

INCRETIN MIMETIC TREATMENT: Enhancing Whole Body Health

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University of Michigan



Two Steps to Receive CME/CE Credit

STEP 1:

Text in CE Code 83336 to 833-256-8390
by 1:00 PM on March 29th

Scan me to open a text message!



This activates your online evaluation in the CE portal (new users follow prompts after texting to set up account).

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Complete the required online evaluation
by April 13, 2025

In the Cloud CME portal at <https://beaumont.cloud-cme.com> [Sign In > select My CME > select Evaluations & Certificates] – or – via the free CloudCME mobile app (organization code *Beaumont*)

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MCT2D Learning Community Series 2025 Incretin Mimetics Nutritional Considerations and More

CME/CE credit is available

In support of improving patient care, this activity has been planned and implemented by Corewell Health Southeast Michigan and Michigan Collaborative for Type 2 Diabetes. Corewell Health Southeast Michigan is jointly accredited by the Accreditation Council for Continuing Medical Education (ACCME), the Accreditation Council for Pharmacy Education (ACPE), and the American Nurses Credentialing Center (ANCC), to provide continuing education for the healthcare team.


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Dietetic CPEU:

 <p>Commission on Dietetic Registration <small>the credentialing agency for the Academy of Nutrition and Dietetics</small></p>	<p>Completion of this RD/DTR profession-specific or IPCE activity awards CPEUs (One IPCE credit = One CPEU).</p> <p>If the activity is dietetics-related but not targeted to RDs or DTRs, CPEUs may be claimed which are commensurate with participation in contact hours (One 60 minute hour = 1 CPEU).</p> <p>RD's and DTRs are to select activity type 102 in their Activity Log. Sphere and Competency selection is at the learner's discretion.</p>
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NO Disclosures



MCT2D



ABOUT ME

- Endocrinologist & obesity medicine specialist since 2010
- Director, Weight Navigation Program
- Director, Post-Bariatric Endocrinology Clinic
- Participating physician, Weight Management Program
- Associate Program Director, Endocrinology Fellowship



Today's Goals

- Review background of incretin mimetics
- Discuss nutritional considerations
- Review sarcopenia
- Review frailty-prevention strategies
- Review case studies to illustrate points



CASE

29-year-old graduate student is establishing care for management of diabetes mellitus.

- Diagnosed at the age of 21
- Started on metformin but started on premixed insulin soon afterwards due to $A1c > 10\%$
- Had persistently poor control, was told she had T1DM, and was switched to basal bolus insulin after which she established in this clinic
- After starting CGM & titrating doses, control improved to mid 8s (on A1c)



CASE

29-year-old with DM

- Though she was pleased with an A1c of 7, she was concerned about the weight gain from 200 to 230 lbs. over the course of 12 months.





According to the American Diabetes Association Standards of Care, for non-acute, chronic care of an individual with diabetes, which health aspect has treatment priority?

2004
(when I started
residency)

Table 6—Summary of recommendations for adults with diabetes

Glycemic control	
A1C	<7.0%*
Preprandial plasma glucose	90–130 mg/dl (5.0–7.2 mmol/l)
Postprandial plasma glucose†	<180 mg/dl (<10.0 mmol/l)
Blood pressure	<130/80 mmHg
Lipids‡	
LDL	<100 mg/dl (<2.6 mmol/l)
Triglycerides	<150 mg/dl (<1.7 mmol/l)
HDL	>40 mg/dl (>1.1 mmol/l)§

Key concepts in setting glycemic goals:

- Goals should be individualized
- Certain populations (children, pregnant women, and elderly) require special considerations
- Less intensive glycemic goals may be indicated in patients with severe or frequent hypoglycemia
- More stringent glycemic goals (i.e. a normal A1C, <6%) may further reduce complications at the cost of increased risk of hypoglycemia (particularly in those with type 1 diabetes)
- Postprandial glucose may be targeted if A1C goals are not met despite reaching preprandial glucose goals

NOTE: no
discussion of ADM
type or order –
not even
metformin!

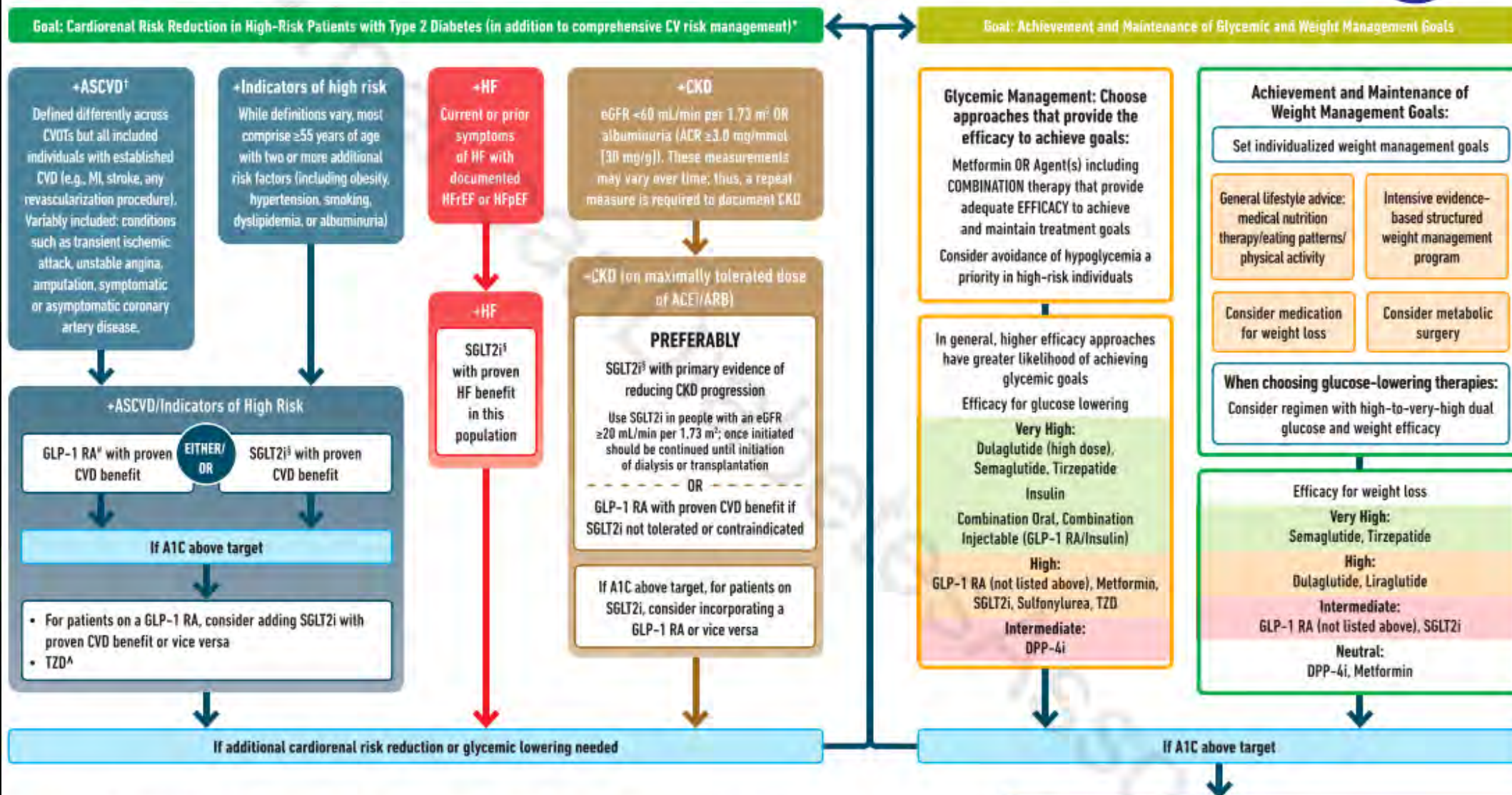
*Referenced to a nondiabetic range of 4.0–6.0% using a DCCT-based assay. †Postprandial glucose measurements should be made 1–2 h after the beginning of the meal, generally peak levels in patients with diabetes. ‡Current NCEP/ATP III guidelines suggest that in patients with triglycerides 200 mg/dl, the “non-HDL cholesterol” (total cholesterol minus HDL) be utilized. The goal is 130 mg/dl (61). §For women, it has been suggested that the HDL goal be increased by 10 mg/dl.

2023

USE OF GLUCOSE-LOWERING MEDICATIONS IN THE MANAGEMENT OF TYPE 2 DIABETES



HEALTHY LIFESTYLE BEHAVIORS; DIABETES SELF-MANAGEMENT EDUCATION AND SUPPORT (DSMES); SOCIAL DETERMINANTS OF HEALTH (SDOH)

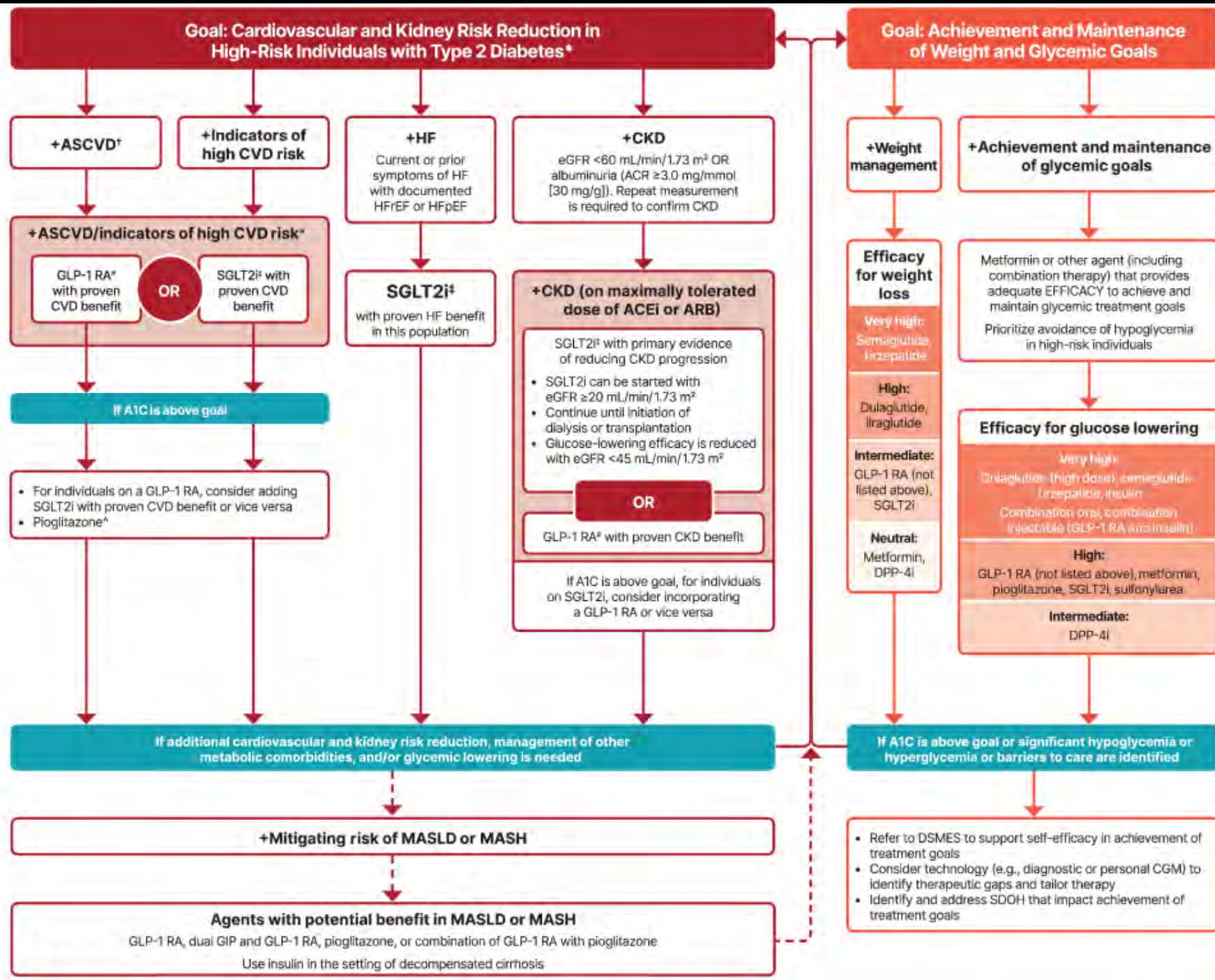


* In people with HF, CKD, established CVD or multiple risk factors for CVD, the decision to use a GLP-1 RA or SGLT2i with proven benefit should be independent of background use of metformin; † A strong recommendation is warranted for people with CVD and a weaker recommendation for those with indicators of high CV risk. Moreover, a higher absolute risk reduction and thus lower numbers needed to treat are seen at higher levels of baseline risk and should be factored into the shared decision-making process. See text for details; ^ Low-dose TZD may be better tolerated and similarly effective; § For SGLT2i, CV/renal outcomes trials demonstrate their efficacy in reducing the risk of composite MACE, CV death, all-cause mortality, MI, HFrEF, and renal outcomes in individuals with T2D with established/high risk of CVD; # For GLP-1 RA, CVOTs demonstrate their efficacy in reducing composite MACE, CV death, all-cause mortality, MI, stroke, and renal endpoints in individuals with T2D with established/high risk of CVD.

Identify barriers to goals:

- Consider DSMES referral to support self-efficacy in achievement of goals
- Consider technology (e.g., diagnostic CGM) to identify therapeutic gaps and tailor therapy
- Identify and address SDOH that impact achievement of goals

2025



CASE

29-year-old with DM

- Though she was pleased with an A1c of 7, she was concerned about the weight gain from 200 to 230 lbs. over the course of 12 months. **She attributed this to insulin and tried to self-reduce resulting in A1c rise back to 8.5%**





CASE

29-year-old with DM

- Was started on pramlintide (Symlin) 60 mcg and titrated to three times daily with meals



Have you heard of pramlintide (Symlin)?

CASE

29-year-old with DM

- Was started on pramlintide (Symlin) 60 mcg and titrated to three times daily with meals
- Noted some hunger reduction and modest weight reduction (230 → 219) but found it burdensome and expensive
- A1c improved from 8.5% to 6.9%



CASE

31-year-old with DM

- She continued with basal bolus insulin + metformin + intermittent use of symlin for another 18 months
- During this time, the pandemic started/persisted, and she had additional weight gain to a peak of 260 lbs. though glycemic control remained at target





CASE

31-year-old with DM

- Decision made to retest c-peptide (previously found to be low). Found to be detectable (though low-normal). Antibody testing for T1DM also negative.
- Decision made to STOP pramlintide and START semaglutide (insurer was covering it without a prior authorization and would be less expensive than pramlintide).



ASIDE from cost/coverage, what are the most common reasons your patients are hesitant to try incretin mimetic/NUSH therapy? (can select multiple)



CASE

IF your patient is hesitant to try a “new” medicine, what do you say?

IF your patient is wondering how these medicines work, what do you say?

IF your patient is skeptical about using these medicines for either diabetes or obesity, what do you say?



(brief)

HISTORY of

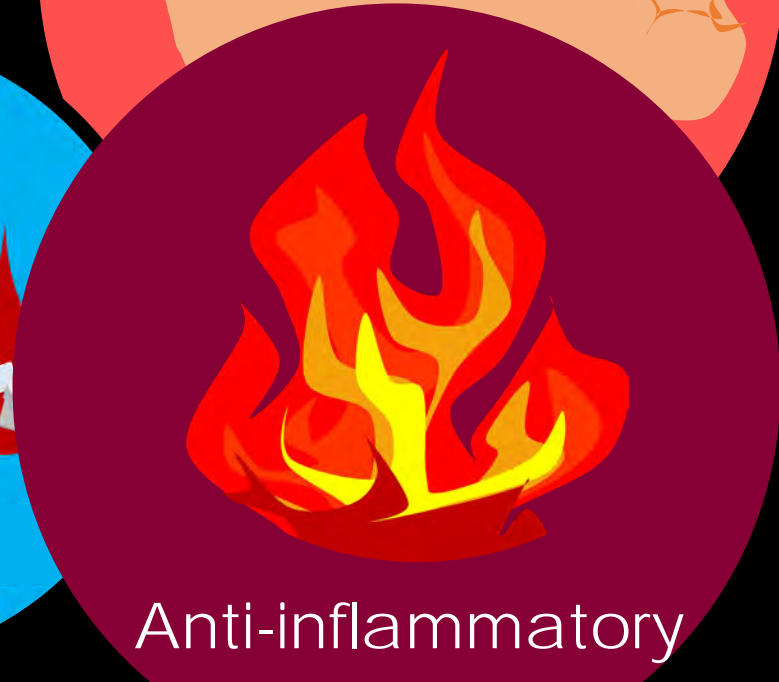
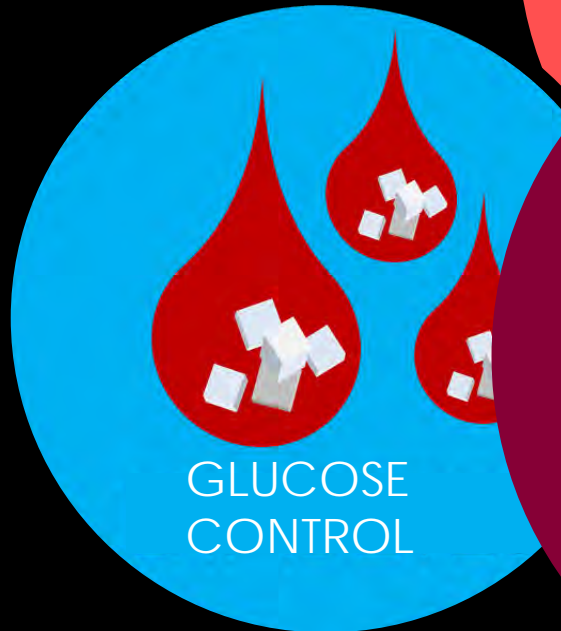
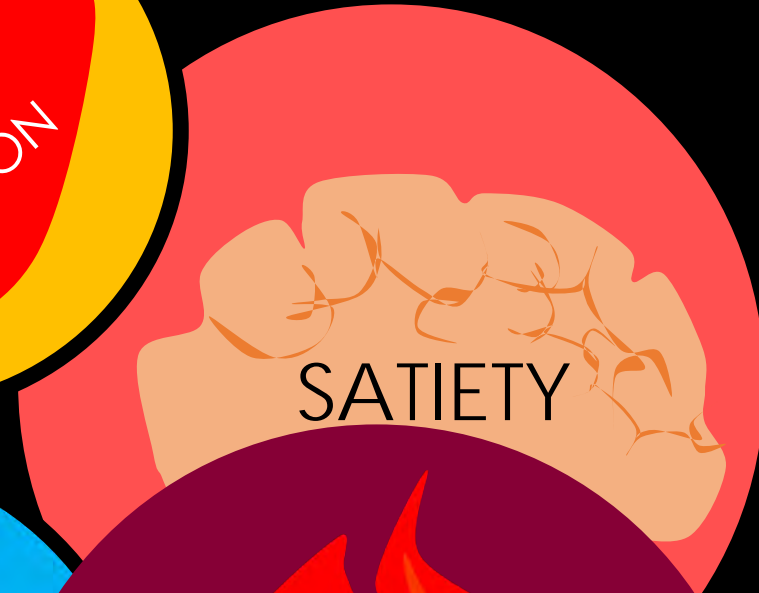
~~GLP-1s~~

Incretin mimetics

a.k.a.

Nutrient-stimulated hormones
(NUSH)s

MAIN DIRECT EFFECTS



“The Incretin Effect” □ first identified in 1987

HORMONES (such as GIP and GLP-1 that...

- respond to oral nutrients
 - carb > fat > protein
- promote satiety via hypothalamus
- decrease gastric emptying
- enhance insulin secretion
- function as “meal termination” signals



Incretin effect is BLUNTED

- In those with diabetes
- In those with obesity
- Especially in those with both
- Therefore: simple “replacement” would be inadequate

INCRETIN EFFECT and weight control: Happy Accident? NO



Lotte Knudsen

Novo Nordisk

Chief Scientific Advisor in Research
and Early Development



Foto: Jeppe Bøje Nielsen / DER SPIEGEL

INCRETIN EFFECT and weight control: Happy Accident? NO



David: Is it correct to say that during the (at least) initial development process, the target market was type 2 diabetics who were not yet insulin-dependent.

Lotte: That's the next level of the conversation that could be really interesting because actually for us, the target market was also obesity from the beginning. Of course, it's impossible to be on top of everything that's being written, but for us it was obesity all the way from the beginning.



INCRETIN EFFECT and weight control: Happy Accident? NO

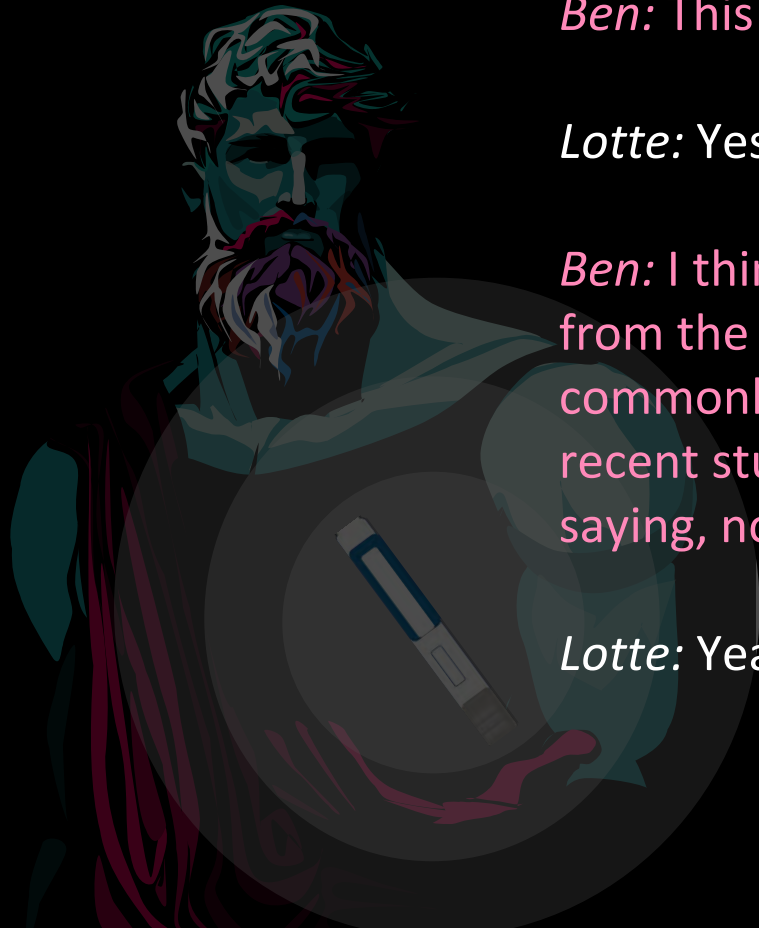


Ben: This was as early as the early mid-90s, this was the thinking?

Lotte: Yes, it was.

Ben: I think this is important to underscore for listeners. This is extremely different from the widely held belief in most media right now that is about Ozempic. What you commonly read is that they were seeking a diabetes drug and oh my god, in this most recent study they stumbled upon this idea that it's amazing for weight loss. You're saying, no, from the very beginning, we thought that this could do both.

Lotte: Yeah, it is a common misunderstanding.





Timeline of Incretin/NUSH therapies

Incretin mimetics / (NuSHs)

STOP TRYING TO MAKE NUSH HAPPEN!

GLP-1 RA

- 2005 Exenatide (2x/d)
- 2010 Liraglutide (1x/d)
- 2012 Exenatide-LA (1x/week)
- 2014 Dulaglutide (1x/week)
- 2014 Albiglutide (1x/week)
- 2016 Lixisenatide (1x/d)
- 2017 Semaglutide (1x/week)
- 2019 Semaglutide (oral, 1x/d)

Amylin mimetic

- 2005 Pramlintide (3x/d)

GLP-1/GIP Dual agonist

- 2022 Tirzepatide (1x/week)



Incretin mimetics / (NuSHs)

GLP-1 RA

- 2005 Exenatide (2x/d)
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- 2012 Exenatide-LA (1x/week)
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- 2016 Lixisenatide (1x/d)
- 2017 semaglutide (1x/week)
- 2019 semaglutide (oral, 1x/d)

Amylin mimetic

- 2005 Pramlintide (3x/d)

GLP-1/GIP Dual agonist

- 2022 Tirzepatide (1x/week)



Lots to digest

2

United States, estimated launch dates of selected weight-loss drug candidates

● Injectable ● Oral

	Manufacturer	Drug	Type
2026	Eli Lilly	orforglipron	●
	Novo Nordisk	cagrisema	●
	Novo Nordisk	semaglutide	●
2027	Boehringer Ingelheim/ Zealand Pharma	servodutide	●
	Eli Lilly	retatrutide	●
2028	Altimune	pemvidutide	●
	Amgen	maritide	●
	Pfizer	danuglipron	●
	Viking Therapeutics	VK2735	●
2029	Eli Lilly	mazdutide	●
	Structure Therapeutics	GSBR-1290	●
	Zealand Pharma	dapiglutide	●
2030 onwards	Novo Nordisk	amycetin	●
	Roche	CT-388	●
	Zealand Pharma	petrelintide	●

Source: Bloomberg Intelligence

Incretin mimetics / (NuSHs) FUTURE directions

- Oral GLP-1 RA
- Small molecular weight GLP-1 RA
- Dual therapy:
 - Amylin-GLP-1 RA
 - GIP R antagonist/ GLP-1 RA
 - GLP-1 RA/ glucagon (GCG)
- Triple therapy:
 - GLP-1 RA/GIP-RA/GCG

Ashwin Kanna Chetty, Ebne Rafi, Natalie J. Bellini, Natalie Buchholz, Diana Isaacs,
A Review of Incretin Therapies Approved and in Late-Stage Development for Overweight and Obesity Management,
Endocrine Practice, Volume 30, Issue 3, 2024, Pages 292-303, ISSN 1530-891X

Campbell JE. Targeting the GIPR for obesity: To agonize or antagonize? Potential mechanisms. Mol Metab. 2021 Apr;46:101139. doi: 10.1016/j.molmet.2020.101139. Epub 2020 Dec 5. PMID: 33290902; PMCID: PMC8085569.



BACK TO CASE

31-year-old with diabetes:

- Ready to start semaglutide.
- Ready to start semaglutide?



Are treatment considerations different/ same when NuSH therapy is used for:

- Diabetes
- Obesity
- CAD
- PAD
- CKD
- OSA

?





When prescribing for diabetes vs obesity vs CAD, do you think differently about how the medicine will affect weight, hunger, cravings? (may select more than one)



Does the indication affect
your nutrition plan?

More attentive?
Less attentive?

Does nutrition get “lost in the
shuffle”?

?



COMPREHENSIVE Nutrition Plan



- Baseline assessment
 - Current practices
 - Dietary/healthy literacy
 - Vitamin/mineral status
- Goal formulation
 - Calories
 - Macronutrients
 - Comorbid management
- Plan
 - RD
 - Commercial
 - Specialized program
 - Self-management



BASELINE Assessment

In the visit:

- The quick-and-dirty
- 24-hr. dietary recall
- **Structured tool** (e.g. The Rapid Eating Assessment for Participants–Shortened (REAP-S) – can be done in < 10-minute)

Pre/post visit:

- Registered Dietitian

BASELINE Assessment

Vitamins/minerals:

- Consider the relative prevalence of deficiencies in the population
- May need pre-intervention screening
- May need a multivitamin +/- other supplements



BASELINE Assessment


Dietary (and health) literacy:

- Will impact treatment recommendations
- If dietary literacy is low, will need more general education, initially
- If high, then general recommendations will be seen as ineffective and a “waste of time”
- Can be discerned, to some degree, at visit



Goal Formulation

Options:

- Simple calorie reduction strategy
- Resting metabolic rate estimation  to guide caloric recommendation
- Macronutrient strategy (e.g. very low carbohydrate)
- Disease specific (diabetes, CHF, renal)





Goal Formulation

For purposes of weight loss, there are numerous, evidenced based strategies:

- DASH
- Diabetes Prevention Program/ Look AHEAD
- Mediterranean
- Low-fat
- Low-carb
- Very-low-carb
- Low-calorie
- Very-low-calorie/ meal replacement
- Intermittent fasting/ time restricted feeding

In addition to patient-specific factors, think about YOUR ecosystem:

What can we do WELL?

Plan/Implementation

Options:

- RD
- Commercial program
- Medical nutrition therapy/
comprehensive program
- Self-managed

*we will move on with the case but will talk about incretin-specific-diet considerations and exercise, later.



Intensive diet (could include partial meal replacement)

Effect of structured lifestyle regimen

- Studies demonstrate effective enhancement of medication use when paired with more intensive dietary intervention

nature medicine

Article

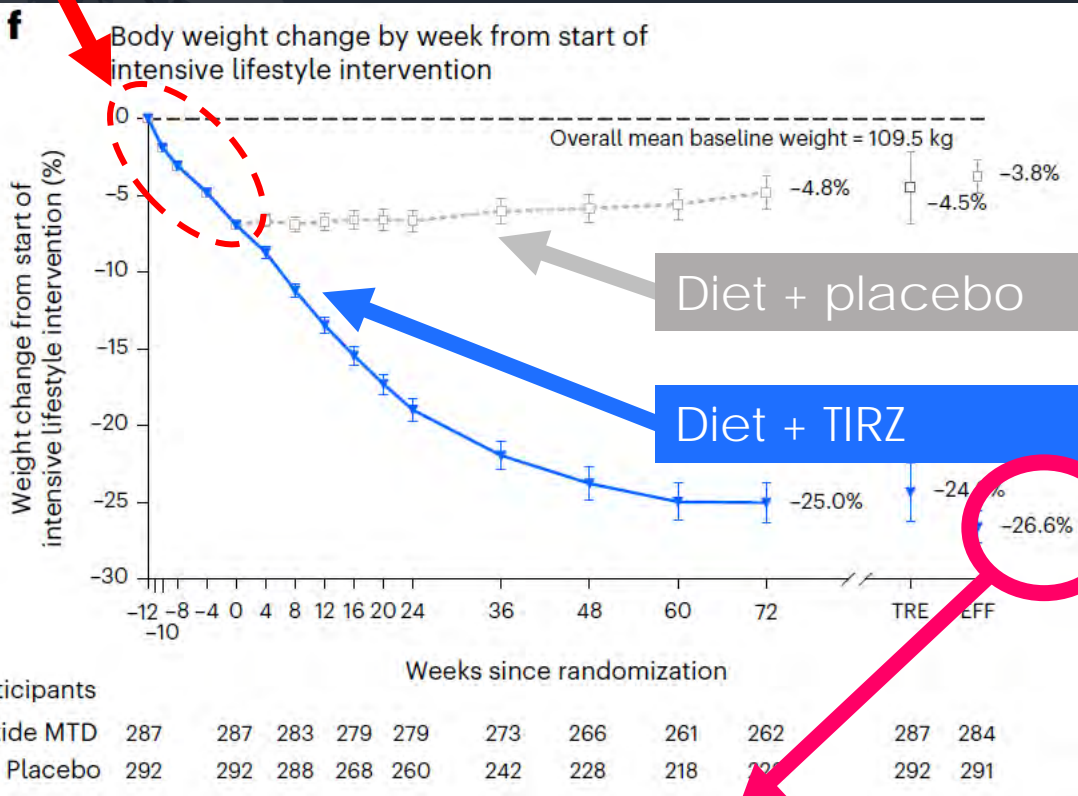
<https://doi.org/10.1038/s41591-023-02597-w>

Tirzepatide after intensive lifestyle intervention in adults with overweight or obesity: the SURMOUNT-3 phase 3 trial

Received: 18 August 2023

Accepted: 15 September 2023

Thomas A. Wadden¹, Ariana M. Chao², Sriram Machineni³, Robert Kushner⁴, Jamy Ard⁵, Gitanjali Srivastava^{6,7}, Bruno Halpern⁸, Shuyu Zhang⁹, Jiaxun Chen⁹, Mathijs C. Bunck⁹, Nadia N. Ahmad⁹ & Tammy Forrester⁹



*Superior results to conventional dietary intervention + TIRZ

CASE

31-33-year-old with DM

- She agreed to start semaglutide –and- agreed to return to see the dietitian.
- She received calorie reduction recommendations and carb moderation advice
- Over the course of two years, went from 260 → 183 lbs. (= 30% weight loss) and A1c ranged between 6-7%. Insulin still required but at lower doses.





CASE

33-year-old with DM

- Happy Ever After?
- LATER, we will revisit this case to discuss some obstacles that arose



NEXT CASE



CASE

62-year-old with DM, MASLD, and OSA

- She would like help with weight control
- Diabetes is managed with metformin monotherapy and control is good
- However, A1c has been rising for the past few years and correlates with an increase in weight
- She started to gain weight in her 30s during pregnancies and had trouble losing weight post-partum
- She is at 240 lbs. with a BMI of 38 kg/m²



CASE

62-year-old

- She has tried commercial weight programs with the last effort a year ago. She was able to lose 15 lbs., but found the diet unsustainable due to hunger/cravings
- She also notes issues with impulse control and emotion-provoked eating
- She has a medical background and believes that her dietary and health literacy is high
- After initial discussion, she was referred to a therapist to discuss disordered eating.
- She also agreed to be seen in the intensive weight management program offered through the institution. This program starts with 12 weeks of total meal replacement followed by a transition to an individualized lifestyle plan.
- However, she changes her mind and returns to discuss obesity modifying medication



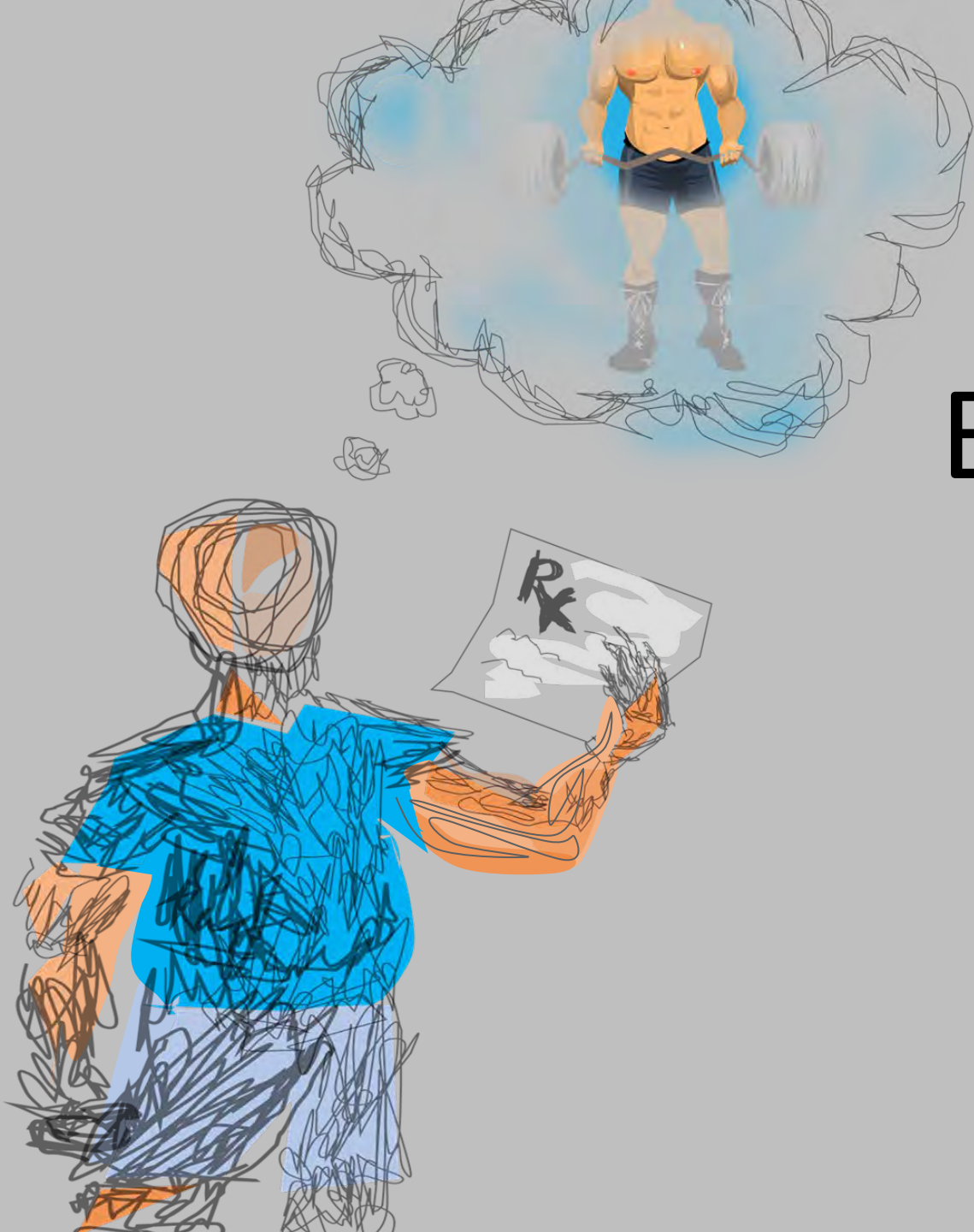
CASE

62-year-old

- Given her history of diabetes, incretin mimetic therapy is being considered
- Baseline bone density study revealed osteopenia with T-scores of -1.2 of L1-L3 and -1.5 of the femoral neck [FRAX score did not warrant prophylactic treatment – even when accounting for diabetes]



For your older patients, are there any particular concerns you have when considering incretin mimetic therapy?



BODY COMPOSITION + INCRETINS

Neeland IJ, Linge J, Birkenfeld AL. Changes in lean body mass with glucagon-like peptide-1-based therapies and mitigation strategies. *Diabetes Obes Metab.* 2024 Sep;26 Suppl 4:16-27. doi: 10.1111/dom.15728. Epub 2024 Jun 27. PMID: 38937282.



The concern for SARCOPENIA

Neeland IJ, Linge J, Birkenfeld AL. Changes in lean body mass with glucagon-like peptide-1-based therapies and mitigation strategies. *Diabetes Obes Metab.* 2024 Sep;26 Suppl 4:16-27. doi: 10.1111/dom.15728. Epub 2024 Jun 27. PMID: 38937282.



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SARCOPENIA:

- loss of muscle mass, strength, and function due to weight loss
- Manifestations: weakness, fatigue, balance disturbance, trouble with walking/standing



WEIGHT LOSS-INDUCED CHANGES

- Reduction in adipose tissue mass
- Reduction in fat-free (a.k.a. lean) mass
 - Muscle
 - Organs
 - Bone
 - Fluids
 - Water

WEIGHT LOSS-INDUCED CHANGES

- Reduction in adipose tissue mass
- Reduction in fat-free (a.k.a. lean) mass
 - Muscle
 - Organs
 - Bone
 - Fluids
 - Water

75%?

Rule of thumb

25%?



55%?

- Semaglutide (STEP-1)
- Semaglutide (SUSTAIN-8)

45%?

INCRETIN-MEDIATED WEIGHT LOSS-INDUCED CHANGES: Different or not?

- Reduction in adipose tissue mass
- Reduction in fat-free (a.k.a. lean) mass



75%?

INCRETIN-MEDIATED WEIGHT LOSS-INDUCED CHANGES: Different or not?

- Reduction in adipose tissue mass
 - Reduction in fat-free (a.k.a. lean) mass
- Tirzepatide (SURMOUNT-1)

25%?



85%?

- Liraglutide + lifestyle

15%?

INCRETIN-MEDIATED WEIGHT LOSS-INDUCED CHANGES: Different or not?

- Reduction in adipose tissue mass
- Reduction in fat-free (a.k.a. lean) mass



100%

?

- Liraglutide + exercise

+15%

INCRETIN-MEDIATED WEIGHT LOSS-INDUCED CHANGES: Different or not?

- Reduction in adipose tissue mass
- **INCREASE** in fat-free (a.k.a. lean) mass



INCRETIN-MEDIATED WEIGHT LOSS-INDUCED CHANGES: Knowledge gaps

- Lack of routine, standardized clinical assessment/ tools
- Heterogeneity of studies
- Small % of participants get body composition
- Lean body mass is not synonymous with muscle mass
- Muscle mass is not the same as muscle function



INCRETIN-MEDIATED WEIGHT LOSS-INDUCED CHANGES:

- Based on the current level of evidence, we cannot definitively conclude that incretin-mediated weight loss results in a **DISPROPORTIONATELY** high degree of lean body mass loss and/or muscle loss



INCRETIN-MEDIATED WEIGHT LOSS-INDUCED CHANGES:

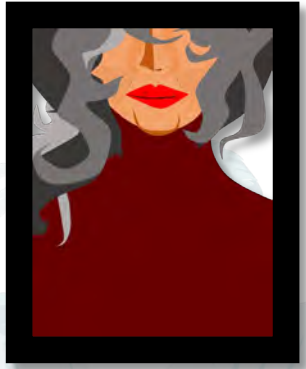
- HOWEVER, we know that some of our patients are at disproportionately high risk of sarcopenia due to:
 - Age
 - Baseline functional status
 - Comorbid conditions
- Therefore, we need to be diligent about **PREVENTING FRAILITY**

What about our patient?





What would you recommend to avoid sarcopenia and frailty?

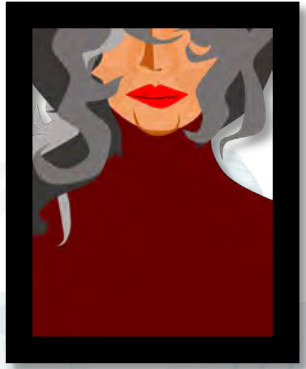


DIET:

Nutrient Dense Foods!

Protein prioritization!

- Common sense, right?
- What does the evidence say?
 - No consensus in the literature



DIET: Protein prioritization

- Challenges:
 - Wide range of recommended amounts
 - Do you use a weight-based calculation? Which weight?
 - How do you select the amount from the range of from 1 – 2 grams of protein/kg/day?
 - High recommended targets can be hard to achieve
 - Easy to recommend, hard for some patients to implement ? do we know what are the practical translations for our recommendations?

1200 - 1500 cal/day (120 gm protein)

Day 1:

Breakfast (400 kcal, 35g protein):

- 3 scrambled eggs (with cooking spray)
- 1 oz shredded cheese
- 1 cup spinach, sautéed
- 2 slices of whole-grain toast

Lunch (350 kcal, 30g protein):

- Grilled chicken breast (4 oz)
- 1 cup quinoa
- 1 cup steamed broccoli
- 1 tbsp olive oil (for cooking or drizzling)

Dinner (450 kcal, 40g protein):

- 6 oz salmon, baked
- 1 cup roasted sweet potatoes
- 1 cup steamed asparagus
- Lemon and herbs for seasoning

Snack (150 kcal, 15g protein):

- Greek yogurt (plain, 3/4 cup)
- 1 tbsp chia seeds

Day 2:

Breakfast (350 kcal, 30g protein):

- Protein oatmeal: 1/2 cup oats + 1 scoop protein powder
- 1 tbsp almond butter
- 1/2 banana, sliced

Lunch (400 kcal, 35g protein):

- Turkey and avocado wrap:
 - 4 oz sliced turkey breast
 - 1 whole wheat tortilla
 - Lettuce, tomato, mustard
- 1/4 avocado

Dinner (450 kcal, 40g protein):

- Grilled shrimp (6 oz)
- 1 cup brown rice
- 1 cup sautéed bell peppers and onions
- Olive oil for cooking (1 tsp)

Snack (150 kcal, 15g protein):

- Low-fat cottage cheese (3/4 cup)
- Sliced cucumber and cherry tomatoes

Day 6 (Vegetarian):

Breakfast (350 kcal, 30g protein):

- Greek yogurt bowl:
 - 1 cup plain Greek yogurt
 - 1 scoop vanilla protein powder
 - 1/2 cup mixed berries
 - 1 tbsp chia seeds

Lunch (400 kcal, 35g protein):

- Lentil and quinoa salad:
 - 1 cup cooked quinoa
 - 1/2 cup cooked lentils
 - Cherry tomatoes, cucumber, bell peppers
 - 2 tbsp feta cheese
 - Lemon-tahini dressing (1 tbsp tahini, lemon juice)

Dinner (450 kcal, 40g protein):

- Paneer stir-fry:
 - 4 oz paneer cheese
 - 1 cup mixed bell peppers, onions, spinach
 - 1/2 cup brown rice
 - Spices: cumin, turmeric, paprika
 - 1 tbsp olive oil

Snack (150 kcal, 15g protein):

- Cottage cheese (3/4 cup)
- Sliced peaches or pineapple

Day 7 (Vegan):

Breakfast (350 kcal, 30g protein):

- Protein smoothie bowl:
 - 1 scoop vegan protein powder
 - 1 cup almond milk
 - 1/2 banana
 - 1 tbsp almond butter
 - Spinach and ice

Lunch (400 kcal, 35g protein):

- Tofu and edamame stir-fry:
 - 4 oz tofu, sautéed
 - 1/2 cup edamame (shelled)
 - 1 cup broccoli and carrots
 - 1 tbsp soy sauce and ginger
 - 1/2 cup brown rice

Dinner (450 kcal, 40g protein):

- Vegan chili:
 - 1 cup black beans
 - 1 cup kidney beans
 - Diced tomatoes, bell peppers, onions
 - Chili powder, cumin, garlic
 - 1/2 avocado for garnish

Snack (150 kcal, 15g protein):

- Roasted chickpeas (1/2 cup)
- Nutritional yeast for seasoning

1200 cal/day (150 gm protein)

🍳 Day 1 (Omnivore - Latino-Inspired)

Breakfast (300 kcal, 50g protein):

- Huevos revueltos con claras y aguacate (1 cup egg whites scrambled with tomatoes, onions, and cilantro)
- 1 small corn tortilla
- ½ small avocado

Lunch (400 kcal, 50g protein):

- Pollo a la plancha (6 oz grilled chicken breast) with lime and garlic
- Arroz con cilantro y limón (½ cup brown rice with cilantro & lime)
- Steamed green beans

Dinner (400 kcal, 50g protein):

- Tilapia Veracruzana (5 oz grilled tilapia with tomatoes, olives, and capers)
- Sautéed bell peppers and zucchini
- ½ cup black beans

Snack (100 kcal, 10g protein):

- 1 scoop whey protein mixed with water

Total: 1200 kcal, 150g protein

🍷 Day 2 (Omnivore - Middle Eastern)

Breakfast (300 kcal, 50g protein):

- Labneh with Za'atar (½ cup low-fat labneh, sprinkled with za'atar)
- 1 boiled egg
- ½ whole wheat pita

Lunch (400 kcal, 50g protein):

- Lean beef kofta (6 oz, seasoned with coriander, cumin, and garlic)
- Grilled zucchini & eggplant
- ½ cup bulgur wheat

Dinner (400 kcal, 50g protein):

- Lemon-garlic shrimp (6 oz, grilled)
- Steamed spinach & sumac
- ½ cup brown rice

Snack (100 kcal, 10g protein):

- Roasted chickpeas (½ cup, spiced with cumin & paprika)

Total: 1200 kcal, 150g protein

🌱 Day 3 (Vegetarian, Indian-Asian Fusion)

Breakfast (300 kcal, 50g protein):

- Besan (chickpea flour) chilla (½ cup besan, water, cumin, turmeric, and chopped vegetables)
- 1 tbsp coconut chutney
- Masala chai (made with almond milk, no sugar)

Lunch (400 kcal, 50g protein):

- Paneer tikka (4 oz grilled paneer, marinated with yogurt, turmeric, and spices)
- Sautéed cabbage & carrots
- ½ cup millet or brown rice

Dinner (400 kcal, 50g protein):

- Soy chunk & tofu stir-fry (½ cup soy chunks, 4 oz tofu, stir-fried with garlic, chili, soy sauce)
- Steamed green beans and mushrooms
- ½ cup red rice

Snack (100 kcal, 10g protein):

- Greek yogurt with cinnamon

Total: 1200 kcal, 150g protein



DIET:

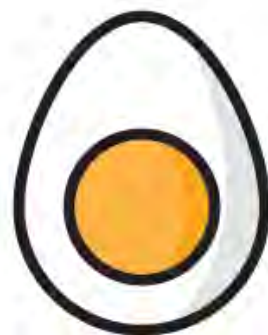
SIMPLIFY!

Maintaining Your Muscle Health with Protein and Hydration



EAT ENOUGH PROTEIN

Protein helps repair and maintain muscle, particularly during weight loss. Eating a greater amount of protein has been shown to help reduce muscle loss while losing weight. Depending on your weight and height, **75-100 gram** of protein is a good place to start for most people.



Protein



Water



Recommendations:



- **Aim for 25-30 grams of protein at each meal.**
- Build your meals and snacks around a protein source such as:
 - Animal proteins (fish, chicken, turkey, beef, pork, venison, and lamb)
 - Plant proteins (beans and lentils, tofu and tempeh)
 - Eggs and dairy (Greek yogurt, high-protein yogurt, cottage cheese, and kefir)



- Try pre-made protein shakes, add protein powders to smoothies, or consider a protein bar. Look for varieties without added sugars.
- Pair your proteins with complex carbs (whole grains, fruits, sweet potatoes), non-starchy vegetables (broccoli, beans, leafy greens), and healthy fats (nuts and seeds, olives, avocado) to make a balanced meal.

These recommendations are for people with normal kidney function. If you have stage 4 or 5 kidney disease, talk with your care team about the right amount of protein for you.

HOW MUCH IS 30 GRAMS OF PROTEIN?

Generally, it takes up $\frac{1}{4}$ of your plate. Here are some examples:



Chicken
1 small breast
1.5 large thighs
2 medium drumsticks



Beef
5 oz steak
 $\frac{1}{2}$ cup or 4 oz cooked
ground beef



Fish
5 oz fillet
canned fish
(1½, 5 oz cans)
Shrimp: 4 oz



Egg
4 whole
1 cup egg whites



**Greek yogurt (plain) or
cottage cheese**
1.3 cups or 10 oz (brands
may vary slightly)



Tofu (firm)
6 oz or 1¼ cups



Cooked Lentils
2 cups or 12 oz



Cooked Beans
(black, pinto, kidney)
2 cups or 12 oz



CASE

62-year-old

- You agree to start an incretin mimetic for weight control with the understand that she will follow with you and a dietitian
- Three months post-treatment initiation, she is doing very well with her diet and tracking/logging
- Weight: 240 → 210 (12.5% loss)
- She still notes some emotion-provoked eating but has not yet established with a therapist. She also does not really notice many changes in the mirror despite the comments of others.
- She is tolerating the medication well. She is on the 3rd dose level. She would like to go up on the medication since she still experiences hunger.
- She is not exercising at all.



Would you increase the dose? Any concerns?



PHYSICAL ACTIVITY
+
EXERCISE



EXERCISE

PHYSICAL
ACTIVITY



EXERCISE: Facts & Myths

- NOT an effective weight LOSS tool, unfortunately.
- Most commonly-used measures of energy. expenditure (ie: fitness trackers; exercise equipment) greatly OVERESTIMATE the number of calories expended during physical activity.
- Exercise IS helpful for **maintaining** a reduced weight state.
- Exercise IS helpful for **preserving lean body mass** during weight loss.
- Exercise SHOULD be performed for **cardiopulmonary health and longevity** - and should be pursued irrespective of weight.
- Therefore, even if weight is not being lost, exercise should be continued for health purposes.

- Lee, L. Djousse, H.D. Sesso, L. Wang, J.E. Buring, "Physical Activity and Weight Gain Prevention," *Journal of the American Medical Association*, vol. 303, no. 12, pp. 1173-1179, 2010
- R.D. Grave, et al. "Cognitive-Behavioral Strategies to Increase the Adherence to Exercise in the Management of Obesity," *Journal of Obesity*, vol. 2011, pp. 1-11.
- D.L. Ballor and E. T. Poehlman, "Exercise-training enhances fat-free mass preservation during diet-induced weight loss: a meta-analytical finding," *International Journal of Obesity*, vol. 18, no. 1, pp.35-40, 1994.
- J.S. Garrow and C.D. Summerbell, "Meta-analysis: effect of exercise, with or without dieting, on the body composition of overweight subjects," *European Journal of Clinical Nutrition*, vol. 49, no. 1, pp. 1-10, 1995.

EXERCISE: Tips

- Individualize initial recommendations based on:
 - Baseline functional status
 - Comorbid conditions; weight-related conditions
 - Life situation
 - Degree of mechanical stress of the excess weight
- Exercise physiology consultation may be prudent for those with significant comorbid challenges
- Even if there are severe limitations, patients can still be encouraged to carve out time in the day to “reserve” space for exercise
- Activities to enhance core strength and balance should be prioritized in those at risk of frailty and those with metabolic bone disease

Maintaining Your Muscle Health with Strength Training



INCORPORATE STRENGTH TRAINING

Strength training (sometimes called resistance training) is exercise using weights, resistance bands, or even your bodyweight to make your muscles work harder. It is the most effective strategy to preserve muscle and boost your metabolism while losing weight.

Recommendations:

- If new to strength training, check with your care team before starting an exercise routine to make sure that it is safe for you.
- Work up to three or four sessions per week. Start with simple bodyweight exercises. Then build on this over time by adding resistance bands or free/hand weights.
- Focus on movements that work several muscle groups and joints.
- Add “movement breaks” (brief 2-5 minute periods of movement) throughout the day to break up sedentary behavior.

Scan below for video examples of exercises for each muscle group!



Lower Body

michmed.org/XWRnR



Upper Body

michmed.org/NYxRw



Back & Core

michmed.org/bQKMm



DON'T FORGET TO STRETCH AND BUILD YOUR BALANCE

Below are basic stretches to complete after strength training exercises.

Calf stretch



Hamstring stretch



Quadriceps stretch



Hip flexor stretch



Iliotibial band stretch



Knee-to-chest stretch



Shoulder stretch



Neck stretch





Future?



Pharmacotherapies are in the development pipeline that act to preserve muscle (e.g., Activin receptor inhibitors)

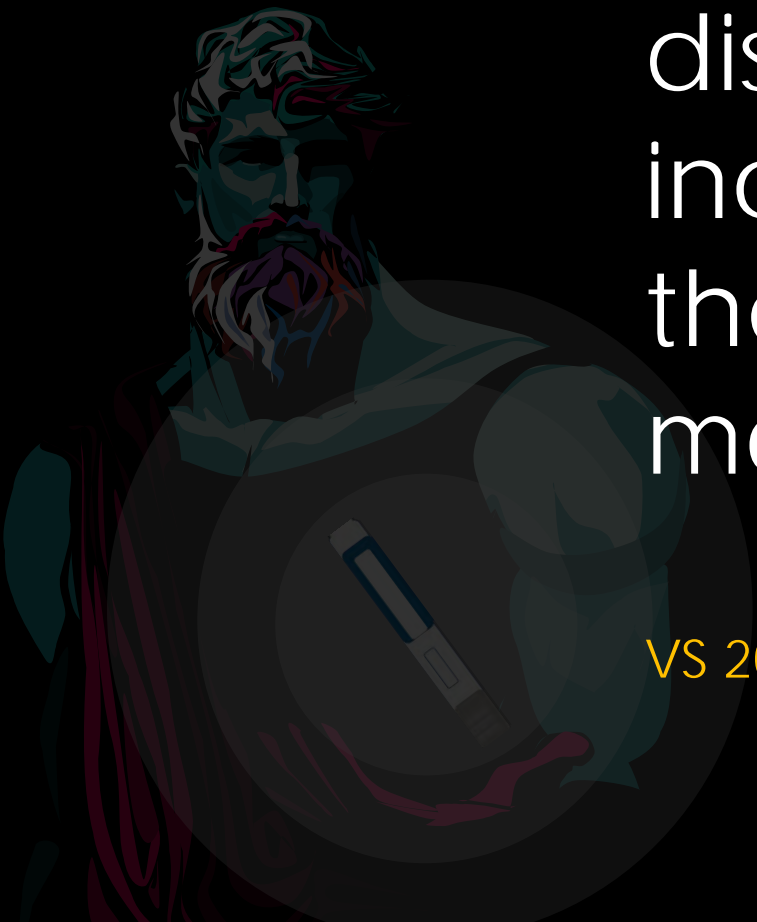


NUSHs/INCRETINS:

Do they make
everything else
obsolete?

...no...



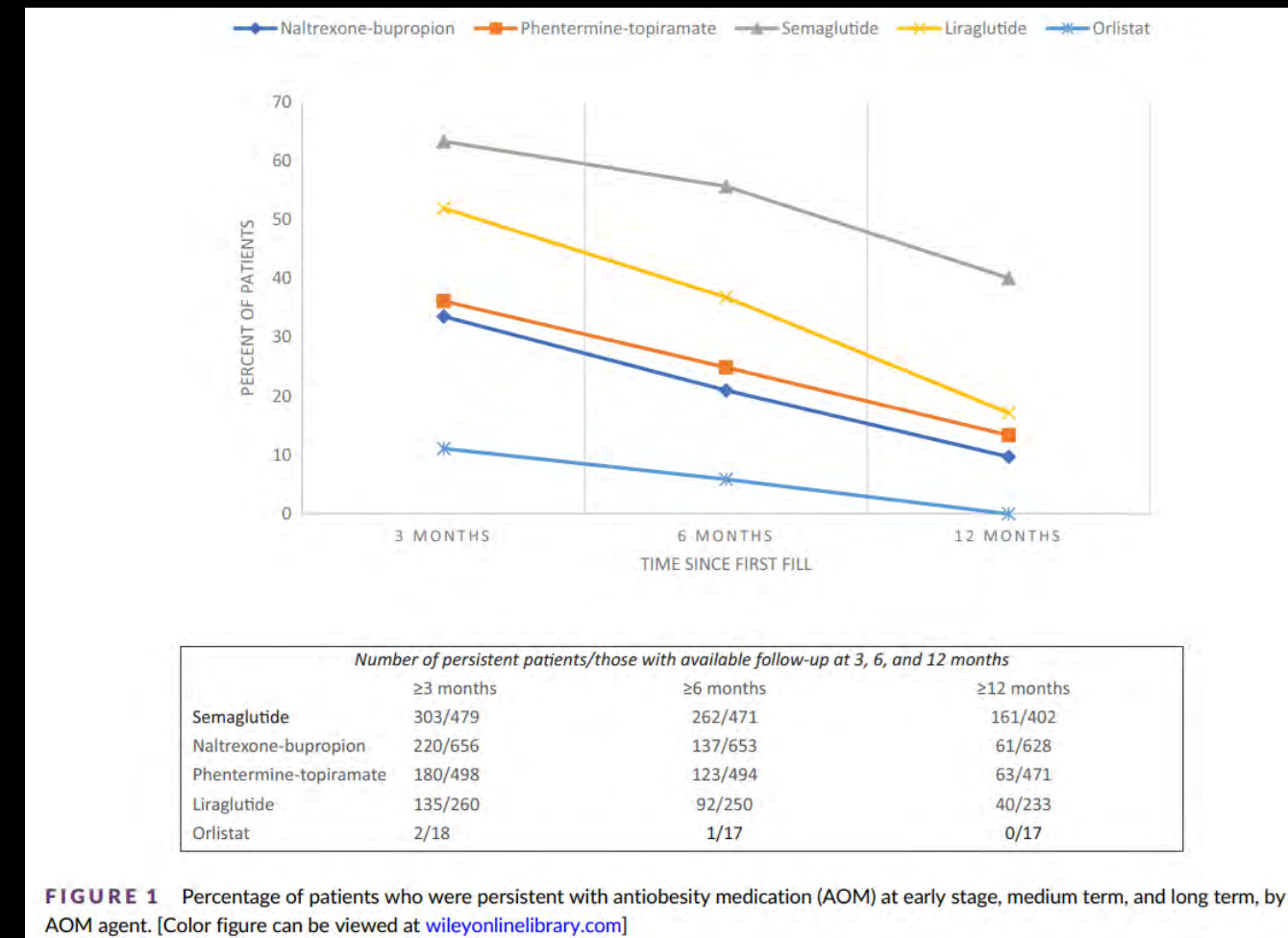


70% “real-world”
discontinuation rate of
incretin mimetic
therapy by 24
months?!?

VS 20-30% clinical trials rates

NOTE: this is true for any weight control medicine

Gasoyan H, Pfoh ER, Schulte R, Le P, Rothberg MB. Early- and later-stage persistence with antiobesity medications: A retrospective cohort study. *Obesity* (Silver Spring). 2023 Dec 6. doi: 10.1002/oby.23952. Epub ahead of print. PMID: 38053443.



WHY?

70% “real-world”
discontinuation rate of
incretin mimetics by 24
months?!?

VS 20-30% clinical trials rates



- Cost
- Coverage
- Tolerance/ side effects
- Shortages
- Lack of true lifestyle changes
- Misaligned expectations
- Lack of education
- Lack of support



CASE

33-year-old with DM

- REVISITED
- Despite improvements in weight and glycemic control, she had to discontinue semaglutide
- The GI side effects (particularly belching) were unmanageable and interfering with her professional singing career





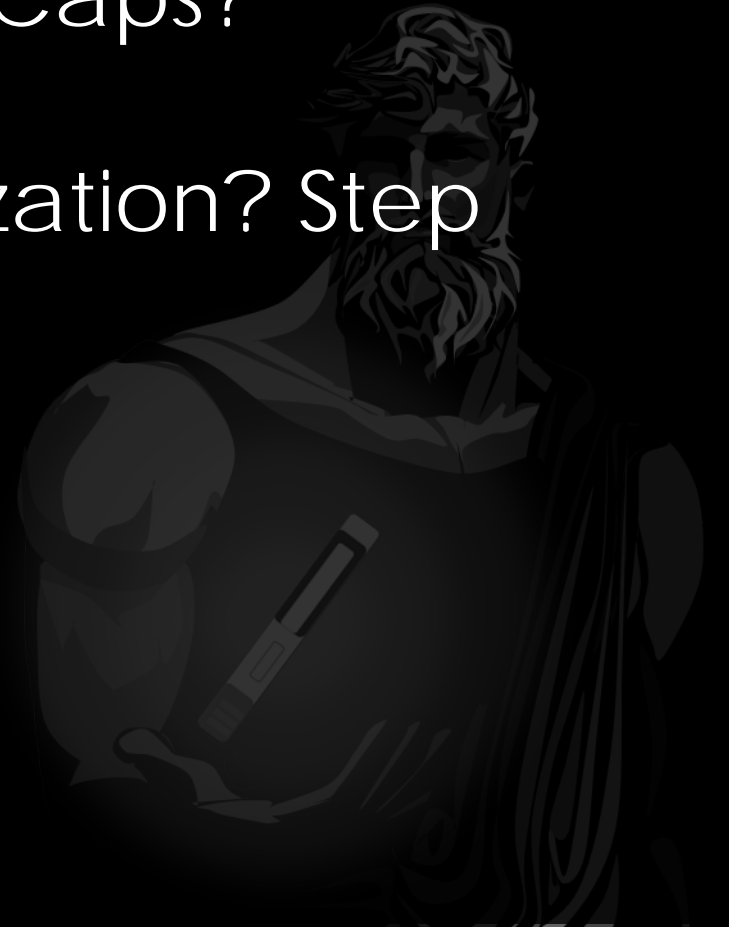
CASE

33-year-old with DM

- REVISITED – part deux
- Recently, we were able to switch to Tirzepatide and she has had return of weight and glycemic control without the GI side effects

COSTS

- Out-of-pocket:
 - INCRETINS: \$900 - \$1300 per month (!)
- Coverage? Caps?
- Prior authorization? Step Therapy?
- Deductible?
- Donut hole?





DIETARY RECALL:

0530 - woke

0700 - coffee (creamer) +
clementines

1200 - 8 chocolate chip
cookies + 2% milk

1930 - flank steak + cheesy
rice + veg

2000 - blueberry pie slice

2130 - bed

CASE

62-year-old

- After being recommended to continue on the same dose, establish with a therapist, and start exercise, the patient did not return to the clinic...
- ...for two years
- During the intervening time, she had received refills through her PCP
- She was now at 155 lbs.
- She returns since she wants to switch therapy since hunger levels are perceived to be too high and she wants to lose another 10 lbs.

MENTAL HEALTH

- Screening should occur for un/der treated mental health issues
 - Uncontrolled mental health issues significantly interfere with weight & health efforts
- Screening should occur for eating disorders (including binge eating)
- May need to postpone weight control efforts unless/until issue(s) are being managed –and- you get permission from treatment team
- Addressing mental health and disordered eating patterns are NECESSARY but NOT SUFFICIENT for weight loss/ maintenance

Rothberg A, McEwen L, Kraftson A, Ajluni N, Fowler C, Miller N, Zuraes K, Herman W. Factors associated with participant retention in a clinical, intensive, behavioral weight management program. BMC Obesity (2015) 2:11

Wing RR, Phelan S. Long-term weight loss maintenance. Am J Clin Nutr. 2005;82:222S–5

Fabricatore AN, Wadden TA, Moore RH, Butryn ML, Hymnsfield SB, Nguyen AM. Predictors of attrition and weight loss success: results from a randomized controlled trial. Behav Res Ther. 2009;47:685–91.

Teixeira PJ, Goings SB, Sardinha LB, Lohman TG. A review of psychosocial pre-treatment predictors of weight control. Obes Rev. 2005;6:43–65

Vocks S, Tuschen-Caffier B, Pietrowsky R, Rustenbach SJ, Kersting A, Herpertz S. Meta-analysis of the effectiveness of psychological and pharmacological treatments for binge eating disorder. Int J Eat Disord. 2010;43(3):205–217

Castellnuovo et al. Cognitive behavioral therapy to aid weight loss in obese patients: current perspectives. Psychology Research and Behavioral Management. 2017(10): 165-173

MENTAL HEALTH

- Help may be needed before/during/after weight change efforts
- Needs can evolve throughout the process
- Do not equate lower weight with increased happiness
- Recognize that weight change impacts the patient and family/friends ? this can lead to positive and negative results
 - Watch for domestic abuse
- Watch for transference of coping behaviors and/or addiction
- Support groups can be helpful
 - Needs facilitation & behavior monitoring
 - Does not replace 1:1 therapy

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MENTAL HEALTH

- Realize there is a poverty of resources
- Advocacy is needed on this front!
- Try and compile a list of local resources
- Some patients may not need mental health support but rather behavioral change support
 - Could involve cognitive behavioral therapy
 - Could utilize tools that are based on CBT principles (e.g., NOOM)

Rothberg A, McEwen L, Kraftson A, Ajluni N, Fowler C, Miller N, Zuraes K, Herman W. Factors associated with participant retention in a clinical, intensive, behavioral weight management program. BMC Obesity (2015) 2:11

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SUMMARY

- Incretin mimetics have a valuable role in the treatment of patients
- Understanding their strengths and weaknesses is vital
- Health is defined by more than just an A1c or the weight on the scale
- Whole health requires comprehensive care including targeted, evidenced-based nutrition and activity recommendations
- We still have much to learn and improve



THANK YOU FOR
LISTENING!

