}$ percentile, 59% had two or more cardiovascular risk factors and 33% had three or more (15).

The presence of cardiovascular risk factors during childhood can lead to an increased incidence of fatal and nonfatal cardiac events in adulthood (16). In this study, a cohort of 276,835 children aged 7-13 years was followed into adult ages of 25-60 years; 10,235 men and 4,318 women suffered either a fatal or a nonfatal cardiovascular event. The risk of a cardiovascular event was associated with the BMI of boys aged 7-13 and girls aged 10-13 years. The relative risk of an event was apparent even with an increase in BMI z-score of just 1 unit. For this Danish population of children a BMI z-score of 0 was equivalent to the 44th percentile on the CDC BMI charts, and a z-score of +1 was equivalent to the 88th percentile. For boys whose BMI z-score was +1 the relative risk of an adult cardiovascular event increased from 1.06 at 7 years of age to 1.17 at 13 years of age. For girls the relative risk increased from the nonsignificant relative risk of 1.02 at 7 years to a significant 1.12 at 13 years. This study illustrates the deleterious effects of even a moderate increase in body weight during childhood and pinpoints the ages between 7 and 13 years as one of the critical periods for intervention and prevention of overweight and obesity (16).

1.2 Pediatric obesity-associated co-morbidities

Concomitant with the greater prevalence of childhood obesity, the prevalence of type 2 diabetes mellitus (T2DM) has increased in children and adolescents (17). T2DM now accounts for 20% of diabetes in children aged 10-19 years (18).

Overweight and obese individuals are at increased risk for dyslipidemia, most commonly a low HDL cholesterol. Lipid abnormalities were found in 12%-17% of overweight and obese children (19).

The overall prevalence of the metabolic syndrome (a constellation of cardiovascular risk factors, comprising abdominal obesity, and two or more of elevated triglycerides, low HDL cholesterol, high blood pressure, increased plasma glucose) (20-22) among 12- to 19-year-olds in the United States is about 4.2% (23). The risk of the metabolic syndrome is nearly 50% in severely obese (BMI \geq 40.6) adolescents (24).

Acanthosis nigricans (AN) is associated with both T2DM and with insulin resistance (IR), but the strongest correlation is with obesity (25). The prevalence of AN is greater in African Americans (51%) than in Caucasians (8%) (25). Skin tags are also associated with T2DM (26).

Obesity, BMI \geq 95th percentile, is associated with hyperandrogenemia and hyperinsulinism in pre- to mid-pubertal girls. Free testosterone concentrations in such girls are higher than in girls with a BMI <85th percentile (27). Taken together, these abnormalities place the obese adolescent girl at risk for polycystic ovary disease (PCOS), which, in turn, is exacerbated by obesity.

Systolic blood pressure correlates with BMI, skinfold thickness, and waist-to-hip ratio in children and adolescents (28). Normal values for blood pressure are standardized to the child's height percentile (29). For children with BMI in the 85th-94th percentile, a blood pressure >95th percentile ranged from 6.6% at 2-5 years to 13.3% at 16-19 years for males; for females the

range was 4.4% at 2-5 years to 16.3% at 16-19 years of age (30). The relative risk for hypertension in obese children (BMI \geq 95th percentile) was 3.26 after three consecutive blood pressure screenings(31).

There is a 10-fold increase in obesity-related glomerulosclerosis among adults (32). Morbidly obese adolescents have also been diagnosed with proteinuria and focal segmental glomerulosclerosis (FSGS), which may progress to end-stage renal disease or remit with weight loss (33).

Obese children are up to six times more likely than lean children to have obstructive sleep apnea (OSA) (34). OSA is independently related to the development of hypertension, cardiovascular diseases, behavioral disorders, poor school performance in children, and poor quality of life in adults (35).

The prevalence of **non-alcoholic fatty liver disease (NAFLD)** is unclear and depends on the detection method. Between 10% and 25% of obese children have elevated transaminases, primarily alanine aminotransferase (ALT); as the degree of obesity increases so does the prevalence of an elevated ALT. Abdominal sonography can detect a fatty liver, which is also associated with a greater BMI, in 52% of obese children. In an autopsy study, the incidence of fatty liver was 38% in obese children 2-19 years of age (36). Morbidly obese children are particularly prone to fatty liver. Obese Hispanic children have a higher incidence of NAFLD. Weight reduction is an effective treatment for NAFLD (37). Although a benign clinical course is typical of NAFLD, it may be associated with increasing fibrosis and, rarely, progression to cirrhosis (38).

Obesity is a major risk factor for gallstones, which were found in 2% of obese children with a BMI >30 compared with an incidence of only 0.6% in non-obese children (39). Additional

risk factors for gallstone development are the metabolic syndrome, hyperinsulinemia, and rapid, significant weight loss (40).

Excess weight is associated with slipped capital femoral epiphysis, genu valga, tibia vara (Blount disease), flat kneecap pressure/pain, flat foot, spondylolesthesis (low back pain), scoliosis, and osteoarthritis (41).

The prevalence of pseudotumor cerebri increases 15-fold with increasing BMI. The risk of intracranial hypertension is not related to the degree of obesity and is increased even in individuals who are just 10% above ideal body weight (41).

Psychosocial problems may become an issue for many overweight and obese children and adolescents (42).

1.3 Remarks

Although not a co-morbidity per se, an earlier onset of pubarche and thelarche is associated with an elevated BMI (43, 44). Whether it is also associated with an earlier onset of menarche is controversial, a question that has been extensively reviewed (45). There is, though, increasing evidence that premature pubarche is linked to the elements of the metabolic syndrome and to polycystic ovarian disease (L Ibáñez, J Dimartino-Nardi, N Potau and P Saenger 2000.

Premature adrenarche – normal variant or forerunner of adult disease? *Endocrine Reviews* 21:671-696) (P Utriainen, J Jääskeläinen, J Romllanen and R Voutilainen 2007. Childhood metabolic syndrome and its components in premature adrenarche. *J Clin Endocr Metab* 92:4282-4285). These children and adolescents require close observation as they mature into early adulthood.

2.0 Diagnosis of overweight and obesity

2.1 Recommendation

We recommend the use of BMI (calculated as weight in kilograms divided by height in meters²), with CDC-derived normative percentiles, as the preferred method for the diagnosis of the overweight or obese child (46). (1|⊕⊕○○)

2.1 Evidence

Although various techniques are available to measure body fat, many are impractical for clinical use. BMI was adopted as the international standard clinical measure of adiposity (47).

An increased BMI is related to morbidity and mortality in adults even if there is imprecision as to the relationship of BMI values to body fat content. Following its widespread use in adults, BMI is now accepted as the standard in children (8). However, the use of BMI in children is more complex than in adults. The BMI standard percentile distribution changes with an individual's age, sex, and, in some populations, ethnicity—thus limiting the utility of the international standard BMI charts for age (48).

2.1 Values and preferences

The Task Force placed a high value on the identification of children and adolescents with high BMI to enable targeting recommended clinical interventions to those individuals and placed a relatively lower value on avoiding the potential psychological and socioeconomic consequences (e.g., labeling and medicalization) of such practice.

2.1 Remarks

Although not ideal, the BMI is the internationally recognized standard for the definition of overweight and obesity, and the CDC standard curves are the most readily available standards for American children. The Task Force elected not to recommend another standard without higher quality evidence.

Stunted populations (49) may have an increased BMI without increased body fat. Taller individuals may have a high BMI even if their body proportions are equivalent to those of shorter individuals (50). In individuals who are not unusually short or tall, a high BMI is likely to predict abnormally high body fat content. However, a normal BMI does not always exclude the presence of increased body fat or increased risk of obesity-associated co-morbidities, (see 2.2 Remarks). Fat tissue and BMI increase naturally with pubertal progression in girls (51), so that matching the value to chronological age may be misleading if the child progresses through puberty outside the average age range.

Much research has focused on using the waist circumference or waist-to-height ratio as a marker of obesity (52-56), as well as an additional marker for IR (23, 52, 57-59). This approach has the advantage of taking into account body fat distribution and the greater cardiovascular risk associated with visceral fat (60). Waist circumference standards for American children of various ethnic groups are available (61). Waist circumference should be recorded because future research may prove this measure to better reflect fatness or morbidity.

BMI (weight in kilograms divided by height in meters²) is plotted according to the standardized CDC growth charts for the United States (46). Waist circumference is measured at the level of the iliac crest and interpreted according to age, sex, and racial standards (61).

2.2 Recommendation

We recommend that a child be diagnosed as overweight if the BMI is $\geq 85^{\text{th}}$ percentile but $< 95^{\text{th}}$ percentile and as obese if the BMI is $\geq 95^{\text{th}}$ percentile for age and sex. (1⊕○○○○)

2.2 Evidence

Adults experience increased morbidity and mortality when the BMI rises over 30, which is about the 95th percentile of BMI at 19½ years of age (62). Clinicians caring for children can use curves generated by tracing the progression of BMI values for age back from the various percentiles at this older age to determine, in an age-adjusted manner, what BMI is equivalent to 25 or 30 for an adult. Although not derived from pediatric data, some experts consider these cutoffs relevant indicators of health and morbidity (63, 64), but others do not (65).

2.2 Remarks

The BMI may be particularly imprecise in children <4 years of age (see section 2.1 Remarks). Thus, clinicians may consider resorting to the available weight and height percentile charts. An increase in weight percentile that is out of proportion to the increase in height percentile should be a warning sign.

Data indicate that individuals of Asian and Native American ethnicity have a greater susceptibility to obesity-associated co-morbidities at a lower BMI than other ethnicities. An adult BMI of ≥ 23 was associated with an increased risk of obesity-associated co-morbidities and a BMI ≥ 27.5 represented high risk (66, 67). There are also differences in the BMI and waist circumference cut-points for Mexican Americans, African Americans, and whites in relation to the likelihood of having a cardiovascular disease risk factor (68). These discrepancies may also apply to children of different ethnic groups although no long-term data are available as yet.

Currently, there are insufficient data to recommend modifying the definition of either overweight or obesity in Asian or Native American children or adolescents.

2.3 Recommendation

We recommend against a routine laboratory evaluation for endocrine causes of obesity in obese children or early- to mid-pubertal obese adolescents unless the child's height velocity, assessed in relation to stage of puberty and family background, is attenuated. (1|⊕⊕○○)

2.3 Evidence

Growth hormone deficiency or other forms of hypopituitarism, hypothyroidism, Cushing disease or syndrome, or pseudohypoparathyroidism are associated with increased BMI, but stature and height velocity are decreased, unless there is pubertal acceleration in height velocity. By contrast, stature and height velocity are usually increased with exogenous obesity. Testing for endocrine disorders is unlikely to be useful unless the patient is short in relation to the family's background or is showing a deceleration in height velocity (69).

There are two uncommon circumstances in which the above rule may seem to be invalid. First is an adrenal tumor with concomitant hypersecretion of androgens and cortisol, but a careful physical examination should reveal signs of virilization. Second is the condition termed "growth without growth hormone" (70). It is rare in patients with idiopathic isolated growth hormone deficiency or idiopathic combined pituitary hormone deficiencies. It can occur in patients whose panhypopituitarism follows removal of a craniopharyngioma or other suprasellar tumor or who have some other CNS abnormality. In 2003, five additional patients with this condition and panhypopituitarism were reported (71). These particular patients were not obese and presented with such features of pituitary deficiency as small penis and delayed puberty.

Hypothalamic obesity may be a related condition as it is associated with CNS insult with hypothalamic damage which may lead to decreased energy expenditure and excessive caloric intake. It is not amenable to most lifestyle alterations (72). (See Section 3.2, discussion of octreotide.)

2.3 Remarks

Deceleration in height velocity can be demonstrated either by using a height velocity curve normalized for age and/or stage of puberty or by observing that the patient is crossing major height percentile lines downward on the standardized height attainment charts (46). Because these height attainment charts are standardized for age and not for stage of puberty, they can be misleading in children with delayed or precocious puberty. Height attainment and velocity charts that have the growth patterns not only of average-maturing children but also of early-maturing and late-maturing children are available (73). Early- and late-maturing children represented on these curves are within the normal variation for the onset of puberty. Children demonstrating either true precocious or true delayed puberty deserve a full evaluation.

2.4 Recommendations

2.4.a *We recommend referral to a geneticist for children whose obesity has a syndromic etiology, especially in the presence of neurodevelopmental abnormalities. (1|⊕○○○)*

2.4.b *We suggest that parents of children who have inexorably gained weight from early infancy and have risen above the 97th percentile for weight by 3 years of age be informed of the availability of MC4R genetic testing. However, the test is positive in only 2%-4% of such*

patients who are above the 97th percentile for weight (69) and currently will not alter treatment.

(2|⊕○○○)

2.4 Evidence

The more common genetic syndromes associated with obesity are also associated with neurodevelopmental abnormalities (Online Mendelian Inheritance in Man, available at: <http://www.ncbi.nlm.nih.gov/omim/>). Children with the various forms of Turner syndrome or SHOX deletions (74), however, are most often developmentally normal and may have few, if any, stigmata. While it is attractive to postulate that human obesity is attributable to a single mutation or group of mutations in candidate genes with prominent roles in the appetite and fuel metabolic pathways, it is likely that most obesity is caused by the accumulation of polymorphisms in multiple genes with relatively small effects on weight gain or quantitative trait loci (75, 76). This concept is supported by a genome-wide scan in participants in the Framingham Heart Study demonstrating that a common polymorphism in a cholesterol-regulating gene is associated with obesity (77).

2.4. Values and preferences

In making the recommendation for referral to a geneticist, the Task Force placed a higher value on the identification of genetic disorders that will provide an explanation of the individual's overweight problem, offer prognostic implications, and spare the family from feelings of guilt that often occur when traditional weight loss interventions are unsuccessful. We also placed a lower value on avoiding the downsides of genetic consultation, namely cost, labeling, and false positive results of testing, when this referral is carefully limited to individuals with a syndromic etiology of obesity.

2.5 Recommendation

We recommend that children with a BMI $\geq 85^{\text{th}}$ percentile be evaluated for associated co-morbidities and complications (see Table 1 and Figure 1). (1|⊕○○○)

2.5 Evidence

In Section 1.1 we discussed the evidence for the increased prevalence of co-morbidities that are associated with a BMI $\geq 85^{\text{th}}$ percentile. Of most concern are the cardiovascular risk factors (hypertriglyceridemia, increased LDL cholesterol, decreased HDL cholesterol, increased fasting blood glucose, and hypertension) predictive of future cardiovascular disease or T2DM or both (see also Section 1.1). Thus, a large enough number of overweight children with a cardiovascular risk factor exist to warrant screening them, as well as obese children. Measurement of ALT is of more importance as the BMI increases into the obese range (discussed in Section 1.2).

2.5. Values and preferences

We placed a relatively high value on the identification of weight-related complications and co-morbidities as this simultaneously focuses our limited treatment resources on those who, as a group, could accrue the greatest potential benefits of treatment and allows the identification and management of common and potentially important conditions. This recommendation places a lower value on avoiding medicalization and its costs and consequences.

2.5 Remarks

The medical history should assess the risk factors for the development of pediatric obesity including maternal diabetes, small for gestational age, large for gestational age, parental obesity (maternal is more important than paternal), early adiposity rebound (9, 79, 80), maternal weight

gain during pregnancy, breast-feeding duration, weight of siblings and more distant relatives, possible consanguinity, as well as all other aspects of a standard pediatric history.

The medical history should also assess the presence of snoring and other manifestations of sleep apnea; polyuria, polydipsia, or weight loss (diabetes); and, in pubertal girls, acne, hirsutism (including the recent use of hair removal techniques that would mask the degree of hirsutism at the time of the examination), and onset and frequency of menses (PCOS). Note the use of antipsychotics associated with weight gain such as clozapine, risperidone, olanzapine, and quetiapine.

Although the various techniques assessing dietary intake are unreliable and subject to error (81), it is still important to estimate the type and quantity of beverage intake, frequency of dining out and where, and frequency and type of snacks, among other dietary issues. The efficacy of food frequency and activity questionnaires is discussed in a 2007 review (82).

Activity history includes the duration, the frequency, and an estimate of the degree of difficulty of exercise performed during school and at-home days, including participation in sports teams or other activities, walking to school and stores, etc. Estimates of screen time (i.e., time in front of a computer, playing video games, or viewing TV) per day may be useful. An environmental history includes safety of parks and neighborhoods and availability of playground equipment, gyms, and pools. All of this information is needed to develop an action plan.

Physical finding should include (a) waist circumference at the level of the iliac crest and interpreted according to age, sex, and racial standards (61), (b) blood pressure, using height percentile-normalized blood pressure tables to interpret the findings (29), (c) acanthosis nigricans and skin tags, (d) severe acne and hirsutism in pubertal age girls, (e) tenderness and range of motion of the knee, leg, or foot, and (f) peripheral edema. Physical findings associated with

syndromic obesity should be identified, particularly if there is a neurodevelopmental abnormality.

Suggested screening tests are listed in Table 1. Other tests may be pertinent when other findings suggest sleep apnea (a sleep study, EKG, echocardiogram) or PCOS (serum testosterone, sex hormone binding globulin [SHBG], dehydroepiandrosterone sulfate [DHEAS] and 17-hydroxyprogesterone [17OHP], and third generation LH and FSH). Testosterone assays in the range found in women and children may be imprecise (83, 84).

The fasting insulin is an optional test to quantify IR by calculation of HOMA-IR or by noting that the fasting insulin is >2 SD above the mean for the laboratory doing the test. There is developing evidence for the association of IR with the development of T2DM in children (85, 86). Most of these children and adolescents are obese (87). Identification of children with IR has been proposed as a strategy for identifying high risk children for targeted diabetes prevention. The gold standard for measuring IR, the hyperinsulinemic-euglycemic clamp, is labor intensive and expensive and, therefore, not recommended for clinical use. A surrogate marker of IR is the HOMA-IR index, based on the measurement of fasting insulin and glucose with higher levels representing greater degrees of IR ($\text{HOMA-IR} = [\text{fasting insulin in } \mu\text{U/mL} \times \text{fasting glucose in mmol/L}] \text{ divided by } 22.5$) (88). In nondiabetic children, HOMA-IR correlates as high as 0.91 with clamp measures (89, 90). Based on a study of 1,802 adolescents aged 12-19 years from the 1999-2002 NHANES, a HOMA-IR >4.39 was recommended as evidence for IR (87). On the basis of that criterion, roughly 46%-52% of the adolescents with a BMI $\geq 95^{\text{th}}$ percentile were insulin resistant; roughly 11%-16% of those with a BMI $\geq 85^{\text{th}}$ percentile but $<95^{\text{th}}$ percentile were insulin resistant; and only 4% of those with a BMI $<85^{\text{th}}$ percentile were insulin resistant (87). In a study of Mexican children aged 10-14 years very similar results were obtained with 3%, 12%, and 47% of the children insulin resistant at BMIs of $<85^{\text{th}}$ percentile, $\geq 85^{\text{th}}$ but $<95^{\text{th}}$

percentile, and $\geq 95^{\text{th}}$ percentile, respectively (91). In a study of Australian children aged 6-13 years (92), increasing BMI z-score was continuously associated with complications of overweight in children. Even in the healthy weight range there was a continuous linear or curvilinear association between increasing BMI z-score and co-morbid risk factors. But there was a subset of children in whom there was no evidence of hyperinsulinism, even at BMI z-scores of +1 to +3. Conversely, some children had evidence for hyperinsulinism even though their BMI z-scores ranged from -2 to 0.

The use of any index of IR is complicated by concerns about the lack of standardized measures for the quantification of insulin (93), the need to be assured that the blood sample is obtained in a truly fasting state, and by the increase in IR from stage 1 to stage 3 of puberty and then a decrease in IR from stage 3 until adulthood (94).

Assessment of HOMA-IR in overweight obese children and adolescents may represent an important strategy for improving the efficacy of treatment for weight loss and chronic disease prevention (95-97). Further, measurement of fasting serum insulin or of HOMA-IR is not necessary to establish a need for weight control or weight loss, especially as the expense of the test limits its potential for broad use in IR screening.

3.0 Treatment of obesity

3.1 Lifestyle recommendations: general considerations

3.1.0 Recommendations

We recommend that clinicians prescribe and support intensive lifestyle (dietary, physical activity, and behavioral) modification to the entire family and to the patient, in an age-

appropriate manner, and as the prerequisite for all overweight and obesity treatments for children and adolescents. (1|⊕○○○)

3.1.0 Evidence

Successful weight management, through lifestyle interventions, confers important intermediate-term health benefits to adults such as reducing the incidence of T2DM (98) and improving cardiovascular fitness (99). Weight management programs in overweight children have also improved body composition and metabolic parameters (100, 101).

Unfortunately, it is commonly thought that lifestyle modification is not efficacious. Weight loss may not occur or, in spite of initial success, weight regain often begins after the active phase of the program has ended (102, 103).

A factor contributing to weight regain may be lack of a continued exercise program. The odds for weight regain are two-fold greater in those patients who are sedentary (102). The importance of exercise in maintaining much of a person's weight loss was one of the conclusions of a meta-analysis of long-term (3- to 5-year) weight maintenance studies (27.2% weight loss retention in the low exercise group and 53.8% weight loss retention in the high exercise group) (104). This meta-analysis also found that a weight loss of >20 kg in adults was associated with a greater degree of weight loss retention as compared with adults losing <10 kg. Additional factors associated with weight regain include Mexican American ethnicity versus non-Hispanic whites (odds ratio 2.0) and losing >20% of body weight versus <15% (odds ratio 2.8) (102). Changes in hormone production and action may also play important roles (95). In socioeconomically disadvantaged families, weight control may not be an important priority when balanced against other problems they face.

Although the long-term outlook may appear bleak, some studies report long-term success in a significant subgroup of patients. The results of population surveys indicate that 25% of adults who had lost >10% of their body weight maintained their weight losses for more than 5 years (105). In another population-based study of individuals who had completed a commercial weight loss program, 18.8% maintained a weight loss of $\geq 10\%$ for 5 years while 42.6% maintained a weight loss of $\geq 5\%$ for 5 years; 19.4% maintained their weights within 5 pounds of their original goals (106). In both papers the authors opined that patients who seek out a health care setting have already tried and failed to lose weight through other means and may represent a more refractory population.

Other factors, besides high levels of physical activity, associated with successful weight maintenance included continued reduced caloric intake, reduced fat intake, and reduced fast food consumption (107). Certain psychological traits also are associated with successful weight loss maintenance (108, 109).

The success of lifestyle modification, at least in a reasonable portion of adults, has prompted its endorsement in guidelines from the U.S. Preventive Services Task Force (USPSTF). That group “recommends that clinicians screen all adult patients for obesity and offer intensive counseling and behavioral interventions to promote sustained weight loss...” (110). The USPSTF emphasizes the concept of intensive counseling, defined as at least one “person-to-person (individual or group) session per month for at least the first 3 months of the intervention” (110).

Emphasis should be on providing an intensive lifestyle modification program. This was most clearly shown by the results of a T2DM prevention trial in which one comparison was between the “standard lifestyle modification”—in which patients received the traditional written

instructions about diet and exercise as well as an annual session discussing the importance of a healthy lifestyle—and an intensive plan covering diet, exercise, and behavior modification for the first 24 weeks followed by monthly follow-up sessions. The initial goal was a 7% weight loss. The average weight loss in the standard lifestyle modification group was 0.1 kg and in the intensive modification group was 5.6 kg ($p < 0.001$). Significant weight loss was maintained over the 4 years of follow-up. The cumulative incidence of diabetes after 3 years was 28.9% in the standard modification group and only 14.4% in the intensive modification group (111).

Although good quality pediatric and adolescent data are scarce (112), there is sufficient evidence that intensive lifestyle modification programs, as in adults, can be an effective tool for pediatric weight control (101, 113). Further, implementation of a formal maintenance program after the treatment phase is completed can be of added importance in maintaining achieved weight loss (113). This fits into a concept of obesity as a chronic disease (114).

A meta-analysis of randomized pediatric trials, commissioned by this Task Force, of combined lifestyle interventions (diet and exercise) for treating obesity showed a modest but significant effect on obesity (equivalent to a decrease in BMI of 1.5, $p < 0.00001$) when these interventions targeted family involvement. There was a decrease in BMI of 0.4 when parents were not specifically included and the effect on weight loss was not significant ($p = 0.13$). While the indirect comparison of effects across studies indicates that treatment is significantly more effective when it targets the family (p value for the interaction was 0.04), the direct comparisons (within the same trial) were inconsistent across trials and showed no significant difference (115). These results suggest involving the family when delivering combined lifestyle interventions. Another meta-analysis that looked specifically at the effects of family-behavioral therapy produced parallel results (116).

An additional meta-analysis of randomized controlled trials of lifestyle interventions, but without an analysis of family involvement, also found moderate positive effects from the interventions when compared with no treatment, wait-list, or information-only controls. These effects persisted for an average follow-up period of 15 months (117). Although there was overlap with the meta-analysis we commissioned, each study contained reports not covered by the other.

An evidence-based position statement of the American Dietetic Association (ADA) supports the utility of family-based lifestyle interventions in children and of similar multicomponent programs for adolescents (118). These recommendations are consistent with the conclusions of an evidence-based review of pharmacological interventions for childhood obesity that highlighted the importance of concomitant intensive lifestyle interventions—dietary, exercise, and family counseling (95)—as well as by a combined CDC and AMA expert committee (119).

3.1.0 Values and preferences

In making this recommendation (and other recommendations associated with specific lifestyle choices), the Task Force placed a relatively higher value on promoting lifestyle choices with their potential wide-reaching benefits and safety and placed a relatively lower value on avoiding the costs of implementation of lifestyle interventions with their potentially limited impact.

Although physicians generally strive to cure the great majority of their patients and may view a long-term success rate of 25% with despair, we should not retreat into a state of therapeutic nihilism. We are at a stage where we must treat overweight and obese patients, accept that perhaps only 25% may respond, but refine our techniques so that lifestyle modification will be effective in an increasing percentage of patients.

3.1.0 Remarks

Weight loss should be encouraged in patients with severe obesity and significant co-morbidities. In this regard, a decrease in BMI of 1.5, as reported in the meta-analysis commissioned by the Task Force, may seem trivial, but, if it is maintained over a longer term, overweight or minimally obese growing children and adolescents without co-morbidities may benefit by simply maintaining weight; BMI will decline as linear growth proceeds, and lifestyle modification may reduce fat mass, increase lean body mass, and improve cardiovascular fitness (120). In the more severely obese or in physically mature patients, moderate weight loss of only 7% was associated with a decrease in the incidence of T2DM (111). This may be a more realistic goal for the severely obese. (See Section 3.1.0 Evidence.) Well-designed, randomized controlled studies of large numbers of patients, employing intensive lifestyle intervention and follow-up maintenance programs, are clearly needed to develop improved techniques. Some of the factors that should be addressed in such studies are described (112).

Successful lifestyle intervention and preventive measures are labor intensive, and even more so if there is a post-treatment maintenance program. The time spent in delivering intensive lifestyle interventions—direct contact with the patient and family (we cannot overemphasize the importance of family involvement) at least once a month for the first 3 months and comprising dietary and nutritional education, a physical activity prescription, and behavioral therapy—is poorly reimbursed, and thus there may be a disincentive to provide these services (121 [pp 221-227]).

3.1.1 Dietary recommendations

3.1.1.a *We recommend that clinicians prescribe and support healthy eating habits such as*

- *avoiding the consumption of calorie-dense, nutrient-poor foods (e.g., sweetened beverages, sports drinks, fruit drinks and juices, most “fast food,” and calorie-dense snacks) (1|⊕⊕○○)*

3.1.1.b *We suggest that clinicians prescribe and support*

- *controlling caloric intake through portion control in accordance with the Guidelines of the American Academy of Pediatrics*

[<http://pediatrics.aappublications.org/cgi/reprint/117/2/544>]. (2|⊕○○○)

- *reducing saturated dietary fat intake for children older than 2 years of age (2|⊕⊕○○)*
- *increasing the intake of dietary fiber, fruits, and vegetables (2|⊕○○○)*
- *eating timely, regular meals, particularly breakfast, and avoiding constant “grazing” during the day, especially after school (2|⊕○○○)*

3.1.1 Evidence

Excessive intake of low-nutrient, calorie-dense, high-fat food, and sugar-sweetened beverages, is a risk factor for obesity (122-124). Since 1965 teens have doubled their consumption of sugared soft drinks and fruit-flavored beverages. The average adolescent boy now consumes 50 ounces of such beverages per day (125). Reducing consumption of sugared beverages (e.g., soda, juices and fruit drinks, and sports drinks) (126, 127) may be an effective way to reduce ingested calories (128, 129). School-based interventions can reduce soda consumption and reduce weight in students at the highest BMI percentiles (130, 131).

Increased portion size parallels the increase in obesity (132, 133). Texas elementary schools are imparting portion control education using educational materials available from the Texas Department of Agriculture (<http://www.squaremeals.org/>).

Decreasing caloric intake by consuming more fruits and vegetables instead of dietary fat can decrease the risk of developing obesity and T2DM (131, 134). Most children do not meet the dietary recommendations needed to maintain a healthy weight and eat less than half the recommended five fruits and vegetables a day (135, 136). Inadequate consumption of dietary fiber may contribute to excessive weight gain (137).

Fast food accounted for only 2% of children's total caloric intake in 1977-1978, but the proportion increased to 10% by 1994-1996 (138). Consumption of food prepared away from home increased from 18% of total calories in 1977-1978 to 32% in 1994-1996 (139). Overweight children are more likely to report eating fast food and to attend a school with vending machines selling chips and candy (140). Vending machine foods are sold separate from the National School Lunch Program and are currently exempt from the Dietary Guidelines; these food sales represent an increasing percentage of total school food sales (141).

Overweight children and adolescents are now more likely to skip breakfast and consume a few large meals per day (142, 143) than their leaner counterparts who are more likely to consume smaller, more frequent meals (144-146). Eating breakfast reduces snacking throughout the remainder of the day (147). One fourth to one third of the energy intake of adolescents is from snacks (148). As snacks tend to be higher in calorie density than meals (149), frequent snacking is associated with high intake of fat, sugar, and calories (150, 151) and with overweight among children (152). Intervention studies to confirm a relationship between meal and snack patterns and overweight are needed.

3.1.1 Values and preferences

See description of values and preferences at 3.1.0.

3.1.1 Remarks

Many different diets have been proposed for weight loss. Currently there is debate about whether a low-fat (usually 30% of calories as fat) or a low-carbohydrate diet is more efficacious. A meta-analysis of randomized controlled studies in adults showed that the low-carbohydrate diet resulted in moderately greater weight loss by 6 months but not after 12 months (153). At the present time there is insufficient pediatric evidence to warrant recommending any one hypocaloric diet over another. But caution should be exercised when using unbalanced hypocaloric diets that may be deficient in essential vitamins and minerals.

3.1.2 Physical activity recommendations

3.1.2.1 *We recommend that clinicians prescribe and support 60 minutes of daily moderate to vigorous physical activity. (1|⊕⊕○○)*

3.1.2.1 Evidence

In the absence of caloric restriction, moderate exercise does not generally cause weight loss. However, in combination with decreased caloric intake, exercise can achieve significant weight loss. Studies performed in the school setting have shown the beneficial effects of exercise in children and youth (154-158). The beneficial effects of both aerobic exercise and resistance training can be short-lived and exercise must be sustained. Time spent in daily vigorous exercise

above in excess of 60 minutes per day provides additional reduction in cardiovascular risk factors (159).

Physical fitness, even without weight loss, may offer some health benefits. Improvement in cardiovascular fitness was associated, in young adults, with improvement in cardiovascular disease risk factors over a 7-year period (160).

School-based interventions have focused on reducing obesity rates (161-171). The Cardiovascular Health in Children (CHIC-I) study was able to improve physiologic outcomes by decreasing body fat and cholesterol concentrations (165). The CHIC-II study was effective in reducing body fat and blood pressure in middle school youth (168). One of the main reasons the short-term CHIC interventions were successful in affecting physiologic variables may be the increased time spent in actual moderate to vigorous physical activity in school (20 minutes in elementary schools and 30 minutes in middle schools). Both school design and adult supervision for physical activity have been shown to affect the amount of physical activity that 6th – 8th graders engage in during their free time (172). School systems are beginning to sponsor after-school lifetime fitness programs (173). Such pilot programs appear successful in controlling the rate of weight gain (174).

A meta-analysis of physical activity interventions, commissioned by this Task Force, found a moderate treatment effect when the outcome measure reflected body composition (e.g., fat percentage) ($p=0.00001$) and no effect when the outcome was BMI; the difference in results was highly significant ($p=0.002$). This would support the concept that exercise may affect cardiovascular risk factors by improving insulin sensitivity and adiposity, and by increasing lean body mass without affecting total body weight. However, there was no significant difference in responses in the six trials reporting both measures (115). This lack of a significant difference in

responses may be due to there being no true difference, to chance (imprecision, i.e., there is a difference but the studies or the effects were not large enough to be detected [a type II error]), or to bias. This probably reflects reporting bias in which the significant outcome measure was more likely to be reported than an insignificant finding, such that when studies reported both measures the effect was similar.

3.1.2.1 Values and preferences

See description of values and preferences at 3.1.0.

3.1.2.1 Remarks

Although current recommendations state that schoolchildren (who spend about half their waking hours in school) should receive a minimum of 30 minutes of moderately vigorous physical activity each school day, only 8% of elementary schools and roughly 6% of junior and senior high schools provide physical education for the entire school year for all students and for all grades (121 [pp 237-284]). Furthermore, only 10%-30% of that time is spent in moderate to vigorous exercise. The limited support for physical education in the schools is unfortunate because exercise has been linked not only to cardiovascular benefits in children (175) but also to improvements in cognitive function and concentration (176, 177). School units on health and hygiene, in which children are taught about nutrition and good health habits, have atrophied. An average of only 9 hours is now spent on health education during the school year, even though pilot studies have demonstrated the efficacy of curricula designed to reduce television-viewing time (121 [pp 237-318]).

Moderate to vigorous exercise is defined as causing “some increase in breathing and heart rate usually associated (in a healthy person) with brisk walking, dancing, swimming, or

cycling on flat terrain.” In exercise physiology terms, the energy expended is ≥ 3 METS (metabolic equivalents) (178).

As discussed under Values and Remarks in Section 3.1.0, studies using either weight loss or the BMI as an endpoint may miss the positive effects of exercise, such as shifting body composition from fat to muscle. The net weight loss under such circumstances may be nil.

3.1.2.2 *We suggest that clinicians prescribe and support a decrease in time spent in sedentary activities, such as watching television, playing video games, or using computers for recreation. Screen time should be limited to 1-2 hours per day, according to the American Academy of Pediatrics (179). (2|⊕○○○)*

3.1.2.2 Evidence

Activities involving TV, videotapes, video games, and computers average 5½ hours per day (180-182). An intervention to reduce TV viewing and meals eaten in front of the TV was successful in a comparatively high-socioeconomic-status population of ethnically homogeneous schoolchildren (183). Results of the meta-analysis, commissioned by the Task Force, of the three randomized trials of interventions focused on reducing sedentary activity were imprecise (i.e., both consistent with favorable and unfavorable impact on obesity outcomes) (115).

3.1.2. Values and preferences

See description of values and preferences at 3.1.0.

3.1.3 Psychosocial recommendations

3.1.3.a *We suggest that clinicians educate parents about the need for healthy rearing patterns related to diet and activity. Examples include parental modeling of healthy habits, avoidance of overly strict dieting, setting limits of acceptable behaviors, and avoidance of using food as a reward or punishment. (2|⊕○○○)*

3.1.3.b *We suggest that clinicians probe for and diagnose unhealthy intra-family communication patterns and support rearing patterns that seek to enhance the child's self-esteem. (2|⊕○○○)*

3.1.3 Evidence

The importance of involving the whole family, and not just the child, in treatment interventions is discussed in section 3.1.0.

The role of parent-child interactions and parenting style in the development of unhealthy lifestyle habits is a subject of investigation (184, 185). An additional factor to overcome before any intervention may be parents' inability to recognize that their child is overweight, particularly for the preschool child (186).

3.1.3. Remarks

It is important to remember that clinician interactions with the family and that all educational materials should be culturally sensitive and in the language best understood by the family.

3.2 Pharmacotherapy recommendations

3.2.a *We suggest that pharmacotherapy (in combination with lifestyle modification) be considered if a formal program of intensive lifestyle modification has failed to limit weight gain or to mollify co-morbidities in obese children. Overweight children should not be treated with pharmacotherapeutic agents unless significant, severe co-morbidities persist despite intensive*

lifestyle modification. In these children, a strong family history of type 2 diabetes mellitus or cardiovascular risk factors strengthens the case for pharmacotherapy. (2|⊕○○○)

3.2.b We suggest that pharmacotherapy be offered only by clinicians who are experienced in the use of anti-obesity agents and are aware of the potential for adverse reactions. (2|⊕○○○)

3.2 Evidence

The utility of pharmacotherapy in adolescents has been reviewed (95, 187), and the use of medication to treat severe obesity can be an additional treatment modality (173, 188-190). Several limitations preclude physicians from early implementation of drug therapies. These include a) the lack of FDA approval for use in preadolescents and younger adolescents; b) reduced efficacy over time, with a plateau after 6 months of treatment due to reduced energy expenditure offsetting the decrease in energy intake—an effect also noted with hypocaloric diets (191); c) the existence of a limited number of well-controlled studies of the safety and efficacy of pharmacologic intervention in obese children; and d) the need to weigh the relative risk of severe adverse events in children against the long-term potential for obesity-related morbidity and mortality. Despite these concerns, the negative health impact of childhood obesity may justify long-term medication but only in combination with lifestyle modification (95, 173, 188-190).

Three pharmacotherapeutic agents—sibutramine, a non-selective reuptake inhibitor appetite suppressant that is most potent for serotonin and norepinephrine, but also blocks dopamine reuptake (192, 193); orlistat, which specifically inhibits intestinal lipase and can reduce fat and cholesterol absorption by approximately 30% (194); and metformin (not FDA approved for the treatment of obesity)—are most commonly used at present. Although metformin reduces hepatic glucose production and plasma insulin, inhibits lipogenesis, increases

peripheral insulin sensitivity, and may reduce appetite by increasing levels of glucagon-like peptide (95), its mechanism of action on weight is unresolved. Only sibutramine (for children >16 years of age) and orlistat (for children >12 years of age) are FDA approved for the treatment of obesity in adolescents (Table 2).

The meta-analysis commissioned by the Task Force (115) showed that sibutramine demonstrated the most effect with a decrease in BMI of 2.4 after 6 months. This effect was statistically significant but patients receiving sibutramine had greater increase in blood pressure and pulse rate than placebo-treated patients. Orlistat was associated with a significant fall in BMI of 0.7, but treatment was associated with increased rates of gastrointestinal side effects including abdominal discomfort, pain and steatorrhea. Side effects are usually mild to moderate and generally decrease in frequency with continued treatment; this decrease may result from patients learning to consume less dietary fat to avoid these side effects. Orlistat must be taken with each meal, thus reducing its utility in children who often are in school during lunchtime.

Additional medications used but not FDA approved for the treatment of obesity

Metformin decreased BMI slightly but significantly in each of the three studies analyzed, but the overall effect did not reach statistical significance in the meta-analysis (115). This outcome may, in part, reflect differences in study design, as one of the studies (199) did not include dietary restrictions. Metformin may be useful in combating the weight gain observed in children taking atypical psychotropic medications, e.g., clozapine, olanzapine, risperidone, quetiapine, aripipazole, and valproate (207). However, cessation of metformin therapy may lead to a rebound hyperinsulinemia and rapid weight gain, whether or not the offending psychotropic medication is continued. Although not approved by the FDA for this indication, metformin has

been successfully used to treat PCOS with and without concomitant obesity and insulin insensitivity (208, 209). Metformin is approved for the treatment of T2DM in children ≥ 10 years of age.

Although not FDA-approved for the treatment of obesity, the FDA-approved product labelling for growth hormone presents data indicating that growth hormone treatment of children with Prader-Willi syndrome decreases body fat percentage and increases lean body mass (210). Growth hormone seemed to be of particular benefit when started before 18 months of age (211). A summary of the benefits and risks of growth hormone treatment of Prader-Willi syndrome was published in 2008 (206). Despite these encouraging preliminary results in Prader-Willi syndrome, a review of the effects of growth hormone treatment in adult obesity failed to reveal any consistent beneficial effects and described the difficulties in assessing body composition (212).

Octreotide acts on the voltage-gated calcium channel of the β -cell coupled to the somatostatin (SSTR₅) receptor (213, 214), and through G_o inhibition limits the opening of this calcium channel, decreasing the magnitude of insulin response to glucose (215). An examination of BMI responses to octreotide in a multivariate analysis in children with hypothalamic obesity demonstrated that insulin hypersecretion with concomitant retention of insulin sensitivity before therapy predicted success (96). In hyperinsulin-secreting obese adults, treatment with octreotide LAR for 6 months resulted in significant weight loss as compared with controls. Greater weight loss correlated with a greater degree of insulin hypersecretion (216).

Mutations of the leptin gene in humans recapitulate the phenotype of the ob/ob leptin-deficient mouse (217). Few such patients (no more than 10 as of 2008) have been described.

These patients manifest hyperphagia from birth, with obesity documentable as early as 6 months of age. The excess insulin, due to vagal overactivity and insulin resistance, may cross-react with the IGF-I receptor to increase height velocity and bone age. Serum leptin is undetectable. The diagnosis is made by unmeasurable serum leptin levels. Leptin therapy in these patients results in extraordinary loss of weight and fat mass (203, 218), along with reduction in hyperphagia, resolution of obesity, induction of puberty, and improvement in T-cell responsiveness (204).

Topiramate is a novel anticonvulsant used in children and adults that blocks voltage-dependent sodium channels, enhances the activity of the GABA_A receptor, and antagonizes a glutamate receptor other than the N-methyl-D-aspartate (NMDA) receptor. Topiramate may induce insulin sensitivity in liver and muscle and directly in adipocytes (219). In a study of adults (205), almost 33% of the subjects dropped out because of adverse events (Table 2). Anorexia and weight loss occur early in 10% to 40% of children treated with topiramate for seizures, an effect that in some cases has led to discontinuing the medication (220). There are no studies of topiramate in childhood obesity. Its use as an anti-obesity agent should be limited as it promotes drowsiness and interferes with cognition, and it should not be used outside of a clinical research study.

Table 2 summarizes the dosage, efficacy, adverse effects, contraindications, and monitoring needs of some of the medications used for the treatment of obesity.

3.2 Values and preferences

The suggestion to limit use of pharmacotherapy in children and adolescents reflects our preference for managing pediatric obesity as a serious lifestyle condition with important life-long consequences and our placing a lower value on achieving short-term success, a higher value on

avoiding drug side effects and costs that accumulate over time, and a higher value on achieving healthy weight through the incorporation of healthy behaviors.

3.2 Remarks

The assessment of drug efficacy presented here is founded only on the ability of medications to reduce BMI or BMI z-score. It must be emphasized that “anti-obesity” drugs may have differential effects on BMI and obesity-associated co-morbidities (95). For example, certain medications (e.g., sibutramine, orlistat) may be more effective for weight loss than for treatment of impaired glucose tolerance, whereas other medications (e.g., metformin) have more potent effects on insulin production and glucose tolerance than on body weight per se. Drug selection should be tailored to the individual patient, with strong attention paid to the family history. The primary objective is to prevent co-morbidities in the obese (BMI $\geq 95^{\text{th}}$ percentile) patient. Most importantly, the benefits of any drug used to treat childhood obesity should clearly outweigh its risks.

In general, children with a BMI $< 95^{\text{th}}$ percentile should not be treated with anti-obesity drugs. Pharmacotherapy for overweight children (BMI $\geq 85^{\text{th}}$ but $< 95^{\text{th}}$ percentile) should be reserved for those with significant, severe co-morbidities who have not responded to lifestyle modification. Although, as mentioned in Section 2.2 Remarks, data suggest that adult Asians (and Native Americans) develop obesity-associated co-morbidities at a lower BMI than do Europeans (66, 67), similar data are not available for children and adolescents, and so we cannot recommend the use of pharmacotherapy at a BMI range differing from the above recommendations.

The use of pharmacotherapeutic agents not yet approved for the treatment of pediatric obesity should be restricted to participation in large, well-controlled clinical studies.

3.3 Bariatric surgery recommendations

3.3.a. *We suggest that bariatric surgery be considered only under the following conditions:*

- 1. The child has attained Tanner 4 or 5 pubertal development and final or near-final adult height.*
- 2. The child has a BMI >50 or has BMI >40 and significant, severe co-morbidities.*
- 3. Severe obesity and co-morbidities persist despite a formal program of lifestyle modification, with or without a trial of pharmacotherapy.*
- 4. Psychological evaluation confirms the stability and competence of the family unit.*
- 5. There is access to an experienced surgeon in a medical center employing a team capable of long-term follow-up of the metabolic and psychosocial needs of the patient and family, and the institution is either participating in a study of the outcome of bariatric surgery or sharing data.*
- 6. The patient demonstrates the ability to adhere to the principles of healthy dietary and activity habits. (2|⊕⊕○○)*

3.3.b *We recommend against bariatric surgery for pre-adolescent children; for pregnant or breast-feeding adolescents, and for those planning to become pregnant within 2 years of surgery; for any patient who has not mastered the principles of healthy dietary and activity habits; for any patient with an unresolved eating disorder, untreated psychiatric disorder, or Prader-Willi syndrome. (1|⊕⊕○○)*

3.3 Evidence

Bariatric procedures for weight loss can be divided into malabsorptive, restrictive, and combination procedures. Purely malabsorptive procedures aim to decrease the functional length or efficiency of the intestinal mucosa through anatomic rearrangement of the intestine. These procedures include the jejunoileal bypass and the biliopancreatic diversion with duodenal switch. Because of the high morbidity and mortality associated with these procedures, they cannot be recommended for use in children.

Laparoscopic adjustable gastric banding (LAGB) is a wholly restrictive procedure. It utilizes a prosthetic band to encircle and compartmentalize the proximal stomach into a small pouch and a large remnant (221). The Roux-en-Y gastric bypass (RYGB) is a combination procedure (221). It is a modification of gastric bypass and has become the most commonly performed bariatric surgical procedure. It involves dividing the stomach to create a small (15- to 30-mL) stomach pouch into which a segment of jejunum approximately 15 to 60 cm inferior to the ligament of Treitz is inserted, while the proximal portion of the jejunum that drains the bypassed lower stomach and duodenum is reanastomosed 75 to 150 cm inferior to the gastrojejunostomy (221). This procedure combines the restrictive nature of gastrectomy with the consequences of dumping physiology as a negative conditioning response when high-calorie liquid meals are ingested. Although the LAGB procedure is considered safer than RYGB, the FDA has not yet approved LAGB for use in adolescents.

Bariatric surgery is an effective weight loss treatment for adults with a BMI >40, and there are sufficient data to calculate complication risks for adults (222). Similar data for adolescents are scarce. A health technology assessment for the Washington State Health Care Authority reviewed the data from 17 studies comprising 553 morbidly obese adolescent patients

(223). A clinically significant weight loss was defined as a loss of 7% of body weight. This amount corresponded to a decrease in BMI of 4 units for patients undergoing RYGB (period of follow-up ranged from 1 to 6.3 years) and a decrease of 3.5 units for patients undergoing LAGB (period of follow-up ranged from 1.7 to 3.3 years). The conclusions reached in this assessment, mostly based on weak evidence, were that a) both procedures resulted in clinically significant weight loss; b) LAGB resolved the co-morbid conditions of diabetes and hypertension whereas RYGB resolved hypertension (there were insufficient data to rate the resolution of other co-morbidities); c) the safety profile (moderate evidence) for LAGB after a follow-up period of 1 to 85 months revealed no operative or postoperative deaths; 26 of 328 patients required re-operation to correct complications (band slippage, intragastric migration, and port/tubing problems); d) the safety profile (moderate evidence) for RYGB after a follow-up period of 2 weeks to 6 years revealed a combination of mild (slight malnutrition) and severe (pulmonary embolism, severe malnutrition, post-operative bleeding, and gastrointestinal obstruction) complications.

We agree with the expert panels (224, 225) that suggest bariatric surgery for adolescents with obesity-related co-morbid conditions that threaten the adolescent's health—a BMI >40 and a severe co-morbidity or a BMI >50 and less severe co-morbidity. These cut-points are the ones generally accepted for adolescents. Others have suggested that we should consider using a BMI $\geq 99^{\text{th}}$ percentile—equivalent to an adult BMI of 35-40—with severe co-morbidities as a cut-point (226). There are insufficient data concerning the complication rates using the current cut-points to warrant suggesting any changes (222, 223, 227).

Gastric pacing studies are being done in adults (228), but no studies in children have been published.

3.3. Values and preferences

Our suggestion for limited use of bariatric surgery places a relatively higher value on avoiding anatomical and functional changes in developing children, on avoiding unforeseen complications associated with life-long exposure to these changes, and on avoiding the costs and perioperative complications of these procedures. It places a relatively lower value on the weight loss and amelioration of obesity-related complications associated with bariatric surgery.

3.3 Remarks

Requirements for patients: It must be clear to the patient and the family that bariatric surgery is an adjunct to a sincere commitment to alteration of lifestyle and behavior rather than a cure. All obese children must first demonstrate their ability to adhere to a family-based dietary and lifestyle modification program.

Requirements for preoperative care: Bariatric surgery in adolescents should be performed in regional pediatric academic centers with programs equipped to handle the data acquisition, long-term follow-up, and multidisciplinary issues of these difficult patients (224). A multidisciplinary team with medical (including endocrine, gastrointestinal, cardiovascular, pulmonary, and otolaryngologic expertise), surgical, nutritional, and psychological expertise should carefully select adolescents who are well informed and motivated as potential candidates for bariatric surgery and should provide preoperative care and counseling. Patients and families must be well informed as to the risks and complications of bariatric surgery.

Requirements for postoperative care: Postoperative attention to the principles of growth, development, and compliance is essential to avoid adverse physical, cognitive, and psychosocial

outcomes after bariatric surgery (224). Adolescents undergoing bariatric surgery require lifelong medical and nutritional surveillance postoperatively (229), especially to ensure adequate vitamin and mineral intake, as well as extensive counseling. Patients lacking such help tend to regain their weight over time (230).

Finally, more data are required to assess the effects of and complications from bariatric surgery in adolescents, and clinicians are encouraged to enroll their patients into a national database. The NIH has funded a multicenter study of bariatric surgery in adolescents (<http://clinicaltrials.gov/ct2/show/NCT00474318>). Criteria for patient enrollment include adolescents who have reached physical and psychological maturity, whose BMI ≥ 40 with significant obesity related co-morbidity, and who were unable to lose weight after 6 months of supervised participation in two separate behavioral or medical weight loss programs.

4.0 Prevention of obesity recommendations

4.1.a *We recommend breast-feeding for a minimum of 6 months. (1|⊕⊕○○)*

4.1.b *We suggest that clinicians promote and participate in efforts to educate children and parents by means of ongoing anticipatory guidance about healthy dietary and activity habits and, further, that clinicians encourage school systems to provide adequate health education courses promoting healthy eating habits. (2|⊕⊕○○)*

4.1.c *We suggest that clinicians promote and participate in efforts to educate the community about healthy dietary and activity habits. (2|⊕○○○)*

4.1 Evidence

The prime objective should be to prevent obesity before it happens, because once it develops, treatment is difficult (231-233). A meta-analysis commissioned by the Task Force (234) of randomized trials of lifestyle interventions to prevent pediatric obesity found significant but modest effects of these interventions on increasing physical activity ($p=0.004$), decreasing sedentary behavior ($p<0.00001$) with a significantly greater effect when directed toward children in contrast to adolescents ($p=0.02$), and reducing unhealthy dietary habits ($p<0.00001$). These beneficial effects did not translate into important changes in BMI (234). But, as discussed under Remarks in Section 3.1.0, weight maintenance in a growing child may be as effective as weight loss in an adult.

Breast-feeding in infancy is associated with a decreased incidence of overweight and obesity in childhood (235-239). Infants exclusively breast-fed for 3 to 5 months are 35% less likely to be obese when they enter school (240). A study of sibling pairs supports this finding (241). A meta-analysis demonstrated an inverse relationship between the duration of breast-feeding and the risk of becoming overweight with a plateau after 9 months of breast-feeding. The odds ratio for becoming overweight declines to 0.81 after 3 months of breast-feeding, to 0.76 after 6 months, and to 0.67 after 9 months of breast-feeding, after which it plateaus (242).

A review of 39 published school-based intervention studies designed to prevent childhood obesity showed that 40% of the 33,852 participating children had a positive effect on weight control, and the remainder had a neutral effect. None of the studies showed a negative effect (243). Exercise can play an important role for adolescents in the context of prevention of young adult obesity (244).

For most children and families, lifestyle patterns related to eating and exercise are established early, affecting children not only when they are young but also throughout life. To

avoid the harmful health consequences of less-than-optimal lifestyle choices, it is incumbent upon health care providers to convey to their patients and their families healthy guidelines, explaining, in a culturally sensitive and language-appropriate manner, the caloric needs and essential nutrient requirements of young children, as well as the importance of physical activity. This is of particular importance when we consider the increased efficacy of prevention trials when directed toward children rather than adolescents (234).

Although healthy infants can differ considerably from one another in their caloric intake, appetite is the most efficient way to determine what an infant needs. Most infants instinctively know how much food they need and will not undereat or overeat unless pressured (245). Babies should be fed when hungry but should not be forced to finish all that they are served. Although low-fat and low-cholesterol diets are in vogue, they are not recommended for children under 2 years of age.

4.1. Values and preferences

In making these recommendations, the Task Force placed a relatively high value on promoting these lifestyle preventive activities with their potential wide-reaching benefits (beyond weight control) and potential safety.

4.1 Remarks

Key players in the prevention and treatment of pediatric obesity are primary health care professionals (121, 231). Although anticipatory guidance is an important facet of pediatric care and it appears capable of having a salutatory effect on excessive weight gain (246), less than half of the recommended guidance is received (247). In some studies only 19% of primary care

physicians were aware of the AAP recommendations on obesity and only 3% complied with all of the recommendations (248). The situation was little better in an academic medical center where charts of obese children documented the obesity only 53% of the time (249). Whereas a diet was prescribed for 71% of the obese children, increased physical activity was prescribed 33% of the time and reduced television time only 5% of the time; the recommended laboratory tests were obtained only 13% of the time. Although obesity was recognized in 76% of the obese adolescents, it was recognized only 31% of the time in preschoolers (249). This latter finding was particularly disappointing because prevention of overweight and obesity should be undertaken while the condition is still mild and more readily treated by weight maintenance rather than weight loss. If a toddler or child is crossing BMI percentiles upwards (or weight-for-height percentiles), the primary care provider should begin to intervene by recommending reduced-fat milk and by restricting the intake of calorie-containing beverages such as juice as an initial step. If progress is not seen, then additional steps need to be undertaken (232, 233).

5.0 Societal barriers to implementation

5.1 Recommendation

We suggest that clinicians advocate for regulatory policies designed to decrease the exposure of children and adolescents to the promotion of unhealthy food choices in the community (e.g., by media advertisements targeting children and adolescents). (2|⊕○○○)

5.1 Evidence

Through advertising, especially on television, the food industry exerts an enormous influence on children that can negate the influence of parents and teachers. Although television can provide

valuable educational programs, its advertisements for sweetened drinks, fast food restaurants, and high-calorie snacks can instill poor eating and leisure-time habits in children. In 2002 the food industry spent \$10-\$12 billion on advertising directed toward children (121 [pp 153-192]). In a survey of television programs aimed at children aged 2-11 and 12-17 years, investigators found that 97.8% and 89.4%, respectively, of the advertisements were for food products consisting predominantly of nutrient-poor, high-sugar ingredients (250). A 30-second ad can affect a child's food choices (251). A 2006 Institute of Medicine report cited these negative factors as influencing children and adolescents to adopt unhealthy lifestyle choices (252).

5.2 Recommendation

We suggest that clinicians advocate that school districts ensure that only nutritionally sound food and drinks are available to children in the school environment, including the school cafeteria and alternative sources of food such as vending machines. (2|⊕○○○)

5.2 Evidence

The average child who participates in the school lunch program consumes one third of the daily-recommended caloric intake in school and three fifths if breakfast is also taken in school (121 [pp 237-284]). Of the 58 million schoolchildren in this country, about 28 million take part in the lunch program and 8 million in the breakfast program. Although federal standards require that these meals meet certain nutritional standards, the total fat composition of lunches exceeded the federal nutrition target of no more than 30% of total calories (121 [pp 237-284]). As children move through the school system, fewer students participate in the school lunch program, and they consume a nutritionally poorer lunch obtained through alternative food sources.

The problem largely lies in the availability of alternative food sources (termed “competitive foods”) in the school via snack bars, vending machines, or school-sponsored fund-raising sales. School districts have been tackling the issue by banning the sale of nutritionally poor food in their cafeterias, snack bars, and vending machines at school-sponsored events. These efforts have met resistance on the part of children, their parents (Weil E 2005 Heavy questions. *New York Times Magazine* January 2), and school administrators who feared the loss of an important revenue source if children were to diminish their purchases of “competitive foods” But in a 5-month post-ban study, schools did not suffer any loss in revenues from changed food and beverage sales (Ryman A 2005 Horne: schools won’t miss junk foods: sales study builds support for ban bill. *Arizona Republic* February 2;1). In several community studies the fat content of meals decreased for those students exposed to a ban on junk food sales (121 [pp 237-284]).

5.3 Recommendation

We suggest that clinicians advocate parental participation in the design of school-based dietary or physical activity programs and that schools educate parents about the rationale for these programs in order to ensure their understanding and cooperation. (2|⊕○○○)

5.3 Evidence

Even if schools institute health promotion programs and restrict the availability of unhealthy foods on site, these efforts may be contravened by cultural concepts and parental actions. Children may not be receptive to new, albeit healthy, foods, and persistence is required on the part of school authorities and parents (Weil E 2005 Heavy questions. *New York Times Magazine*

January 2). Adults, as well as children, may resist efforts to restrict unhealthy foods in schools. For example, when pastries and candies for at-school celebrations and birthdays were banned in one Texas school district, parents objected and the Texas state legislature passed a measure prohibiting such a ban (Mui YO 2005 At many elementary schools, the party's over. *Washington Post* October 30;sect C:1).

It is important that parents, as the primary caretakers, be educated about proper nutrition because the likelihood of the family's consuming the recommended amount of fruits and vegetables increases not only with family income and general educational level, but also with the amount of nutrition education received (253).

5.4 Recommendation

We suggest that clinicians advocate that community master planners design, redesign, and organize communities to maximize opportunities for safe walking or cycling to school, recreational activity and athletic events, and neighborhood shopping as means to encourage greater physical activity. (2|⊕○○○)

5.4 Evidence

The physical and social organization of communities plays a role in the prevention and treatment of childhood obesity. Today, only 25% of children walk or bike to school as compared with 48% in 1969 (121 [pp 193-221]).

Communities need to “provide places where children can play outside, particularly within their residential neighborhoods, and where they can safely walk, bike, or travel by other self-propelled means to destinations such as the park, playground, or school” (121 [pp 193-221]).

Some communities are now tackling this task as they adopt street layouts that suppress vehicular speed.

Parental concern about neighborhood crime and, therefore, their children's safety may cause children to remain at home after school and to engage in sedentary rather than physical activities. Although the data on this particular correlation are inconsistent (121 [pp 193-221]), studies show a correlation between parents' perceptions of neighborhood safety and childhood obesity (254, 255).

Some localities have made an effort to promote the "Safe Routes to School" (SR25) initiative to increase the percentage of students walking or riding bicycles to school. Comprehensive SR25 programs have been successful where tried (256, 257).

5.5 Recommendation

We suggest that clinicians advocate that policymakers provide incentives to ensure that retailers can offer affordable, high-quality fresh fruits and vegetables to all. (2|⊕○○○)

5.5 Evidence

For the poor, food expense represents an important item in a family's budget (121 [pp 193-221]). At least one study showed that a diet rich in fresh fruits, vegetables, whole grains, and protein costs more than a diet based on refined grains, added sugars, and fats (258). But a diet richer in fruits and vegetables can reduce caloric intake and lead to weight control (259). There are fewer conveniently located supermarkets in poorer neighborhoods but many "convenience stores" selling calorie-dense foods. Urban stores tend to stock fewer of the healthier foods and have less of a variety of foods (260). Even if supermarket availability were to improve, prices are still

lower in the poorer neighborhoods for the higher fat, calorie-dense foods (261). Some policies (including the federal government farm subsidy program) have encouraged production of high fructose corn syrup and other commodities used extensively in processed foods (Quaid L 2005 Feds aren't subsidizing recommended foods. *Washington Post* August 11). Even when restaurants advertise and make healthy food items available, consumer habits may continue to prefer burgers, fries, and other high-calorie, high-fat items (Pressler MW 2005 Hold the health, serve that burger. *Washington Post* August 18;sect A:1; Mitchell D 2005 Supersize comeback for fast food. *New York Times* November 12:sect B:5).

Summary of evidence-based guidelines for the prevention and treatment of pediatric obesity

The Task Force used systematic reviews of research evidence to formulate its key recommendations.

The number **1** indicates a strong recommendation and is associated with the phrase “we recommend”; **2** denotes a weak recommendation and is associated with the phrase “we suggest.”

Evidence grading: ⊕○○○ denotes very low quality evidence, ⊕⊕○○ low quality, ⊕⊕⊕○ moderate quality, and ⊕⊕⊕⊕ high quality.

Making strong recommendations with low quality evidence

For many issues in which the evidence base is of low or very low quality, the Task Force has, nonetheless, elected to make strong recommendations The strength of any recommendation depends on the tradeoff between benefits, and risks and burdens; and the quality of the evidence. A strong recommendation can be made when the tradeoff is clear enough so that most patients would make the same choice. A weak recommendation (suggestion in this guideline) is made when the tradeoff is less clear and the individual patient's values would affect the different choices (3).

“Clinical practice guidelines are guides only and may not apply to all patients and all clinical situations. Thus, they are not intended to override clinicians’ judgment” (4).

1.0 The problem with obesity

The objective of interventions in overweight and obese children and adolescents is the prevention or amelioration of obesity-related co-morbidities, e.g., glucose intolerance and T2DM, metabolic syndrome, dyslipidemia, and hypertension.

2.0 Diagnosis of overweight and obesity

2.1 We recommend the use of the body mass index (BMI, calculated as weight in kilograms divided by height in meters²), with CDC-derived normative percentiles, as the preferred method for the diagnosis of the overweight or obese child (46) **(1|⊕⊕○○)**

2.2 We recommend that a child be diagnosed as overweight if the BMI is $\geq 85^{\text{th}}$ percentile but $< 95^{\text{th}}$ percentile for age and sex and as obese if the BMI is $\geq 95^{\text{th}}$ percentile for age and sex. **(1|⊕○○○)**

2.3 We recommend against a routine laboratory evaluation for endocrine causes of obesity in obese children or early- to mid-pubertal obese adolescents unless the child’s height velocity, assessed in relation to stage of puberty and family background, is attenuated. **(1|⊕⊕○○)**

2.4.a We recommend referral to a geneticist for children whose obesity has a syndromic etiology, especially in the presence of neurodevelopmental abnormalities. **(1|⊕○○○)**

2.4.b We suggest that parents of children who have inexorably gained weight from early infancy and have risen above the 97th percentile for weight by 3 years of age be informed of the availability of MC4R genetic testing. However, the test is positive in only 2%-4% of such

patients who are above the 97th percentile for weight (69) and currently will not alter treatment.

(2|⊕○○○)

2.5 We recommend that children with a BMI \geq 85th percentile be evaluated for associated comorbidities and complications [see Table 1, Figure 1]. **(1|⊕○○○)**

3.0 Treatment of obesity

3.1 Lifestyle recommendations

3.1.0 We recommend that clinicians prescribe and support intensive lifestyle (dietary, physical activity and behavioral) modification to the entire family and to the patient, in an age-appropriate manner, and as the prerequisite for all overweight and obesity treatments for children and adolescents. **(1|⊕○○○)**

3.1.1 Dietary recommendations

3.1.1.a We recommend that clinicians prescribe and support healthy eating habits such as

- avoiding the consumption of calorie-dense, nutrient-poor foods (e.g., sweetened beverages, sports drinks, fruit drinks and juices, most “fast food,” and calorie-dense snacks) **(1|⊕⊕○○)**

3.1.1.b We suggest that clinicians prescribe and support

- controlling caloric intake through portion control in accordance with the Guidelines of the American Academy of Pediatrics
[<http://pediatrics.aappublications.org/cgi/reprint/117/2/544>] **(2|⊕○○○)**
- reducing saturated dietary fat intake for children older than 2 years of age **(2|⊕⊕○○)**
- increasing the intake of dietary fiber, fruits, and vegetables **(2|⊕○○○)**
- eating timely, regular meals, particularly breakfast, and avoiding constant “grazing” during the day, especially after school **(2|⊕○○○)**

3.1.2 Physical activity recommendations

3.1.2.1 We recommend that clinicians prescribe and support 60 minutes of daily moderate to vigorous physical activity. **(1|⊕⊕○○)**

3.1.2.2 We suggest that clinicians prescribe and support a decrease in time spent in sedentary activities, such as watching television, playing video games, or using computers for recreation. Screen time should be limited to 1-2 hours per day, according to the American Academy of Pediatrics (179) **(2|⊕○○○)**

3.1.3 Psychosocial recommendations

3.1.3.a We suggest that clinicians educate parents about the need for healthy rearing patterns related to diet and activity. Examples include parental modeling of healthy habits, avoidance of overly strict dieting, setting limits of acceptable behaviors, and avoidance of using food as a reward or punishment. **(2|⊕○○○)**

3.1.3.b We suggest that clinicians probe for and diagnose unhealthy intra-family communication patterns and support rearing patterns that seek to enhance the child's self-esteem. **(2|⊕○○○)**

3.2 Pharmacotherapy recommendations

3.2.a We suggest that pharmacotherapy (in combination with lifestyle modification) be considered if a formal program of intensive lifestyle modification has failed to limit weight gain or to mollify co-morbidities in obese children. Overweight children should not be treated with pharmacotherapeutic agents unless significant, severe co-morbidities persist despite intensive lifestyle modification. In these children, a strong family history of type 2 diabetes mellitus or cardiovascular risk factors strengthens the case for pharmacotherapy. **(2|⊕○○○)**

3.2.b We suggest that pharmacotherapy be offered only by clinicians who are experienced in the use of anti-obesity agents and are aware of the potential for adverse reactions. **(2|⊕○○○)**

3.3 Bariatric surgery recommendations

3.3.a . We suggest that bariatric surgery be considered only under the following conditions:

1. The child has attained Tanner 4 or 5 pubertal development and final or near-final adult height.
2. The child has a BMI >50 or has BMI >40 and significant, severe co-morbidities.
3. Severe obesity and co-morbidities persist despite a formal program of lifestyle modification, with or without a trial of pharmacotherapy.
4. Psychological evaluation confirms the stability and competence of the family unit.
5. There is access to an experienced surgeon in a medical center employing a team capable of long-term follow-up of the metabolic and psychosocial needs of the patient and family, *and the institution is either participating in a study of the outcome of bariatric surgery or sharing data.*
6. The patient demonstrates the ability to adhere to the principles of healthy dietary and activity habits. **(2|⊕⊕○○)**

3.3.b We recommend against bariatric surgery for pre-adolescent children; for pregnant or breast-feeding adolescents, and for those planning to become pregnant within 2 years of surgery; for any patient who has not mastered the principles of healthy dietary and activity habits; for any patient with an unresolved eating disorder, untreated psychiatric disorder, or Prader-Willi syndrome. **(1|⊕⊕○○)**

4.0 Prevention of obesity recommendations

4.1.a We recommend breast-feeding for a minimum of 6 months. **(1|⊕⊕○○)**

4.1.b We suggest that clinicians promote and participate in efforts to educate children and parents by means of ongoing anticipatory guidance about healthy dietary and activity habits and, further, that clinicians encourage school systems to provide adequate health education courses promoting healthy eating habits. **(2|⊕⊕○○)**

4.1.c We suggest that clinicians promote and participate in efforts to educate the community about healthy dietary and activity habits. **(2|⊕○○○)**

5.0 Societal barriers to implementation

We suggest that clinicians advocate

5.1 For regulatory policies designed to decrease the exposure of children and adolescents to the promotion of unhealthy food choices in the community (e.g., by media advertisements targeting children and adolescents). **(2|⊕○○○)**

5.2 That school districts ensure that only nutritionally sound food and drinks are available to children in the school environment, including the school cafeteria and alternative sources of food such as vending machines. **(2|⊕○○○)**

5.3 For parental participation in the design of school-based dietary or physical activity programs and that schools educate parents about the rationale for these programs in order to ensure their understanding and cooperation. **(2|⊕○○○)**

5.4 That community master planners design, redesign, and organize communities to maximize opportunities for safe walking or cycling to school, recreational activity and athletic events, and neighborhood shopping as means to encourage greater physical activity. **(2|⊕○○○)**

5.5 That clinicians advocate that policymakers provide incentives to ensure that retailers can offer affordable, high quality fresh fruits and vegetables to all. **(2|⊕○○○)**

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Figure 1 legend:

* See Section 3.2 and the footnote to Table 2 for the criteria for the selection of candidates for pharmacotherapy.

† See Section 3.3 for the criteria for selection of candidates for bariatric surgery.

TABLE 1. SCREENING TESTS FOR THE MORE COMMON OBESITY CO-MORBIDITIES

Co-morbidity	Case Detection Tests (Abnormal Values*)
<p align="center">Pre-diabetes</p> <p>1) Impaired fasting plasma glucose (Verify fasting status)</p> <p>2) Impaired glucose tolerance (if OGTT is used)</p>	<p align="center">Fasting plasma glucose (>100 mg/dL),</p> <p align="center">2 hr glucose >140 but <200</p>
<p align="center">Diabetes mellitus</p>	<p align="center">Fasting plasma glucose >126 mg/dL, or random value >200 mg/dL (if OGTT used, 2 hr glucose >200) If asymptomatic, must have repeat abnormal values on another occasion</p>
<p align="center">Dyslipidemia</p>	<p align="center">Fasting (12-14 hours) lipids—</p> <p align="center">Triglycerides >110 mg/dL (75th percentile); ≥160 mg/dL (90th percentile)</p> <p align="center">LDL cholesterol ≥110 mg/dL (75th percentile); ≥130 mg/dL (90th percentile)</p> <p align="center">Total cholesterol ≥180 mg/dL (75th percentile); ≥200 mg/dL (90th percentile)</p> <p align="center">HDL cholesterol ≤35 mg/dL (10th percentile); ≤40 mg/dL (25th percentile)</p> <p align="center">(76) **</p>
<p align="center">Hypertension</p>	<p align="center">Blood pressure >90th percentile (standardized according to sex, age, and height percentile)</p> <p align="center">(32)</p>
<p align="center">Non-alcoholic fatty liver</p>	<p align="center">Alanine aminotransferase (ALT) >2 SD above the mean for the laboratory</p>

* To convert mg/dL to mmol/L, multiply by 0.0555 for glucose, 0.0259 for cholesterol, and 0.0113 for triglycerides.

** A proposed refinement of these abnormal lipid levels has the potential advantage of linking adolescents' lipid levels to those of adults (78).

TABLE 2: MEDICATIONS PROPOSED FOR THE TREATMENT OF OBESITY*

Drug	Dosage	Side Effects	Monitoring and Contraindications
Sibutramine (95, 195, 196) Not FDA approved for <16 years of age	5-15 mg PO qd	Tachycardia, hypertension, palpitations, insomnia, anxiety, nervousness, depression, diaphoresis	Monitor HR, BP. Do not use with other drugs, MAO inhibitors.
Orlistat (95, 197, 198) Not FDA approved for <12 years of age	120 mg PO tid	Borborygmi, flatus, abdominal cramps, fecal incontinence, oily spotting, vitamin malabsorption	Monitor 25OHD ₃ levels. MVI supplementation is strongly recommended. A lower dose preparation has been approved for over-the-counter sale.
Metformin [†] Not FDA approved for treatment of obesity Approved for ≥10 years of age for T2DM (95, 96, 199, 200)	250-1000 mg PO bid	Nausea, flatulence, bloating, diarrhea; usually resolves. Lactic acidosis not yet reported in children.	Do not use in renal failure or with intravenous contrast. MVI supplementation is strongly recommended.
Octreotide [†] Not FDA approved for treatment of obesity (96, 201, 202)	5-15 mcg/kg/d SQ ÷ tid	Cholelithiasis (can be prevented by concurrent ursodiol), diarrhea, edema, abdominal cramps, nausea, bloating, reduction in thyroxine concentrations, decreased growth hormone but normal IGF-I.	Monitor fasting glucose, FT ₄ , HbA _{1c} . Useful only for hypothalamic obesity. Ursodiol co-administration strongly recommended.
Leptin [†] Not approved by FDA (203, 204)	Titration of dose to serum levels, SQ	Local reactions	Useful <i>only</i> in leptin deficiency.
Topiramate [†] Not FDA approved for treatment of obesity (205) Data in adults only	96-256 mg PO qd	Paresthesias, difficulty with concentration/attention, depression, difficulty with memory, language problems, nervousness, psychomotor slowing	No pediatric data.
Growth hormone [†] Not FDA approved for treatment of obesity	1-3 mg/m ² SQ daily	Edema, carpal tunnel syndrome, death in patients with pre-existing obstructive	FDA –approved only in Prader-Willi syndrome to increase

(206)		sleep apnea	height velocity. It should be used only after screening to rule out obstructive sleep apnea. Must closely monitor pulmonary function, glucose, HbA _{1c} .
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RCT, randomized controlled trial; MVI, multivitamins

* Pharmacotherapy is not usually considered if the BMI is <95th percentile, but there are additional factors to consider. If we initiate pharmacotherapy early in the course of obesity, we may prevent severe weight gain and metabolic complications, but we may treat an excess of children, raise the rate of unwarranted side effects, and increase the costs to individuals and to society. Alternatively, if we begin medication late in the course of obesity, we run the risk of runaway weight gain and long-term morbidity. One approach that reconciles these difficulties is to act aggressively with lifestyle intervention in overweight and mildly obese patients to prevent severe obesity and to consider pharmacotherapy when the risk of complications is high or soon after complications emerge. The tipping point for pharmacotherapy could be if the family history is strongly positive for a major co-morbidity. **Lifestyle intervention should precede pharmacotherapy and should be maintained during pharmacotherapy.**

† The use of these non-FDA-approved agents should be restricted to large, well-controlled studies.

Figure 1. Diagnosis and Management Flow Chart

