

Obesity Drugs: The Next Wave of GLP-1 Competition

New drugs from public and private companies will challenge Novo and Lilly's dominance.

Morningstar Equity Research

September 2024

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First Wave of Obesity Drugs Led by Novo and Lilly

New public and private entrants are emerging to challenge the leading positions of Novo Nordisk and Eli Lilly in a potential \$200 billion total GLP-1 market, which we expect will see 68% of sales from weight loss indications by 2031. Given first-mover advantages and innovation with next-generation products, we expect Novo and Lilly to retain two thirds of the total market by 2031, reinforcing their wide moats.

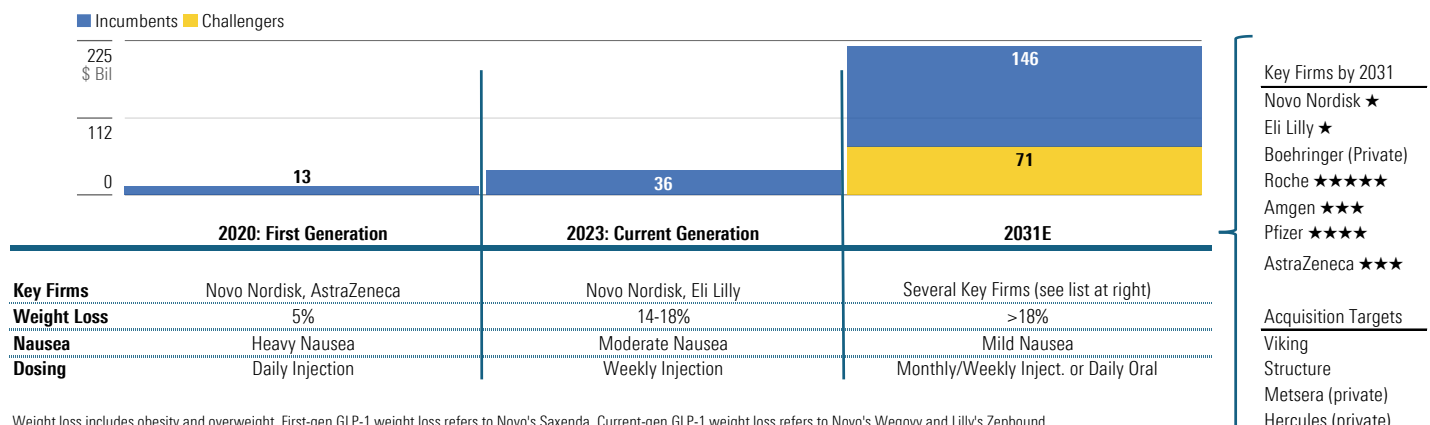
Next Wave of Obesity Drugs

Midstage cardiometabolic pipelines are filling up across public and private biopharma as Novo and Lilly's success revitalizes drug development. We expect 16 new obesity drugs could launch by 2029, with roughly \$70 billion of the GLP-1 market coming from these new challengers by 2031.

Key Takeaways

- ▶ Novo Nordisk and Eli Lilly are launching leading obesity drugs that will support a \$200 billion total GLP-1 market. While we assume they will retain nearly 70% of the market, we view the stocks as overvalued.
- ▶ Over the next three to four years, next-generation obesity drugs from Roche (FVE: \$55), Amgen (FVE: \$317), Pfizer (FVE: \$42), AstraZeneca (FVE: \$78), private firm Boehringer, and acquisition targets including Viking and Structure will likely take market share and pressure drug pricing.
- ▶ Led by Metsera and Hercules (both acquisition targets), venture capital/private obesity drug development will layer in new drugs. On pages 19-20, we rank the most likely acquirers and targets.

Exhibit 1 GLP-1 Evolution: Key Private and Public Companies Challenge Incumbents Novo and Lilly in a \$200 Billion Potential Market



Weight loss includes obesity and overweight. First-gen GLP-1 weight loss refers to Novo's Saxenda. Current-gen GLP-1 weight loss refers to Novo's Wegovy and Lilly's Zepbound.



GLP-1 Drugs Represent Paradigm Shift for Obesity Treatment, but Valuations Look Steep

GLP-1 drugs act similarly to the hormone glucagon-like peptide 1 (GLP-1), which helps the body produce insulin to help treat diabetes and slows the movement of food from the stomach to help treat obesity.¹ While initially approved for diabetes, GLP-1 drugs are gaining new indications, including the large opportunity in obesity following decades of failed attempts to create an effective and safe drug.

Novo Nordisk's semaglutide (Ozempic and Rybelsus for diabetes and Wegovy for obesity) and Eli Lilly's tirzepatide (Mounjaro for diabetes and Zepbound for obesity) are the clear leaders in the GLP-1 class. Novo Nordisk and Eli Lilly hold a significant first-mover advantage, as most other firms largely reduced cardiometabolic research a decade ago due to a lack of innovation, side effects, and lower pricing power.

However, new entrants are emerging, enticed by an expected \$200 billion market. We expect significant market share penetration in obesity but also pricing pressure, multiple new drug launches, and several potential acquisitions to make the market more competitive than what investors are currently assuming (Exhibit 2).

Exhibit 2 PitchBook/Morningstar Analysis Offers a Different Take on a Rapidly Growing Obesity Market

 Market Perspective	 PitchBook/Morningstar Take	Reference Pages
Novo Nordisk's and Eli Lilly's obesity drugs face limited competitive risks.	Competitive obesity drugs will likely begin to launch in 2026.	Page: 3 Exhibit: 3
The obesity driven GLP-1 market is very large, but complex to quantify.	We quantify key assumptions to support a \$200 billion annual market.	Page: 4 Exhibit: 4
Obesity drug penetration will be high.	Relative to other blockbuster markets like atopic dermatitis, asthma and psoriasis, we expect obesity drugs to post higher market penetration.	Page: 5 Exhibit: 5
Obesity drug pricing will remain fairly strong.	Pricing will likely fall in the near term to gain expanded payor access, and price declines will accelerate to over 10% annually by 2027 as more competition enters the market.	Page: 5 Exhibit: 6
Secondary indications beyond obesity will drive significant sales.	The high overlap between obesity and other cardiovascular disease will largely just help market penetration into obesity.	Page: 6 Exhibit: 7
Eli Lilly and Novo Nordisk look undervalued.	Eli Lilly and Novo Nordisk look overvalued, with high market share retention and stable pricing needed to support current price and undervalued calls.	Page: 7 Exhibit: 8
The next wave of obesity drugs is building, but too far away to value.	The next wave of drugs is poised to launch in the 2026-2029 timeframe and could improve on current mechanisms of action and bring novel targets.	Page: 7 Exhibit: 9
The efficacy of Eli Lilly and Novo Nordisk obesity drugs sets a high bar.	The next wave of obesity drugs is surpassing Eli Lilly and Novo Nordisk drugs, likely on efficacy, convenience, and tolerability.	Pages: 8-10 Exhibit: 10-11
The market recognizes new strategies are emerging to treat obesity.	We highlight the major evolutions of obesity drug treatment strategies.	Page: 11 Exhibit: 12
Investments are increasing into earlier stage obesity drugs.	We highlight the key buyers and sellers in the public and private markets.	Pages: 14-20 Exhibits: 21-23




















Source: Morningstar.

One the most dynamic parts of this market is the next wave of obesity drugs in the pipeline, with potential launches from new challengers potentially starting in 2026 with Boehringer Ingelheim and

¹ Mayo clinic.

Zealand's survodutide. Several launches in 2027 (Amgen, Altimmune), 2028 (Pfizer, Roche, Viking, and Structure), and 2029 (Roche, Viking, AstraZeneca, and Zealand) stand out to us as potentially competitive with Novo Nordisk and Eli Lilly. That said, Novo and Lilly are also advancing their next-generation pipelines, beginning with Novo's cagrisema and Lilly's orforglipron in 2026.

Exhibit 3 Potential Launch Timeline for New GLP-1 Therapies and Combinations: Competition to Novo and Lilly Begins in 2026

		Approved	2025E	2026E	2027E	2028E	2029E+
GLP-1	injectables	Wegovy (2021) 					
	orals		Rybelsus 	orforglipron 		danuglipron  GSBR-1290 	CT-996  AZD5004 
+GIP	injectables	Zepbound (2023) 			maritide 	CT-388  VK2735 	
	orals						VK2735 
+amylin	injectables			cagrisema 			petrelintide 
	orals						amycretin  amycretin
+glucagon (injectable)				Boehringer Ingelheim survodutide 	altimmune pemvidutide  retatrutide*		
+CB1 (oral)							monlunabant 
+activin/myostatin (injectable)						bimagrumab 	

Source: Morningstar, company reports. *triple GLP-1/GIP/glucagon.

\$200 Billion GLP-1 Market Size to Attract New Entrants

We estimate the GLP-1 drug market to reach annual sales of more than \$200 billion and highlight the key assumptions behind the \$200 billion annual market (Exhibit 4).

Exhibit 4 GLP-1 Drug Market Projections With Key Assumptions

	2023	2028E	2031E	2031 Market Assumptions
				Patients Treated with GLP-1
US Diabetes Patients (mil)	5.2	13	16.7	41%
US Obesity Patients* (mil)	0.9	11.2	21.1	23%
US Overweight Patients** (mil)	0.2	3	4.8	13%
US Total GLP-1 Patients (mil)	6.3	27.2	42.6	
Percentage of US Adults	2%	10%	16%	
International Diabetes Patients (mil)	12.2	41.6	57.6	19%
International Obesity Patients* (mil)	0.4	8.3	23.9	7%
International Overweight Patients** (mil)	0.1	4.3	10.6	1%
International Total GLP-1 Patients	12.7	54.2	92.1	
Percentage of Intl Adults	0%	1%	2%	
				Insured Versus Out-of-Pocket Pricing
Price (USD)	2023	2028E	2031E	
US Diabetes	3,739	2,817	2,415	Primarily net insured price
US Obesity	7,107	4,253	2,921	Weighted average of insured/out-of-pocket price
International Diabetes	749	579	497	Primarily net insured price
International Obesity	3,300	2,553	2,189	Mostly out-of-pocket
				Novo Nordisk/Eli Lilly Combined Market Share
Total Market (USD Bil)	36	150	217	67% (2031E)

Source: Morningstar, company reports. *Non-diabetic. **Includes overweight patients treated for cardiovascular risk and kidney disease.

Solid Market Penetration and Strong, but Weakening, Pricing Support the Large Obesity Market

Two key assumptions drive our outlook: 1) market penetration rates, and 2) obesity drug pricing. On US market penetration, we project 41% of diabetics and close to one quarter of nondiabetic obesity patients will be on a GLP-1 drug by 2031 based on the strong efficacy of the drugs and likely robust insurance coverage. As shown in Exhibit 5, our assumed 2031 GLP-1 penetration rates are ahead of advanced drug penetration in psoriasis and atopic dermatitis, but the high media coverage of the GLP-1 class likely means higher market share gains. Internationally, we project much lower market penetration rates in obesity (especially outside of developed markets) due to lack of robust insurance coverage causing pricing to remain too high for most patients to utilize the drugs.

Exhibit 5 Market Penetration by Major Diseases Relative to Projected GLP-1 Market Penetration

	2022 Market Data (US/EU5)				2031E (US)	
	Atopic Dermatitis	Asthma	Psoriasis	Rheumatoid Arthritis	GLP-1 Obesity	GLP-1 Diabetes
Treated with advanced therapy	9%	17%	23%	41%	23%	41%
Years with advanced therapy on the market	7	21	17	26	10	21

Source: Morningstar, [Sanofi](#), [CDC](#), company reports.

Note: Markets are US, EU5 (France, Germany, UK, Spain, Italy) except GLP-1 obesity and diabetes, which are US.

Note: Wegovy's 2021 approval is used for the 10-year marker for obesity and Victoza's 2010 approval is used for diabetes.

Obesity Drug Pricing in the US Will Likely Fall Faster Over the Next Decade

Over the next two years, we expect Novo and Lilly to concede price declines to expand insurance coverage over larger patient groups. By 2027 and beyond, we expect new entrants to cause the annual pricing declines to accelerate toward 10%-15% as competitors work to gain insurance coverage. While the current GLP-1s haven't faced as steep pricing declines as other diabetes medicines such as DPP-4s and SGLT-2s, recent pricing declines have accelerated, as we believe Novo and Lilly increased pricing competition when both firms launched a weekly GLP-1 drug. We expect the pricing pressure to accelerate as new GLP-1s reach the market with similar efficacy, as seen with the cardiometabolic drug class of DPP-4s and SGLT-2s (Exhibit 6). Additionally, we expect forced Medicare price negotiations for Wegovy (as part of the Inflation Reduction Act) to also pressure GLP-1 pricing starting in 2027.

Exhibit 6 Historical and Projected Pricing Dynamics in Cardiometabolic Drugs

Cardiometabolic Drug Class	DPP-4s	SGLT-2s	GLP-1s: 2010-2026E	GLP-1s: 2027E-2031E
Variable Response Rate/Dosing	No: largely similar	No: largely similar	Yes: Novo and Lilly GLP-1 drugs offered better cardiovascular outcomes and easier weekly dosing.	No: Likely interchangeability in Obesity
More than two companies	Yes: 4 key competitors	Yes: More than 5 key competitors	No: Largely only Novo and Lilly	Yes: Likely to have more than 5 key competitors
Estimated peak annual price declines	>10%	>10%	Slight pricing gains initially, followed by moderate pricing declines	>10% in US Obesity Market

Source: Morningstar, [Health Affairs](#), and company reports.

Note: DPP-4s (Dipeptidyl peptidase-4) include Januvia, Onglyza, Tradjenta and others. SGLT-2s (sodium glucose cotransporter 2) include Farxiga, Jardiance, Invokana, and others.

New GLP-1 Indications Should Primarily Support Market Penetration Into Obesity

The most important market for GLP-1 drugs is obesity due the large prevalence rate and limited prescription options. However, additional indications in obesity-related diseases should help with obesity market penetration and insurance coverage. As shown in Exhibit 7, obese patients significantly outnumber other obesity-related indications. Further, the overlap is very high in obesity-related indications and obesity. As a result, we expect new obesity-related indications to largely help with market penetration within the obesity market and to a lesser extent into the overweight market.

Exhibit 7 Overlap of Obesity Patients and Obesity-Related Indications

Disease	Sales Opportunity	US Patients (millions)	Also Have Obesity	Other Prescription Options	GLP-1 Status
Obesity	Major	115	100%	Few	Approvals for Novo's Saxenda (2014) and Wegovy (2021) and Eli Lilly's Zepbound (2023).
Type 2 Diabetes	Major	35	60%	Many	Many GLP-1 approvals, first was exenatide in 2005.
MASH	Minor	13	85%	Few	Boehringer's survodutide and Lilly's Zepbound posted positive Phase 2 data in MASH in 2024.
Heart Failure	Minor	7	50%	Several	Novo's semaglutide saw a 41% reduced risk of worsening HF in post-hoc analysis (2026E approval), while Lilly's trial showed a 38% reduction in HF outcomes (2025E approval).
Atherosclerotic Disease	Minor	21	48%	Many	Novo's Ozempic and Lilly's Trulicity have shown 26% and 12% reduction in cardiovascular events for diabetics, respectively. Novo's Wegovy showed 20% reduction in similar events in obesity.
Chronic Kidney Disease	Moderate	17	41%	Several	Novo's Wegovy reduced adverse kidney disease events by 22%, likely approval in early 2025.

Source: Morningstar, Novo Nordisk Capital Markets Day 2024.

Valuation Outlook for Obesity Drug Leaders Novo and Eli Lilly

We view Novo Nordisk and Eli Lilly as overvalued with the market likely projecting too much market penetration for obesity drugs or expecting obesity drug pricing to remain stable. In Exhibit 8, we highlight the unlikely scenarios needed to support the current valuations of Novo and Lilly as well as the very unlikely outcome that the stocks are currently undervalued. While Novo and Lilly have many drugs currently marketed and in the pipeline, we have isolated the scenarios to the obesity market as we believe that is the largest factor that can influence the firms' valuations.

To support the current valuations of Eli Lilly and Novo Nordisk, US GLP-1 market penetration into obesity would need to reach close to 75% with international obesity market penetration reaching close to 25% with only low-single-digit pricing declines. These high penetration rates with more moderate pricing declines seem very unlikely. Further, for the scenario to support the stocks as undervalued, Lilly and Novo would also have to retain close to 95% of the market by 2031, which seems highly unlikely given the increasing competitive pressures.

Exhibit 8 2031 Valuation Scenarios for Eli Lilly and Novo Nordisk

Scenario	Most Likely Outcome	Potential Bullish Outcome	Unlikely Bullish Outcome
	Base Case	Increased Market Penetration Less Pricing Pressure	Increased Market Penetration Stable Pricing Increased Market Share
U.S. GLP-1 Obesity Market Penetration	25%	60%	60%
International GLP-1 Obesity Market Penetration	7%	25%	25%
Obesity Drug Annual Pricing Change 2024-2031	Low-Double Digit Declines	Mid-Single Digit Declines	Stable
Novo Nordisk and Lilly Combined Market Share	Close to 70%	Close to 70%	85%
Valuation of Novo Nordisk and Lilly	Overvalued	Fairly Valued	Undervalued

Source: Morningstar, company reports. Note: Assumes largely successful transitions to next-generation obesity drugs for Novo and Lilly as well as some moderate mix shift to Lilly from Novo by 2031.

New Drugs to Challenge Novo Nordisk’s and Eli Lilly’s Obesity Drug Leadership

Novo Nordisk and Eli Lilly have set the bar for efficacy and side effect profiles with obesity drugs Wegovy and Zepbound, but we expect the next generation of drugs will likely reach or exceed these drug profiles and potentially increase the convenience of dosing through oral dosing or monthly injection dosing. In Exhibit 9, we highlight the most important obesity drugs and drug candidates.

Exhibit 9 Comparing Obesity Incumbents to Challengers: Key Potential Differentiators

Firm	Drug	Mechanism of Action					Completed Phase	Oral?	Small Molecule?	Efficacy	Tolerability
		GLP-1	GIP	Glucagon	Amylin	CB1					
Incumbents: Approved											
Novo Nordisk	Wegovy	✓					3	No	No	Lagging	Solid
Eli Lilly	Zepbound	✓	✓				3	No	No	Solid	Solid
Incumbents: Next Generation											
Novo Nordisk	Rybelsus 50 mg	✓					3	Yes	No	Lagging	Solid
Novo Nordisk	cagrisema	✓			✓		1b	No	No	Leading	Solid
Novo Nordisk	oral amycretin	✓			✓		1	Yes	No	Leading^	Solid^
Novo Nordisk	monlunabant					✓	1	Yes	Yes	Leading^	Leading^
Eli Lilly	retatrutide	✓	✓	✓			2	No	No	Leading	Lagging
Eli Lilly	orforglipron	✓					2	Yes	Yes	Lagging	Lagging
Eli Lilly	mazdutide	✓		✓			3	No	No	Lagging	Solid
Challengers											
Viking	VK2735	✓	✓				2	Both*	No	Leading	Lagging
Boehringer Ingelheim/Zealand	survodutide	✓		✓			2	No	No	Lagging	Lagging
Structure Therapeutics	GSBR-1290	✓					2	Yes	Yes	Solid	Solid
Altimune	pemvidutide	✓		✓			2	No	No	Lagging	Lagging
Pfizer	danuglipron^^	✓					2b	Yes	Yes	Lagging	Lagging
Roche (Carmot)	CT-388	✓	✓				1	No	No	Leading	Solid^
Roche (Carmot)	CT-996	✓					1	Yes	Yes	Leading	Solid^
Amgen	maritide**	✓	✓				1	No	No	Leading	Lagging
Zealand	petrelintide				✓		1	No	No	Solid	Leading

Source: Morningstar, company reports. Phase 3 data from Step 1 (Wegovy), Surmount 1 (Zepbound), and Oasis 1 (Rybelsus). *Data for VK2735 is for injectable version (oral data only available for four weeks in phase 1). **maritide is a GIP antagonist, other GIP-targeting drugs are agonists. ^Minimal published data to support drug’s potential profile. ^^ Danuglipron data is for twice-daily dosing; new program is once-daily dosing.

Obesity Drug Candidates Can Differentiate on Efficacy, Convenience, Supply, and Tolerability

We see several ways that new firms can differentiate from incumbents Novo Nordisk and Eli Lilly. We expect efficacy (speed, duration, and quality of weight loss), gastrointestinal tolerability, injectable versus oral dosing, and availability of supply will all be key determinants of differentiation. There is no head-to-head data available for these drugs, although Lilly will have Zepbound and Wegovy head-to-head data later this year. Obesity trials enroll different populations and have different rules for how patients are introduced to effective dose levels, which affects both speed of efficacy and tolerability. This limits our ability to compare trial data, although we think there is sufficient data to sort drugs based on available data.

Some Drug Candidates Could Have Convenience and Supply Advantages

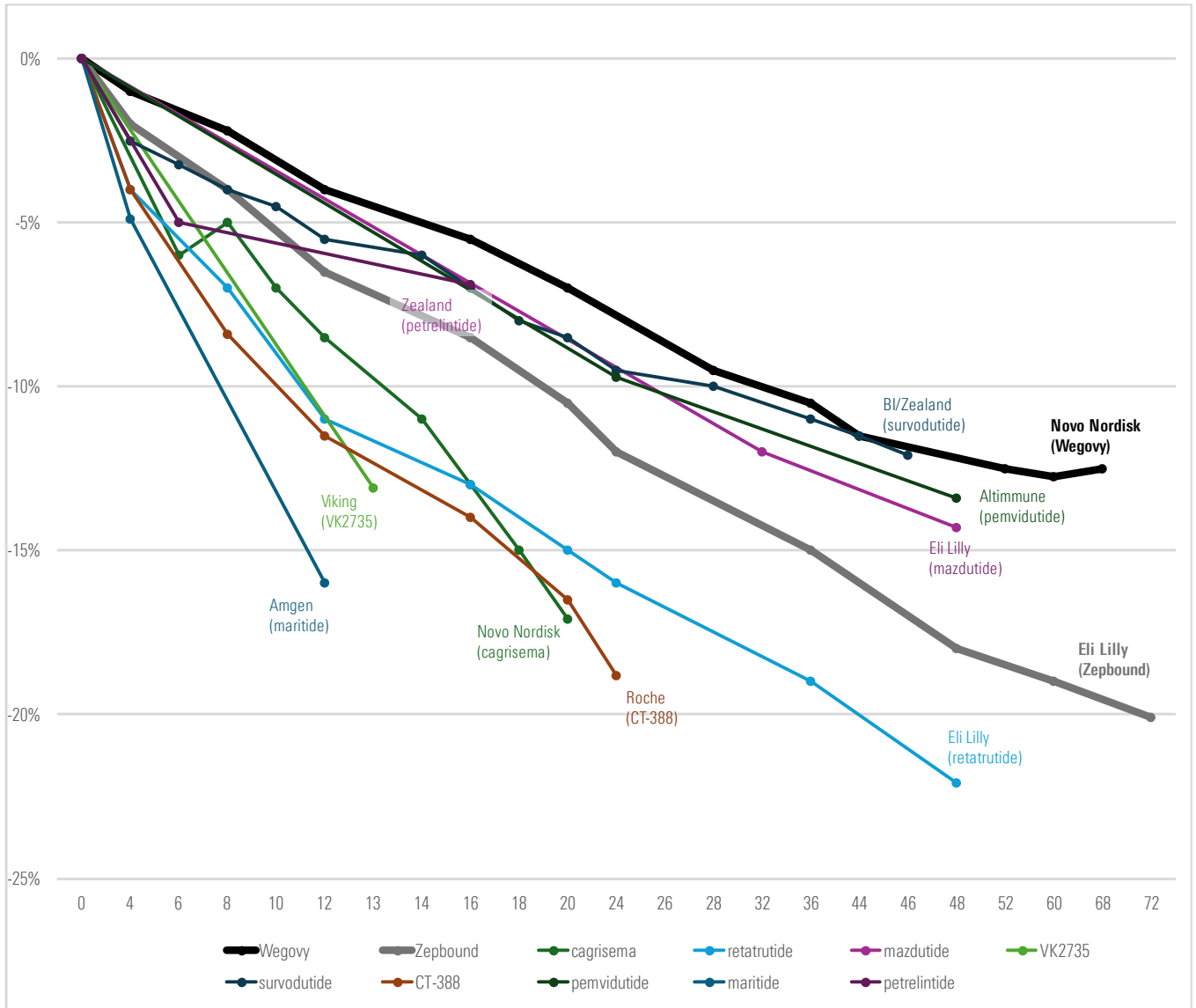
Novo Nordisk and Eli Lilly are struggling to provide enough supply of their peptide-based drugs, which require complex manufacturing and face limitations in global infrastructure for such drugs. We think oral versions of peptides, including high-dose Rybelsus and Viking's VK2735, could be even harder to supply, as dosages need to be even higher to reach therapeutic levels when digested (although the fill/finish process would be simplified). We think Amgen's injectable maritide could be slightly easier to scale, particularly if dosing can be extended to monthly or even quarterly (instead of weekly Wegovy and Zepbound injections). By far the easiest to scale would be oral small molecule drugs, such as Lilly's orforglipron, Novo Nordisk's monlunabant, Pfizer's danuglipron, Roche's CT-996, and Structure's GSB-1290. For example, Structure management noted in a June 2024 presentation that its current manufacturing capacity could supply more than 120 million patients, far surpassing what we estimate as the roughly 20 million patients (diabetes and obesity) who drove the \$36 billion in GLP-1 global sales in 2023.

Injectable Efficacy: Amgen, Viking, and Roche Could Challenge Incumbents on Efficacy

Several controlled trials and real-world data point to the conclusion that Lilly's Zepbound has stronger efficacy than Novo's Wegovy. That said, we think Wegovy's longer time on the market, head start with label expansions, and Novo's solid pipeline will continue to keep the playing field relatively level between these firms.

Looking beyond Wegovy and Zepbound at injectable drug candidates, we see a split in efficacy between those that look more similar to Wegovy and those that could even surpass Zepbound's efficacy. For example, GLP-1 and glucagon-targeting drug candidates survodutide (Boehringer Ingelheim/Zealand), pemvidutide (Altimmune), and mazdutide (Lilly and Innovent) all appear to have efficacy closer to Wegovy than Zepbound. With launches unlikely prior to 2026, we don't have high expectations for sales of these therapies, as entrenchment of Novo and Lilly will only increase over this time. Zealand's amylin drug petrelintide has shown encouraging early data and still has a chance at differentiation, particularly if tolerability stays strong. We see stronger potential for Novo's GLP-1/amylin cagrisema, Roche's GLP-1/GIP CT-388, and Lilly's triple agonist retatrutide, although tolerability issues and potential cardiovascular side effects could hold back retatrutide. Other GLP-1/GIP targeting drug candidates from Viking and Amgen have also generated promising data, although data released so far is shorter term.

Exhibit 10 Placebo-Adjusted Percentage Weight Loss Over Time: Wegovy and Zepbound Versus Injectable Obesity Drug Candidates (weeks)

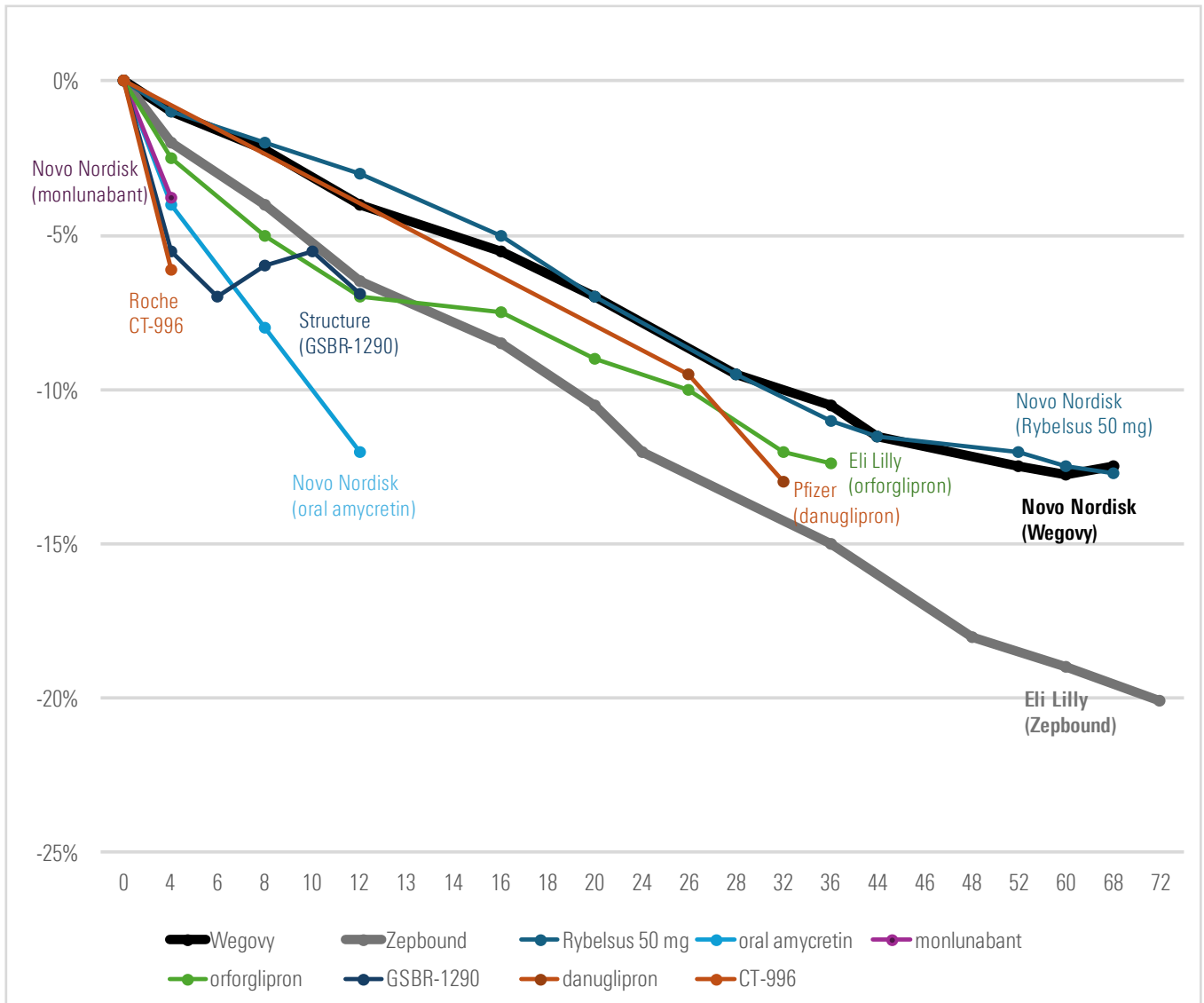


Source: Morningstar, company reports, Lancet, New England Journal of Medicine. Data varies slightly depending on whether data is reported based on intent to treat (treatment regimen estimand all randomized patients in trial) or per protocol (efficacy estimand, only those following treatment schedule). Novo Nordisk (Wegovy, cagrisema) and Viking (VK2735) efficacy data is intent to treat, so appears slightly less effective. Exact reporting methods vary and not all trials are published with disclosed protocols.

Oral Efficacy: Structure Could Compete With Novo's Next-Generation Drug Candidates

Among oral drug candidates, we think high-dose Rybelsus could be the first to launch, but supply issues could end up delaying the launch further. We think orforglipron stands a better chance at generating stronger efficacy than injectable Wegovy, given the long-term data generated in phase 2. In addition, if Pfizer's once-daily danuglipron can generate similar efficacy as the twice-daily program, its prospects look similar to orforglipron's. That said, Novo's CB1 monlunabant and a next-generation GLP-1/amylin drug (amycretin) as well as Structure's GSBR-1290 all have the potential to generate data that surpasses Zepbound, although data so far is relatively short term.

Exhibit 11 Placebo-Adjusted Percentage Weight Loss Over Time: Wegovy and Zepbound Versus Oral Obesity Drug Candidates (weeks)



Source: Morningstar, company reports, Lancet, New England Journal of Medicine. Data varies slightly depending on whether data is reported based on intent to treat (treatment regimen estimand, all randomized patients in trial) or per protocol (efficacy estimand, only those following treatment schedule). Novo Nordisk (Wegovy, Rybelsus) efficacy data is intent to treat, so appears slightly less effective. Exact reporting methods vary and not all trials are published with disclosed protocols. Danuglipron data is phase 2b study of discontinued twice-daily dose (ongoing as once-daily program).

Quality of Weight Loss: Fat Mass Loss Is Better Than Lean Mass Loss

Quality of weight loss is increasingly in focus, as 40% of weight loss for Wegovy patients in Step 1 was lean mass, and roughly 25% of weight loss for Zepbound patients in Surmount-1 was lean mass. Natural weight loss (with diet and exercise alone) is typically roughly 20% lean mass (when total weight loss is roughly 10%),² and we expect new targets and combinations could help reduce the loss of lean mass.

In fact, among the several strategies in obesity clinical development, a combination strategy with myostatin/activin-targeting therapies is being pursued by multiple firms (Exhibit 12).

Exhibit 12 Strategies to Build Differentiation Beyond Wegovy

Strategy	Rationale	Example Firms (Drug Candidate)
GLP-1 small molecule	Me-too efficacy but much simpler manufacturing, convenience of oral	Lilly (orforglipron), Structure (GSBR-1290) AstraZeneca (ECC5004), Terns (TERN-601) Pfizer (danuglipron), Roche (CT-996)
GIP combos	Incretin that adds to GLP-1 efficacy without increasing tolerability issues	Lilly (Zepbound), Viking (VK2735), Roche (CT-388), Amgen (maritide)
Lean Mass Preservation (myostatin/activin)	Could combine with GLP-1 therapies to reduce weight loss from muscle	<u>Public:</u> Lilly (bimagrumab), Roche (R07204239), Regeneron (trevogrumab, garetosmab), Keros Therapeutics (KER-065), Scholar Rock (apitegromab), Biohaven (taldefgrobep) <u>Private:</u> SixPeaks 35Pharma (HS-235)
Gene Therapy	Could remove need for chronic therapy	Fractyl Health (to clinical trial in 2025)
Me-Too GLP-1	Expand supply in various geographies	<u>Public:</u> Hanmi (Korea, efpeglenatide) Jiangsu HengRui (China, noiiglutide) <u>Private:</u> Sciwind (China, ecnoglutide)
Amylin Targeting	Another mechanism to increase satiety, either with GLP-1 or as monotherapy	Novo (cagrisema, amycretin) Zealand (petrelintide) Lilly (eloralintide) Structure (entering phase 1)
CB1	Peripheral CB1 targeting drugs aim to avoid depression side effect of Sanofi's rimonabant	Novo (monlunabant) Skye (nimacimab) Corbus (CRB-913-CB1) Agenix (AGTX-2004)
NLRP3	Blocking "inflammasome" activation in obese patients could lead to weight loss	<u>Public:</u> Ventyx (VT3232) <u>Private:</u> Nodthera (NT-0249,0976) Ventus (VENT-02)
Glucagon	Another incretin that could be used in combo, this time increasing energy expenditure	Lilly (retatrutide, mazdutide) Altimmune (pemvidutide) BI/Zealand (survodutide)

Source: Morningstar, company reports. Maritide is a GIP antagonist, while other GIP-target drugs are agonists.

² Turicchi, Jake et al. Associations between the rate, amount, and composition of weight loss as predictors of spontaneous weight regain in adults achieving clinically significant weight loss: A systematic review and meta-regression. *Obesity Reviews*, 20 (7). pp. 935-946. ISSN 1467-7881.

In addition, Altimmune's GLP-1/glucagon-targeting therapy pemvidutide yielded 22% lean mass loss in phase 2, showing a potential slight improvement to standard of care. Pipeline candidates that avoid GLP-1 altogether, such as Zealand's amylin drug petrelintide, could also see less lean mass loss, although this lean mass data is preclinical and the drug is just entering phase 2.

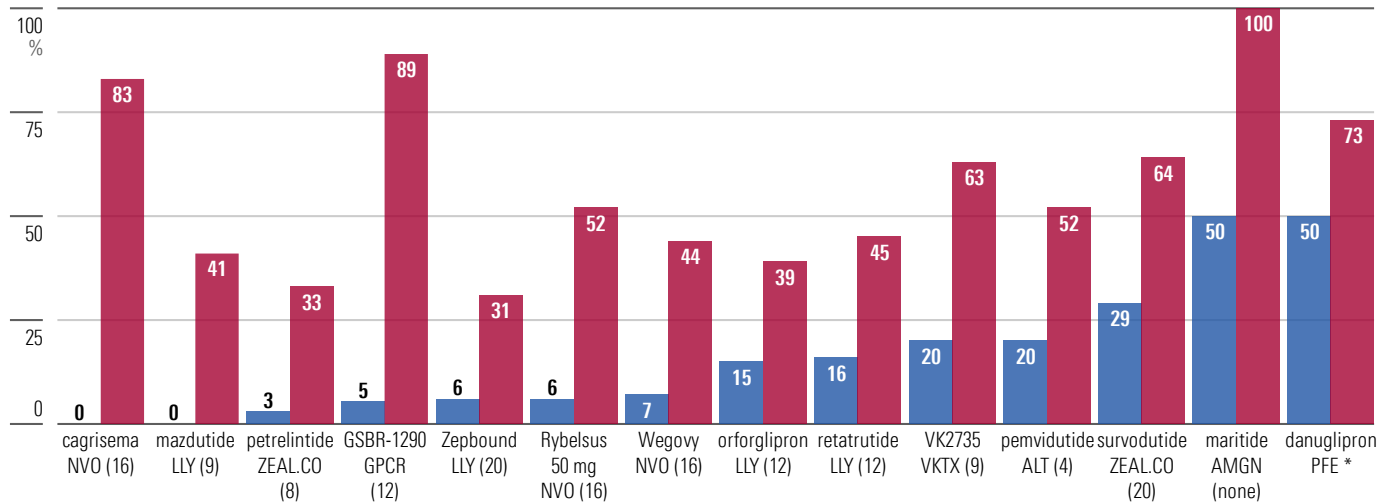
Tolerability Looks Promising for Several New Drugs; Better Dosing Could Improve Data

Aside from efficacy, convenience, and supply, we think tolerability remains a key potential differentiator for new obesity medicines. Exhibit 13 sorts data on one measure of tolerability: discontinuations due to adverse events. A longer dose escalation period (in weeks in parentheses) should result in better tolerability. We think phase 1 data from Zealand's amylin targeting petrelintide, phase 2 data for Structure's oral GLP-1 GSBP-1290, and phase 1b data for Novo Nordisk's GLP-1/amylin injectable cagrisema bode well for solid or perhaps leading tolerability, if phase 3 data also supports this profile. We also think there is room for improved tolerability for several pipeline candidates, particularly Lilly's oral GLP-1 orforglipron and Amgen's maritide, with slower dose escalation to avoid nausea (maritide patients at the highest dose in the phase 1 study were not titrated at all). We have excluded Roche's CT-388 and Novo's monlunabant and oral amycretin from this analysis, as we don't have sufficient details to compare their tolerability.

Exhibit 13 Obesity Drugs Sorted by Adverse Event-Related Discontinuation Rates at Key Efficacy Dose

Dose escalation in weeks in parentheses

■ Adverse Event-Related Discontinuations ■ Nausea



Source: Morningstar, company reports, Lancet, New England Journal of Medicine. NVO: Novo Nordisk. LLY: Eli Lilly. GPCR: Structure Therapeutics. VKTX: Viking Therapeutics. ALT: Altimmune. ZEAL.CO: Zealand Pharma. AMGN: Amgen. PFE: Pfizer. *weekly, biweekly, and monthly titration at various doses, discontinuation rate at least 50% in all arms.

Upcoming Catalysts Will Mean a Dynamic Market for Obesity Firms

Aiming to differentiate on all these measures, several firms will be reporting key data in the near term that could position them to compete with incumbents Lilly and Novo, as shown in Exhibit 14. Novo and Lilly also expect next-generation data for their obesity pipelines.

Exhibit 14 Upcoming Catalysts for Incumbents and Potential New, Differentiated Obesity Drugs

Firm	Drug	Status	Next Data	Notes
Incumbents: Approved				
Eli Lilly	Zepbound	Approved	H2H v Wegovy (2024) High Dose (Q4 2024) MASH (2024)	Zepbound has strong data in sleep apnea and is likely to prove more effective than Wegovy in the head-to-head study. Higher dose could improve Wegovy's efficacy, but at risk of more tolerability issues.
Novo Nordisk	Wegovy	Approved	Alzheimer's (2025)	
Incumbents: Next Generation				
Eli Lilly	retatrutide	Phase 3	2026	Lilly is aiming for leading efficacy and includes patients with sleep apnea and knee osteoarthritis.
Eli Lilly	orforglipron	Phase 3	2025	Lilly is aiming for similar efficacy to Zepbound, oral convenience.
Eli Lilly	bimagrumab	Phase 2	H2 2024	Data for this activin type II blocker (activin/myostatin) with semaglutide could lead to Zepbound combo to reduce muscle loss.
Eli Lilly	eloralintide	Phase 2	2025	Lilly has moved its own amylin targeting molecule to phase 2, perhaps as an option to defend against cagrisema competition.
Eli Lilly/Innovent	mazdutide	Phase 3	Higher 9mg dose (2025)	Innovent's phase 3 has a high bar to surpass Zepbound and advance in Lilly pipeline. Oral semaglutide requires more manufacturing capacity, could lead to further delays.
Novo Nordisk	Rybelsus 50 mg	Filing	EU Launch (2025)	
Novo Nordisk	cagrisema	Phase 3	First Ph 3 studies (Q4 2024) H2H v Zepbound (2025)	Aiming for leading efficacy with potential for better tolerability and less lean mass loss than Wegovy.
Novo Nordisk	oral amycretin	Phase 1	Subcu (2025)	Despite solid early oral data, Novo could move forward with subcu to conserve supply.
Novo Nordisk	monlunabant	Phase 2	Q3 2024	Could have better tolerability and less muscle loss than GLP-1, and peripherical action could prevent psychiatric side effects.
Challengers				
Altimune	pemvidutide	Phase 3	Trial start (2024)	Encouraging phase 2 muscle loss and CV data could differentiate, and better titration in phase 3 could improve tolerability.
Amgen	maritide	Phase 3	Phase 2 data (2024)	Amgen is entering phase 3 after an encouraging look at phase 2 data, although details are still coming.
AstraZeneca	AZD5004	Phase 2b	Trial start (H2 2024)	Phase 1 data should be revealed as this oral GLP-1 moves to mid-stage development. Could focus on lower BMI (overweight).
AstraZeneca	AZD9550 AZD6234	Phase 1	Phase 1 data (2024)	GLP-1/glucagon and amylin programs round out Astra's obesity pipeline and add potential for combination regimens.
Boehringer Ingelheim/Zealand	survodutide	Phase 3	Phase 3 data (Late 2025)	Late 2025 Phase 3 data at higher dose with longer dose titration to minimize side effects
Pfizer	danuglipron	Phase 1	Dose optimization data (Q1 2025)	Once daily optimization is taking place ahead of pivotal trials for this GLP-1, which saw tolerability setbacks at twice daily dose.
Roche	CT-388	Phase 2	T2D Phase 1b data (2024)	Roche is testing multiple doses/titration in a new phase 2 study to differentiate from Zepbound, hoping to start phase 3 in 2025.
Roche	CT-996	Phase 2	Phase 2 start (2025)	After positive 4-week data in phase 1, additional phase 1 arms are starting this year and phase 2 likely next year.
Structure Therapeutics	GSBR-1290	Phase 2b	Tablet data (late 2025)	Structure is aiming for differentiated tolerability, oral convenience.
Terns	TERN-601	Phase 1	2H 2024	Inspired by the structure of Pfizer's danuglipron, Terns is developing a once-daily oral.
Viking	VK2735	Phase 2	Higher dose oral data and phase 2 start (Q4 2024) Phase 3 injectable start (2025)	Viking is moving its injectable to phase 3 next year, possibly including a monthly dose option, but peptide structure is a manufacturing challenge, especially for oral version.
Zealand	petrelintide	Phase 2b	Trial start (2024)	Phase 1 data showed encouraging tolerability data and early efficacy data similar to Wegovy and Zepbound.
Zealand	dapiglutide	Phase 2b	Higher dose data (2024/2025) Trial start (H1 2025)	Higher doses, longer-term data could support this GLP-1/GLP-2.

Public Markets: Large Biopharma Firms Interested in Buying Into Obesity Market

While cardiometabolic drug development was a key area of focus for the biopharma industry up until the early 2010s, most biopharma firms (except Novo and Eli Lilly) significantly reduced research and development efforts in the therapeutic area as pricing power declined and innovation stalled. The industry heavily pivoted more toward oncology, immunology, and rare disease drugs, where pricing power remained strong and innovative scientific advancements were striving.

Given the huge obesity market opportunity combined with an increased understanding of treatment, we expect large biopharma firms to accelerate development efforts into obesity drugs through acquisitions. On the public side, close to \$5 billion in upfront payments for obesity drug-related acquisitions have occurred since 2023 (Exhibit 15). This is a relatively small investment into acquisitions for one of the most innovative therapeutic areas, as the industry's largest (>\$20 billion) annual acquisitions alone typically total over \$50 billion.³ In addition, obesity-based collaborations in 2023 between large public firms and smaller private innovators, like AstraZeneca/Eccogene (focused on oral GLP-1 ECC5004) and Novo Nordisk/EraCal (focused on Era-379), show strong appetite among big biopharma firms for access to novel obesity drug candidates.

Exhibit 15 Top Obesity Drug-Related VC Exits, by Exit Size, 2020-24

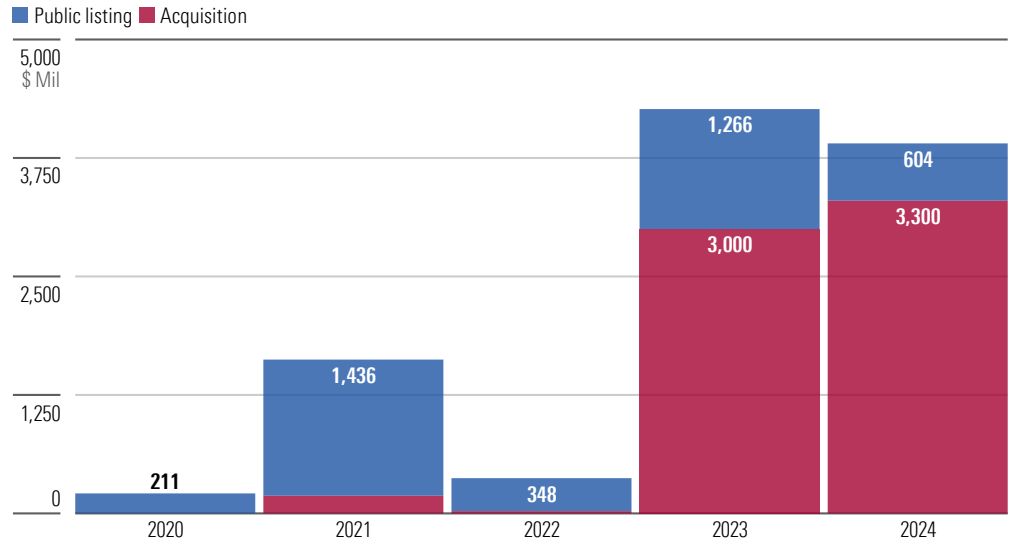
Company name	Exit type	Acquiring Firm	Close date	Exit size (\$M)
Carmot Therapeutics	Merger/Acquisition	Roche	January 26, 2024	\$3,300
Versanis Bio	Merger/Acquisition	Eli Lilly	August 14, 2023	\$1,925
Inversago Pharma	Merger/Acquisition	Novo Nordisk	September 8, 2023	\$1,075
Cutia Therapeutics	IPO	N/A	June 9, 2023	\$789
Epitomee Medical	IPO	N/A	December 15, 2021	\$632
Fractyl Health	IPO	N/A	February 2, 2024	\$604
VectivBio	IPO	N/A	April 9, 2021	\$451
Structure Therapeutics	IPO	N/A	February 4, 2023	\$414
Terns Pharmaceuticals	IPO	N/A	February 5, 2021	\$280
Keros Therapeutics	IPO	N/A	April 8, 2020	\$211
Lobesity	Merger/Acquisition	9 Meters	July 19, 2021	\$113
Doer Biologics	Merger/Acquisition	Huadong Medicine	April 27, 2021	\$75

Source: Morningstar, PitchBook. Data as of July 17, 2024.

On the private side, exits are increasingly focused on acquisitions, rather than IPOs (Exhibit 16), although BioAge Labs filed for an IPO in early September.

³ Morningstar Biopharma Industry Landscape, April 2024, slide 20.

Exhibit 16 Obesity Drug Development VC Exit Value (in \$ millions) by Exit Type

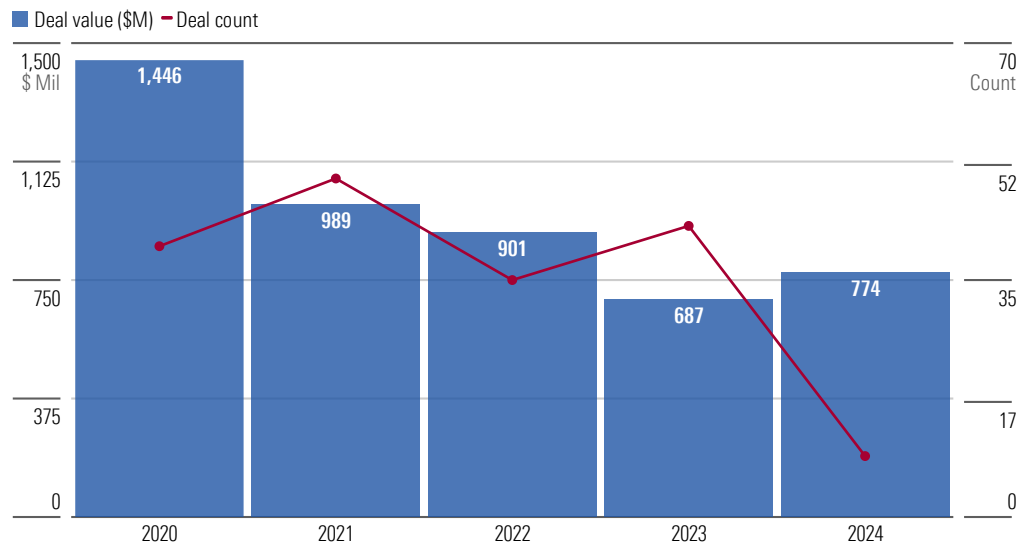


Source: PitchBook, Data as of July 17, 2024.

Private Data Trends: Emerging Startups in the Obesity Treatment Landscape

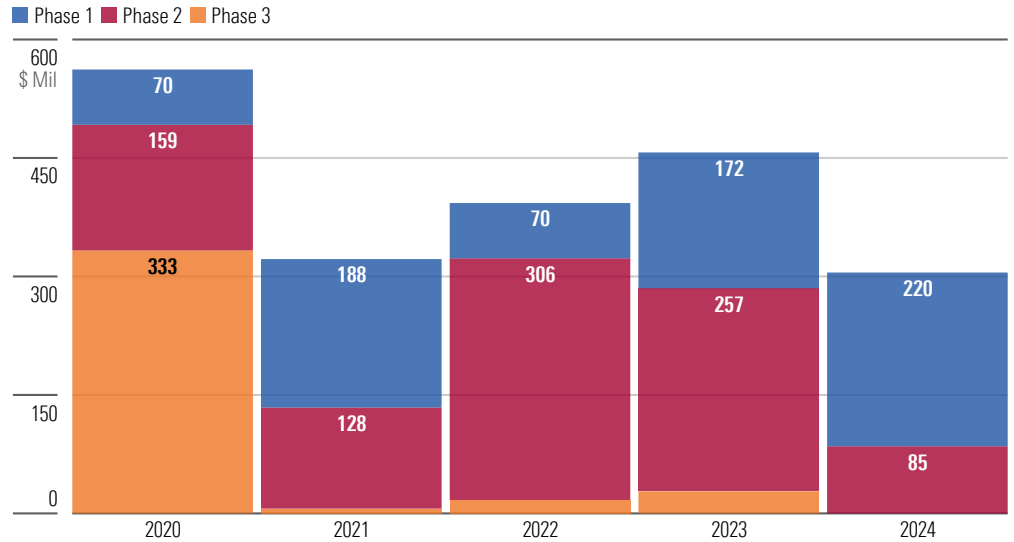
The obesity treatment sector is experiencing a surge of innovative startups, each bringing unique approaches to tackle this global health crisis. These emerging companies are attracting significant investment and exploring novel mechanisms to compete with established players like Novo Nordisk and Eli Lilly.

Exhibit 17 Obesity Drug Development: VC Deal Activity, 2020-24



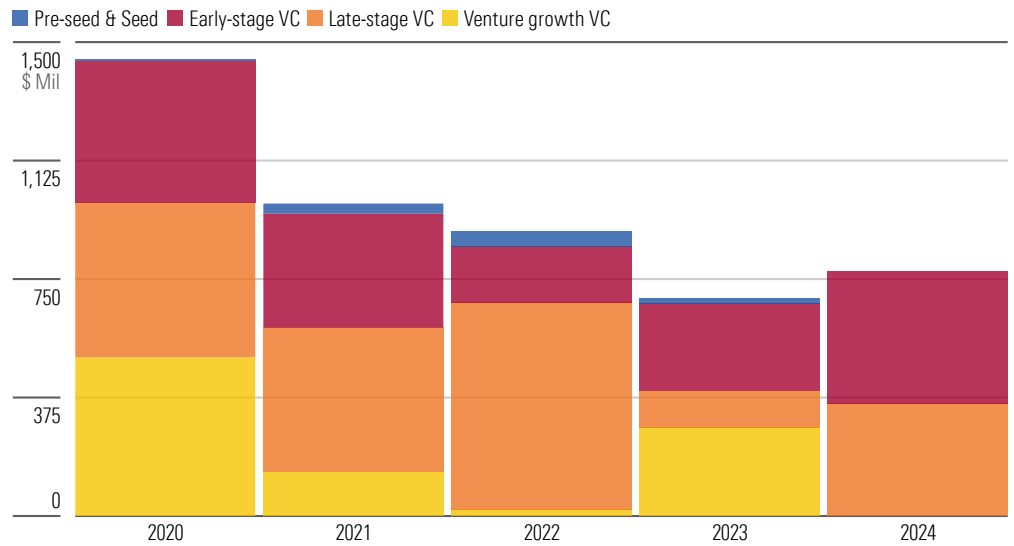
Source: PitchBook, Data as of July 17, 2024.

Exhibit 18 Obesity Drug Development VC Deal Size (in \$ millions) by Highest Phase



Source: PitchBook, Data as of July 17, 2024. * Determined as the highest phase of a trial that started prior to the round closing. * Combines in-between phases with the highest phase.

Exhibit 19 Obesity Drug Development VC Deal Value (in \$ millions) by stage



Source: PitchBook, Data as of July 17, 2024.

Metsera, backed by \$290 million from Arch Venture Partners and Population Health Partners, is developing a portfolio of oral and injectable weight loss drugs targeting multiple gut hormones. Similarly, Hercules CM NewCo, a \$400 million venture supported by Bain Capital, RTW Investments, and Atlas Venture, has licensed a portfolio of incretin drugs from Chinese pharmaceutical company Jiangsu Hengrui. SixPeaks Bio, incubated by Versant Ventures with \$30 million in Series A funding, is partnering with AstraZeneca to develop new approaches to obesity treatment. Additionally, Deep Apple Therapeutics focuses on small-molecule drugs for metabolic disorders, leveraging structural and computational biology to target G-protein coupled receptors (GPCRs) involved in metabolism.

OrsoBio, with a total of \$97 million in funding, is advancing a diverse portfolio including mitochondrial protonophores, ACC2 inhibitors, and LXR inverse agonists for treating obesity and associated metabolic disorders.

CinRx Pharma has raised \$73 million to support obesity drug development through a hub-and-spoke model. One of its ventures, CinFina, is developing four obesity drugs licensed from Johnson & Johnson, emphasizing muscle preservation alongside weight loss. CinRx continues to explore new obesity therapies and create additional biotech ventures. ProFound, in partnership with Pfizer, is exploring next-generation obesity assets using proteomics to identify new drug targets. This collaboration is part of a \$50 million initiative with Flagship Pioneering, aiming to expand Pfizer's obesity drug pipeline with innovative treatments.

Many other private investments in the space are internationally based, focusing on "me-too" drugs primarily in China, Korea, and Japan. These companies are potential merger and acquisition targets for Big Pharma or startups attempting to license them in a roll-up strategy similar to Metsera or Hercules CM. Notable examples include Sciwind Biosciences, Gmax Bio, Regor Therapeutics, and Sohia Pharma, all of which are based on GLP-1 biology.

Emerging trends in the field include a focus on muscle preservation, oral alternatives to injectables, and innovative drug delivery systems. Companies like CinFina and BioAge Labs are developing molecules to help patients maintain muscle mass. Syntis Bio is developing SYNT-101, an oral daily pill that mimics the effects of gastric bypass, while Rivus Pharmaceuticals works on controlled metabolic accelerators as once-daily oral doses. i2o Therapeutics, with \$46 million in Series A funding, is developing a drug-device combination licensed from Intarcia, including an implantable device called "Medici" that can deliver GLP-1 medication for up to one year. Companies are also exploring multimodal approaches to address not only weight loss but also related complications such as diabetes and cardiovascular diseases, with Rivus Pharmaceuticals aiming to treat cardiovascular conditions rather than targeting the weight loss market directly.

Lastly, the IPO market is heating up for obesity-related startups. BioAge Labs is raising \$100 million, potentially serving as a bellwether for the sector and catalyzing further investment. Meanwhile, MBX Biosciences has filed for a \$100 million IPO. While its lead asset targets hypoparathyroidism, MBX also has a preclinical long-acting GLP-1/GIP receptor co-agonist peptide, MBX 4291, aimed at improving obesity treatment dosage and user-friendliness. These IPOs highlight continued investor interest in metabolic disease therapies and could pave the way for more startups to access public markets.

Exhibit 20 Top VC Deals in Obesity Since 2020 (>\$75 Million)

Company	Close date	Deal size (\$M)	Deal type	Stage	Lead investor(s)
Hercules CM	May 16, 2024	\$400	Early Stage VC		
Metsera	April 3, 2024	\$350	Early Stage VC	Series A	ARCH Venture Partners, Population Health Partners
Kallyope	January 27, 2022	\$236	Later Stage VC	Series D	Mubadala Investment Company, The Column Group
BioAge Labs	February 1, 2024	\$194	Later Stage VC	Series D	Sofinnova Investments
Ventus Therapeutics	February 9, 2022	\$140	Later Stage VC	Series C	RA Capital Management, SoftBank Investment Advisers
Rivus Pharmaceuticals	August 19, 2022	\$132	Early Stage VC	Series B	RA Capital Management
Kallyope	March 25, 2020	\$112	Later Stage VC	Series C	
Ventus Therapeutics	April 8, 2021	\$100	Early Stage VC	Series B	RA Capital Management
BioAge Labs	December 3, 2020	\$90	Later Stage VC	Series C	Andreessen Horowitz, Elad Gil
Regor Therapeutics	February 18, 2021	\$90	Early Stage VC	Series B	Lilly Asia Ventures
Aardvark Therapeutics	May 9, 2024	\$85	Later Stage VC	Series C	Decheng Capital
OrsoBio	November 7, 2023	\$85	Early Stage VC	Series A	Enavate Sciences, Longitude Capital
Neurogastrx	July 12, 2021	\$80	Later Stage VC	Series B	
SixPeaks Bio	August 23, 2024	\$80	Early Stage VC	Series A	Versant Ventures
GMAX Bio	March 5, 2021	\$78	Later Stage VC	Series C	China Merchants Group
ProFound Therapeutics	May 26, 2022	\$75	Early Stage VC		

Source: PitchBook, Data as of Sept. 1, 2024.

Most Likely to Acquire: Incumbents Novo and Lilly, Also New Obesity Players J&J and Merck

To catch up to the underinvestment in obesity drug development, we expect several big biopharma firms to make acquisitions over the next 18 months, targeting smaller focused obesity drug development firms. In Exhibit 21, we rank the most likely large cap buyers of obesity drug development firms. We rank the companies on strategic fit and balance sheet strength to enable acquisitions.

Exhibit 21 Ranking of Big Biopharma Firms Most Likely to Make Obesity Drug Acquisitions

Ranking	Company	Strategic Fit	Balance Sheet Strength
1	Merck	High: Solid entrenchment in cardiometabolic but largely missed the GLP-1 innovation, which could drive acquisitions, especially with the approaching Keytruda 2028 patent loss.	High: With only limited recent acquisitions, Merck is poised to make a major deal.
2	Novo	High: A top leader in cardiometabolic and likely always looking to augment internal efforts.	High: Recent Catalent deal will reduce cash but major windfalls from obesity drug sales will replenish cash quickly.
3	Johnson & Johnson	Medium/High: Recent cardiometabolic setbacks (Xarelto, Invokana, apicitentan) could trigger the need for external innovation.	High: Very strong balance sheet (even after the Shockwave acquisition) enable solid potential acquisitions.
4	Eli Lilly	High: Potentially the best positioned firm in cardiometabolic and willingness to buy additional related assets (such as Versanis) sets up future deals.	Medium/High: Very robust cashflows from obesity drugs and limited patent losses could enable major acquisitions.
5	AstraZeneca	Medium/High: Licensing of ECC5004 (GLP-1) and strong position of cardiometabolic drug Farxiga with future combinations shows increasing interest.	Medium/High: Despite Alexion acquisition, the robust cashflows set up solid potential.
6	Boehringer Ingelheim	High: With a major focus in cardiometabolic drugs and with a lead drug (suvodutide) that could have side effect issues, the appetite for acquisitions is likely high.	Medium: Limited recent deals leaves the firm optional
7	Roche	Medium/High: Recent acquisition of Carmot shows interest in GLP-1 drugs and limited pipeline suggests potential for more deals.	Medium/High: Limited recent deals leaves the firm optionality.
8	Daiichi Sankyo	Medium: While historically a area of focus for the company, cardiometabolic drugs seem less important than cancer drugs.	High: Recent antibody drug conjugate deals have increased cashflows opening the potential for more acquisitions.
9	Sanofi	Medium: The firm has pivoted to immunology, but the historical entrenchment in cardiometabolic disease could entice the firm to make an acquisition.	High: Limited recent deals leaves the firm optionality.
10	Novartis	Medium: While holding a solid portfolio of cardiometabolic drugs, the focus on cancer, immunology and rare disease makes future obesity deals less likely.	High: Only moderate recent deal flow opens the potential for more acquisitions.
11	Chugai	Medium: Royalty on Lilly's orforglipron could entice the firm to gain greater exposure to cardiometabolic drugs.	Medium/High: Could likely finance a mid-sized acquisition, but aversion to debt may limit larger deals.
12	Regeneron	Low/Medium: PCSK9 drug Praluent has been disappointing in cardiometabolic and focus is more on eye-care, immunology and cancer.	Medium/High: Potential to make acquisitions.
13	Amgen	Medium/High: The recent clinical advancement of MariTide in obesity opens a newer therapeutic area that could use more follow up drugs.	Low: The major acquisition of Horizon leaves the cash account more depleted.
14	GSK	Low: Largely moved away from cardiometabolic over the last decade.	Medium/High: Cash from Haleon divestment, lower dividend and only moderate sized recent acquisition allow for more deal flow.
15	Pfizer	Medium/High: GLP-1 drug daniglupron's delay likely elevates external deal options.	Low: Major acquisition of Seagen likely limits ability to make major deals.
16	Bayer	Medium/High: A solid entrenchment in cardiometabolic disease would be a natural fit for obesity drug development.	Low: Glyphosate litigation hampers ability to make large acquisitions.
17	Gilead	Low/Medium: Focused largely on virology and cancer, although liver disease research did bring a GLP-1 to preclinical development.	Medium: The large acquisition of Immunomedics and recent tuck in acquisitions make large near-term deals less likely.
18	Bristol Myers	Low/Medium: While Eliquis is a leader in atrial fibrillation, the firm has largely exited the cardiometabolic landscape.	Low/Medium: Despite the major acquisition of Celgene and recent mid-sized deals for Karuna and RayzeBio, the strong cashflows would enable acquisitions.
19	AbbVie	Low/Medium: While not a major competitor in cardiometabolic drugs, the strong entrenchment in the aesthetics industry could make an obesity drug a good match.	Low/Medium: Cash availability is lower following the large acquisition of Allergan and recent mid-sized acquisitions of ImmunoGen and Cerevel.
20	Takeda	Low/Medium: With a heavy immunology focus, obesity drug interest seems low.	Low: The major acquisition of Shire likely limits big deals.
21	Biogen	Low: Focused on neurology, oncology and rare diseases, not cardiometabolic.	Low: Recent Reata acquisitions limits major deal potential.

Source: Morningstar, company reports.

Acquisition Targets for Obesity Drugs Span Public and Private Markets

On the public side, we have seen several firms with strong early-stage obesity drug data. We expect these firms are ready to sell to a larger firm as late-stage development, manufacturing, and marketing will likely be too costly for smaller firms to manage independently. In Exhibit 22, we rank the most likely acquisition targets for obesity drugs.

Exhibit 22 Potential Acquisition Targets in Obesity: The Public Market

Rank	Target	Key Drug/Product(s)	Current Valuation (\$B)	Rationale	Challenge
1	Structure	GSBR-1290 (Daily Oral GLP-1) Phase 2 data reported	2.1	Phase 2 efficacy data looks similar to Lilly's oral orforglipton with potentially a lower discontinuation rate. Non-peptide structure could make manufacturing much easier than Rybelsus.	Likely at least a year behind orforglipton.
2	Viking	VK2735 (Weekly Injection GLP-1/GIP) Phase 3 ready VK2735 (Daily Oral Version GLP-1/GIP) Phase 2 data reported	5.8	Phase 2 injectable data looks competitive with Zepbound/Wegovy. An oral formulation is testing higher doses to see if efficacy can also be improved.	20% discontinuation rate is concerning. The oral version also looks less effective and is peptide based, making manufacturing more challenging.
3	Altimune	Pemvidutide (Weekly Injection GLP-1/Glucagon) Phase 3 Ready	0.4	Long-term weight loss data look slightly better than Wegovy's.	20% discontinuation rate could be a limitation, but allowing for dose declines as patients ramp in phase 3 should help.
4	Zealand	Survodutide (Weekly Injection GLP-1/Glucagon) Phase 3 with partner Boehringer Dapiglutide (GLP-1/GLP-2) and Petrelintide (amylin) in phase 2	8.5	High-single digit/low-double digit percentage royalty on global sales of survodutide, among first challengers in obesity market. Unique mechanism of action for mid-stage pipeline with GLP-1/GLP-2 and amylin targeting drugs.	Only get royalties on survodutide, which had a high dropout rate of 29%. Dapiglutide dosage still being raised to achieve competitive efficacy.
5	Fractyl Health	Revita and Rejuva	0.1	Offers the potential for long-term maintenance.	Unique treatment could have side effects.

Source: Morningstar, company reports. Valuation data as of Aug. 5, 2024.

On the private side, names like NodThera, Corteria, and Diasome stand out as having a greater than 50% probability of being acquired, according to PitchBook data (Exhibit 23). ■■

Exhibit 23 Top VC- Backed Obesity Companies, by Total VC Raised to Date (in \$ millions)

Company Name	VC (\$M) raised to date	IPO probability	M&A probability	No Exit probability
Human Longevity	\$1,087.8	93%	5%	2%
Kallyope	\$479.0	95%	3%	2%
Hercules CM	\$400.0	N/A	N/A	N/A
Metsera	\$350.0	N/A	N/A	N/A
BioAge Labs	\$318.2	95%	3%	2%
Ventus Therapeutics	\$300.0	95%	3%	2%
CinRx Pharma	\$176.0	32%	18%	50%
Rivus Pharmaceuticals	\$167.0	93%	3%	4%
OrsoBio	\$139.6	76%	13%	11%
Sciwind Biosciences	\$138.2	N/A	N/A	N/A
Aardvark Therapeutics	\$129.0	69%	22%	9%
Neurogastrx	\$127.8	3%	21%	76%
Raynovent Biotech	\$115.7	N/A	N/A	N/A
NodThera	\$102.2	26%	60%	14%
GMAX Bio	\$94.3	N/A	N/A	N/A
Regor Therapeutics	\$90.0	N/A	N/A	N/A
Diasome	\$89.7	15%	78%	7%
Corteria Pharmaceuticals	\$88.4	42%	51%	7%
ProFound Therapeutics	\$87.7	15%	57%	28%
SixPeaks Bio	\$80.0	N/A	N/A	N/A

Source: PitchBook, Data as of Sept. 1, 2024.

Appendix

Exhibit 24 Obesity Development Pipeline—Private Firms

Firm	Drug	Mechanism	Administration	Small Molecule?	Phase
35Pharma	HS235	Activin/GDF ligand trap	injectable	no	preclinical
Aardvark Therapeutics	ARD-101	bitter taste receptor agonist	oral	yes	2
Antag therapeutics	-	GIP antagonist	injectable	no	preclinical
Aphaia Pharma	APH-012	nutrient sensing cell stimulator	oral	yes	2
Beijing QL Biopharmaceutical	ZT002	GLP-1 agonist	injectable	no	1c
BioAge	BGE-100	NLRP3 inhibitor	oral	yes	preclinical
BioAge	BGE-105 (azelaprag)	G-protein coupled apelin receptor agonist	oral	yes	2
Biorestorative Therapies	ThermoStem	brown fat derived stem cells	injectable	no	preclinical
Boehringer Ingelheim	BI1820237	NPY2R	injectable	no	1
Boehringer Ingelheim/Gubra	BI3034701	undefined "triple agonist"	injectable	no	1
Boehringer Ingelheim/Zeland	survodutide/BI456906	GLP-1/glucagon agonist	injectable	no	3
CinFina	CIN-109	GDF15 analog	injectable	no	1
CinFina	CIN-110	PYY analog	injectable	no	1
CinFina	CIN-209	GLP-1/GDF15	injectable	no	preclinical
CinFina	CIN-210	GLP-1/PYY	injectable	no	preclinical
Corteria	COR-1389	CRF2 agonist	injectable	no	1
Cyta	CYTA-001	thyroid hormone receptor beta agonist	undisclosed	no	preclinical
Cytoki	CK-0045	IL-22 analog	injectable	no	1
Diasome	HDV incretins	hepatocyte-directed vesicles	injectable	no	preclinical
Enterin	ENT-03	PTP1B inhibitor	injectable	no	1
ERX Pharmaceuticals	ERX-1000	leptin sensitizer	oral	yes	1
Gila Therapeutics	GT-001	PYY analog	oral	no	1
Glaceum	HSG4112	glabridin analog	oral	yes	2
Glyscend Therapeutics	GLY-200	polymer GI coating	oral	no	2
Gmax	GMA102	GLP-1 agonist	injectable	no	3
Gmax	GMA105 (glutazumab)	GLP-1 agonist	injectable	no	2
Gmax	GMA106	GLP-1/GIP agonist	injectable	no	1
Hercules (via Jiangsu HengRui)	HRS-7535	GLP-1 agonist	oral	yes	1/2
Hercules (via Jiangsu HengRui)	HRS-4729	next-gen incretin agonist	not disclosed	undisclosed	preclinical
Hercules (via Jiangsu HengRui)	HRS-9531	GLP-1/GIP agonist	injectable	no	2
Kallyope	K-757	nutrient receptor agonist	oral	yes	2
Kallyope	K-833	nutrient receptor agonist	oral	yes	2
Metsera (via D&D)	-	GLP-1	injectable	no	1
Metsera (via D&D)	DD03	GLP-1/GIP/GCGR	oral	no	preclinical
Metsera (via D&D)	DD07	amylin agonist	oral	no	preclinical
Metsera (via D&D)	DD02S	GLP-1 agonist	oral	no	preclinical
Metsera (via D&D)	DD14	GLP-1/GIP agonist	oral	no	preclinical
Metsera (via D&D)	DD15	GLP-1/GIP/glucagon	injectable	no	preclinical
MBX Biosciences	MBX 4291	GLP-1/GIP	injectable	no	preclinical
Mindrak AI	MDR-001	GLP-1 agonist	oral	yes	1
NodThera	NT-0249, NT-0976	NLRP3 inhibitor	oral	yes	2
Olatec	OLT1177	NLRP3 inhibitor	oral	yes	2
OrsoBio	TLC-1235	liver targeted protonophore	oral	yes	preclinical
OrsoBio	TLC-6740	liver targeted protonophore	oral	yes	1
Pep2Tango	PTT-A	GLP-1/GIP/amylin/calcitonin	injectable	no	preclinical
QL Biopharmaceutical	ZT003	GLP-1/FGF21 agonist	injectable	no	preclinical

Source: Morningstar, PitchBook, company reports.

Exhibit 24 (continued) Obesity Development Pipeline — Private Firms

Firm	Drug	Mechanism	Administration	Small Molecule?	Phase
Regor Therapeutics	RGT-075	GLP-1 agonist	oral	yes	2
Regor Therapeutics	RGT-028	GLP-1 agonist	oral	yes	preclinical
Regor Therapeutics	RGT-274	GLP-1 agonist	oral	yes	preclinical
Resalis	RES-010	miR-22 inhibitor	injectable	no	1
Resalis	RES-020	miR-22 inhibitor	injectable	no	preclinical
Rivus	HU6	controlled metabolic accelerator	oral	yes	2
Sciwind	XW003 (ecnoglutide)	GLP-1 agonist	injectable	no	3
Sciwind	XW014	GLP-1 agonist	oral	yes	1
Sciwind	XW004	GLP-1 agonist	oral	no	1
Sciwind	XW017	GIP agonist	injectable	no	preclinical
Scohia Pharma	SCO-267	GPR40	oral	yes	1
Scohia Pharma	SCO-094	GLP-1/GIP agonist	undisclosed	undisclosed	1
SixPeaks	dual specific antibody	activin type IIA/IIB receptors	injectable	no	preclinical
SixPeaks	dual specific antibody	GLP-1 with activin	injectable	no	preclinical
Supercede	undisclosed	ACTR2	oral	yes	preclinical
Syntis Bio	SYNT-101	polydopamine coating	oral	no	1
Ventus	VENT-02	NLRP3 inhibitor	oral	yes	1
Zhejiang Doer (Doer Biologics)	DR10624	GLP-1/glucagon/FGF21R	injectable	no	1

Source: Morningstar, PitchBook, company reports.

Exhibit 25 Obesity Development Pipeline — Public Firms**Public Firms**

Firm	Drug	Mechanism	Administration	Small Molecule?	Phase
Agentix	AGTX-2004	CB1R antagonist	oral	yes	preclinical
Alnylam	ALN-INHBE	INHBE inhibitor	injectable	no	preclinical
Alnylam	ALN-Gene C	undisclosed	injectable	no	preclinical
Alnylam	ALN-Gene D	undisclosed	injectable	no	preclinical
Altimmune	pemvidutide/ALT-801	GLP-1/glucagon agonist	injectable	no	3
Amgen	AMG133/MariTide	GLP-1 agonist/GIP inhibitor	injectable	no	2
Arrowhead	ARO-INHBE	INHBE inhibitor	injectable	no	preclinical
Arrowhead	ARO-ALK7	ALK7 inhibitor	injectable	no	preclinical
AstraZeneca	AZD9550	GLP-1/glucagon agonist	injectable	no	1
AstraZeneca	AZD6234	amylin analog	injectable	no	1
AstraZeneca/Eccogene	ECC5004/AZD5004	GLP-1	oral	yes	1
Biohaven	taldefgrobep	myostatin/activin A inhibitor	injectable	no	2
Corbus	CRB-913	CB1 inverse agonist	oral	yes	preclinical
D&D Pharmatech	DD01	GLP-1/glucagon agonist	injectable	no	1
D&D Pharmatech	NLY12	GLP-1 agonist	injectable	no	preclinical
D&D Pharmatech	DD13	amylin agonist	injectable	no	preclinical
Elevai Labs	EL-22	myostatin probiotic	oral	no	preclinical
Elevai Labs	EL-32	myostatin/activin A probiotic	oral	no	preclinical
Eli Lilly	retatrutide	GLP-1/GIP/glucagon	injectable	no	3
Eli Lilly	orforglipron	GLP-1	oral	yes	3
Eli Lilly	mazdutide/LY3305677*	GLP-1/glucagon agonist	injectable	no	3
Eli Lilly	dacra QW II (LY3541105)	amylin/calcitonin agonist	injectable	no	1
Eli Lilly	GLP-1 NPA1	GLP-1	injectable	yes	2
Eli Lilly	GIP/GLP coagonist 3	GLP-1/GIP	injectable	no	1
Eli Lilly	LY3841136/eloralintide	amylin receptor agonist	injectable	no	2
Eli Lilly	bimagrumab	activin receptor type IIA/IIB	injectable	no	2
Fractyl Health	GLP-1 agonist gene therapy	GLP-1	injectable	no	preclinical
Gan & Lee	GZR18	GLP-1 agonist	injectable		1/2
Gilead	GS-4571	GLP-1	oral	yes	preclinical
Glyscend	GLY-200	mucin-complexing polymer	oral	yes	2
Gubra	GUB014295 (GUBamy)	amylin agonist	injectable	no	1
Gubra	UCN2	CRHR2 agonist	injectable	no	preclinical
Hanmi	efpeglenatide	GLP-1	injectable	no	3
Hanmi	HM15275	GLP-1/GIP/GCGR	injectable	no	1
Hanmi	efpegerglucagon	glucagon	injectable	no	2
Jiangsu HengRui	Noiiglutide	GLP-1 agonist	injectable	no	3
Keros Therapeutics	KER-065	TGF-beta (myostatin/Activin A)	injectable	no	1
Laekna (China)	LAE102	ActRIIA (activin receptor)	injectable	no	1
Lipocine	LPCN 2401	androgen receptor agonist	oral	yes	2
Merck	efinopegdutide	GLP-1/glucagon agonist	injectable	no	2a
Novo Nordisk		GLP-1/GIP	injectable	no	1
Novo Nordisk	cagrisema	GLP-1/amylin	injectable	no	3
Novo Nordisk	amycretin	GLP-1/amylin	oral or subcu	1 oral, 1 subcu	1
Novo Nordisk	tri-agonist	undisclosed	undisclosed	no	preclinical
Novo Nordisk	amylin monotherapy	amylin	undisclosed	no	preclinical
Novo Nordisk	INV-202/monlunabant)	CB1 receptor blocker	oral	yes	2
Novo Nordisk	INV-347 (next-gen)	CB1 receptor blocker	oral	yes	1
Novo Nordisk	EMB1 (peptide)	EMB1 agonist	injectable	no	preclinical
Novo Nordisk	EMB2 (small molecule)	EMB2 agonist	oral	yes	preclinical
Novo Nordisk	EraCal pipeline	EraCal Therapeutics	oral	yes	preclinical

Source: Morningstar, PitchBook, company reports.

Exhibit 25 (continued) Obesity Development Pipeline — Public Firms

Firm	Drug	Mechanism	Administration	Small Molecule?	Phase
Palatin	bremelanotide	MC4R	injectable	no	2
Pfizer	danuglipron	GLP-1	oral	yes	1
Pfizer	PF-07976016	undisclosed	oral	yes	1
Pfizer/Nxera	PF-06954522	GLP-1	oral	yes	1
Rani/ProGen	PG-102	GLP-1/GLP2	oral	no	1
Regeneron	trevogrumab	myostatin Ab	injectable	no	2
Regeneron	garetosmab	activin A Ab	injectable	no	2
Regeneron	GPR75	GPR75 inhibitor	undisclosed	undisclosed	preclinical
Regeneron	mibavademab	LEPR (leptin)	injectable	no	2
Roche	RO7204239	myostatin Ab	injectable	no	preclinical
Roche/Carmot	CT-388	GLP-1/GIP agonist	injectable	no	1/2
Roche/Carmot	CT-996	GLP-1	oral	yes	1
Scholar Rock	apitegromab	myostatin	injectable	no	2
Scholar Rock	SRK-439	myostatin	injectable	no	preclinical
Shionogi	S-309309	MGAT2 inhibitor	oral	yes	2
Skye Bioscience	nimacimab	CB1 antibody	injectable	no	2
Structure Therapeutics	GSR-1290	GLP-1	oral	yes	2
Structure Therapeutics	TBD	amylin analog	oral	yes	preclinical
Structure Therapeutics	TBD	GIP agonist	oral	yes	preclinical
Structure Therapeutics	TBD	glucagon agonist	oral	yes	preclinical
Structure Therapeutics	ANPA-0073	apelin (APJR)	oral	yes	1
Sun Pharma	GL0034	GLP-1 agonist	injectable	no	1
Terns Pharmaceuticals	TERN-601	GLP-1	oral	yes	1
Terns Pharmaceuticals	TERN-501 combo	THR-beta agonist	undisclosed	undisclosed	preclinical
Ventyx	VT3232	NLRP3 inhibitor	oral	yes	2
Veru	enobosarm	selective androgen receptor modulator	oral	yes	2
Viking	VK2735	GLP-1/GIP agonist	injectable	no	2
Viking	VK2735	GLP-1/GIP agonist	oral	no	1
Viking	DACRA (dual amylin/calcitonin receptor agonists)		injectable	no	preclinical
Vivani	NPM-115	high-dose GLP-1 (exenatide)	implant	no	preclinical
Vivani	NPM-139	GLP-1 (semaglutide)	implant	no	preclinical
Wave Life Sciences	RNAi INHBE	INHBE inhibitor	injectable	no	preclinical
Yunovia (spin off of public Ildong)		GLP-1 agonist	oral	yes	1
Zealand	dapiglutide	GLP-1/GLP-2 agonist	injectable	no	2
Zealand	petrelintide/ZP8396	amylin analog	injectable	no	2
Zealand	ZP 6590	GIPR agonist	injectable	no	preclinical

Source: Morningstar, PitchBook, company reports.

Research Methodology for Valuing Companies

Overview

At the heart of our valuation system is a detailed projection of a company's future cash flows, resulting from our analysts' research. Analysts create custom industry and company assumptions to feed income statement, balance sheet, and capital investment assumptions into our globally standardized, proprietary discounted cash flow, or DCF, modeling templates. We use scenario analysis, in-depth competitive advantage analysis, and a variety of other analytical tools to augment this process. We think analyzing valuation through discounted cash flows presents a better lens for viewing cyclical companies, high-growth firms, businesses with finite lives (mines, for example), or companies expected to generate negative earnings over the next few years. That said, we don't dismiss multiples altogether but rather use them as supporting cross-checks for our DCF-based fair value estimates. We also acknowledge that DCF models offer their own challenges (including a potential proliferation of estimated inputs and the possibility that the method may miss short-term market-price movements), but we believe these negatives are mitigated by deep analysis and our long-term approach.

Morningstar's Equity Research Group ("we," "our") believes that a company's intrinsic worth results from the future cash flows it can generate. The Morningstar Rating for stocks identifies stocks trading at a discount or premium to their intrinsic worth — or fair value estimate in Morningstar terminology. Five-star stocks sell for the biggest risk-adjusted discount to their fair values, whereas 1-star stocks trade at premiums to their intrinsic worth.

Four key components drive the Morningstar rating:

- ▶ our assessment of the firm's economic moat.
- ▶ our estimate of the stock's fair value.
- ▶ our uncertainty around that fair value estimate.
- ▶ the current market price.

This process ultimately culminates in our single-point star rating.

Economic Moat

The Morningstar Economic Moat Rating is a structural feature that Morningstar believes positions a firm to earn durable excess profits over a long period of time, with excess profits defined as returns on invested capital above our estimate of a firm's cost of capital. The economic moat rating is not an indicator of the investment performance of the investment highlighted in this report. Narrow-moat companies are those that Morningstar believes are more likely than not to achieve normalized excess returns for at least the next 10 years. Wide-moat companies are those that Morningstar believes will earn excess returns for 10 years, with excess returns more likely than not to remain for at least 20 years. Firms without a moat, including those that have a substantial threat of value destruction-related risks related to environmental, social, and governance; industry disruption; financial health; or other idiosyncratic issues, are more susceptible to competition. Morningstar has identified five sources of economic moats: intangible assets, switching costs, network effect, cost advantage, and efficient scale.

Fair Value Estimate

Each stock's fair value is estimated by using a proprietary discounted cash flow model, which assumes that the stock's value is equal to the total of the free cash flows of the company is expected to generate in the future, discounted back to the present at the rate commensurate with the riskiness of the cash flows. As with any DCF model, the ending value is highly sensitive to Morningstar's projections of future growth.

Fair Value Uncertainty

The Morningstar Uncertainty Rating represents the analysts' ability to bound the estimated value of the shares in a company around the fair value estimate, based on the characteristics of the business underlying the stock, including operating and financial leverage, sales sensitivity to the overall economy, product concentration, pricing power, exposure to material ESG risks, and other company-specific factors. Based on these factors, analysts classify the stock into one of several uncertainty levels: Low, Medium, High, Very High, or Extreme. Our recommended margin of safety—the discount to fair value demanded before we'd recommend buying or selling the stock—widens as our uncertainty of the estimated value of the equity increases.

Market Price

The market prices used in this analysis and noted in the report come from exchanges on which the stock is listed, which we believe is a reliable source.

Morningstar Rating for Stocks

The Morningstar Rating for Stocks is a forward-looking, analyst-driven measure of a stock's current price relative to the analyst's estimate of what the shares are worth. Stock star ratings indicate whether a stock, in the equity analyst's educated opinion, is cheap, expensive, or fairly priced. To rate a stock, analysts estimate what they think it is worth (its "fair value"), using a detailed, long-term cash flow forecast for the company. A stock's star rating depends on whether its current market price is above or below the fair value estimate. Those stocks trading at large discounts to their fair values receive the highest ratings (4 or 5 stars). Stocks trading at large premiums to their fair values receive lower ratings (1 or 2 stars). A 3-star rating means the current stock price is close to the analyst's fair value estimate.

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