



### **Forward-looking Statements**

This presentation contains "forward-looking statements", as that term is defined in the Private Securities Litigation Reform Act of 1995 in the United States, as amended, even though no longer listed in the United States this is used as a definition to provide Zealand Pharma's expectations or forecasts of future events regarding the research, development and commercialization of pharmaceutical products, the timing of the company's pre-clinical and clinical trials and the reporting of data therefrom and the company's significant events and potential catalysts in 2025 and any financial guidance published by the company, as applicable. These forward-looking statements may be identified by words such as "aim," "anticipate," "believe," "could," "estimate," "expect," "forecast," "goal," "intend," "may," "plan," "possible," "potential," "will," "would" and other words and terms of similar meaning. You should not place undue reliance on these statements, or the scientific data presented.

The reader is cautioned not to rely on these forward-looking statements. Such forward-looking statements are subject to risks, uncertainties and inaccurate assumptions, which may cause actual results to differ materially from expectations set forth herein and may cause any or all of such forward-looking statements to be incorrect, and which include, but are not limited to, unexpected costs or delays in clinical trials and other development activities due to adverse safety events, patient recruitment or otherwise; unexpected concerns that may arise from additional data, analysis or results obtained during clinical trials; our ability to successfully market both new and existing products; changes in reimbursement rules and governmental laws and related interpretation thereof; government-mandated or market-driven price decreases for our products; introduction of competing products; production problems at third party manufacturers; dependency on third parties, for instance contract research or development organizations; unexpected growth in costs and expenses; our ability to effect the strategic reorganization of our businesses in the manner planned; failure to protect and enforce our data, intellectual property and other proprietary rights and uncertainties relating to intellectual property claims and challenges; regulatory authorities may require additional information or further studies, or may reject, fail to approve or may delay approval of our drug candidates or expansion of product labeling; failure to obtain regulatory approvals in other jurisdictions; exposure to product liability and other claims; interest rate and currency exchange rate fluctuations; unexpected contract breaches or terminations; inflationary pressures on the global economy; and political uncertainty.

If any or all of such forward-looking statements prove to be incorrect, our actual results could differ materially and adversely from those anticipated or implied by such statements. The foregoing sets forth many, but not all, of the factors that could cause actual results to differ from our expectations in any forward-looking statement. All such forward-looking statements speak only as of the date of this presentation and are based on information available to Zealand Pharma as of the date of this presentation. We do not undertake to update any of these forward-looking statements to reflect events or circumstances that occur after the date hereof.

Information concerning pharmaceuticals (including compounds under development) contained within this material is not intended as advertising or medical advice.



### **Table of contents**

- **About Zealand Pharma**
- **Obesity**

Petrelintide (amylin analog)

Dapiglutide (GLP-1/GLP-2 receptor dual agonist)

Survodutide (glucagon/GLP-1 receptor dual agonist)

Rare diseases

Dasiglucagon for congenital hyperinsulinism

Glepaglutide for short bowel syndrome

**Inflammation** 

ZP9830 (Kv1.3 blocker)

**Additional company information** 

Profit & Loss (Q1 2025)

Cash position

2025 financial guidance

Management Team

- 14
- 32
- 38
- **49**
- **53**
- **57**
- **59**
- 61

- 62 63 64
- **65**



### **About Zealand Pharma**

**Corporate Presentation** 

### We are an international biotech company headquartered in Denmark





Headquarters and labs outside Copenhagen, Denmark

#### Founded in 1998

- Peptide platform validated through two approved products marketed by Sanofi and Novo Nordisk
- ~440 employees globally as of June 10, 2025

### Listed on NASDAQ CPH (ZEAL.CO)

- Market cap on June 10, 2025: USD ~4.8B (DKK ~31.8B)
- 71.0M Shares Outstanding as of June 7, 2025

### Cash position<sup>a</sup>

USD ~1.3B (DKK ~8.5B) as of March 31, 2025

### **OPEX guidance for 2025**<sup>b</sup>

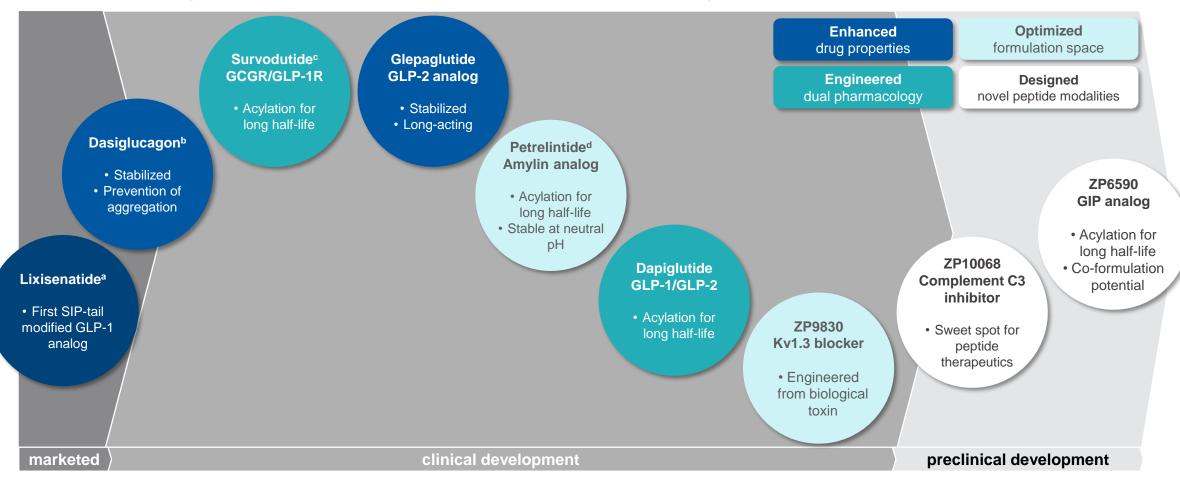
Net operating expenses are expected to be DKK 2,000-2,500M

aCash position includes cash, cash equivalents and marketable securities. Zealand also has a Loan facility with the European Investment Bank (EIB) of EUR 90 million in three tranches (Tranche A of EUR 50 million was disbursed in Q1 2024; Tranches B and C are subject to pre-specified milestones being met). Financial considerations from the collaboration and licensing agreement with Roche, announced in March 2025, is not included in these figures. <sup>b</sup>Financial guidance for 2025 excludes transaction-related costs associated with the Roche collaboration.

Based on foreign exchange rates as of June 10, 2025 (DKK 6.6 = USD \$1). OPEX=Operating Expenses.

### We strive to be the world's best peptide drug discovery and development company





<sup>&</sup>lt;sup>a</sup>Marketed globally by Sanofi.

bLicensed to Novo Nordisk: DKK 227.5 million outstanding in potential development, regulatory, manufacturing and sales milestones + high single to low double digit % royalties on global sales.

cLicensed to Boehringer Ingelheim: Boehringer solely responsible for development and commercialization globally. EUR 315 million outstanding potential development, regulatory and commercial milestones + high single to low double digit % royalties on global sales.

<sup>&</sup>lt;sup>d</sup>Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe.

GCGR=glucagon receptor; GIP=gastric inhibitory polypeptide; GLP-1=glucagon-like peptide-1; GLP-1R=glucagon-like peptide-1 receptor; GLP-2=glucagon-like peptide-2; SIP=structure-inducing probe.

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



|                                    | Product candidate <sup>a</sup>                        | Partnered               | Pre-clinical     | Phase 1    | Phase 2 | Phase 3 | Registration |
|------------------------------------|---|-------------------------|------------------|------------|---------|---------|--------------|
| Obesity and related co-morbidities | Petrelintide (amylin analog) <sup>b</sup>             | Roche                   | Obesity          |            |         |         |              |
|                                    | Petrelintide/CT-388 (amylin + GLP-1/GIP) <sup>b</sup> | Roche                   | Obesity          |            |         |         |              |
|                                    | Dapiglutide (GLP-1R/GLP-2R dual agonist)              |                         | Obesity          |            |         |         |              |
|                                    | ZP6590 (GIP receptor agonist)                         |                         | Obesity          |            |         |         |              |
|                                    | Survodutide (GCGR/GLP-1R dual agonist)°               | Boehringer<br>Ingelheim | Obesity          |            |         |         |              |
|                                    | Survodutide (GCGR/GLP-1R dual agonist) <sup>c</sup>   | Boehringer<br>Ingelheim | MASH             |            |         |         |              |
| V                                  |   | Ü                       |                  |            |         |         |              |
| Rare diseases                      | Dasiglucagon: SC continuous infusion                  |                         | Congenital hyper | insulinism |         | :       |              |
|                                    | Glepaglutide (GLP-2 analog)                           |                         | Short bowel synd | rome       |         |         |              |
| u                                  |   |                         |                  |            |         |         |              |
| Inflammation                       | ZP9830 (Kv1.3 ion channel blocker)                    |                         | Undisclosed      |            |         |         |              |
|                                    | ZP10068 (complement C3 inhibitor)                     |                         | Undisclosed      |            |         |         |              |
| ٤                                  |   |                         |                  |            |         |         |              |

<sup>a</sup>Investigational 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.

bCollaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe; Survodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally. EUR 315 million outstanding in 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.

### Exciting news flow with many potential catalysts in the next ~12 months



NON-EXHAUSTIVE

Q2 2025

#### **Petrelintide**<sup>a</sup>

Subgroup analysis from Part 2 of Ph1b trial (16wks) at ADA 2025

#### **Dapiglutide**

Presentation of results from Ph1b dose-titration trial (13wks) at ADA 2025

#### Dapiglutide

Topline results from Part 2 of Ph1b dose-titration trial (28wks)

H<sub>2</sub> 2025

#### Survodutide<sup>b</sup>

Completion of Ph3 obesity trials (SYNCHRONIZE™-1 and 2)

#### **Dapiglutide**

Initiation of Ph2 trial (overweight/obesity)

#### Glepaglutide (SBS)

Initiation of additional Ph3 trial (EASE-5)

### Glepaglutide (SBS)

Submission of MAA to EMA

**Zealand Pharma Capital Markets Day** 

H1 2026

#### **Petrelintide**<sup>a</sup>

Topline results from Ph2 ZUPREME-1 trial

#### **Petrelintide**<sup>a</sup>

Completion of Ph2 ZUPREME-2 trial

#### Petrelintide/CT-388a

Initiation of Ph2 trials

#### Survodutide<sup>b</sup>

Topline results from Ph3 obesity trials

**ZP9830 (Kv1.3 Ion Channel Blocker)**Topline results from Ph1 SAD trial

Legend:

Obesity

Rare diseases

Inflammation

#### Potential partnership agreements across therapeutic areas

<sup>&</sup>lt;sup>a</sup>Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe.

bSurvodutide is licensed to Boehringer Ingelheim, with Boehringer solely responsible for development and commercialization globally. Primary completion of SYNCHRONIZE™-1 and 2 is expected in H2 2025, ClinicalTrials.gov (NCT06066515; NCT06066528), accessed April 2025.

SAD=single ascending dose; SBS=short bowel syndrome; MAA=marketing authorization application; EMA=European Medicines Agency; ADA=American Diabetes Association



### **Obesity**

**Corporate Presentation** 

# The obesity pandemic represents one of the greatest healthcare challenges of our time





For **300,000 years**, human beings maintained a relatively **stable BMI**...



The obesity pandemic has evolved in only 50 years

**50% of adults** globally are expected to have **overweight or obesity** by 2030<sup>1</sup>



Today, more than 5 million deaths globally are ascribed to overweight and obesity every single year<sup>1</sup>

Early days in the evolution of this market...





There is a significant unmet medical need for more and better treatment options

Sources: <sup>1</sup>World Obesity Atlas 2024; <sup>2</sup>Almandoz et al. (2024) Nutritional considerations with antiobesity medications, Obesity (Silver Spring), 32(9): 1613-1631; <sup>3</sup>American Medical Association 2024: https://www.ama-assn.org/topics/obesity.

BMI=body mass index; CVD=cardiovascular disease.

# We believe in a shift from maximizing weight loss to improved tolerability and effects on comorbidities...



### Segment characteristics and key focus areas



#### Prescriberdriven

### **High-risk obesity**

Specialist-driven prescriptions with focus on benefits on co-morbidities and health impact of weight loss.

#### Focus on:

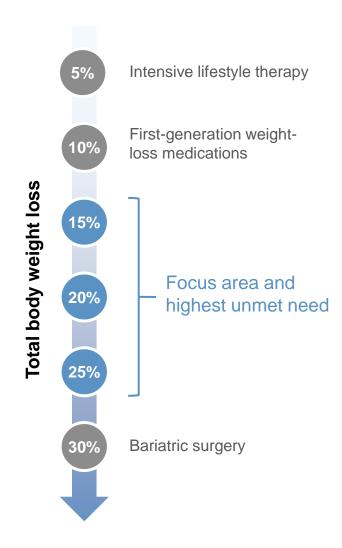
- 1. Comorbidity risk reduction
- 2. Health outcomes
- 3. Relative weight loss
- 4. Tolerability and convenience of treatment

### **Moderate-risk obesity**

Patient-driven primary care prescriptions with focus on quality of weight loss and convenience of treatment.

#### Focus on:

- 1. Desired weight loss
- 2. Tolerability and convenience of treatment
- 3. Quality of weight loss, incl. muscle preservation
- 4. Health outcomes





### ...and that success of future weight-loss medications relies on differentiation on multiple fronts



**GLP-1-based** 

GLP-1 mono

GLP-1/GIP

GLP-1/GCG<sup>a</sup>

GLP-1/amylin

GLP-1/GLP-2a

**Amylin-based** 

Amylin mono<sup>a</sup>

Amylin/GLP-1/GIP<sup>a</sup>

Amylin/other

Other drug classes

### **Examples of differentiation factors**



Unique **non-incretin mechanisms**, addressing quality of weight loss for weight maintenance (incl. preservation of muscle mass)



Improved tolerability by addressing GI side effects



Effects on obesity-related comorbidities



Offer **greater convenience** through dosing regimen and/or delivery method



Develop loose 'flexible-use' or **fixed-dose combinations** for patient segments who need the highest weight loss

### Zealand Pharma has a rich clinical-stage pipeline of differentiated product candidates for obesity



#### **Petrelintide**<sup>a</sup>

Long-acting amylin analog

Ph2 trials in obesity ongoing

Potential best-in-class alternative to GLP-1RA-based therapy



### Petrelintide/CT-388a

Amylin + GLP-1/GIP fixed-dose combination

Ph2 initiation in obesity expected in H1 2026

Potential best-in-disease weight loss efficacy and glycemic control



### **Survodutide**<sup>b</sup>

Glucagon/GLP-1 receptor dual agonist

Ph3 programs in obesity and MASH ongoing

Potential best-in-class therapy for obesity and MASH



### **Dapiglutide**

GLP-1/GLP-2 receptor dual agonist

Ph2 in obesity planned for initiation in H2 2025

Potential first-in-class therapy for obesity and inflammation-related comorbidities

<sup>&</sup>lt;sup>a</sup>Collaboration and license agreement with Roche, including co-development and co-commercialization in the U.S. and Europe.

bSurvodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally. EUR 315 million outstanding potential development, regulatory and commercial milestones + high single to low double digit % royalties on global sales.

 $GLP-1RA=glucagon-like\ peptide-1\ receptor\ agonist;\ GLP-1=glucagon-like\ peptide-1;\ GIP=gastric\ inhibitory\ polypeptide;\ GLP-2=glucagon-like\ peptide-2;\ MASH=metabolic\ dysfunction-associated\ steatohepatitis.$ 

# **GLP-1RA-based therapies effectively reduce body** weight but are associated with GI tolerability issues<sup>1</sup>



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<sup>4,5</sup>



GLP-1RAs are commonly associated with GI side effects, including constipation, nausea, vomiting and diarrhea<sup>4,5</sup>



Up to 30% of patients with obesity discontinue GLP-1RA treatment within 1 month<sup>6</sup>



Up to 60–70% of patients discontinue GLP-1RA treatment within 12 months<sup>7</sup>

Petrelintide<sup>b</sup> is a long-acting amylin analog with the following target product profile:



**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

<sup>a</sup>For chronic weight management: Wegovy and Zepbound; <sup>b</sup>Collaboration and license agreement with Roche, including co-development and co-commercialization in the U.S. and Europe.

Sources: <sup>1</sup>Wang et al. Front Endocrinol (Lausanne) 2023;14:1085799; <sup>2</sup>Wegovy (semaglutide) US Pl. Available from: https://www.accessdata.fda.gov/drugsatfda\_docs/label/2024/215256s011lbl.pdf, accessed July 2024; <sup>3</sup>Zepbound (tirzepatide) US Pl. Available from: https://www.accessdata.fda.gov/drugsatfda\_docs/label/2024/217806s003lbl.pdf, accessed July 2024; <sup>4</sup>Wilding et al. N Engl J Med 2021;384(11):989–1002; <sup>5</sup>Jastreboff et al.

N Engl J Med 2022;387(3):205–216; <sup>6</sup>Blue Health Intelligence. Real-world trends in GLP-1 treatment persistence and prescribing for weight management. May 2024; <sup>7</sup>Gasoyan et al. Obesity (Silver Spring) 2024;32(3):486–493.

Gl=gastrointestinal; GLP-1RA=glucagon-like peptide-1 receptor agonist; QW=once-weekly.

Intellectual property: Composition of matter, patent expiry in 2037. Patent-term extension up to 5 years, i.e. 2042. Potential rights beyond 2042 based on patent applications and additional elements.

# Petrelintide is a long-acting, potential best-in-class amylin analog designed with stability at neutral pH



**Petrelintide** (ZP8396) is a 36-amino-acid acylated peptide, based on the peptide sequence of **human amylin**<sup>1</sup>





Long-acting amylin analog due to acylation (half-life of 10 days), suitable for once-weekly administration<sup>1,2</sup>



Chemical and physical stability with no fibrillation around neutral pH, allowing for co-formulation and co-administration with other peptides<sup>3,4</sup>



Potent balanced agonist effect on amylin and calcitonin receptors<sup>1,5</sup>

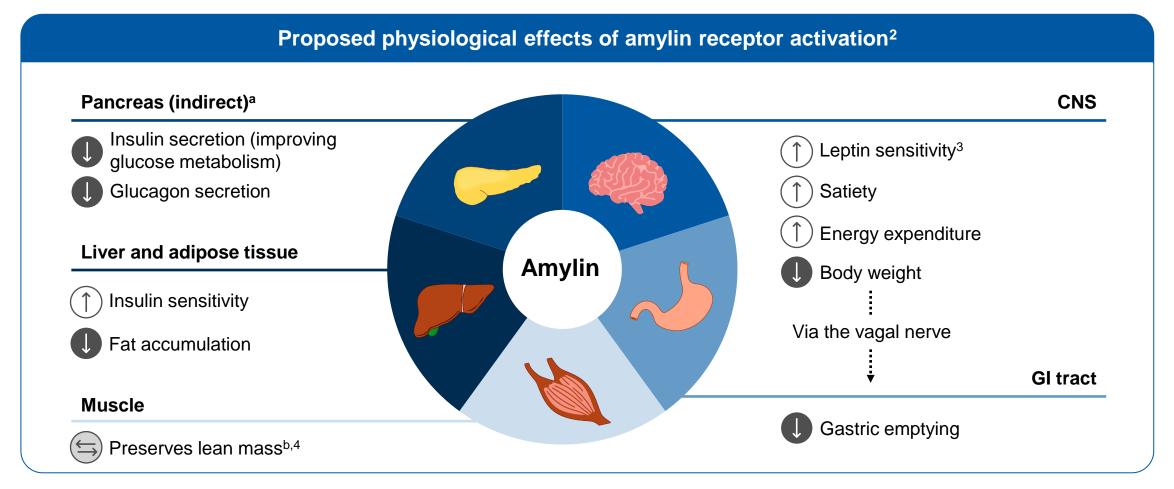
Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe.

Sources: ¹Data on file; ²Brændholt Olsen et al. Poster 92-LB. Presented at ADA 83<sup>rd</sup> Scientific Sessions, June 23–26, 2023, San Diego, CA; ³Skarbaliene et al. Poster 1406-P. Presented at ADA 82<sup>nd</sup> Scientific Sessions, June 3–7, 2022, New Orleans, LA; ⁴Eriksson et al. Poster 532. Presented at ObesityWeek, November 1–4, 2022, San Diego, CA; ⁵Eriksson et al. Presentation at ObesityWeek, November 1–4, 2022, San Diego, CA.

Intellectual property: Composition of matter, patent expiry in 2037. Patent-term extension up to 5 years, i.e. 2042. Potential rights beyond 2042 based on patent applications and additional elements.

# Native amylin, a non-incretin hormone<sup>1</sup>, increases satiety, unlike GLP-1, which reduces appetite

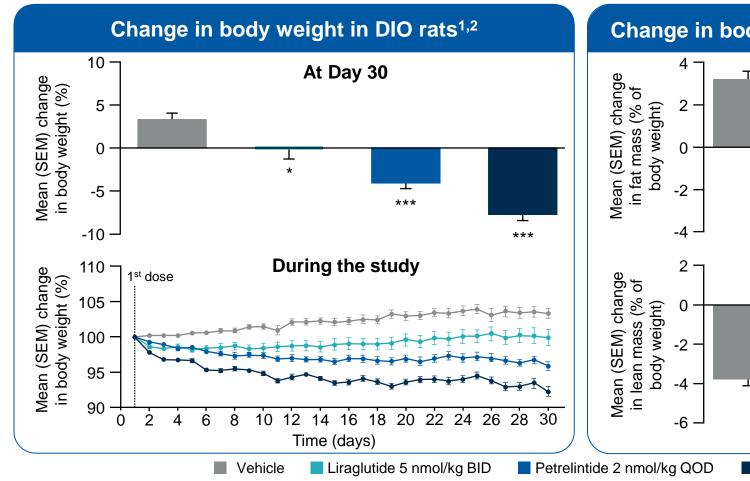


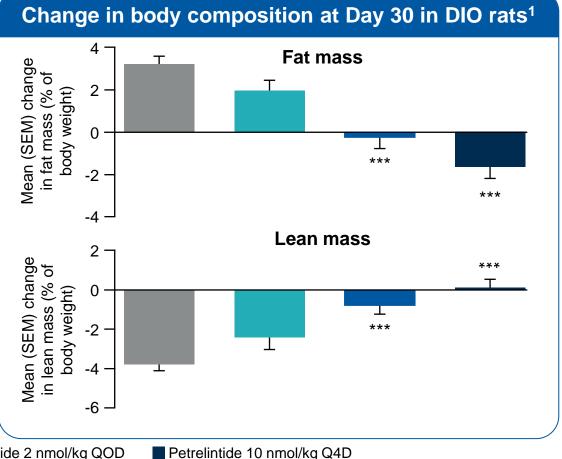


<sup>a</sup>Mediated by the effect of amylin on the CNS; <sup>b</sup>Demonstrated pharmacologically with several amylin analogs in pre-clinical studies, including with petrelintide. Sources: <sup>1</sup>Hayes et al. Annu Rev Nutr 2014;34:237–260; <sup>2</sup>Figure adapted from Mathiesen et al. Eur J Endocrinol 2022;186(6):R93–R111; <sup>3</sup>Roth et al. Proc Natl Acad Sci U S A 2008;105(20):7257–7262; <sup>4</sup>Vestergaard et al. Poster presented at ADA 84<sup>th</sup> Scientific Sessions, June 21–24, 2024, Orlando, FL. [1662-P]. CNS=central nervous system; Gl=gastrointestinal; GLP-1=glucagon-like peptide-1.

### In DIO rats, petrelintide reduced fat mass and preserved lean mass vs vehicle and a GLP-1RA





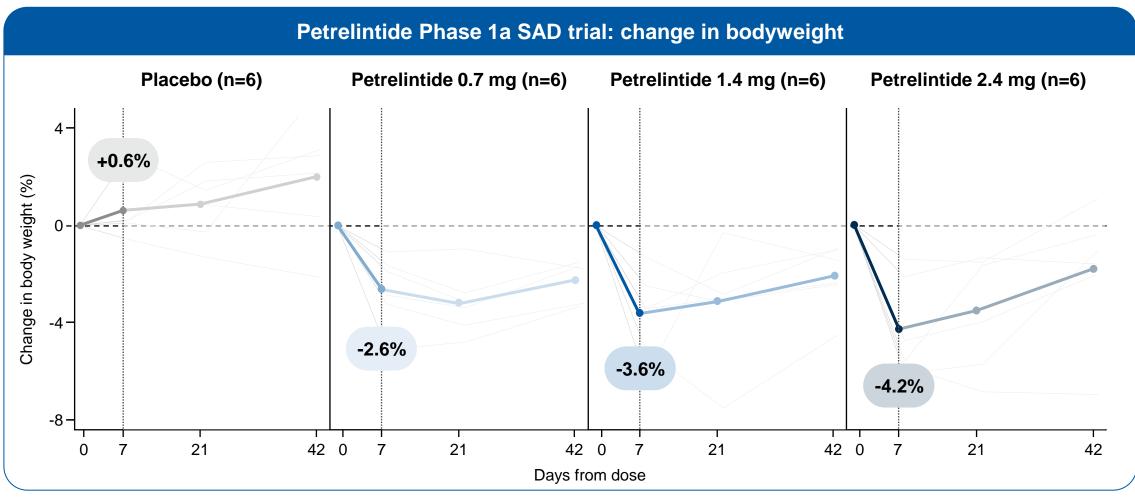


Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe. Sources: <sup>1</sup>Vestergaard et al. Poster presented at ADA 84<sup>th</sup> Scientific Sessions, June 21–24, 2024, Orlando, FL. [1662-P]; <sup>2</sup>Data on file. BID=twice daily; DIO=diet-induced obese; GLP-1RA=glucagon-like peptide-1 receptor agonist; Q4D=every 4 days; QOD=every other day; SEM=standard error of the mean.

<sup>\*</sup>p<0.05, \*\*\*p<0.001 vs vehicle.

# Dose-dependent weight loss was observed with single doses of petrelintide

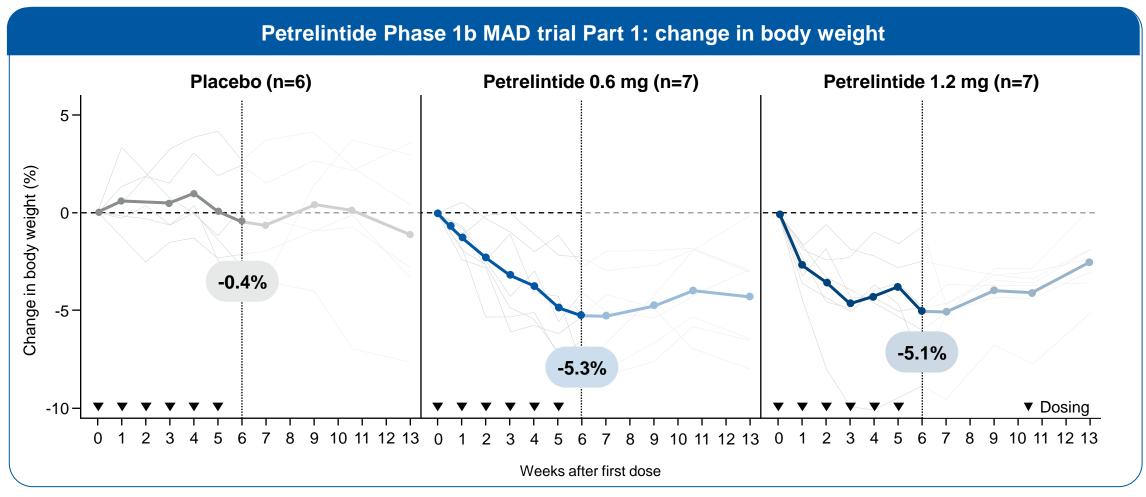




Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe. Source: Figure adapted from Brændholt Olsen et al. Poster 92-LB. Presented at ADA 83<sup>rd</sup> Scientific Sessions, June 23–26, 2023, San Diego, CA. SAD=single ascending dose.

# Consistent weight loss was observed after 6 weeks of treatment, with no dose escalation





Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe. Source: Figure adapted from Brændholt Olsen et al. Poster presented at ObesityWeek, October 14–17, 2023, Dallas, TX. MAD=multiple ascending dose.

### Petrelintide was well tolerated with no serious or severe TEAEs and no withdrawals from the trial



### Phase 1b MAD trial Part 1: TEAEs reported with petrelintide and placebo

| Number of participants (events)    | Placebo<br>(n=6) | Petrelintide 0.6 mg<br>(n=7) | Petrelintide 1.2 mg<br>(n=7) |
|------------------------------------|------------------|------------------------------|------------------------------|
| Total AEs                          | 5 (28)           | 6 (23)                       | 7 (29)                       |
| Mild                               | 5 (24)           | 6 (23)                       | 7 (27)                       |
| Moderate                           | 3 (4)            | 0                            | 1 (2)                        |
| Severe                             | 0                | 0                            | 0                            |
| Serious                            | 0                | 0                            | 0                            |
| Metabolism and nutrition disorders | 1 (1)            | 6 (9)                        | 6 (8)                        |
| GI disorders                       | 3 (7)            | 2 (6)                        | 5 (9)                        |

- All drug-related TEAEs were mild and transient, and most had an onset within two days of the first dose
- Nausea occurred in three participants on petrelintide, with one also reporting vomiting; no other participants reported vomiting
- No injection-site reactions were reported, and no participants developed anti-drug antibodies



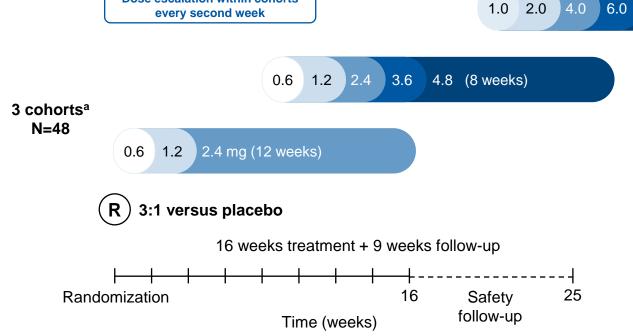
### **Trial design: Petrelintide Phase 1b MAD Part 2**

### A randomized, double-blind, placebo-controlled, Phase 1b, MAD trial of petrelintide – Part 2<sup>1,2</sup>

Aim: to evaluate the safety, tolerability, PK and PD of multiple SC doses of petrelintide, with dose escalation



- Adults 18–64 vears
- Body weight ≥70 kg
- HbA1c <6.5%
- BMI 27.0–39.9 kg/m<sup>2</sup>



### **Primary endpoint:**

9.0 (6 weeks)

Incidence of TEAEs

### **Key secondary endpoints:**

- PK parameters of petrelintide
- PD parameters related to an MTM
- Body weight related endpoints

Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe.

Dose escalation within cohorts

<sup>a</sup>Safety evaluation occurred after 4 weeks of treatment at the target dose for each cohort. Initiation of the next, higher dose cohort only occurred following safety evaluation for the previous cohort. Sources: <sup>1</sup>ClinicalTrials.gov (NCT05613387), accessed October 2024: <sup>2</sup>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.

# Most participants were male and had a BMI at the lower end of the eligible range



### Petrelintide Phase 1b MAD trial Part 2: baseline characteristics<sup>1,2</sup>







Age



Weight



**BMI** 

**79%** of participants were **male** 

Median 49 years

Median 92.4 kg

Median 29.2 kg/m<sup>2</sup>

(eligible range: 27.0–39.9 kg/m<sup>2</sup>)

Baseline characteristics were balanced across the dose cohorts<sup>2</sup>

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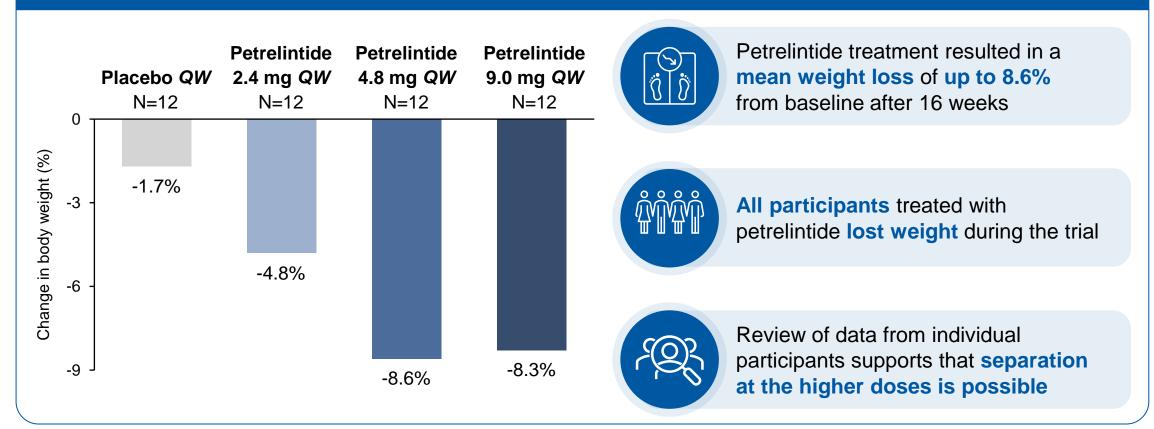
Sources: ¹Zealand Pharma. Press release 20 June 2024. Available from: https://www.globenewswire.com/news-release/2024/06/20/2901879/0/en/Zealand-Pharma-announces-positive-topline-results-from-the-Phase-1b-16-week-multiple-ascending-dose-clinical-trial-with-long-acting-amylin-analog-petrelintide.html, accessed July 2024; ²Data on file.

BMI=body mass index; MAD=multiple ascending dose.

### Substantial weight loss was observed at 16 weeks...







N represents cohort size at randomization.

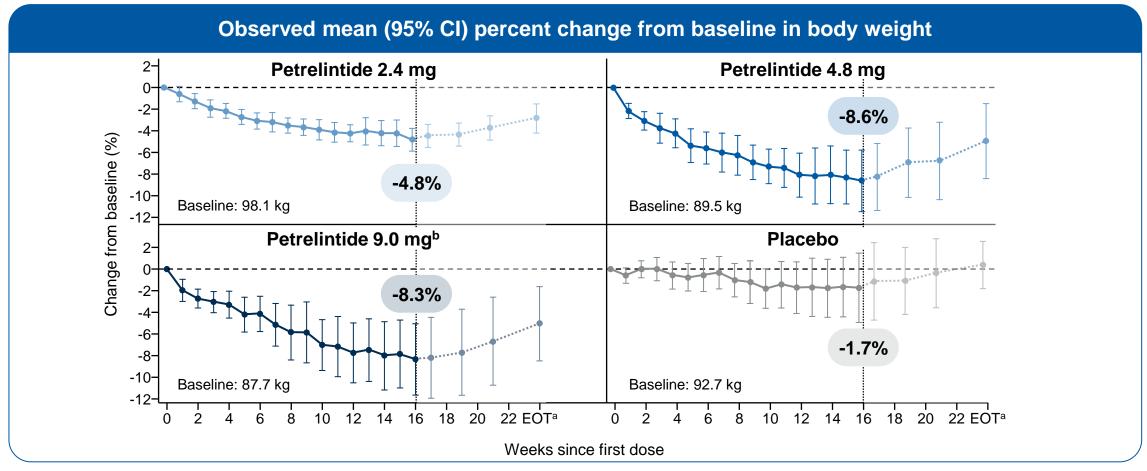
Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe.

Sources: ¹Zealand Pharma. Press release 20 June 2024. Available from: https://www.globenewswire.com/news-release/2024/06/20/2901879/0/en/Zealand-Pharma-announces-positive-topline-results-from-the-Phase-1b-16-week-multiple-ascending-dose-clinical-trial-with-long-acting-amylin-analog-petrelintide.html, accessed July 2024; ²Data on file.

DG=dose group; MAD=multiple ascending dose; QW=once-weekly.

### ...with continued body weight loss expected with longer treatment duration





Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe.

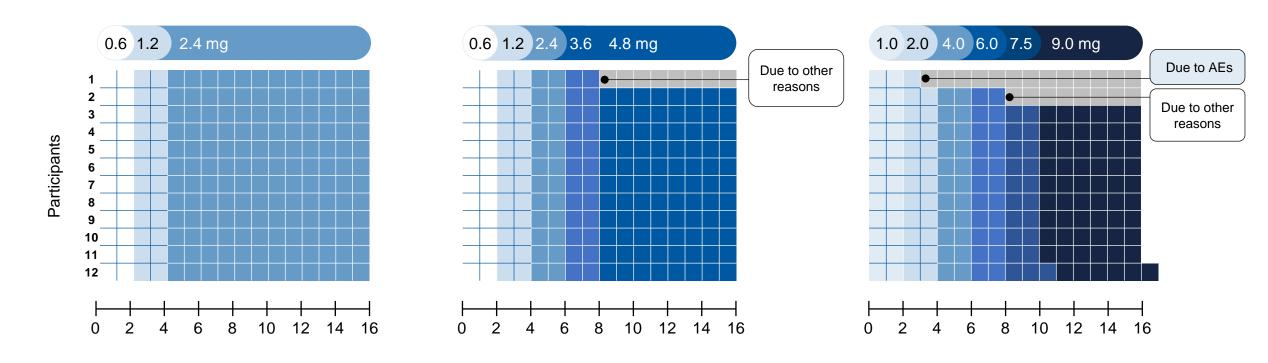
<sup>a</sup>EOT includes measurements at the EOT visit, performed at 24 or 25 weeks after dosing, and also performed for participants discontinuing treatment early; <sup>b</sup>One participant had one extra week at 7.5 mg, and thereby only five weeks on maintenance dose at Week 16. After Week 16, this 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 rates of study treatment completion and adherence to dose escalation within cohorts

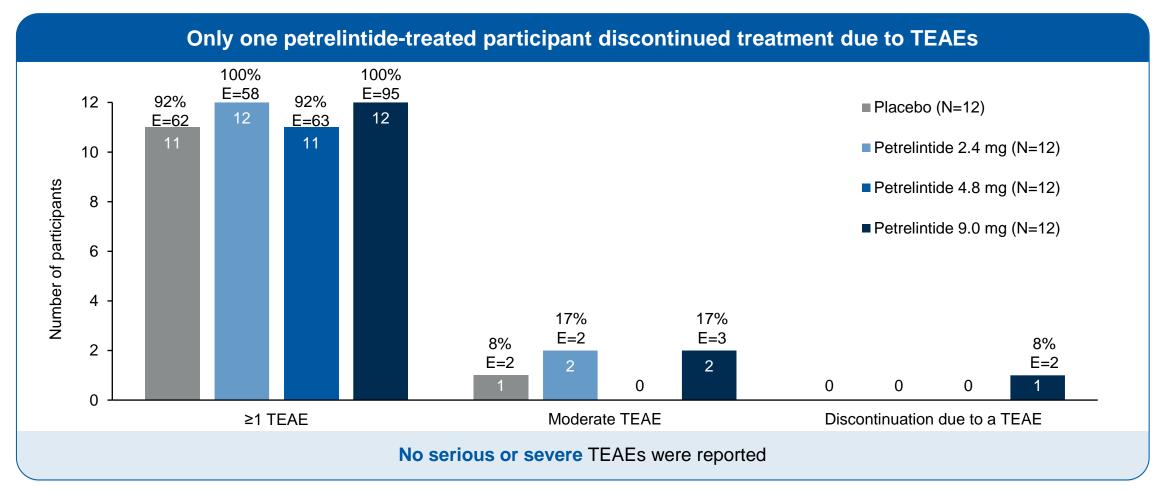




- Three participants discontinued petrelintide: one due to AEs, one to focus on recovery from a cold, and one due to personal reasons
- One participant in the 9.0 mg arm had an extra week at 7.5 mg (due to tolerability)
- The remaining participants followed dose escalation steps within cohorts

### Vast majority of TEAEs reported by petrelintidetreated patients were mild





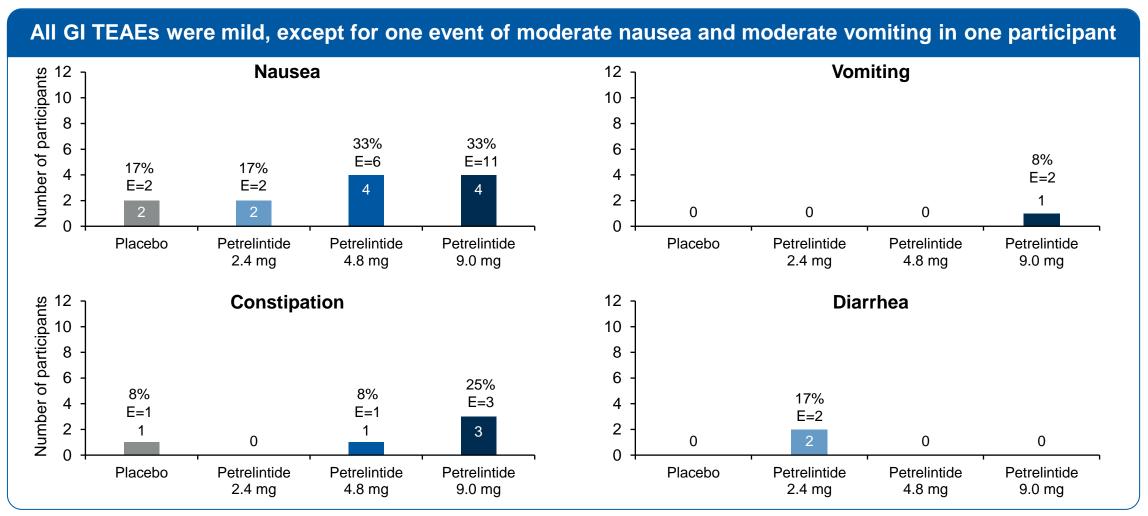
Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe. Source: Data on file. Data presented at ObesityWeek 2024 in San Antonio, Texas.

5 moderate AEs reported by petrelintide exposed participants: pausea, vomiting, pasopharyngitis, acute sinusitis, back pain

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





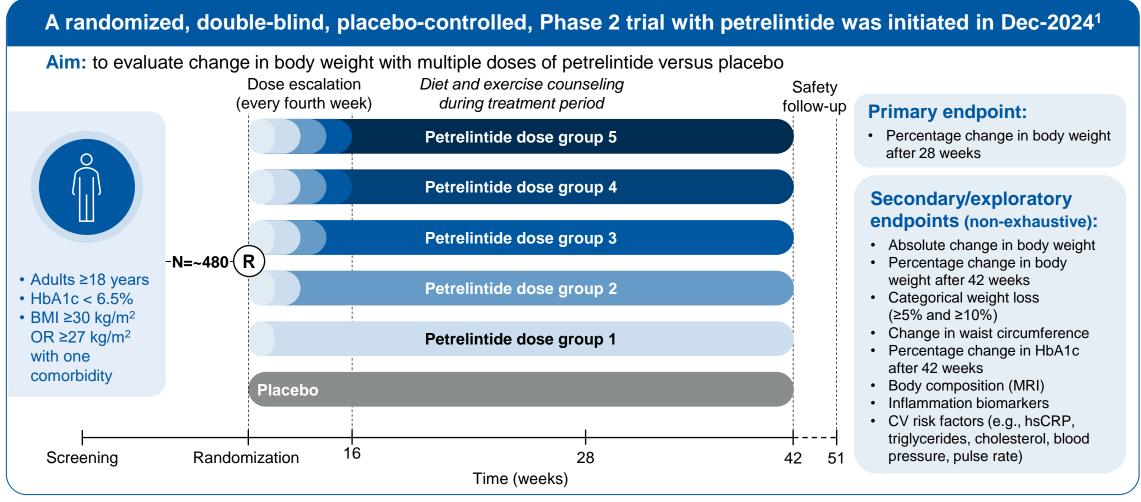
Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe. Source: Data on file. Data presented at ObesityWeek 2024 in San Antonio, Texas.

N=12 in each treatment group.

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

# Petrelintide is in development as monotherapy in a comprehensive Phase 2 trial (ZUPREME-1)...



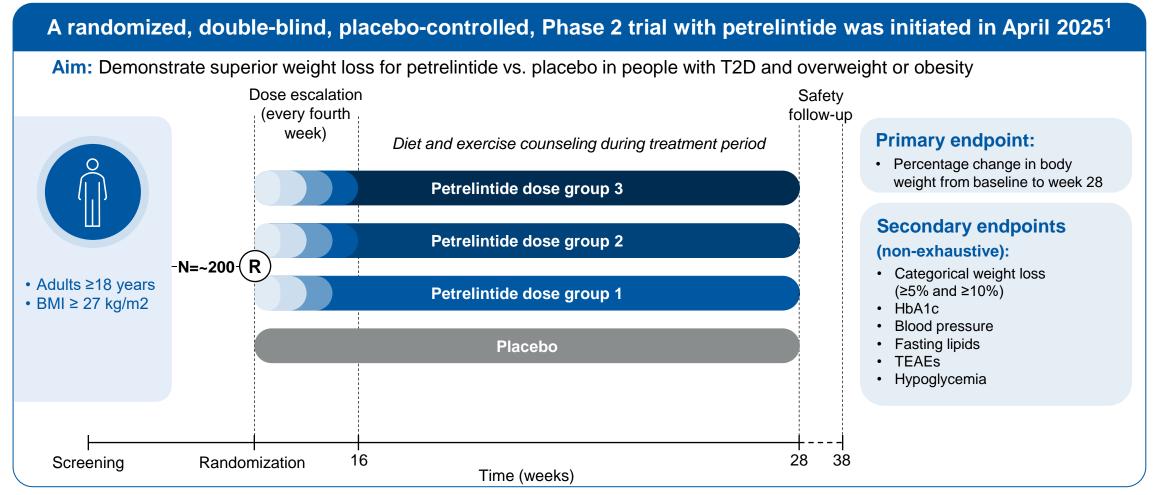


Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe. Source: ¹ClinicalTrials.gov (NCT06662539), accessed April 2025

 $BMI = body \ mass \ index; \ HbA1c = glycated \ hemoglobin; \ MRI = magnetic \ resonance \ imaging; \ hsCRP = high-sensitivity \ C-reactive \ protein.$ 

# ... and in a Phase 2 trial in people with overweight/obesity and T2D (ZURPREME-2)





Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe. Source: ¹ClinicalTrials.gov (NCT06926842), accessed April 2025

T2D=type 2 diabetes; BMI=body mass index; QW=once-weekly; HbA1c=hemoglobin A1C (glycated hemoglobin); TEAE=treatment emergent adverse event.



### Our vision is to establish the leading amylin-based weight management franchise with petrelintide

Potential to address the unmet medical needs of the majority of people with overweight and obesity

### Unmet medical needs...



Alternative mechanisms of action to provide new treatment options



**Improved GI tolerability** for a better patient experience and treatment persistence



Improved effect on obesity-related comorbidities



**Greater** weight loss **efficacy** for the segment of patients who need most weight loss

### ...being targeted with petrelintide

Petrelintide monotherapy as a foundational therapy targeting:

- ~15-20% weight loss
- non-incretin mechanism
- substantially improved GI tolerability
- muscle preservation

#### Petrelintide in combinations:

 with CT-388 for people who need more weight loss and/or better glycemic control

# related indications Rapidly expanding

Collaboration and license agreement with Roche for petrelintide, including co-development and co-commercialization in the U.S. and Europe.

### A transformative collaboration and license agreement<sup>1</sup> to unlock the full potential of petrelintide





| True partnership agreement                            | Shared vision for petrelintide as a future foundational therapy for weight management Co-development and co-commercialization (up to 50% in U.S. and Europe) |            |
|---|--|------------|
| Important synergies and complementary capabilities    | Combining Zealand's >25 years of peptide expertise with Roche's global R&D, manufactu<br>commercial capabilities   | uring, and |
| Maximizing the full value potential of petrelintide   | Addressing different high unmet medical needs, both as monotherapy and in combination agents (e.g., CT-388), to reach as many patients as possible           | with other |
| position of positional                                | Accelerating and expanding the opportunities with petrelintide in weight management and ndications   | d related  |
| Up to \$5.3 billion in total consideration to Zealand | 61.65 billion in upfront (of which \$1.4 billion due at closing and \$250 million in anniversary   | у          |
|   | Jp to \$1.2 billion in development milestone payments  |            |
|   | Jp to \$2.4 billion in sales-based milestone payments  |            |
| Economics and upside further                          | 50/50 profit sharing in U.S. and Europe  |            |
| enhanced  | Royalties on net sales in the rest of the world  |            |
|   | 3350 million to Roche from Zealand Pharma for CT-388 in the first combination product  |            |

# A potential first-in-class GLP-1R/GLP-2R dual agonist for obesity and low-grade inflammation



### **Design of molecule**

**Dapiglutide** is derived from a GLP-2 peptide backbone with amino acid substitutions to 'dial in' GLP-1R activity



GLP-1 component reduces body weight and GLP-2 has potential for additional anti-inflammatory effects<sup>1</sup>



Designed with higher potency towards the GLP-1R while retaining activity on the GLP-2R<sup>2</sup>



**Long-acting** with a half-life (123–129 hours) that is suitable for **once-weekly administration**<sup>3</sup>

### Positioning opportunities and differentiation



Obesity – pursuing ≥20% weight loss



**Safety and tolerability** – similar to other GLP-1RA-based weight-loss medications



Cardiovascular benefits – potential cardioprotective benefits from GLP-1 agonism and additional antiinflammatory effect from GLP-2 agonism



**Comorbidities** – potential for regenerative effects to address organ damage associated with low-grade inflammation, such as MASH and Alzheimer's disease

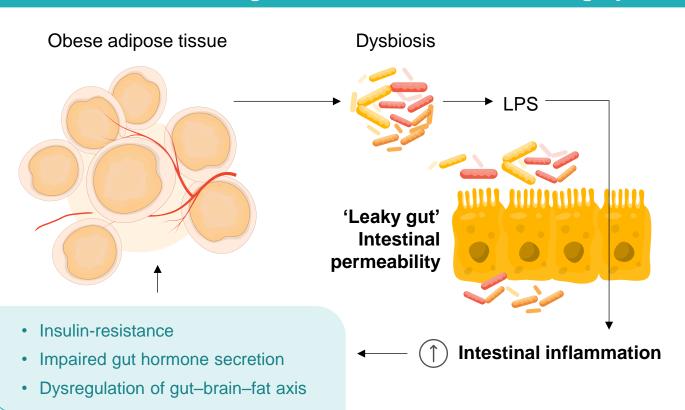
Sources: ¹Drucker & Yusta. Annu Rev Physiol 2014;76:561–583; ²Reiner et al. JPEN J Parenter Enteral Nutr 2022;46(5):1107–1118; ³Data presented by Agersnap at the 82<sup>nd</sup> ADA Scientific Sessions, June 3–7, 2022. GLP-1=glucagon-like peptide-1; GLP-1R=glucagon-like peptide-1 receptor; GLP-2=glucagon-like peptide-2; GLP-2R=glucagon-like peptide-2 receptor; MASH=metabolic dysfunction-associated steatohepatitis (formerly, non-alcoholic steatohepatitis, or NASH).

Intellectual property: Composition of matter, patent expiry in 2037. Patent-term extension up to 5 years, i.e. 2042. Potential rights beyond 2042 based on patent applications and additional elements.

### People with obesity have increased low-grade inflammation, which drives several comorbidities



Excess fat storage can trigger low-grade systemic inflammation through reduced intestinal barrier integrity<sup>1</sup>



### Obesity-related low-grade inflammation can result in:



CVD as increased inflammation drives residual risk in people with CVD<sup>2</sup>



Liver disease due to abnormal accumulation of triglycerides in the liver<sup>3</sup>



#### **Neuro-inflammation**

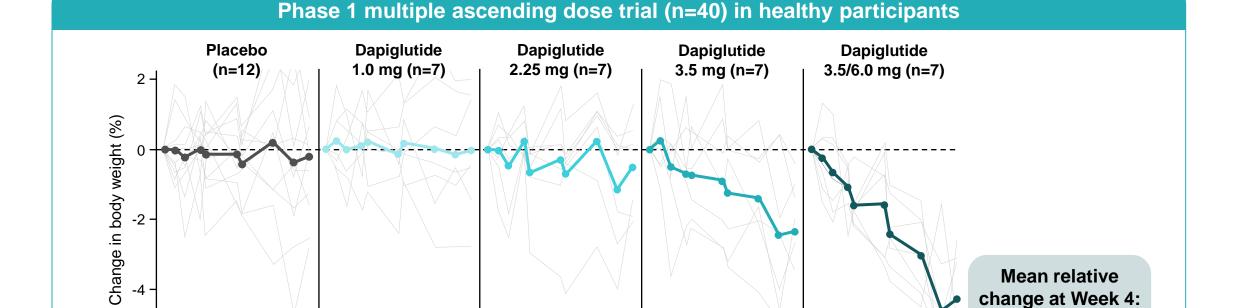
due to excess circulating proinflammatory cytokines and changes in the integrity of the blood–brain barrier<sup>4</sup>

Sources: <sup>1</sup>Figure adapted from Vetrani et al. Nutrients 2022;14(10):2103, used under the Creative Commons Attribution (CC BY 4.0) license (https://creativecommons.org/licenses/by/4.0/). The figure has been reformatted. The publication is available at https://doi.org/10.3390/nu14102103; <sup>2</sup>Ridker et al. Lancet 2023;401(10384):1293–1301; <sup>3</sup>Luo & Lin. Immun Inflamm Dis 2021;9(1):59–73; <sup>4</sup>Salas-Venegas et al. Front Integr Neurosci 2022;16:798995. CVD=cardiovascular disease; LPS=lipopolysaccharides.

3

# Dapiglutide showed dose-dependent mean weight loss of up to 4.3% over 4 weeks





Dapiglutide was generally well-tolerated with no severe or serious AEs, no withdrawals due to AEs, and no observation of anti-drug antibodies

Weeks from first dose

Source: Figures adapted from data presented by Agersnap at the 82<sup>nd</sup> ADA Scientific Sessions, June 3–7, 2022, New Orleans, LA. AE=adverse event.

-4.3%

▼ Dosing

### Doses investigated in DREAM trial were at lower end of therapeutic range



Investigator-led trial DREAM evaluating effects on body weight, gut permeability and inflammation<sup>1</sup>

### **Trial design**

- N=54, men and women aged 18-75 years (BMI ≥30 kg/m²)
- Duration = 12 weeks
- Dose strengths = up to 6.0 mg (similar to 4-week MAD)<sup>2</sup>
- No lifestyle interventions, such as diet or exercise

### **Purpose of trial**

- Gain key mechanistic insights into the effects of low doses of GLP-1/GLP-2 receptor dual agonist dapiglutide
- Evaluate the potential for weight loss and assess the safety and tolerability profile of dapiglutide

### Low doses investigated at the lower end of the therapeutic window in obesity setting

### **Topline data**

- Mean weight loss of up to 4.3% with dapiglutide treatment
- Results in line with outcomes observed with shorter term treatment using lower doses of incretin-based therapies
- Dapiglutide assessed to be well-tolerated with fewer TEAEs related to the GI system compared to what have been reported from other trials with incretin-based therapies
- No TEAEs related to the GI system led to treatment discontinuation



Data on inflammatory markers, as well as data from gut biopsies, to be presented at future scientific meeting

### We expect to report topline results from the 28week Phase 1b trial in Q2 2025...



The 13-week Phase 1b trial<sup>1</sup> evaluated higher doses of dapiglutide than prior clinical trials<sup>a</sup>

Data from Part 1 of Phase 1b trial to be presented at ADA in June 2025



 Mean placebo-adjusted weight loss of up to 8.3% after 13 weeks



- Total of 54 participants, 85% male with a median baseline BMI of 30.0 kg/m<sup>2</sup>
- 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
- 2/54 discontinued treatment due to GI AEs

Next steps include topline results from 28-week trial<sup>2</sup> and initiation of Phase 2b trial

Topline results from Part 2 of Phase 1b trial in Q2 2025



- Part 2 of the Phase 1b trial includes a higher dose cohort with 28 weeks treatment
- Monthly dose escalation

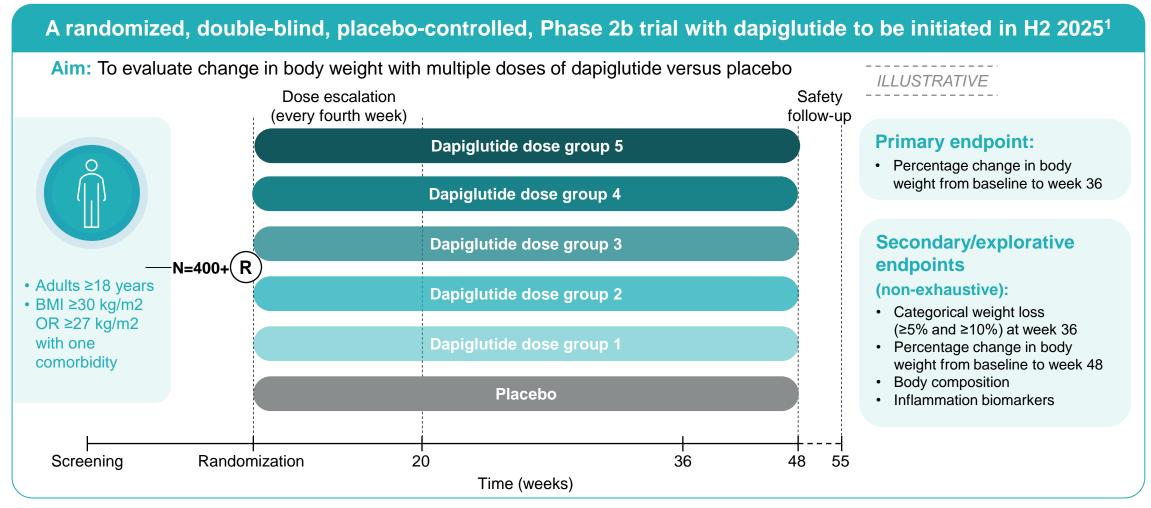


Phase 2 trial expected to initiate in H2 2025

<sup>a</sup>DREAM, an investigator-led mechanistic trial with dapiglutide: ClinicalTrials.gov (NCT05788601), and the 4-week Phase 1a clinical trial: ClinicalTrials.gov (NCT04612517). Sources: <sup>1</sup>Zealand Pharma company announcement No. 44/2024, September 9, 2024; <sup>2</sup>ClinicalTrials.gov (NCT06000891), accessed March 2025. BMI=body mass index; GI=gastrointestinal; AE=adverse event.

# ...and initiate a Phase 2 trial in people with overweight/obesity





Source: 1Data on file.

BMI=body mass index; EOT=end of treatment

## Survodutide<sup>a</sup> is in Phase 3 development for obesity and MASH with first- and best-in-class potential





### **Design of molecule**

**Survodutide** is a 29-amino-acid peptide, based on the hormone **oxyntomodulin** with dual agonism at GCG and GLP-1 receptors



MoA reduces body weight by increasing energy expenditure and regulating appetite<sup>1</sup>



Deliberately designed with **strong bias towards GLP-1** receptor (8:1 receptor bias vs glucagon)<sup>2</sup>



Extended half-life for **once-weekly administration** achieved by amino acid substitutions<sup>2</sup>

### Positioning opportunities and differentiation



Obesity – potential for ~20–25% weight loss and improved glycemic control



Safety and tolerability – similar to other GLP-1RA-based weight-loss medications



**Cardiovascular benefits** – potential benefits driven by GLP-1RA



MASH – potential for important benefit in MASH with direct effect of glucagon on the liver

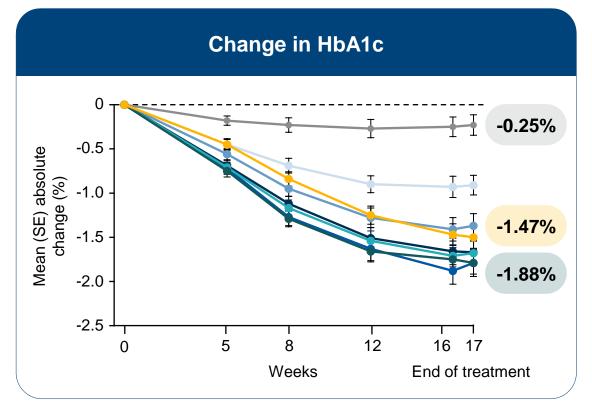
<sup>&</sup>lt;sup>a</sup>Survodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally. Sources: <sup>1</sup>Wynne et al. Int J Obes (Lond) 2006;30(12):1729–1736; <sup>2</sup>Zimmermann et al. Mol Metab 2022;66;101633.

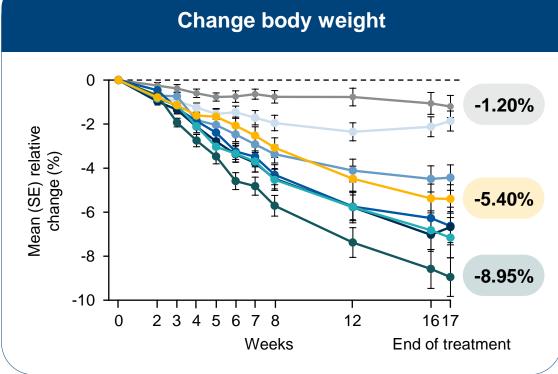
GCG=glucagon; GLP-1=glucagon-like peptide-1; GLP-1RA=glucagon-like peptide-1 receptor agonist; MoA=mechanism of action; MASH=metabolic dysfunction-associated steatohepatitis (formerly, non-alcoholic steatohepatitis, or NASH). Intellectual property: Composition of matter, patent expiry in 2034. Patent-term extension up to 5 years, i.e. 2039. Potential rights beyond 2039 based on patent applications and additional elements.

## In a 16-week Phase 2 trial in T2DM, survodutide effectively reduced HbA1c and body weight









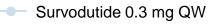
The safety and tolerability

profile was as expected and in line with

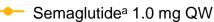


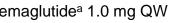
Survodutide 1.8 mg QW

Survodutide 1.8 mg BIW



Survodutide 2.7 mg QW





Survodutide 1.2 mg BIW Survodutide 0.9 mg QW increasing doses of GLP-1R agonists

Body weight at baseline was 93.0-100.1 kg and HbA1c at baseline was 7.9-8.2%.

Survodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally. Source: Figures adapted from Rosenstock. Presentation at ObesityWeek, November 1-4, 2022, San Diego, CA.

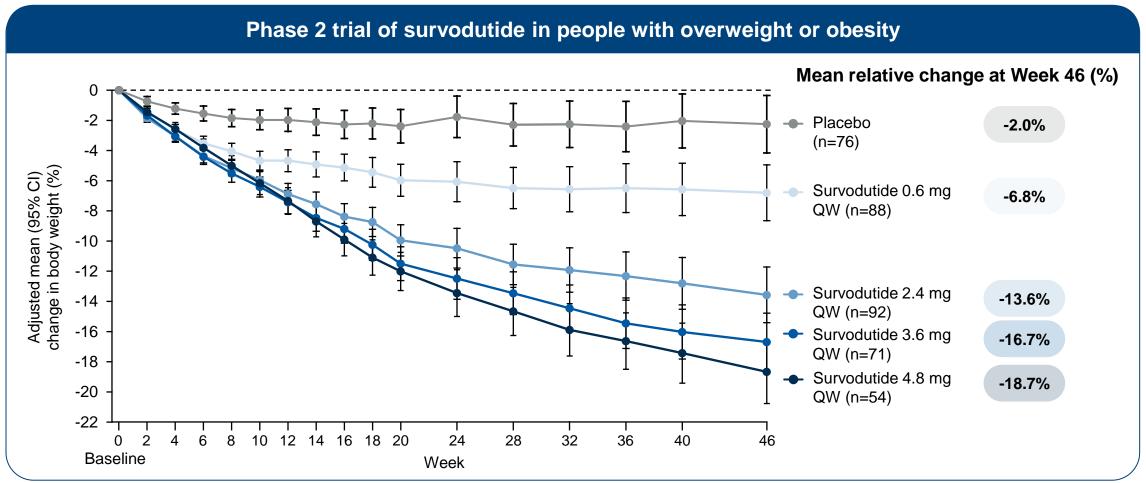
BIW=twice-weekly; GLP-1R=glucagon-like peptide-1 receptor; HbA1c=hemoglobin A1c; QW=once-weekly; SE=standard error; T2DM=type 2 diabetes mellitus.

<sup>&</sup>lt;sup>a</sup>The semaglutide arm was open-label.

## In a Phase 2 trial in obesity, survodutide dosedependently reduced body weight by up to 18.7%







Survodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally. Source: Figure adapted from Le Roux et al. Oral presentation (51-OR) at ADA 83<sup>rd</sup> Scientific Sessions, June 23–26, 2023, San Diego, CA. Analysis based on dose reached at the end of treatment regardless of the dose assigned at randomization. CI=confidence interval; QW=once-weekly.

## No unexpected safety findings in the Phase 2 obesity trial with survodutide





- As expected, GI disorders were the most frequent drug-related AEs
- Most treatment discontinuations occurred during the rapid dose escalation phase (up to Week 20) and may be mitigated with more gradual dose-escalation

| TEAE, n (%) <sup>a</sup>                   | Survodutide<br>0.6 mg (n=77) | Survodutide<br>2.4 mg (n=78) | Survodutide<br>3.6 mg (n=77) | Survodutide<br>4.8 mg (n=77) | Survodutide<br>total (n=309) | Placebo<br>(n=77) |
|--|------------------------------|------------------------------|------------------------------|------------------------------|------------------------------|-------------------|
| Any TEAE                                   | 70 (90.9)                    | 70 (89.7)                    | 71 (92.2)                    | 70 (90.9)                    | 281 (90.9)                   | 58 (75.3)         |
| Nausea <sup>b</sup>                        | 26 (33.8)                    | 51 (65.4)                    | 48 (62.3)                    | 49 (63.6)                    | 174 (56.3)                   | 15 (19.5)         |
| Vomiting <sup>b</sup>                      | 7 (9.1)                      | 23 (29.5)                    | 26 (33.8)                    | 27 (35.1)                    | 83 (26.9)                    | 4 (5.2)           |
| Diarrheab                                  | 14 (18.2)                    | 22 (28.2)                    | 18 (23.4)                    | 15 (19.5)                    | 69 (22.3)                    | 8 (10.4)          |
| Constipationb                              | 9 (11.7)                     | 17 (21.8)                    | 19 (24.7)                    | 20 (26.0)                    | 65 (21.0)                    | 4 (5.2)           |
| Leading to treatment discontinuation       | 15 (19.5)                    | 20 (25.6)                    | 19 (24.7)                    | 22 (28.6)                    | 76 (24.6)                    | 3 (3.9)           |
| GI-related                                 | 5 (6.5)                      | 13 (16.7)                    | 13 (16.9)                    | 20 (26.0)                    | 51 (16.5)                    | 1 (1.3)           |
| Serious                                    | 1 (1.3)                      | 2 (2.6)                      | 6 (7.8)                      | 4 (5.2)                      | 13 (4.2)                     | 5 (6.5)           |
| Investigator defined,<br>drug-related TEAE | 47 (61.0)                    | 66 (84.6)                    | 62 (80.5)                    | 62 (80.5)                    | 237 (76.7)                   | 29 (37.7)         |
| Serious, drug-related TEAE                 | 0 (0.0)                      | 0 (0.0)                      | 2 (2.6)                      | 0 (0.0)                      | 2 (0.6)                      | 0 (0.0)           |

<sup>&</sup>lt;sup>a</sup>Survodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally.

Source: Table adapted from Le Roux et al. Oral presentation (51-OR) at ADA 83rd Scientific Sessions, San Diego, June 23–26, 2023.

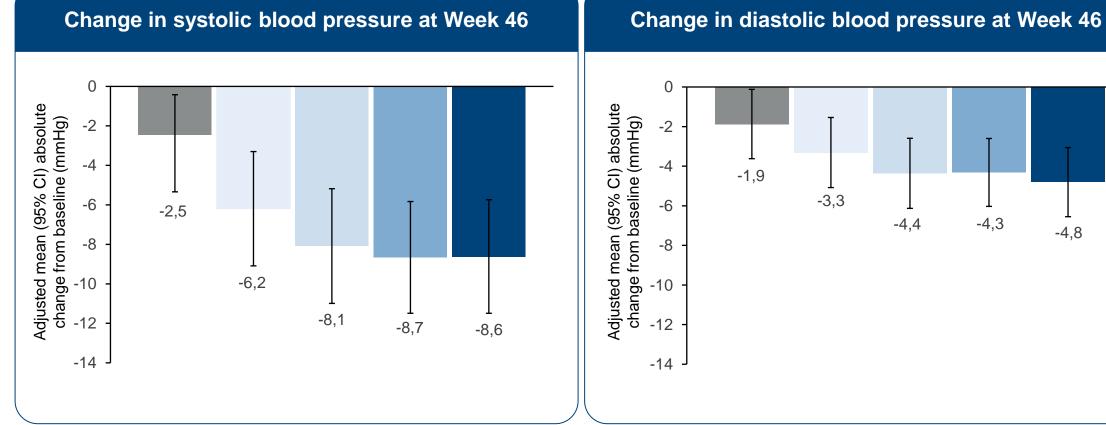
AE=adverse event; GI=gastrointestinal; TEAE=treatment-emergent adverse event.

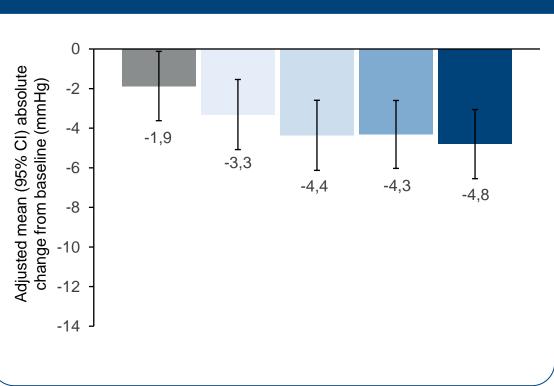
<sup>&</sup>lt;sup>b</sup>Based on the treated set and presented according to planned treatment; <sup>b</sup>TEAEs listed according to preferred term and occurred in ≥15% patients in any treatment arm.

## Survodutide reduced blood pressure by up to 8.6 mmHg (systolic) and up to 4.8 mmHg (diastolic)









Placebo

Survodutide 0.6 mg QW

Survodutide 2.4 mg QW

Survodutide 3.6 mg QW

Survodutide 4.8 mg QW

Mean blood pressure at baseline across cohorts: 122.6-127.5 mmHg for systolic blood pressure; 80.5-82.4 mmHg for diastolic blood pressure. Survodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally. Source: Figures adapted from Le Roux. Presentation at the 59th EASD Annual Meeting, October 2–6, 2023, Hamburg, Germany. CI=confidence interval; QW=once-weekly.

# The Phase 3 program with survodutide<sup>a</sup> in obesity is ongoing and is fully enrolled





#### Inclusion criteria

### Study design

### **Primary endpoint**

### SYNCHRONIZETM-11

Efficacy and safety in patients with obesity without T2DM

- HbA1c <6.5% (no history of diabetes)</li>
- BMI ≥30 or BMI ≥27 with comorbidities<sup>b</sup>

- N=727
- 1:1:1 ratio (3.6 mg, 6.0 mg, or placebo)
- · Trial duration: 76 weeks

### **SYNCHRONIZE**<sup>TM</sup>**-2**<sup>2</sup>

Efficacy and safety in patients with obesity and T2DM

- HbA1c ≥6.5% and <10%</li>
- BMI ≥27
- T2DM managed with diet and exercise alone or with stable pharmacological treatment
- N=756
- 1:1:1 ratio (3.6 mg, 6.0 mg or placebo)
- Trial duration: 76 weeks
- Percentage change in body weight from baseline to Week 76
- Achievement of body weight reduction
   ≥5% from baseline to Week 76

### SYNCHRONIZETM-CVOT3

Long-term CV safety in patients with obesity and established CVD/CKD or risk factors for CVD

- BMI ≥27 with CVD and/or at least two weight-related risk factors for CVD, or
- BMI ≥30 with CVD/CKD and/or at least two weight-related factors for CVD
- N=5,550
- 1:1:1 ratio (3.6 mg, 6.0 mg or placebo)
- Trial duration: up to 114 weeks
- Time to first occurrence of any of five major adverse cardiac events (5P-MACE) to demonstrate noninferiority

Inclusion criteria for all three trials include age ≥18 years. 5P-MACE includes cardiovascular death, non-fatal stroke, non-fatal myocardial infarction, ischemia-related coronary revascularization or heart failure.

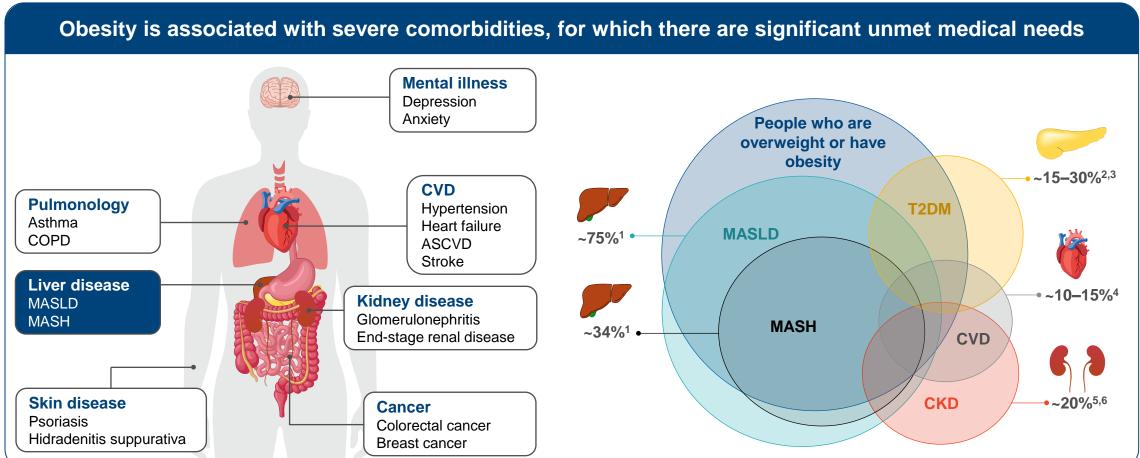
aSurvodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally; bComorbidities comprise dyslipidemia, hypertension, obstructive sleep apnea, and others.

Sources: ¹ClinicalTrials.gov (NCT06066515), accessed April 2025; ²ClinicalTrials.gov (NCT06066528), accessed April 2025; ³ClinicalTrials.gov (NCT06066515), accessed April 2025; ³ClinicalTrials.gov (NCT06066528), accessed April 2025; accessed April 2025;

## There is a significant overlap between obesity and liver disease







Estimates of overlap of co-morbidities are not available in literature; approximation in figure is based on individual prevalence estimates.

Survodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally.

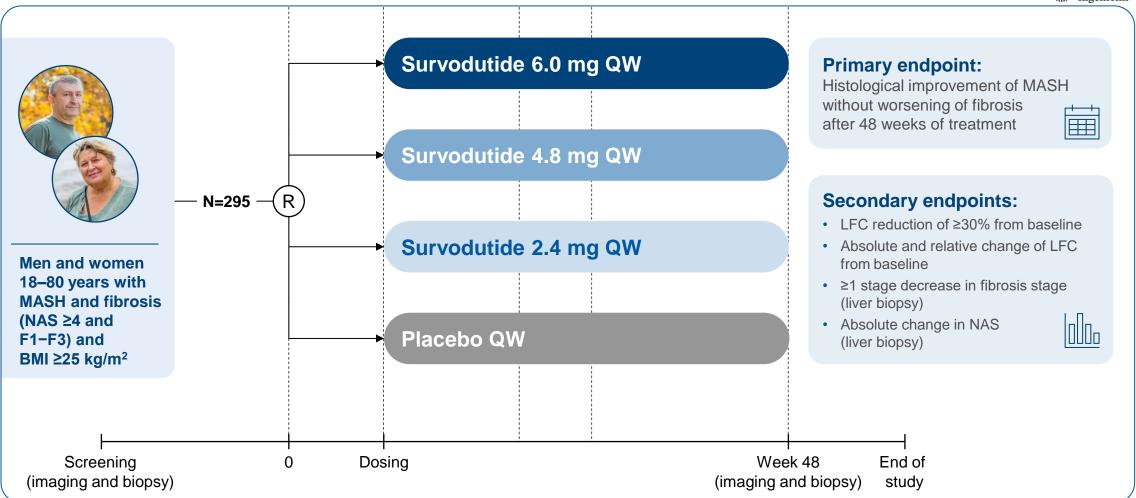
Sources: ¹Quek et al. Lancet Gastroenterol Hepatol 2023;8(1):20–30; ²Vinciguerra et al. Acta Diabetol 2013;50(3):443–449; ³Pantalone et al. BMJ Open 2017;7(11):e017583; ⁴Schienkiewitz et al. BMC Public Health 2012;12:658; ⁵Arinsoy et al. J Ren Nutr 2016;26(6):373–379; ⁶Yim & Yoo. Clin Exp Pediatr 2021;64(10):511–518.

ASCVD=atherosclerotic cardiovascular disease; CKD=chronic kidney disease; CVD=chronic obstructive pulmonary disease; CVD=cardiovascular disease; MASLD=metabolic dysfunction-associated steatotic liver disease (formerly, NAFLD, or non-alcoholic fatty liver disease); MASH=metabolic dysfunction-associated steatohepatitis (formerly, non-alcoholic steatohepatitis, or NASH); T2DM=type 2 diabetes mellitus.

## Survodutide<sup>a</sup> has been evaluated in a Phase 2 trial in MASH







<sup>&</sup>lt;sup>a</sup>Survodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally. Source: ClinicalTrials.gov (NCT04771273), accessed February 2024.

BMI=body mass index; LFC=liver fat content; NAS=NAFLD activity score; MASH=metabolic dysfunction-associated steatohepatitis (formerly NASH, or non-alcoholic steatohepatitis); QW=once-weekly.

## Survodutide<sup>a</sup> shows best-in-class potential in MASH Phase 2 trial





### 48-week biopsy-driven Phase 2 MASH trial<sup>1</sup>



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)

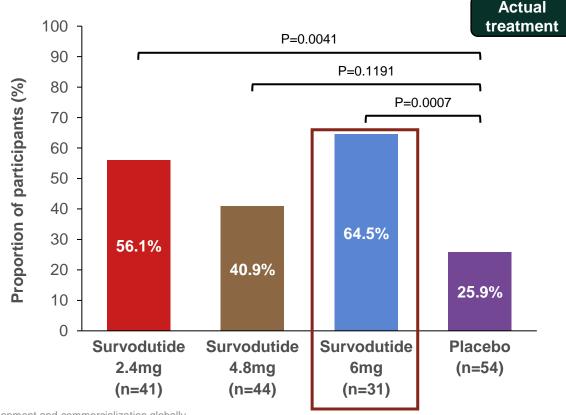


Survodutide treatment did not show unexpected safety or tolerability issues, including at the higher dose of 6.0 mg



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

## Improvement in liver fibrosis with no worsening of MASH Paired biopsy results (F2/F3)<sup>2</sup>



<sup>&</sup>lt;sup>a</sup>Survodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally.

Sources: ¹Boehringer Ingelheim press release June 7, 2024. Data presented at the EASL Congress 2024 in Milan, Italy. ²A sensitivity analysis based on participants with paired biopsy results at baseline and end of treatment. ³Boehringer Ingelheim press release October 8, 2024.

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

# The largest Phase 3 program in MASH with an incretin-based therapy is ongoing





#### Inclusion criteria

#### Diagnosis of MASH<sup>a</sup> and biopsyproven fibrosis stage F2-F3

Granted Breakthrough Therapy Designation by the U.S. FDA<sup>2</sup>

### Study design

- N=1,800
- 1:1 ratio (6.0 mg or placebo)
- Trial duration
  - Part 1: 52 weeks
  - Part 2: Up to 7 years

### **Primary endpoint**

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 liverrelated events or all-cause mortality

#### LIVERAGE-Cirrhosis<sup>3</sup>

LIVERAGE<sup>1</sup>

Efficacy and safety in patients

with MASH and fibrosis (F2/F3)

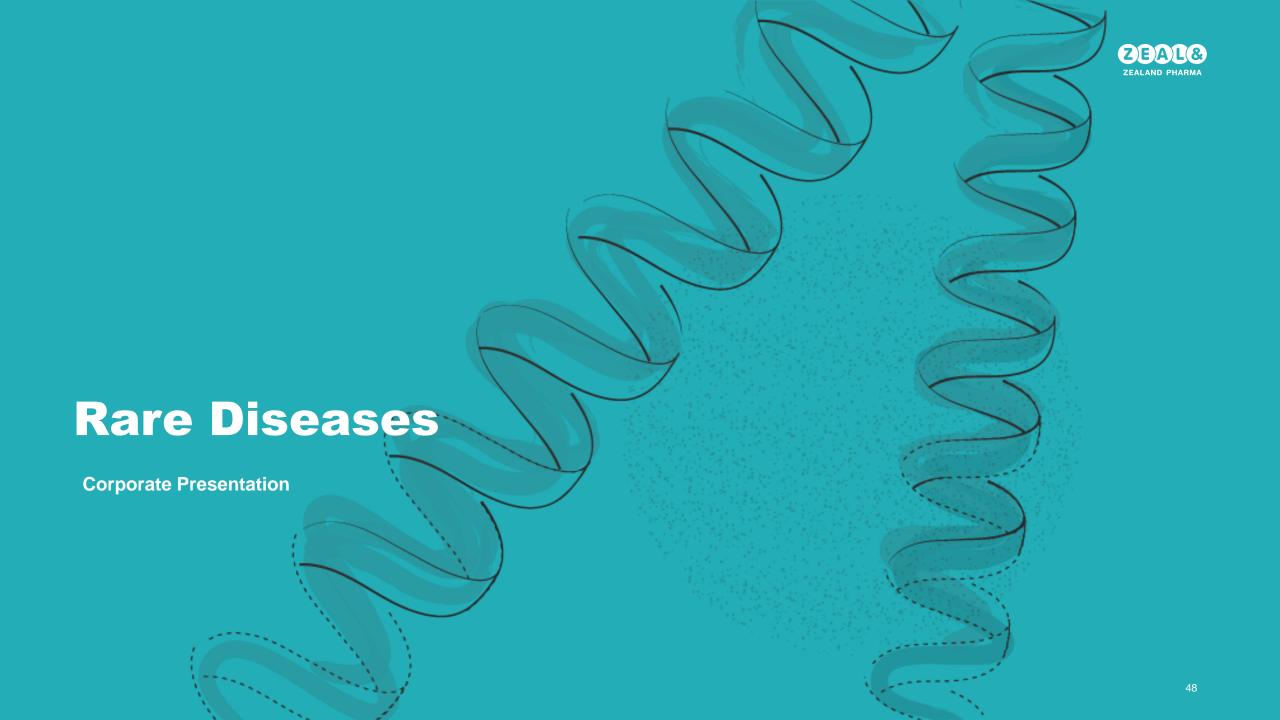
Efficacy and safety in patients with MASH and cirrhosis (F4)

- Diagnosed compensated MASH cirrhosis<sup>b</sup>
- 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

<sup>a</sup>MASH diagnosis defined by a NAS score≥4, with at least 1 point in inflammation and ballooning each. <sup>b</sup>Diagnosed 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 Survodutide is licensed to Boehringer Ingelheim from Zealand Pharma, with Boehringer solely responsible for development and commercialization globally.

Sources: ¹LIVERAGE. ClinicalTrials.gov (NCT06632444), accessed April 2025; ²Boehringer Ingelheim press release October 8, 2024; ³LIVERAGE-Cirrhosis. ClinicalTrials.gov (NCT06632447), accessed April 2025; N=number of participants; BMI=body mass index; MASH = metabolic-associated steatohepatitis; MELD = Model for End-stage Liver Disease; CSPH = clinically significant portal hypertension



## CHI is a severe, ultra-rare genetic disorder with significant impact on patients' QoL



## There is a significant unmet need for an effective treatment



#### CHI is an ultra-rare disease in newborns and children

- 1 in 28-50,000 newborns per year are diagnosed with genetically determined CHI in the US and EU<sup>1,2</sup>
- CHI can cause serious episodes of hypoglycemia during childhood<sup>2,3</sup>



## Persistent episodes of hypoglycemia may result in brain damage

- Hypoglycemia can cause seizures in ~50% of the patients<sup>4</sup>
- Lack of proper management within days can lead to permanent brain injury and neurocognitive impairment<sup>3,4</sup>



### Significant impact on the patient and caregivers' quality of life

- Complex care requirements can cause lengthy and frequent hospitalizations and make daily social activities difficult<sup>4,5</sup>
- Severe CHI requires continuous enteral feeding, making transfer to other caregivers difficult (e.g., school)<sup>4</sup>
- More than 50% of CHI patients may be unresponsive to current medical treatment options<sup>6</sup>



Sources: <sup>1</sup>Arnoux JB et al. 2011 Orphanet J Rare Dis;6:63; <sup>2</sup>Yau et al. Plos One 2020;15(2):e0228417; <sup>3</sup>Thornton PS et al., J Pediatr. 2015;167(2):238-45. <sup>4</sup>Banerjee I et al., Orphanet J Rare Dis. 2022;17:61; <sup>5</sup>Pasquini TLS et al. Front Endocrinol 2022:13;876903; <sup>6</sup>Yorifuji et al. Clin Pediatr Endocrinol 2017;26(3):127-152. QoL=quality of life; CHI=congenital hyperinsulinism

# Dasiglucagon has potential to address shortcomings of current management of CHI



## **Current treatments for CHI are associated with significant limitations and clinical barriers**

#### Cited by healthcare providers as greatest limitations<sup>5</sup>

- Lack of responsiveness or incomplete response
- Adverse effects or intolerable side effects

| Maver                                    | Adverse effects of intelerable side effects   |  |  |  |  |
|--|---|--|--|--|--|
| Treatment                                | Current usage (availability varies by country)  | Clinical barriers  |  |  |  |
| Diazoxide                                | Approved for hyperinsulinism due to<br>various underlying conditions in the<br>US and certain ex-US regions <sup>2</sup>  | <ul> <li>FDA-issued warning on pulmonary hypertension in infants in 2015<sup>2,3</sup></li> <li>Lack of adequate response<sup>1</sup></li> <li>Hypertrichosis<sup>2</sup></li> <li>Fluid retention, acute heart failure, pulmonary hypertension<sup>2</sup></li> </ul> |  |  |  |
| Glucagon                                 | • Used off-label in CHI¹  | <ul> <li>Requires daily reconstitution of<br/>lyophilized glucagon</li> <li>Precipitates in the infusion tube (cannot<br/>use long-term)¹</li> </ul>   |  |  |  |
| Somatostat<br>in analogs<br>(octreotide) | <ul> <li>Used off-label in CHI¹</li> <li>Short acting: 3-4 daily s.c. injections/continuous infusion¹,⁴</li> <li>Long-acting: intramuscular injection every 28 days⁵</li> </ul> | <ul> <li>Hepatotoxicity<sup>1,4</sup></li> <li>Tachyphylaxis, QT prolongation<sup>4</sup></li> <li>Necrotizing enterocolitis (can be fatal in children with CHI)<sup>1,4</sup></li> </ul>  |  |  |  |
| Pancreatic<br>surgery                    | Total/near-total pancreatectomy in<br>diffuse CHI if medical management<br>fails <sup>1</sup>   | <ul> <li>Patients develop lifelong insulin<br/>dependent diabetes mellitus<sup>5</sup></li> <li>Patients develop lifelong severe<br/>exocrine insufficiency<sup>5</sup></li> </ul>   |  |  |  |

### Dasiglucagon for subcutaneous infusion\*

#### Wearable s.c. infusion pump system<sup>6</sup>

 Glucagon analog designed to allow for continuous subcutaneous (s.c.) infusion via pump



Dasiglucagon is a glucagon receptor agonist that works by causing the liver to release stored sugar to the blood



Two Phase 3 trials in neonates and children up to 12 years of age demonstrated potential in management of CHI



Zealand is prepared to resubmit Part 1 of the original NDA related to dosing of up to three weeks<sup>7</sup>



Submission of Part 2 of the original NDA related to dosing beyond three weeks is planned for after Part 1 resubmission<sup>7</sup>

IP exclusivity: compound patent US 2035 and EU 2039

Sources: ¹Yorifuji et al. Clin Pediatr Endocrinol 2017;26(3):127-152; ²Proglycem. Package insert. Teva Pharmaceuticals; 2015; ³Gray KD et al. J Perinatol. 2018;38(11):1496-1502; ⁴Haris et al. Therapeutic Adv Endocrinology Metabolism 2020;11:1-23; ⁵Zealand Pharma, Physician Market Survey, 2020; ⁶Zealand Pharma has entered a collaborative development and supply agreement with DEKA Research & Development Corporation and affiliates for infusion pump system; ¬FDA 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.

<sup>\*</sup>Investigational compound and device whose safety and efficacy have not been evaluated or approved by the FDA or any other regulatory authority

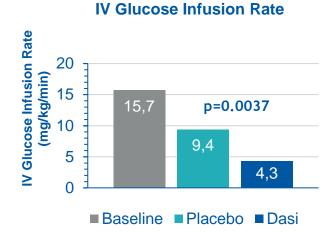
### Three Phase 3 trials form the basis of our NDA submission to the U.S. FDA





Trial 17103: Dasiglucagon significantly reduced the requirement for IV glucose in a hospital setting

### Part 1: Placebo control, crossover x 48 hours<sup>1</sup>

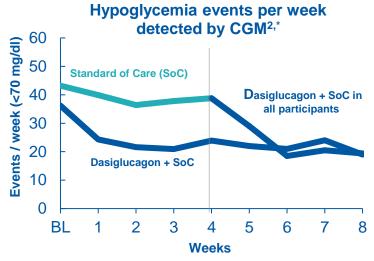


### Part 2: 21-days open-label treatmentc<sup>1</sup>

- 10 of 12 patients weaned off IV glucose >12 hours
- 7 patients weaned off IV glucose without need for pancreatectomy



Trial 17109: Dasiglucagon reduced time in hypoglycemia by ~50% and hypoglycemic events by 37-40% in a homecare setting



\*Primary endpoint comparing rates of hypoglycemia detected by SMPG demonstrated no difference between dasiglucagon and SoC CGM = continuous glucose monitoring; SMPG = self-measured plasma glucose

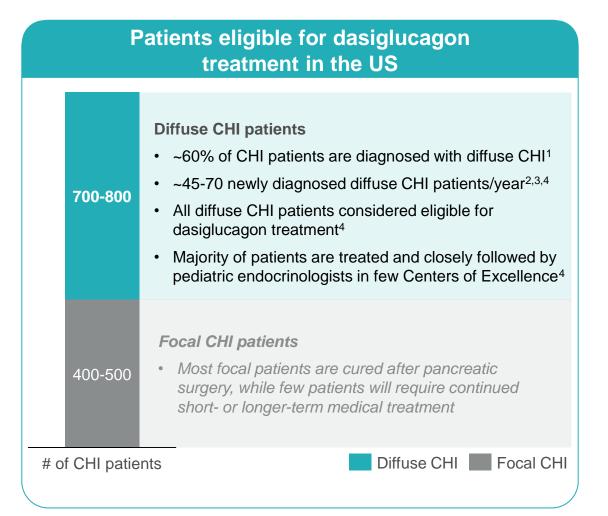
- Assessed as generally well tolerated in both trials
- Skin reactions and gastrointestinal disturbances most frequently reported adverse events

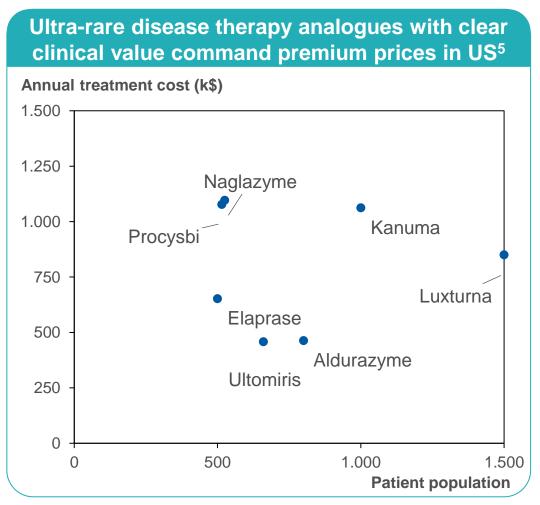
### 42 of 44 participants continued into long-term extension trial 17106

17103 Phase 3 clinical trial enrolled patients aged 7 days to 12 months, who were newly diagnosed and dependent on IV glucose in hospital setting: https://clinicaltrials.gov/ct2/show/NCT04172441 17106 is an open label long-term safety study that enrolled 17109 and 17103 participants with ongoing positive benefit / risk aged >1 month: https://clinicaltrials.gov/ct2/show/NCT03941236 17109 Phase 3 clinical trial enrolled children aged 3 months to 12 years being treated with standard of care (+/- surgery) with persistent hypoglycemia: https://clinicaltrials.gov/ct2/show/NCT03777176 Sources: 1De Leon et al. J Clin Endocrinol Metab, November 2024 (published online ahead of print); 2Thornton et al. J Clin Endocrinol Metab. 2023 Nov 1;109(4):1071–1079.

## Opportunity to treat up to 800 patients at ultrarare disease price levels in the U.S.







Sources: <sup>1</sup>Arya et al. Plos One 2014;9:e98054; <sup>2</sup>Arnoux JB et al. 2011 Orphanet J Rare Dis;6:63; <sup>3</sup>Yau et al. Plos One 2020;15(2); <sup>4</sup>Based on KOL interviews (2022); <sup>5</sup>Zealand Pharma Payer & Pricing Research, December 2022 <a href="Indications by product">Indications by product</a>: Procysbi (nephropathic cystinosis); Naglazyme (Maratolamy syndrome); Ultomoris (atypical hemolytic uremic syndrome); Kanuma (lysosomal acid lipase deficiency); Luxturna (biallelic RPE65 mutation-associated retinal dystrophy); Elaprase (Hunter syndrome); Aldurazyme (Hurler and Hurler-Scheie forms of Mucopolysaccharidosis I).

## **Short Bowel Syndrome (SBS) is a rare, chronic and debilitating condition**



#### High unmet need

- SBS is a rare, chronic and debilitating condition resulting in impaired intestinal absorptive capacity<sup>1, 2</sup>
- SBS is associated with significant medical complications including liver and renal failure, metabolic complications, chronic fatigue, and life-threatening infections<sup>3</sup>

#### Life-long dependency on parenteral support (PS)

- SBS patients experience chronic dependence on complex PS to provide necessary nutrition and fluid intake and balance<sup>3</sup>
- PS management is associated with a significant burden on health care systems and reduction in the patients' and caregivers' quality of life<sup>4,5</sup>

#### **Need for improved treatment options**

More effective and convenient treatments to further reduce PS is needed, with the ultimate goal of enteral autonomy<sup>3</sup>



Sources: ¹Jeppesen P., Expert Opinion on Orphan Drugs; 1:515-25, 2013; ²Pironi, L, et al. Definitions of intestinal failure and the short bowel syndrome. Best Practice & Research Clinical Gastroenterology. 30(2), 173-185 (2016); ³Cueda C et al. ESPEN Practical Guideline: clinical nutrition in chronic intestinal failure. Clin Nutrition 40; 5196-5120 (2021); ⁴Belza et al. Stress, Anxiety, Depression and Health-Related Quality of Life in Caregivers of Children with Intestinal Failure on Parenteral Nutrition: A Cross-sectional Survey Study. JPEN J Parenter Enteral Nutr. 2022 Nov 6. doi: 10.1002/jpen.2461; ⁵Winkler et al. Clinical, social, and economic impacts of home parenteral nutrition dependence in short bowel syndrome.

## Glepaglutide has best-in-class potential as a next-generation GLP-2 therapy for SBS patients



## Gattex® (teduglutide): only currently available GLP-2 treatment



Effective half-life of 1.3 hours at steady state



0.05 mg/kg daily subcutaneous dosing via vial/syringe



Multi-step reconstitution process<sup>1</sup>



## Glepaglutide: a long-acting stable GLP-2 analog<sup>a</sup>



Effective half-life of ~88 hours at steady state<sup>2</sup>



Expected 10 mg twice-weekly subcutaneous dosing



Ready-to-use auto-injector with needle protection

• Forms depot at the injection site



Second Phase 3 trial to be initiated in 2025 (EASE-5)

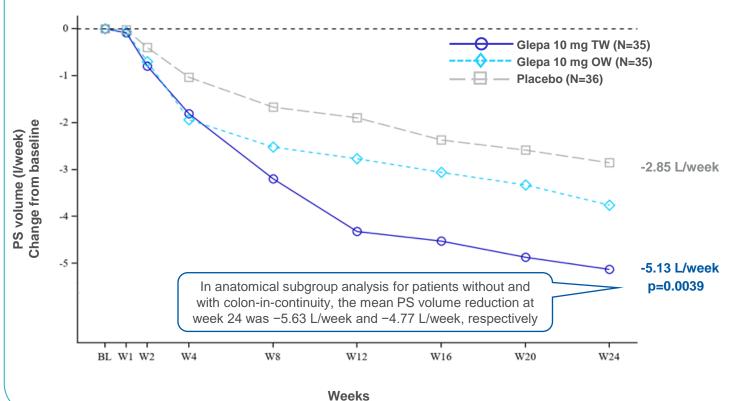
- Further confirmatory evidence for US resubmission
- Support regulatory submissions outside US and EU

<sup>a</sup>Investigational product, not approved for distribution; IP exclusivity: Compound patent 2026 + 5 years PTE; Dosing regime (pending) 2038, Clinical formulation (pending) 2039 Sources: <sup>1</sup>https://www.gattex.com/resources-and-support/; <sup>2</sup>Agersnap M. et al, 2022, Clin Drug Investigation; 42(12):1093-1100; <sup>3</sup>The U.S. FDA issued a Complete Response Letter for the glepaglutide New Drug Application for the treatment of short bowel syndrome

## Glepaglutide significantly reduced weekly PS volume at Week 24 in the EASE SBS-1 trial<sup>1</sup>







## Clinical response was significantly higher with twice weekly glepaglutide compared to placebo (p=0.0243)

- 65.7% twice weekly glepaglutide
- 45.7% once weekly glepaglutide
- 38.9% placebo group

## 9 patients treated with glepaglutide discontinued PS during the trial

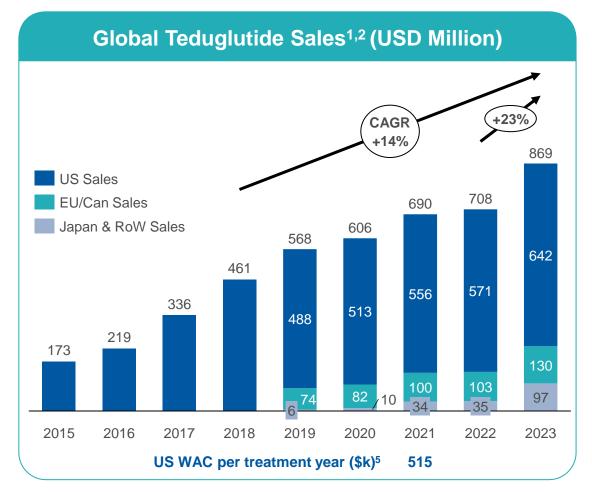
- 14% (n=5) twice weekly glepaglutide
- 11% (n=4) once weekly glepaglutide
- No patients receiving placebo

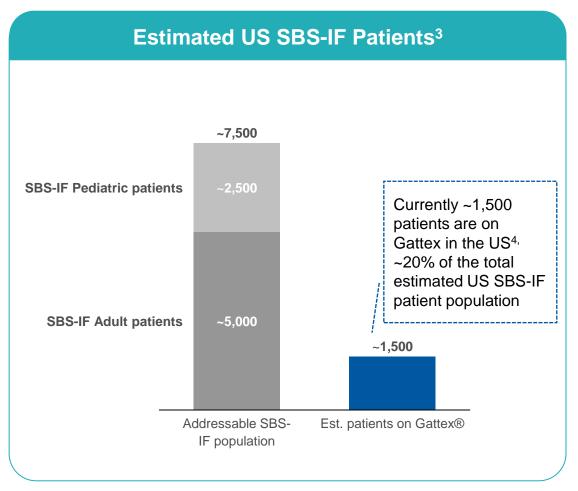
#### Glepaglutide appeared to be well-tolerated in the trial

 Most frequently reported adverse events were injection site reactions and gastrointestinal events

# Global teduglutide sales of >\$850M with significant room for market expansion





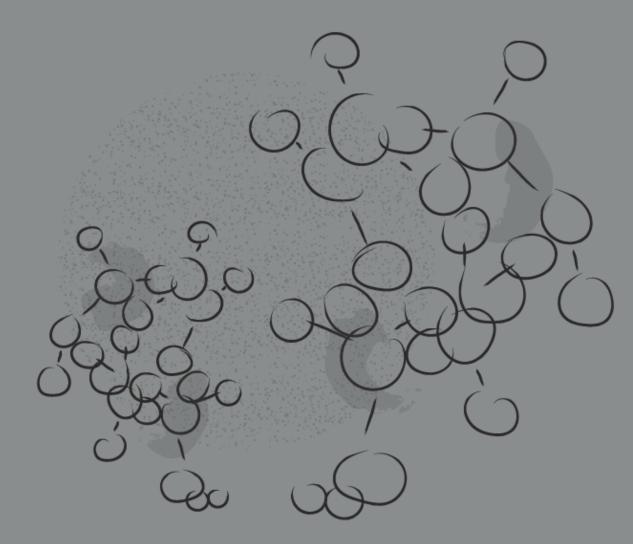


Sources: ¹2015-2018: Carnegie ZEAL research report, 24 February 2020; ²2019-23: Gattex/Revestive sales data as reported in Takeda SEC filings, following fiscal financial year April to Mar. Converted to USD per Yearly Average Currency Exchange Rates, IRS.gov; ³SBS Intestinal Failure patient estimates based on Zealand Pharma claims analysis, 2020 and Mundi et al, Characteristics of Chronic Intestinal Failure in the USA Based on Analysis of Claims Data, JPEN in Press 2022; ⁴ZP estimate based on US Gattex sales and net price estimate; ⁵WAC at end of year, <a href="https://app.prospectorx.com">https://app.prospectorx.com</a>. WAC=wholesaler acquisition cost; SBS=short bowel syndrome; IF=intestinal failure



## **Chronic Inflammation**

**Corporate Presentation** 



## Early-stage pipeline targets chronic inflammatory diseases with significant unmet medical needs



### **Kv1.3 blocker (ZP9830)**

- Potent and selective inhibitor of Kv1.3 with the potential to treat a broad range of cell-mediated autoimmune diseases<sup>1</sup>
- Concentration-dependent inhibition of proinflammatory cytokine release (including IFN-g, IL-2 and IL17A) from stimulated human whole blood<sup>1</sup>
- First-in-human clinical trial investigating the safety and tolerability of ZP9830 has been initiated<sup>2</sup>

### **Complement C3 inhibitor (ZP10068)**

- Investigational, long-acting inhibitor of Complement C3, which has the potential to treat a broad range of complement-mediated diseases
- In 2023, Zealand completed activities to support advancing ZP10068 into the first-in-human clinical trials
- Zealand will assess the potential next steps for ZP10068<sup>3</sup>

## ZP9830 is a Kv1.3 inhibitor designed to treat cell-mediated immune disorders



ZP9830 inhibits Kv1.3, the main K+ channel of leukocytes from the innate and adaptive immune system1



Kv1.3. channels are essential for the activation, proliferation, migration and cytokine production of leukocytes<sup>2</sup>



T effector memory and class-switched memory B cells play a **key role** in **autoimmunity** and **chronic inflammation** and are **dependent** on Kv1.3 for function<sup>3</sup>



Inhibition of Kv1.3 channels preserves the protective effects of the rest of the immune system, making it an attractive pharmaceutical target



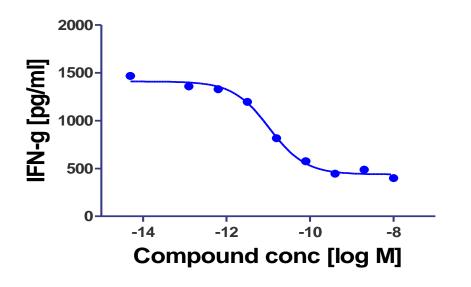
ZP9830 is a potent and selective Kv1.3 inhibitor with potential to treat a broad range of cell-mediated autoimmune diseases. First-in-human clinical trial initiated in Q4 2024<sup>4</sup>

## First-in-human clinical trial investigating safety and tolerability of ZP9830 is ongoing





Concentration-dependent inhibition of pro-inflammatory cytokine release (including IFN-g, IL-2 and IL17A) from stimulated human whole blood<sup>1</sup>



### First-in-human SAD clinical trial<sup>2</sup>

The first-in-human Phase 1 single ascending dose trial include 10 dose cohorts and is expected to enroll 92 healthy men.

Cohorts 1-3 (SC): Assess safety and PK

Cohorts 4-9 (SC): Assess safety, PK, and PD

Cohort 10 (IV): Assess safety and PK

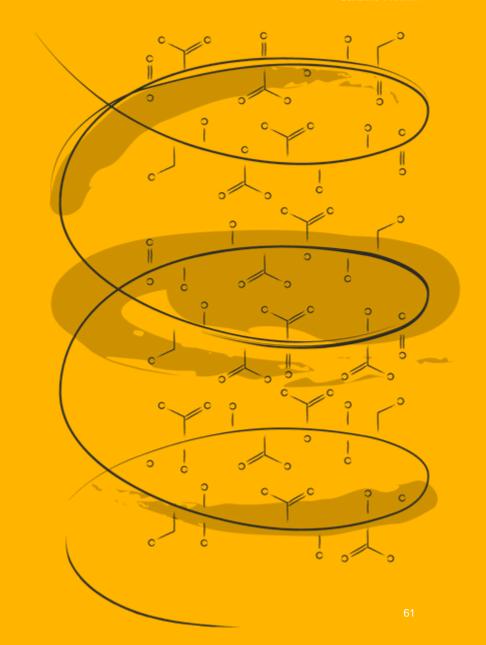
The trial will investigate:

- Safety and tolerability profile with single ascending doses
- PK profile to determine the appropriate dose level(s)
- Effect of ZP9830 on the body's immune system



# Additional company information

**Corporate Presentation** 



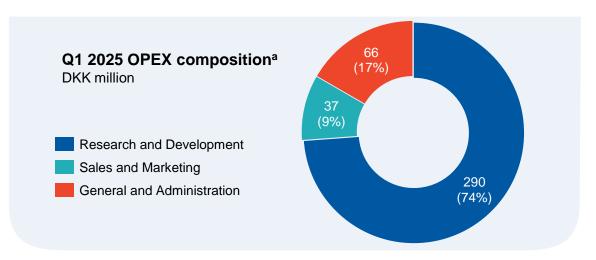


### Q1 2025 Profit & Loss

| DKK million                         | Q1 2025 | Q1 2024 |
|-------------------------------------|---------|---------|
| Revenue                             | 8.1     | 15.1    |
| Gross profit                        | 7.7     | 10.5    |
| Research and development expenses   | -290.3  | -190.9  |
| Sales and marketing expenses        | -37.4   | -9.2    |
| General and administrative expenses | -65.5   | -66.2   |
| Net operating expenses              | -393.1ª | -266.3  |
| Operating result                    | -385.5ª | -255.8  |
| Net financial items                 | 70.3    | 25.8    |
| Result before tax                   | -315.1ª | -230.0  |
| Tax                                 | 1.4     | 1.4     |
| Net result for the period           | -313.8ª | -228.6  |

## P&L reflecting Zealand's investments in its differentiated R&D assets and organization

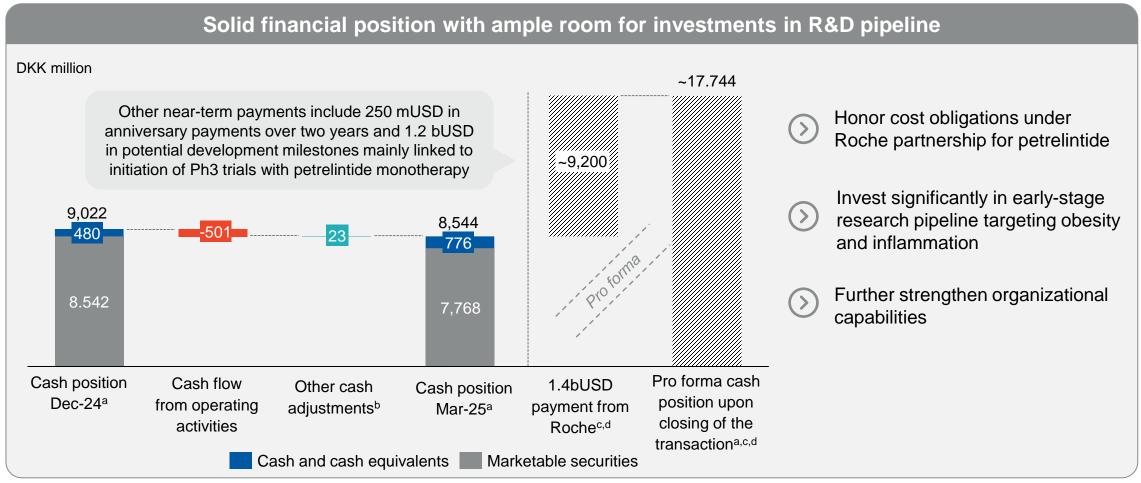
- Revenue of DKK 8 million is mainly driven by the license and development agreement with Novo Nordisk for Zegalogue<sup>®</sup>.
- R&D expenses of DKK 290 million are mainly driven by clinical costs associated with our mid-stage obesity assets, whereas S&M expenses of DKK 37 million are driven by pre-commercial activities for the rare disease assets. G&A expenses are mainly driven by strengthening of organizational capabilities as well as investments in IT infrastructure and patent portfolio.
- Net financial items of DKK 70 million are mainly driven by interest income from the excess liquidity invested in marketable securities.



<sup>&</sup>lt;sup>a</sup>Excluding transaction costs of DKK 21.6 million related to the Roche partnership agreement. Net operating expenses including transaction-related costs amount to DKK 414.7 million..

# Well-funded with no need to raise additional capital towards expected profitability





<sup>&</sup>lt;sup>a</sup>Cash 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;

<sup>b</sup>Other cash adjustments include proceeds from sale of shares of Beta Bionics, Inc; <sup>c</sup>Based on foreign exchange rates as of May 7, 2025 (DKK 6.6 = USD \$1); <sup>d</sup>Zealand Pharma anticipates receiving USD 1.65 billion in upfront payments from Roche, of which USD 1.4 billion due at closing of the transaction and USD 250 million in anniversary payments over two years. Pro forma financial figures are subject to uncertainties including, inter alia, exchange rate fluctuations and regulatory delays.

EIB = European Investment Bank



## 2025 financial guidance

| DKK million  | 2025<br>Guidance <sup>a</sup> | 2024<br>Actuals |
|--|-------------------------------|-----------------|
| Revenue anticipated from existing and new license and partnership agreements | No guidance                   | 63              |
| Net operating expenses <sup>b</sup>  | 2,000 – 2,500                 | 1,327           |

<sup>&</sup>lt;sup>a</sup>Financial guidance on net operating expenses for 2025, published on February 20, 2025, is confirmed excluding transaction-related costs related to the Roche collaboration announced on March 12, 2025. Total transaction fees related to the Roche partnership agreement are expected to be approximately DKK 200 million in 2025; <sup>b</sup>Net operating expenses consist of R&D, S&M, G&A and other operating items. Financial guidance based on foreign exchange rates as of May 7, 2025.



## **Corporate management team**











