



Corporate presentation

July 2022

An integrated orphan drug company, focusing on late-stage development for commercialization

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# Agenda



- 1. An integrated orphan drug company, focusing on late-stage development for commercialization
- 2. Emcitate®
  - Clinical development program
  - Commercial opportunity
- 3. Aladote®
  - Clinical development program
  - Commercial opportunity
- 4. The orphan drug segment and path to market
- 5. Summary
- A. Appendix

# Significant progress towards Emcitate® marketing applications in the US and Europe in 2023



Strong momentum continues in 2022

- Fruitful regulatory interactions clarify the regulatory path forward for Emcitate
  - Targeting Emcitate EU MAA submission H1 2023.
  - Targeting Emcitate US NDA submission mid-2023 under the Fast Track Designation.
  - FDA acknowledges that effects on T3 levels and the manifestations of chronic thyrotoxicosis could provide a basis for Emcitate approval.
  - For the US submission, a 30-day, placebo-controlled study in 16 patients will be conducted to verify the results on T3 levels seen in previous clinical trials and publications.
- The outcome from the regulatory interactions increases the likelihood of success for Emcitate and the probability to receive *Priority Review Voucher* (PRV) in the US.
- Triac Trial II study with Emcitate fully recruited results Q1 2024.
- Orphan drug designation (ODD) for Emcitate for RTH-β in the US and EU.
- SEK 180 million fully guaranteed preferential rights issue approved at EGM.

# An integrated orphan drug company, focusing on late-stage development for commercialization

- Dedicated orphan drug development company with two late-stage orphan drug assets: Emcitate® and Aladote®
- Target MAA/NDA submissions for Emcitate in 2023 and for Aladote in 2024/2025
- Highly attractive **orphan drug segment** with potential >\$1Bn annual sales opportunity
- Plan to launch through small inhouse commercial organization in the EU and US
- Combined core expertise in late-stage orphan clinical development, registration and commercialization with experience from:



Listed on NASDAQ Stockholm (EGTX) HQ in Stockholm, Sweden

















# Orphan drug segment – a highly attractive opportunity



### Shorter clinical development time<sup>1</sup>

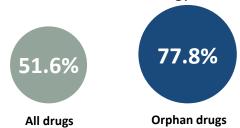
Phase II to launch Average # of years



### Higher probability of success<sup>3</sup>

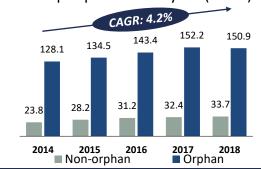
### Phase III to approval

POS in metabolic/endocrinology indications



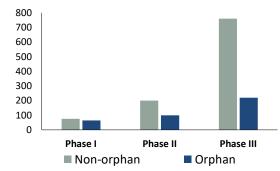
### Higher attainable prices<sup>2</sup>

Mean cost per patient and year (USDk)

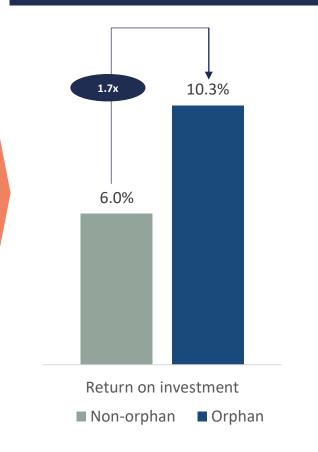


### Fewer patients for clinical trials<sup>4</sup>

### Patients per trial

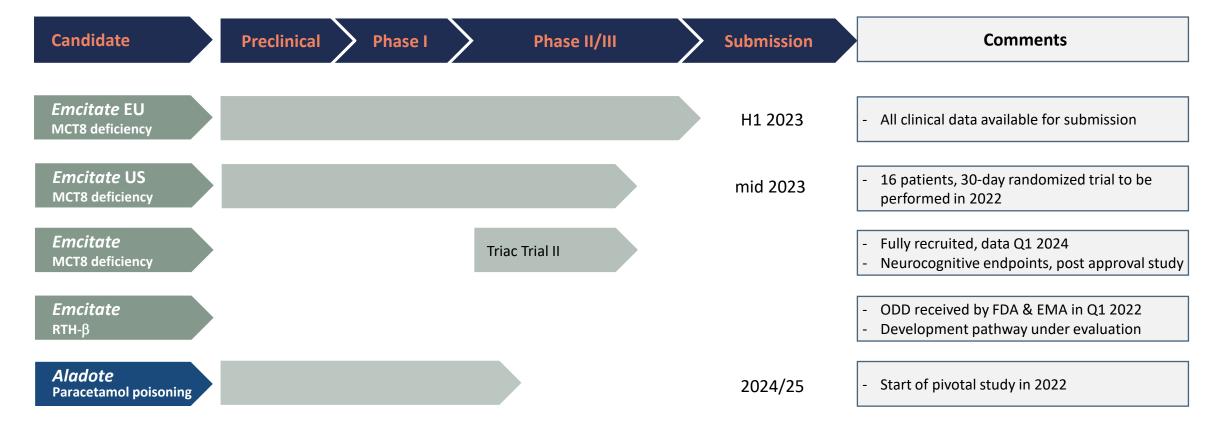


### **Orphan drugs attractive returns**<sup>5</sup>



## **Pipeline overview**

Planned Emcitate EU and US filings in 2023





## Two highly promising orphan drug candidates

### Emcitate® – Therapy for MCT8 deficiency

- MCT8 deficiency affects ~1:70,000 males: high unmet medical need, no available treatment. No competing sponsored products in clinical development
- ODD in EU & US
- US Rare Pediatric Disease Designation, eligible for Priority Review Voucher. Fast track designation granted by FDA
- Triac Trial I (Phase IIb) completed with **significant** and **clinically** relevant effects on **T3 levels** and the manifestations of **chronic thyrotoxicosis**
- Real-world data published 2021 confirms long-term efficacy and safety of Emcitate
- MAA based on existing clinical data in H1 2023
- NDA in mid 2023, after conducting a 30 days placebo-controlled study in 16 patients to verify the results on T3
- Triac Trial II fully recruited; to establish the effects of early intervention on neurocognitive development, previously seen in Triac Trial I. Results expected in Q1 2024
- More than 150 patients are being treated with Emcitate on a named patient basis

# Aladote® – To prevent acute liver injury caused by paracetamol poisoning

- Paracetamol poisoning is one of the most common overdoses with >175,000 hospital admissions globally per annum
- No adequate treatment exists for increased risk patients
- Orphan drug designation (ODD) granted in 2019 in the US
- Ongoing dialogue with EMA on the appropriate scope of the indication for an ODD in the EU
- Successful results from Phase Ib/IIa study in paracetamol overdosed patients
- Pivotal Phase IIb/III study planned for marketing authorization application in both US and EU, targeting study start in 2022
- No competing products in clinical development

resentation | Egetis Therapeutics | 2022-07-12

# MCT8 deficiency: a detrimental condition with significant unmet medical need



#### What is MCT8 deficiency?

- · Genetic X-linked disorder
- Impaired thyroid hormone trafficking across cellular membranes
- MCT8 is a key thyroid hormone transporters in the body
- Prevalence 1:70,000 males



Patients with MCT8 Deficiency1)

#### What does it mean?

- Non-functional MCT8 protein: T3 cannot cross blood-brainbarrier
- Low amounts of thyroid hormone in the brain & CNS
- Disrupted feedback loop results in a compensatory increase in circulating thyroid hormone

 Simultaneous too high & too low thyroid hormone in different tissues

### What are the challenges?

- Patients appear normal at birth
- Initial symptoms within the first months of life
- Severe intellectual disability
- Most patients never able to sit or walk; limited ability to communicate
- Life-long morbidity: agitation, CV symptoms, wasting & impaired life expectancy

 Heavily dependant on caregivers resulting in very high disease burden

## How do you manage the disease?

- No available therapy
- Easy diagnosis once considered with readily available, low-cost lab-test
- Large proportion of patients remain undiagnosed with significant delay to diagnosis



 Significant unmet medical need: humanitarian, health economic, societal

### Quick facts from natural history<sup>2</sup>

Median onset of symptoms: 4 months

Median age of diagnosis: 24 months

Patients surviving into adulthood: 70%

Severe intellectual disability: 100%

Ability to sit independently: 8%

Hypotonia, hypertonia

& persistence of primitive reflexes: 90%

Severe underweight: 75%

Cardiac arrythmias (PAC): 76%

Median life expectancy: 35 years

Life long 24-hour care: 100%

## Orphan drug candidate

with clear scientific and mechanistic rationale and established safety profile

### Difference normal MCT8 and deficiency of MCT8

 Thyroid hormone T3 requires transporters such as MCT8 to enter the target cells

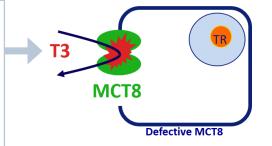
### Normal MCT8

- Functional thyroid gland producing
- Production of functional MCT8
- → T3 cross cell membrane and enters target cell

# MCT8 Norma

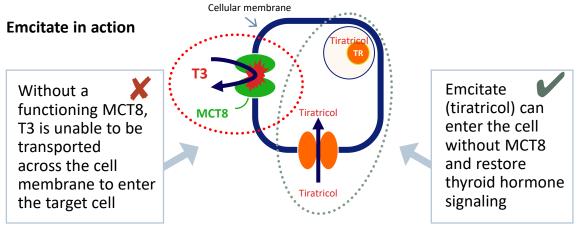
### Mutated MCT8 X

- Functional thyroid gland producing T3
- Absence or loss of function of MCT8 on the cell surface
- → T3 cannot cross cell membrane and fails to enter cells



### **Emcitate (tiratricol) – Addressing the MCT8 deficiency**

- Tiratricol is a thyroid hormone analogue with high chemical and structural similarity to T3
- Unlike T3, tiratricol can cross cellular membranes without a functional MCT8 transporter
- Tiratricol can bypass the problem in patients with MCT8 deficiency, enter MCT8 deficient cells and restore thyroid hormone signalling
- Experience from 40 years on the French market in a different indication, owned and controlled by company



### **Emcitate® Overview**



Lead candidate for addressing MCT8 deficiency, a condition with high unmet medical need and no available treatment



- Triac Trial I completed with significant and clinically relevant effects
- Erasmus Medical Center cohort study confirms long-term efficacy and safety in MCT8 deficiency patients for up to 6 years (Oct 2021)
- Triac Trial II, early intervention trial in young subjects to establish the effect on the neurocognitive development, previously seen in Triac Trial I. Fully recruited April 2022. Results expected Q1 2024



- Orphan drug designation in EU & US, US Rare Pediatric Disease Designation eligible for Priority Review Voucher
- Fast track designation granted by FDA (Oct 2021)
- Intend to submit MAA to the EMA based on existing clinical data H1 2023
- US **NDA submission planned mid-2023**: A 30-day, placebo-controlled study in 16 patients will be conducted to verify the results on T3 levels seen in previous clinical trials and publications



- Estimated 10k 15k MCT8 deficiency patients (1:70k males), no sponsor-initiated trials ongoing in MCT8 deficiency
- Analogue orphan drugs priced at premium
- Launched disease awareness initiatives to support diagnosis of MCT8 deficiency
- More than 150 patients are being treated with Emcitate on a named patient or compassionate use basis, following individual regulatory approvals from national regulatory agencies
- Expected market exclusivity is 12y in EU (ODD 10y, pediatric ext. 2y), 7.5y in US (ODD 7y, pediatric 0.5y)

# Overview of completed Phase IIb – Triac Trial I



- Evaluate the efficacy and safety of oral administration of tiratricol in male patients with MCT8 deficiency of all ages
- Highly significant primary outcome Change in T3 serum concentrations
- Safe and tolerable
- Results published in The Lancet 2019

Secondary objective and results

- Change in other thyroid hormone function tests, thyrotoxic symptoms and markers
- Significant and clinically relevant effects observed across secondary endpoints

**Description** 

- An international, single-arm, open-label, Phase II trial
- ClinicalTrials.gov identifier: NCT02060474

# of patients

46 MCT8 patients in 9 countries

**Timetable** 

- Initiated in October 2014 (first patient in)
- Completed in June 2018

### THE LANCET

**\ (1)** 

Effectiveness and safety of the tri-iodothyronine analogue Triac in children and adults with MCT8 deficiency: an international, single-arm, open-label, phase 2 trial

Stefan Groeneweg, Robin P Peeters, Carla Moran, Athanasia Stoupa, Françoise Auriol, Davide Tonduti, Alice Dica, Laura Paone, Mara Rozenkova, Jana Malikova, Adri van der Walt, Irenaeus F.M. de Coa, Anne McGowan, Gret a Lyons, Fernke K. Aarsen, Diana Barca, Ingrid M. van Beynum, MariekeM van der Knoop, Jurgen Jansen, Martien Manshande\*, Roelineke J. Lunsing, Stan Nowak, Corstiaan A. den Uil, M. Carola Zillikens, Frank E Visser, Paul Vrijmoet h. Marie Clair e Y de Wit, Nicole I Wolf, Angelique Zandstra, Gautam Ambegoonkar, Yogen Singh, Yalanda B de Rijke, Marco Medici, Enrico S Bertini, Sylvia Deposeter, Jan Lebl, Marco Cappa, Linda De Meideir\*, Heiko Krude, Dana Craiu, Federica Zibordi, Isabelle Oliver Petit, Michel Polak, Krishna Chatterjee, TheoJ Visser\*, W Edward Visser

Background Deficiency of the thyroid hormone transporter monocarboxylate transporter 8 (MCT8) causes severe LinuxiDiabetesEnd intellectual and motor disability and high serum tri-iodothyronine (T<sub>i</sub>) concentrations (Allan-Herndon-Dudley Patiented Online syndrome). This chronic thyrotoxicosis leads to progressive deterioration in bodyweight, tachycardia, and muscle 109/31, 2019 wasting, predisposing affected individuals to substantial morbidity and mortality. Treatment that safely alleviates peripheral thyrotoxicosis and reverses cerebral by pothyroidism is not yet available. We aimed to investigate the effects of treatment with the T, analogue Triac (3.3', 5-tri-iodothyroacetic acid, or tiratricol), in patients with MCT8 deficiency.

Methods In this investigator-initiated, multicentre, open-label, single-arm, phase 2, pragmatic trial, we investigated the "OrMantanae ded nAcque effectiveness and safety of oral Triac in male paediatric and adult patients with MCT8 deficiency in eight countries in 2018, Prof OrMantanae ded nAcque Europe and one site in South Africa. Triac was administered in a predefined escalating dose schedule-after the initial dose of once-daily 350 µg Triac, the daily dose was increased progressively in 350 µg increments, with the goal of attaining serum total T<sub>3</sub> concentrations within the target range of 1-4-2-5 nmol/L. We assessed changes in several clinical and biochemical signs of hyperthyroidism between baseline and 12 months of treatment. The prespecified Foota Freeton MO. primary endpoint was the change in serum T, concentrations from baseline to month 12. The co-primary endpoints MMMediciMO, Prof T) Vaser Ph were changes in concentrations of serum thyroid-stimulating hormone (TSH), free and total thyroxine (T.), and total WEVERSHIE, Septia reverse T, from baseline to month 12. These analyses were done in patients who received at least one dose of Triac and had at least one post-baseline evaluation of serum throid function. This trial is registered with Clinical Trials.gov, number (M. van Beynum MD).

Findings Between Oct 15, 2014, and June 1, 2017, we screened 50 patients, all of whom were eligible. Of these patients, four (896) patients decided not to participate because of travel commitments. 46 (9296) patients were therefore enrolled MMYAN det Knoop MSC. in the trial to receive Triac (median age 7-1 years [range 0-8-66-8]) . 45 (98%) participants received Triac and had at MCY deWit MD), Department least one follow-up measurement of thyroid function and thus were included in the analyses of the primary endpoints. Of these 45 patients, five did not complete the trial (two patients withdrew [travel burden, severe pre-existing comorbidity], one was lost to follow-up, one developed of Graves disease, and one died of sepsis). Patients required a mean dose of 38.3 µg/kg of bodyweight (range 6.4-84.3) to attain T, concentrations within the target range. Serum T, (Poet's 50 RIBE PRO) concentration decreased from 4-97 nmol/L (SD 1-55) at baseline to 1-82 nmol/L (0-69) at month 12 (mean decrease Medicine 3-15 nmol/L, 95% CI 2-68-3-62; p<0-0001), while serum TSH concentrations decreased from 2-91 mU/L (SD 1-68)

PROME Z BREETS MOL BEARMS to 1-02 mU/L (1-14; mean decrease 1-89 mU/L, 1-39-2-39; p<0-0001) and serum free T, concentrations decreased Medical Centre, Rottestams, from 9.5 pmol/L (SD 2.5) to 3.4 (1.6; mean decrease 6.1 pmol/L (5.4-6.8; p<0.0001). Additionally, serum total T. Netherlands, Welkome Trust concentrations decreased by 31 · 6 nmol/L (28 · 0 - 35 · 2; p<0 · 0001) and reverse T, by 0 · 08 nmol/L (0 · 05 - 0 · 10; p<0 · 0001). Seven treatment-related adverse events (transiently increased perspiration or irritability) occurred in six (13%) patients. 26 serious adverse events that were considered unrelated to treatment occurred in 18 (39%) patients (mostly hospital Cambridge UK (C MOTOL MS) admissions because of infections). One patient died from pulmonary sepsis leading to multi-organ failure, which was unrelated to Triac treatment

Interpretation Key features of peripheral thyrotoxicosis were alleviated in paediatric and adult patients with MCT8 Necestation News Tolkhorts University deficiency who were treated with Triac. Triac seems a reasonable treatment strategy to ameliorate the consequences of untreated peripheral thyrotoxicosis in patients with MCT8 deficiency.

Funding Dutch Scientific Organization, Sherman Foundation, NeMO Foundation, Wellcome Trust, UK National and Genetic, Children's Institute for Health Research Cambridge Biomedical Centre, Toulouse University Hospital, and Una Vita Rara ONLUS. Hospital Toulouse University

of Paedlatric Cardiology

Neurology (FFM de Coo M.C)

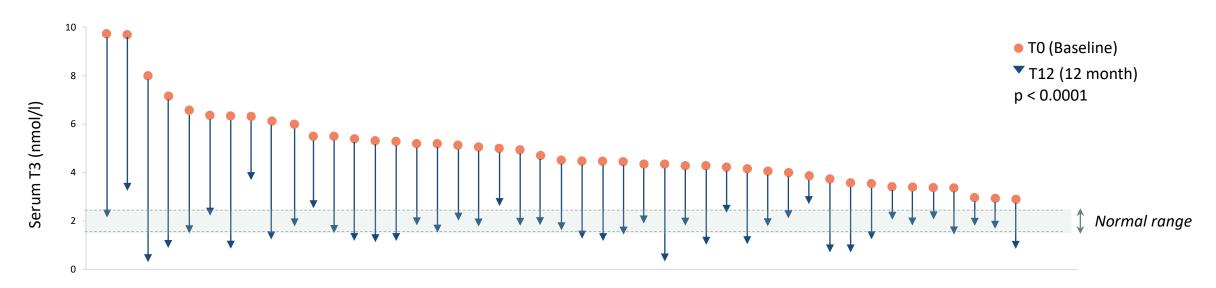
K Chatteriee FRCP:- Paediatric Prof M Possi M.Di. Departmen

www.thelancet.com/diabetes-endocrinology Published online July 31, 2019 http://dx.doi.org/10.1016/52213-8587(19)30155-X

Corporate presentation | Egetis Therapeutics | 2022-07-12 Source: Groeneweg et al, Lancet D&E 2019;

# Consistent, clinically relevant and highly significant results

Reached target level serum T3 in completed Phase IIb trial (Triac Trial I)



Endpoints	Baseline mean ( $\pm$ SD)	12 months mean ( $\pm$ SD)	Difference in means (95% CI)	p-value
Serum T3 (nmol/L)	4.97 (± 1.55)	1.82 (± 0.69)	-3.15 (-3.62, -2.68)	<0.0001
Weight to age (z score)	-2.98 ( $\pm$ 1.93)	-2.71 ( $\pm$ 1.79)	0.27 (0.03, 0.50)	0.025
Resting heart rate (bpm)	112 ( $\pm$ 23)	104 ( $\pm$ 17)	-9 <i>(-16, -2)</i>	0.01
Mean heart rate 24 h (bpm)	102 ( $\pm$ 14)	97 ( <i>±</i> 9)	-5 <i>(-9, -1)</i>	0.012
SHBG (nmol/L)	212 ( $\pm$ 91)	178 ( $\pm$ 76)	-35 <i>(-55, -15)</i>	0.0013
Total cholesterol (mmol/L)	3.2 ( $\pm$ 0.7)	3.4 <i>( ± 0.7)</i>	0.2 (0.0, 0.3)	0.056
CK (U/L)	108 ( $\pm$ 90)	161 ( $\pm$ 117)	53 <i>(27, 78)</i>	<0.0001

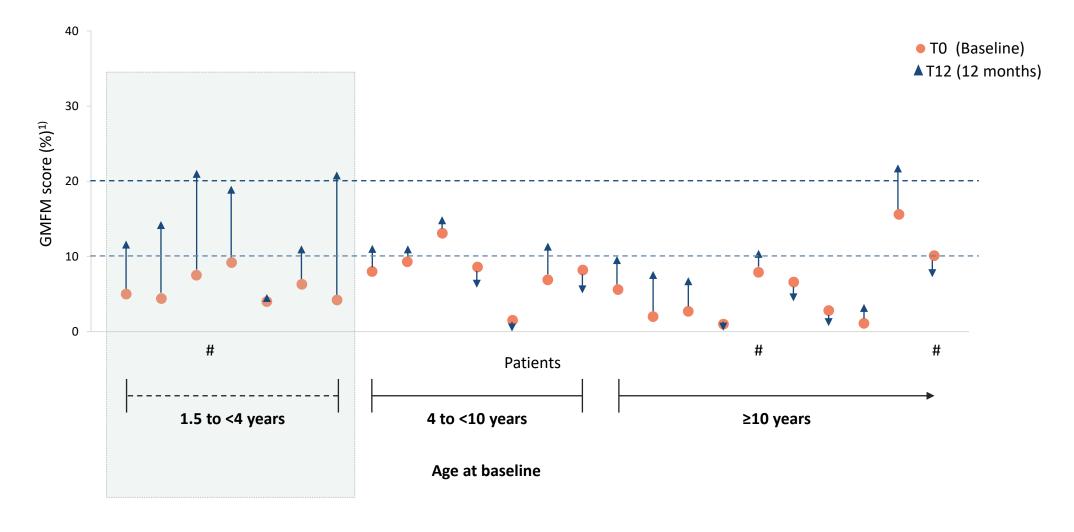
Source: Groeneweg et al; Lancet D&E 2019

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# Indication of positive effect on neurocognitive development

In the youngest patients which is further studied in ongoing Triac Trial II





# New data confirms long-term efficacy and safety of Emcitate® in MCT8 deficiency patients



Published in October, 2021

ACCEPTED MANUSCRIPT

Long-term efficacy of T3 analogue Triac in children and adults with MCT8 deficiency: a real-life retrospective cohort study 3

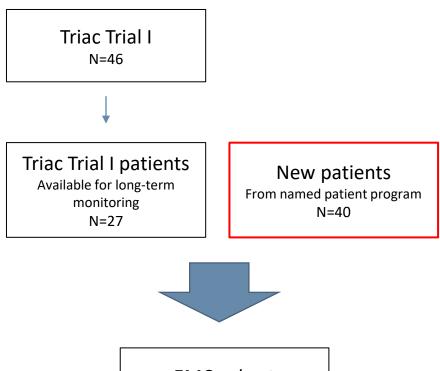


Ferdy S van Geest, Stefan Groeneweg, Erica L T van den Akker, Iuliu Bacos, Diana Barca, Sjoerd A A van den Berg, Enrico Bertini, Doris Brunner, Nicola Brunetti-Pierri, Marco Cappa ... Show more Author Notes

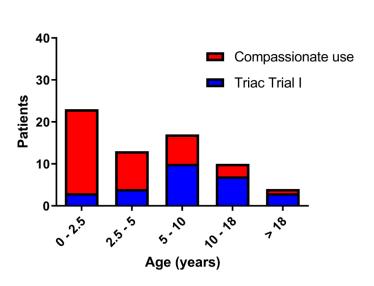
- Investigator-initiated real-world cohort study at 33 sites conducted by the Erasmus Medical Center
- Investigated efficacy and safety of Emcitate in 67 patients with MCT8 deficiency
  - Median baseline age of 4.6 years (range: 0.5–66 years) and were treated with tiratricol for up to 6 years, with a median of 2.2 years (range 0.2 – 6.2 years)
  - The primary endpoint in the study was the change in serum T3 concentration from baseline to last-available measurement
  - The pre-specified secondary endpoints were key measurements of clinical complications of chronic peripheral thyrotoxicosis

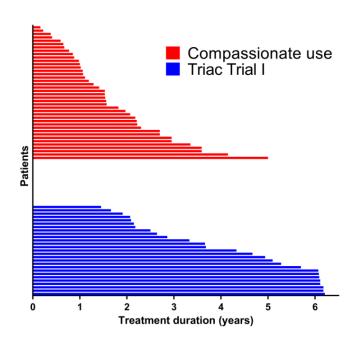
# New patient cohort of equal size to the Triac Trial I

Long term follow up up to >6 years



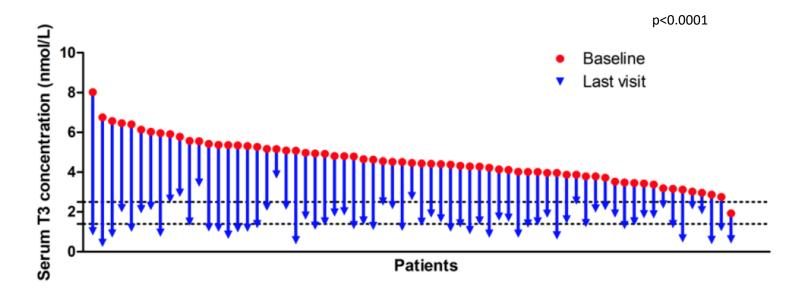
EMC cohort Published in JCEM N=67

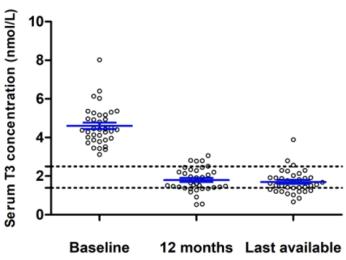




# New cohort confirms primary endpoint results in Triac Trial I

Fast and durable normalization of T3 values in almost all patients





# Consistent, clinically relevant and highly significant results

- across endpoints
- Data confirm the positive results from previous study, Triac
   Trial I
- Normalization of serum T3 corresponds to improvement in thyroid hormone status in end target tissues
- Beneficial effects are maintained or continue to improve over time, up to six years
- Consistent efficacy seen across key clinical and biochemical parameters that were sustainably alleviated in patients with MCT8 deficiency regardless of age

Table 2: Changes from baseline to last visit in predefined outcomes

	Baseline mean (SD)	Last visit mean (SD)	Mean change (95% CI)	P value
Primary outcome				
T3 (nmol/L; n=67)	4.58 (1.11)	1.66 (0.69)	-2.92 (-3.23 to -2.61)	<0.0001
Secondary outcomes				
Anthropometric parameters and heart rate				
Body weight (kg; n=58)	17.8 (12.1)	23.6 (14.5)	5.7 (4.2 to 7.2)	
Weight-for-age Z score (n=58)	-2.81 (1.94)	-2.64 (1.81)	0.17 (-0.18 to 0.53)	0.3263
Δ Weight-for-age – predicted weight-for-age Z score (n=55)	0.07 (1.83)	0.79 (1.92)	0.72 (0.36 to 1.09)	0.0002
Height (cm; n=44)	101 (21)	116 (23)	15 (12 to 19)	
Height-for-age Z score (n=44)	-1.84 (1.77)	-1.92 (1.51)	-0.09 (-0.50 to 0.32)	0.6705
Δ Height-for-age – predicted height-for-age Z score (n=43)	-0.44 (1.38)	0.14 (1.41)	0.58 (0.12 to 1.05)	0.0139
Weight-for-height Z score (n=44)	-2.02 (2.49)	-1.50 (2.44)	0.52 (-0.35 to 1.39)	0.2358
Heart rate (bpm; n=48)	113 (21)	97 (20)	-17 (-24 to -10)	<0.0001
Heart rate-for-age Z score (n=48)	1.59 (0.89)	0.96 (1.01)	-0.64 (- 0.98 to -0.29)	0.0005
Thyroid function tests				
TSH (mU/L; n=62)*	3.32 (2.30)	0.95 (0.73)	-2.38 (-2.98 to -1.77)	<0.0001
Free T4 (pmol/L; n=64)	9.5 (2.3)	3.4 (1.6)	-6.1 (-6.7 to -5.4)	<0.0001
T4 (nmol/L; n=63)	54.2 (11.8)	18.1 (9.8)	-36.1 (-39.5 to -32.7)	<0.0001
Peripheral markers				
Sex hormone-binding globulin (nmol/L; n=48)	245 (99)	209 (92)	-36 (-57 to -16)	0.0008
Creatinine (µmol/L; n=47)	32 (11)	39 (13)	7 (6 to 9)	<0.0001
Creatine kinase (U/L; n=47)*	110 (87)	128 (80)	18 (-8 to 45)	0.2166

All outcomes were assessed in all patients who received Triac treatment longer than the mean time to optimal dose (5.0 months; N=64). Data are mean. Body weight-for-age Z scores were calculated using TNO growth calculator and heart rate-for-age Z scores were calculated using the Boston Z score calculator. Abbreviations: T3=tri-iodothyronine. TSH=thyroid-stimulating hormone. T4=thyroxine. \*TSH and creatine kinase concentrations were log-transformed to ensure a normal distribution before paired t tests were done (non-transformed means [SDs] and mean changes [95% CIs] are presented for the sake of interpretability).



# Egetis intends to submit MAA for Emcitate® to EMA in H1 2023 based on existing clinical data



Press release issued Dec 13, 2021

- Based on regulatory interactions, Egetis concludes that available data from Triac Trial I and recently published long-term data are sufficient for a Marketing Authorisation Application (MAA) in Europe
- Having all clinical data required for regulatory submission already at hand significantly reduces the remaining risk for Emcitate
- The ongoing Triac Trial II will continue to further establish the effects of early intervention on the neurocognitive development aspects of the disease

# Egetis intends to submit a marketing authorisation application for Emcitate® to the European Medicines Agency based on existing clinical data

- Egetis concludes, based on recent regulatory interactions, that available Triac Trial I data together
  with recently published long-term data are sufficient for a Marketing Authorisation Application in
  Europe
- Having all clinical data required for regulatory submission already at hand significantly reduces the remaining risk for Emcitate
- Revised submission timelines will be communicated as soon as all parts of the regulatory dossier are confirmed
- Egetis will host a webcast today at 15:00 CET (9:00am ET)

Stockholm, Sweden, December 13, 2021 - Egetis Therapeutics AB (publ) (Nasdaq Stockholm: EGTX) today announced that after a pre-submission meeting held last week with concerned European regulatory agencies (EMA's Rapporteur and Co-Rapporteur), the Company concludes that the clinical data from the Triac Trial I (Groeneweg et al. 2019), together with the data from long-term treatment with Emcitate (tiratricol) for up to six years in 67 patients (van Geest et al. 2021) will be sufficient for a regulatory review of a Marketing Authorisation Application (MAA) to the European Medicines Agency for the treatment of monocarboxylate transporter 8 (MCT8) deficiency. Thus, all clinical data necessary for regulatory submission is already available. The ongoing Triac Trial II will continue to further establish the effects of early intervention on the neurocognitive development aspects of the disease.

"We are delighted with the outcome of the pre-submission meeting, giving us a clear path to our MAA submission, and subsequent regulatory review, based on existing clinical data. Having all clinical data required for regulatory submission already at hand significantly reduces the remaining risk for Emcitate and could also potentially enable an earlier submission in Europe than we had previously expected. This is a substantial opportunity for us and the European patients suffering from MCT8 deficiency. In parallel, as part of our efforts to make Emcitate available as soon as possible, we continue our dialogues with regulatory authorities in other jurisdictions to obtain their views on the available clinical data and its implications for regulatory submissions" said Nicklas Westerholm, CEO, Egetis Therapeutics.

# Treatment effects on T3 levels in MCT8-deficiency could provide a basis for marketing approval in the US – NDA targeted in mid 2023



Press release issued Jan 18, 2022

- FDA acknowledges that a treatment effect on T3 levels and the manifestations of chronic thyrotoxicosis in MCT8-deficiency could provide a basis for marketing approval also in the US.
- A small, 30-day, placebo-controlled study in 16 treated patients, to be identified primarily through our existing named patient program, will be conducted to verify the results on T3 levels seen in previous clinical trials and publications in a randomized controlled setting.
- An NDA in the US is targeted to be submitted in mid 2023 under the Fast Track Designation.
- A major step towards marketing authorization and increases the likelihood of success for *Emcitate* and the probability to receive a US Rare Pediatric Disease **Priority Review Voucher** (PRV).

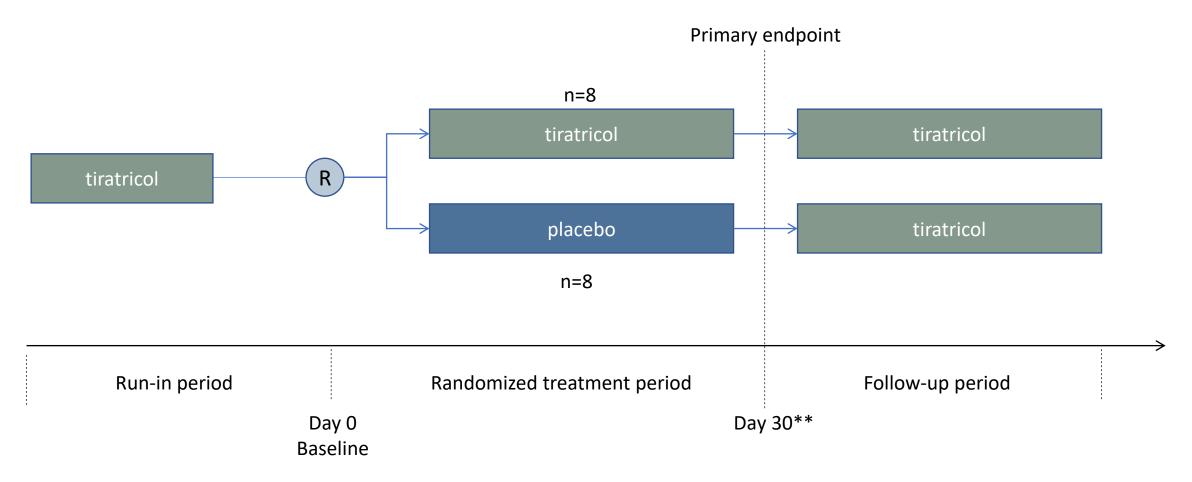
# Egetis concludes that demonstrating treatment effects on T3 levels in MCT8-deficiency could provide a basis for marketing approval for Emcitate® in the US

- Emcitate® (tiratricol) is the first potential treatment of MCT8 deficiency, a rare genetic disease with high unmet medical need and no available treatment
- In recent positive regulatory interactions, FDA acknowledges that a treatment effect on T3 levels and the manifestations of chronic thyrotoxicosis in MCT8-deficiency could provide a basis for marketing approval also in the US.
- An NDA in the US is targeted to be submitted in mid-2023 under the Fast Track Designation.
- A small, 30-day, placebo-controlled study in 16 treated patients, to be identified through the existing
  named patient program, will be conducted to verify the results on T3 levels seen in previous clinical
  trials and publications in a randomized controlled setting
- This is a major step towards a marketing application and increases the likelihood of success for Emcitate and the probability for Egetis to receive a US Rare Pediatric Disease Priority Review Voucher (PRV).
- Egetis will host a webcast today at 15:00 CET (9:00am ET)

Stockholm, Sweden, January 18, 2022 - Egetis Therapeutics AB (publ) (Nasdaq Stockholm: EGTX) (the "Company") today announced that in recent regulatory interactions, the US Food and Drug Administration (FDA) acknowledges that demonstrating a treatment effect on thyroid hormone T3 levels and the manifestations of chronic thyrotoxicosis could provide a basis for marketing approval also in the US. Consequently, the Company now has an aligned regulatory strategy for EU and US. The Company intends to submit a New Drug Application (NDA) in the US for Emcitate® (tiratricol) for the treatment of monocarboxylate transporter 8 (MCT8) deficiency in mid-2023 under the Fast Track Designation granted by the FDA in October 2021. This follows the announcement in December 2021 of intention to submit the Marketing Authorisation Application (MAA) for Emcitate to the European Medicines Agency (EMA) based on existing clinical data on the manifestations of chronic thyrotoxicosis in MCT8 deficiency.

## **Controlled Study - design**

Primary endpoint: Serum T3 levels, measured as the proportion of patients meeting T3 ≥ULN\* within the randomized treatment period



<sup>\*</sup> ULN: Upper Limit of Normal

<sup>\*\*</sup> Randomized treatment period end after 30 days or when rescue criterion (T3 ≥ULN) is met, whichever comes first Egetis Therapeutics | 2022-

# **Emcitate regulatory pathway to submissions in EU and US**



The first potential treatment for MCT8 deficiency, a rare genetic disease with high unmet medical need and no available treatment

Included in MAA in EU

Included in NDA in US

### Triac Trial I

- Completed 2018 (Groeneweg, 2019)
- Open-label, international, multicentre study
- N= 46

### **EMC** cohort study

- New data 2021 (van Geest, 2021)
- N= 27 from Triac Trial I & N=40 new pts from compassionate use

### **Natural history**

- Retrospective data, 2003 to 2019 (Groeneweg, 2020)
- N= 151

### **Controlled study**

- To be started in 2022
- N= 16
- Pts from named patient/ compassionate use program

To be added post approval when data available

### **Triac Trial II**

- Open-label, international, multi-centre study
- Pts ≤ 30 months of age
- Focus on neurocognition
- N= 15-20 planned
- Full 96 weeks data, expected in Q1 2024

Data already available

# Triac Trial II fully recruited: to establish effects of early intervention on neurocognitive development



Market approval not dependent on Triac Trial II data

# Primary objective

Confirm findings from Triac Trial I in youngest age group

 Improvement in neurocognitive development as measured by GMFM<sup>1</sup> and BSID-III<sup>2</sup> compared to natural history controls

# Secondary objective

Achievement of motor milestones (e.g. hold head, sit independently)

• Normalization of thyroid hormone function tests and markers of thyrotoxicosis

### **Description**

Open label, multi-centre trial in very young children with MCT8 deficiency

International trial with 10 centres in CZ, DE, IT, UK, FR, NL, US

Design discussed and anchored with EMA and FDA

ClinicalTrials.gov identifier: NCT02396459



15-20 children, 0-30 months of age



- First Patient First Visit in Dec 2020, recruitment target met in April 2022
- Results from 96 week read out expected in Q1 2024 and data is expected to be submitted postapproval to regulatory authorities shortly thereafter and available for HTA interactions
- Market approval not dependent on Triac Trial II data



# **Emcitate® clinical development timeline**



- Acquisition of all rights and assets relating to Teatrois in France
- Triac Trial I completed
- EMA Protocol Assistance EU

- CTA approvals in EU
- IND submission
- IND open (SMP)
- US Rare Pediatric Disease Designation (RPD)
- FPFV<sup>2)</sup> pivotal Phase IIb/III early intervention trial

- US & EU ODD RTH-B
- Completed recruitment in Triac Trial II
- Conduct 16 pts randomized 30 day study for US NDA
- Filing EU MAA H1 '23 & US NDA mid '23 under Fast **Track Designation**
- Launch preparations

2022/23 2017 2018 2019 2020 2021 2024

- Licensing of IP, data and knowhow within thyroid hormone signaling disorders from Erasmus Medical Centre<sup>1)</sup>
- **EU** Orphan Drug Designation (ODD)

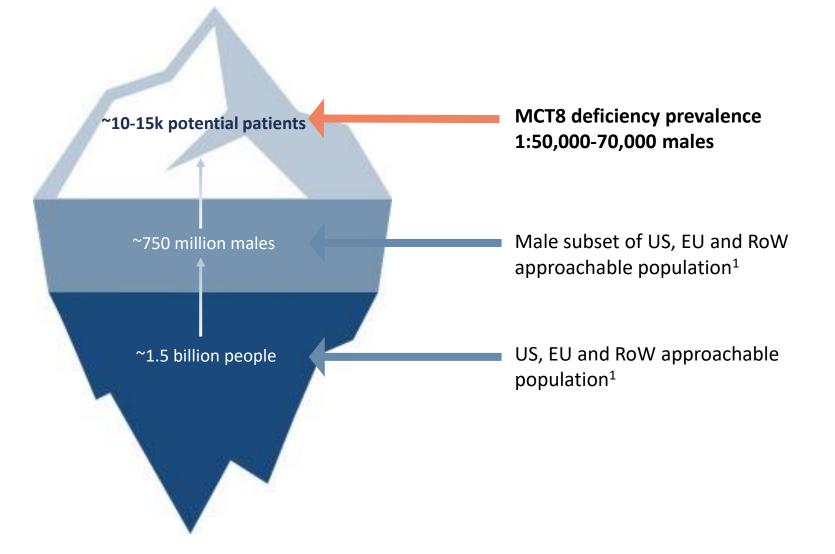
- US Orphan Drug **Designation MCT8**
- Publication Triac Trial I
- FDA pre-IND meeting
- CTA submissions EU

- Fast Track Designation US
- Data published confirms long-term efficacy and safety of Emcitate®

- EU and US approvals, pricing and launch
- US Priority Review Voucher
- Publication of complete Triac Trial II data (96 weeks)

# Estimating 10-15k addressable patients globally

No approved treatment for MCT8 deficiency



### MCT8 deficiency epidemiology

- At least one new-born diagnosed per 140,000 live births in the Netherlands in past years, corresponding to 1:70,000 males
- Actual number of patients could be higher:
  - Screening study suggests prevalence of 1:50,000 males<sup>2</sup>
  - Once treatment is available, more patients tend to be diagnosed

# Emcitate® – alleviating patient and societal burden

Aiming to provide value for both patients and society



**Patients** 

- Median life-expectancy of MCT8 patients is 35 years<sup>1</sup>
- Patients underweight for age or without ability to hold head have an even increased risk of premature death.

Society

- All MCT8 patients have significant neurocognitive disability from early childhood and typically require constant, life-long supportive care
- A recent study in a condition with similar severity (SMA) estimated total healthcare cost (excluding treatment cost) to USD 138k per patient and year<sup>2</sup>



**Emcitate** holds potential to become the **first approved therapy** to address the root cause of MCT8 deficiency, restore thyroid hormone signaling and thereby **prevent disease progression**, alleviate symptoms and **prolong lives** 



# Supporting diagnosis through disease awareness initiatives

MCT8 deficiency awareness and educational activities launched through various channels

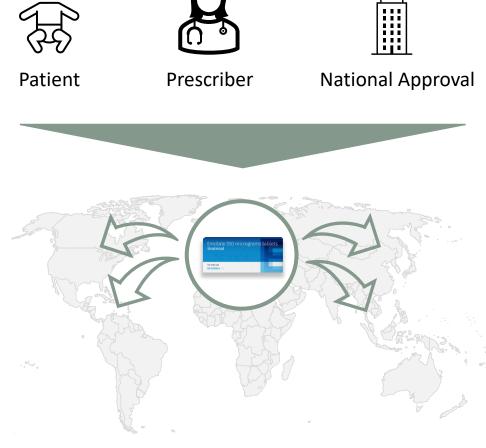
- Disease awareness and educational efforts aim to
  - increase number of physicians who understand how to diagnose and manage MCT8 deficiency
  - speed up diagnosis
- Collaborating with patient advocacy groups and KOLs
- Exhibit at scientific/medical conferences 2022:
  - European Paediatric Neurology Society, April, Glasgow
  - European Thyroid Association, Sept, Brussels
  - European Society of Pediatric Endocrinology, Sept, Rome
  - American Thyroid Association, Annual Meeting, Oct, Montreal
- Several channels for efficient reach.
  - mct8deficiency.com
  - Mailings
  - Social media
  - Publications



# **Emcitate supplied globally on a named patient basis**

The named patient use (NPU) confirms the significant unmet medical need in MCT8 deficiency and the view of Emcitate's potential to address it

- NPU and compassionate use programs
  - mechanisms to allow early access to a medicine prior to regulatory marketing approval
  - granted to pharmaceuticals under development for situations with high unmet medical needs and where no available treatment alternatives exist or are suitable
- Emcitate is being supplied on a named patient basis, following individual approval from the national medicines agencies, to
  - more than 150 patients
  - in over 25 countries



**MEDICINES AGENCY** 

## Analogue orphan drugs priced at premium

Rapid market penetration with considerable sales already 3rd year in market

- Payers in general accept higher prices for orphan drugs compared to traditional drugs and especially if they;
  - Address an ultra rare disease, e.g. prevalence less than 1:50,000 people
  - Target a severe disease, i.e. life threatening/debilitating
  - Provide **health gain**, rather than just condition stabilization

 Emcitate fulfills these criteria, no other drugs available or being developed for MCT8 deficiency

### **Analogue orphan drugs**

	<b>Vimizim®</b> Recombinant enzyme	<b>Kalydeco®</b> Small molecule	<b>Spinraza®</b> Antisense oligonucleotide	<b>Brineura®</b> Recombinant enzyme
Disease	MPS IVA	CF with specific mutations	SMA	CLN2
Rarity - less than 1:50,000 people	✓	✓	✓	✓
Severity – life threatening/debilitating	✓	✓	✓	✓
Health gain	✓	✓	✓	✓
Global annual treatment cost	> \$400k	> \$250k	> \$350k	> \$600k
Year of 1st approval	2011	2012	2016	2017
Global sales 3rd year in market	\$354mn	\$464mn	\$1.7bn	\$110m
Global sales 2020	\$544mn	\$803mn	\$2.1bn	\$110m



# FDA granted Rare Pediatric Disease designation to Emcitate®

US Rare Pediatric Disease Priority Review Voucher (PRV) provides a ~\$100m opportunity

### **Overview PRV**

- The FDA grants Rare Pediatric Disease designation (RPD) to therapies for serious or life-threatening diseases affecting fewer than 200,000 people in the USA.
- Sponsors holding a RPD can apply to receive a US Rare Pediatric Disease Priority Review Voucher (PRV) up on approval.
- PRV program recently prolonged until FY 2026
- Provides accelerated FDA review of a new drug application for another drug candidate, in any indication, shortening time to market in the US.
- The voucher may be sold or transferred to another sponsor.
- By end 2019 22 PRVs for rare pediatric diseases had been awarded by FDA, 12 were sold with individual voucher sale prices ranging from \$67m to \$350m.

### **Examples of PRVs sold**

Seller	Buyer	Value	Year
Bavarian Nordics	Undisclosed	\$95M	2019
SOBI	AstraZeneca	\$95M	2019
Bayer Healthcare	argenx	\$100M	2020
Lumos Pharma	Merck	\$100M	2020
Sarepta Therapeutics	Gilead	\$125M	2020
Albireo	Undisclosed	\$105M	2021
BioMarin	Undisclosed	\$110M	2022

Aladote® - clinical development program

resentation | Egetis Therapeutics | 2022-07-12

# Paracetamol/acetaminophen poisoning

no adequate treatment for increased-risk patients

What is paracetamol/ acetaminophen poisoning?

Minimum toxic dose of paracetamol/acetaminophen in adults is only 7.5g

- Risk factors include malnutrition, alcoholism and consumption of other medications
- Paracetamol/acetaminophen poisoning can lead to acute liver failure, liver transplant or death

How many does it affect?

• 19 billion units of paracetamol /acetaminophen packages are sold in the US alone every year

- >175,000 patients hospitalised globally per annum driven by 89,000 cases/year of paracetamol overdose in the US and 105,000 cases/year in the UK (~ 50% hospitalised)
- ~50% of paracetamol overdose cases are unintentional

Why is current treatment inadequate?

- Efficacy of current NAC (N-acetylcysteine) treatment decreases with time
- Approximately 25% of patients are late arrivals to hospitals (>8h) –
  late arrivals are at increased risk
- There is no effective treatment option for patients at increased risk

A new standard of care is needed

 Aladote® aims to become a new standard of care for patients with increased risk for liver injury in combination with NAC



## Orphan drug candidate

with clear scientific and mechanistic rationale

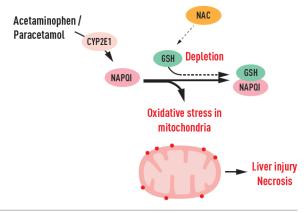
### Early presenters (<8h) NAC treatment effective against liver injury

 Liver glutathione (GSH) replenished by NAC, toxic NAPQI metabolite excreted as GSH conjugate

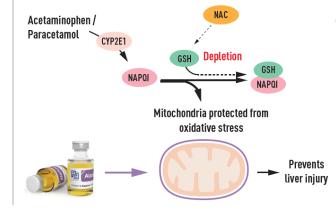


Late presenters (>8h) are at increased-risk for liver injury NAC treatment + Aladote® to prevent liver injury

**Under NAC treatment alone** liver GSH stores depleted by the toxic NAPQI metabolite -> oxidative stress, mitochondrial dysfunction and liver injury (necrosis)



 In most cases NAC effectively prevents liver injury i.e. limited need for Aladote®



Reactive nitrogen species (RNS), Reactive Oxygene Species (ROS)

 Aladote<sup>®</sup> (calmangafodipir) prevents ROS and RNS formation, restores mitochondrial energy production and prevents liver injury

### Overview of completed Phase Ib/IIa



 Met primary endpoint of safety tolerability in the combination of Aladote® and NAC

- Results presented at the 58th Annual Meeting of the Society of Toxicology, EASL ILC in April, Vienna and published in Lancet's journal EBioMedicine in 2019
- Presented at, American College of Medical Toxicology (ACMT) and Society of Toxicology (SOT), as novel emerging treatments for acetaminophen/ paracetamol toxicity in 2021

Secondary objectives and results

 Measurements of Alanine transaminase (ALT), international normalised ratio (INR), keratin-18, caspase-cleaved keratin-18 (ccK18) and microRNA-122 (mir122) and glutamate dehydrogenase (GLDH) indicates that Aladote® reduce liver injury

Description

- An open label, rising-dose, randomized study exploring safety and tolerability of Aladote<sup>®</sup> co-treatment with NAC
- ClinicalTrials.gov identifier: NCT03177395

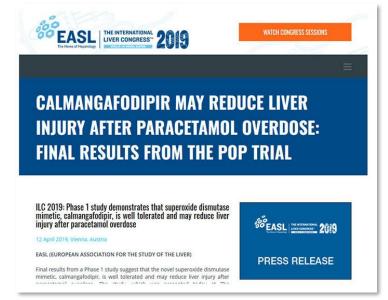
# of patients

 Single ascending dose study in 3 dosing cohorts of 8 patients (N=24) as add-on to NAC regime

**Timetable** 

- Initiated in June 2017 (first patient in)
- Completed in September 2018





### Positive proof-of-principle Phase Ib/IIa results

Indicates that Aladote may reduce liver injury



#### Safety & tolerability

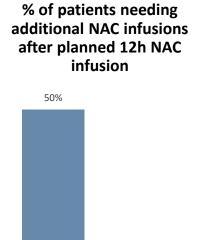
Event	NAC alone	NAC + 2 μmol/kg Aladote	NAC + 5 μmol/kg Aladote	NAC + 10 μmol/kg Aladote
Any AE	6 (100%)	6 (100%)	6 (100%)	6 (100%)
Any SAE	2 (33%)	4 (67%)	2 (33%)	3 (50%)
SAE Starting within 7 days	1 (17%)	1 (17%)	1 (17%)	2 (33%)

- Met primary endpoint of safety tolerability in the combination of Aladote® and NAC
- No AE or SAE probably or definitely related to Aladote®

### Liver injury – ALT<sup>1</sup> pre-defined secondary outcome

Event	NAC alone	NAC + 2 μmol/kg Aladote	NAC + 5 μmol/kg Aladote	NAC + 10 μmol/kg Aladote
50% ALT increase	2 (33%)	0 (0%)	0 (0%)	1 (17%)
100% ALT increase	1 (17%)	0 (0%)	0 (0%)	1 (17%)
ALT >100 U/L at 10 hours	2 (33%)	0 (0%)	0 (0%)	0 (0%)
ALT >100 U/L at 20 hours	2 (33%)	0 (0%)	0 (0%)	0 (0%)

• ALT >100 U/L is the indication to stay in hospital

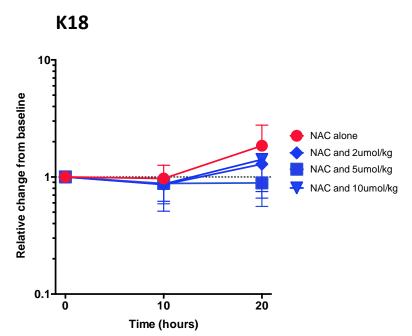


(n=2 out of 18

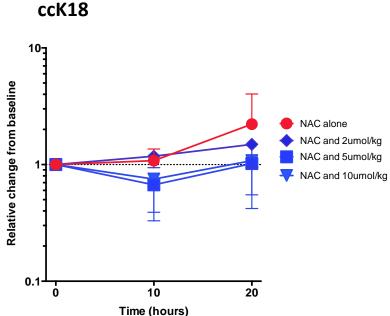
11%

# Aladote® demonstrates consistent results of reduced liver injury as measured by exploratory biomarkers

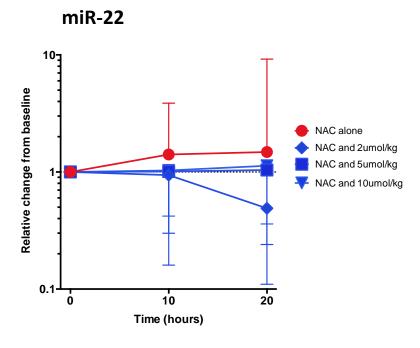




K18 is a measure of cell death and correlate with peak ALT activity during the hospital stay



ccK18, is a measure of cell death and correlate with peak ALT activity during the hospital stay



miR-122 is a liver specific early marker (micro-RNA) for acute liver injury which predicts a rise in ALT activity following paracetamol overdose

### Pivotal Phase IIb/III study for US/EU regulatory submission



Patient population

Patients who have overdosed on paracetamol with increased risk of liver damage due to late arrival at hospital (> 8h) who need treatment with NAC

**NAC** regimen

Approved 21 hours NAC regimen

Treatment groups

 4 groups in combination with NAC: Aladote® high dose; Aladote® middle dose; Aladote® low dose; Placebo

Initiation of active treatment

• IV (bolus) as soon as possible after randomization and after starting NAC treatment (but no later than 4 hours after starting NAC treatment)

**Interim analysis** 

 Interim analysis after 35 patients per treatment group, which includes a futility analysis, dose selection and analysis of continued study size (number of patients)

**Study size** 

250 patients planned

Efficacy endpoints

- Primary: Combination of ALT and INR
- Number (%) of patients who need extended NAC treatment after 21 hours
- Length of hospital stay
- Explorative biomarkers: K18, miR-122 and GLDH



**Study countries** 

EU, UK and USA

### Aladote® clinical development timeline



- **US ODD granted**
- Results presented at Society of Toxicology, **EASL ILC and Lancet EBiomedicine**
- Regulatory interactions with FDA and EMA

- Orphan Drug **Designation EU**
- Initiate pivotal Phase IIb/III study
- Interim analysis after 50% of patients included
- Recruitment completed

2024/25 2020/21 2022/23 2032<sup>1</sup> 2018 2019 Design of pivotal study Phase Ib/IIa study fully Regulatory submissions finalized Europe/US recruited Initial Phase Ib/IIa results Regulatory interactions Europe/US approval FDA, EMA & MHRA and launch Established Scientific **Advisory Board** Regulatory submissions **ROW** Full Phase Ib/IIa results Submission of ODD

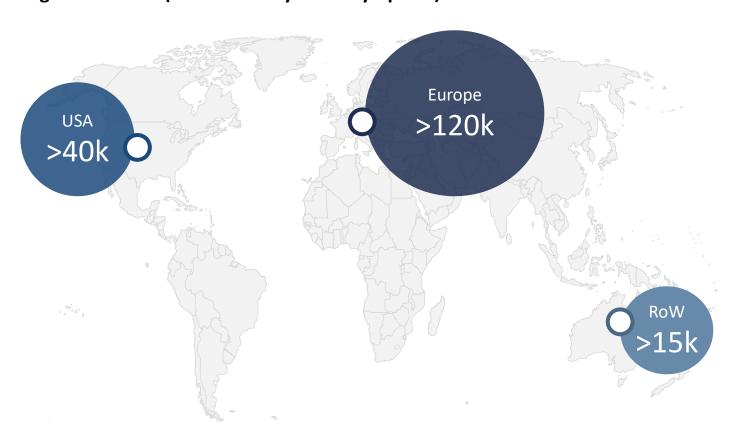
Aladote® - Commercial opportunity

resentation | Egetis Therapeutics | 2022-07-12

### Estimating at least 175k addressable patients globally



Annual number of POD (paracetamol/acetaminophen overdose) cases hospitalized and receiving i.v. antidote (NAC currently the only option)



#### **POD** epidemiology

89,000 cases/year of paracetamol overdose in the US and 105,000 cases/year in the UK

- ~50% hospitalized and receive i.v. antidote treatment
- ~25% are late arrivals

Global paracetamol/acetaminophen exposure varies, leading to POD incidence being different between countries

### Aladote® – alleviating patient and societal burden

Aiming to provide value for both patients and society

#### POD is a life threatening condition with remaining medical needs

#### **Patients**

- POD (paracetamol/acetaminophen overdose) can lead to acute liver failure, liver transplant or death
- In US and UK together, yearly > 500 deaths due to POD and more people registered for liver transplantation

#### Society

- In the US the annual cost in 2010 was estimated at > \$1bn to treat patients with POD1
- The POD Emergency Department and inpatient cost is approximately USD 13-40k<sup>1</sup>
- The average POD inpatient length of stay is 3.1 days with a variance of +4.4 days for the most severe cases<sup>1</sup>
- US liver transplant costs USD 125-473k<sup>1</sup>



With **Aladote**, the ambition is to **reduce hepatic injury** of POD and thereby contribute to **fewer hospitalization days**, **prevent need** for liver transplantation and **increase survival** 

### Analogue antidotes priced at \$ 3.5k - 50k



National emergency hospital stocking guidelines - opportunity for rapid market penetration

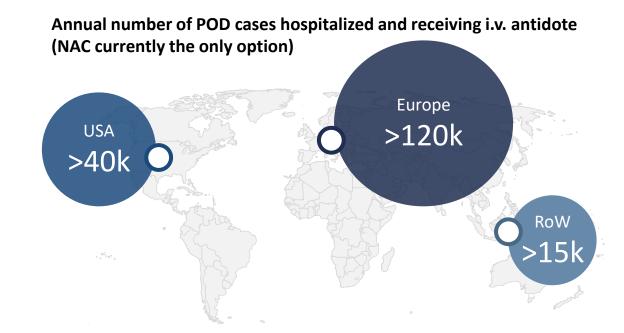
- Various antidotes, e.g. vs. drug overdosing, metal poisoning, snake bites and reversal of anticoagulant treatment effects
- Limit morbidity/mortality when used within appropriate time
- National recommendations for stocking of antidotes at hospitals providing emergency care
  - For getting payer/formulary committee acceptance to be stocked, antidotes are in general priced lower than traditional orphan drugs, despite
     often having orphan status
  - Getting included provides great opportunity for rapid market penetration
    - Praxbind stocked in 3,200 US hospitals < 3 years from launch</li>
    - Andexxa sales \$112mn in US alone second year on market
- Analogue antidotes for comparable settings as Aladote have global average costs of \$ 3.5k 50k per treatment

	Naloxone hydrocloride	Praxbind	Andexxa/Ondexxya	Aladote (target profile)
Year of first approval	1971	2015	2018	NA
Poisoning indication	Opioid toxicity	Reversal of anticoagulant effects of the NOAC dabigatran	Reversal of anticoagulant effects of the factor Xa inhibitors apixaban & rivaroxaban	Paracetamol/ acetaminophen toxicity
Cost per treatment	Low since generic	\$ 3.5k – 4.5k	\$ 25k – 50k	TBD

### Aladote® commercial opportunity

- Addressing unmet needs in antidote market create substantial opportunity

- POD is a life-threatening condition with remaining medical needs
- No effective treatments for high-risk patients, e.g. patients arriving > 8h after ingestion
- No other companies developing drugs for POD
- Opportunity for rapid sales uptake due to national emergency hospital stocking guidelines
- Anologue antidotes priced at \$3.5k 50k





### >\$350mn annual sales opportunity assuming:

- Global average annual treatment cost per patient: \$5k
- Addressable patients: >175,000
- Market penetration: 40%

The orphan drug segment and path to market resentation | Egetis Therapeutics | 2022-07-12

### Orphan drug segment – a highly attractive opportunity



Orphan drug designation is awarded to products targeting limited disease populations<sup>1</sup>

More than 7,000 known rare diseases

Approx. 10% of the general population may be affected by a rare disease

Substantial unmet medical need for patients, only 5% of rare diseases have an approved therapy

• Less extensive clinical trials
• Agile and faster development process
• Lower costs
• Lower development risk

• Free regulatory advice
• Reduced fees
• Expedited review
• Market exclusivity

• No or few competitors
• Highly focused target groups

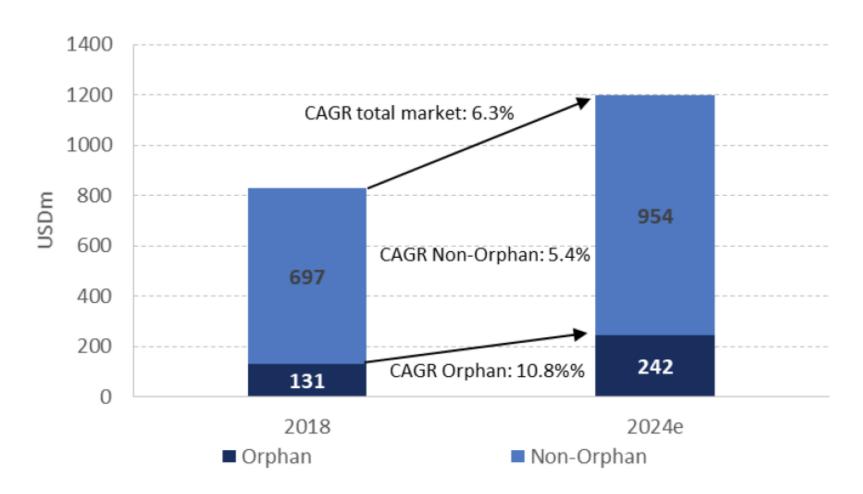
Premium pricing

Well-defined patient populations with substantial unmet medical need

### CAGR estimates of total pharmaceutical market vs orphan



The global orphan or rare disease market size was valued at an estimated USD 140 - 150 bn and is expected to grow at 10-14% CAGR over the coming five years.



### Commercialisation of *Emcitate & Aladote*

### Commercial infrastructure build up initiated

#### Strong success factors...

- High unmet medical need without competing compounds
- Centralized, focused target groups of specialists
- Top-down scientific sales approach
- Leading KOL support
- Treatment algorithms highly protocol driven

...for sustainable, profitable & lean commercialisation

- Building inhouse commercial capabilities for launch of Emcitate® and Aladote® in EU and US
- Small and focused footprint with an estimated < 50 FTEs considered sufficient for both assets
- Retain larger share of product revenues over time within Company
- Commercialisation in other territories through partners

**5.**Summary

Corporate resentation | Egetis Therapeutics | 2022-07-12 51

### Two highly promising orphan drug candidates

Emcitate® – Therapy for genetic disturbance in thyroid hormone signaling with life-long severe disability

- Lead candidate for addressing MCT8 deficiency which affects ~1:70,000 males, a condition with high unmet medical need and no available treatment. No competing sponsored products in clinical development
- Obtained Orphan drug designation in the EU and US 2017 and 2019, respectively. US Rare Pediatric Disease Designation received in Nov 2020, eligible for Priority Review Voucher. Fast track designation granted by FDA in Oct 2021
- Triac Trial I (Phase IIb) completed with significant and clinically relevant effects on T3 levels and the manifestations of chronic thyrotoxicosis
- Real-world data published in Oct 2021 confirms long-term efficacy and safety of Emcitate<sup>®</sup> in MCT8 deficiency patients
- Intend to submit MAA to EMA based on existing clinical data in H1 2023
- Intend to NDA submission in mid 2023 based on treatment effect on T3 levels and the manifestations of chronic thyrotoxicosis in MCT8deficiency. A placebo-controlled study in 16 treated patients will be conducted to verify the results on T3
- Triac Trial II fully recruited; to establish the effects of early intervention on neurocognitive development, previously seen in the Triac Trial I.
   Results are expected in Q1 2024
- More than 150 patients are being treated with Emcitate on a named patient basis, following individual regulatory approvals from the national regulatory agencies

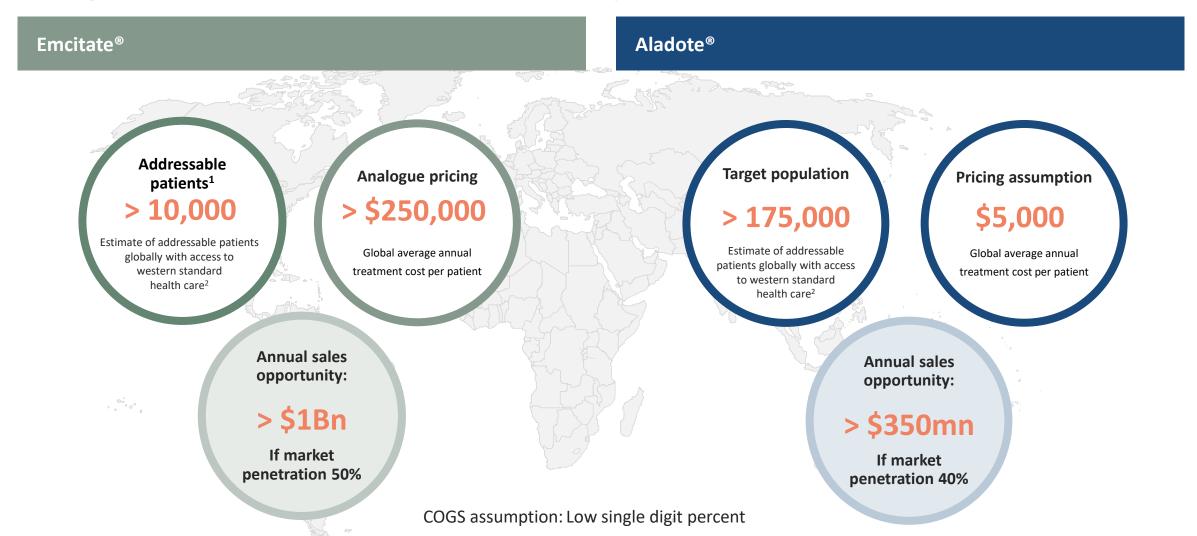
## Aladote® – Prevents acute liver injury caused by paracetamol/acetaminophen poisoning

- Paracetamol poisoning is one of the most common overdoses with >175,000 hospital admissions globally per annum
- No adequate treatment exists for increased risk patients
- Orphan drug designation (ODD) granted in 2019 in the US
- Ongoing dialogue with EMA on the appropriate scope of the indication for an ODD in the EU
- Successful results from Phase Ib/IIa study in paracetamol overdosed patients
- Pivotal Phase IIb/III study planned for marketing authorization application in both US and EU, targeting study start in 2022
- No competing products in clinical development

### Late-stage orphan drug pipeline, \$1Bn+ annual sales opportunity



Analogue benchmarks indicate substantial market potential



### **Upcoming pipeline milestones**





**Emcitate**®

- ✓ Fast Track Designation US
- Data published confirms long-term efficacy and safety of Emcitate
- ✓ US & EU ODD RTH-β
- ✓ Recruitment target met in Triac Trial II, Apr 2022
- Conduct 16 pts randomized 30 day study for US NDA
- Filing EU MAA H1 '23 & US NDA mid '23 under Fast Track Designation
- EU approval and launch
- US approval and launch
- US Rare Pediatric Disease Priority Review Voucher
- Results Triac Trial II

2021

2022/23

2024/25

Aladote®

- Orphan Drug designation EU
- Initiate pivotal Phase IIb/III study\*

- Interim analysis
- LPFV pivotal Phase IIb/III study
- Filing EU/US
- EU/US approval and launch



### An integrated orphan drug company, focusing on late-stage development for commercialization

- Dedicated orphan drug development company with two late-stage orphan drug assets: Emcitate® and Aladote®
- Target MAA/NDA submissions for Emcitate in 2023 and for Aladote in 2024/2025
- Highly attractive **orphan drug segment** with potential >\$1Bn annual sales opportunity
- Plan to **launch** through niche inhouse commercial organization in the EU and US
- Combined core expertise in late-stage orphan clinical development, registration and commercialization with experience from:



Listed on NASDAQ Stockholm (EGTX) HQ in Stockholm, Sweden





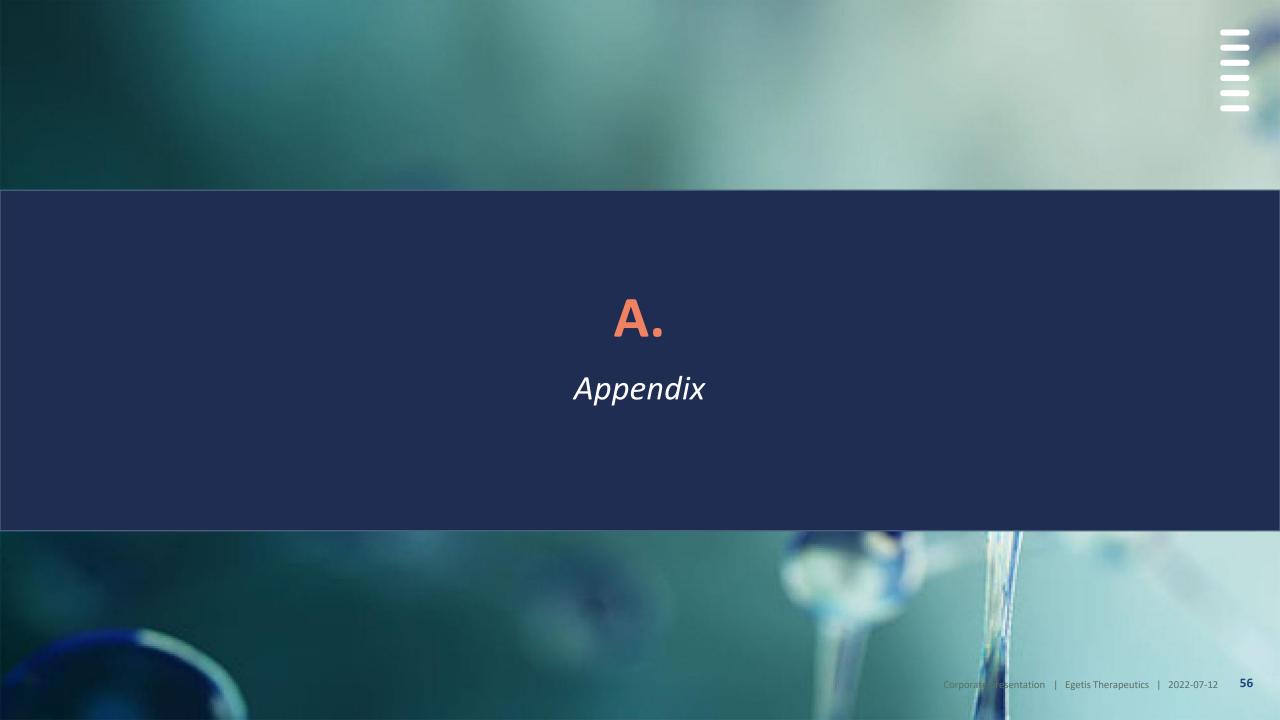












### **Leadership team**



#### Nicklas Westerholm

CEO

- Took office in June 2017 and has previously worked in the AstraZeneca Group since 1995 in several global roles in various business areas, most recently as VP Project & Portfolio Management. Prior Nicklas has held positions such as Executive Officer & VP Japan Operations, Director Investor Relations, Head of Global API Supply and Head of Development Manufacture. He has studied Analytical and Organic Chemistry at Stockholm University and Chemical Engineering at KTH, as well as studies at University of Warwick, INSEAD and Harvard Business School.
- Ownership: 109,873 shares and 2,400,000 employee stock options



#### Yilmaz Mahshid

CFO

- Yilmaz has experience from different senior positions in the life science sector, including Investment Manager & Controller at Industrifonden, and CFO at PledPharma between 2017 and 2020, as well as healthcare analyst at Pareto Securities and Öhman Fondkommission. Prior to joining Egetis Therapeutics, Yilmaz was CEO of the listed biotech company Medivir. Yilmaz also has a solid academic background with a PhD from the Department of Medical Biochemistry and Biophysics at Karolinska Institutet, Stockholm.
- Ownership: 191,000 shares and 1,150,000 employee stock options



#### **Henrik Krook**

#### **VP Commercial Operations**

- Appointed VP Commercial Operations in December 2020. He has a broad experience from over 15 years in commercial leadership settings, including both big pharma and biotechs. He has previously held senior corporate and commercial advisory roles for biotech companies such as Affibody and senior managerial positions at e g Alexion, Novartis and Roche. Henrik has a PhD in immunology from Uppsala University and an Executive MBA from Stockholm School of Economics.
- Ownership: 170,000 shares and 1,150,000 employee stock options



#### Kristina Sjöblom Nygren

CMO

- Took office in May 2020 and has previously worked as CMO and Head of Development at Santhera, where she oversaw activities in late-stage clinical development, registration, post-approval commitments and managed access-programs within rare diseases in different therapeutic areas. Previously, Kristina spent 18 years at SOBI, Wyeth and AstraZeneca, where she held a number of senior positions. She has been involved in many different interactions with regulatory bodies such as the US FDA and the EMA including scientific advice and orphan drug applications. Before joining the industry, she worked as a licensed physician in several clinical positions. She holds a Diploma in Pharmaceutical Medicine, and an MD from the Karolinska Institute, Stockholm.
- Ownership: 6,000 shares and 650,000 employee stock options



#### **Christian Sonesson**

#### VP Product Strategy & Development

- Appointed VP Product Strategy & Development in August 2017 following 13 years at Astra Zeneca. He has broad experience within drug development, including successfully leading products during Phase 3 (FORXIGA® in type 1 diabetes) and of regulatory submissions and defense, bringing new drug candidates to market in different regions (e.g. FORXIGA® in type 2 diabetes, MOVANTIK®, ONGLYZA®-SAVOR, BRILINTA®-PEGASUS and QTERN®). PhD in Biostatistics from Gothenburg University and an Executive MBA from Stockholm School of Economics.
- Ownership: 12,000 shares and 1,150,000 employee stock options



#### Karl Hård

#### VP, Head of Investor Relations & Communications

- Appointed in February 2022. He has 25 years experience within the pharma and biotech sector, incl. 10 years in R&D and 9 years in Investor Relations at AstraZeneca, latterly as VP Investor Relations. Previously, Head of IR and Communications at Kiadis Pharma (The Netherlands) and Redx Pharma (UK). PhD in Bio-organic Chemistry from Utrecht University. Former Assistant Professor of Chemistry at Leiden University.
- Ownership: 0 shares and 100,000 employee stock options

### **Board of directors**





**Thomas Lönngren** *Chairman of the board* 

- Board member since: 2021
- MSc in social and regulatory pharmacy and a degree in Pharmacy, University of Uppsala.
- Other assignments: Board member at Compass Pathsways PLC and NDA group. Director at own company PharmaExec Consulting AB. Advisor to NDA group, Artis Venture, Baren Therapeutics, Centre for Innovation in Regulatory Science (CIRS) and ScientificMed AB. Faculty member of GLG Institute
- Ownership: 165,219 shares



Mats Blom
Board member

- Board member since: 2021
- BA, Business Administration and Economics, University of Lund and MBA, IESE University of Navarra.
- Other assignments: CFO NorthSea Therapeutics and Board member of Hansa Biopharma and Auris Medical
- Ownership: 2,257,512 shares



Peder Walberg
Board member

- Founder and CEO of Rare Thyroid Therapeutics
- MD and BSc in international economy and business administration, Uppsala University
- · Other assignments: Board Member of Immedica Pharma AB,
- Previous assignments: Founder and CEO, Medical Need, Head of Business Development and Strategy, Swedish Orphan International and SOBI. BoD of Wilson Therapeutics and identified Decuprate for treatment of Wilson disease
- Ownership: 31,858,414 shares (through Cetoros AB)



Gunilla Osswald

Board member

- Board member since: 2017
- · Ph.D. in biopharmacy and pharmacokinetics
- Other assignments: CEO BioArctic AB
- Ownership: -



Elisabeth Svanberg
Board member

- Board member since: 2017
- MD, Ph.D., Assoc Professor in surgery
- Other assignments: Chief Development Officer Ixaltis SA. Board member Swedish Orphan Biovitrum (SOBI), Amolyt Pharma and Galapagos
- Ownership: -

### **Share Register and Market Cap**



#### **Shareholders**

Source: Monitor by Modular Finance. Compiled and processed data from various sources, including Euroclear, Morningstar and the Swedish Financial Supervisory Authority (Finansinspektionen). The verification date may vary for certain shareholders.

#### 10 largest shareholders

Name	Capital	Votes	Num. of shares	Verified
Peder Walberg	19.30%	19.30%	31 858 414	2021-12-31
Peter Lindell	10.37%	10.37%	17 124 820	2021-12-31
Avla Holding AB	10.04%	10.04%	16 572 442	2021-12-31
Fjärde AP-fonden	8.67%	8.67%	14 311 300	2021-12-31
RegulaPharm AB	5.97%	5.97%	9 846 730	2021-12-31
Avanza Pension	2.67%	2.67%	4 406 802	2021-12-31
Thomas Eldered	1.79%	1.79%	2 953 462	2021-09-30
Carl Rosvall	1.64%	1.64%	2 707 914	2021-12-31
Mats Blom	1.37%	1.37%	2 257 512	2021-09-30
Unionen	1.28%	1.28%	2 120 165	2021-12-31
Total 10	63.10%	63.10%	104 159 561	
Total number of owners	6,895			2021-12-31
Total number of shares	165,068,560			2021-12-31

- Cash position: SEK 107M (~EUR 11M)\*
- Number of outstanding shares: 165M
- MCap: ~SEK 800M\*\*
- Listing venue: Nasdaq Stockholm Main Market

# Egetis Therapeutics resolves on a fully guaranteed preferential rights issue of approximately SEK 180 million



Announced March 21, 2022

- To finance the preparations for regulatory submissions for market approval in EU and US, initiate the establishment of the Company's commercial infrastructure in Europe and US for Emcitate® and pre-launch activities, as well as general corporate purposes, in addition to providing financial flexibility
- Existing shareholders, including Cetoros AB (Peder W), Cidro Förvaltning AB (Peter L), Avla Holding AB (Kennet R), Fjärde APfonden, RegulaPharm AB (Gudrun H), Flerie Invest AB (Thomas E) and Unionen, as well as members of management and the Board of Directors, have undertaken to subscribe for shares representing approximately 39.3 percent of the Rights Issue
- The Company further strengthens its specialist investor base. New investors, including Linc AB, as well as existing shareholder Flerie Invest AB and members of management and the Board of Directors, have undertaken to subscribe for shares representing approximately 27.8 percent of the Rights Issue through assuming subscription rights of select existing shareholders free of cost
- In total, subscription undertakings represent approximately 67.2 percent of the Rights Issue, corresponding to approximately SEK 121.4 million.
- A consortium of existing shareholders, including Fjärde AP-fonden and members of management, as well as the new investor Linc AB, have undertaken to guarantee approximately 32.8 percent of the Rights Issue, corresponding to approximately SEK 59.4 million. Consequently, the Rights Issue is secured in its entirety.
- Approved at an extraordinary general meeting, on April 13, 2022



**Acquisition of Rare Thyroid Therapeutics on 5 November 2020** 

### The combination will drive synergies

PledPharma and Rare Thyroid Therapeutics merged to launch a new company



#### **PledPharma**

- Team with profound late-stage drug development experience and strong trackrecord
- Listing on Nasdag Stockholm provides access to public markets and capital as well as visibility
- Desired prospective partner in project collaborations. Previous major license agreement with Solasia
- Efficient internal organisation and strong corporate governance

#### Rare Thyroid Therapeutics

- Team with strong track-record of identifying and developing ODDs and creating shareholder value
- Strong network of external project advisors with specialist knowledge. Collaboration with Erasmus Medical Center in Rotterdam
- Founding team with experience from international launch and commercialisation of orphan drugs

#### **Synergistic orphan drug focus**

- 2020 accelerated PledPharma's strategic review
- Lead asset Aladote® facilitates the new pronounced strategic focus on orphan drug segment
- Emcitate® and RTT's capabilities fit well with the new strategy
- Build critical mass, generate synergies and improve operational effectiveness for projects in the orphan segment
- Size, vicinity and complementary capabilities allow for a fast and smooth integration



### The acquisition and rights issue

Institutional investor base broadened

#### **Acquisition**

- On 5 November 2020, PledPharma acquired all outstanding common shares in Rare Thyroid Therapeutics
- Consideration consisted of a combination of PledPharma common shares and cash
- An upfront cash payment of SEK 60m
- 63.8 million shares representing approx 39% of the total number of outstanding shares in PledPharma post rights issues
- Owners of Rare Thyroid Therapeutics will receive a royalty of 3% of net sales generated through Emcitate<sup>®1</sup>
- Owners of Rare Thyroid Therapeutics will also be granted 50% of the net proceeds from a potential sale of US Rare Pediatric Disease Priority Review Voucher related to Emcitate®

#### **Rights issue**

- Successfully raised SEK 250 million in oversubscribed rights issue (c. SEK 200m) and utilized overallotment option (c. SEK 50m)
  - Subscription price of SEK 5.25 per share corresponding to a 2.5 percent premium to close 2 October 2020
- Institutional investor base broadened
  - Overallotment Option, allocated to the Fourth Swedish National Pension Fund ("AP4"), NYIP (Nyenburgh Holding BV) and Nordic Cross
- The proceeds will be used to finance: (i) the development of Emcitate® and Aladote® to market approval in Europe and USA (60%); (ii) initial commercial preparations (20%); (iii) general corporate purposes and financial flexibility (20%)







### Thank you!

Egetis Therapeutics egetis.com