

Corporate Presentation

Forward-Looking Statements

Some of the information in this presentation relates to future events or future business and financial performance. Such statements constitute forward-looking information within the meaning of the Private Securities Litigation Act of 1995. Such statements can be only predictions and the actual events or results may differ from those discussed due to, among other things, the risks described in the public filings and other publications of Cyclo Therapeutics, Inc. Forward-looking statements are identified by words such as "anticipates", "projects", "expects", "plans", "intends", "believes", "estimates", "target", and other similar expressions that indicate trends and future events.

The market data and certain other statistical information used throughout this presentation are based on independent industry publications, governmental publications, reports by market research firms or other independent sources. Some data are also based on the Company's good faith estimates. In addition, this presentation includes summaries of scientific activities and outcomes that have been condensed to aid the reader in gaining general understanding.

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Factors that could cause the Company's results to differ materially from those expressed in forward looking statements include, without limitation, the Company's need for additional capital; the Company's reliance on its Trappsol[®] Cyclo[™] product, which may never receive regulatory approval; the Company's ability to commercialize any of its proposed drug products if it receives regulatory approval; the outcome of the Company's clinical trials, which may not support the Company's product claims or may result in adverse side effects; the cost and timing of the Company's clinical trials; the Company's reliance on third parties to conduct clinical trials and to produce its products; and other risks associated with being a clinical stage biotechnology company.

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Company Snapshot

<u>Who:</u> In 1990, the company was formed as Specialty Fine Chemical business specializing in cyclodextrins. In 2014, the business was expanded into a biotechnology company dedicated to developing life-changing medicines through science and innovation for patients and families living with challenging diseases.

<u>What:</u> Trappsol[®] Cyclo[™] is a proprietary formulation of hydroxypropyl beta cyclodextrin and in multiple clinical studies has shown encouraging results to effectively manage the transportation of cholesterol.

Why: Because cholesterol is so important to the normal function of our cells, its synthesis and degradation is tightly controlled by an array of cellular processes. When there is an imbalance in cholesterol synthesis or metabolism, cells and organs may not function properly, leading to disease or death.

How: Trappsol[®] Cyclo[™], with its cyclic structure, facilitates the transport of accumulated cholesterol out of cellular lysosomes so it can be further processed and excreted out of cells.

Currently Targeting 2 Serious Diseases with Unmet Medical Need

Niemann-Pick Disease Type C Fatal and progressive genetic disorder Orphan indication affecting >9,000 in 80 countries (~400 in U.S. / 320 EU5)¹

Alzheimer's Disease 6th leading cause of death affecting 5 million people in the U.S.²

Platform technology has potential to fuel pipeline expansion opportunities

2. https://www.alz.org/alzheimers-dementia/facts-figures



^{1.} April 2021, Tessellon Inc. (former Kantar Health experts with 25+ years of epidemiology and forecasting experience), (<u>www.Tessellon.com</u>); Exhaustive literature search with a broad range of MESH terms in United States + 79 other countries.

Platform Technology Pipeline:

Trappsol[®] Cyclo[™] allows for a multiple shots on goal model



Ongoing Collaboration with University of the Witwatersrand, Johannesburg to Advance Trappsol[®] Cyclo[™] Platform and Explore Pipeline Expansion Opportunities

Orphan Drug Designation in U.S. | Fast Track Status in U.S. | Potential for Priority Review Voucher (PRV) in U.S Orphan Designation in EU | EMA Pediatric Investigational Plan Adopted



Leadership Team with Proven Experience









Lise Lund Kjems, MD, PhD Chief Medical Officer Albireo aldeyra Intarcia Bylvay. (odevixibat) UNOVARTIS





ECE



Jeffrey L. Tate, Ph.D. Chief Operating Officer, Chief Quality Officer & Director 孤 natural biologics

UNIVERSITY OF MINNESOTA







Lori McKenna Gorski **Global Head of Patient Advocacy**

SANOFI GENZYME 🌍





Scientific Advisory Board



Rita Colwell, Ph.D.

Co-Chair

Internationally recognized scientist, microbiologist and founder of CosmosID, a privately held bioinformatics firm.

Distinguished University Professor at U. Maryland and Johns Hopkins University. Former Director, National Science Foundation (1998 - 2006). National Medal of Science awardee. Member, US National Academy of Sciences.



NATIONAL ACADEMY OF SCIENCES



Benny Liu, M.D.

Key Opinion Leader in Niemann-Pick Disease Type C

Gastroenterologist at Alameda Healthy System, CA and Highland Hospital. Globally recognized expert in lipid metabolism.

First to discover that cyclodextrins release cholesterol from cells using an animal model. Assistant Clinical Professor, UCSF.

ALAMEDA HEALTH SYSTEM



Gerald F. Cox, M.D., Ph.D.

Internationally Renowned for Clinical Drug Development Seasoned biotechnology executive with 20-year successful track record of drug development for rare genetic diseases and extensive worldwide regulatory experience





Sharon H. Hrynkow, Ph.D.

Co-Chair

Neuroscientist with more than 25 years' experience in global health arena, public and private sectors.

Senior executive at NIH.

First president of non-profit Global Virus Network.

Former Member of President's Council of Advisors on Science and Technology.5 years at Cyclo Therapeutics leading clinical and scientific programs.





Caroline Hastings, M.D.

Key Opinion Leader in Niemann-Pick Disease Type C

Pediatric hematologist oncologist, Director of NeuroOncology, and Professor of Pediatrics, UCSF Benioff Children's Hospital Oakland.

First physician in US to use cyclodextrins for treatment in NPC, compassionate use. Advisor to US and Australian NPC Advocacy organizations and to physicians globally on NPC.

Benioff Children's Hospitals

Niemann-Pick Disease Type C

Ongoing Pivotal Transport NPC Phase 3 Study



NPC: A Debilitating Disease with Fatal Outcomes

- Rare, fatal and progressive genetic disorder affecting the brain, liver, spleen and lungs
- Characterized by a defect in the NPC1 protein
- Cholesterol and lipids accumulate in cells of major organs and tissues
- Leading to cell and tissue dysfunction

) U.S. Approved NPC Therapies

1 EU Approved Therapy with No Systemic Effects

Market Opportunity¹

United States: \$300 Million | Worldwide: \$600 Million

Incidences

1/100,000 (~35 per year in U.S.)

Of Diagnosis

- ~ 3% are age 3 and below
- ~ 97% are age 3 and above
- ~ 60% age 16 and above

Median Survival

Early Infantile (2m-2): 4.6y Late Infantile (3-6): 9.4y Juvenile (7-15): 15.4y Adolescent/Adult (16+): 12.2y



1: Data on file Cyclo Therapeutics

*Scope: United States + 79 other countries; *Commissioned Tessellon Inc – former Kantar Health experts with 25+ years of epidemiology and forecasting experience, (<u>www.Tessellon.com</u>); *Exhaustive literature search with a broad range of MESH terms.



Trappsol[®] Cyclo[™]

Enables the Effective Transport of Cholesterol Out of Cells



Cholesterol as measured by Filipin staining at Baseline and after 7 doses over 14 weeks



The lack of light blue represents the clearing of cholesterol from cells



Trappsol[®] Cyclo[™] Summary of Completed Clinical Studies in NPC

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Study 101

Phase 1 study in NPC patients age 18 years and older showed Trappsol[®] Cyclo[™] was welltolerated with an acceptable safety and tolerability profile

- After IV infusion, the drug is detectable in the cerebrospinal fluid within hours after the start of infusion
- Cholesterol synthesis and metabolism affected, and cholesterol cleared from cells, mimicking effects from nonclinical studies (*in vitro* and *in vivo*) in NPC models

Study 201

Consistent pharmacodynamic effects and safety profile observed in a 48-week Phase 1/2 study in NPC patients aged 2 years and older

- 100% of patients assessed by treating physicians to be either stable or improved
- 88% (8 of 9 patients who completed the study), experienced clinically meaningful improvements in one or more efficacy endpoints, assessed by the 17 Domain NPC Severity Scale
- Based on totality of data from the Phase 1 and Phase 2 studies, the 2000 mg/kg dose was selected for the Phase 3 study



Transport NPC Ongoing Pivotal Phase 3 Study in Niemann-Pick Disease Type C



Double-blind, randomized, placebo-controlled, parallel-group study and is currently the most advanced clinical research program underway to identify a treatment for NPC

Number of Subjects	93					
Current Sites	23 across 9 countries	United States, United Kingdom, Italy, Germany, Spain, France, Poland, Israel, and Australia				
Duration	96-week trial, with interim analysis at 48 weeks					
Dose	2000 mg/kg via IV infusion					
Primary Endpoint	NPC Composite Severity Score					
Secondary Endpoints	SCAFI, Swallow, Vineland-2					
Exploratory Endpoints	Inclusive of speech, liver and lung function					



We Have the Only Active Late-Stage Clinical Program in NPC

Company	Indication	Preclinical	Phase 1	Phase 2	Phase 3	Potential Indication (Systemic and Neurological)	Safety Profile	Summary
Cyclo therapeutics	Trappsol [®] Cyclo™ (Intravenous every 2 weeks, home infusions)							Met all primary endpoints of the Ph1 and Ph1/2 showing favorable safety and efficacy. Ph3 currently enrolling and additional site activation ongoing.
ACTELION	Zavesca* (Oral 3 times daily)							FDA: Data did not support benefit risk. Off-label in US. EMA: Approved January 2009.
ORPHAZYME	Arimoclomol (Oral 3 times daily)				*			FDA: CRL received June 18, 2021, noting additional data required to support benefit risk. EMA: 23-Feb-2022, CHMP has notified of a negative trend vote following Oral Explanation. CHMP's current orientation is to not approve arimoclomol when it convenes by the end of March 2022.
IntraBio	IB1001 (Oral 3 times daily)							Met with FDA, EMA and UK, Phase 3 study required, 6-month placebo- controlled crossover trial with approximately 50 patients (4 years of age and above). Study enrollment anticipated H1 2022.
Mandos Health	Adrabetadex (VTS-270) (Intrathecal every 2 weeks)				*			MNK concluded program 20-Jan-2021 noting the benefit risk is negative. Mandos received Court Approval to buy from MNK 29-Jun-2021. FDA has not found drug to be safe and effective for use to treat NPC1 or for any other use.
🚑 ESCAPE BIO	ESB1609 (Oral, brain- penetrant			•		TBD	TBD	Selective sphingosine-1-phosphate 5 (S1P5) receptor agonist.
	ſH					Differ	rentiation Key:	- Optimal - Moderate - Weak

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Alzheimer's Disease

Open IND for Phase 2 study





Alzheimer's Disease The Most Common Form of Dementia

An irreversible, progressive neurologic disorder that slowly degrades memory, thinking and social skills that affects a person's ability to function independently.

Similarities with NPC

- Cognitive decline
- Elevated levels of tau
- Amyloid plaques



- Affects more than 5 million people in the U.S.¹
- 6th leading cause of death in the U.S.¹
- 500,000 new cases every year²
- 13.8 million cases projected by 2050¹

Alzheimer's Disease

Secondary Tauopathy



Biologic Similarities

Cholesterol Accumulation in Regions of Brain Elevated Levels of Tau in CSF

Amyloid Plaques in the Brain

Niemann-Pick Disease Type C

Primary Tauopathy

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Disease Manifestation

Cognitive decline / dementia Premature death Clumsiness Progressive motor symptoms Ataxia, dystonia, dysarthria, dysphasia Psychiatric signs: psychosis, depression Weight loss

Disease Manifestation

Progressive cognitive decline / early dementia Premature death Clumsiness, gait disturbance Delayed motor milestones Progressive: ataxia, dystonia Seizures Weight loss



Trappsol[®] Cyclo[™] for the Potential Treatment of Alzheimer's Disease Targeting Reduction of Amyloid Beta and Tau

Received IND Clearance from the U.S. FDA to Advance Phase 2 Study

Preeminent Neuroscientist and World-Renowned Researcher, Cynthia A. Lemere, PhD Senior Advisor for Advancement of Alzheimer's Disease Asset

Positive Results in Alzheimer Patient Under Compassionate Use Program

FDA authorized use of Trappsol[®] Cyclo[™] in geriatric patient

18 months of monthly IV infusion

Disease did not progress

Family reported less volatility and greater word-finding ability

18 months of data has led to development of Phase 2 protocol



Alzheimer's Mini-Mental State Evaluation Performance¹

Cyclo Patient Cognitively Normal = 30 ··· O·· Expected Decline of AD Patients

"The patient has shown cognitive and neurologic stability in serial examinations during this study that indicates possible benefit as there would be an expected measurable cognitive and functional decline over an 18-month period in persons with Alzheimer's disease dementia, "Treating Physician



Corporate Overview

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Financial Snapshot - Nasdaq: CYTH

Cash Balance¹ \$16.6 Market Cap² ~\$22M

Shares Outstanding **8.4M** Average Volume² ~49K



Investment Summary

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Leveraging over 3 decades of experience with cyclodextrins to advance clinically de-risked programs towards approval in diseases with unmet medical need

Platform technology has demonstrated to be safe and effective with over 10 years of patient exposure

Transport NPC

Enrolling and dosing patients in Pivotal Phase 3 study in Niemann-Pick Disease Type C

Significant market opportunity with no approved therapy to treat both systemic and neurological manifestations of NPC

FDA: Orphan Drug Designation (ODD), Fast-Track, Rare Pediatric Disease Designation, potential PRV; EMA: ODD and adopted PIP Pipeline expansion into Alzheimer's Disease (AD), patent filed globally and is currently being prosecuted

Received Study May Proceed from FDA December 2021, FPI targeted H2 2022



Multiple value-driving milestones expected

Platform technology with opportunity to expand into multiple indications Leadership team with proven track-record in execution and value creation

therapeutics

Thank you!