

# Forward Looking Statements

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Founder & Chief Executive Officer

# PALSONIFY: A New Era in Acromegaly Treatment





**V** 

Broad Label with Strong Clinical Data on Biochemical and Symptom Control Sets the Foundation for a Strong Commercial Launch of PALSONIFY

### **Our Mission:**

To be the world's leading endocrine company that consistently pioneers new therapeutics to help patients better control their disease and improve their daily lives





# Palsonify Launch: Early Momentum and **Upcoming Key Metrics**

### **Strong Start of Palsonify Launch**



### What We Will Share in January

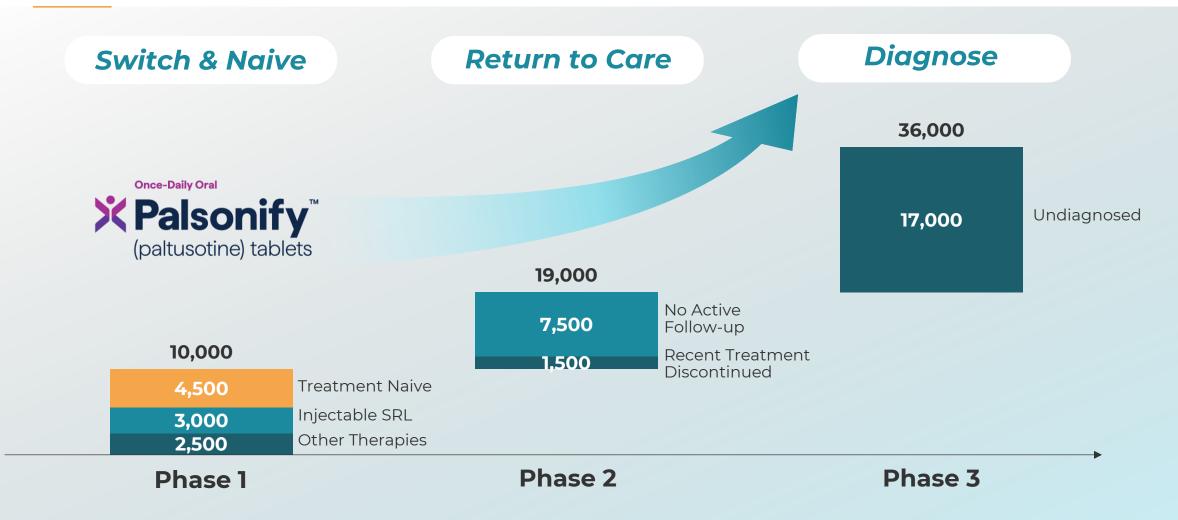








# Three-Phase Strategy for Helping More People with Acromegaly Get the Care They Need



# Continued Value Creation with Deep Pipeline of Transformative Drug Candidates

| Program                      |   | Preclinical                      | Phase 1                | Phase 2 | Phase 3    | Registration | Upcoming Milestones              |
|------------------------------|---|----------------------------------|------------------------|---------|------------|--------------|----------------------------------|
|                              | PATHFNDR'   | Acromegaly (US)                  |                        |         |            |              | Approved September 25, 2025      |
| Paltusotine (SST2 agonist)   | pathindra   | Acromegaly (EU)                  |                        |         |            |              | CHMP Opinion (1H 2026)           |
|                              | CAREFNDR<br>Featuring a Monimasive<br>Daily Regimen | Carcinoid syndrome               |                        |         |            |              | Phase 3 (2H 2025)                |
| Atumelnant (ACTH antagonist) | Calm-CAH  | Congenital adrenal h             | nyperplasia (adult)    |         |            |              | Phase 3 in Adult (2H 2025)       |
|                              |   | Congenital adrenal h             | nyperplasia (pediati   | ric)    |            |              | Phase 2/3 in Pediatric (2H 2025) |
|                              |   | ACTH dependent Cus               | shing's syndrome       |         |            |              | Phase 2/3 (1H 2026)              |
| CRN09682 (SST2+ NDC)         | BRAVESST2:  | NETs and SST2-expressolid tumors | essing                 |         |            |              | Phase 1/2                        |
| TSH antagonist               |   | Graves' disease*                 |                        |         |            |              |                                  |
| SST3 agonist                 |   | ADPKD                            |                        |         |            |              |                                  |
| PTH antagonist               |   | Hyperpara-<br>thyroidism         |                        |         |            |              |                                  |
| Oral GLP-1 nonpeptide        |   | Obesity                          |                        |         |            |              |                                  |
| Oral GIP nonpeptide          |   | Obesity                          |                        |         |            |              |                                  |
|                              | Dartners  | § SANWA KAGA                     | AKU KENKYUSHO CO., LTD | ı. Ra   | adionetics | loyal        |                                  |



Licensee of CRN01941 for

veterinary use

Licensee of targeted, nonpeptide

radiopharmaceuticals

Japan Development and Commercialization

Partner for Paltusotine

**Partners** 

# Carcinoid Syndrome: CAREFNDR Phase 3 for Paltusotine

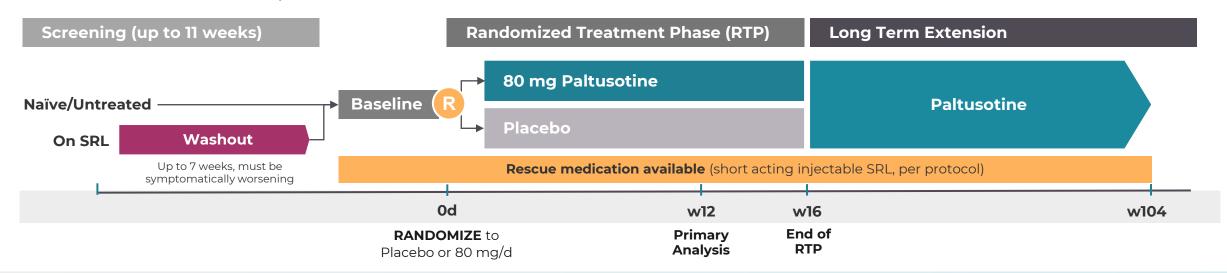
#### **Trial Size:**

141 patients, 2:1 randomization

#### **Key Eligibility Criteria:**

- Treatment naïve or currently untreated and actively symptomatic -OR- controlled on SRL therapy and symptom worsening upon washing out of treatment
- Grade 1 or 2 NET, Positive SSTR expression





**Primary Endpoint** 

Change from baseline in frequency of flushing

**Key Secondary Endpoint** 

Change from baseline in bowel movement frequency



**Additional Efficacy Endpoints** 

Flushing severity, bowel movement urgency. OLE to include assessment of tumor control (PFS)

# NETs & SST2+ Tumors: CRN09682 Phase 1/2 Study Will Be Proof of Concept for NDC Platform

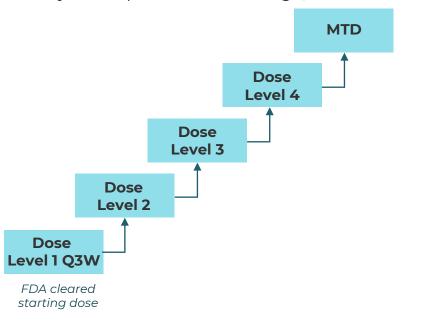


#### **Key Eligibility Criteria:**

- Metastatic or locally advanced inoperable NETs, NECs or other solid tumors
- Tumor progression on or after last line of therapy
- Positive SSTR expression by FDA approved SSTR PET/CT
- No carcinoid syndrome

#### Ph 1: Dose Escalation

• Bayesian Optimal Interval design, n=3-6/cohort



#### Ph 2: Dose Expansion\*

n=approximately 25/cohort

**Cohort 1: Pancreatic NET** Well-differentiated **Cohort 2: ex-Pancreatic NET** 

**Cohort 3: NEC Poorly Differentiated** (includes SCLC)

Well-differentiated

**Cohort 4: Other Solid Tumors** (e.g. Breast, Head-Neck)

\*representative of potential cohorts



#### **Key Endpoints**

- Safety & tolerability of CRN09682
- Define DLT/MTD and select Expansion Dose
- PK of CRN09682 and MMAF



commended

Expansion

#### **Efficacy Endpoints**

 Measure preliminary anti-tumor activity of CRN09682: ORR, DOR, PFS by RECIST v1.1

# Adult CAH: Design and Status, Phase 2 Atumelnant

#### **Key Eligibility Criteria**

#### N = 34 - 40

- Male or female participants ≥18 to 75 years. Age: ≥16 years (US)
- Classic 21-hydroxylase deficiency
- On ≥15mg
   Hydrocortisone
   equivalent daily dose
- A4 > 1.5 x ULN

#### **Treatment Arms:**

4 cohorts, each 12 weeks (N=6-12)

40 mg Once Daily (PM Dosing) (n=11)

80 mg Once Daily (PM Dosing) (n=11)

120 mg Once Daily (PM Dosing) (n=6)

80 mg Once Daily (AM Dosing) + GC Reduction (n=6-12)

**Open-Label Extension includes Patients from All Cohorts** 

Pre-trial glucocorticoid therapy (dose and regimen) maintained throughout the trial for first 3 cohorts

#### Objectives: Evaluate the Safety, Efficacy, and Pharmacokinetics of Atumelnant

Primary Endpoint: Change from baseline in morning serum A4 at week 12

**Secondary Endpoint:** Change from baseline in morning serum 17-OHP at week 12

**Primary Safety Assessment:** Incidence of TEAEs throughout the study



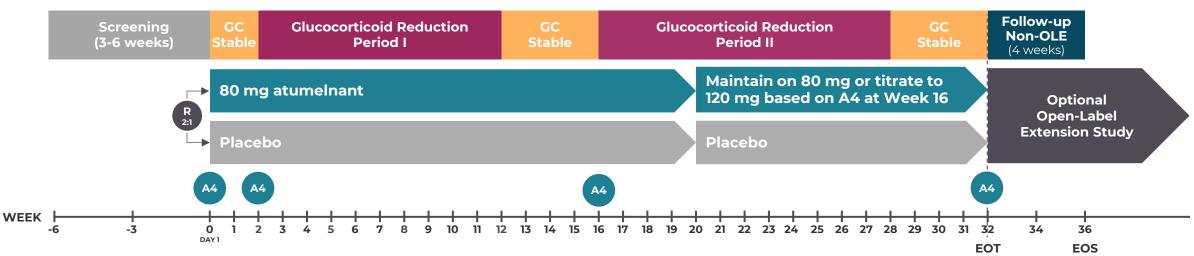
# Adult CAH: Global Phase 3 Trial Designed to Assess Normalization of Androgen and Glucocorticoids

#### **Key Eligibility Criteria (N = 150):**

- Male or female participants ≥ 18 to 75 years.
- Classic 21-hydroxylase deficiency
- Stable GC dose for 2 months

- A4 > ULN¹ with supraphysiologic GC dose (≥11 mg/m²/day)
- A4 > ULN<sup>1</sup> with physiologic GC dose (<11 mg/m<sup>2</sup>/day)
- Normal A4<sup>2</sup> with supraphysiologic GC dose (≥15 mg/m²/day)





### 1 Primary Endpoint

Proportion of participants with morning **post-GC** A4 ≤ ULN who are on physiologic GC replacement at Week 32

### **2** Key Secondary Endpoints

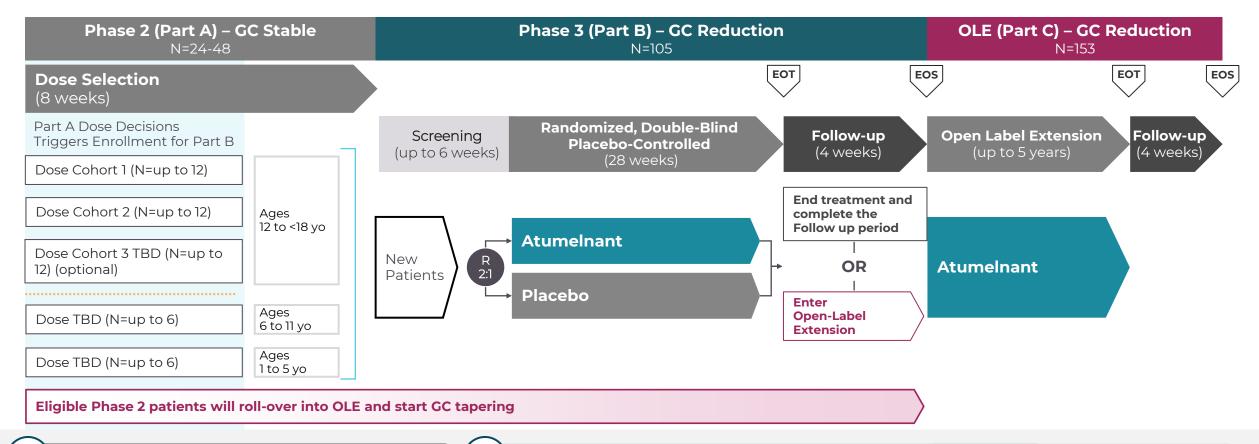
- Percent change from baseline in serum morning pre-GC A4 at week 2
- Percent change from baseline in serum morning pre-GC 17-OHP at week 32
- Proportion of participants with morning pre-GC A4 ≤ ULN who are on physiologic GC replacement at Week 32
- Percent change from baseline in GC daily dose when **post-GC** A4 ≤ ULN at week 32

#### **3** Other Secondary Endpoints

Defined to evaluate the impact of atumelnant on the clinical signs, symptoms, co-morbidities and outcomes of CAH

# Pediatric CAH: Global Phase 2/3/OLE Balance-CAH Operationally Seamless Trial





#### **Primary Endpoint**

- Phase 2: Change from baseline in morning serum A4 at Week 8
- Phase 3: Percent change from baseline in GC daily dose at Week 28 while serum early morning A4 ≤ ULN
- OLE: Change from baseline in morning serum A4 over time

#### **Key Secondary Endpoints (Phase 3)**

- Change from baseline in morning serum A4 at Week 4
- Change from baseline in morning serum 17-OHP at Week 4
- Proportion of participants with physiologic GC dose while serum early morning A4 <ULN at Week 28

#### **Key Eligibility Criteria**

- Male or female participants 1 to <18 years.
- Classic 21-hvdroxvlase deficiency
- Stable GC dose for 1 month
- A4 > ULN with supraphysiologic GC dose (≥11 mg/m<sup>2</sup>/day)

R=Randomization; EOT=End of Treatment; EOS=End of Study. CAH=congenital adrenal hyperplasia; GC=glucocorticoid.



# Executing on Our Strategy to Make PALSONIFY the Foundation of Acromegaly Care

Leverage synergistic field teams and targeted marketing to educate on best-in-class profile and streamline access

#### **PATIENTS**

Activate patients to start, switch, or resume treatment

#### **PROVIDERS**

Make PAI SONIFY the therapy of choice

#### **PAYERS**

Demonstrate value proposition to facilitate access

ACTIVATE

ADOPT

ACCESS

ADHERE

# Activating Both Naïve and Switch Patients

### Reinforcing IGF-1 and Symptom Control in a Once-Daily Oral

#### **PATIENTS**

Activate patients to start, switch, or resume treatment

### **All 22**

U.S. OLE patients in process of transitioning onto commercial product

95% | 5%

Initial mix of filled Rx medically switch | naïve patients

### **PATIENT ACTIVATION**

#### **ACROMEGALY REALITY**

Disease education campaign to raise awareness of the lived realities of acromegaly and support earlier recognition and diagnosis



Comprehensive patient support services to assist with access, adherence and personalized care throughout the treatment journey

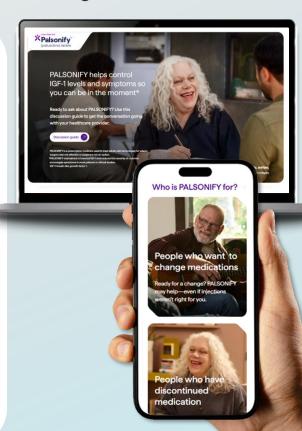
### Branded campaign launched

Launched patient website, omnichannel media campaign, patient discussion guide and tools. Activated our CRM system. Nurse educators will directly support patient education.









# Leading with Efficacy First to Engage Providers

#### **PROVIDERS**

Make PALSONIFY the therapy of choice

95%

Top prescriber targets reached

70% | 30%

Prescriber setting Community | PTC

### Key messages that have resonated among active prescribers:

- Strong efficacy data in both switch and naïve patients
- IGF-1 normalization and symptom improvement
- Ease of administration with a once-daily oral
- CrinetiCARE services and access support

"PALSONIFY could change the face of acromegaly treatment."

- Endocrinologist at PTC center in Massachusetts

"PALSONIFY is great for patients with uncontrolled symptoms and breakthrough symptoms."

Endocrinologist at PTC center in California

# Positive Initial Response from Payers: Rapid Access

Strong value proposition to payers – expect formulary coverage in most plans within 6-9 months

#### **PAYERS**

Demonstrate value proposition to facilitate access

### 50% | 50%

Reimbursed | Quickstart for filled Rx

### **Up to 12 Months**

Duration of approved prior authorizations



#### **Unprecedented Safety and Efficacy**

Ability to achieve rapid biochemical and symptom control based on Phase 3 data



#### **Maintain Control**

Limit patient and societal burden of uncontrolled acromegaly



#### **Optimize Treatment Paradigm**

Ensure patients getting intended clinical benefit



**Improve Patient Adherence and Outcomes** 

Once-daily oral dosing



## Financial Results

|               | Three Months Ended |               |  |  |  |
|---------------|--------------------|---------------|--|--|--|
| (in millions) | September 30, 2025 | June 30, 2025 |  |  |  |
| Revenues      | \$ 0.1             | \$ 1.0        |  |  |  |
| R&D Expenses  | (90.5)             | (80.3)        |  |  |  |
| SG&A Expenses | (52.3)             | (49.8)        |  |  |  |
| Net Loss      | \$ (130.1)         | \$ (115.6)    |  |  |  |

|                            | October 28, 2025 |
|----------------------------|------------------|
| Common Stock Outstanding   | 94.9 Million     |
| Fully Diluted Share Count* | 111.9 Million    |

## \$1.1 Billion Cash Balance Funds Current Operating Plan into 2029

### \$1.1 Billion

Cash, cash equivalents, & investments as of September 30, 2025

### Into 2029

Cash runway based on current operating plan

### **\$340 Million - \$370 Million**

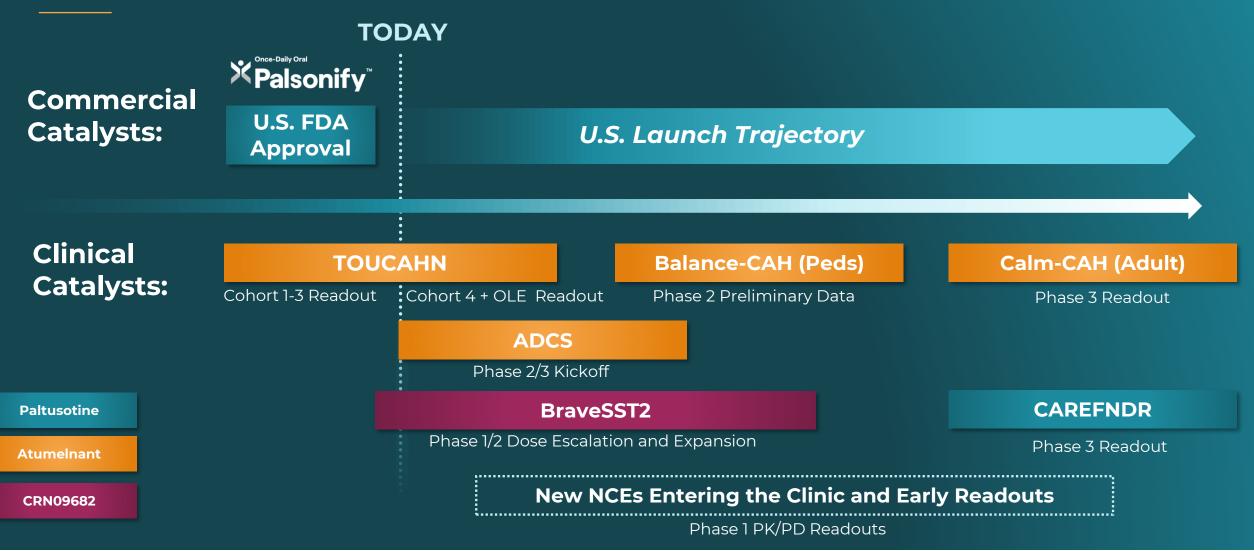
2025 operating cash burn guidance

### **Supports Strategic Initiatives Including:**

- Commercialization of PALSONIFY
- Pipeline programs and innovation from discovery
- Optionality to prioritize or pursue opportunities to enhance value across our portfolio



# Poised to Deliver Multiple Commercial & Clinical Catalysts in the Next 24+ Months



## **Q&A Session**



**SCOTT STRUTHERS** Founder and Chief Executive Officer



**STEPHEN BETZ** Founder and Chief Scientific Officer



**ISABEL KALOFONOS** Chief Commercial Officer



**TOBY SCHILKE** Chief Financial Officer



**DANA PIZZUTI** Chief Medical Officer



**ALAN KRASNER** Chief Endocrinologist