Pediatric Obesity Treatment with Pharmacotherapy

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Objectives

- Identify many factors contributing to making obesity a progressive, chronic, and complex disease
- Describe anti-obesity medications for pediatric obesity
- Explain monitoring, side effects, and contraindications for anti-obesity medications

Case

- 16 yo male had been coming to Wellness Works for almost 2 years
- He has been working on healthy lifestyle changes drinking more water, going to the gym 2-3 times/week, weight-lifting but trying to do more cardio
- Reports he doesn't eat a lot during meals but snacks throughout the day
- Admits he eats when he is bored at home currently summer, trying to stay busy
- Doesn't feel he eats large amounts of food, no guilt associated with eating
- Sleeping 11p/12a-9am, +snoring, good energy level during the day
- No symptoms of anxiety, depression, or ADHD

Case Continued

- Blood Pressure: 117/89, 118/77
- BMI: 170% of 95th percentile (Class III obesity)
- HbA1c 5.8 prediabetes range
- ALT 52 elevated
- Lipid panel normal
- Patient and his mother are interested if there is any medication that he could take to help with weight loss.

Percentage of Children Ages 2 to 19 Who Are Obese, by Age: Select Years, 1971-2016



surce: Data for 1971-2014: Fryar, C. D., Carroll, M. D., & Ogden, C. L. (2016). Prevalence of overweight and obesity nong children and adolescents aged 2-19 years: United States, 1963-1965 through 2013-2014. Hyattsville, MD: U.S. epartment of Health and Human Services, Centers for Disease Control and Prevention, National Center for Health atistics. Retrieved from https://www.cdc.gov/nchs/data/hestat/obesity_child_13_14/obesity_child_13_14.pdf. ata for 2015-2016: Hales, C. M., Carroll, M. D., Fryar, C. D., & Ogden, C. L. (2017). Prevalence of obesity among adults uf youth: United States, 2015-2016 (NSCH Data Brief No. 288). Hyattsville, MD: U.S. Department of Health and uman Services, Centers for Disease Control and Prevention, National Center for Health Statistics. Retrieved from tps://www.cdc.gov/nchs/products/databriefs/db288.htm.

Public Health Concern

- NHANES data from 2017-2018 reported prevalence of 19.3% in youth and severe obesity as high as 6%
 - 12-19 years old 21.2%
 - 25.6% in Hispanic youth; 26.9% Mexican American youth; 24.2% non-Hispanic Black youth
 - Children with obesity are 5 times more likely to have obesity as adults than children without obesity
 - Obesity is a chronic & progressive disease.

CLINICAL PRACTICE GUIDELINE Guidance for the Clinician in Rendering Pediatric Care

American Academy of Pediatrics



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Clinical Practice Guideline for the Evaluation and Treatment of Children and Adolescents With Obesity

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Treatment Experience of Obesity as a Chronic Disease



FIGURE 1

Treatment experience of obesity as a chronic disease (this figure illustrates how longitudinal care is important to help address this chronicity and to address and buffer the social and contextual factors that influence a person's health).

Hampl, Pediatrics, 2023.

Managing Obesity in Primary Care

• Top 3 chronic conditions of childhood

asthma - 8.3%

ADHD - 9.8%

Obesity – 19%

Need to dispel many myths of obesity.

- Children will not outgrow obesity due to so many prevalent risk factors.
- Obesity is a disease, not a lifestyle choice.

It's not so simple!

- Fat mass our body's defense against starvations
- Brain/body defends the fat mass set point
- Fat mass set point is regulated by the brain



Jania Jastreboff, Weill-Cornell Obesity Medicine course, 2022. Figure Adapted from Zheng & Berthoud et al. Physiology. 2008;23 75-83.



Jania Jastreboff, Weill-Cornell Obesity Medicine course, 2022.



•Genetics

- •Socioeconomic Status/Food Insecurity
- •Sleep
- •Early-life risk factors
- •Microbiome
- •Children with special healthcare needs
- •Psychosocial stress/Mental Health

KAS 2: Pediatricians and other PHCPs should evaluate children 2 to 18 y of age with overweight (BMI \ge 85th percentile to <95th percentile) and obesity (BMI \ge 95th percentile) for obesity-related comorbidities by using a comprehensive patient history, mental and behavioral health screening, SDoH evaluation, physical examination, and diagnostic studies.

Obesity & Covid

- COVID disrupted children and adolescents structured routines.
- Families who were disproportionately affected by obesity pre-pandemic were more likely to experience an impact on income, food, and other social determinants of health
- The estimated proportion of persons aged 2-19 years old with obesity was 19.3% in August 2019 and 22.4% in August 2020
- The monthly rate of increase in BMI nearly doubled during the COVID-19 pandemic compared with a pre-pandemic period
- 6-11 year old children experienced largest increase in rate of BMI change; pandemic rate was 2.5 times as high as pre-pandemic rate



Lange, et al, MMWR,2021



Estimated Body Mass Index Before and During the COVID-19 Pandemic Jan 2018 – November 2020, Lange, MMWR, 2021, Lange, MMWR, 2021.

APPENDIX 1 Algorithm for Screening, Diagnosis, Evaluation, and Treatment of Children and Adolescents with Obesity



Treatment of Pediatric Obesity Refer early to Intensive Health Behavior & Lifestyle Treatment, no watchful waiting

3% reduction in BMI in 1 month predicted 5% reduction in BMI in 1 year

USPSTF ≥ 26 contact hours of behavioral interventions over 12 months

Wellness Works Program

- 2-21 years old
- Offer visits in Mount Pleasant, North Charleston, Summerville, and virtual visits throughout South Carolina, also available with School-based Health Program
- Dietician visits every month and physician visit every 3-6 months, Physical Therapist, & Psychologist too!
- Exercise sessions everyday
- Group sessions once/week
- Curriculum includes individual sessions on healthy eating, safe exercising, reading food labels, limiting tempting foods and screen time, goal setting, self-monitoring, rewards, and problem solving
- Place referral to Heart Health in epic

KAS 11: Pediatricians and other PHCPs should provide or refer children 6 y and older (Grade B) and may provide or refer children 2 through 5 y of age (Grade C) with overweight (BMI \ge 85th percentile to <95th percentile) and obesity (BMI \ge 95th percentile) to intensive health behavior and lifestyle treatment. Health behavior and lifestyle treatment is more effective with greater contact hours; the most effective treatment



Bariatric Surgery

Roux-en-Y bypass, vertical sleeve gastrectomy

Drug Therapy

Orlistat, Phentermine, Liraglutide

Behavioral Intervention

Behavioral change strategies, dietitian counseling



Anti-Obesity Pharmacotherapy

- There haven't been many options until recently
- Offer all treatment at once, no need to wait
- New FDA approved medications is the dawn of a new era
- Remember: obesity is a chronic disease. Treat obesity like you would treat any other chronic disease.

KAS 12: Pediatricians and other PHCPs should offer adolescents 12 y and older with obesity (BMI \ge 95th percentile) weight loss pharmacotherapy, according to medication indications, risks, and benefits, as an adjunct to health behavior and lifestyle treatment

Weight Promoting Medications:

Antihistamines Steroids Amitriptyline Paroxetine Sertraline Carbamazepine Gabapentin Valproate Aripiprazole Clozapine Haloperidol Mirtazapine Olanzapine Quetiapine Risperidone Ziprasidone Propranalol

Weight Neutral Medications:

Inhaled nasal steroids Montelukast **Bupropion Buspirone** Citalopram Trazadone Venlafaxine Escitalopram Fluoxetine Lamotrigine Topiramate Levetiracetam Zonisamide Amphetamine Methylphenidate



Currently FDA Approved Adolescent Anti-Obesity Medication (AOM) Table

Created by Katie Queen, MD, FAAP, DABOM katiequeen3@gmail.com, revised 10.7.23

Medication & Dosing	Mechanism of action	Mean BMI Reductio n	Contraindicatio ns	Precautions	Good Choice For:	Side effects
Semaglutide (Wegovy) - 0.35 mg weekly x 1 month 0.5 mg weekly x 1 month 1:7 mg weekly x 1 month 1:7 mg weekly x 1 month 2:4 mg weekly Liraglutide (Saxenda) 0.6 mg daily x 1 week, then 1:8 mg daily x 1 week, then 2:4 mg daily x 1 week, then 3:4 mg daily x 1 week, then 3:5 mg daily x 1 week, then	GLP1 Class; - Binds and activates GLP- 1 receptors in brain, gut, pancreas, heart - Reduces hunger and cravings - Improves satiety - Improves insulin sensitivity	~16%	~ Personal or FHx MEN2, medullary thyroid cancer ~ Pregnancy ~ Nursing	- Pancreatitis - Gallbladder Dz - Insulin - GERD	 Prediabetes/Diabetes Strong hunger & cravings (hungry brain) Poor Satiety (hungry gut) NAFLD 	 Nause/Vomiting Diarrhea/Constipation Headaches Dsypepsia/GERD Abd Pain Fatigue/Dizziness Injection site rxn
Phentermine/ Topiramate (Qsymia) • 3.75 mg Ph/23 mg T QAM • Increase to 7.5 mg Ph/46 mg T after 2 weeks • If <3 % wt loss at 3 months, increase to 8.35 mg Ph/69 mg T x 2 wks then • Max 15 mg Ph/92 mg T	 Sympathomimetic, GABA receptor modulation Controlled Substance (class IV) Carbonic anhydrase inhibition Reduces hunger and cravings Improved satiety Can NOT stop abruptly, must wean off higher doses or seizures can occur 	~8-10%	 Cardiovascular Dz Uncontrolled HTN MAOI Use Hyperthyroid Glaucoma Hx Drug Abuse Agitated State Pregnancy Nursing Hypersensitivity Hx 	 Insomnia Other stimulant use SSRIs Insulin Renal impairment HTN (controlled) Alcohol Use Kidney stones Metabolic acidosis Poor cognitive function, academic problems Developmental Delay Severe Depression/Anxiety 	 Strong Hunger & Cravings (hungry brain) Poor Satiety (hungry gut) Low Energy (slow burner) Binge eating symptoms Atypical anti-psychotic wt gain Migraines/Headaches Night eating Hx seizures 	- HTN - Tachycardia - Insomnia - Dry Mouth, - unpleasant taste - Paresthesias - Fatigue, Dizziness - Constipation - Cognitive challenges - Depression/Anxiety - Arthralgias, - Ligament sprain

Semaglutide (Wegovy)

 GLP-1 agonist, enhances glucose-dependent insulin secretion, suppresses glucagon secretion, slows gastric emptying, improved glucose control, decreased food intake and enhanced satiety

5-16% decrease in BMI

- FDA approved for children 12 years and older with obesity
- Weekly injections, Start at low dose and titrate up as tolerated
- Month 1: 0.25mg/Month 2: 0.50mg/Month 3: 1.0mg/Month 4: 1.7mg/Month 5: 2.4mg



Liraglutide (Saxenda)

- GLP-1 agonist, enhances glucose-dependent insulin secretion, suppresses glucagon secretion, slows gastric emptying, improved glucose control, decreased food intake and enhanced satiety
- 8% decrease in BMI
- FDA approved children 12 years and older Daily injections
- Dosing Increases each week 0.6mg, 1.2mg, 1.8mg, 2.4mg, 3.0mg



Semaglutide (Wegovy)/Liraglutide (Saxenda)

- Side effects: n/v/constipation, abdominal pain
 - watch for signs of pancreatitis, gallbladder disease, hypoglycemia, and depression
- Contraindications: Patients with aversion to needles, history of pancreatitis, personal or family history of medullary thyroid cancer or MEN type 2
- Contraception
- Insurance coverage is an issue
- Encourage high protein diet

Phentermine/ Topiramate ER (Qsymia)

- Norepinephrine reuptake inhibitor
- Modulation of GABA receptors
- 8-10% decrease in BMI
- FDA approved for adolescents 12 years and older with obesity
- Dosing: 3.75mg/23mg, 7.5mg/46mg, 11.25mg/69mg, 15mg/92mg



Phentermine/Topiramate ER (Qsymia)

- At 12 weeks after escalating to top dose: discontinue if patient hasn't experienced a reduction of at least 5% of baseline BMI
- Discuss contraception
- Insurance

Phentermine



- Schedule IV medication; ADHD stimulants are schedule II medications
- FDA approved for 16 years and older for short-term use (12 weeks), approved in 1959!
- Dosing: 8mg, 15mg, 37.5mg
- Side effects: Dry mouth, increased heart rate and blood pressure, insomnia, anxiety
- Contraindications: Cardiovascular disease, uncontrolled HTN, glaucoma, anxiety

Topiramate

- Topiramate is not FDA approved for obesity by itself
- Dosing: 25mg, 50mg, 100mg (target dose is 75-100mg)
- Side effects: Paresthesia, cognitive slowing
- Contraindications: Nephrolithiasis
- If discontinuing, taper dose due to possibility of precipitating seizures with abrupt cessation
- Can take phentermine and topiramate together or separate to help with hunger during different times of the day

Orlistat

sto

- Intestinal lipase inhibitor that blocks fat absorption
- 3% body weight loss or $<1 \text{kg/m}^2$ in BMI reduction
- FDA approved for obesity in adolescents 12 years and older
- Side effects: steatorrhea, fecal urgency, flatulence
- Contraindicated with chronic malabsorption syndrome, cholestasis, liver disease or renal impairment

Setmelonotide

- Melanocortin 4 receptor agonist in patients with POMC, PCSK1, leptin receptor deficiency or Bardet-Biedl syndrome
- FDA approved for children 6 years and older with specific genetic mutation
- 8-10% decrease in BMI
- Consider genetic testing in specific patients
- Side effects: Injection site reaction, skin hyperpigmentation, GI symptoms, priapism
- Contraindicated in depression

Metformin & other options

- FDA approved for type 2 DM in patients 10 years and older
- Modest effects on weight loss
- Meta-analysis of 6 studies shows mean BMI reduction of -0.86kg/m²
- May add with antipsychotic to help prevent weight gain/diabetes
 - Important to screen for mental health concerns including binge eating disorder, anxiety, depression, & ADHD





Key Takeaways

- Obesity is a chronic disease. Once medication is started, it is likely needed long term.
- First step of pharmacotherapy is to look at current medications and see if any can be stopped or changed to weight neutral medications.
- Anti-obesity medication is a supplement to healthy lifestyle changes.
- Start low and go slow, one medication at a time, but polypharmacy is likely.

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