

Semaglutide Patient Information

Introduction

Managing weight is a crucial aspect of overall health. Being overweight or obese is not just about appearance; it's a medical condition that significantly increases the risk of chronic diseases. This information sheet provides an overview of three medications—Tirzepatide, Semaglutide, and Retatrutide—designed to aid in weight loss and improve related health conditions.

Health Risks of Overweight and Obesity

Understanding the health risks associated with excess weight underscores the importance of weight management.

Cardiovascular Diseases

- Individuals with obesity have a **32% higher risk** of developing coronary artery disease compared to those with normal weight.
- Approximately 75% of hypertension cases are related to obesity.

Type 2 Diabetes

- Around 90% of individuals with type 2 diabetes are overweight or obese.
- Each 1-point increase in Body Mass Index (BMI) is associated with a 7% increase in the risk of developing diabetes.

Cancer Risk:

- Obesity is linked to an increased risk of at least 13 types of cancer, including breast (post-menopause), colorectal, endometrial, kidney, liver, and pancreatic cancers.
- In the United States, about 40% of all cancers diagnosed are now associated with overweight and obesity.

Sleep Apnea

- An estimated **70%** of individuals with obstructive sleep apnea are obese.
- Weight gain of 10% can increase the risk of developing sleep apnea by six times.

Osteoarthritis

• For every **5 point** increase in BMI, the risk of osteoarthritis increases by **35%**.

Non-Alcoholic Fatty Liver Disease (NAFLD)

• Among obese individuals, the prevalence of NAFLD can be as high as 90%.

Reproductive Health

- Women: Obesity can cause menstrual irregularities and infertility. Approximately 30% of cases of
 infertility are related to weight issues.
- Men: Obesity is associated with lower testosterone levels, affecting fertility and sexual function.

Mental Health

• Individuals with obesity have a **55% increased risk** of developing depression over time. There is a bidirectional relationship; those with depression have a **58% increased risk** of becoming obese.

Mortality and Life Expectancy

• Each **5-point** increase in BMI above 25 is associated with a **31% higher risk** of premature death.

Weight Loss Medications

Semaglutide, tirzepatide, and retatrutide belong to a class of diabetes and weight loss medications commonly referred to as GLP-1 (Glucagon-like Peptide) agonists. All of these medications affect GLP-1. However, tirzepatide also affects GIP (Glucose-dependent Insulinotropic Polypeptide), and retatrutide affects GLP-1, GIP, and Glucagon.

Receptors

- GLP-1 (Glucagon-like peptide-1): It enhances the release of insulin, which helps lower blood sugar levels, and slows down gastric emptying, which contributes to a feeling of fullness.
- GIP (Glucose-dependent insulinotropic polypeptide): This receptor helps enhance insulin secretion in response to meals, further supporting glucose regulation. It also contributes to metabolic processes that promote weight loss.
- Glucagon receptors: These are involved in regulating energy expenditure and promoting lipolysis (the breakdown of fats). Activation of glucagon receptors can increase metabolic rate, which may aid in weight reduction.

Remember that overweight and obesity are chronic medical conditions and frequently require chronic medication use. Contrary to popular belief, it's NOT just about eating less and exercising more. Many times, your body causes you to store fat easier, to not burn fat for energy, mistakenly tells you that you are hungry when you're not, and fights your attempts at losing weight. It is not "cheating" to use medication to control your weight any more than it is "cheating" to use blood pressure medication to control your blood pressure. Yes, your weight may increase if you stop the medication, the same way your blood pressure increases if you stop your blood pressure medication.

You absolutely have to have good eating habits and monitor calorie intake, make sure you are being active, and get enough sleep. It is also imperative that you consume an adequate amount of protein. You not only require protein to maintain your muscle mass, but all of your vital organs require a significant amount of protein. Try to consume about 1 gram of protein for every pound of body weight. For this, you use your GOAL body weight, not your current body weight. For example, if you currently weigh 200 lbs, but your goal/healthy weight is 140; you will aim for a daily intake of 140 grams. If you have any kidney dysfunction, please consult your nephrologist or primary care provider for daily protein recommendations. You have to track your protein intake. Most people significantly overestimate how much protein they consume. All of these things will increase your success with weight loss and help you keep the weight off!

I recommend taking these medications for at least 6 months. Your body wants to stay at the weight it is. You have a "set point" for a weight that your body will try to maintain. It takes a long time to lower that set point. If you want to keep the weight you have lost off, you have a better chance if you take the medication for 6 months or longer. You also have to make lifestyle and diet changes while you are taking the medication.

Semaglutide (Brand Names Ozempic® and Wegovy®)

What is Semaglutide?

Semaglutide (Pronounced Sem-uh-glue-tide) is a once-weekly injectable medication used primarily for the treatment of type 2 diabetes and chronic weight management. It belongs to a class of drugs known as glucagon-like peptide-1 (GLP-1) receptor agonists. By mimicking the action of the natural hormone GLP-1, semaglutide helps regulate blood sugar levels and reduces appetite.

How Does Semaglutide Work?

- **Stimulates Insulin Secretion:** Semaglutide enhances the pancreas's ability to secrete insulin in response to elevated blood glucose levels, helping to lower blood sugar.
- **Suppresses Glucagon Release:** It reduces the secretion of glucagon, a hormone that raises blood sugar levels, thereby contributing to better glycemic control.
- Slows Gastric Emptying: Semaglutide delays the rate at which food leaves the stomach, leading to a
 more gradual absorption of glucose and prolonged feelings of fullness.

• Reduces Appetite:

By acting on appetite centers in the brain, it decreases hunger and food intake, which can lead to weight loss.

It appears to affect leptin, the hormone that tells you that you are full, and ghrelin, the hormone that tells you that you are hungry.

Potential Health Benefits

- Improved Blood Sugar Control: Semaglutide stimulates insulin secretion and suppresses glucagon release in a glucose-dependent manner. This leads to better regulation of blood glucose levels, helping individuals with type 2 diabetes achieve optimal glycemic control.
- Weight Loss: One of the significant benefits of semaglutide is its ability to promote weight loss. It slows gastric emptying and reduces appetite by acting on appetite centers in the brain. Clinical trials have shown substantial weight reduction in both diabetic and non-diabetic individuals using semaglutide.

 Obese people who took injectable semaglutide lost 5-25% of their body weight within 6 months.
- Cardiovascular Benefits: Semaglutide has been associated with a reduction in major adverse cardiovascular events (MACE) such as heart attack and stroke in patients with type 2 diabetes at high cardiovascular risk. It may improve heart health by lowering blood pressure and improving lipid profiles.
- Reduced Risk of Kidney Disease Progression: By improving glycemic control and reducing cardiovascular risk factors, semaglutide may slow the progression of diabetic nephropathy, thus offering renal protective effects.
- Improved Lipid Profile: Treatment with semaglutide has been shown to lower levels of triglycerides and LDL cholesterol (bad cholesterol) while increasing HDL cholesterol (good cholesterol), contributing to a healthier lipid profile.
- Potential Benefits on Blood Pressure: Some studies have noted modest reductions in systolic and diastolic blood pressure among patients treated with semaglutide, which can further reduce cardiovascular risks.

Potential Side Effects

- The most common side effects are gastrointestinal (GI) and include nausea, vomiting, constipation, acid reflux, stomach pain, and diarrhea. Cyanocobalamin (B12) is added to reduce GI side effects.
- May cause loss of muscle if you are not consuming an adequate amount of protein and doing resistance exercise. Glycine supplementation can help prevent muscle loss
- Significant weight loss may cause sagging skin and more pronounced wrinkles. This is not specific to GLP-1 medications. ANYTHING that causes significant weight loss will cause more pronounced wrinkles and sagging skin because the underlying supporting structure of your skin has been stretched out from being overweight. The more overweight you are, the more pronounced this effect is when you lose weight.
- RISK OF THYROID C-CELL TUMORS. If you or any family members have been diagnosed with Multiple Endocrine Neoplasia Syndrome Type 2 or Medullary thyroid cancer you should not take Semaglutide.

- Semaglutide may increase the risk of developing pancreatitis. However, previous pancreatitis has not been shown to increase future risk of pancreatitis with Semaglutide.
- The risk of serious side effects increases in patients with hypoglycemia, kidney problems, and risk of allergic reactions. Increasing at longer intervals helps decrease side effects.
- This medication is not insulin and should not be used if you have type 1 diabetes or if you develop diabetic ketoacidosis. Semaglutide should not be used with any other GLP-1 medications.
- If you develop intolerable nausea or vomiting, go back to the previous dose for a few more weeks and try
 to increase it again later. You do not have to increase to the maximum dose. If you are losing weight and
 not having significant side effects, you can maintain at that dose. If you develop severe constipation or
 abdominal pain, stop the medication and get evaluated by your PCP or Emergency Dept ASAP. If you
 develop swelling in your neck or difficulty swallowing, get evaluated by your PCP or Emergency Dept.

Dosage Guidelines

See dosing schedule at the end of this document.

Semaglutide is given as a weekly subcutaneous injection. The starting dose is 0.25mg weekly, and the maximum dose is 2.5mg weekly. Very few patients require the maximum dosage. The optimal dosage is one in which you are losing weight and don't have intolerable side effects. This is different for each patient.

Medication is stored in the refrigerator.

If you experience intolerable side effects, like nausea and vomiting, go back to the dose you were at before for a week or two and try increasing it again. If intolerable nausea or vomiting persists, just stay at the dose you tolerate.

Cost

Strive Pharmacy - Direct shipping to the following states: (AZ, CO, DC, DE, FL, GA, HI, IA, ID, KS, MA, MD, ME, MO, MS, MT, ND, NE, NH, NM, NY, OH, OK, OR, PA, PR, RI, SD, TN, TX, UT, WA, WY)

Semaglutide/Glycine/B12 – 5mg/5mg/1mg/ml.

\$330 per 2ml. Shipping is \$30. May order up to 4ml at a time.

Strive Pharmacy - Shipping to the following states: (AL, AK, AR, CT, IN, KY, LA, MI, MN, NC, NJ, SC, VA, VT, WI, WV)

Semaglutide/Glycine/B12 - 5mg/5mg/1mg/ml.

\$330 per 2ml. Shipping is \$72. May order up to 4ml at a time.

Research Peptide. Please see document titled "Research Peptide Information" in the Education Folder under Records in the patient portal.

5mg Vial (5mg/ml) - \$183.50 (Includes shipping and bacteriostatic water for reconstitution)

10mg Vial (5mg/ml) - \$298.50 (Includes shipping and bacteriostatic water for reconstitution)

Important Disclosures

- These statements have not been evaluated by the US Food and Drug Administration (FDA).
- Not intended to diagnose, treat, cure, or prevent any disease.
- Research peptides and some compounded drugs are not FDA-approved but are produced under strict quality control measures.

THE FOLLOWING RECONSTITUTION INFORMATION ONLY PERTAINS TO MEDICATIONS IN POWDER FORM.

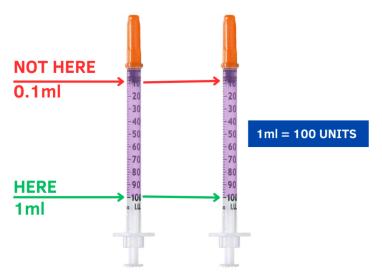
Reconstitution Instructions

IMPORTANT:

- Follow the instructions below regarding the amount of bacteriostatic water to use when reconstituting the peptide. DO NOT follow the instructions that come with the peptide.
- Do NOT throw away the vial of bacteriostatic water!!! It is a multiuse vial and can be used for your next order!

For a 5mg Vial: Inject 1 ml of bacteriostatic water into the vial of powder (1ml = 100 units).

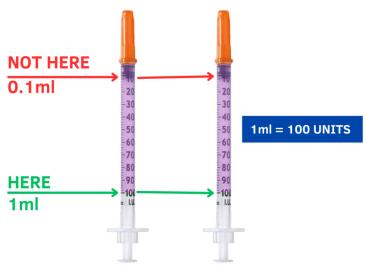
1 ml - 100 units (Not 10 UNITS)



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For a **10mg Vial**: Inject **2 ml of bacteriostatic water** into the vial of powder (2ml = 200 units). You will need to inject 2 full 1ml syringes of water into the vial.

1 ml - 100 units (Not 10 UNITS)



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• See the **document** titled "Reconstituting Medications in Powder Form" in the Education Folder in the patient portal.

- See the following **Instructional videos** in the Education Folder in the patient portal:
 - "Reconstituting Powdered Medications"
 - "Injection Video Introduction"
 - "Injection Video Drawing Up the Medication"
 - "Injection Video Administering the Medication"

Storage and Stability for Research Peptides

- Vials are shipped as **lyophilized powder**, requiring no refrigeration during shipping.
- In Lyophilized Form:
 - Stable for up to 3 years in the freezer and 2 years in the refrigerator.
 - Protect from light.
- Once Reconstituted:
 - Stable for 6 weeks.
 - Must be refrigerated and kept away from light.
 - Avoid placing vials in the refrigerator door to prevent degradation from frequent temperature changes.

Quality Assurance

- All research peptides are subjected to third-party testing with publicly available Certificates of Analysis (COA).
- Testing includes:
 - RP-HPLC (Reversed-Phase High-Performance Liquid Chromatography)
 - Mass Spectrometry (MS)
 - Sterility Testing
 - Additional tests meeting or exceeding U.S. Pharmacopeia (USP) and USP-National Formulary (NF) regulations.

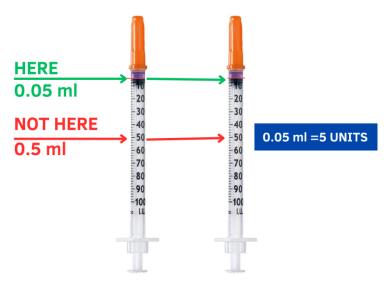
The manufacturer ensures quality, safety, and efficacy, complying with regulatory standards.

Dosing Schedule for Strive Compounded Semaglutide and Research Peptide

Weeks 1 through 4

Dose: 0.25mg

How much you inject: 0.05ml (or 5 units on insulin syringe) SQ once a week for 4 weeks

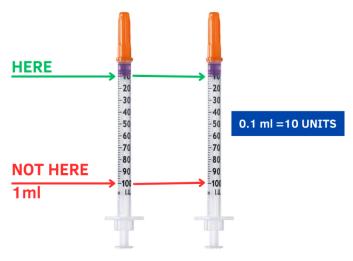


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Weeks 5 through 8

Dose: 0.5mg

How much you inject: 0.1ml (or 10 units on insulin syringe) SQ once a week for 4 weeks

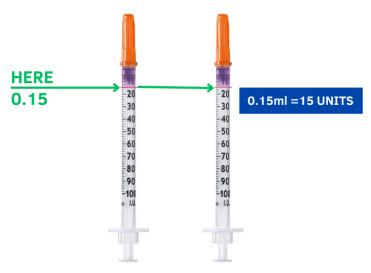


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Weeks 9 through 12

Dose: 0.75mg

How much you inject: 0.15ml (or 15 units on insulin syringe) SQ once a week for 4 weeks

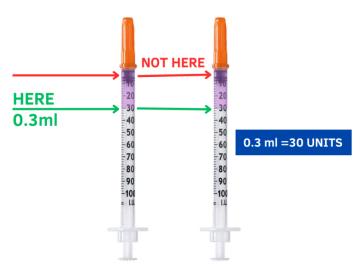


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Weeks 13 through 16

Dose: 1.5mg

How much you inject: 0.3ml (or 30 units on insulin syringe) SQ once a week for 4 weeks



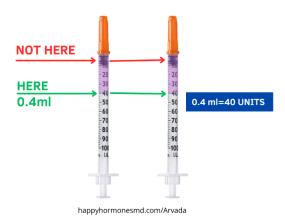
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Weeks 17 through 20

Dose: 2mg

How much you inject: 0.4ml (or 40 units on insulin syringe) SQ once a week for 4 weeks

0.4ml - 40 units (Not 4 UNITS)



Weeks 21 and on

Dose: 2.5mg

How much you inject: 0.5ml (or 50 units on insulin syringe) SQ once a week for 4 weeks

0.5ml - 50 units (Not 5 UNITS)

