

Q32024 Presentation

Zealand Pharma



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Significant progress across obesity pipeline

Petrelintide (amylin analog)	Dapiglutide (dual GLP-1/GLP-2 receptor agonist)	Survodutide ^b (dual GCG/GLP-1 receptor agonist)
Presented extremely encouraging weight loss and tolerability data from Phase 1b trial (MAD Part 2) ^a	Reported positive topline data from Part 1 of the Phase 1b trial	Boehringer Ingelheim announced BTD for survodutide in MASH and initiated two Phase 3 trials
Potential best-in-class alternative to GLP-1RA-based therapies	Potential first-in-class therapy for obesity and inflammation-related co-morbidities	Potential best-in-class therapy for obesity and MASH

^aTopline results from the Phase 1b trial were announced on June 20, 2024. Detailed results were presented at ObesityWeek 2024 on November 5, 2024.

^bSurvodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally (subject to Zealand's co-promotion rights in the Nordic countries). MAD=multiple ascending dose; GLP-1RA=glucagon-like peptide-1 receptor agonist; GLP-2=glucagon-like peptide-2; GCG=glucagon; MASH=metabolic dysfunction-associated steatohepatitis (formerly NASH, or nonalcoholic steatohepatitis); BTD=Breakthrough Therapy Designation

GLP-1RA-based therapies are effective at reducing weight (in PwO, but are associated with GI tolerability issues¹



There is a significant unmet need for alternative treatment options with different mechanisms of action

Today, **two QW GLP-1RA-based therapies are approved**,^{a,2,3} offering ~15–21% mean weight loss^{4,5}



GLP-1RAs are commonly associated with GI side effects, including constipation, nausea, vomiting and diarrhea^{4,5}



Up to **30%** of patients with obesity **discontinue GLP-1RA treatment** within **1 month**⁶



Up to 60–70% of patients discontinue GLP-1RA treatment within 12 months⁷

Petrelintide represents an alternative to GLP-1RA-based therapies targeting:



15–20% mean weight loss and high-quality weight loss with potential for preservation of lean mass



Reduced food intake via a **non-incretin mechanism** that increases satiety and restores leptin sensitivity



Significantly improved GI tolerability with both lower frequency and severity of adverse events

^aFor chronic weight management: Wegovy and Zepbound.

Sources: 1. Wang et al. Front Endocrinol (Lausanne) 2023;14:1085799; 2. Wegovy (semaglutide) US PI. Available from: https://www.accessdata.fda.gov/drugsatfda_docs/label/2024/215256s011lbl.pdf, accessed July 2024; 3. Zepbound (tirzepatide) US PI. Available from: https://www.accessdata.fda.gov/drugsatfda_docs/label/2024/217806s003lbl.pdf, accessed July 2024; 4. Wilding et al. N Engl J Med 2021;384(11):989–1002; 5. Jastreboff et al. N Engl J Med 2022;387(3):205–216; 6. Blue Health Intelligence. Real-world trends in GLP-1 treatment persistence and prescribing for weight management. May 2024; 7. Gasoyan et al. Obesity (Silver Spring) 2024;32(3):486–493. Gl=gastrointestinal; GLP-1RA=glucagon-like peptide-1 receptor agonist; PwO=people with obesity; QW=once-weekly.

Our R&D pipeline addresses unmet medical needs across several therapeutic areas





alnvestigational compounds whose safety and efficacy have not been evaluated or approved by the U.S. Food and Drug Administration (FDA) or any other regulatory authority.

^bSurvodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally (subject to Zealand's co-promotion rights in the Nordic countries): EUR 315 million outstanding potential development, regulatory and commercial milestones + high single to low double digit % royalties on global sales.

GCGR=glucagon receptor; GIP=gastric inhibitory polypeptide; GLP-1R=glucagon-like peptide-1 receptor; GLP-2=glucagon-like peptide-2; GLP-2R=glucagon-like peptide-2 receptor; MASH=metabolic dysfunction-associated steatohepatitis (formerly NASH, or nonalcoholic steatohepatitis); SC=subcutaneous; T1DM=type 1 diabetes mellitus.

Petrelintide

Design of Phase 1b MAD trial Part 2 with petrelintide





^aSafety evaluation occurred after 4 weeks of treatment at the target dose for each cohort.² Initiation of the next, higher dose cohort only occurred following safety evaluation for the previous cohort.²

Sources: 1. ClinicalTrials.gov (NCT05613387), accessed October 2024; 2. Data on file.

BMI=body mass index; HbA1c=glycated hemoglobin; MAD=multiple ascending dose; MTM=mixed test meal; PD=pharmacodynamics; PK=pharmacokinetics; SC=subcutaneous; TEAE=treatment-emergent adverse event.

Petrelintide

Substantial weight loss was observed with petrelintide at 16 weeks in the Phase 1b MAD trial Part 2

ZEAL&

Observed mean (95% CI) percent change from baseline in body weight



^aEOT includes measurements at the EOT visit, performed at 24 or 25 weeks after dosing, and also performed for participants discontinuing treatment early.

^bOne participant had one extra week at 7.5 mg, and thereby only five weeks on maintenance dose at Week 16. After Week 16, this participant is included with weeks after last dosing.

Source: Data on file. Data presented at ObesityWeek 2024 in San Antonio, Texas.

CI=confidence interval; EOT=end of trial; MAD=multiple ascending dose

High levels of study treatment completion and adherence with dose escalation within cohorts





- Three participants discontinued petrelintide: one due to AEs, one to focus on recovery from a cold, and one due to personal reasons
- One participant in the 9.0 mg arm had an extra week at 7.5 mg (due to tolerability)
- · The remaining participants followed dose escalation steps within cohorts

Obesity

Petrelintide

Petrelintide

Vast majority of TEAEs reported by petrelintidetreated patients were mild





Source: Data on file. Data presented at ObesityWeek 2024 in San Antonio, Texas.

5 moderate AEs reported by petrelintide exposed participants: nausea, vomiting, nasopharyngitis, acute sinusitis, back pain

E=number of events; N=number of participants; TEAE=treatment-emergent adverse event.

Petrelintide

Petrelintide treatment appeared safe and was well-tolerated at all dose levels in the 16-week trial



All GI TEAEs were mild, except for one event of moderate nausea and moderate vomiting in one participant



Source: Data on file. Data presented at ObesityWeek 2024 in San Antonio, Texas.

N=12 in each treatment group.

E=number of events; TEAE=treatment-emergent adverse event; GI=Gastrointestinal

Petrelintide

Continuing development of petrelintide as monotherapy through a comprehensive Phase 2b trial

A randomized, double-blind, placebo-controlled, Phase 2b trial with petrelintide to be initiated in Q4 2024



Source: Data on file. BMI=body mass index; HbA1c=glycated hemoglobin; MRI=magnetic resonance imaging

Dapiglutide

Placebo-adjusted mean weight loss of up to 8.3% with dapiglutide in 13-week Part 1 of Phase 1b trial



The 13-week Phase 1b trial evaluated higher doses of dapiglutide than prior clinical trials^a Next steps include topline results from 28-week trial and initiation of Phase 2b trial



Placebo-adjusted weight loss of up to a mean of 8.3% after 13 weeks



- Total of 54 participants, 85% male with a median baseline BMI of 30.0 kg/m²
- No lifestyle modifications, such as diet or exercise, were included in trial



- Dapiglutide up to 13 mg assessed to be safe and well-tolerated with GI AEs consistent with profile of incretin-based therapies
- Two participants out of 54 discontinued treatment due to GI AEs



- Part 2 of the Phase 1b trial includes a higher dose cohort (up to 26 mg) with 28 weeks treatment using monthly dose escalation
- Topline results expected in H1 2025



Large, comprehensive Phase 2b trial to be initiated in H1 2025

^aDREAM, an investigator-led mechanistic trial with dapiglutide: ClinicalTrials.gov (NCT05788601), and the 4-week Phase 1a clinical trial: ClinicalTrials.gov (NCT04612517). BMI=body mass index; GI=gastrointestinal; AE=adverse event.

MASH

Survodutide

Survodutide^a GCG/GLP-1 receptor dual agonist shows best-in-class potential in MASH Phase 2 trial



Boehringer Ingelheim

Phase 2 biopsy-driven trial in people with MASH¹



Participants showing **improvement in MASH** without worsening of fibrosis (stages F1-F3): **83.0% with survodutide** vs 18.2% with placebo (p<0.0001)



Participants showing **improvement in liver fibrosis** with no worsening of MASH (stages F2-F3): **64.5% with survodutide** vs 25.9% with placebo (p=0.0007)



Survodutide treatment **did not show unexpected safety or tolerability issues**, including at the higher dose of 6.0 mg



Boehringer received **U.S. FDA BTD for survodutide** in MASH and fibrosis (stages F2-F3), and **launched two Phase 3 trials**³ in MASH for survodutide

Improvement in liver fibrosis with no worsening of MASH

Paired biopsy results (F2/F3)²



^aSurvodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally (subject to Zealand's co-promotion rights in the Nordic countries). Sources: 1. Boehringer Ingelheim press release June 7, 2024. Data presented at the EASL Congress 2024 in Milan, Italy. 2. A sensitivity analysis based on participants with paired biopsy results at baseline and end of treatment. 3. Boehringer Ingelheim press release October 8, 2024.

MASH= metabolic dysfunction-associated steatohepatitis (formerly NASH=non-alcoholic steatohepatitis); Cl=confidence interval; QW=once-weekly; GCG=glucagon; GLP-1=glucagon-like peptide-1; BTD= Breakthrough Therapy Designation

MASH

Survodutide

Phase 3 program with survodutide in MASH has been initiated





	Inclusion criteria	Study design	Primary endpoint
LIVERAGE ^{TM 1} • Efficacy and safety in patients with MASH and fibrosis (F2/F3)	 Diagnosis of MASH^a and biopsy- proven fibrosis stage F2-F3 Granted Breakthrough Therapy Designation by the U.S. FDA² 	 N=1,800 1:1 ratio (6.0 mg or placebo) Trial duration Part 1: 52 weeks Part 2: Up to 7 years 	 Part 1: 52 weeks MASH resolution without worsening of liver fibrosis, and Improvement in fibrosis stage with no worsening of MASH Part 2: Time to first occurrence of liver-related events or all-cause mortality
LIVERAGE TM -Cirrhosis ³ Efficacy and safety in patients with MASH and cirrhosis (F4)	 Diagnosed compensated MASH cirrhosis^b 	 N=1,590 1:1 ratio (6.0 mg or placebo) Trial duration: Up to 4.5 years 	Time to first occurrence of liver-related events or all-cause mortality

^aMASH diagnosis defined by a NAS score≥4, with at least 1 point in inflammation and ballooning each. ^bDiagnosed according to modified Liver Forum criteria (Noureddin et al, Gastroenterology 2020;159:422-427) Inclusion criteria for both trials include age ≥18 years. Further inclusion criteria apply.

Liver-related events include progression to cirrhosis (LIVERAGE), liver transplant, hepatic decompensation event(s), worsening of MELD score to ≥15, and progression to CSPH

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Rare diseases

Regulatory status in the US for our rare disease programs



Dasiglucagon in CHI: Regulatory submissions expected in Q4 2024



Glucagon receptor agonist designed to allow for continuous subcutaneous infusion via a wearable pump system^a



Due to the timing of a third-party manufacturing facility reinspection, the U.S. FDA issued a CRL for dasiglucagon in CHI for up to three weeks of dosing^b.



Zealand expects to resubmit Part 1 of NDA related to dosing of up to three weeks by the end of 2024



Zealand expects to submit Part 2 of NDA related to dosing beyond three weeks by the end of 2024^b

^aZealand Pharma has entered a collaborative development and supply agreement with DEKA Research & Development Corporation and affiliates for infusion pump system.

^bFDA issued a Complete Response Letter (CRL) to Part 1 of the NDA due to inspection findings at a third-party manufacturing facility that were not specific to dasiglucagon; Part 2 to be supported by additional analyses from existing CGM datasets included as a secondary outcome measure in the Phase 3 program.

CHI=congenital hyperinsulinism; SBS-IF=short bowel syndrome with intestinal failure; NDA=new drug application; PDUFA=Prescription Drug User Fee Act; FDA=Food and Drug Administration; CGM=continuous glucose monitoring; GLP-2=glucagon-like peptide-2; CRL=complete response letter

Glepaglutide in SBS-IF: PDUFA date December 22, 2024



GLP-2 receptor agonist designed to be administered in **ready-to-use auto-injector** with needle protection



Late-cycle meeting with US FDA completed

Rare diseases

Near-term commercial opportunities through rare disease franchise



Congenital hyperinsulinism (CHI)

- Ultra-rare disease in newborns and children
- CHI can cause serious episodes of hypoglycemia
- >50% of CHI patients may be unresponsive to current treatment options¹
- Focus is on bringing the product to patients as quickly as possible
- Preparing for U.S. launch in 1H 2025 contingent on regulatory approval
 - Growing commercialization capabilities prior to potential FDA approval
 - All effort is focused on setting up the necessary architecture to serve the patients in the best way possible

Short bowel syndrome (SBS)

- Rare disease resulting in impaired intestinal absorptive capacity, resulting in dependency on parenteral support^{2,3}
- One product is marketed for the disease, but many patients remain uncontrolled and untreated
- Unmet need for improved treatment options which may allow patients the potential to ease the burden of complex disease management⁴
- Undertaking pre-commercial activities to enable launch post approval
 - Evaluated market dynamics
 - Established brand and go-to-market strategy
 - Remain focused on finding a partner to maximize the opportunity

Sources: 1. Yorifuji et al. Clin Pediatr Endocrinol 2017;26(3):127-152; 2. Jeppesen P., Expert Opinion on Orphan Drugs; 1:515-25, 2013; 3. Pironi, L, et al. Definitions of intestinal failure and the short bowel syndrome. Best Practice & Research Clinical Gastroenterology. 30(2), 173-185 (2016); 4. Cueda C et al. ESPEN Practical Guideline: clinical nutrition in chronic intestinal failure. Clin Nutrition 40; 5196-5120 (2021); Dasiglucagon and glepaglutide are investigational compounds whose safety and efficacy have not been evaluated or approved by the FDA or any other regulatory authority.

Opportunity for establishing a new foundational therapy for weight management



Clear societal impact



The obesity pandemic represents the greatest healthcare challenge of our time



50% of adults globally are expected to have overweight or obesity by 2030¹



More than **5 million deaths** globally are **ascribed to overweight and obesity** every single year¹





Eligible patients in the US receiving prescriptions for weight loss therapy²



Up to **30%** of patients with obesity **discontinue GLP-1RA treatment** within **1 month**³



More than **80% of** patients **discussed how to maintain muscle** or **address muscle loss**⁴

Opportunities

Patient perceptive



~2/3 of adults want to lose up to 20% of current weight⁴



More than **50%** of patients are **NOT willing to accept GI AEs**, including nausea, vomiting and diarrhea⁵

PCP perspective



Limited time for patients and **no** time for follow-ups + limited knowledge of weight management result in call backs

Sources: 1. World Obesity Atlas 2024; 2. Novo Nordisk H1 2024 Investor Presentation: <u>https://investor.novonordisk.com/q2-2024-presentation/?page=45.;</u> 3. Blue Health Intelligence. Real-world trends in GLP-1 treatment persistence and prescribing for weight management. May 2024; LifeSci Capital Survey May 2024 (N=4995); 4. Bloomberg Intelligence – Obesity Prescriber Survey, April 2024 (N=100); 5. Kmodo Claims Database (2023); HCP=healthcare professional; GLP-1RA=glucagon-like peptide-1 receptor agonist; WL=weight loss; NCD=non-communicable diseases; AOM =Anti-obesity market; GI AEs=gastrointestinal adverse events

Petrelintide

The target product profile of petrelintide holds potential to address the needs of patients and HCPs



Establishing petrelintide as the future foundational therapy for weight management Target product profile: Market benefit: 1 GLP-1RA-like weight loss (~15-20%) Vast majority of patients seek weight loss of 10-20% 2 Significantly improved GI tolerability profile Lower frequency and severity of GI AEs 3 Potential for higher quality of weight loss Potential to preserve lean muscle mass Potential for better patient experience ("feeling better") 4 Unique MoA (satiety) with ability to eat but feel full (no "food aversion") Sweet spot for PCPs 5 Simplicity (fewer call-backs)

HCP=healthcare professional; GLP-1RA=glucagon-like peptide-1 receptor agonist; GI=gastrointestinal; MoA=mechanism of action; AE=adverse event; PCP=primary care physician.

Q3 2024 YTD Profit & Loss



DKK million	Q3-24 YTD	Q3-23 YTD
Revenue	53.6	319.6
Cost of goods sold	-7.5	-5.2
Gross profit	46.2	314.4
Research and development expenses	-665.9	-494.7
Sales and marketing expenses	-50.2	-17.8
General and administrative expenses	-199.8	-134.4
Other operating Items	-3.1	13.8
Net operating expenses	-919.1	-633.2
Operating result	-872.9	-318.8
Net financial items	81.1	-124.8
Result before tax	-791.8	-443.5
Тах	4.0	4.6
Net result for the period	-787.8	-439.0

P&L reflecting Zealand's investment in its differentiated assets targeting obesity

- Revenue of DKK 54 million is mainly driven by the license and development agreement with Novo Nordisk for Zegalogue[®].
- Total operating expenses of DKK 919 million are higher than last year, primarily driven by the increase in R&D expenses due to clinical advancement of the obesity pipeline and activities supporting the regulatory review by the US FDA of the late-stage rare disease assets. S&M expenses are mainly driven by pre-commercial activities for the rare disease assets while the increase in G&A expenses reflect additional legal expenses related to our patent portfolio and strengthening of organizational capabilities.
- Net financial items of DKK 81 million are mainly driven by interest income from marketable securities.



Strong cash position of DKK 9.2 billion enables significant investments in our obesity programs



Secured cash of DKK 8.6 billion in 2024 through capital raises and the EIB loan facility^a



^aThe EUR 50 million Tranche A of the EIB loan facility was disbursed in March 2024. ^bCash position includes cash, cash equivalents and marketable securities. EIB loan Tranches B and C (EUR 20 million each) are excluded from this chart. The two tranches are subject to pre-specified milestones being met.

EIB = European Investment Bank



2024 financial guidance

DKK million	2024 Guidance	2023 Actuals
Revenue anticipated from existing and new license and partnership agreements	No guidance due to uncertain size and timing	343
Net operating expenses ^a	1,250 – 1,350	896

Exciting news flow with many potential catalysts expected in the near term



NON-EXHAUSTIVE

Q4 2024 H1 2025 Petrelintide Petrelintide Initiate Phase 2b trial (overweight/obesity without T2D) Complete enrollment in Phase 2b trial (overweight/obesity without T2D) **Dasiglucagon (CHI)** Petrelintide Resubmit Part 1 of NDA to US FDA Initiate Phase 2b trial (overweight/obesity with T2D) **Dasiglucagon (CHI)** Dapiglutide Report topline results from Part 2 of Phase 1b dose-titration trial (28wks) Submit analyses supporting chronic use to US FDA Dapiglutide **Glepaglutide (SBS)** Initiate Phase 2b trial (overweight/obesity) Gain US regulatory NDA decision ZP9830 (Kv1.3 Ion Channel Blocker) Dapiglutide Initiate first-in-human clinical trialsre Present results from Phase 1b dose-titration trial **Dasiglucagon (CHI)** Gain US regulatory decision for Part 1 and 2 of NDA Legend: Obesity Inflammation Rare diseases Potential partnership agreements across therapeutic areas





Q3 2024 November 7, 2024