

Obesity Management

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Designed for healthcare professionals, this interactive virtual course equips participants with comprehensive strategies for the medical treatment of obesity. The curriculum is tailored to those committed to mitigating the health impacts of obesity, such as prescribers, pharmacists, and allied health workers.

COURSE DESCRIPTION

By the end of this course, learners will be able to:

- ▶ Describe the prevalence of obesity in US adults, citing statistical data and epidemiological trends.
- ▶ Explain evidence-based treatment options for obesity, including efficacy and safety of current therapeutic interventions.
- ▶ Develop a comprehensive management plan for treatment of patients with obesity, including considerations for behavioral counseling, obesity pharmacotherapy, and surgical interventions.
- ▶ Construct a multidisciplinary team approach, integrating various healthcare professionals to support patient-centered care.

LEARNING OBJECTIVES

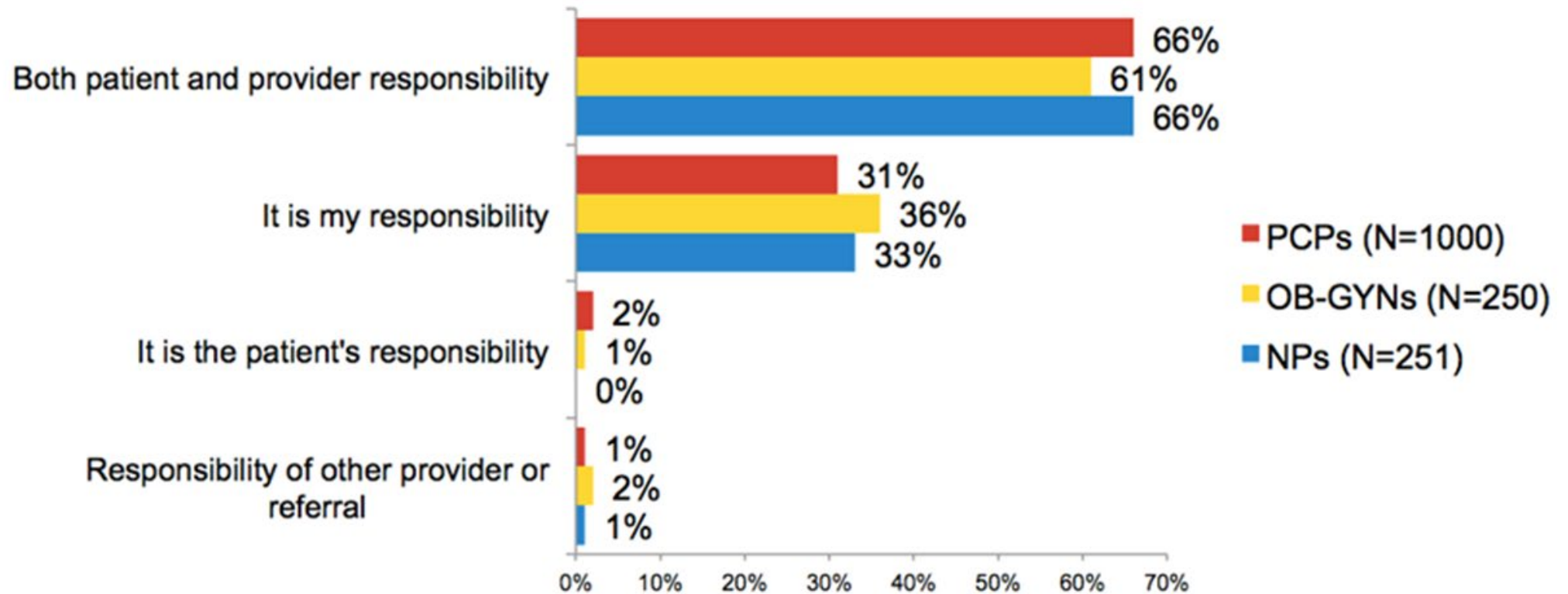
INTRODUCTION TO OBESITY

Which of these best characterizes your beliefs about obesity treatment?

- A. Obesity is a medical condition; it is the healthcare provider's responsibility to ensure that patients are counseled and receive treatment for obesity
- B. Obesity is a personal issue; it is the patient's responsibility to ensure that he/she gets the help they need
- C. Obesity is both a medical and personal issue; the responsibility for addressing obesity is shared between healthcare providers and patients
- D. Obesity is an issue of personal responsibility and willpower; patients should take better care of themselves and not burden the healthcare system

An Obesity Paradox

Whose responsibility is it to ensure that a patient is counseled about obesity?
(% Responding, single mention)



An Obesity Paradox

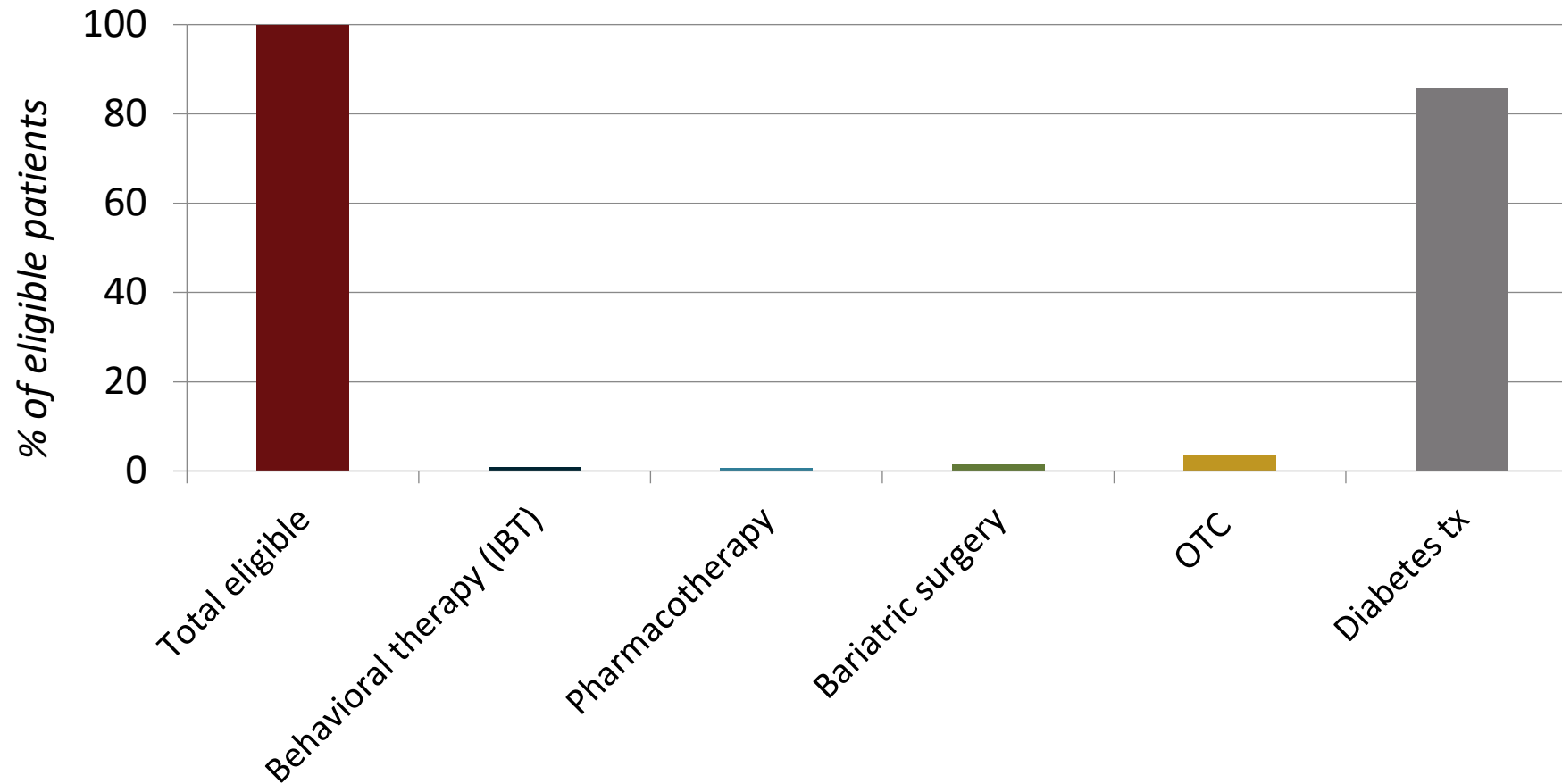
When Health Care Providers (HCPs) Talk to Patients About Weight...

- 6-8x increased likelihood of correctly perceiving excess weight status
- 2-3x increased likelihood of having obesity management plan in place
- 3-4x increased likelihood of attempting weight loss
- 2x increased likelihood of losing >5% body weight
- Improved weight loss, weight loss maintenance, weight-related behaviors, weight-related comorbidities

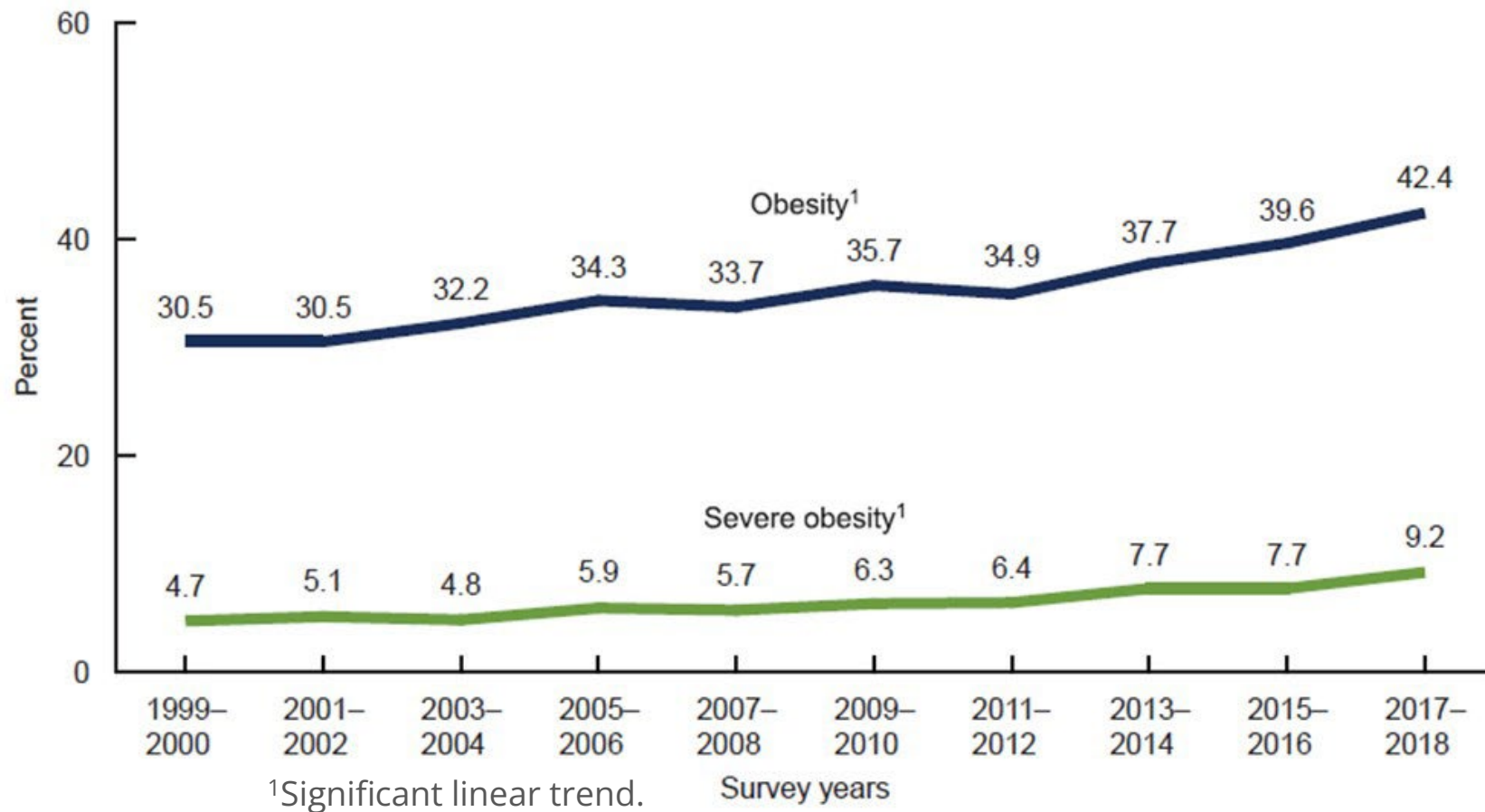
Clinical Obesity Practice

- Under-diagnosis
 - BMI 30-35 kg/m²: 10.2% diagnosed
 - BMI >50 kg/m² : 56.8%
- Under-documentation
 - 34% of 33,718 patients with severe obesity
- Under-discussion
 - 54% with BMI >25 kg/m² told of excess weight
 - 2% of PCPs discussed recorded BMI with patients
- Under-counseling
 - 67% with severe obesity receive weight loss advice
 - Weight discussions last as little as 55 seconds

Clinical Obesity Practice

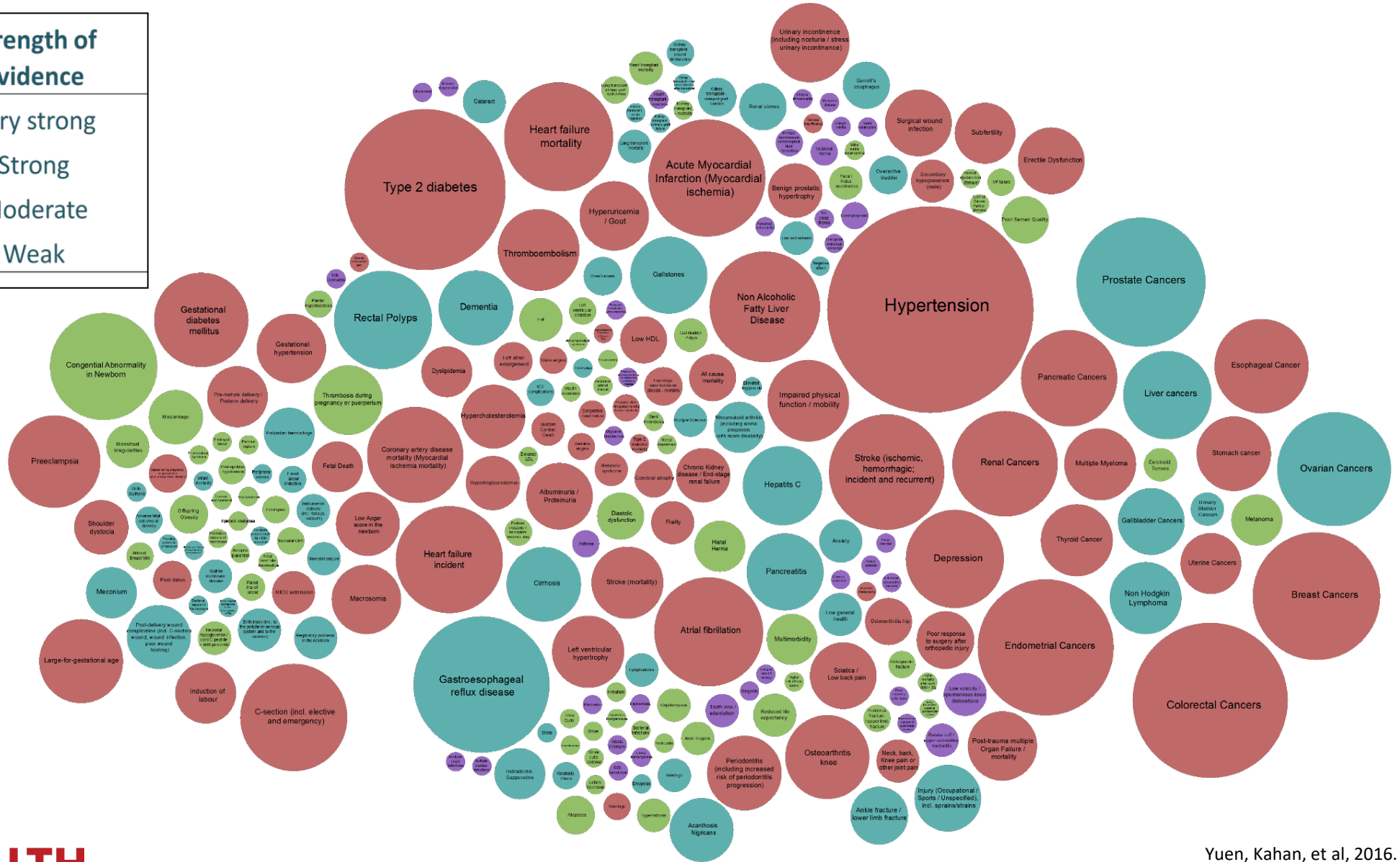


Trends in Obesity among Adults in the United States

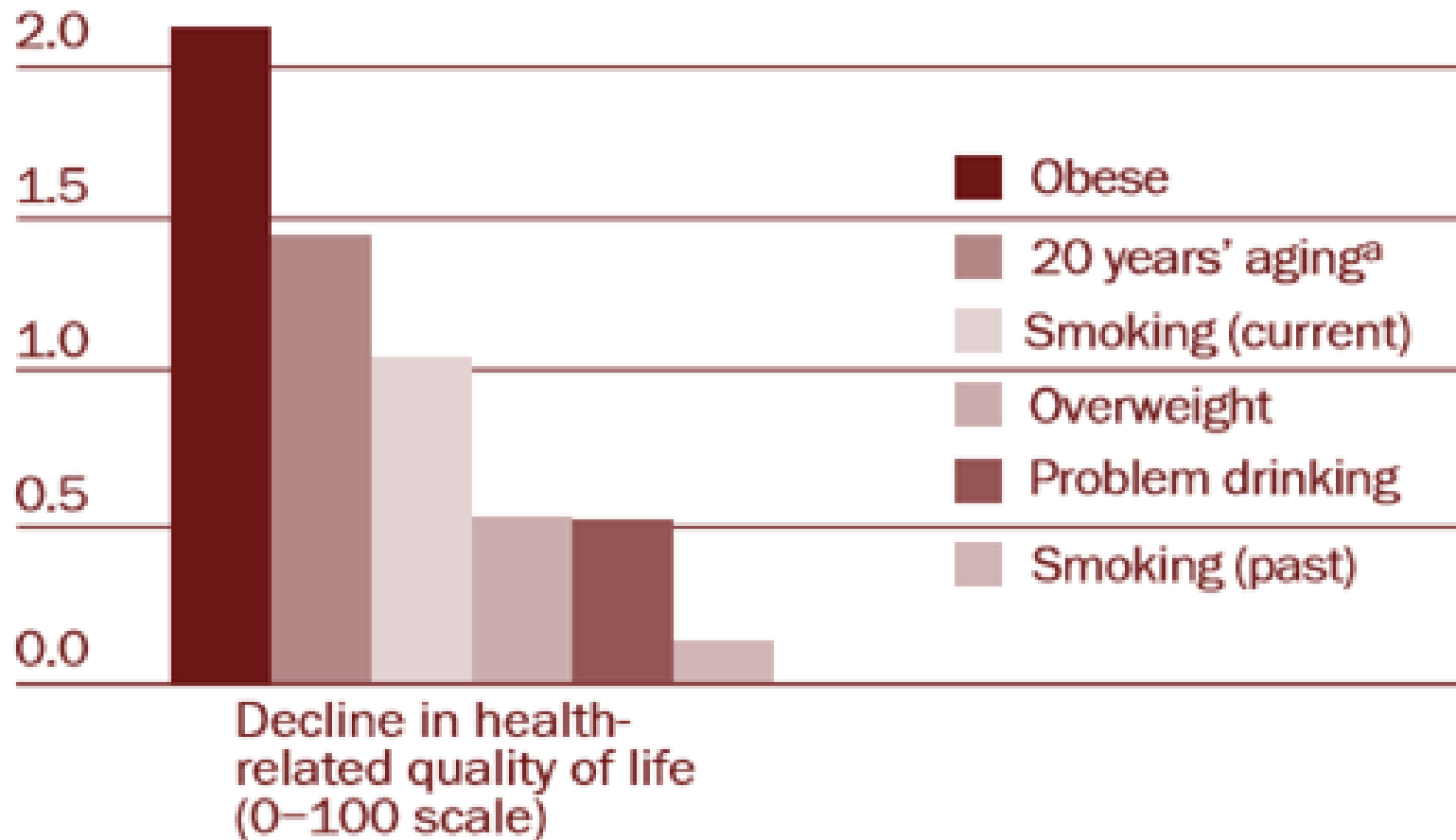


Obesity Comorbid Conditions

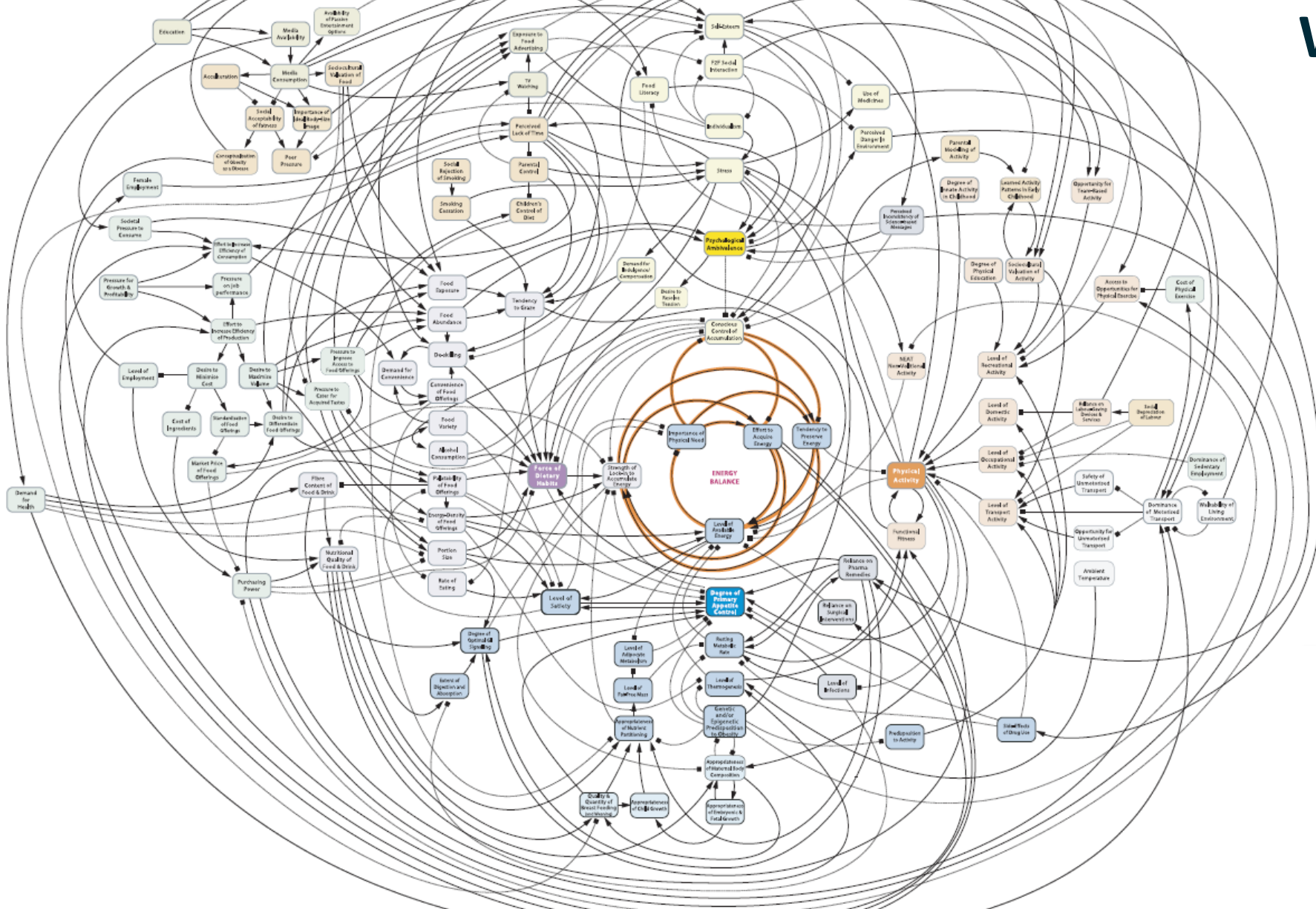
Color	GRADE	Strength of evidence
Red	4	Very strong
Teal	3	Strong
Light Green	2	Moderate
Purple	1	Weak



Obesity Strongly Impairs Quality of Life



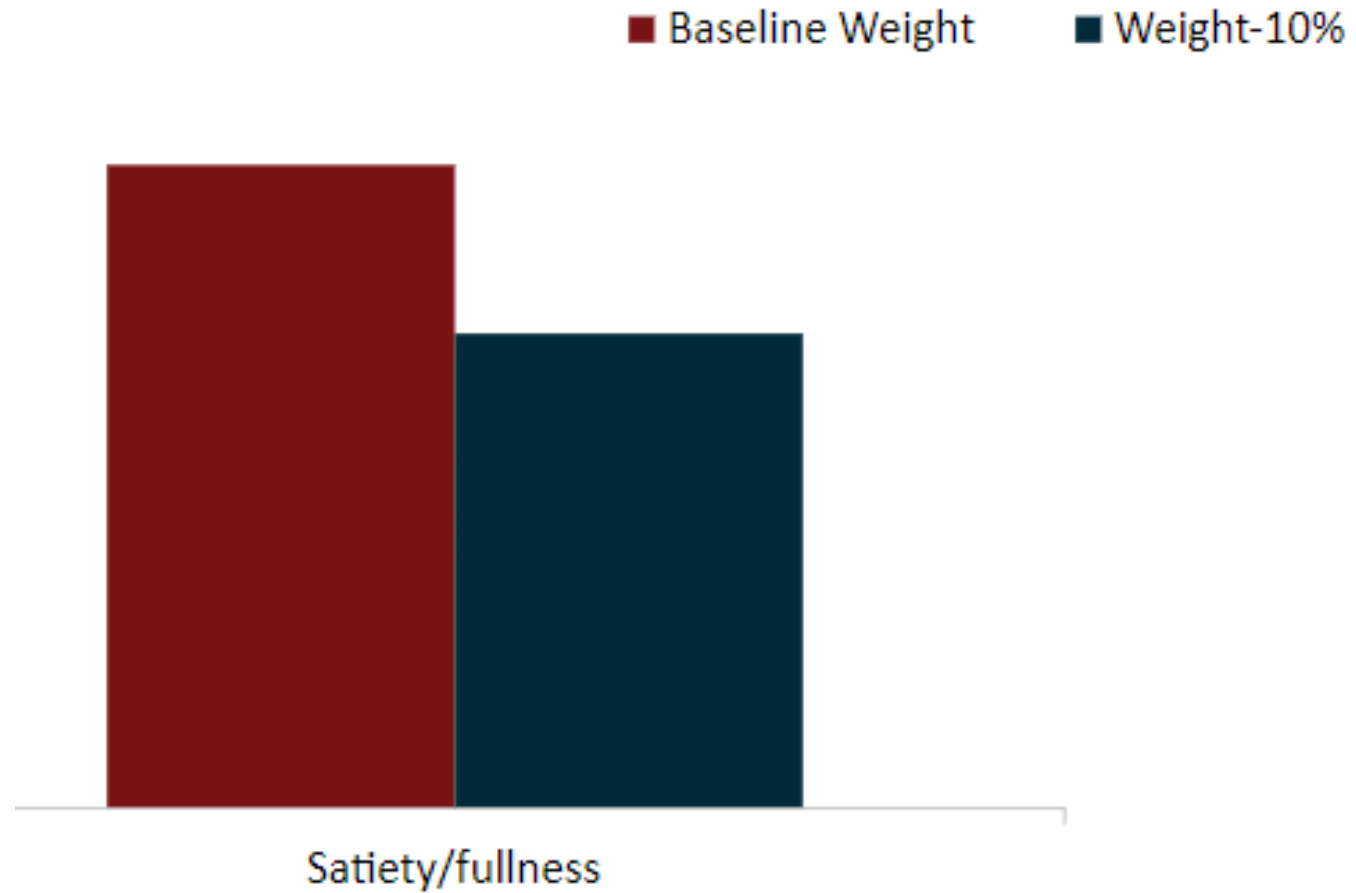
Why Is It So Hard To Manage Weight?



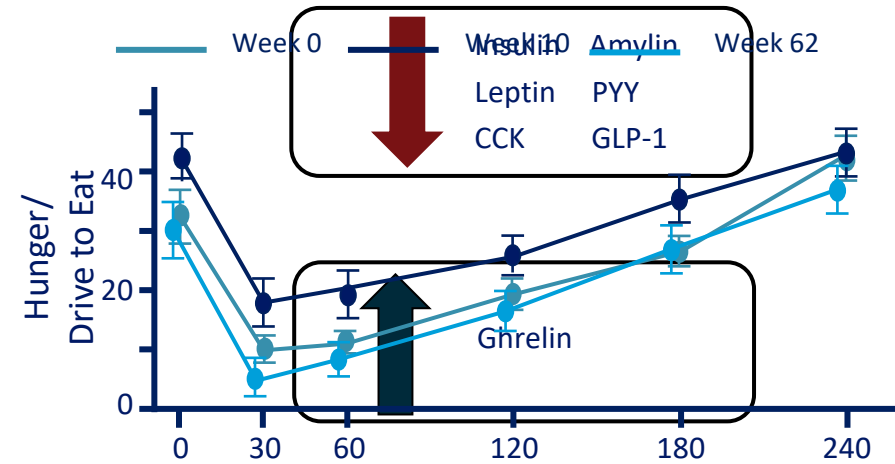
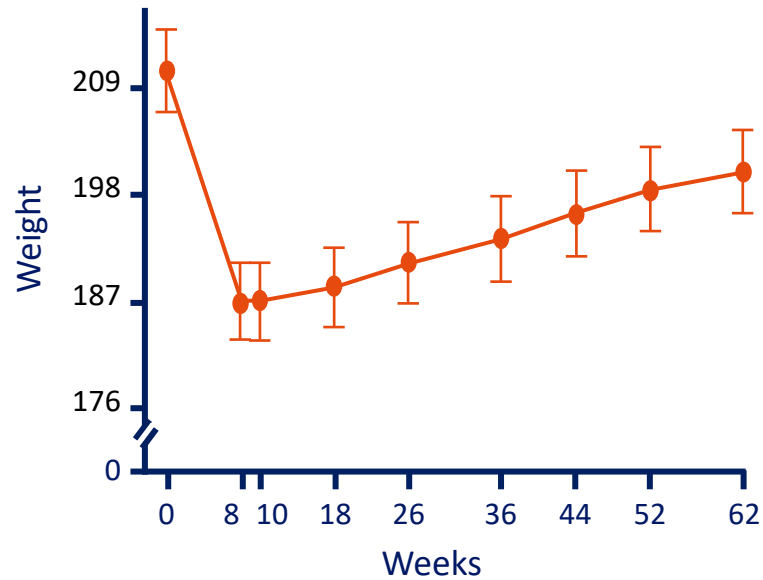
- Media
- Social
- Psychological
- Economic
- Food
- Activity
- Infrastructure
- Developmental
- Biological
- Medical

Positive Influence →
Negative Influence ←

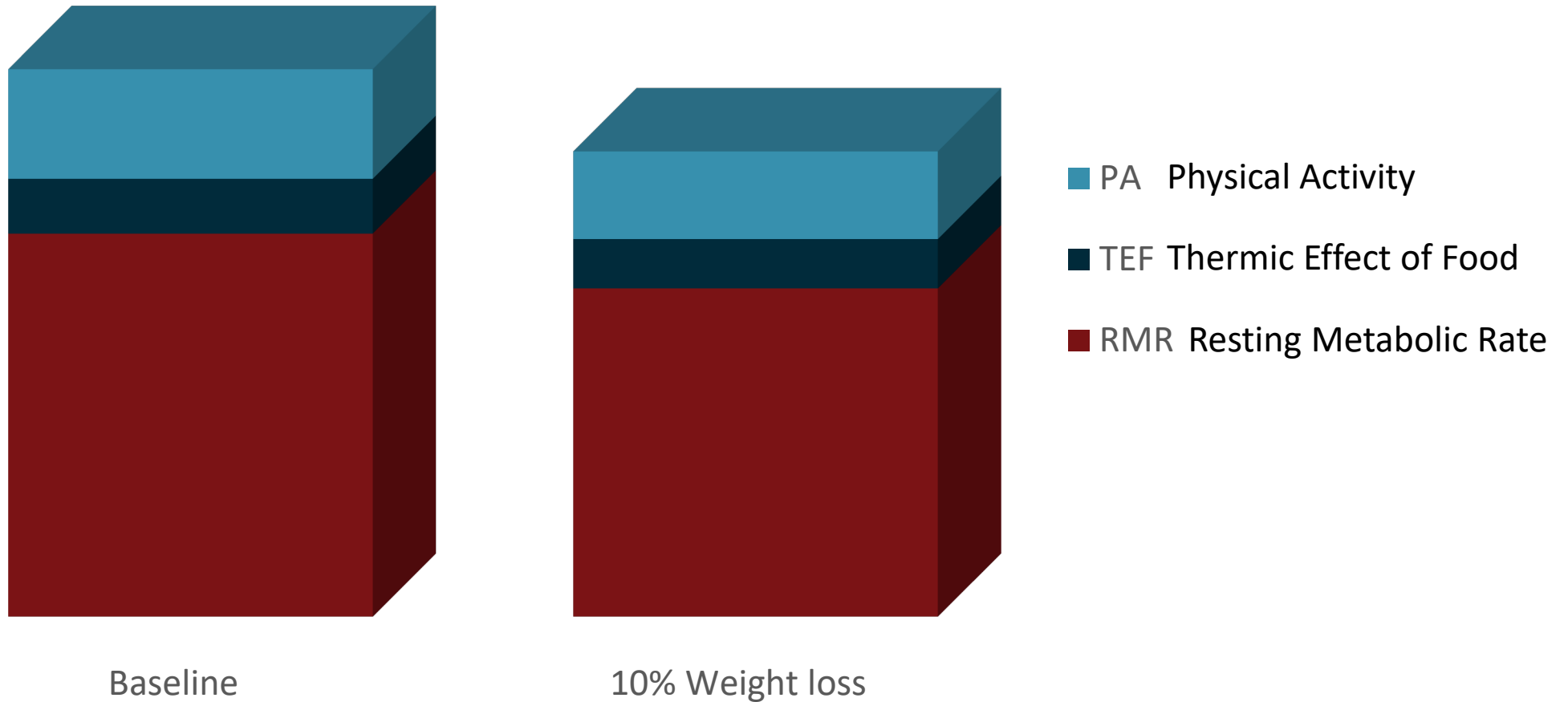
Effect of Weight Loss on Satiety



Hormone Changes and Hunger Persistently Oppose Weight Loss

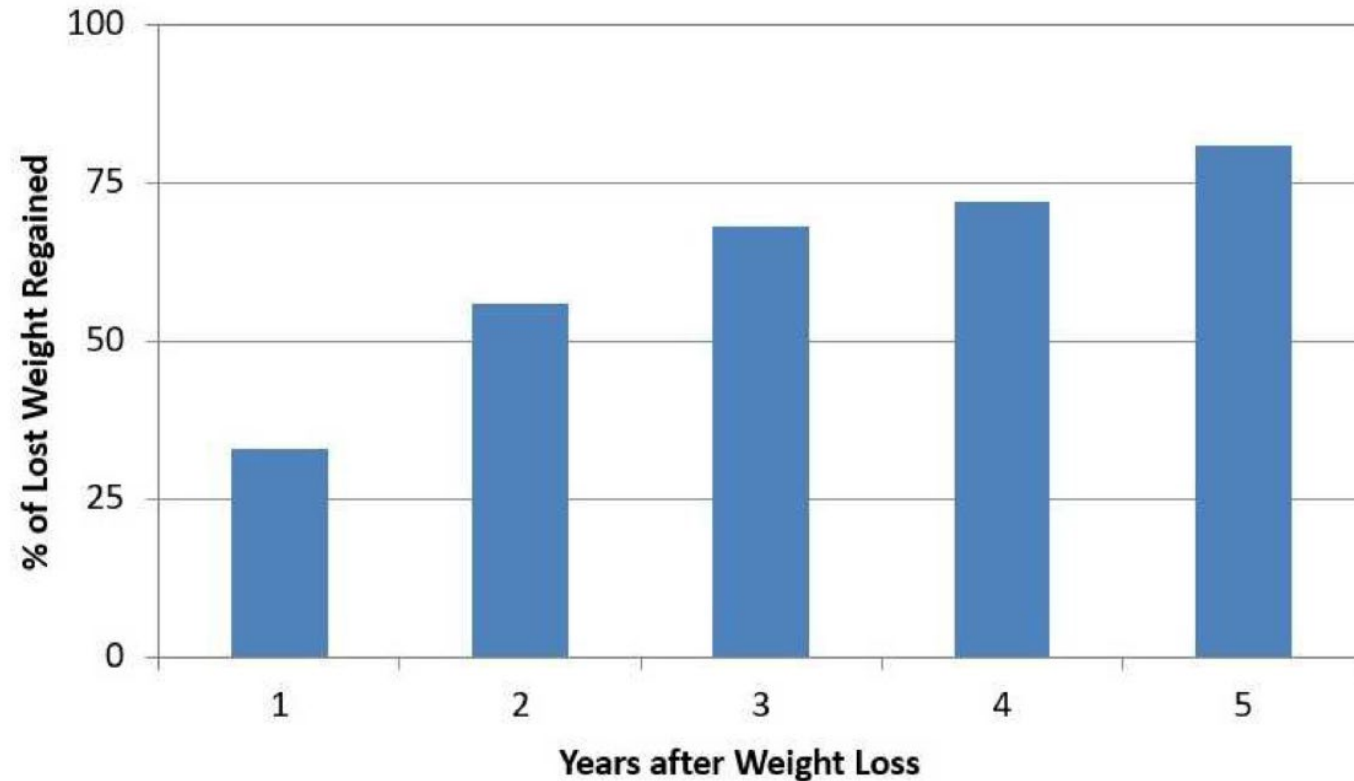


Effect of Weight Loss on Energy Expenditure (EE)



Weight Regain is Common

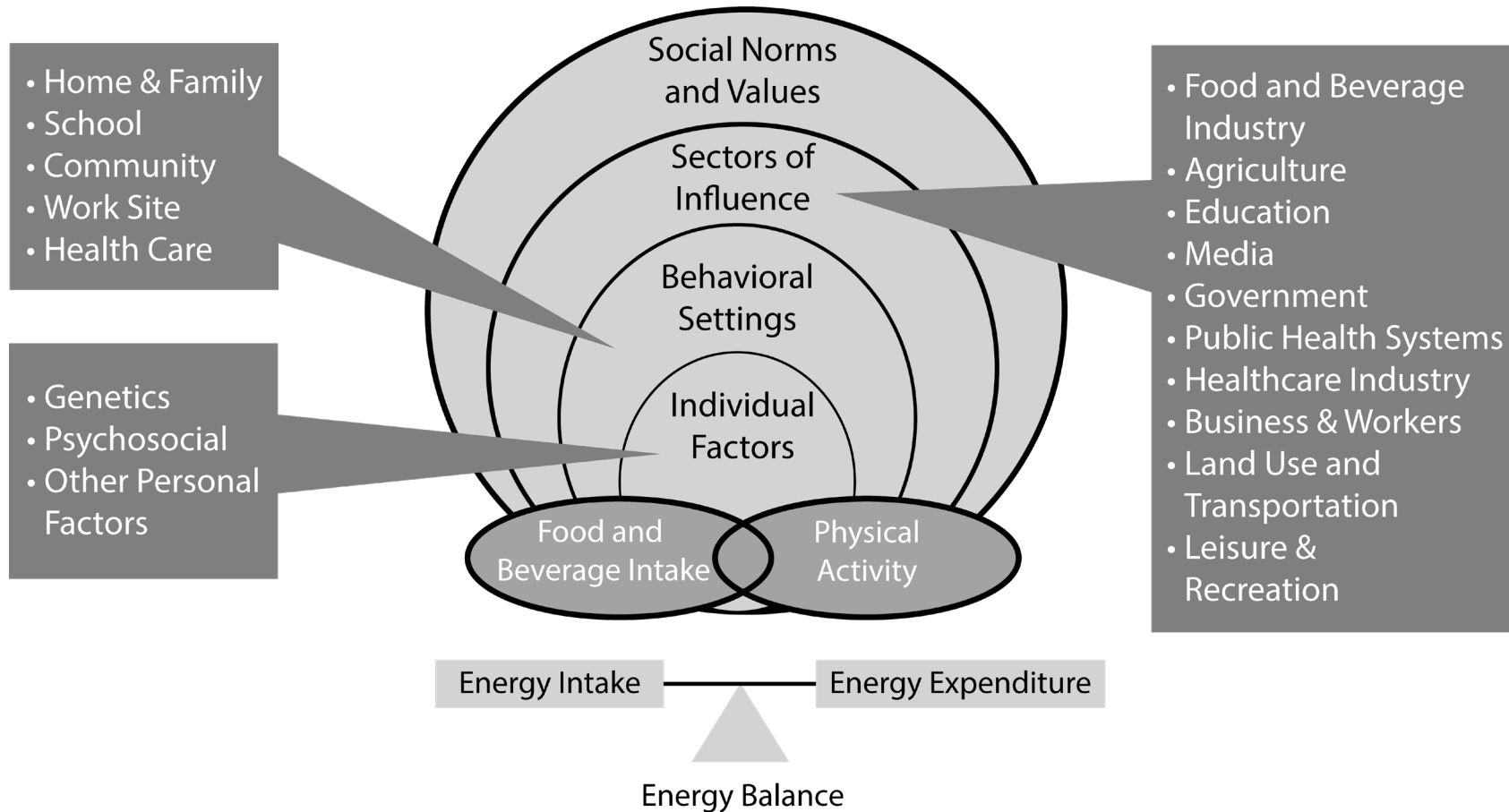
Meta-Analysis: Average time course of weight regain after weight loss intervention (n = 29 studies)



ADDRESSING OBESITY

Guidelines and Benefits

CDC Framework for Addressing Obesity



Modest Weight Loss Improves Health and Health Risks

Benefits of 5–10% weight loss

Reduced risk
of T2DM



Reduced CV
risk factors



Improved
lipid profile



Improved
blood pressure



Improved
sleep apnea



Improved
quality of life



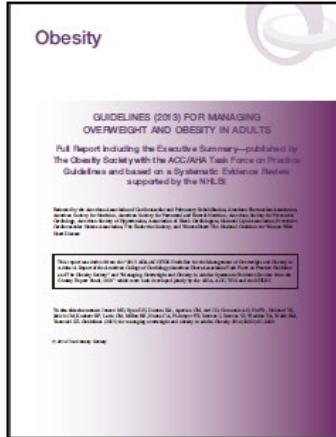
Modest Weight Loss Improves Health and Health Risks

Obesity Complication	% of Weight Loss for Benefit	Notes	References
T2DM Prevention	3-10%	Maximum benefit at 10%	DPP (Lancet, 2009) SEQUEL (Garvey et al, 2013)
T2DM (HbA1c)	3-15%	HA1c still decreasing at >15%	Look AHEAD (Wing, 2011)
Dyslipidemia (TG/HDL)	3-15%	TG still decreasing at >15%	Look AHEAD (Wing, 2011)
HTN	5-15%	BP still decreasing at >15%	Look AHEAD (Wing, 2011)
NAFLD	10%	Improved steatosis, inflammation, mild fibrosis	Assy et al, 2007; Dixon et al, 2004; Anish et al, 2009
Sleep Apnea	10%	Little benefit at ≤ 5%	Sleep AHEAD (Foster, 2009) Winslow et al, 2012
Osteoarthritis	5-10%	Improved symptoms and joint stress mechanics	Christensen et al, 2007 Felson et al, 1992; Aaboe et al, 2011
Stress Incontinence	5-10%		Burgio et al, 2007 Leslee et al, 2009
GERD	5-10% (F) 10% (M)		Singh et al, 2013 Tutuian R, 2011
PCOS	10-15%	Lower androgens, improved ovulation, increased insulin sensitivity	Panidis D et al, 2008 Norman et al, 2002 Moran et al, 2013

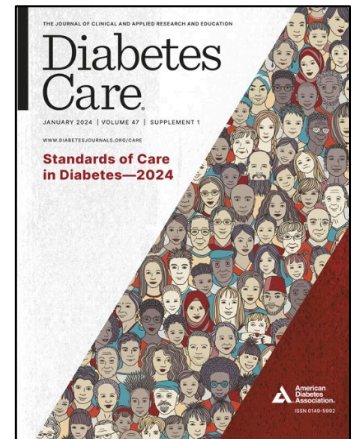
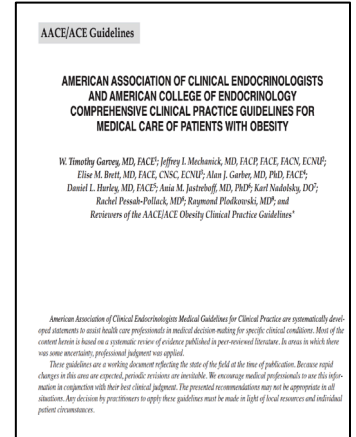
OBESITY TREATMENT

Modalities

Key Obesity Guidelines



- AHA/ACC/TOS Guidelines for Managing Overweight and Obesity in Adults (2013)
- Pharmacologic Management of Obesity: An Endocrine Society Clinical Practice Guideline (2016)
- AACE/ACE Comprehensive Clinical Practice Guidelines for Medical Care of Patients with Obesity (2016)
- ADA Obesity and Weight Management for the Prevention and Treatment of Type 2 Diabetes: Standards of Care in Diabetes (2024)
- AGA Clinical Practice Guideline on Pharmacological Interventions for Adults With Obesity (2022)



Case example: Cassie

- Cassie is a 47-year-old woman whose weight and several cardiovascular disease risk factor levels have been increasing for several years.
- Medical history includes hypertension, hyperlipidemia, metabolic syndrome, depression, obstructive sleep apnea, and GERD
- BMI: 34 kg/m² BP: 137/84 mmHg HR: 88 bpm Waist Circumference (WC): 35 in
- Physical exam is unremarkable
- Notable lab values:
 - Glucose 118 mg/dL A1c: 6.3 mg/dL ALT: 55 U/L eGFR: >60 mL/min/1.73m²
 - LDL: 115 mg/dL TG: 122 mg/dL HDL: 53 mg/dL
- Medications: hydrochlorothiazide, metoprolol, atorvastatin, paroxetine, metformin
- She has tried numerous diets but says she doesn't lose much weight and always regains weight quickly.

Clinical Obesity Treatment Modalities

- Intensive behavioral therapy
- Structured or medically monitored diets
- Pharmacotherapy
- Surgical therapy

Clinical Obesity Treatment Modalities

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Guidelines For Behavioral Therapy

- Patients who need to lose weight should receive a comprehensive behavior management program for at least 6 months (Level A)
- Gold standard is **on-site**, high-intensity (14+ sessions during initial 6 months) comprehensive intervention, either individually or in a group setting, **delivered by trained interventionist** and **persisting for at least 1 year** (Level A)
- Low-moderate intensity primary care interventions have not been shown to be effective (Level A)
- Other approaches (e.g., web- or phone-based) lead to less weight loss and health improvement (Level B)

Who is a Behavioral Therapy Trained Interventionist

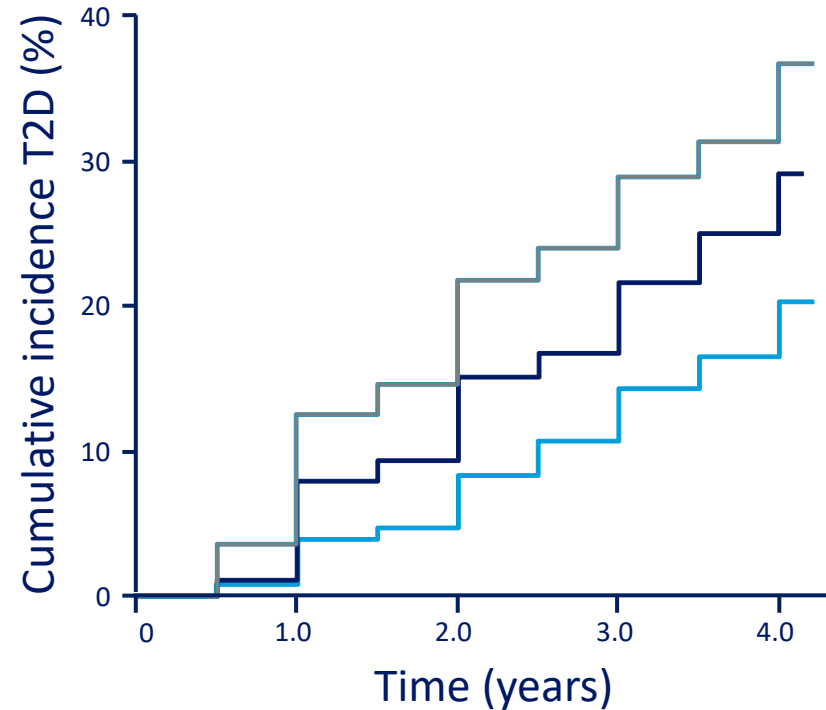
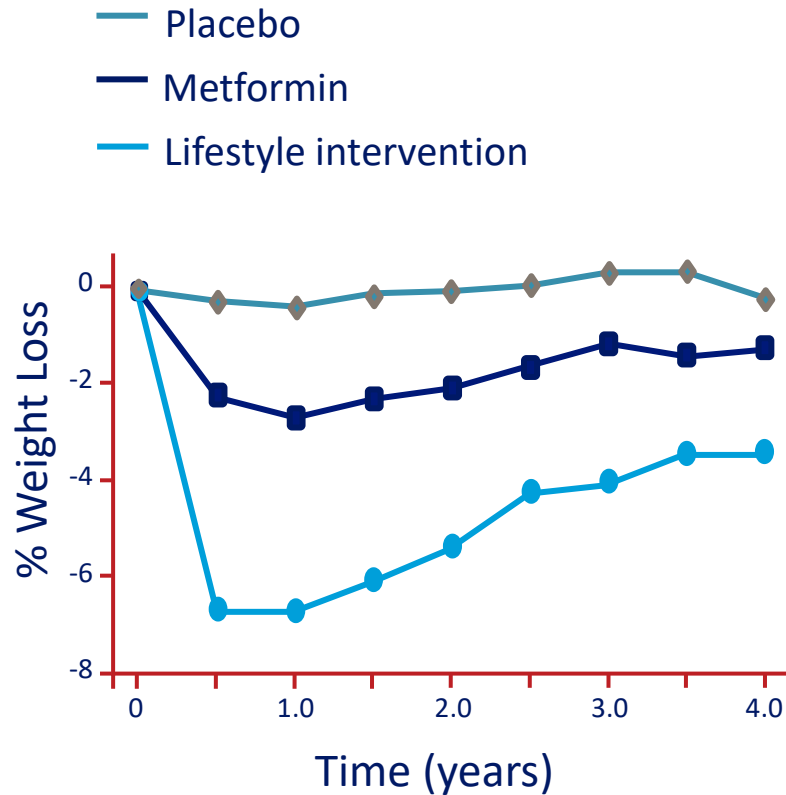
Gold standard is on-site, high-intensity comprehensive intervention, either individually or in a group setting, delivered by trained interventionist....

*“Trained interventionists included mostly health professionals (eg, registered **dietitians, psychologists, exercise specialists, health counselors,** or professionals in training) who adhered to **formal protocols in weight management**. In a few cases, lay persons were used as trained interventionists; they received instruction in weight management protocols (designed by health professionals) in programs that have been validated in high-quality trials published in peer-reviewed journals.”*

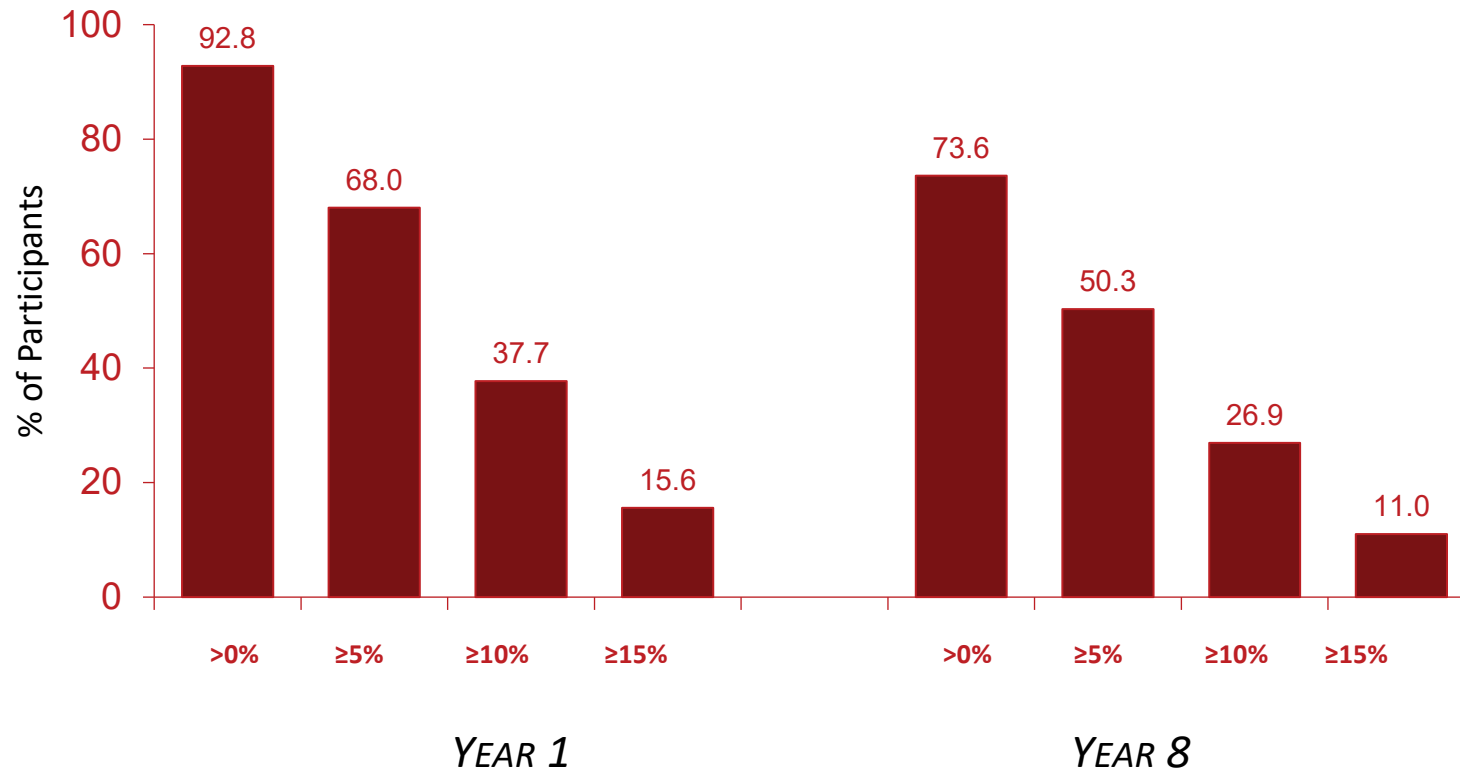
Behavioral Therapy for Obesity

Counseling	<ul style="list-style-type: none">• Regular interaction via group or individual contact<ul style="list-style-type: none">– Intensive initial counseling frequency
Diet	<ul style="list-style-type: none">• Calorie-reduced diet<ul style="list-style-type: none">– 1200-1500 kcal for <250 lb; 1500-1800 kcal for ≥250 lb
Physical Activity	<ul style="list-style-type: none">• 150 minutes/week of moderate activity• Strength training desirable
Behavioral Strategies	<ul style="list-style-type: none">• Structured curriculum of behavior change education, including identifying target behaviors and building skills to achieve target behaviors• Self-monitoring of food intake, physical activity, and/or weight• Goal setting, problem solving, stimulus control• Addressing barriers to change• Behavioral resources (e.g., portion-controlled meals)• Regular feedback and guidance from an interventionist• Weight maintenance strategies and relapse prevention• Nutritional supplements have not been shown to be effective for weight loss

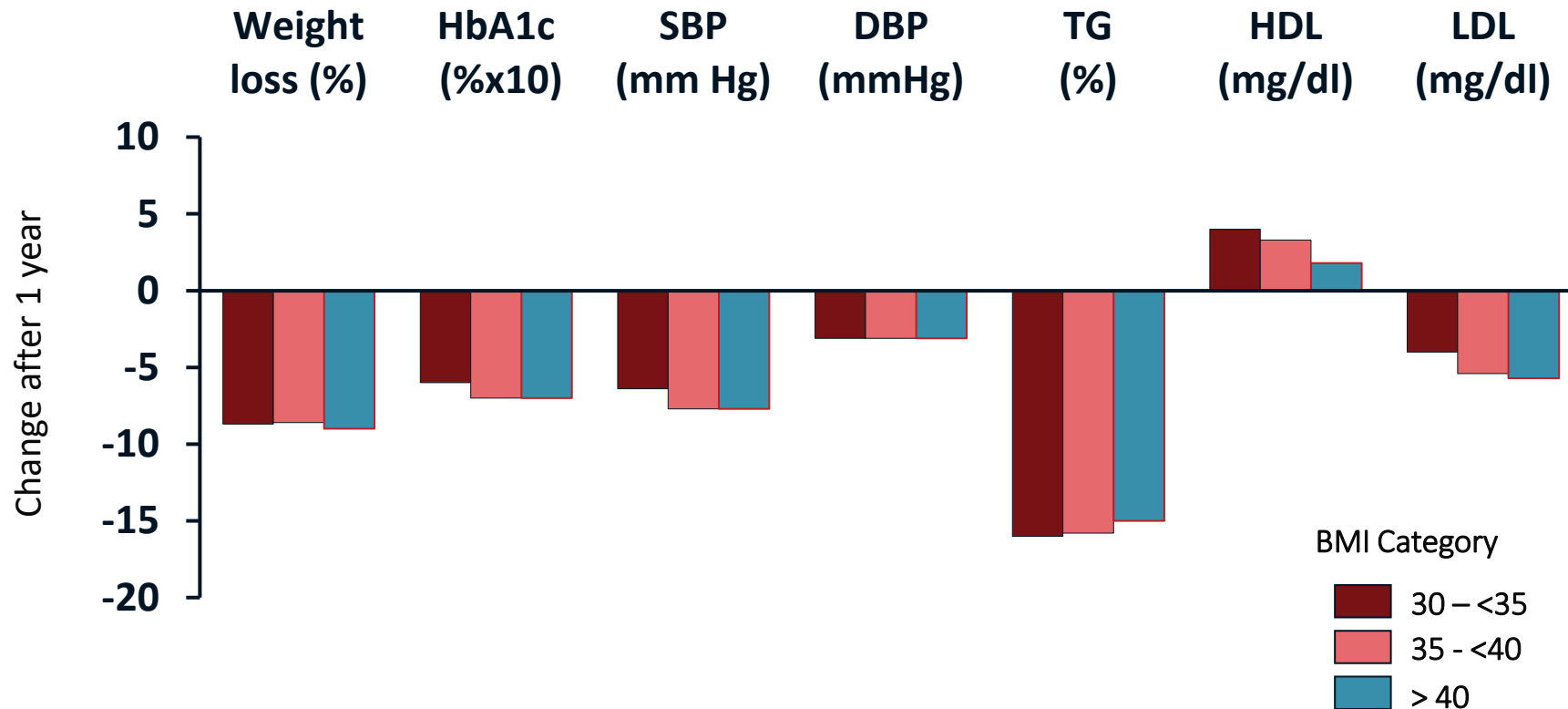
Modest Weight Loss Improves Health and Health Risks



Behavioral Therapy in Patients with Obesity and Diabetes



Comorbidity Improvements With Behavioral Therapy



Case example: Cassie, 6 months later

- Cassie continued attention to healthful dietary intake and exercise, worked one-on-one with a dietitian, and engaged at a local Diabetes Prevention Program group.
- Weight was down just 2 lbs. Blood pressure and heart rate was similar.
- Repeat labs showed glucose 111 mg/dL and A1c 6.2 mg/dL.
- She's frustrated with the lack of weight loss and concerned that even if she loses more weight that she'll regain it, as in past weight loss attempts.

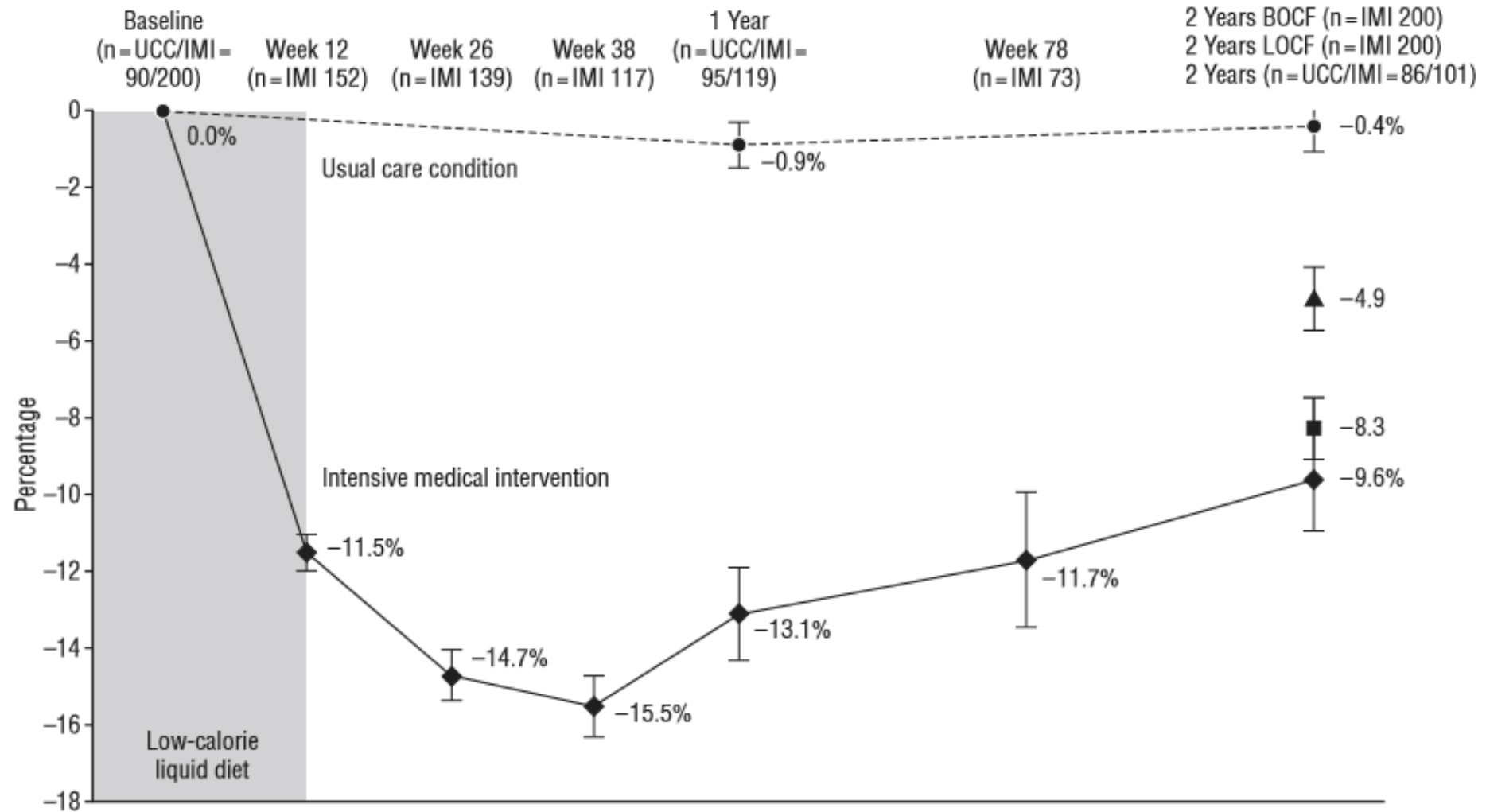
When Standard Behavioral Therapy Isn't Enough

- How do we escalate treatment for those who don't respond to standard behavioral therapy?
- How do we enhance initial weight loss for those who don't achieve sufficient weight loss to improve health status/risks?
- How do we enhance longer-term weight maintenance and minimize regain?

Clinical Obesity Treatment Modalities

- Intensive behavioral therapy
- **Structured or medically monitored diets**
- Pharmacotherapy
- Surgical therapy

Very Low-Calorie Diet with Meal Replacement Products



Clinical Obesity Treatment Modalities

- Intensive behavioral therapy
- Structured or medically monitored diets
- **Pharmacotherapy**
- Surgical therapy

Prioritize Weight-Losing or Weight-Neutral Medications

Whenever possible, minimize medications for comorbid conditions that are associated with weight gain.

	Weight Gain Associated With Use	Alternatives (Weight Reducing in Parentheses)
Diabetes	Insulin, sulfonylureas, TZDs, mitiglinides	(Metformin), (acarbose), (miglitol), (pramlintide), (GLP1 agonists/GIP), (SGLT2 inhibitors)
Hypertension medications	β -blocker	ACE inhibitors, calcium channel blockers, angiotensin-2 RAs
Antidepressants and mood stabilizers	Amytriptyline, doxepin, imipramine, nortriptyline, trimipramine, mirtazapine, paroxetine	(Bupropion), nefazodone, fluoxetine
Oral contraceptives	Progestational steroids	Barrier methods, intrauterine devices

Guidelines For Pharmacotherapy

- Use pharmacotherapy as adjunct to diet, exercise, and behavioral counseling for adults... (Level 1, strong evidence)
 - with BMI 30+; or 27+ with comorbidity;
 - who are unable to lose and successfully maintain weight;
 - who meet label indications
- Assess efficacy and safety monthly for the first 3 months, then every 3 months thereafter (Level 2, weak evidence)
- At 3 months, if loss is 5% or more, continue; if not, discontinue and seek alternative approaches (Level 1, strong evidence)
- Use medications to promote long-term weight loss maintenance (Level 2, weak evidence)
- Use weight-losing and weight-neutral medications as first and second line therapy and discuss weight effects of medications with patients (Level 1, strong evidence)

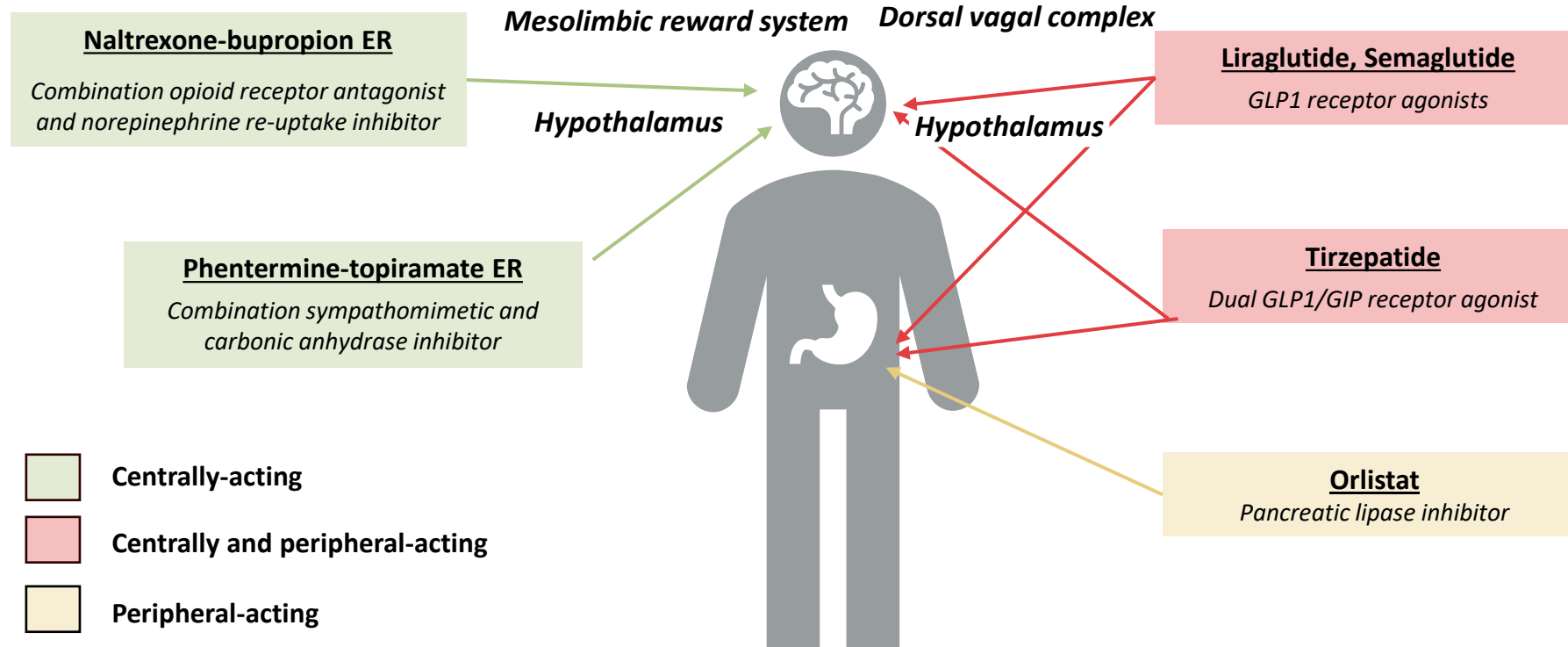
General Considerations for Pharmacotherapy

- BMI ≥ 30 kg/m² or BMI ≥ 27 kg/m² with comorbidity
- Unable to improve weight/health sufficiently with guided lifestyle changes alone
- Not pregnant, trying to conceive, or breastfeeding
- In general, without unstable cardiac disease, uncontrolled hypertension (SBP >180 mm Hg, DBP >110 mm Hg), or eating disorder
- Note specific contraindications per medication

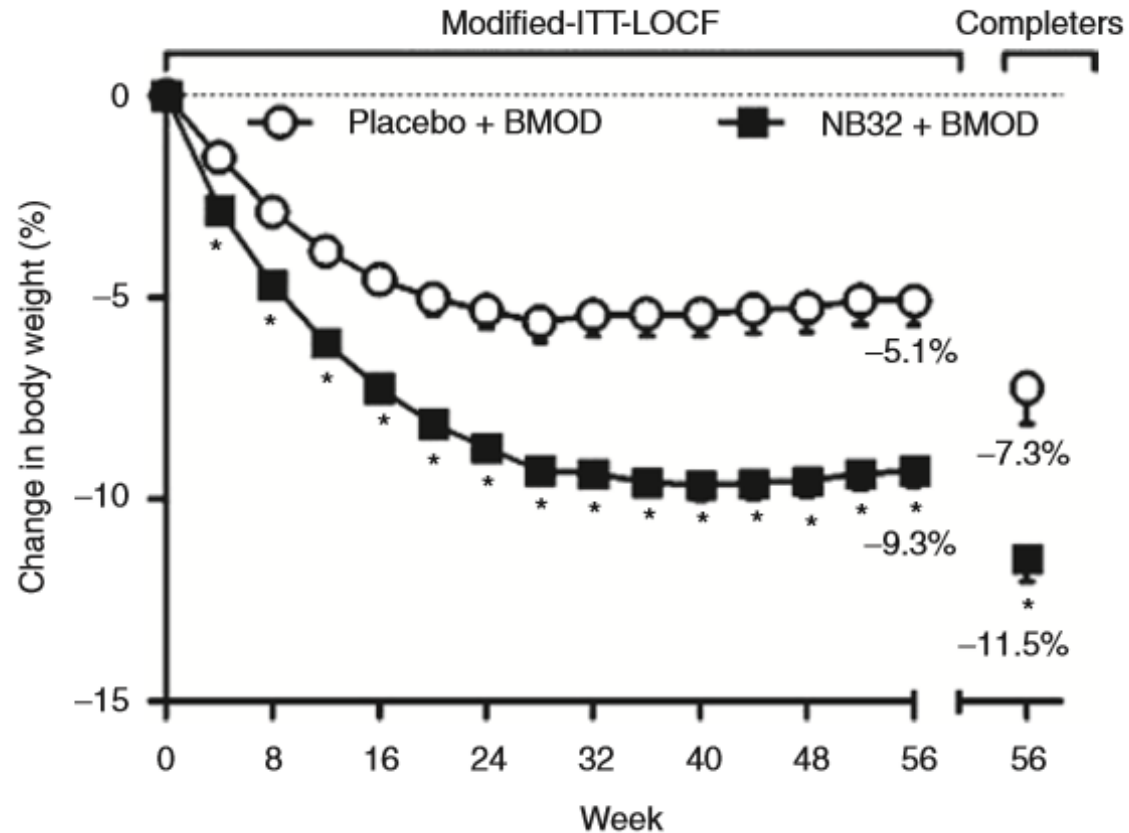
Obesity Pharmacotherapy

- 4 FDA-approved short-term medications
 - Phentermine, benzphetamine, diethylpropion, and phendimetrazine
- 6 FDA-approved long-term medications
 - Orlistat (Xenical/Alli)
 - Phentermine/topiramate ER (Qsymia)
 - Naltrexone/Bupropion SR (Contrave)
 - Liraglutide 3.0 mg (Saxenda)
 - Semaglutide 2.4 mg (Wegovy)
 - Tirzepatide (Zepbound)

Long Term Obesity Pharmacotherapy Sites of Action



Pharmacotherapy Increases Magnitude and Likelihood of Weight Loss



Abbreviation Key:

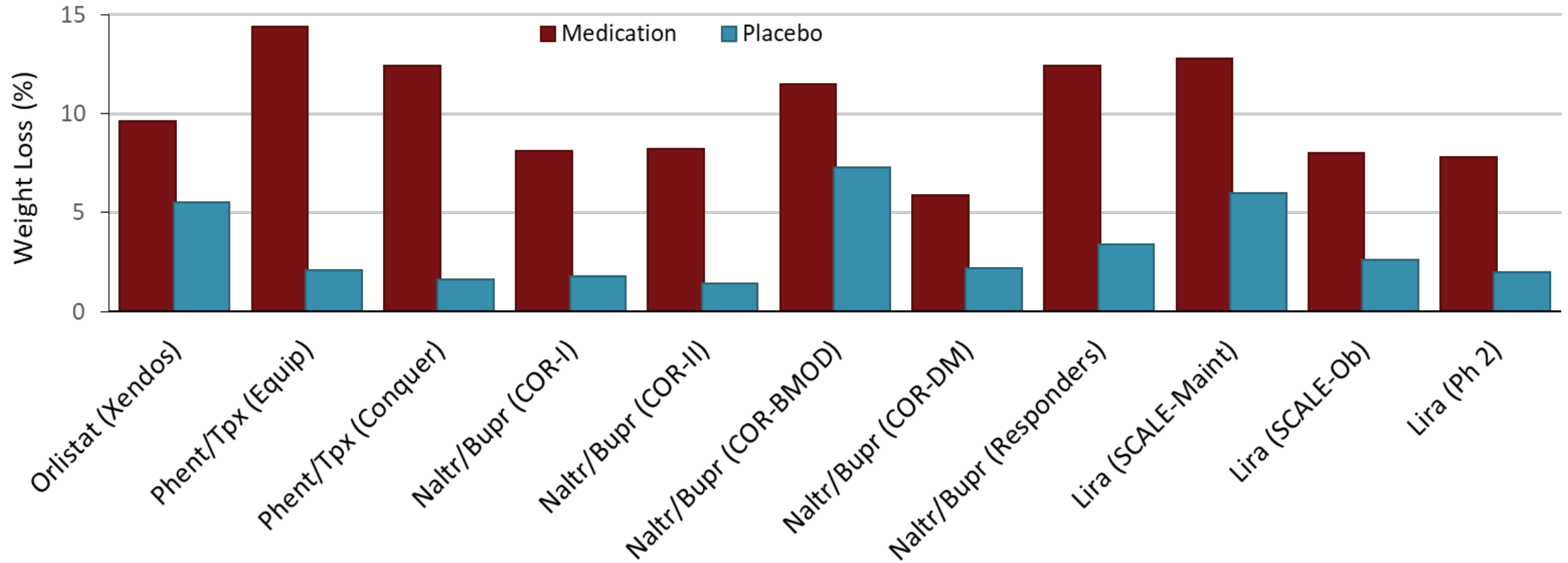
BMOD behavior modification

LOCF last observation carried forward

ITT intent to treat

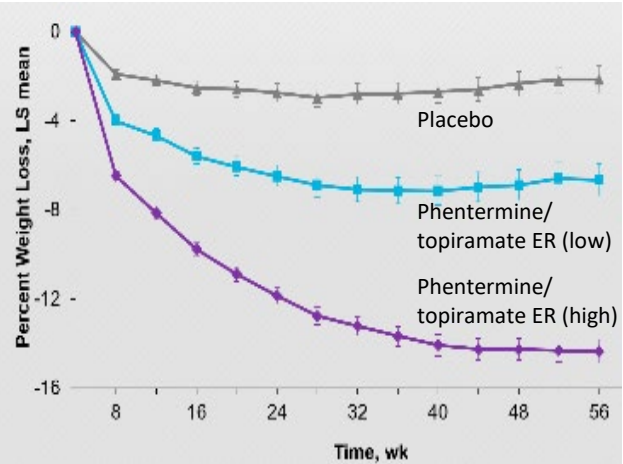
NB32 Naltrexone SR 32mg/day plus bupropion SR 360 mg/day

Pharmacotherapy Increases Magnitude and Likelihood of Weight Loss

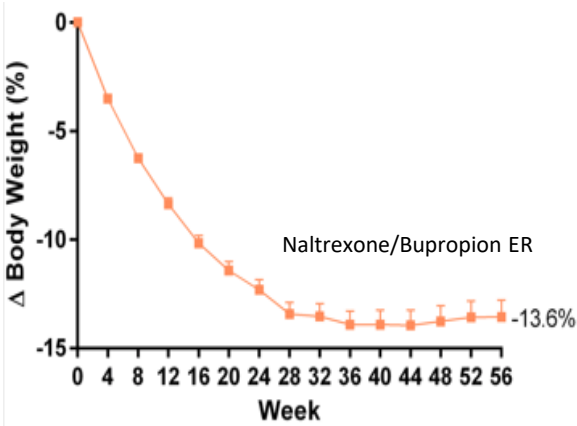


Newer Meds Lead to 10-15%+ Weight Loss in Responding Patients

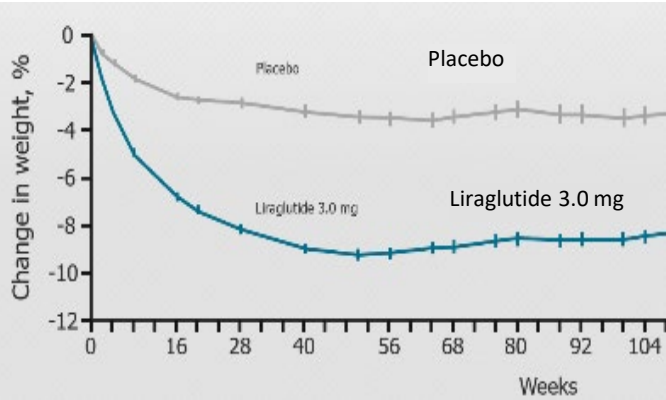
Phentermine/topiramate ER



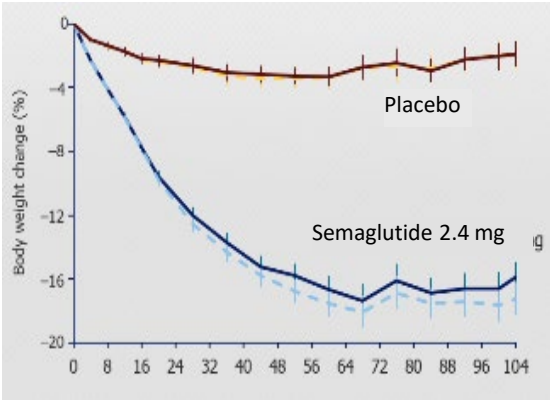
Naltrexone/bupropion ER



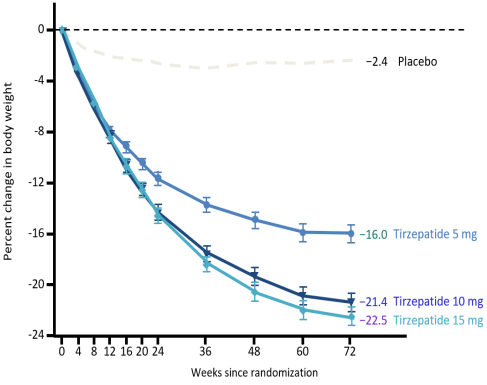
Liraglutide 3.0 mg



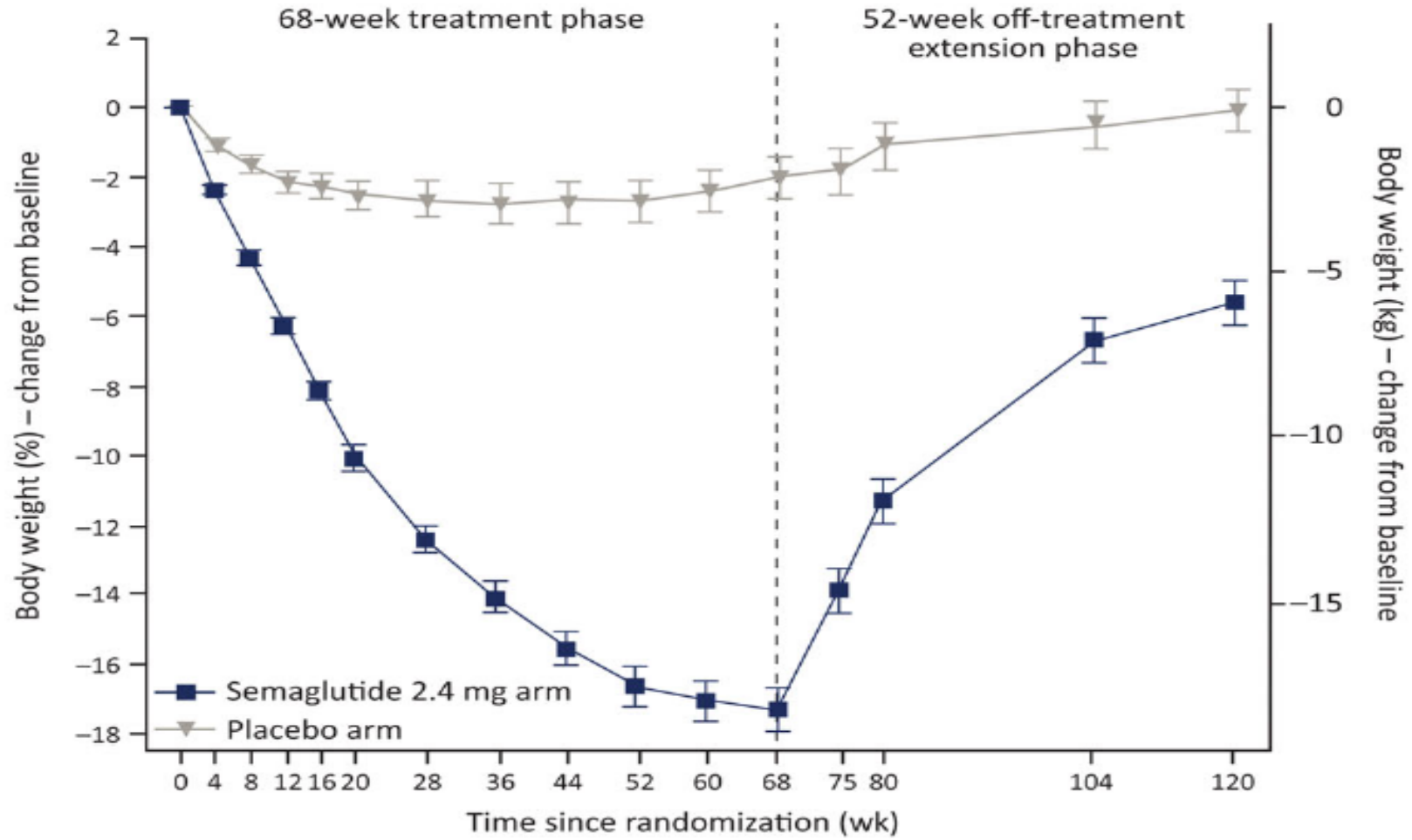
Semaglutide 2.4 mg



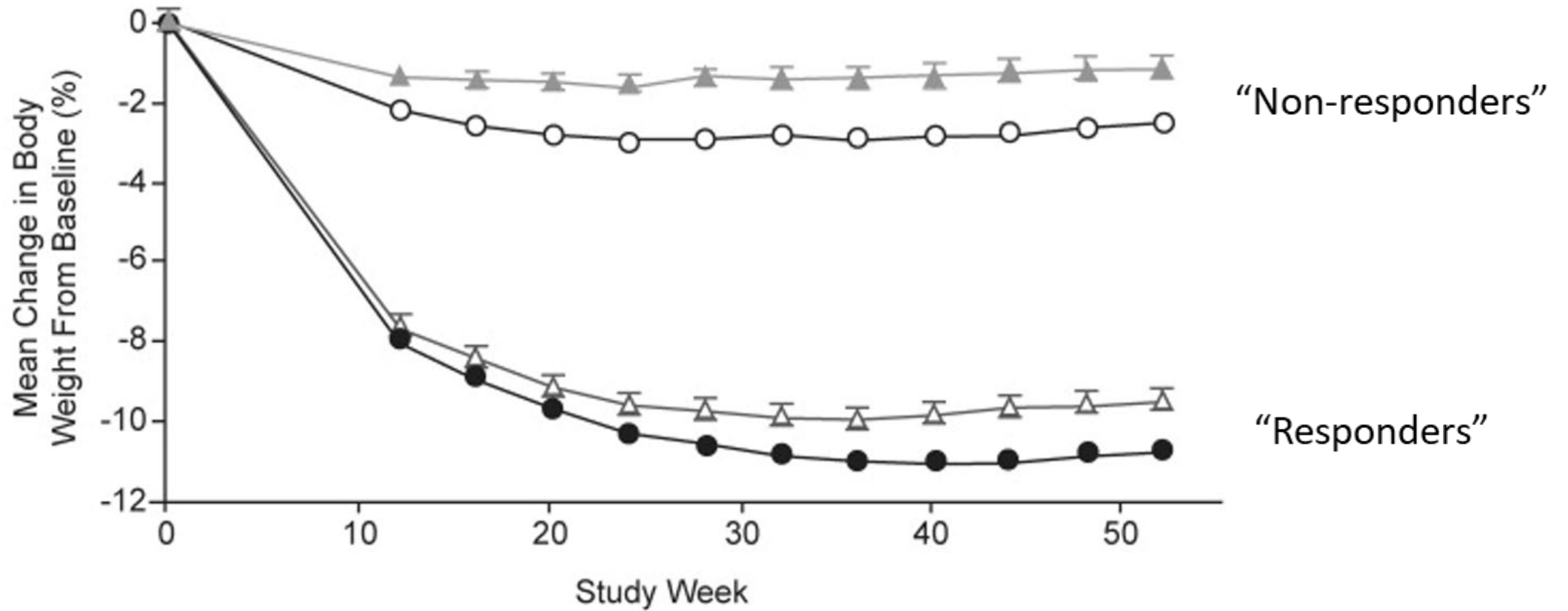
Tirzepatide



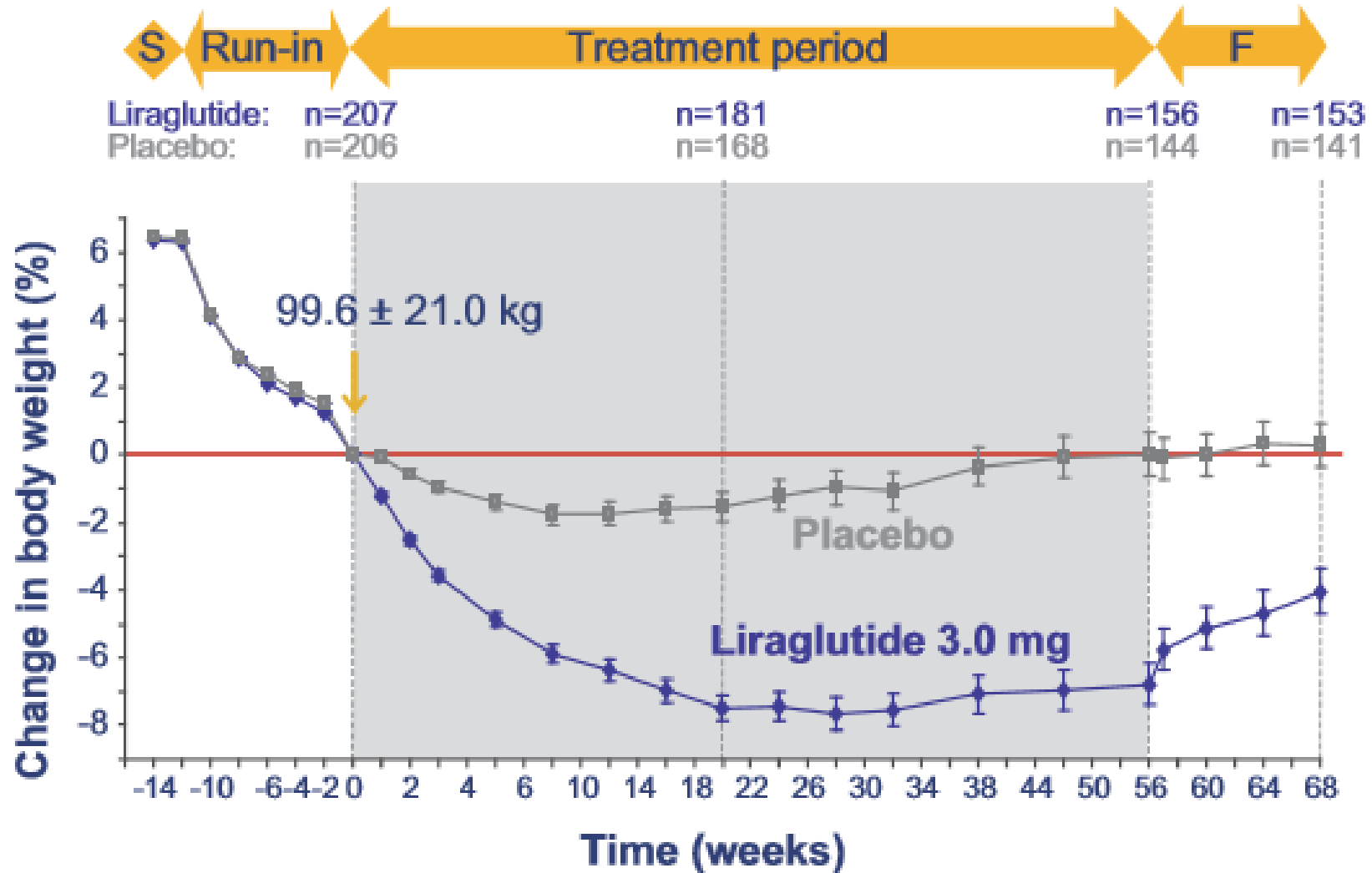
Long-Term Use Is Standard



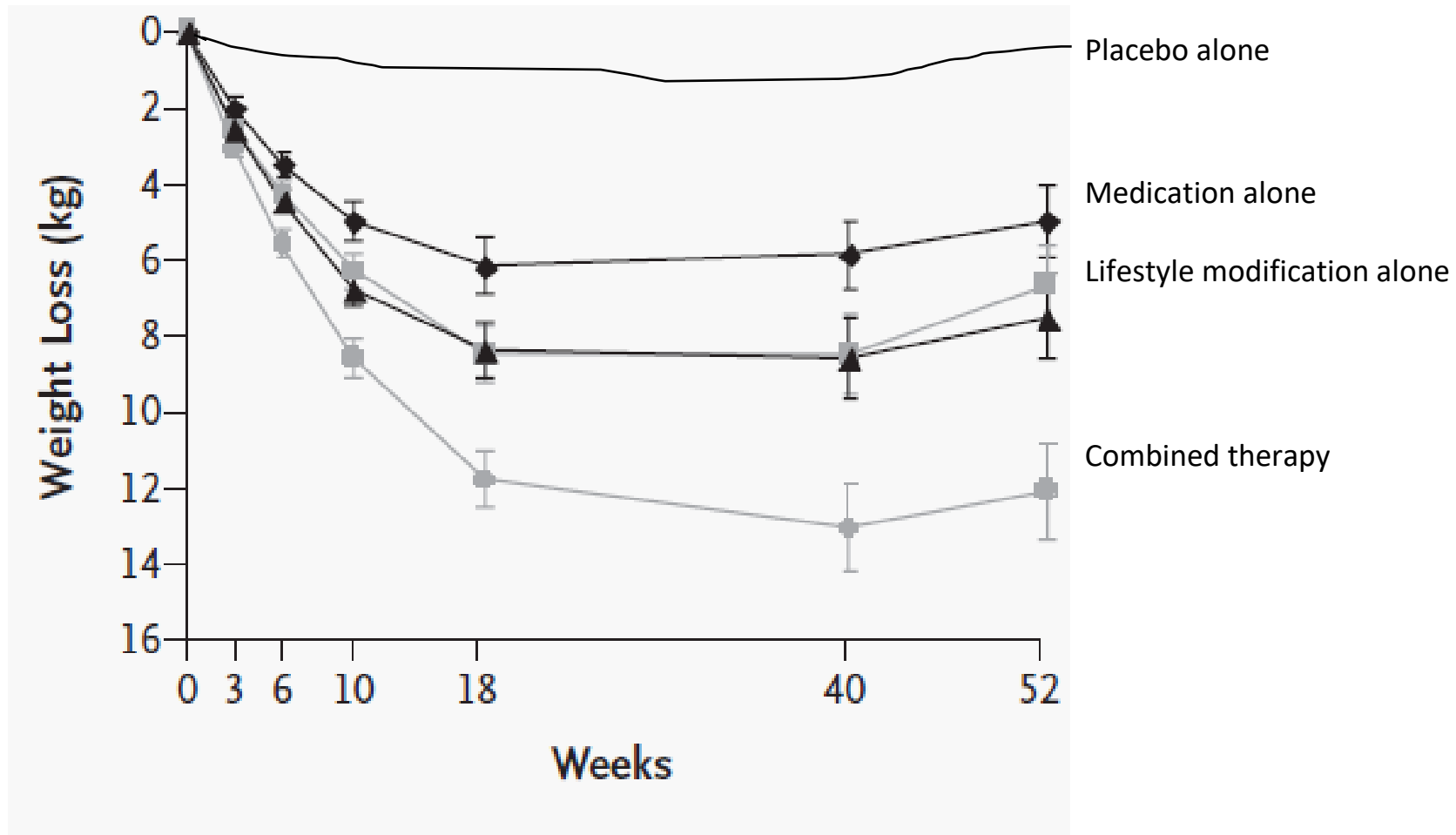
Outcomes by Responder Status



Pharmacotherapy Improves Weight Maintenance



Lifestyle Modification and Pharmacotherapy

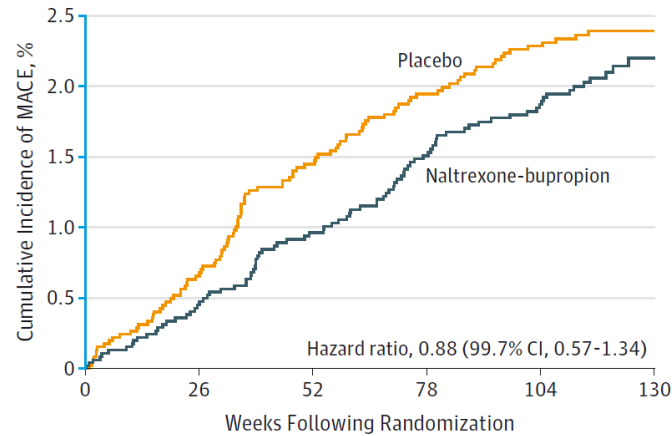


Pharmacotherapy Improves Risk Factors and Comorbidities

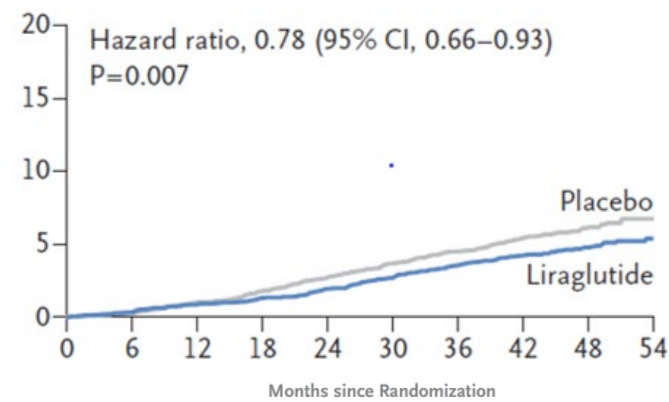
	Orlistat	Phentermine/ Topiramate ER	Naltrexone/ Bupropion ER	Liraglutide 3.0 mg	Semaglutide 2.4 mg	Tirzepatide
WC	↓	↓	↓	↓	↓	↓
BP	↓	↓	↑	↓	↓	↓
HR	↓	–	↑	↑	↑	↑
LDL-C	↓	↓	↓	↓	↓	↓
HDL-C	↑	↑	↑	↑	↑	↑
TG	↓↓↓	↓↓↓	↓↓↓	↓↓↓	↓↓↓	↓↓↓
A1C	↓	↓	↓	↓↓↓	↓↓↓	↓↓↓
Diabetes	↓	↓	↓	↓↓↓	↓↓↓	↓↓↓

Long-Term Cardiovascular Safety

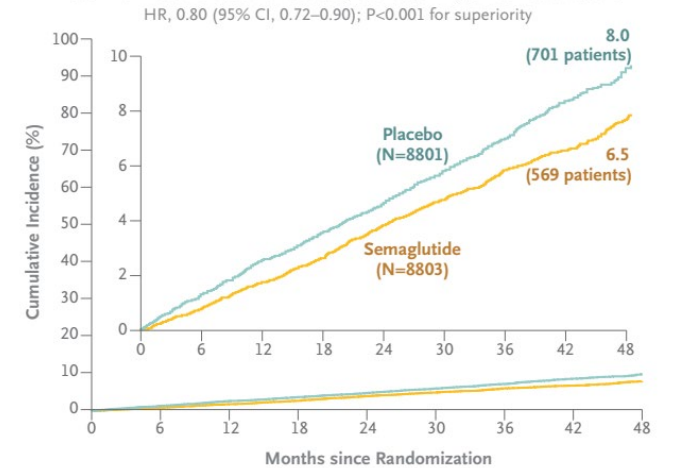
Time to MACE in 50% interim analysis



Death from Cardiovascular Causes



Death from Cardiovascular Causes, Nonfatal MI, or Nonfatal Stroke



Obesity Pharmacotherapy

- 4 FDA-approved short-term medications
 - Phentermine
 - Less commonly utilized benzphetamine, diethylpropion, and phendimetrazine
- 6 FDA-approved long-term medications
 - Orlistat (Xenical/Alli)
 - Phentermine/topiramate ER (Qsymia)
 - Naltrexone/Bupropion SR (Contrave)
 - Liraglutide 3.0 mg (Saxenda)
 - Semaglutide 2.4 mg (Wegovy)
 - Tirzepatide (ZepBound)

Phentermine

- Sympathomimetic amine
- Approved in 1959 for short-term use (schedule IV)
- **Dosing:** 8 to 37.5 mg PO QD 2 hours after breakfast
- **Side effects:** Dry mouth, insomnia, dizziness, irritability, increased blood pressure, elevated heart rate
- **Contraindications:** pregnancy, MAOIs, glaucoma, drug abuse history, hyperthyroidism
- **Relative contraindications:** CVD, CHF, CVA, uncontrolled hypertension, tachycardia, arrhythmia

CHF = congestive heart failure; CVA = cerebrovascular accident; CVD = cardiovascular disease; MAOI = monoamine oxidase inhibitor

Orlistat

- Lipase inhibitor
- Approved in 1999 for long-term use
- **Dosing:** 120 mg TID PO with meals (prescribed) or 60 mg TID (OTC)
 - Use MVI with fat-soluble vitamins at bedtime
 - May affect absorption of some medications
- **Side effects:** abdominal pain, flatulence, fecal urgency
- **Contraindications:** pregnancy, chronic malabsorption syndrome, cholestasis, and severe renal impairment

MVI = multivitamin; OTC = over-the-counter; TID = 3 times daily.

Phentermine/Topiramate ER

- Sympathomimetic amine (phentermine) and carbonic anhydrase inhibitor (topiramate)
- Approved in 2012 for long-term use (schedule IV)
- **Dosing:** Titration from 3.75/23 mg PO QAM for the 1st two weeks, to 7.5/46 mg PO QAM, titration of dosing up through week 13 to 15/92 mg PO QAM max dose titrated for WL >3% of baseline BMI
 - Discontinue if less than 3% weight loss after 12 weeks
- **Side effects:** Constipation, paresthesia, insomnia, nasopharyngitis, xerostomia, increased blood pressure
- **Contraindications:** pregnancy, glaucoma, MAOIs, hyperthyroidism

Naltrexone/Bupropion ER

- Opioid receptor antagonist (naltrexone) and dopamine/norepinephrine reuptake inhibitor (bupropion)
- Approved in 2014 for long-term use
- **Dosing:** 32/360 mg tablet, titration from 1 PO QD to 2 PO BID over the course of 4 weeks (not to exceed 2 tablets BID)
 - Discontinue if <5% weight loss after 16 weeks
- **Warning for suicidal thoughts in adolescents**
- **Side effects:** Constipation, nausea, headache, xerostomia, insomnia, elevated heart rate and blood pressure
- **Contraindications:** pregnancy, MAOI, uncontrolled hypertension, seizure disorder, chronic opioid use

BID = twice daily.

Liraglutide 3.0 mg

- Glucagon-like peptide 1 (GLP-1) receptor agonist
- Approved in 2014 for long-term use
- **Daily** subcutaneous injection
- **Dosing:** weekly escalation from 0.6 mg to 3.0 mg SQ daily
 - Discontinue if <4% weight loss at 16 weeks
- **Hypoglycemia possible and more likely when combined with other diabetic agents**
- **Side effects:** Gastrointestinal side effects (nausea, vomiting, diarrhea, esophageal reflux), injection site reactions, elevated heart rate, hypoglycemia
- **Contraindications/warnings:** pregnancy, medullary thyroid carcinoma, pancreatitis

Semaglutide 2.4 mg

- GLP-1 receptor agonist
- Approved in 2021 for long-term use
- **Once-weekly** subcutaneous injection
- **Dosing:** monthly escalation from 0.25 mg to 2.4 mg weekly
- **Hypoglycemia possible and more likely when combined with other diabetic agents**
- **Side effects:** Gastrointestinal side effects (nausea, vomiting, diarrhea, esophageal reflux), injection site reactions, elevated heart rate, hypoglycemia
- **Contraindications/warnings:** pregnancy, medullary thyroid carcinoma, pancreatitis

Tirzepatide

- Dual GLP-1/GIP receptor agonist
- Approved in 2023 for long-term use
- **Once-weekly** subcutaneous injection
- **Dosing:** starting dose of 2.5 mg SQ weekly increased at increments 2.5 mg – 15 mg weekly till 5mg SQ once weekly, then may increase by 2.5 mg every 4 weeks up to 15 mg as needed.
- **Hypoglycemia possible and more likely when combined with other diabetic agents**
- **Side effects:** stomach pain, anxiety, bloating, blurred vision
- **Contraindications/warnings:** pregnancy, medullary thyroid carcinoma, pancreatitis

Case example: Cassie, 3 months later

- After starting an obesity medication, Cassie's weight has come down 13 lbs (6%) over 3 months. Blood pressure has improved and plan to repeat pertinent blood tests in the coming months.
- She feels more confident with her progress, though still feels the weight loss is slow and wonders if another medication may be better.

Clinical Obesity Treatment Modalities

- Intensive behavioral therapy
- Structured or medically monitored diets
- Pharmacotherapy
- **Surgical therapy**

Guidelines For Bariatric Surgery

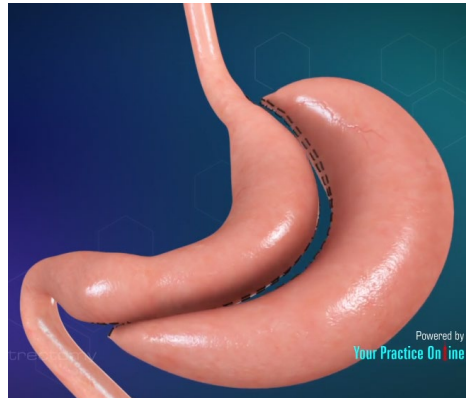
- Advise patients with BMI ≥ 40 kg/m² (or ≥ 35 kg/m² with comorbidity) that bariatric surgery may be an appropriate option to improve health (Grade A)
- Offer referral to an experienced bariatric surgeon for consultation and evaluation (Grade A)
- Insufficient evidence to recommend for or against surgery for BMI < 35 kg/m²
- No clear guidance for medical devices

Bariatric Surgery

Gastric Band



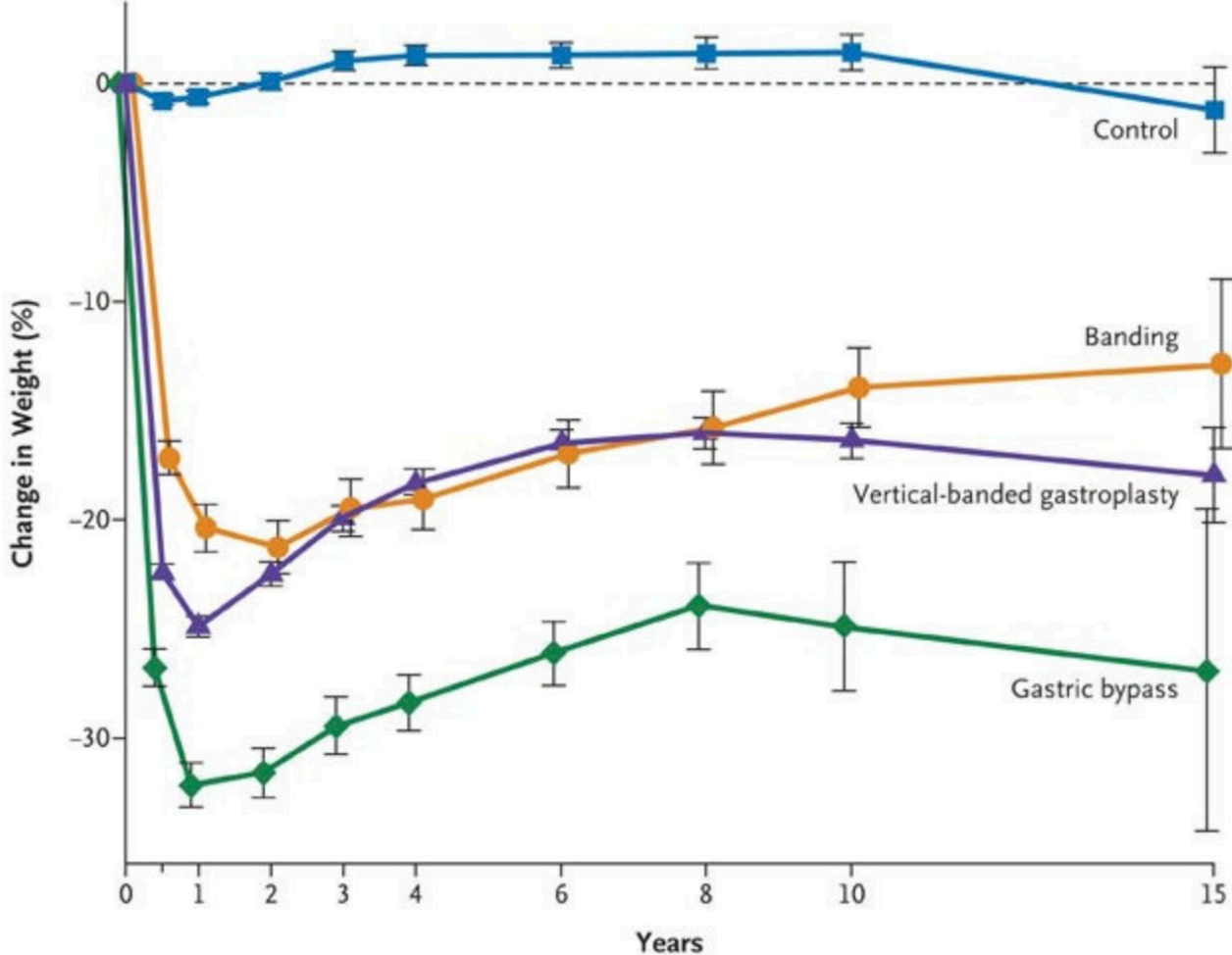
**Sleeve
Gastrectomy**



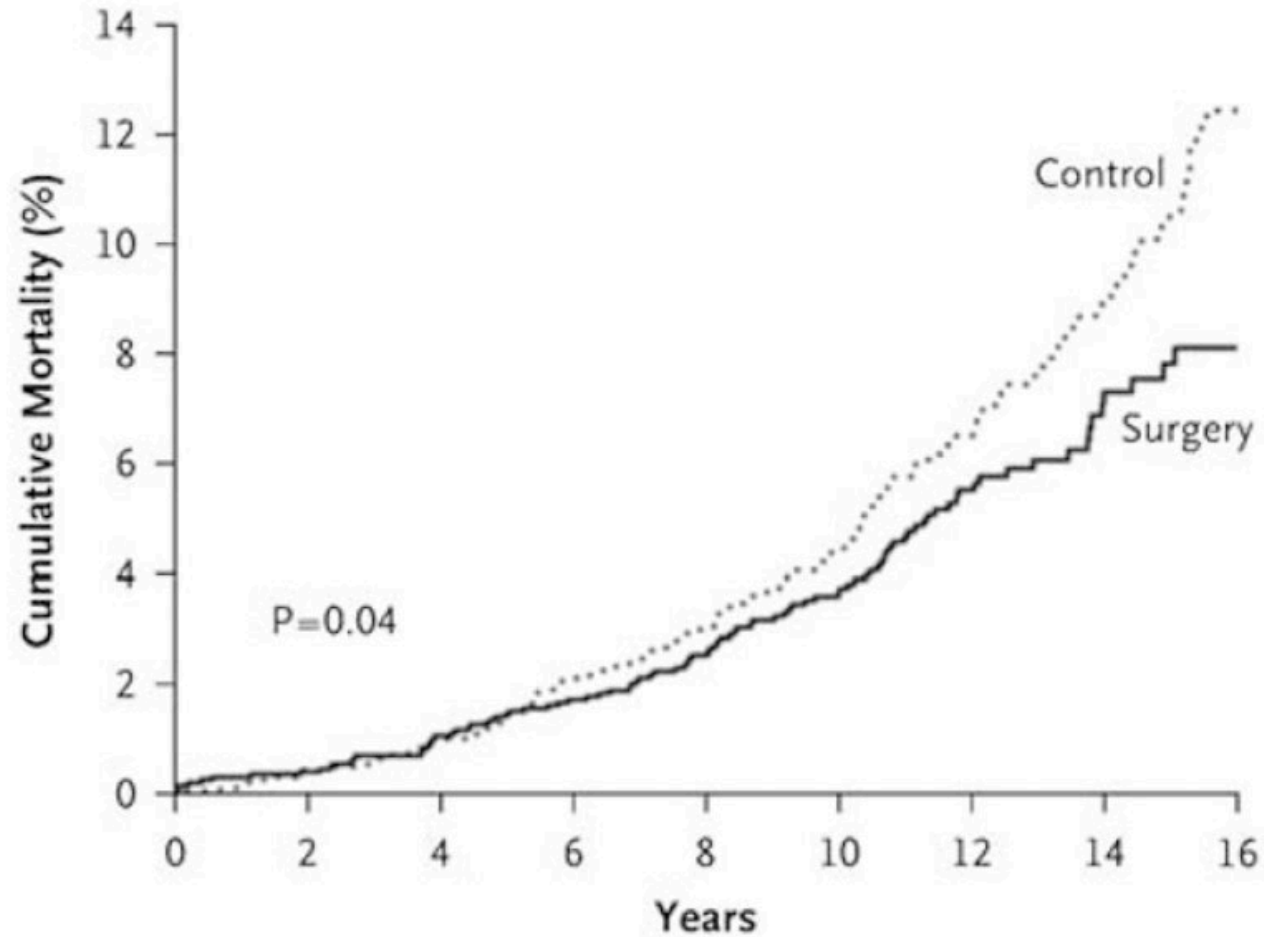
**Roux-en-Y
Gastric Bypass**



Bariatric Surgery Improves Long-term Weight Management

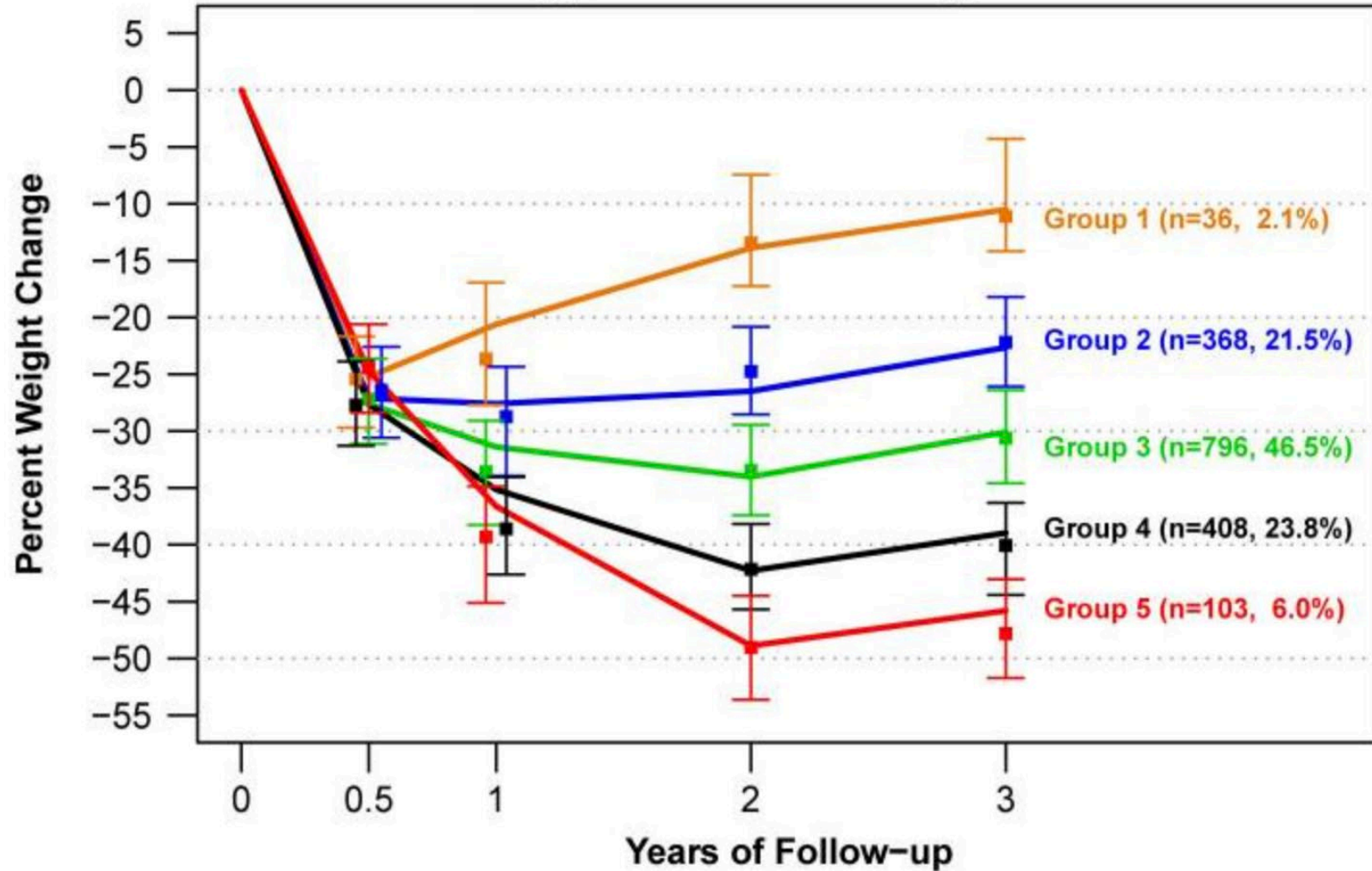


Bariatric Surgery Improves Mortality

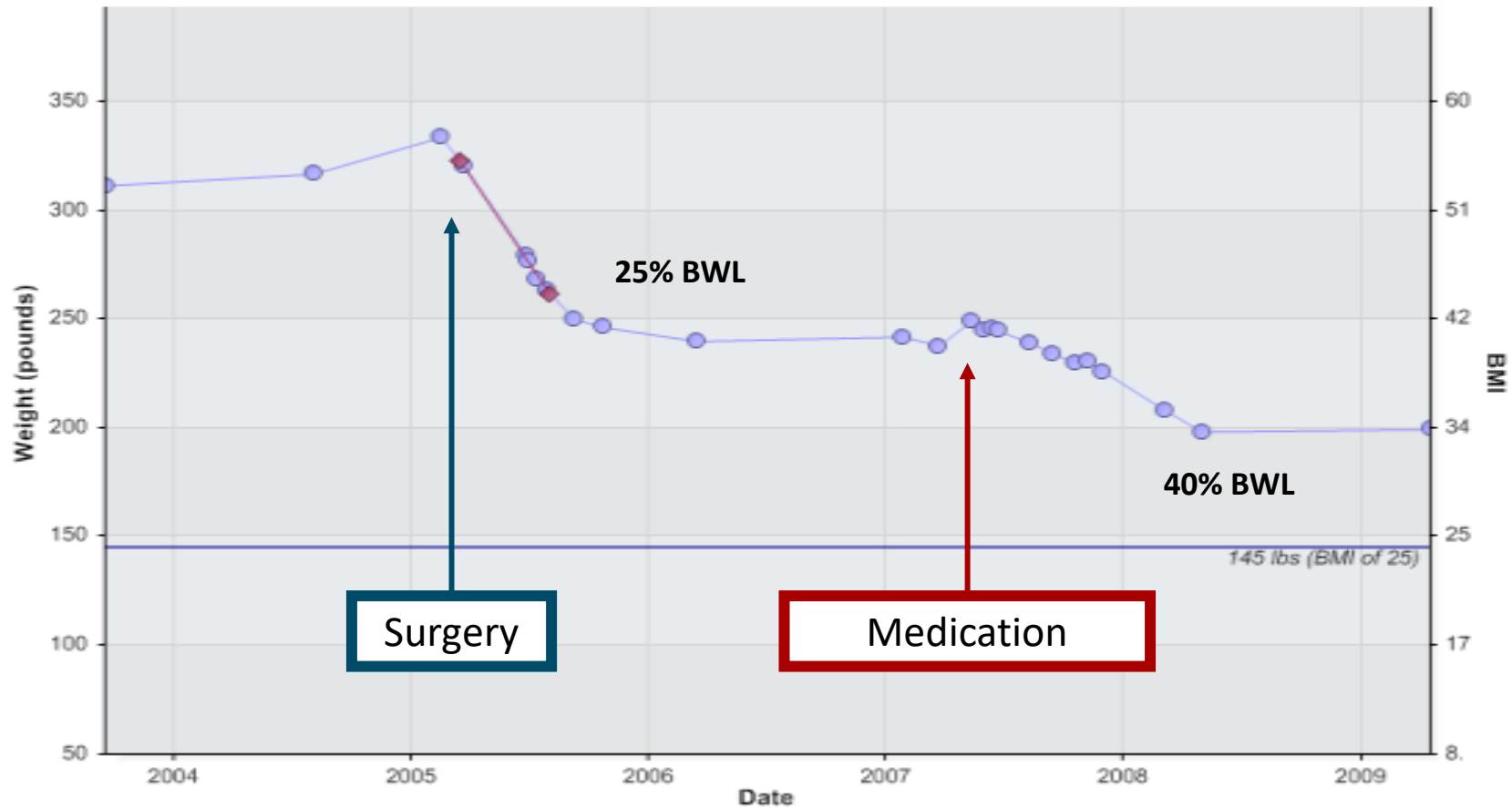


Trajectories of Weight Change After Surgery

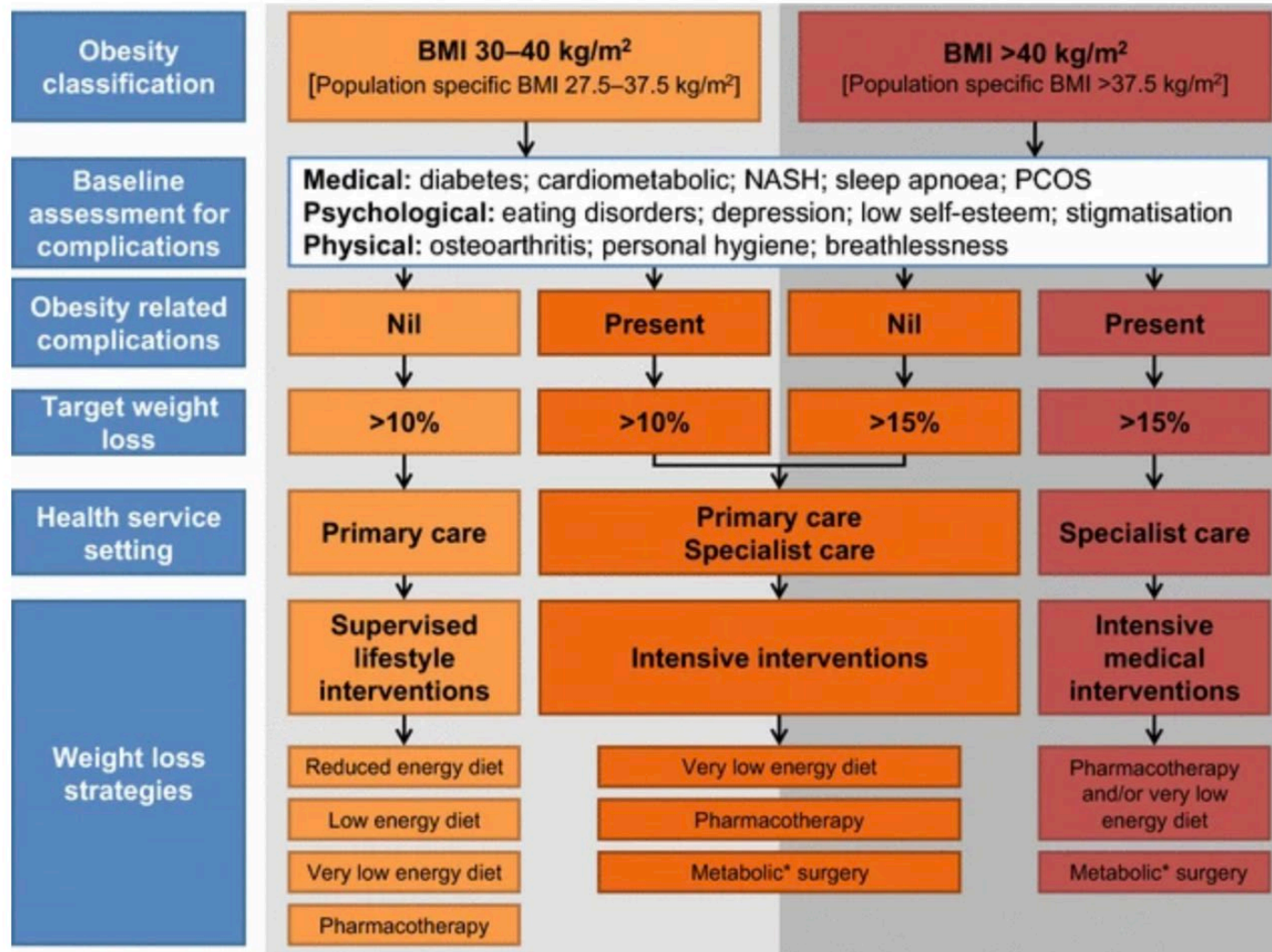
(a) Roux-en-Y Gastric Bypass



Combining Modalities Works Better



Example of Escalation of Services



Case example: Cassie, 4 months later

- Cassie's weight is now down 25 lbs (12%), her blood pressure has normalized, and lab tests show A1c now 5.7 mg/dL and lipids much improved.
- She has restarted working with a dietitian and has also been enjoying an exercise class at local YMCA club.

Conclusion

- **Pharmacological management of obesity** can significantly improve outcomes and reduce obesity-related complications.
- **Lifestyle intervention** is the foundation but often insufficient alone for most individuals.
- **Medications** should be considered for adults with a BMI ≥ 30 kg/m², or ≥ 27 kg/m² with weight-related complications, who do not respond adequately to lifestyle interventions.
- A **multidisciplinary team multipronged therapeutic approach** is often required to support patients in their weight loss journey.

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