

# THE GLP-1 / GPCR GOLD RUSH

A strategic analysis of a market reshaping medicine, from blockbuster drugs to the vast therapeutic frontier ahead.

# Executive Summary: The Next Strategic Frontiers

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The GLP-1 market is a \$36B duopoly set to exceed \$200B by 2031. The smartest path forward is not head-to-head competition, but capturing the immense value in solving for the tolerability, muscle loss, and convenience gaps left by today's blockbusters.

## Opportunity 1: Improved Tolerability & Adherence

High discontinuation rates (47-65%) due to GI side effects create a massive opportunity for next-generation drugs with better tolerability profiles, achieved through novel mechanisms like biased agonism or allosteric modulation.

## Opportunity 2: Muscle Mass Preservation

Significant muscle loss (up to 25% of total weight) is a key concern with current GLP-1s. Combination therapies with muscle-preserving agents (e.g., myostatin inhibitors) represent a critical and valuable area of R&D.

## Opportunity 3: Convenient Oral Formulations

Strong patient preference for oral drugs over injectables makes the development of effective, once-daily small molecules a top strategic priority for expanding market access, improving adherence, and capturing market share.

## Opportunity 4: Novel GPCR & Non-incretin Targets

With hundreds of GPCRs still undrugged, there is immense potential for first-in-class therapies targeting novel pathways beyond GLP-1/GIP to treat obesity and related metabolic disorders, opening up new therapeutic avenues.

## Opportunity 5: Expanded Indications

The proven benefits of GLP-1s in cardiovascular, kidney, and liver disease are just the beginning. Repurposing these drugs for a wide range of new indications is a massive value-creation opportunity.

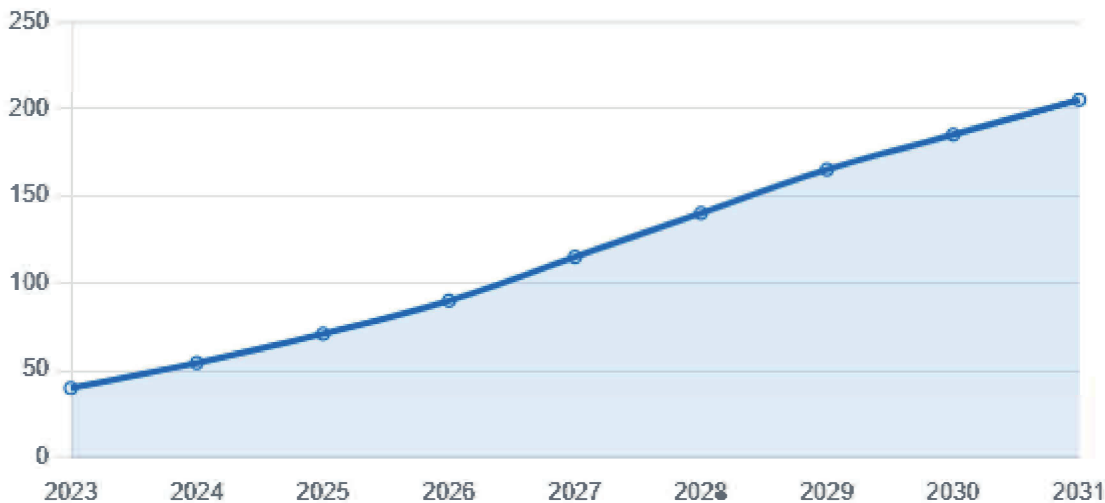
## Opportunity 6: Digital Health & Wraparound Services

A clear need exists for digital tools to support patients on GLP-1 therapy, helping with adherence, managing side effects, tracking progress, and providing lifestyle coaching for a holistic approach to care.

# An Exploding Market Dominated by a Duopoly

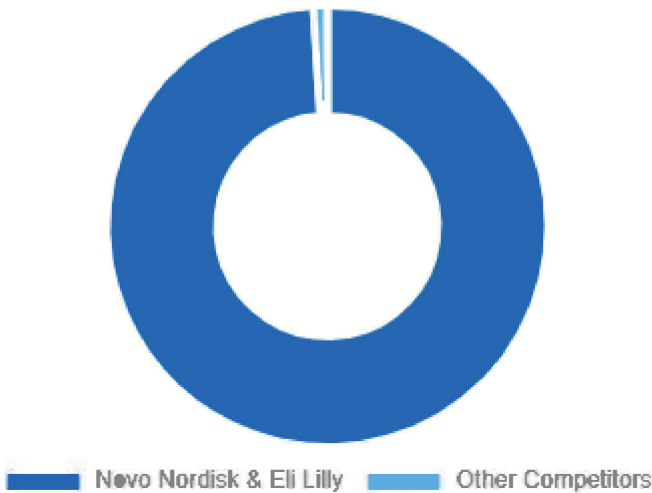
The GLP-1 market is experiencing unprecedented growth, driven by transformative therapies for diabetes and obesity. This expansion has created a landscape where two key players, Novo Nordisk and Eli Lilly, hold a commanding position.

Projected GLP-1 Market Growth (USD Billions)



Source: Morningstar Equity Research

2023 GLP-1 Market Share



Source: Fierce Pharma, GlobalData

**\$36 Bn**

Total GLP-1 Global Sales (2023)

Source: Fierce Pharma

**~38%**

Annual Rx Growth (2022-2024)

Source: Medscape

**25-50M**

Projected US Users by 2030  
(23-46% of US obese population)

Source: Goldman Sachs

# The Macro Environment: A Perfect Storm of Opportunity & Risk

The GLP-1 phenomenon does not exist in a vacuum. It is being shaped by powerful external forces, from intense political pressure on drug pricing in the US to a broader societal shift towards a comprehensive, multi-trillion dollar wellness market.

## Pricing & Political Headwinds

In the US, which accounts for ~40% of global pharma sales, the administration has set targets to slash drug prices. Measures like the Inflation Reduction Act (IRA), which allows Medicare to negotiate prices for key drugs (including semaglutide), and the threat of tariffs create significant long-term pricing pressure and market access risk.

## Strategic Response: Reshoring Production

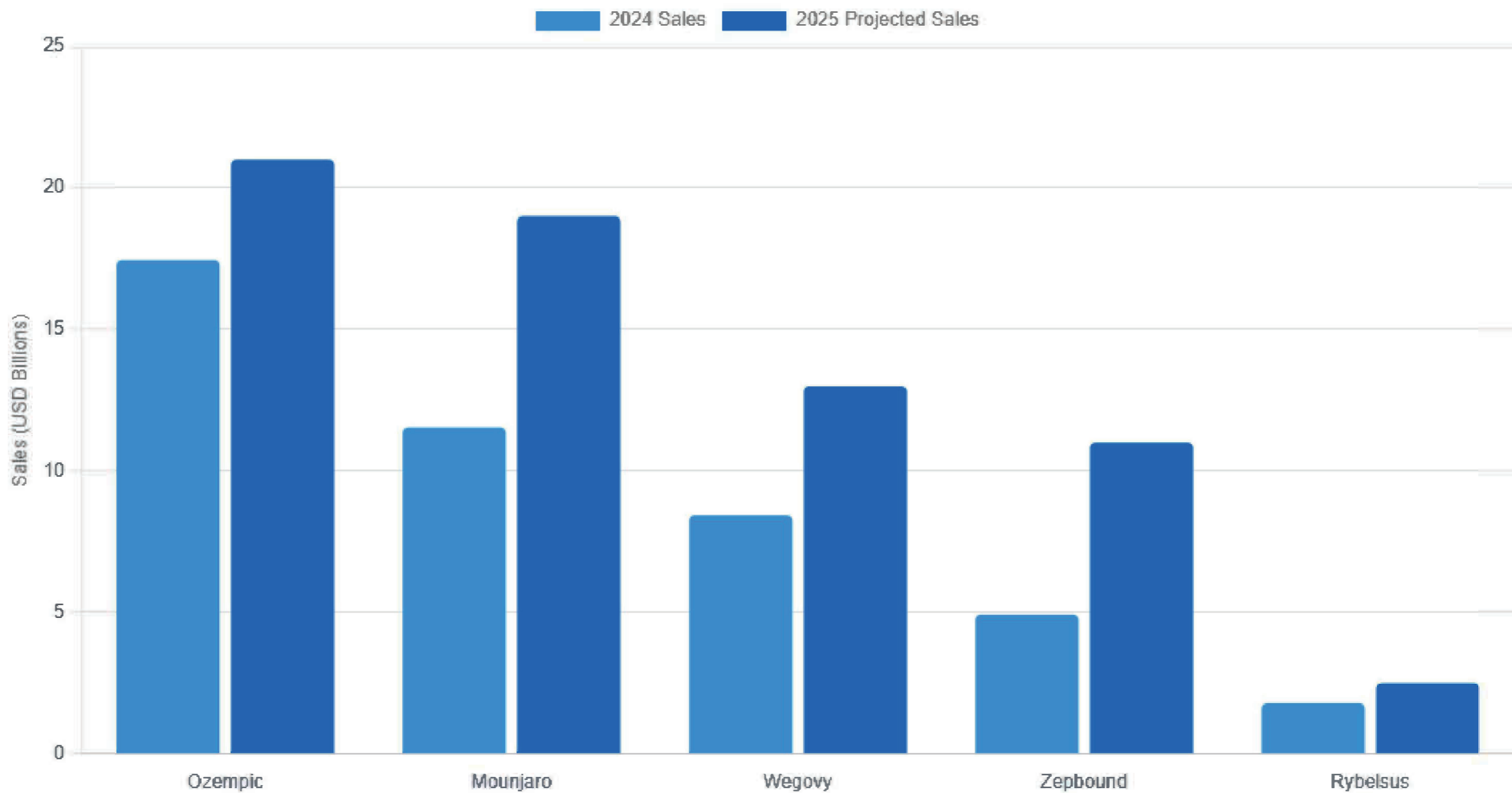
As a direct response to potential tariffs and supply chain vulnerabilities, major pharmaceutical companies are announcing massive investments to increase US-based R&D and manufacturing. These multi-billion dollar commitments (e.g., Novo's acquisition of Catalent) are a key strategic hedge against geopolitical risk.

## The Expanding Wellness Ecosystem

The rise of GLP-1s is fueling a broader \*\*\$2 trillion global wellness market\*\*. This creates a "halo effect," boosting demand for complementary products and services like medical aesthetics, specialized nutrition (protein-fortified foods), and fitness programs focused on preserving muscle mass during weight loss.

# The Billion-Dollar Blockbusters

A handful of key drugs are driving revenue, with sales projections for 2025 indicating sustained and massive growth for the leading products.



Source: Statista, Bloomberg Intelligence, Eli Lilly Q4 2024 Earnings

# The GLP-1/GPCR Pipeline: A Competitive Snapshot

The pipeline is rich with multi-receptor agonists, oral formulations, and novel mechanisms aiming to improve upon the efficacy and tolerability of current market leaders.

## On the Market

### Wegovy/Ozempic/Rybelsus (Semaglutide)

Novo Nordisk | GLP-1RA. Approved for T2D, Obesity, CV Risk Reduction.

### Zepbound/Mounjaro (Tirzepatide)

Eli Lilly | Dual GLP-1/GIP Agonist. Approved for Obesity, T2D, Sleep Apnea.

### Saxenda (Liraglutide)

Novo Nordisk | GLP-1RA. Approved for weight loss, lost exclusivity end of 2024.

## In Clinical Trials

### Retatrutide (LY3437943)

Eli Lilly | Triple GLP-1/GIP/Glucagon Agonist. Phase 3; showed 24.2% weight loss.

### CagriSema

Novo Nordisk | GLP-1/Amylin Combo. Phase 3; showed 22.7% weight loss.

### Orforglipron (LY3502970)

Eli Lilly | Oral GLP-1RA. Phase 3; showed ~8% weight loss.

### Survodutide (BI 456906)

Boehringer/Zealand | Dual GLP-1/Glucagon. Phase 3; ~15% weight loss.

### VK2735

Viking Therapeutics | Dual GLP-1/GIP. Phase 2; showed 14.7% weight loss.

### Mazdutide (IBI362)

Innovent Bio | Dual GLP-1/Glucagon. Phase 3 (China); 15.4% weight loss.

### Danuglipron

Pfizer | Oral GLP-1RA. Reformulated, entering Phase 1.

### GSBR-1290

Structure Therapeutics | Oral GLP-1RA. Phase 2; showed 6.2% weight loss.

### Petrelintide (ZPR8396)

Roche/Zealand | Amylin Analog. Phase 2b trial initiated Dec 2024.

## Pre-Clinical & Novel Mechanisms

### Amycretin

Novo Nordisk | Oral GLP-1/Amylin. In pipeline, targeting high efficacy oral option.

### Bimagrumab Combos

Eli Lilly | Myostatin Inhibitor. For muscle preservation alongside GLP-1s. Phase 2.

### Non-incretin GPCRs

Novo/Deep Apple | Oral Small Molecules targeting novel pathways for cardiometabolic diseases.

Source: Company filings & press releases (Eli Lilly, Novo Nordisk, Roche, Pfizer), PitchBook, Morningstar, Medscape, Statista, Fierce Pharma.

## The Patient Experience Gap & Tolerability

Despite incredible efficacy, high patient discontinuation rates due to side effects reveal a critical unmet need for therapies with improved tolerability and muscle preservation.

**47%**

### Discontinuation Rate (T2D)

among patients with Type 2 Diabetes within one year, largely due to gastrointestinal side effects.

Source: American Diabetes Association Scientific Sessions 2023

**65%**

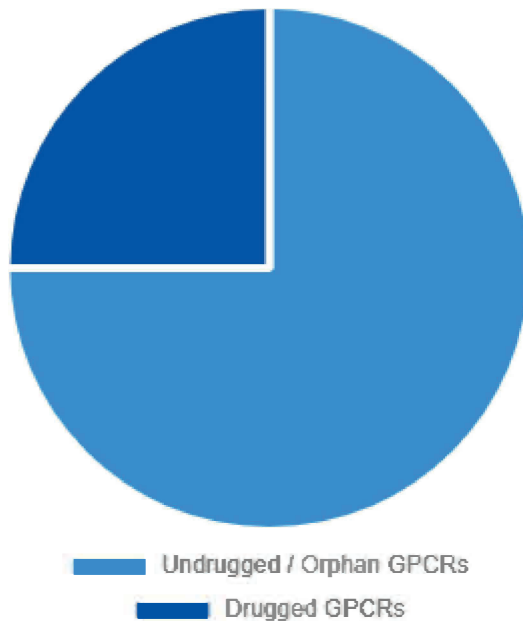
### Discontinuation Rate (Obesity)

among patients with obesity within one year, highlighting the trade-off between efficacy and real-world adherence.

Source: American Diabetes Association Scientific Sessions 2023

# The GPCR Frontier: A Vast Untapped Opportunity

The GLP-1 receptor is part of the vast G Protein-Coupled Receptor (GPCR) family, the most successful drug target class in history. Yet, most GPCRs remain undrugged, representing an immense frontier for future innovation.



Source: Nature, Trends in Pharmacological Sciences

## Advanced GPCR Targeting Strategies

- **Biased Agonism:** Tailors signaling pathways to achieve specific outcomes, minimizing negative effects (e.g., Roche's CT-388).
- **Allosteric Modulators:** Bind to distinct sites to fine-tune receptor activity, offering advantages in specificity and reduced desensitization.
- **Bivalent & Bitopic Ligands:** Emerging strategies involving ligands designed to bind to multiple sites on a receptor for improved selectivity.
- **Cryptic Binding Pockets:** Computational tools are identifying previously unknown pockets on GPCRs, leading to novel therapeutic approaches.

• Source: Nature, Trends in Pharmacological Sciences, Roche



# Drug Modalities: The Strategic Toolkit

The GLP-1/GPCR pipeline employs diverse drug modalities, each with distinct advantages in terms of patient convenience, efficacy, and target specificity.

## Peptides & Proteins

These include the core GLP-1s and next-generation multi-agonists (GIP, glucagon, amylin). Their modular nature allows for precise tuning of receptor binding and activity, enabling superior weight loss and cardiometabolic benefits.

Source: Nature, Medscape, Eli Lilly, Novo Nordisk

## Small Molecules

Comprising 71% of drugs in clinical trials, small molecules offer advantages in oral bioavailability and cost-effective manufacturing. Oral small molecule GLP-1s (e.g., orforglipron) are a "next competitive frontier" for convenience.

Source: Nature, Eli Lilly, Pfizer

## Administration Routes

While injectables currently dominate, there's a strong patient preference (3:1) for oral options due to convenience. The shift to oral formulations is a key strategic goal to expand market access and improve long-term adherence.

Source: American Diabetes Association Scientific Sessions 2023

## Blockbuster M&A: The Race for Pipeline & Capacity

Major players are executing multi-billion dollar deals to acquire next-generation assets and secure the manufacturing capabilities required to meet unprecedented demand.

Manufacturing & Capacity

**Novo Holdings Acquires  
Catalent**

**\$16.5B**

A transformational acquisition to secure critical fill-finish manufacturing for GLP-1 production at scale (2024).

Pipeline Expansion

**Roche Acquires Carmot**

**\$2.7B+**

Strategic entry into metabolic disease with dual GLP-1/GIP and oral programs (2024).

Pipeline Enhancement

**Eli Lilly Acquires VersanisBio**

**\$1.93B**

Strengthens pipeline with bimagrumab, a muscle-preserving agent for obesity (2023).

# Following the Money: Fundraising in the GLP-1/GPCR Space

Significant venture capital and private equity investment is flowing into the next generation of companies developing novel therapies and enabling technologies, signaling strong confidence in the sector's future.

## Market Leaders (Total Raised)

Pfizer	\$34.5B (2025)
Amgen	\$31.8B (2025)
Merck & Co.	\$30.0B (2025)
AstraZeneca	\$23.9B (2025)
Novartis	\$13.1B (2025)
Eli Lilly & Co.	\$11.85B (2025)
Novo Nordisk	\$11.85B (2025)
GSK plc	\$8.85B (2025)
Sanofi	\$1.5B (2025)

## Key Innovators & Biotechs (Total Raised)

Human Longevity	\$1.09B (2024)
Viking Therapeutics	\$985.10M (2025)
Structure Therapeutics	\$937M+
Metsera	\$625.00M (2025)
Isomorphic Labs	\$579.06M (2025)
Kallyope	\$479.0M (2024)
MBX Biosciences	\$403.08M (2025)
BioAge Labs	\$318.2M (2024)
Ventus Therapeutics	\$300.0M (2024)

# White Space Analysis: The Next Therapeutic Frontiers

The success of first-generation GLP-1s has illuminated significant unmet needs and vast opportunities for next-generation therapies and complementary solutions.

## Improved Tolerability & Adherence

High discontinuation rates due to GI side effects create a major opportunity for drugs with better tolerability profiles, achieved through biased agonism, novel targets, or formulation strategies.

## Muscle Mass Preservation

Significant muscle loss is a key concern. Co-therapies with muscle-preserving agents (e.g., myostatin inhibitors) or novel agonists with different metabolic effects represent a critical area of R&D.

## Convenient Oral Formulations

The strong patient preference for oral drugs over injectables makes the development of effective, once-daily small molecules a top strategic priority for expanding market access and long-term adherence.

## Novel GPCR & Non-incretin Targets

With hundreds of GPCRs still undrugged, there is immense potential for first-in-class therapies targeting novel pathways beyond GLP-1/GIP to treat obesity and related metabolic disorders.

## Expanded Indications

The proven benefits of GLP-1s in cardiovascular, kidney, and liver disease are just the beginning. Repurposing these drugs for a wide range of new indications is a massive value-creation opportunity.

## Digital Health & Wraparound Services

A clear need exists for digital tools to support patients on GLP-1 therapy, helping with adherence, managing side effects, tracking progress, and providing lifestyle coaching for a holistic approach to care.

# Competitive Dynamics & Strategic Shifts

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The market is defined by intense competition, aggressive M&A, strategic partnerships, evolving regulatory landscapes, and the looming impact of patent expirations.

## M&A Activity

**Roche acquires Carmot Therapeutics (\$2.7B+):** Strategic entry into metabolic disease with dual GLP-1/GIP and oral programs (2024).

**Novo Holdings acquires Catalent (\$16.5B):** Secures critical fill-finish manufacturing capacity for GLP-1 production (2024).

**Eli Lilly acquires VersanisBio (\$1.93B):** Strengthens pipeline with muscle-preserving agent bimagrumab (2023).

## Strategic Partnerships

**Roche & Zealand Pharma (\$5.3B):** Co-development of non-GLP1 amylin analog, petrelintide (2025).

**Novo Nordisk & Deep Apple (\$812M):** Collaboration for novel oral small molecule drugs targeting non-incretin GPCRs (2025).

**Eli Lilly & Scribe Therapeutics:** Partnership to develop in vivo CRISPR-based therapeutics.

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## Regulatory & Market Access

**Medicare Coverage:** Part D now covers Wegovy for its cardiovascular indication, but not solely for weight loss. Semaglutide is set for IRA price negotiations in 2027.

**Patent Expirations:** Novo Nordisk's Saxenda lost exclusivity at the end of 2024 (generics in 2025). Semaglutide's patent expires in 2031, opening the door for biosimilars.

**Global Generics:** China launched its first domestic GLP-1 (beinaglutide) in 2023 and is actively pursuing semaglutide generics. The first exenatide generic launched in Nov 2024.

# Sources & Methodology

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## Methodology

This report was produced through a hybrid analytical approach. An AI-powered large language model (LLM) was used as a research assistant to rapidly query, extract, and synthesize key data points from a large corpus of source documents. This included market research reports, company financial filings, press releases, and scientific journals. All quantitative data and key facts were then verified and curated by the author to build the core narrative and strategic insights presented in this document. This method allows for a comprehensive and efficient analysis that combines the speed of machine learning with the critical thinking of human expertise.

## Major Sources & Data Providers

- PitchBook
- Morningstar Equity Research
- GlobalData
- Fierce Pharma
- Statista
- Bloomberg Intelligence
- American Diabetes Association (ADA)
- The New England Journal of Medicine (NEJM)
- Nature
- The Lancet
- JAMA
- Economist Intelligence Unit (EIU)
- McKinsey & Company
- Company Filings & Press Releases