

Obesity and Prostate Cancer What's the link and what should I do about it?

Louis J. Aronne, MD, FACP, FTOS, DABOM September 10, 2023



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Disclosures

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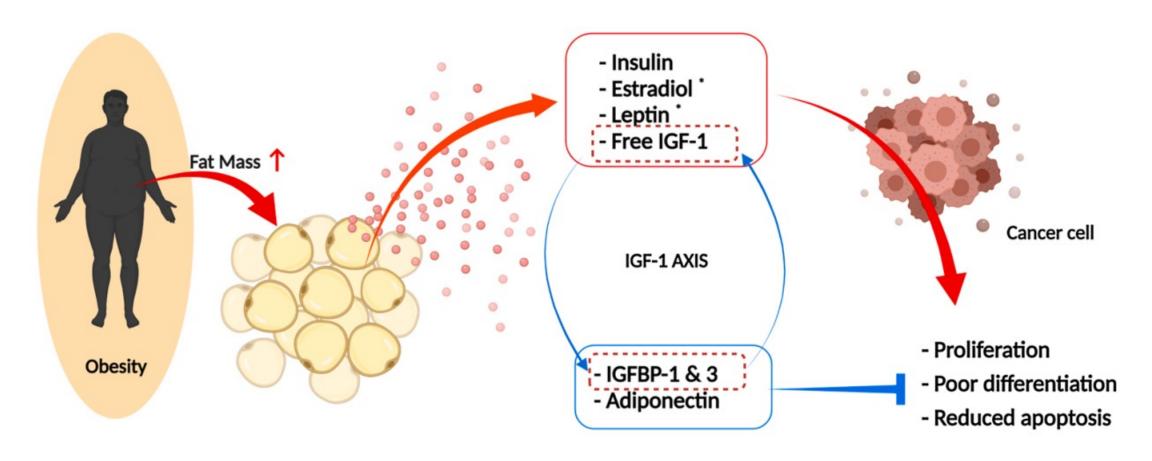
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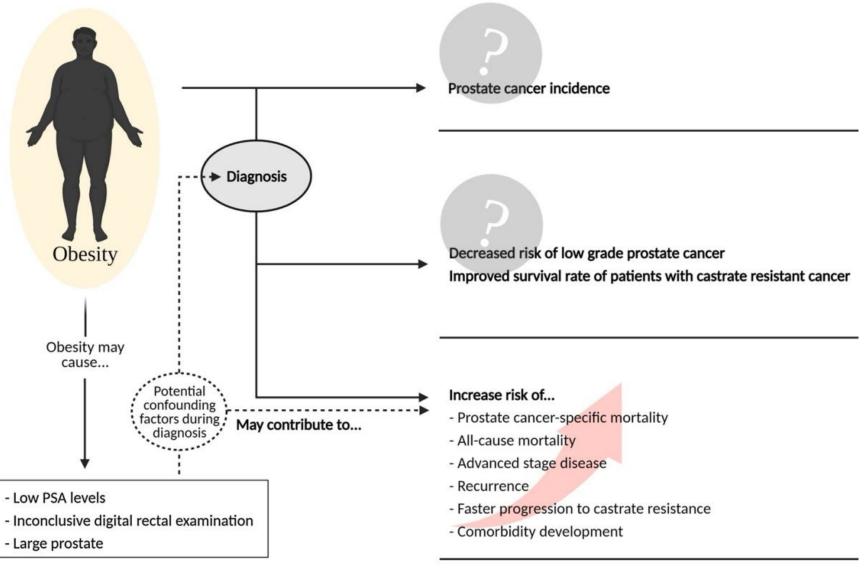


Proposed mechanisms for the association between fat mass and prostate cancer progression





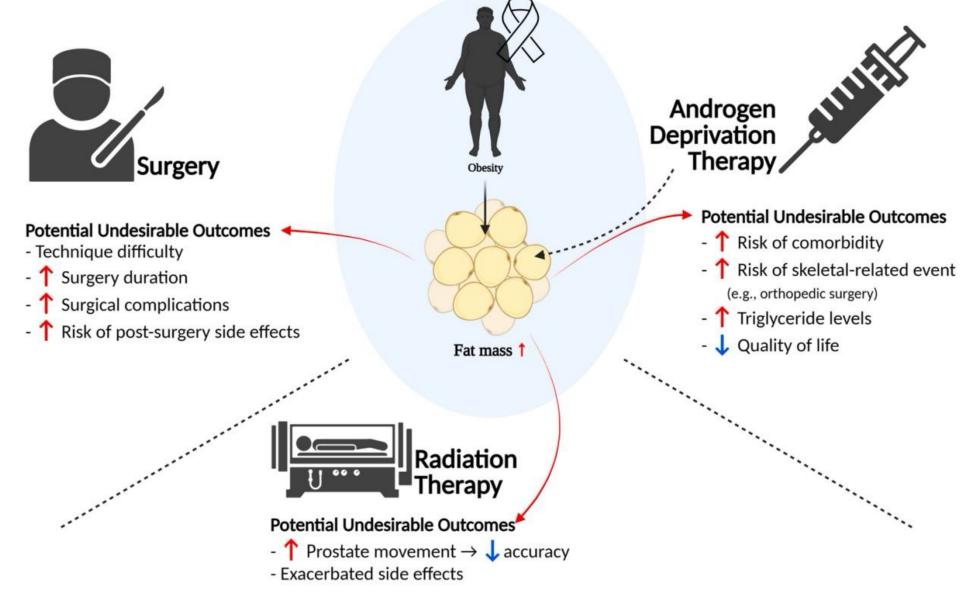
Men with obesity and prostate cancer: at increased risk of poor prognosis and general health





Wilson RL, et al. Crit Rev Oncol Hematol. 2022 Jan;169:103543.

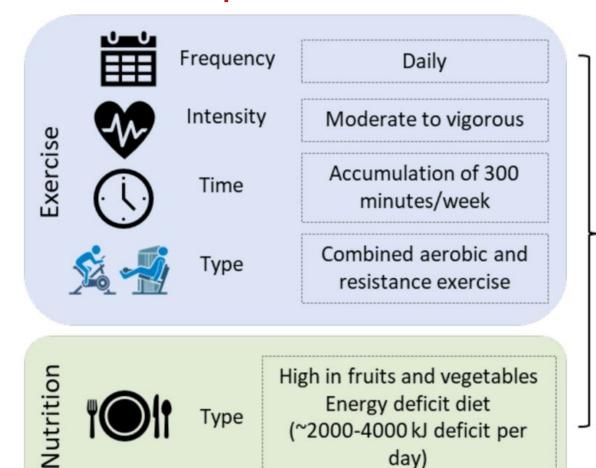
Undesirable outcomes of prostate cancer treatment





Wilson RL, et al. Crit Rev Oncol Hematol. 2022 Jan;169:103543.

Exercise and nutrition weight loss recommendations for prostate cancer patients



Considerations:

- Predominately fat mass loss.
- Timing of intervention.
- Treatment type.
- Severity of side effects and comorbidities.
- Adequate calcium and vitamin D intake and inclusion of impact training for bone maintenance.
- Adequate protein intake and resistance training for lean mass maintenance.

Weill Cornell Medicine

Does Sugar Feed Cancer? Maybe!

Diabetes and Cancer

- Epidemiology studies suggest that diabetes is strongly associated with several cancers
- High insulin, fat cell hormones, growth factors, changes of diabetes all contribute to the increase
- Metformin, a diabetes treatment, reduces the risk of cancer in people with diabetes

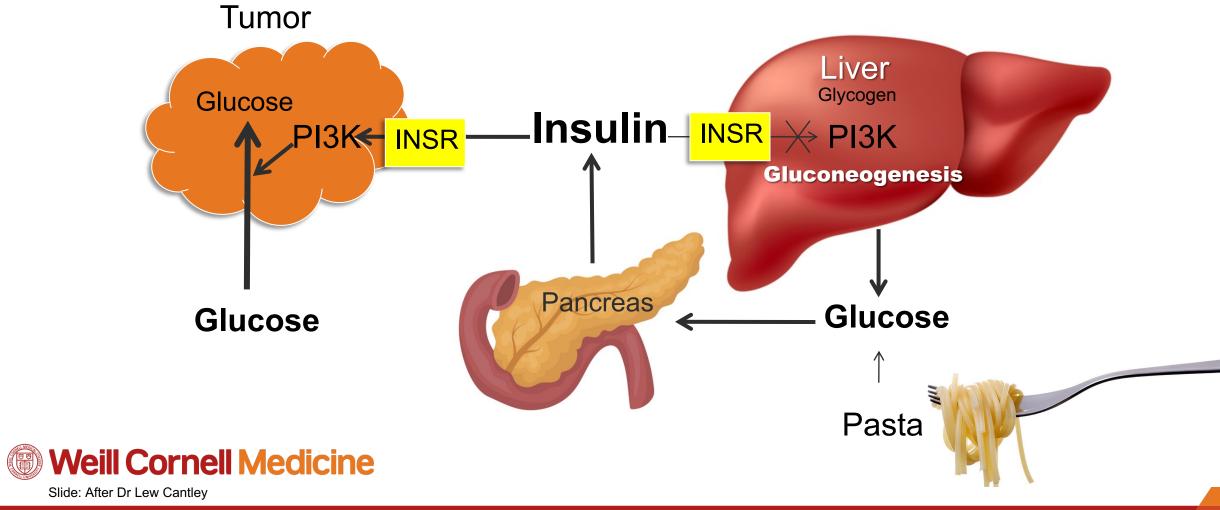


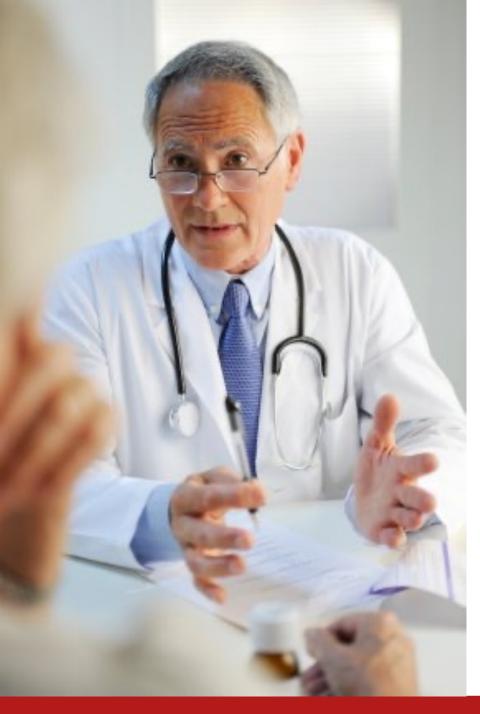


Insulin Resistance and Tumor Growth

The liver, muscle and fat are insulin resistant... but the tumor is insulin sensitive!!

It sucks up all the sugar!

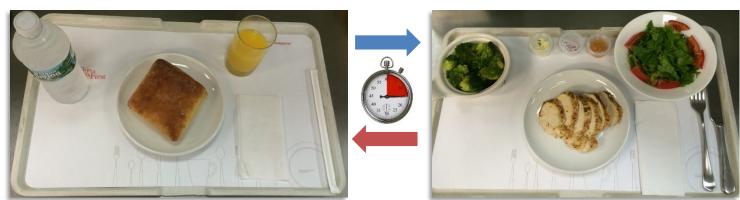




Q: "Doctor, I get the overwhelming urge to eat carbs."

A: "The order in which you consume food has a significant impact on blood sugar. So...

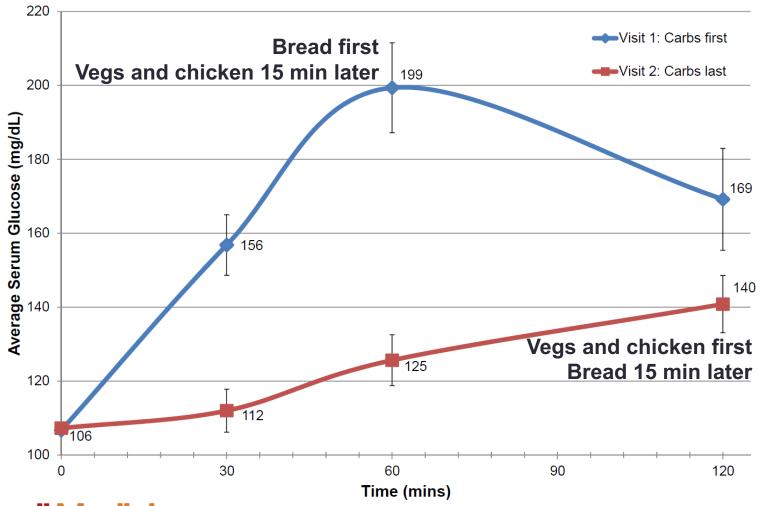
...eat your vegetables and protein first!"



Eating protein and vegetables before carbs lowers glycemic index!!

The order in which food is consumed impacts blood sugar and insulin after the meal!

Post-prandial Glucose Response





Caloric Density of Fresh vs Processed Foods





What's the best exercise?

Interval Training/Resistance Training



Examples: weight-lifting, Pilates, other activities (besides aerobics)

Twice per week provides greatest weight loss

Interval Training: Intensity of activity varies

Provides greatest weight loss

Examples: hill climbing program on bike or cross trainer

The highest level of exercise intensity sets your "metabolism"

You can maintain twice the weight loss exercising for just one-half the time if you increase the intensity



What is Metformin, and should I be taking it?

Metformin: first-line drug for treating diabetes⁵

There is increasing consensus that metformin has an anti-cancer activity

- Inhibits cellular transformation and selectively kills cancer stem cells in breast cancer cell lines⁵
- As metformin alters energy metabolism in diabetics, we speculate that metformin may block a metabolic stress response that stimulates the inflammatory pathway associated with a wide variety of cancers⁵
- "Our findings suggest that metformin could be a useful adjuvant agent, with the greatest benefits seen in colorectal and prostate cancer, particularly in those receiving radical radiotherapy, and randomised, controlled trials which investigate dose and duration, alongside efficacy, are advocated."



^{1.} Pollak MN. Cancer Discov. 2012 Sep;2(9):778-90. 2. Anisimov VN. Aging (Albany NY). 2010 Nov;2(11):760-74.

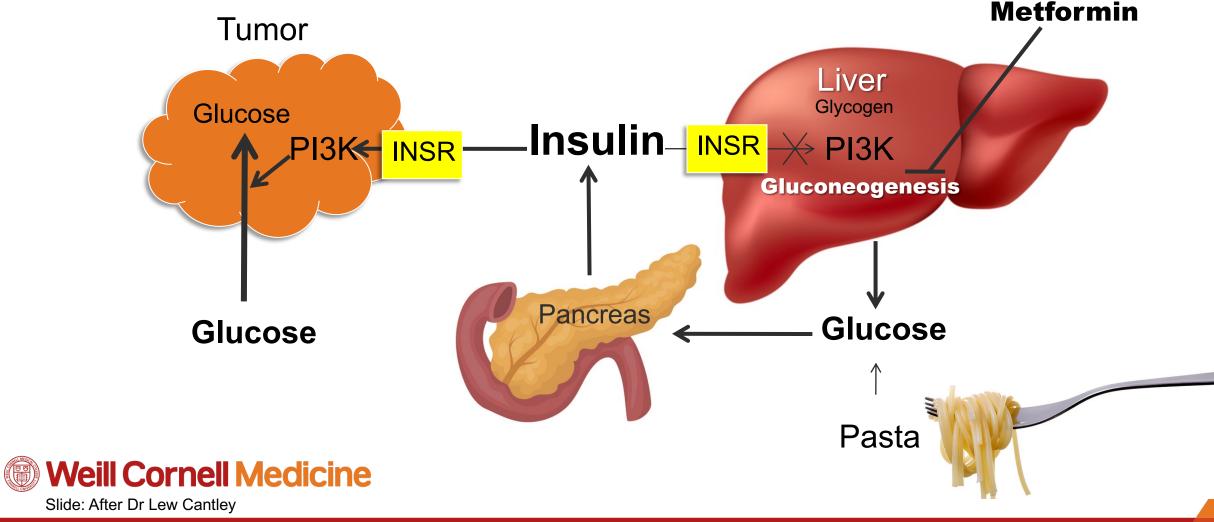
Del Barco S, et al. *Oncotarge*t. 2011 Dec;2(12):896-917. 4. Martin-Castillo B, et al. *Cell Cycle*. 2010 Mar 15;9(6):1057-64.

Hirsch HA, et al. *Proc Natl Acad Sci U S A*. 2013 Jan 15;110(3):972-7. 6. Coyle C, et al. *Ann Oncol*. 2016 Dec;27(12):2184-2195.

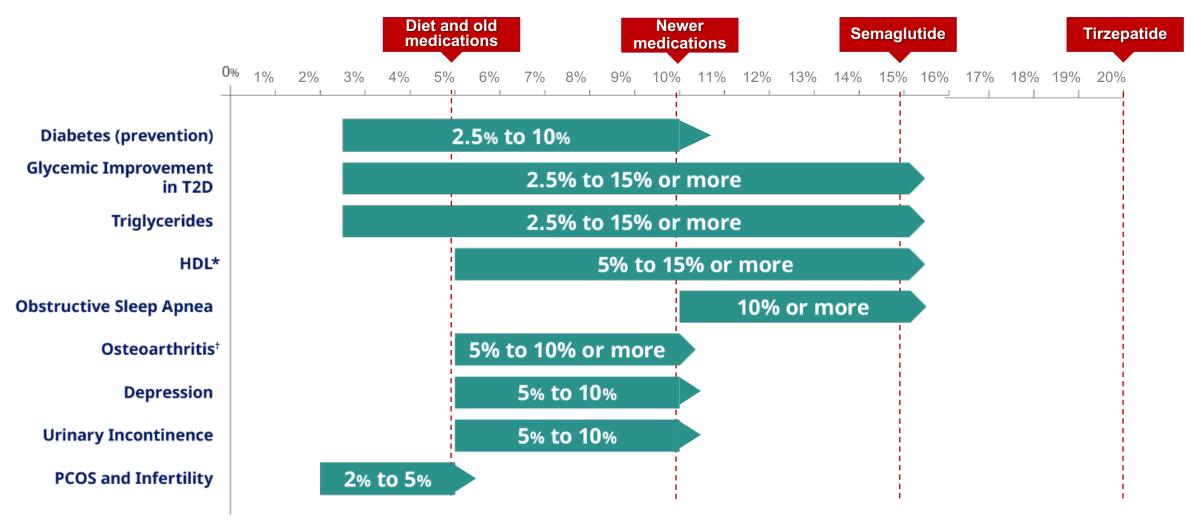
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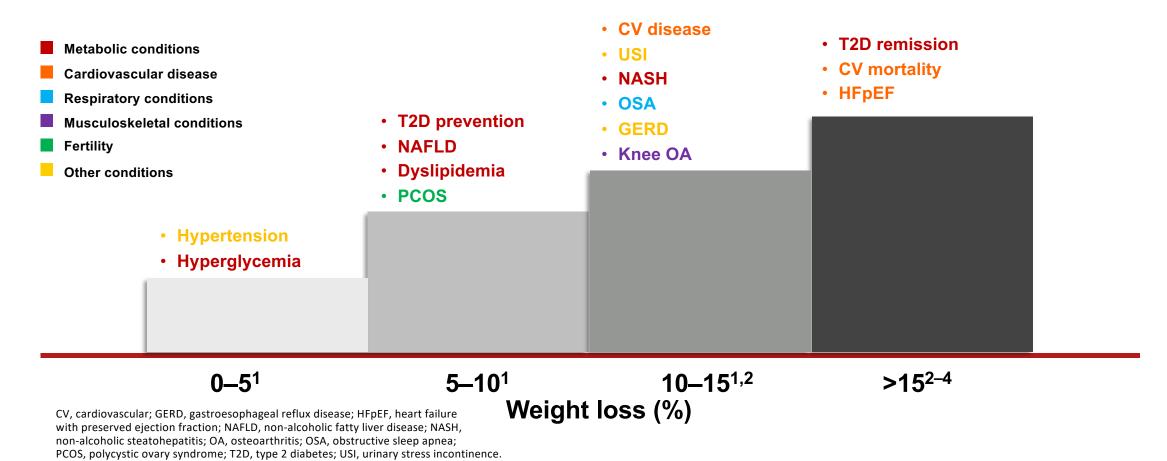
What about Weight Loss: More Weight Loss Provides More Clinical Benefit²





Ryan DH, Yockey SR. Curr Obes Rep. 2017;6(2):187-194. 2.Garvey WT, Mechanick JI, Brett EM, et al. Endocr Pract. 2016;22(suppl 3):1-203.
 Wing RR, Lang W, Wadden TA, et al. Diabetes Care. 2011;34(7):1481-1486.

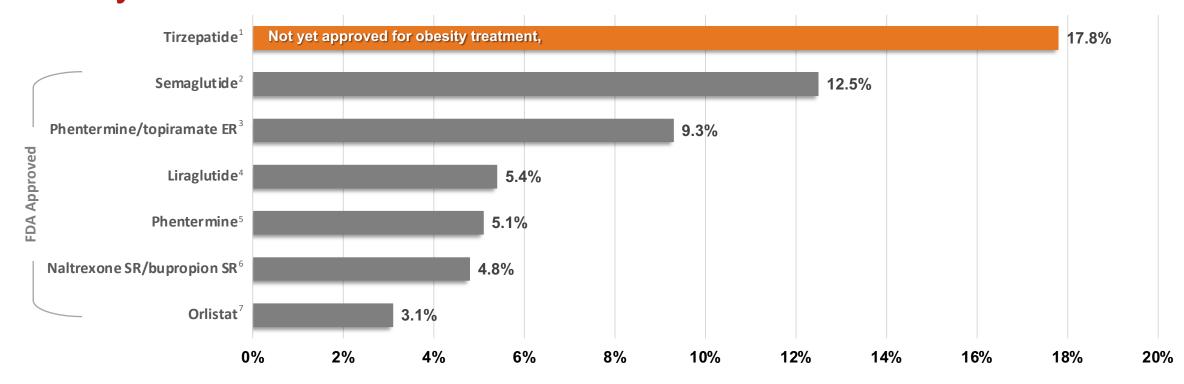
Greater weight loss improves obesity-related complications: Goal is 15% or greater weight loss





^{1.} Garvey WT et al. Endocr Pract 2016;22(Suppl. 3):1–203; 2. Look AHEAD Research Group. Lancet Diabetes Endocrinol 2016;4:913–21; 3. Lean ME et al. Lancet 2018;391:541–51; 4. Benraoune F and Litwin SE. Curr Opin Cardiol 2011;26:555–61.

Placebo-subtracted Efficacy of Current and Pending Antiobesity Medications



ITT Placebo-substracted weight loss

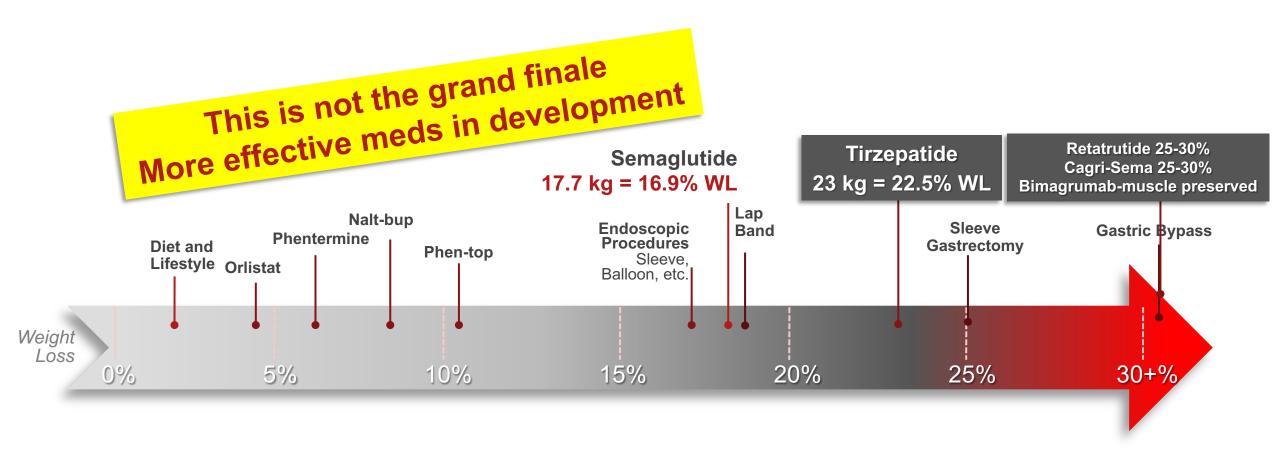
- 1. Jastreboff AM, et al. N Engl J Med. 2022 Jul 21;387(3):205-216.
- Wilding JPH, et al. N Engl J Med. 2021 Mar 18;384(11):989.
- 3. Allison DB, et al. EQUIP Trial. Obesity (Silver Spring). 2012 Feb;20(2):330-42.
- 4. Pi-Sunyer X, et al. N Engl J Med. 2015 Jul 2;373(1):11-22.
- 5. Aronne LJ, et al. Obesity (Silver Spring). 2013 Nov;21(11):2163-71.
- 6. Greenway FL, et al. COR-I Trial. Lancet. 2010 Aug 21;376(9741):595-605.
- 7. Finer N, et al. Int J Obes Relat Metab Disord. 2000 Mar;24(3):306-13.

Add 3-4% for a behavioral intervention



Current Treatment Landscape with Tirzepatide

New drugs and devices can reduce weight and weight-related comorbidities

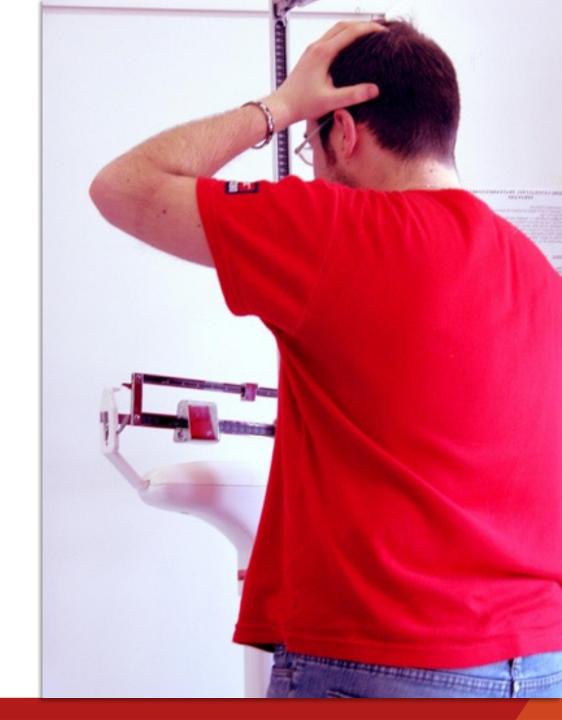




What should you do?

- Maintain a healthy body weight
- If overweight, lose some weight
- If higher and no progress, consider medical treatment
- Maintain or increase muscle mass
- Limit sugar and starch intake
- Limit alcohol to 5-7 drinks per week
- You may want to talk to your doctor about whether metformin is appropriate



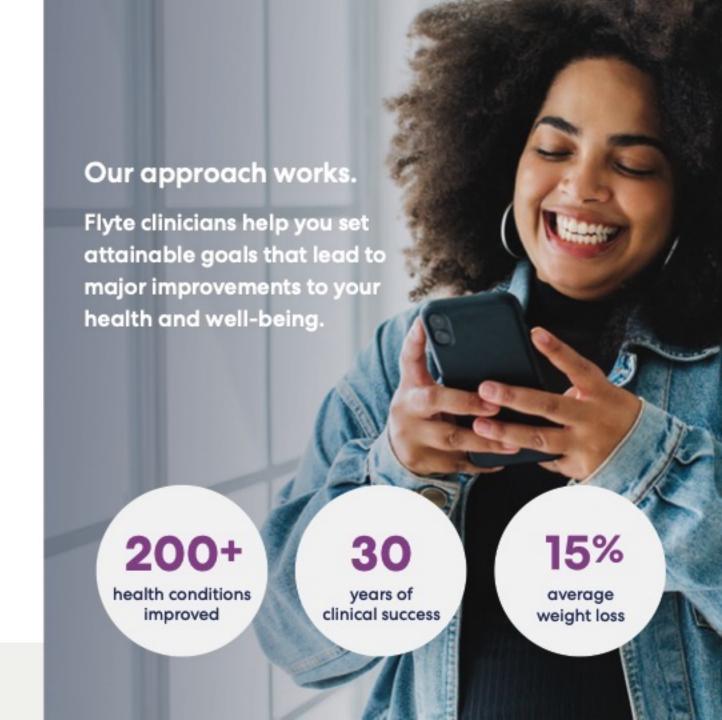




Flyte is a unique combination of medical guidance and behavior-change support that leads to significant and sustainable weight loss.

The Flyte program includes:

- A dedicated team of medical weight-loss experts who understand you
- Tools to track your progress and connect to your team every step of the way
- Medications and eating plans tailored to your unique biology and weight-loss goals





Thank you!