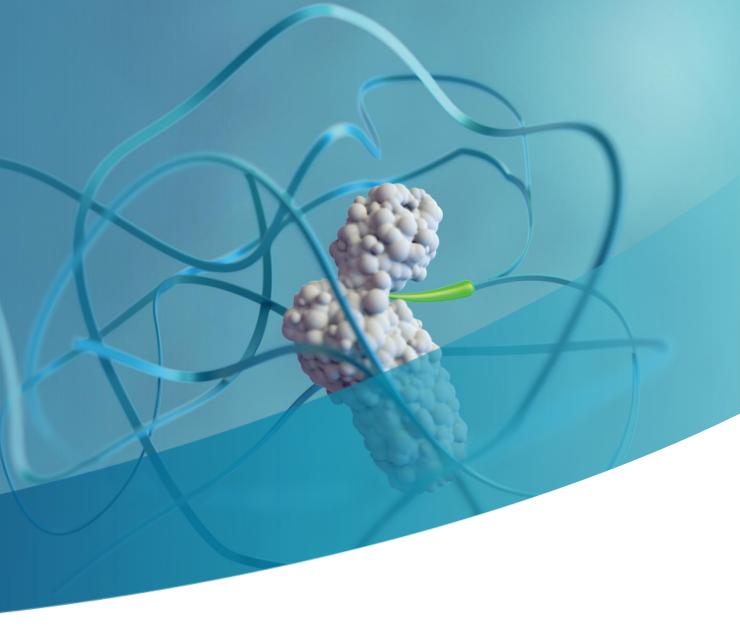
## Ascendis Pharma A/S

J.P. Morgan Healthcare Conference San Francisco January 2025





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This presentation contains forward-looking statements. All statements other than statements of historical facts contained in this presentation, such as statements regarding our products and prospective product candidates; revenue and growth expectations; PDUFA goal dates; clinical trial results; the expected timing of future clinical trial results, regulatory filings and feedback from regulatory authorities; the scope, progress, results and costs of developing our product candidates or any other future product candidates; timing and likelihood of success; patient enrollment; timing and expansion of commercial launches, pipelines, and investments; potential payments and royalties relating to investments and partnerships; plans and objectives of management for future operations and commercialization and manufacturing activities; and future results of current and/or anticipated products and product candidates are forward-looking statements. These forward-looking statements are based on our current expectations and beliefs, as well as assumptions concerning future events. These statements involve known and unknown risks, uncertainties and other factors that could cause our actual results to differ materially from the results discussed in the forward-looking statements. These risks, uncertainties and other factors are more fully described in our reports filed with or submitted to the SEC, including without limitation, our prospectus supplement filed on September 20, 2024 and our current and future reports filed with or submitted to the SEC, including our most recent Annual Report on Form 20-F filed with the SEC on February 7, 2024, particularly in the sections titled "Risk Factors" and "Operating and Financial Review and Prospects." In light of the significant uncertainties in our forward-looking statements, you should not place undue reliance on these statements or regard these statement made by us in this presentation or warranty by us or any other person that we will achieve our objectives and plans in any specif

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### Ascendis Positioned to Drive Rapid Revenue Growth



- Three clinically validated Endocrinology Rare Disease medicines:
  - SKYTROFA<sup>®</sup> Approved for pediatric growth hormone deficiency (GHD)<sup>1</sup>
    - Full-year 2024 SKYTROFA revenue expected to be ~€202 million<sup>2</sup>, excluding sales deductions related to prior years
    - sBLA submitted for the treatment of adults with GHD, with a PDUFA goal date of July 27, 2025
  - YORVIPATH<sup>®</sup> Approved for the treatment of hypoparathyroidism in adults<sup>1</sup>
    - Launched January 2024 in Germany/Austria; available in the U.S. since late December 2024
    - Full-year 2024 YORVIPATH revenue expected to be ~€29 million<sup>3</sup>
  - TransCon<sup>™</sup> CNP Successful pivotal data in achondroplasia
    - Following pre-NDA meeting, planned filings for treatment of children with achondroplasia in U.S. Q1 2025, EU in Q3 2025
- Expanding TransCon technology platform with broad therapeutic applicability
  - Formed Eyconis for Ophthalmology and partnered with Novo Nordisk for Metabolic and Cardiovascular diseases
  - $\circ\,$  Internal development for Oncology with focus on TransCon IL-2  $\beta/\gamma\,$
  - New TransCon protein degrader technology to expand pipeline with additional potential blockbusters

### Business model built on fast, successful drug development and commercial therapeutic synergies

1. Approved in the U.S., EU, and other territories, including Norway, Iceland, Liechtenstein, and Great Britain (covering England, Wales, Scotland). 2. Calculated as unaudited preliminary estimate of full year 2024 SKYTROFA revenue of ~€197 million plus ~€5 million of sales deductions related to prior years. 3. Unaudited preliminary estimate.





PASSIC

### Achieve blockbuster status for multiple products and expand our engine for future innovation

#### Be the Leading Endocrinology Rare Disease Company

- Achieve blockbuster status (>\$1B) for each of TransCon PTH, TransCon hGH, and TransCon CNP through worldwide commercialization
- Be the leader in Growth Disorders and Hypoparathyroidism, pursuing clinical conditions, innovative LCM, and complementary patient offerings
- Expand pipeline with Endocrinology Rare Disease blockbuster product opportunities

#### **Create Value in Additional Therapeutic Areas through Innovative Business Models**

- Obtain accelerated approval in Oncology with registrational trials ongoing
- Pursue TransCon product opportunities in >\$5B indications
- · Maximize value creation of these product opportunities through collaboration with therapeutic area market leaders

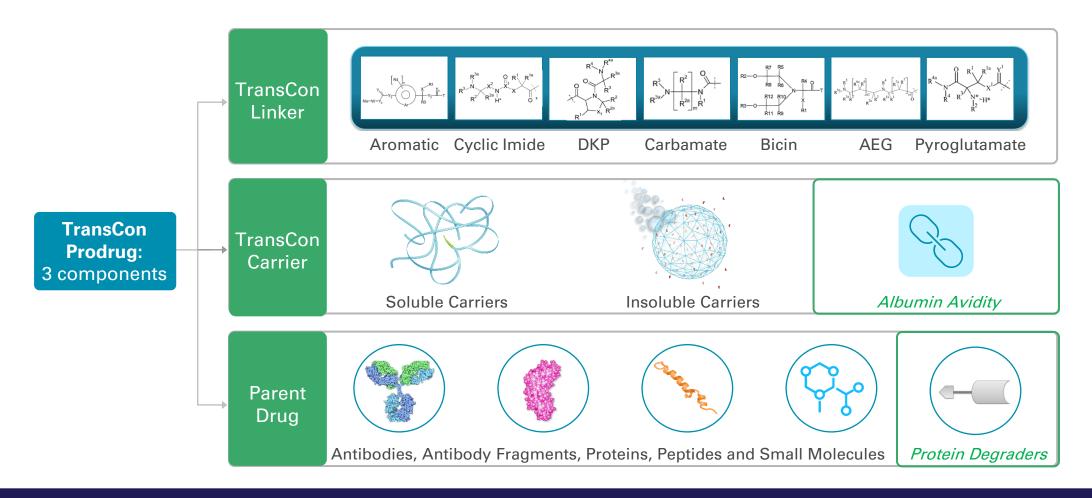
#### **Differentiate with Ascendis Fundamentals**

- · Outperform industry drug development benchmarks with Ascendis' product innovation algorithm
- · Remain independent as a profitable biopharma through lean and flexible ways of working
- Let our values Patients, Science, Passion drive our decisions to success

Ascendis Pharma's 2025 - 2030 strategic roadmap

### Overview of TransCon Technologies

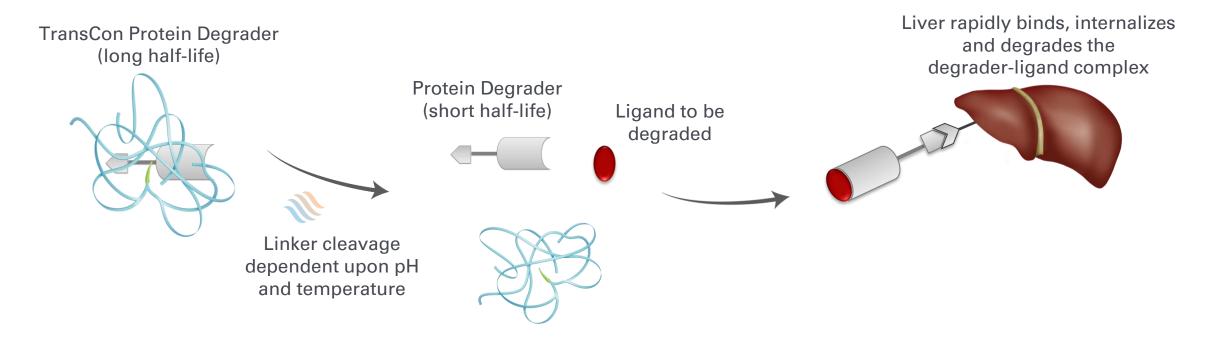




#### Continued expansion of TransCon technologies, enabling new product candidates

### TransCon Protein Degrader Technology

 TransCon protein degrader technology designed to enable efficient clearance of hormones, cytokines, and other targets by utilizing hepatic scavenger receptors



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Enabling product opportunities for diseases characterized by excess hormones, cytokines, etc.

### TransCon Clinical Development Pipeline



Endocrinology Rare Diseases		Indication	Status	Region
Lead Indication	TransCon CNP	Achondroplasia (children aged 2–11)	<ul> <li>Pivotal<sup>1</sup></li> </ul>	Multinational
Label Expansion	TransCon hGH	Adult Growth Hormone Deficiency	<ul> <li>sBLA submitted<sup>2</sup></li> </ul>	Multinational
		<ul> <li>Turner Syndrome (children aged 1–10)</li> </ul>	• Phase 2 <sup>3</sup>	• U.S.
	TransCon CNP	Achondroplasia (infants)	<ul> <li>Phase 2<sup>4</sup></li> </ul>	Multinational
	TransCon CNP	<ul> <li>Achondroplasia (adolescents)</li> </ul>	• Phase 2 <sup>5</sup>	Multinational
	TransCon CNP + TransCon hGH	<ul> <li>Achondroplasia (children aged 2–11)</li> </ul>	• Phase 2 <sup>6</sup>	Multinational
Partner Programs	TransCon hGH	Pediatric GHD	• BLA submitted <sup>7</sup>	• China
	TransCon hGH	Pediatric GHD	<ul> <li>Phase 3<sup>8</sup></li> </ul>	• Japan
	TransCon PTH	Hypoparathyroidism in adults	<ul> <li>Phase 3<sup>9</sup></li> </ul>	China
	TransCon PTH	Hypoparathyroidism in adults	<ul> <li>J-NDA submitted<sup>10</sup></li> </ul>	• Japan
	TransCon CNP	Achondroplasia	• Phase 2 <sup>11</sup>	• China
Oncology		Indication	Status	Region
Lead Indication	TransCon IL-2 β/γ	<ul> <li>Various tumor types</li> </ul>	• Phase 2 <sup>12,13</sup>	Multinational

1. Pivotal ApproaCH Trial (NCT05598320). 2. sBLA submitted to U.S. FDA, PDUFA goal date July 27<sup>th</sup> 2025. 3. New InsiGHTS Trial (NCT05690386). 4. reACHin Trial (NCT06079398). 5. teACH (NCT06732895). 6. COACH Trial (NCT06433557). 7. VISEN Pharmaceuticals' Phase 3 trial. 8. Japanese riGHt Trial. 9. PaTHway China. 10. PaTHway Japan. 11. ACcomplish China. 12. BelievelT-201 Trial (NCT05980598). 13. IL-Believe Trial (NCT05081609).

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# Growing Worldwide Commercialization in Endocrinology Rare Disease<sup>1</sup>

Multiple approaches:

- Ascendis direct commercialization in the U.S. and select European countries
- Through partners with local expertise and infrastructure outside of U.S. and Europe Direct<sup>2</sup>

Ascendis direct commercialization

- Exclusive distribution agreement
- Strategic investment and exclusive license agreement
- Exclusive license agreement

As of December 31, 2024.
 DACH, France & BeNeLux, Iberia, Italy, Nordics, UK & Ireland.



# SKYTROFA® TransCon hGH

Once-weekly growth hormone therapy

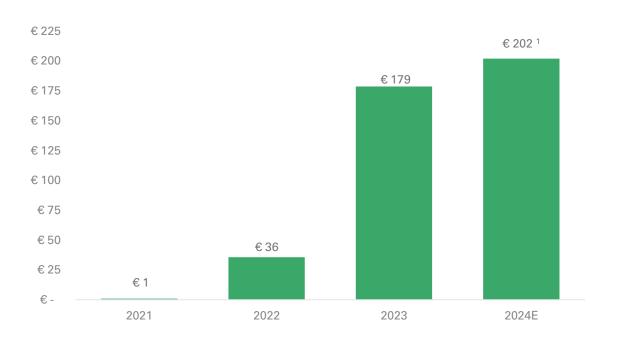


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### SKYTROFA - High Value Growth Hormone Brand

- Launched for pediatric growth hormone deficiency in U.S. in Q4 21, Germany in Q3 23
  - Successful Phase 3 riGHt Trial in Japan
  - Planned expansion across multiple countries
- Planned label expansion to drive growth
  - sBLA accepted for adult growth hormone deficiency; PDUFA goal date of July 27, 2025
  - Positive topline results from Phase 2 New InsiGHTS Trial in Turner syndrome
  - Submit IND or similar for a basket trial evaluating additional indications<sup>2</sup> in Q3 2025

#### SKYTROFA Revenue (€ million)



#### Committed to making SKYTROFA the global leading product in value in a growing growth hormone market

sBLA = supplemental Biologics License Application.

1. Calculated as unaudited preliminary estimate of full year 2024 SKYTROFA revenue of ~€197 million plus ~€5 million of sales deductions related to prior years. 2. Planned for small for gestational age without catch-up growth ["SGA"]; Idiopathic short stature ["ISS"]; SHOX deficiency (including Turner syndrome).

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### TransCon hGH Program Summary and Outlook



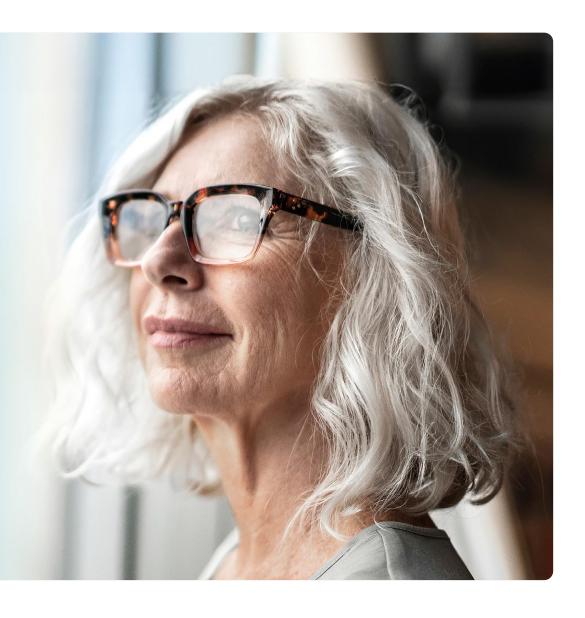
- 2024 SKYTROFA revenue, with single indication in pediatric GHD, expected to be ~€202 million<sup>1</sup>
  - $\circ\,$  High-value brand in the U.S., revenue per patient of around 3 times that of daily growth hormone
  - Volume (mg) increased 84% in 2024 resulting in 6.5% market share of total U.S. growth hormone market<sup>2</sup>
- Adult GHD
  - $^\circ\,$  sBLA submitted to FDA; PDUFA goal date of July 27, 2025
  - U.S. commercial launch planned in Q4 2025, pending approval
- Turner syndrome
  - Phase 2 New InsiGHTS Trial achieved primary objective at Week 26
- Plan to submit IND or similar for basket trial evaluating additional indications<sup>3</sup> in Q3 2025
- Planned commercial launches across multiple indications and countries

### Committed to making SKYTROFA a blockbuster product

Calculated as unaudited preliminary estimate of full-year 2024 SKYTROFA revenue of ~€197 million plus ~€5 million of sales deductions related to prior years.
 Based on third party prescription data for 2024.
 Planned for small for gestational age without catch-up growth ["SGA"]; Idiopathic short stature ["ISS"]; SHOX deficiency (including Turner syndrome).

# YORVIPATH<sup>®</sup> TransCon PTH

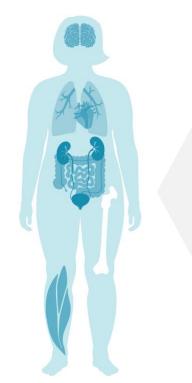
Treatment of hypoparathyroidism in adults





### Unmet Medical Need in Hypoparathyroidism: One Disease, Multiple Consequences<sup>1-4</sup>





### Absence or deficiency of parathyroid hormone (PTH) is linked to multi-organ manifestations<sup>5,6</sup>

#### Central Nervous System

- Seizures
- Basal Ganglia Calcifications
- Parkinsonism Or Dystonia

#### Respiratory

• Laryngospasm

#### Cardiovascular

- Vascular Calcifications
- Cardiac Arrhythmias
- Hypocalcemia-associated Dilated Cardiomyopathy

#### Renal

- Nephrocalcinosis\*
- Nephrolithiasis\*
- Chronic Kidney Disease\*

#### **Peripheral Nervous System**

- Paresthesia
- Muscle Cramps
- Pain
- Tetany

#### Neuropsychiatric

- Anxiety & Depression
- Cognitive Impairment ("Brain Fog")

#### **Ophthalmological**

- Cataracts
- Papilledema

#### Dental

Altered Tooth Morphology

#### Dermatological

- Dry Skin
- Thinning Hair
- Brittle Nails

#### **Musculoskeletal**

- Myopathy
- Spondyloarthropathy

\* These manifestations are mostly the result of management with active vitamin D and calcium rather than of the disease itself.

1. Underbjerg L, Sikjaer T, Mosekilde L, et al. Cardiovascular and renal complications to postsurgical hypoparathyroidism: a Danish nationwide controlled

- historic follow-up study. J Bone Miner Res. 2013;28(11):2277-2285. doi:10.1002/jbmr.1979.
- 2. Underbjerg L, Sikjaer T, Mosekilde L, et al. The epidemiology of nonsurgical hypoparathyroidism in Denmark: a nationwide case finding study. J Bone Miner Res. 2015;30(9):1738-1744. doi:10.1002/jbmr.2501.
- 3. Shoback DM, Bilezikian JP, Costa AG, et al. Presentation of hypoparathyroidism: etiologies and clinical features. J Clin Endocrinol Metab. 2016;101(6):2300-2312. doi:10.1210/jc.2015-3909.
- 4. Underbjerg L, Sikjaer T, Mosekilde L, et al. The epidemiology of nonsurgical hypoparathyroidism in Denmark: a nationwide case finding study. *J Bone Miner Res.* 2015;30(9):1738-1744. doi:10.1002/jbmr.2501.
- 5. Mannstadt M, Bilezikian JP, Thakker RV, et al. Hypoparathyroidism. *Nat Rev Dis Primers*. 2017;3:17055. doi:10.1038/nrdp.2017.55.

6. Shoback DM, Bilezikian JP, Costa AG, et al. Presentation of hypoparathyroidism: etiologies and clinical features. J Clin Endocrinol Metab. 2016;101(6):2300-2312. doi:10.1210/jc.2015-3909.

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Current Clinical Practice Guideline

- Consider PTH replacement therapy in patients not adequately controlled on conventional therapy
- Inadequate control is considered to be <u>any one</u> of the following:
  - Symptomatic hypocalcemia
  - Hyperphosphatemia
  - Renal insufficiency
  - Hypercalciuria
  - Poor quality of life
- In addition, individuals with poor compliance, malabsorption, or who are intolerant of large doses of calcium and active vitamin D may also benefit from PTH replacement therapy

### Vast majority of patients with hypoparathyroidism fulfill guideline for PTH replacement therapy

Khan AA, Bilezikian JP, Brandi ML, et al. Evaluation and Management of Hypoparathyroidism Summary Statement and Guidelines from the Second International Workshop. *J Bone Miner Res.* Dec 2022;37(12):2568-2585. doi:10.1002/jbmr.4691

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Evaluation and Management of Hypoparath Summary Statement and Guidelines from th International Workshop	

TASK FORCE

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**TRMR**<sup>®</sup>

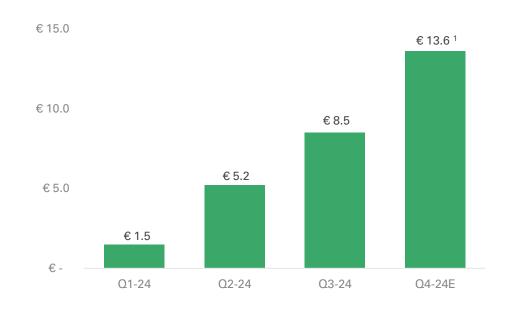
### YORVIPATH Commercial Update



### • Europe Direct

- Commercial launch in Germany and Austria in January 2024, with initial list price of €105,000 per patient per year
- Outside Germany and Austria, providing commercial product through early access routes, such as 'named patient,' until commercial reimbursement established
- Expect multiple commercial launches across Europe in 2025
- International Markets
  - 8 exclusive distribution agreements signed for 50+ countries
  - Revenue recognized from Er-Kim (Central & Eastern Europe), Neopharm (Israel), and Canada
- ~700 patients on YORVIPATH treatment in Europe Direct and International Markets at the end of 2024

### YORVIPATH Revenue (€ million)



### YORVIPATH on track for additional commercial launches in Europe Direct and International Markets in 2025

<sup>1</sup>Unaudited preliminary estimate.

# YORVIPATH<sup>®</sup> U.S. FDA Approved and Now Commercially Available



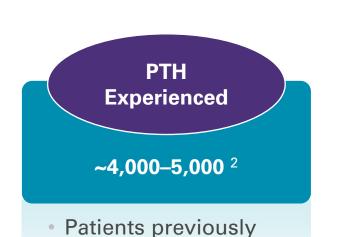


- YORVIPATH is the first and only product indicated for the treatment of hypoparathyroidism in adults<sup>1</sup>
- Commercially available in the U.S. December 2024

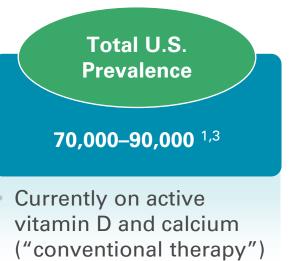
1. Limitations of Use: YORVIPATH was not studied for acute post-surgical hypoparathyroidism. YORVIPATH's titration scheme was only evaluated in adults who first achieved an albumin-corrected serum calcium of at least 7.8 mg/dL using calcium and active vitamin D treatment. YORVIPATH [package insert]. Princeton, NJ: Ascendis Pharma Endocrinology, Inc. August 2024.

### Hypoparathyroidism: U.S. Patient Population





treated with PTH





Newly

### YORVIPATH annual WAC price of ~\$285,000

WAC = Wholesale Acquisition Cost.

1. U.S. prevalence literature review (Powers, Clarke). 2. Internal estimates and Symphony Metys data. 3. U.S. prevalence literature review and epi meta-analysis (Powers, Clarke, Milliman project, ipm.ai claims project; HCUPnet, Healthcare Cost and Utilization Project. Agency for Healthcare Research and Quality, Rockville, MD for surgical cohort projection).

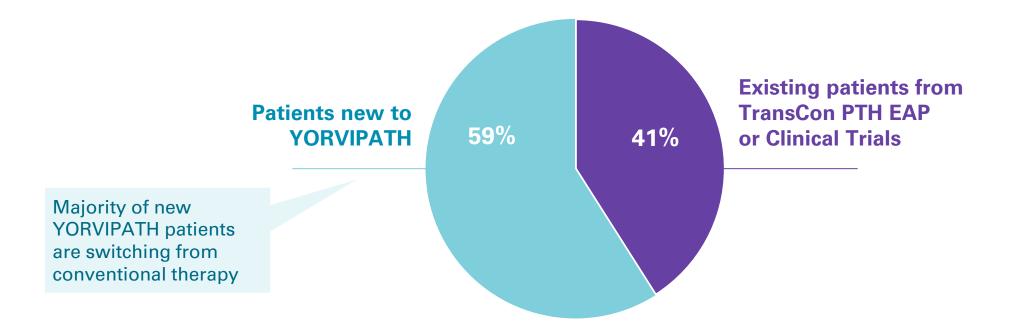
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• New prescriptions enrolled in A.S.A.P. or direct with specialty pharmacy (n=324) as of January 9, 2025

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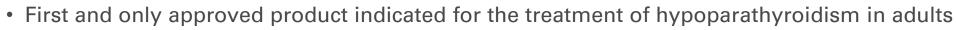


#### Strong early enrollments across all segments from ~150 unique prescribers

A.S.A.P. = Ascendis Signature Access Program; EAP = Expanded Access Program.

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### TransCon PTH Program Summary and Outlook



- Clinical trial results consistent across geographies, ethnicities, and etiologies
- TransCon PTH under regulatory review for market authorization in Australia, Israel, Japan, and Switzerland
- YORVIPATH 2024 revenue expected to be ~€29 million<sup>1</sup>
  - ~700 patients on YORVIPATH treatment in Europe Direct and International Markets at the end of 2024
  - Commercial launch expected in ≥5 additional Europe Direct countries in 2025
- U.S. commercial launch status as of January 9, 2025
  - Strong initial enrollments with 324 patients, including patients switching from conventional therapy
  - ~150 unique prescribers from 38 states



<sup>1.</sup> Unaudited preliminary estimate.

# TransCon CNP (navepegritide)

Investigational prodrug of CNP designed to provide sustained release and continuous exposure of active CNP for the treatment of pediatric achondroplasia





TransCon CNP is an investigational product candidate. For investor communication only. Not for use in product promotion. Not for further distribution.

### Pivotal ApproaCH Trial Summary



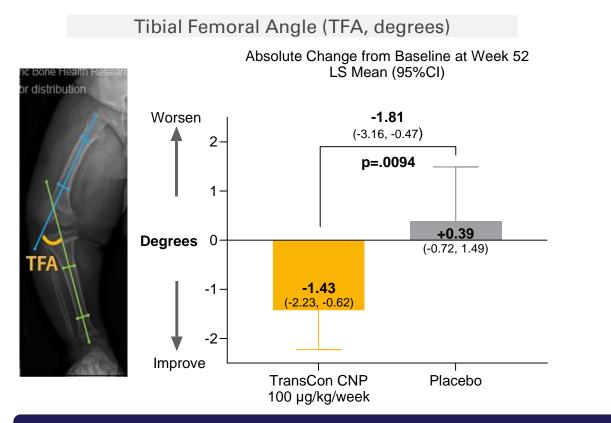
- The pivotal ApproaCH Trial in children aged 2-11 years achieved primary objective
  - TransCon CNP demonstrated LS mean AGV of 5.89 cm/year with LS mean treatment difference of 1.49 cm/year at Week 52 compared to placebo (p<0.0001)</li>
  - For children aged 5-11 years TransCon CNP demonstrated LS mean AGV of 5.79 cm/year with a change from baseline AGV superior to placebo with LS mean treatment difference of 1.78 cm/year at Week 52 (p<0.0001)</li>
- Other endpoints support that TransCon CNP may provide benefits beyond linear growth
  - Treatment with TransCon CNP resulted in numerical improvements in health-related quality of life compared to placebo as observed in several ACEM domains
  - Treatment benefit in muscle functionality demonstrated in sub-group of children aged 5-8 years
  - Treatment with TransCon CNP showed improvement in body proportionality, leg bowing, and other parameters
- TransCon CNP was generally well-tolerated, with low frequency of injection site reactions (all mild), and no evidence of hypotensive effect

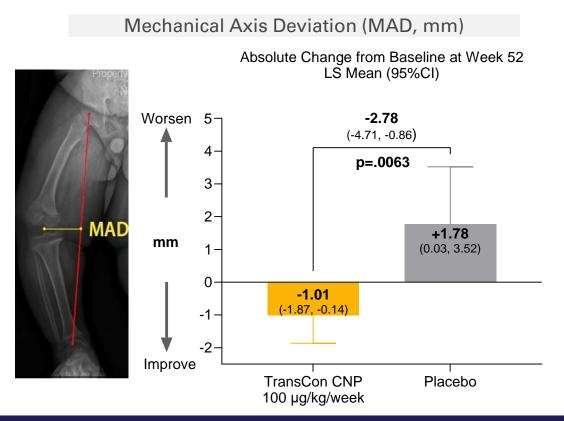
### Once-weekly TransCon CNP may address the need for an efficacious, safe, tolerable, and convenient treatment

Data on file. Ascendis Pharma. ACEM = Achondroplasia Child Experience Measure.

### Impact of TransCon CNP on Leg Bowing

Leg bowing is a common complication in achondroplasia, resulting in pain, impaired physical functioning, need for corrective surgery, and a negative impact on quality of life<sup>1,2,3</sup>





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TransCon CNP demonstrated significant improvements in leg bowing

Data on file. Ascendis Pharma.

1. Hunter et al. J Med. Genet. 35(9) (1998): 705–712. 2. Matsushita et al. Calcif. Tissue Int. 104 (2019) 364–372. 3. Nahm et al. Orphanet Journal of Rare Diseases 18 (2023) 139.

### TransCon CNP Program Summary and Outlook

- In pivotal ApproaCH Trial, TransCon CNP demonstrated significant improvements in linear growth and body proportionality, as well as benefits beyond linear growth
  - Following pre-NDA meeting, planned filings for treatment of children with achondroplasia in U.S. Q1 2025, EU in Q3 2025
- Comprehensive development plans continue with ongoing and planned trials to support TransCon CNP in additional patient populations
- COACH Trial first combination trial of TransCon hGH and TransCon CNP to further accelerate growth in achondroplasia; topline Week 26 data in children aged 2-11 expected in Q2 2025
- Plan to submit IND or similar for the treatment of hypochondroplasia in Q4 2025

### With SKYTROFA and TransCon CNP, Ascendis is well-positioned to become the leader in growth disorders

## Expanding the Endocrine Rare Disease Pipeline and Beyond



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### X-Linked Hypophosphatemia



### Background

- Characterized by excess FGF-23 production leading to excessive renal phosphate wasting<sup>1,2</sup>
- Affecting ~1:20000



### **Limitations of Existing Therapy**

- Increase in serum phosphorous from baseline plateaus at 1.0-1.5 mg/dL, irrespective of burosumab drug exposure<sup>3</sup>
- At the recommended doses, serum phosphorous remained in the lower limit of normal<sup>3</sup>
- Significant unmet medical need remains within the XLH patient population<sup>4,5</sup>

### Clear biological rationale for applying TransCon protein degrader technology to normalize FGF-23 levels

FGF-23 = Fibroblast growth factor 23. XLH = X-linked hypophosphatemia. <u>1. International XLH alliance</u>. **2** Beck-Nielsen *Orphanet J Rare Dis*. 2019. **3**. FDA Multi-Disciplinary Review BLA 761068. **4**. Padidela *Calcif Tissue Int* 2021 **5**. Linglart *JCEM* 2022.

### Collaboration with Novo Nordisk



- Development and commercialization of TransCon Technology-based products in metabolic (including obesity and type 2 diabetes) and cardiovascular diseases
- Highlights of the multi-product collaboration:
  - Upfront fee of \$100 million for the exclusive license and funding of the programs by Novo Nordisk
  - Lead program: once-monthly TransCon Semaglutide product candidate initially targeting obesity and type 2 diabetes
    - Potential to receive additional payments of up to \$185 million in development and regulatory milestone payments
    - Potential to receive escalating tiered, mid-single digit royalties on global net sales and sales milestone payments
  - Novo Nordisk responsible for clinical development, regulatory, commercial manufacturing, and commercialization

### Reflects our Vision 2030 to create value in additional large therapeutic areas

### Other Investments and Partnerships



- Eyconis (exclusive global rights for TransCon based ophthalmology products)
  - Formed Eyconis in January 2024 with a \$150 million commitment from external investors
- Teijin (exclusive rights for TransCon hGH, PTH, and CNP in Japan)
  - In December 2024, submitted a J-NDA to the Japanese medical authorities for TransCon PTH for the treatment of adults in Japan with hypoparathyroidism
- VISEN Pharmaceuticals (exclusive rights for TransCon hGH, PTH, and CNP in Greater China)
  - Announced in September 2024, Phase 3 PaTHway China Trial success for TransCon PTH, achieving primary and key secondary endpoints in treating adults with hypoparathyroidism

### Creating value through innovative business models

# Financial Update & 2025 Selected Milestones



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### 2024 Financial Update



- Total full-year 2024 product revenue expected to be ~€226<sup>1</sup> million, driven by:
  - SKYTROFA full-year 2024 revenue expected to be ~€202<sup>2</sup> million, excluding sales deductions related to prior years
  - YORVIPATH full-year 2024 revenue expected to be ~€29<sup>1</sup> million
- Full-year 2024 total revenue expected to be ~€364<sup>1</sup> million
  - Includes \$100 million Novo Nordisk milestone payment as non-product revenue
- December 31, 2024 pro forma cash balance of ~€655<sup>3</sup> million
- Looking ahead...
  - Investment in YORVIPATH launch in the U.S. and Europe Direct and TransCon CNP pre-launch activities
  - Stable R&D investment as current portfolio matures and early-stage pipeline renews

### Plan to be well-capitalized through cash flow breakeven and beyond

Unaudited preliminary estimate.
 Calculated as unaudited preliminary estimate of full year 2024 SKYTROFA revenue of ~€197 million plus ~€5 million of sales deductions related to prior years.
 Calculated as unaudited preliminary estimate of December 31, 2024 cash balance of €560 million plus expected payment from Novo Nordisk of \$100 million.

### Selected Milestones Expected in 2025



Q1 2025	Q2 2025	Q3 2025	Q4 2025
<b>TransCon CNP</b> Submit NDA to the FDA	<b>TransCon CNP</b> Topline Week 26 data from COACH	<b>TransCon CNP</b> Submit MAA to the EMA	<b>TransCon CNP</b> Submit IND or similar for hypochondroplasia
		<b>TransCon hGH</b> Adult GHD FDA Decision	<b>TransCon hGH</b> Adult GHD U.S. commercial availability, if approved
		<b>TransCon hGH</b> Submit IND or similar for basket trial in additional indications <sup>1</sup>	<b>TransCon PTH</b> Commercial launch expected in ≥5 additional Europe Direct countries

1. Planned for small for gestational age without catch-up growth ["SGA"]; Idiopathic short stature ["ISS"]; SHOX deficiency (including Turner syndrome).

## Thank you

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