

General Corporate & Gene Therapy Overview:

*At the Forefront of the Human
Genome Medicine Revolution*

April 2020



Forward-Looking Statements

This presentation contains "forward-looking statements" within the meaning of the Private Securities Litigation Reform Act of 1995 relating to preclinical and clinical development of our product candidates, the timing and reporting of results from preclinical studies and clinical trials, the prospects and timing of the potential regulatory approval of our product candidates, commercialization plans, manufacturing and supply plans, financing plans, and the projected revenues and cash position for the Company. The inclusion of forward-looking statements should not be regarded as a representation by us that any of our plans will be achieved. Any or all of the forward-looking statements in this presentation may turn out to be wrong and can be affected by inaccurate assumptions we might make or by known or unknown risks and uncertainties. For example, with respect to statements regarding the goals, progress, timing, and outcomes of discussions with regulatory authorities, and in particular the potential goals, progress, timing, and results of preclinical studies and clinical trials, actual results may differ materially from those set forth in this release due to the risks and uncertainties inherent in our business, including, without limitation: the potential that results of clinical or preclinical studies indicate that the product candidates are unsafe or ineffective; the potential that it may be difficult to enroll patients in our clinical trials; the potential that regulatory authorities, including the FDA, EMA, and PMDA, may not grant or may delay approval for our product candidates; the potential that we may not be successful in commercializing Galafold in Europe, Japan, the US and other geographies or our other product candidates if and when approved; the potential that preclinical and clinical studies could be delayed because we identify serious side effects or other safety issues; the potential that we may not be able to manufacture or supply sufficient clinical or commercial products; and the potential that we will need additional funding to complete all of our studies and manufacturing. Further, the results of earlier preclinical studies and/or clinical trials may not be predictive of future results. In addition to financial information prepared in accordance with U.S. GAAP, this presentation also contains adjusted financial measures that we believe provide investors and management with supplemental information relating to operating performance and trends that facilitate comparisons between periods and with respect to projected information. These adjusted financial measures are non-GAAP measures and should be considered in addition to, but not as a substitute for, the information prepared in accordance with U.S. GAAP. With respect to statements regarding projections of the Company's revenue and cash position, actual results may differ based on market factors and the Company's ability to execute its operational and budget plans. In addition, all forward-looking statements are subject to other risks detailed in our Annual Report on Form 10-K for the year ended December 31, 2019 to be filed today. You are cautioned not to place undue reliance on these forward-looking statements, which speak only as of the date hereof. All forward-looking statements are qualified in their entirety by this cautionary statement, and we undertake no obligation to revise or update this news release to reflect events or circumstances after the date hereof.

In addition to financial information prepared in accordance with U.S. GAAP, this presentation also contains adjusted financial measures that we believe provide investors and management with supplemental information relating to operating performance and trends that facilitate comparisons between periods and with respect to projected information. These adjusted financial measures are non-GAAP measures and should be considered in addition to, but not as a substitute for, the information prepared in accordance with U.S. GAAP. We typically exclude certain GAAP items that management does not believe affect our basic operations and that do not meet the GAAP definition of unusual or non-recurring items. Other companies may define these measures in different ways. Full reconciliations of GAAP results to the comparable non-GAAP measures for the reported periods appear in the financial tables section of this presentation. When we provide our expectation for non-GAAP operating expenses on a forward-looking basis, a reconciliation of the differences between the non-GAAP expectation and the corresponding GAAP measure generally is not available without unreasonable effort due to potentially high variability, complexity and low visibility as to the items that would be excluded from the GAAP measure in the relevant future period, such as unusual gains or losses. The variability of the excluded items may have a significant, and potentially unpredictable, impact on our future GAAP results.

A RARE COMPANY

A leading fully-integrated, global rare disease biotechnology company



First Oral Precision
Medicine for Fabry Disease



Gene Therapy
PLATFORM
Protein Engineering
& Glycobiology



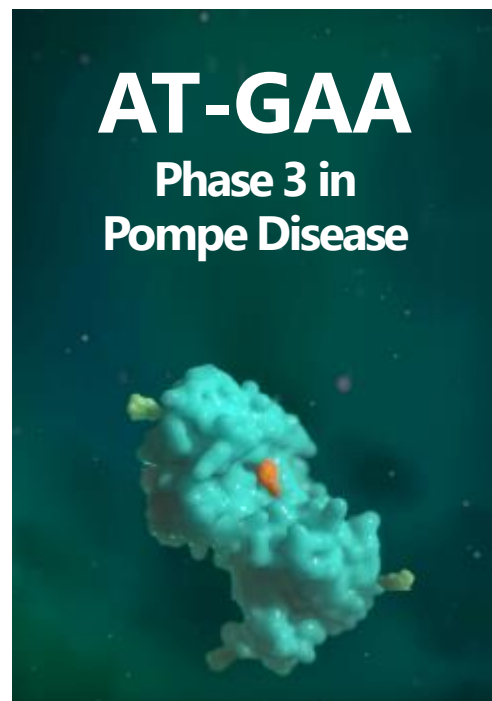
World Class
BIOLOGICS
Capabilities



EMPLOYEES
in 27 Countries



AT-GAA
Phase 3 in
Pompe Disease



\$450M+
Cash
as of 12/31/19

**Two Clinical-
Stage Gene
Therapies**

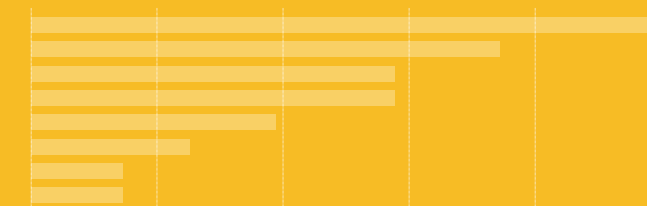


**GLOBAL
COMMERCIAL
ORGANIZATION**



**Robust R&D
Engine**

Nearly 50+ Lysosomal
Disorders and More
Prevalent Rare Diseases



Key Takeaways

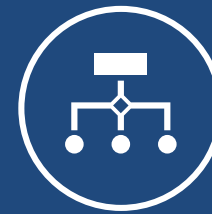
Recent successes across our science, clinical, regulatory and commercial efforts position us for the future



Galafold Continues
Strong Launch
Performance &
Cornerstone of
Amicus Success



AT-GAA for Pompe
Advances Toward
Approval as “Crown
Jewel” of Amicus
Portfolio



Portfolio of Gene
Therapy Programs
and Technologies
Provides Foundation
for Future



Strong Financial
Outlook with Current
Cash Well into 2022

A RARE PORTFOLIO

	DISCOVERY	PRECLINICAL	PHASE 1/2	PHASE 3	REGULATORY	COMMERCIAL
Fabry Franchise						
Galafold® (migalastat) Monotherapy ODD						
Fabry Gene Therapy	PENN					
Pompe Franchise						
AT-GAA (Novel ERT + Chaperone) ODD						
Pompe Gene Therapy	PENN					
Batten Franchise – Gene Therapies						
CLN6 Batten Disease ODD RPD	NCH					
CLN3 Batten Disease ODD RPD	NCH					
CLN8 Batten Disease	NCH					
CLN1 Batten Disease	NCH					
Next Generation Research Programs and CNS Gene Therapies						
CDKL5 Deficiency Disorder GTx / ERT	PENN					
Niemann-Pick Type C (NPC)	NCH / PENN					
Others	NCH / PENN					
MPS Franchise						
Mepsevii™ (vestronidase alfa) <i>(Japan Only)*</i>						
Next Generation MPSIIIA	PENN					
MPSIIIB	PENN					

LEGEND

ODD - Orphan Drug Designation

RPD - Rare Pediatric Disease Designation

*Exclusive license from Ultragenyx for Japanese rights to Mepsevii™, investigator-sponsored trial in Japan underway



Galafold[®] (migalastat) Global Launch...

...taking a leadership role in
the treatment of Fabry disease

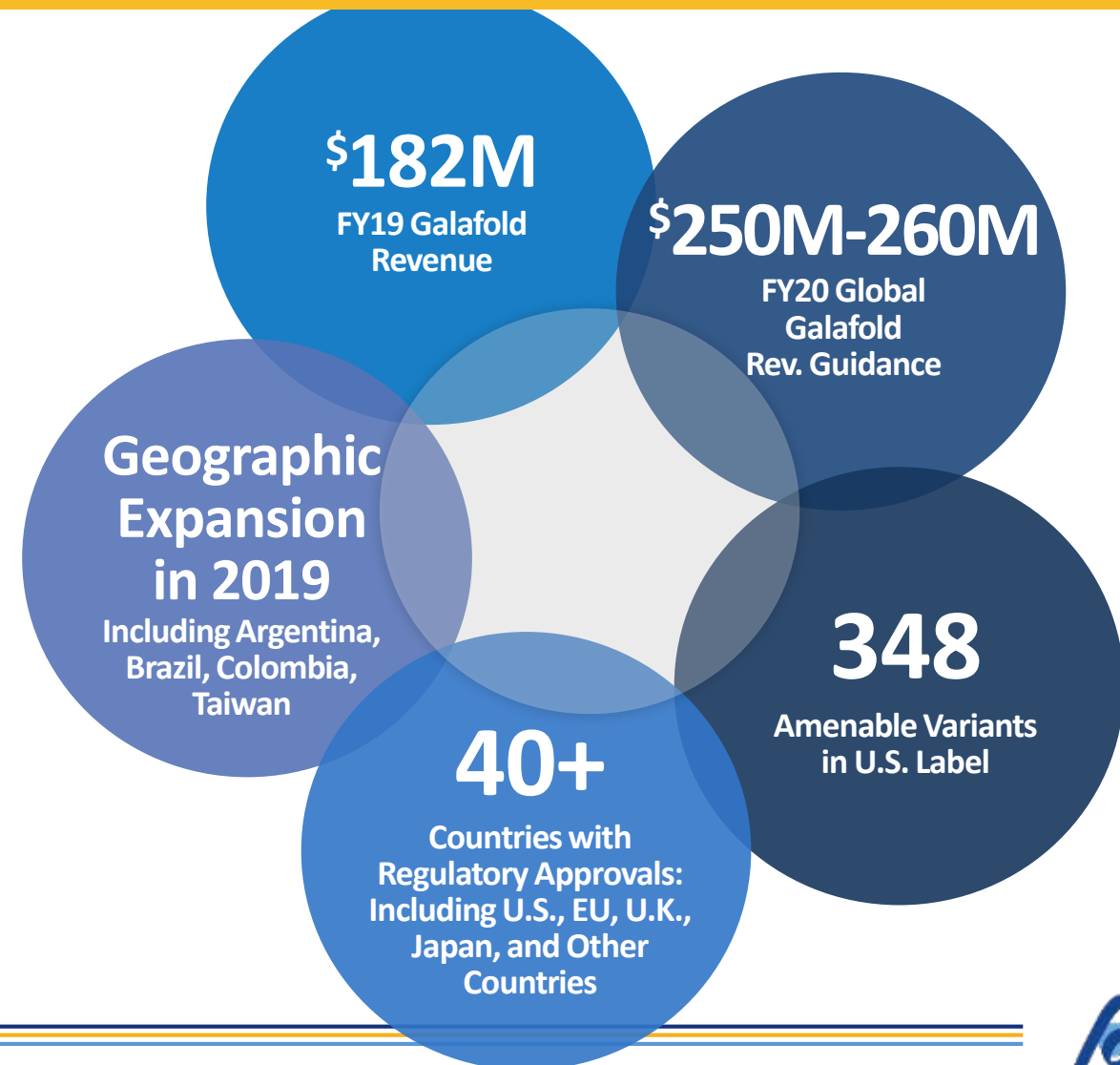
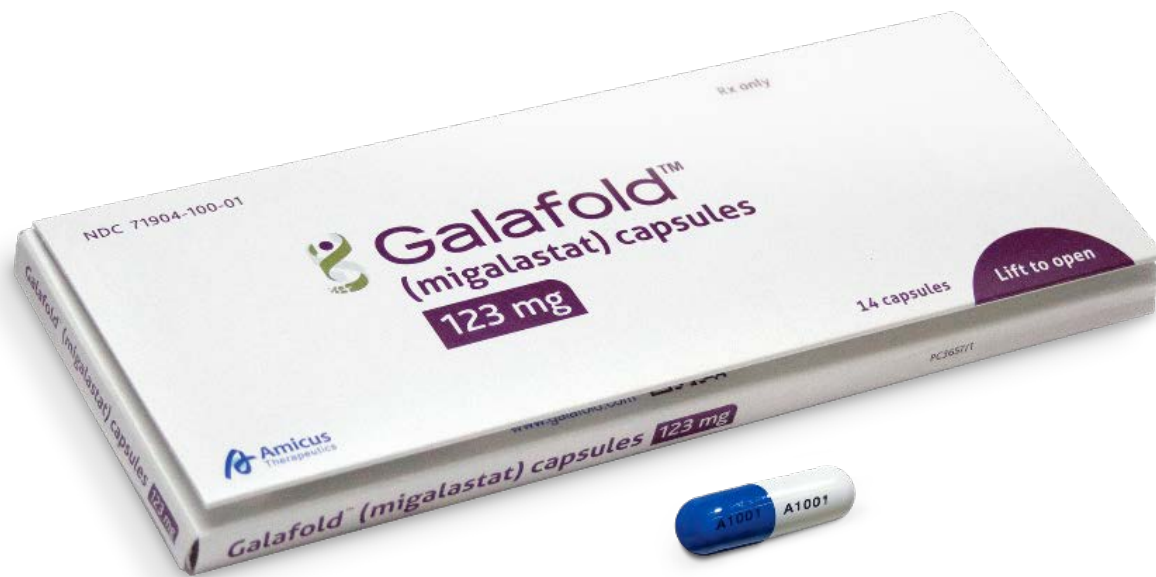
“We push ideas as far and as fast as possible”

- Amicus Belief Statement

Galafold Snapshot (as of December 31, 2019)

Galafold is the cornerstone of Amicus' success. It is an orally delivered small molecule precision medicine with a unique mechanism of action for Fabry patients with amenable variants that replaces the need for intravenously delivered enzyme replacement therapy

One of the Most Successful Rare Disease Launches

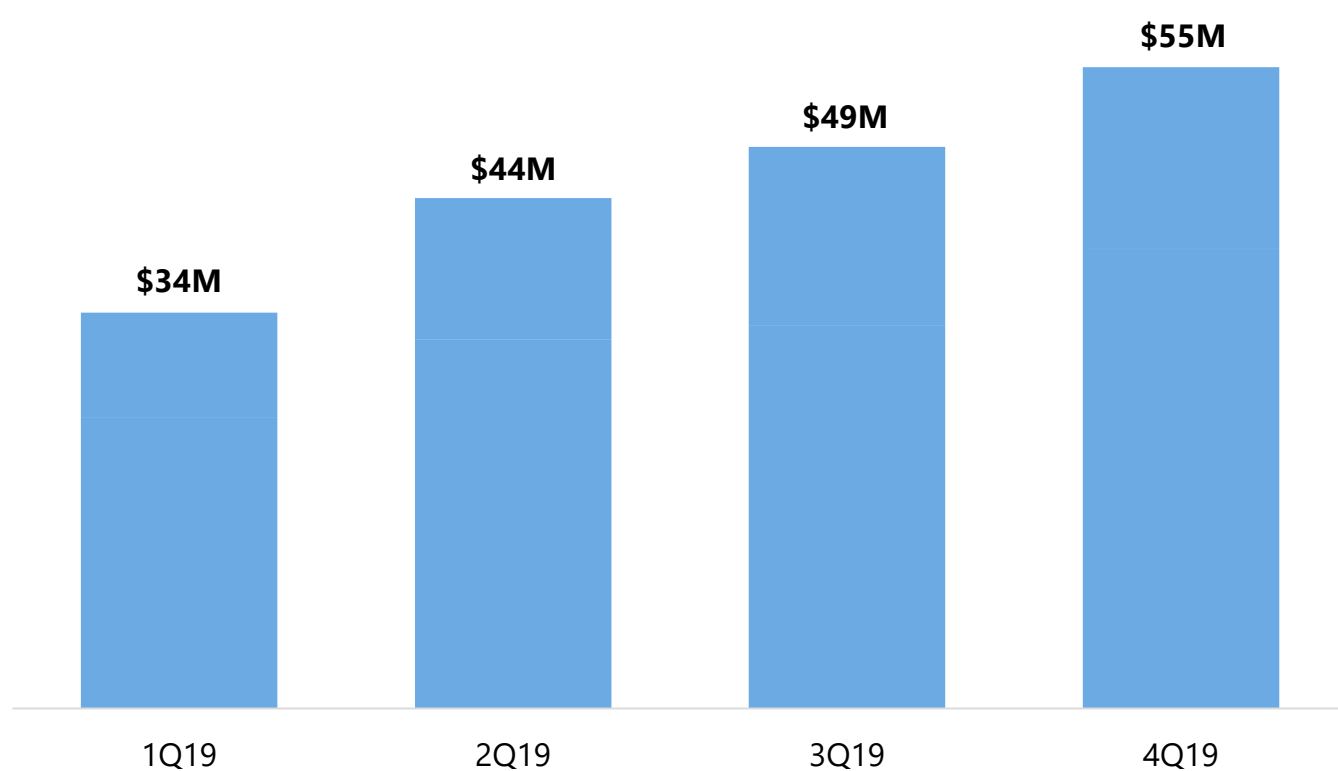


Galafold is indicated for adults with a confirmed diagnosis of Fabry Disease and an amenable mutation/variant. The most common adverse reactions reported with Galafold (≥10%) were headache, nasopharyngitis, urinary tract infection, nausea and pyrexia. For additional information about Galafold, including the full U.S. Prescribing Information, please visit <https://www.amicusrx.com/pi/Galafold.pdf>. For further important safety information for Galafold, including posology and method of administration, special warnings, drug interactions and adverse drug reactions, please see the European SmPC for Galafold available from the EMA website at www.ema.europa.eu.

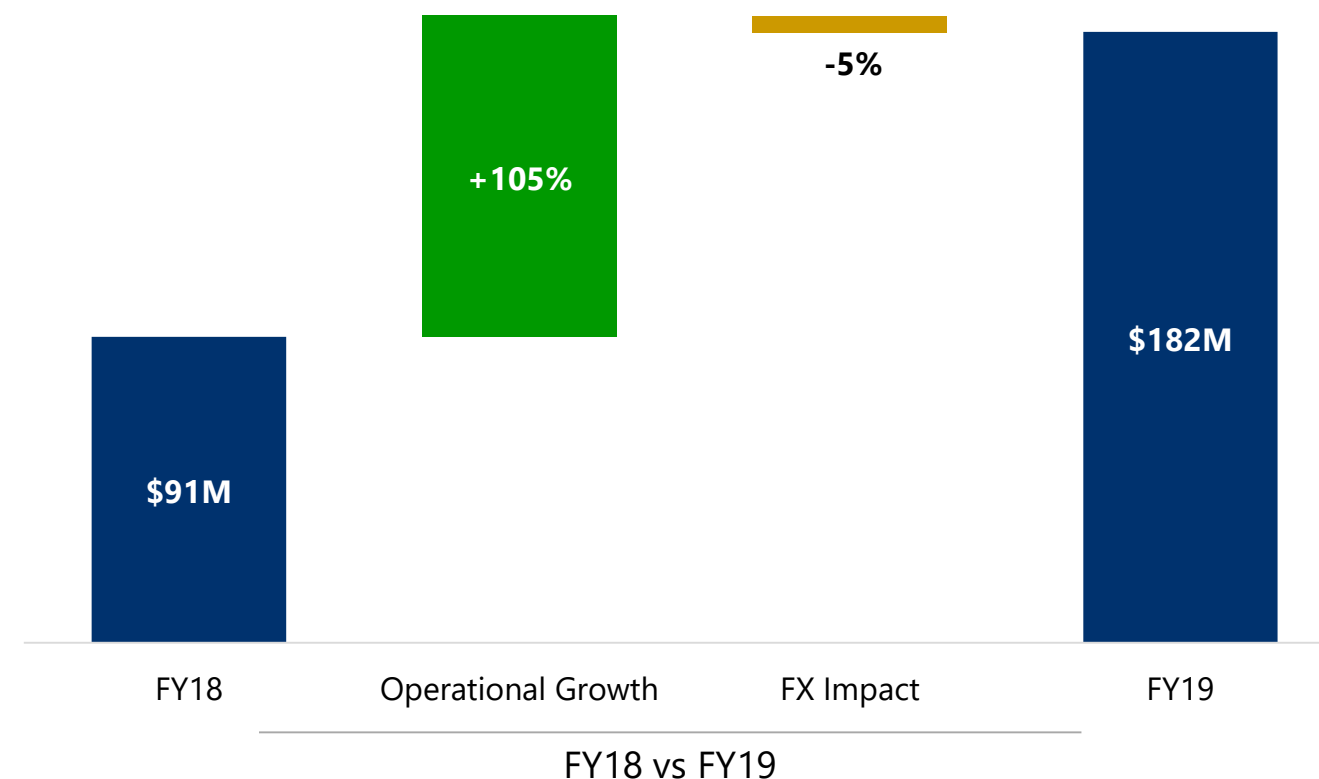
2019 Galafold Success

Strong full-year revenue performance of \$182M, exceeding guidance of \$170M-\$180M

Quarterly Galafold Sales



Year-over-Year Galafold Sales Growth



Galafold Global Launch Momentum (as of December 31, 2019)

Global commercial metrics continue to be very strong with >90% compliance and adherence, 30% global market share of treated amenable patients and continued broad market access

FY19 Strength Reflects Positive Momentum Across All Key Global Commercial Metrics and 1,000+ Treated Patients

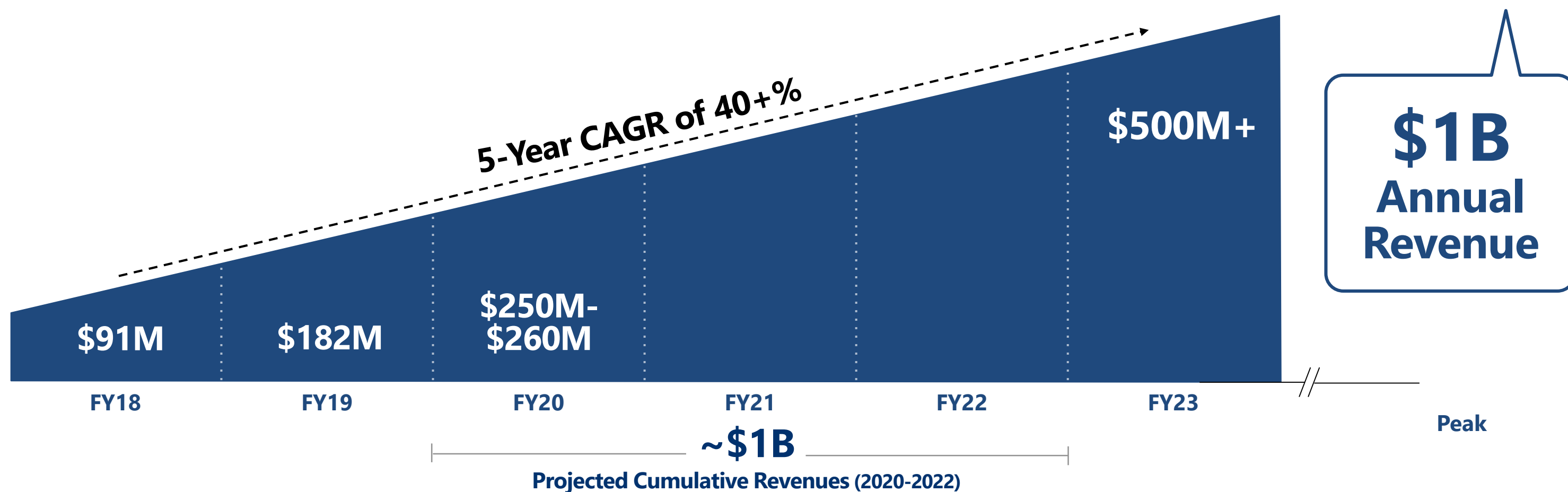
- **Global:** 30%+ estimated global market share of treated amenable patients (as of 12/31/19)*
- **U.S.:** Steady growth in adoption from 100+ prescribers and broad reimbursement coverage
- **EU:** Accelerated patient growth in new and established markets throughout 2019
- **APAC:** Continued strong contribution from Japan and Australia
- **LATAM:** New approvals in Brazil, Colombia and Argentina lay strong foundation for future growth
- **Demographics:** Global mix of switch (65%) and previously untreated patients (35%)



*Market share based on reported global Fabry sales for the calendar year ending 4Q19 and assumes a 35% amenability rate.

Galafold Growth Trajectory

Galafold is on track to generate \$1B+ in projected cumulative revenues from 2020-2022 and is on an anticipated path to \$500M+ in annual sales in 2023 and \$1B+ annual sales at peak





AT-GAA: Next Potential Standard of Care for Pompe Disease

“We encourage and embrace constant innovation”

- Amicus Belief Statement

Pompe Disease Overview

Pompe disease is a severe and fatal muscular dystrophy and one of the most prevalent lysosomal disorders with very high unmet medical need



5,000 – 10,000+ patients diagnosed WW¹; newborn screening suggests underdiagnosis

Age of onset ranges from infancy to adulthood

Patients on current standard of care decline after ~2 years

Respiratory and cardiac failure are leading causes of morbidity and mortality

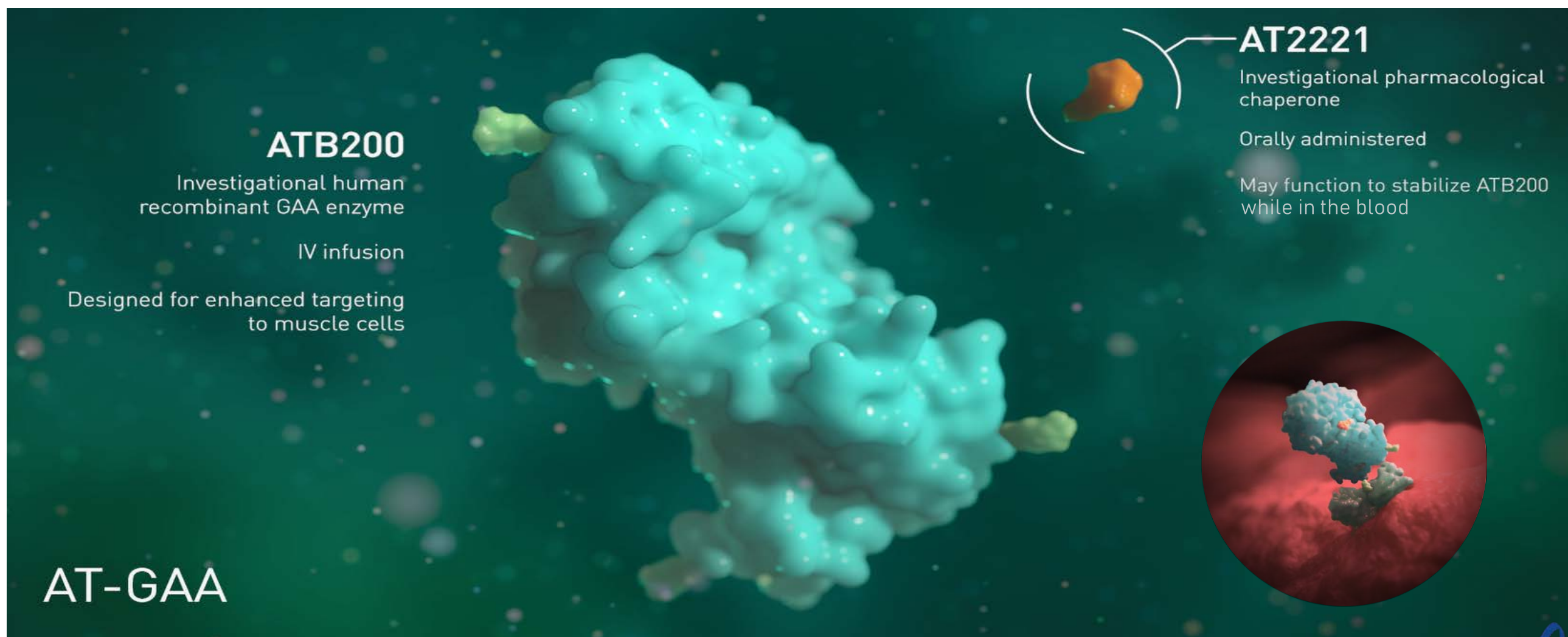
Deficiency of GAA leading to glycogen accumulation and cellular dysfunction

Symptoms include muscle weakness, respiratory failure, and cardiomyopathy

~\$1B+ global Pompe ERT sales²

AT-GAA: Foundation in Protein Engineering

Amicus scientists specializing in protein engineering and glycobiology created a uniquely glycosylated and highly phosphorylated ERT (AT-GAA) that significantly enhances targeting to key muscles affected in patients



U.S. FDA Granted BTDD to AT-GAA in Late-Onset Pompe Disease (LOPD)

AT-GAA is the first ever second-generation product for any lysosomal disorder to earn FDA Breakthrough Therapy Designation (BTDD)

Plans to apply for and initiate a rolling BLA submission for AT-GAA in LOPD in 2020



AT-GAA BTDD Based on Ph 1/2 Clinical Efficacy

- Improvements in 6-minute walk distance
- Comparison to natural history of treated patients



BTDD Features

- Intensive guidance on an efficient drug development program
- Organizational commitment involving senior agency staff
- All Fast Track program features including rolling submission



BTDD Criteria

- Intended to treat a serious or life-threatening disease or condition
- Preliminary clinical evidence indicates drug may demonstrate substantial improvement over existing therapies on one or more clinically significant endpoints

Pompe Patient Experience in Phase 1/2 Clinical Study (ATB200-02)

Phase 1/2 results showed strong and durable effects in patients out to two years, leading to dramatic improvements in muscle strength and function, as well as significant improvements in key biomarkers of disease



6-Min Walk Test (m)	Cohort	Baseline (n=10)	Change at Month 6 (n=10 ^a) Mean (SD)	Change at Month 12 (n=10 ^a) Mean (SD)	Change at Month 24 (n=9 ^{a,b}) Mean (SD)
	Cohort 1 ERT-Switch Ambulatory	397.2 (96.8)	+23.9 (52.2)	+42.2 (46.5)	+36.4 (60.5)
	Cohort 3 ERT-Naïve	399.5 (83.5)	+41.8 (29.4)	+63.1 (29.1)	+60.7 (36.5)

FVC (% Predicted)	Cohort	Baseline (n=9 ^c)	Change at Month 6 (n=9 ^c) Mean (SD)	Change at Month 12 (n=9 ^c) Mean (SD)	Change at Month 24 (n=8 ^{b,c}) Mean (SD)
	Cohort 1 ERT-Switch Ambulatory*	52.6 (14.7)	-1.2 (4.0)	-3.0 (6.0)	+0.9 (4.9)
	Cohort 3 ERT-Naïve	53.4 (20.3)	+4.4 (5.6)	+4.6 (8.8)	+6.8 (6.8)

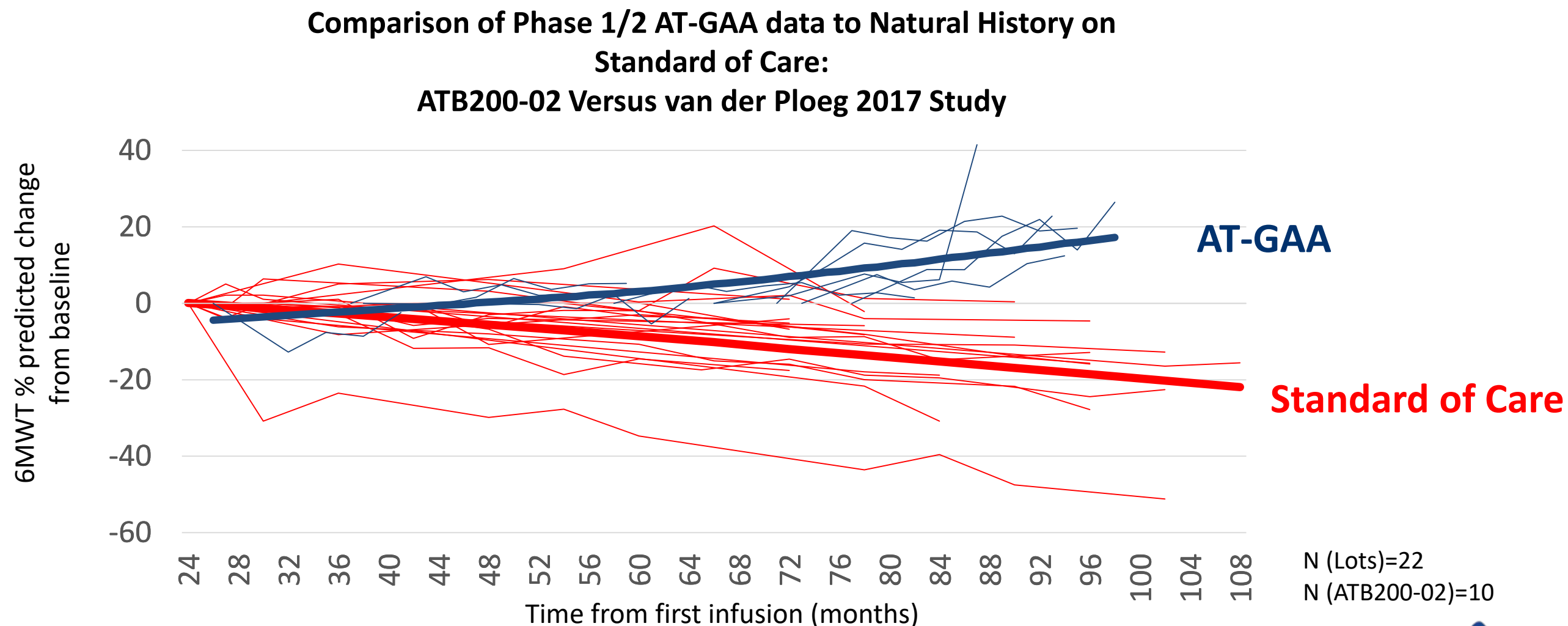
Data from interim analysis 8.

^a One patient in Cohort 1 discontinued after 18 weeks due to burden of travel; baseline value is not shown for this patient. ^b One patient in Cohort 1 discontinued from study before Month 24. ^c Baseline FVC not available for 1 patient in Cohort 1

6MWT Natural History: Phase 1/2 AT-GAA Data vs. Medical Literature

van der Ploeg 2017

Improvement in percentage predicted 6MWD seen in all patients who switched from alglucosidase alfa to AT-GAA

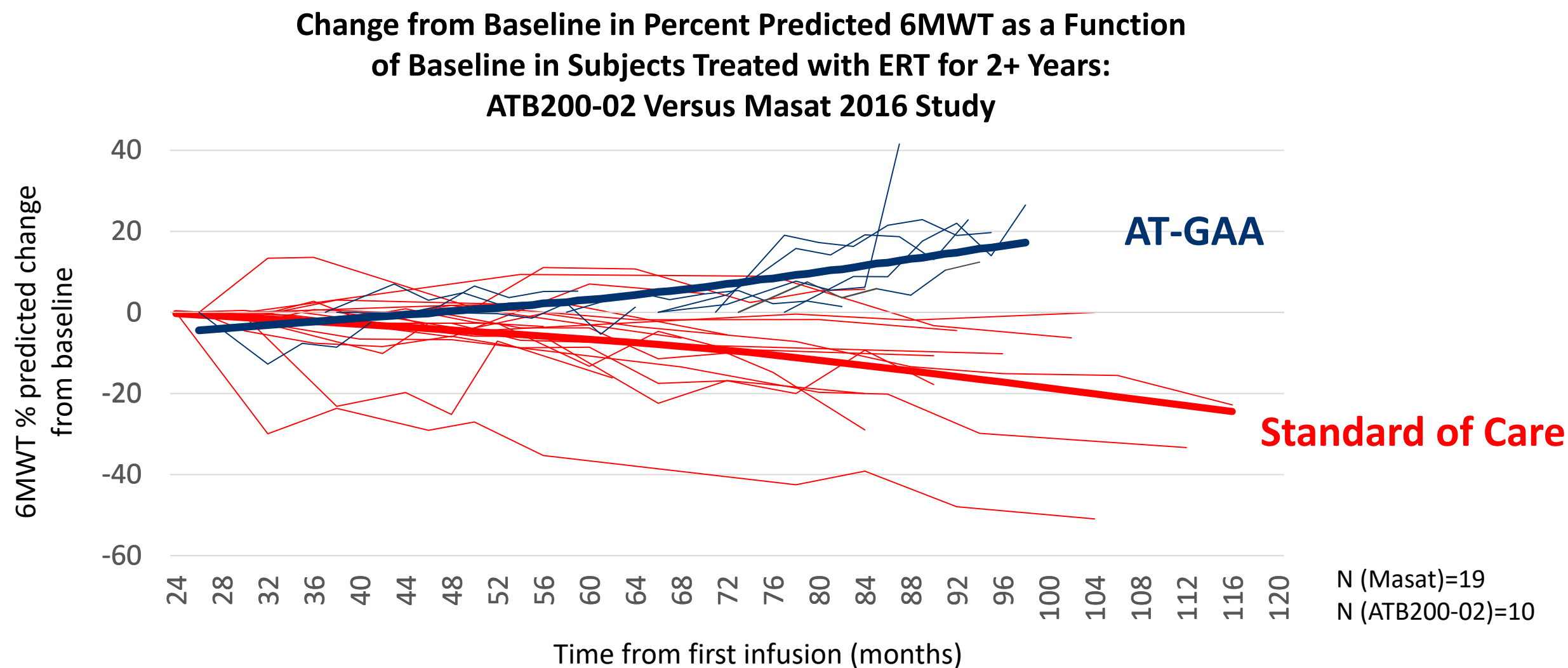


1. Data for AT-GAA represent time from first infusion of SOC ERT and change from baseline at the time of switching from SoC to AT-GAA
2. Source: ATB200-02 IA#7; Ans T. van der Ploeg et al. Poster presented at the 13th Annual WORLD Symposium™ 2017, February 13–17, 2017, San Diego, CA, USA

6MWT Natural History: Phase 1/2 AT-GAA Data vs. Medical Literature

Masat 2016

Second natural history data set confirms the large treatment effect of AT-GAA



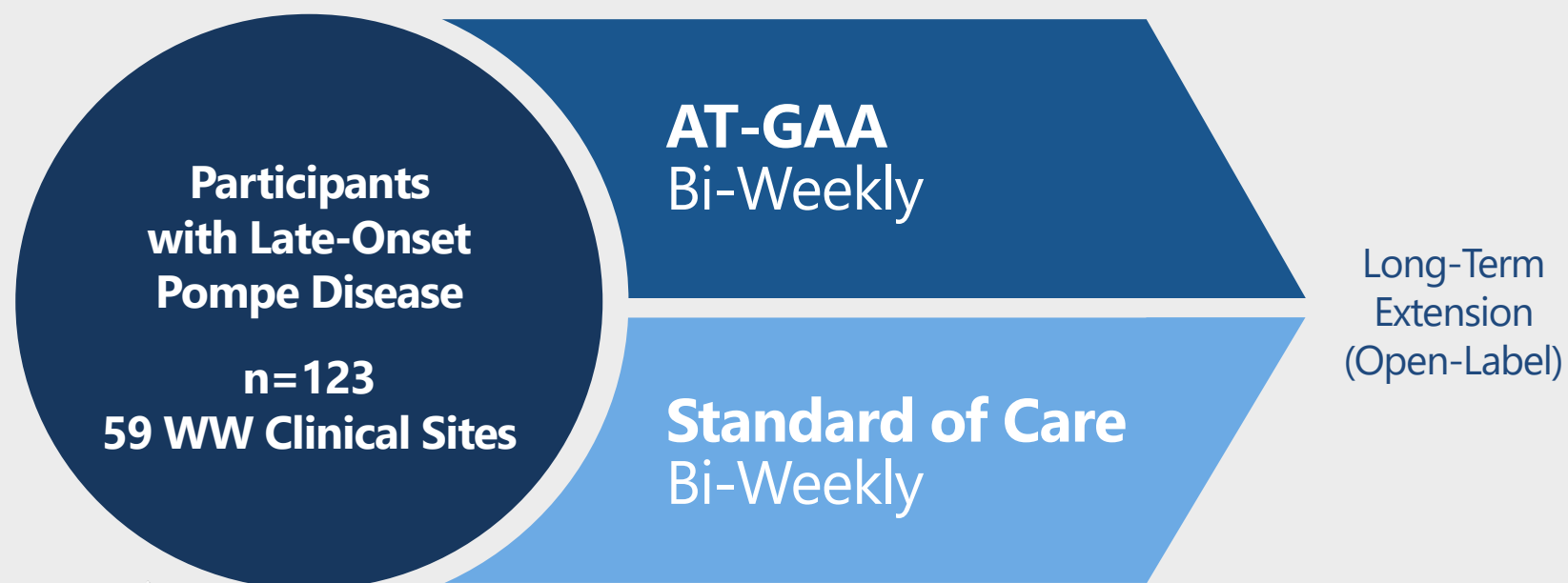
PROPEL (ATB200-03) Study Design

PROPEL 

Phase 3 exceeded enrollment with data expected in 1H2021. The study is highly powered for success and supports a broad label, with FDA and EMA agreement on study design and primary endpoint (6MWT)

52-Week Primary Treatment Period (Double-Blind)

2:1 Randomization



Primary Endpoint: 6-Minute Walk Test at Week 52;
Multiple Secondary Endpoints

- PROPEL pivotal study over-enrolled with data expected in 1H2021
- Study includes ERT-Switch and ERT-Naïve Patients
- FDA and EMA agreed upon primary endpoint of 6MWD, an integrated measure of disease progression that evaluates both cardiopulmonary and musculoskeletal systems

AT-GAA: Key Takeaways



AT-GAA for Pompe
Advances Toward
Approval as “Crown
Jewel” of Amicus
Portfolio

- PROPEL pivotal study exceeded enrollment with data expected 1H2021
- Breakthrough Therapy Designation and the Promising Innovative Medicine designation highlight unmet need in Pompe disease today
- Plan to submit and initiate rolling submission of Biologics License Application in 2020
- Manufacturing PPQ runs at WuXi biologics on track
- Peak revenue potential of \$1B-\$2B, with exclusivity well into 2030s



Next Generation Gene Therapy Platform

“We have a duty to obsolete our own technologies”

- Amicus Belief Statement

A Natural Evolution: Chaperones to Optimized ERT to Gene Therapy

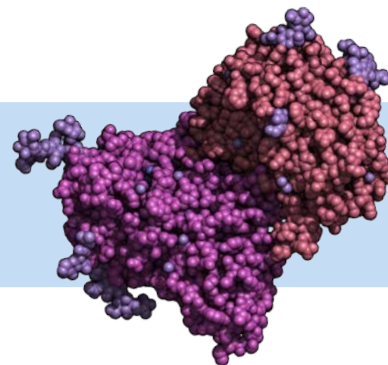
Amicus' expansion into gene therapy is built upon years of experience in developing genetic medicines designed to deliver deficient proteins to target cells and organelles

Pharmacological Chaperones



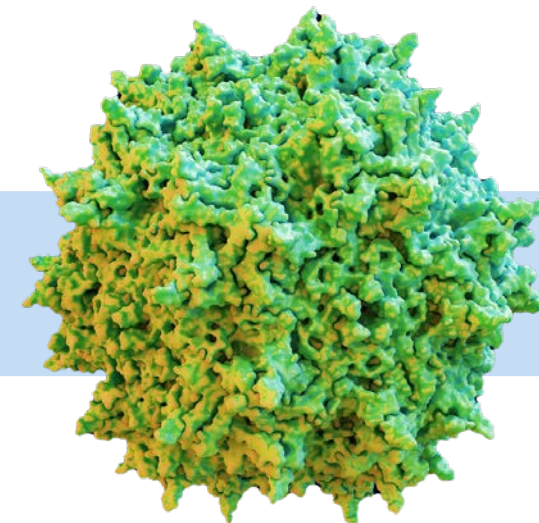
Stabilize
"naturally produced" enzymes

Next-Generation ERTs



Stabilize and target
"externally produced" enzymes

Gene Therapies



Stabilize and target
"internally produced" enzymes

Validated Gene Therapy Platform for CNS

The Amicus Batten programs leverage AAV technologies and platforms utilized in the neuromuscular space at Nationwide Children's Hospital/Sanford and have robust preclinical and now clinical proof of concept

Clinically validated AAV gene therapy approach at NCH and Sanford

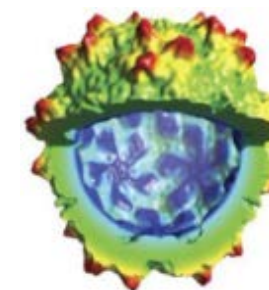
- Nationwide Children's Hospital Center for Gene Therapy (NCH)
- Intrathecal delivery with robust expression throughout CNS

Preclinical safety and efficacy studies replicated across multiple diseases at NCH

- SMA, CLN6, CLN3, CLN8

Amicus applying platform to multiple types of Batten disease and other Neurologic LSDs

- Two clinical programs in CLN6 and CLN3 Batten disease show initial safety in 15 patients; promising efficacy results in first 8 patients in CLN6
- Active preclinical programs in CLN8 and CLN1 Batten disease with other neurologic LSDs in earlier preclinical development



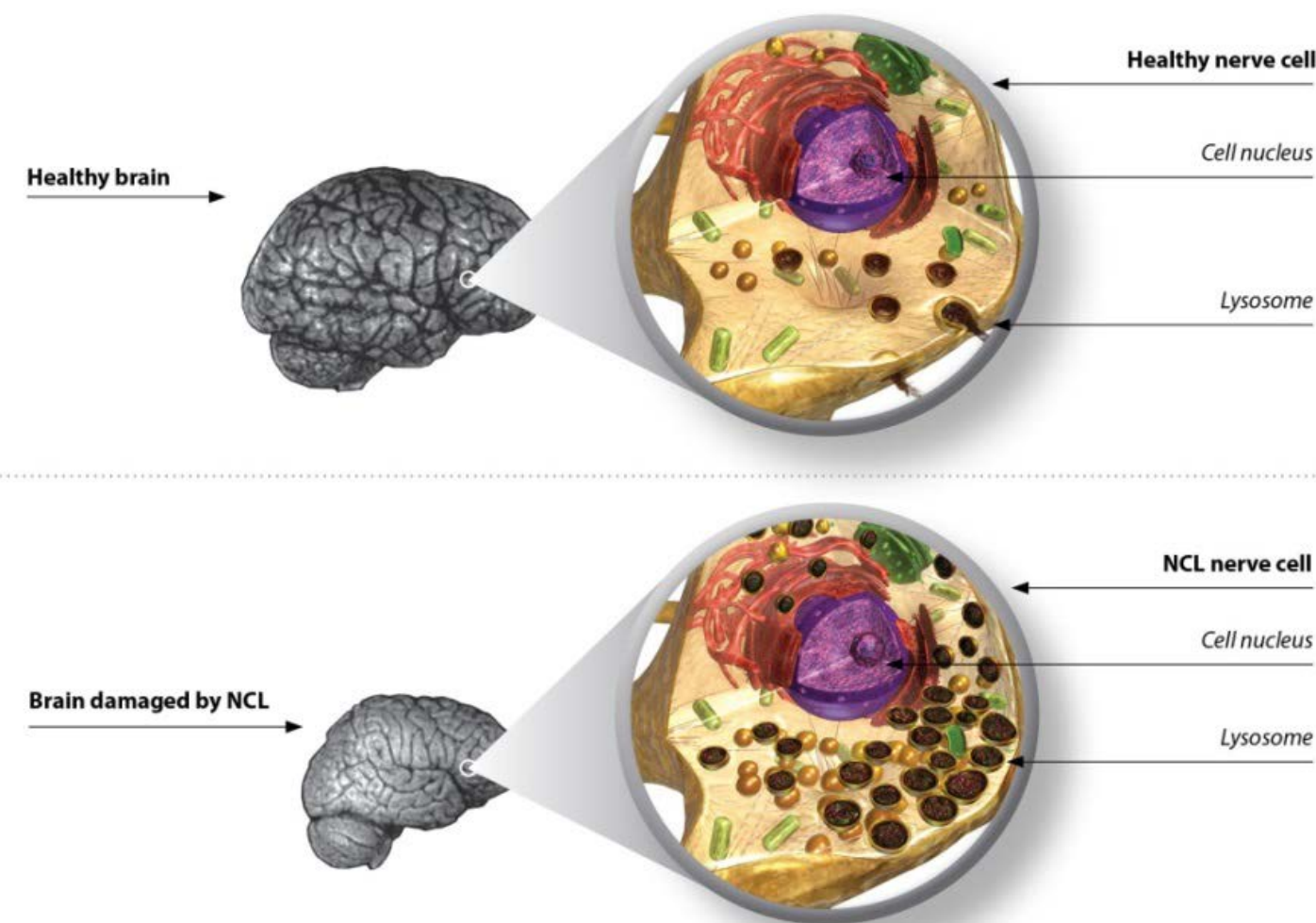
Foust, Kaspar et al, 2009

Batten Disease Overview

Batten disease is a devastating early childhood disease that is 100% fatal in children

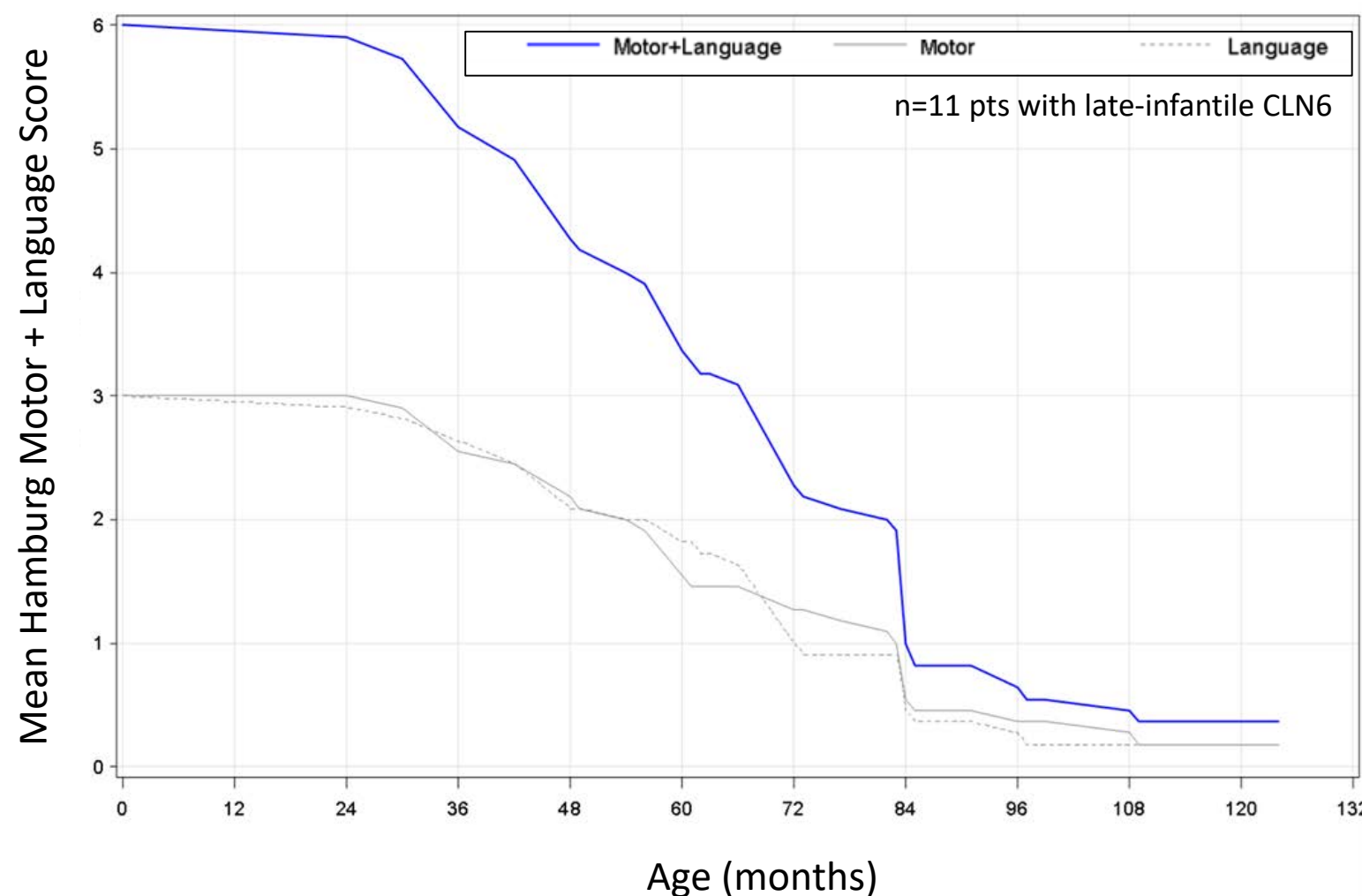
Disease Overview

- A group of disorders known as neuronal ceroid lipofuscinoses (NCLs), collectively referred to as Batten disease
- Mutation in one of 13 different CLN genes leads to lysosomal dysfunction
- Signs and symptoms include loss of speech, ambulation, vision and cognition



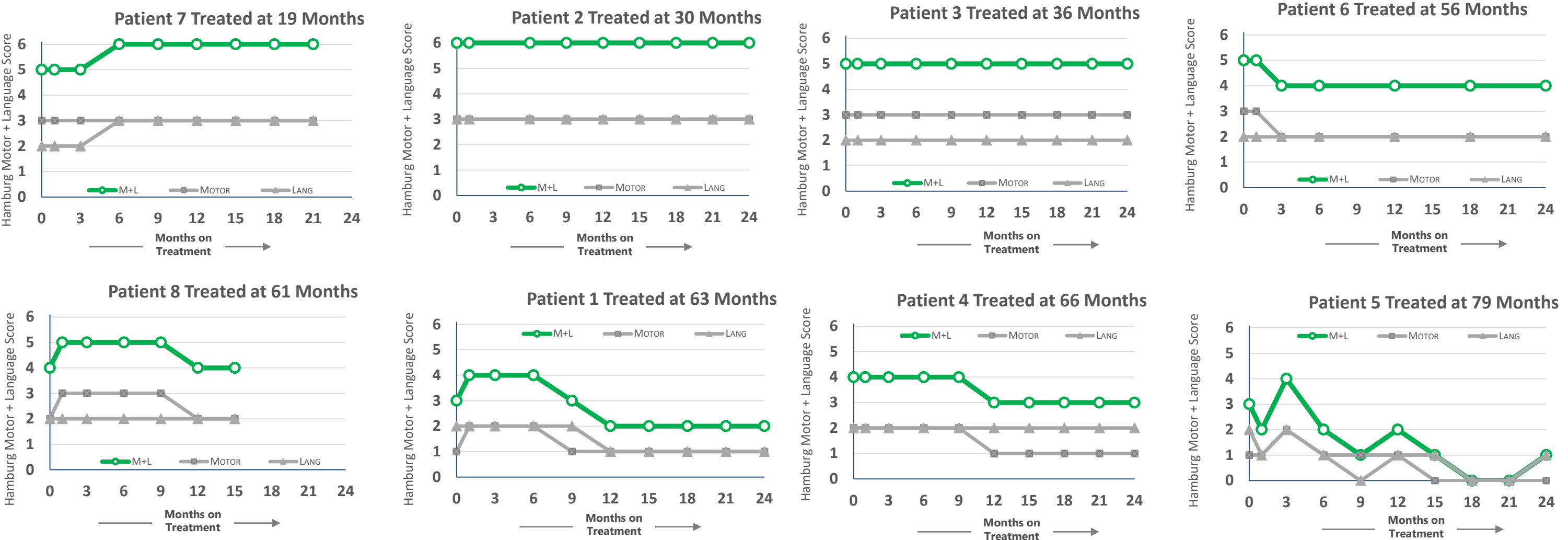
CLN6 Natural History

CLN6 natural history shows a progressive decline of approximately one point per year in the Hamburg score from age two onwards with similar decline in motor and language



Clinical Efficacy: Combined and Individual Hamburg Scores (n=8)

CLN6 gene therapy halts or substantially slows progression of disease with a positive impact on motor and language function in 7 out of 8 patients



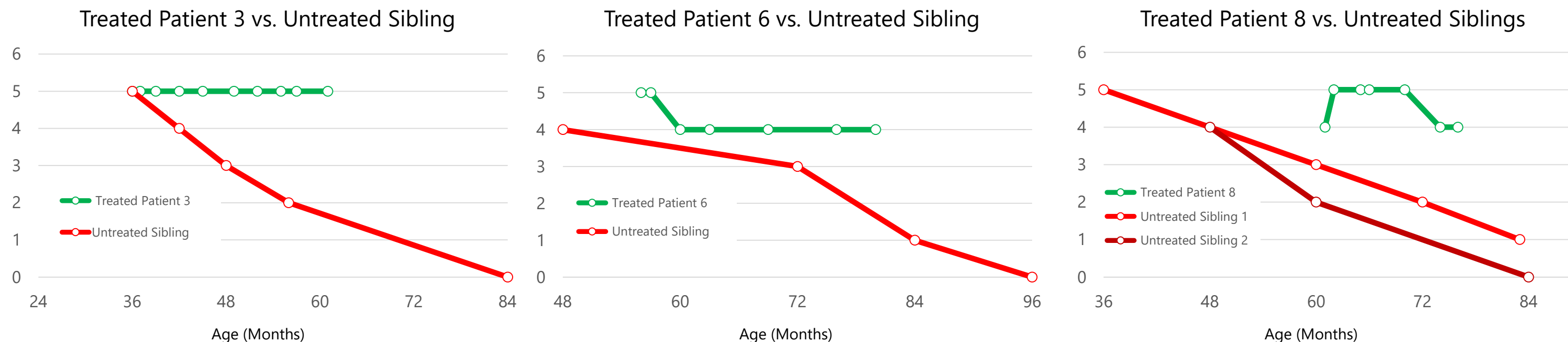
Separate Motor and Language Scores are Consistent with the Respective Combined Score.

Source: Data on file

CLN6 Clinical Efficacy Data: Sibling Comparisons (Natural History)

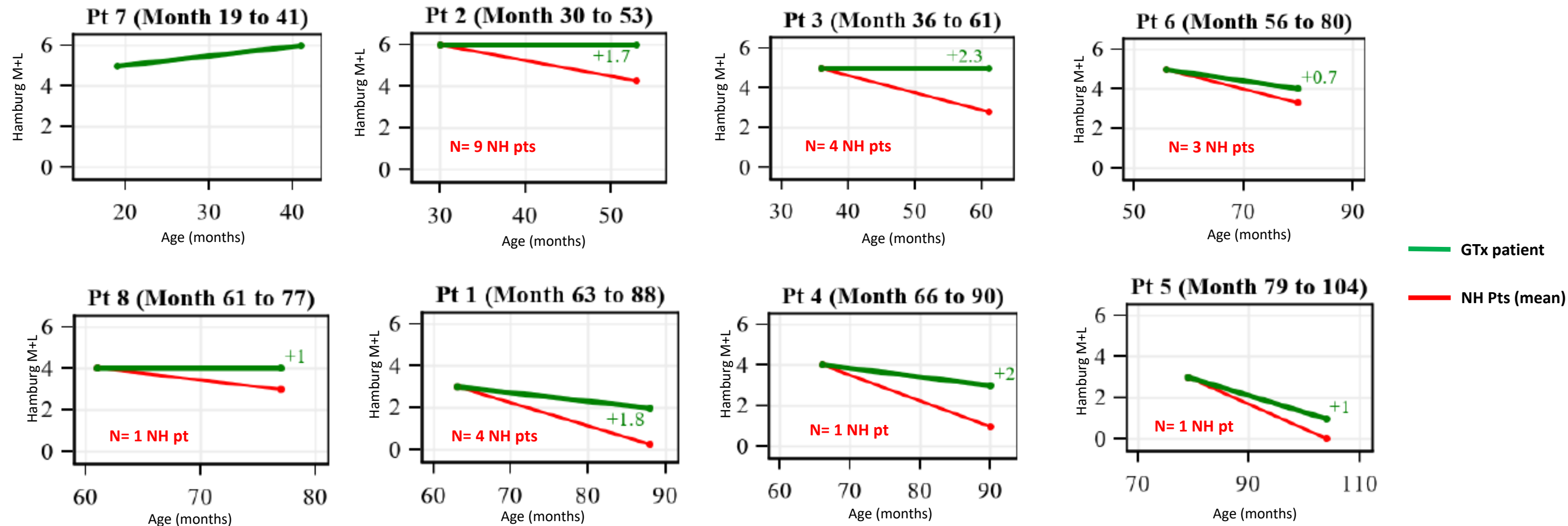
Treated patients demonstrated stabilization relative to untreated siblings in the natural history data set who experienced substantial declines in motor and language ability

Treated AAV-CLN6 Patients vs Natural History Sibling with CLN6 (Hamburg Score: Motor + Language over time)



CLN6 Clinical Efficacy Data: Natural History Matched Comparisons

New analysis of treated patients demonstrate improvement compared to natural history patients matched for age and baseline Hamburg M+L score*

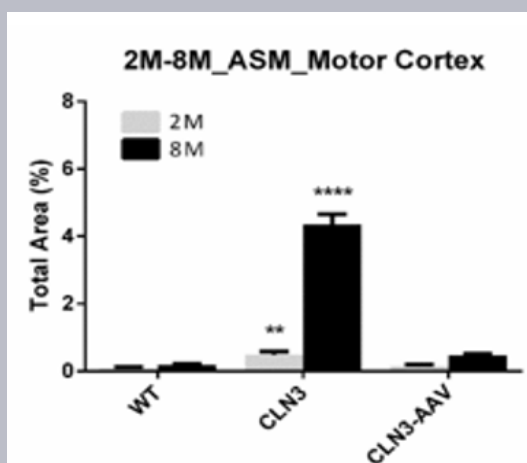


*Matched for age and exact baseline Hamburg score. No current match (for age and exact M+L score at baseline) for youngest patient (pt. 7)

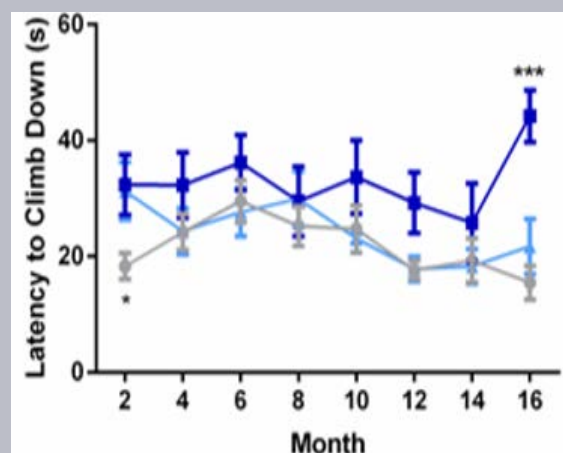
CLN3 Batten Disease: Preclinical and Clinical Summary

Amicus' second clinical stage gene therapy in CLN3 Batten disease has successfully completed dosing in three children in Cohort 1 (low dose) with dosing of additional Cohort 2 (high dose) patients in 2H 2019

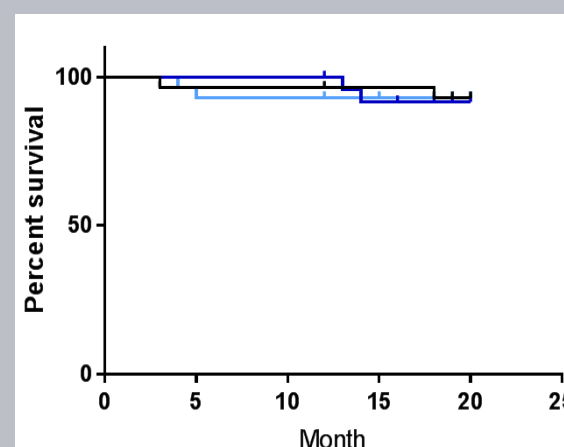
Preclinical Data in KO Mice



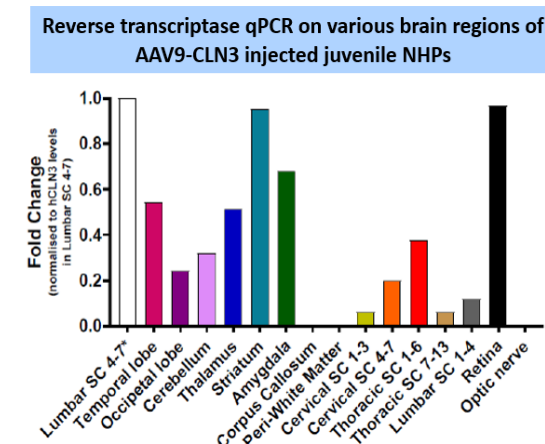
Reduction of storage material in mouse model



Improvement of motor function and cognitive behavior in mouse model



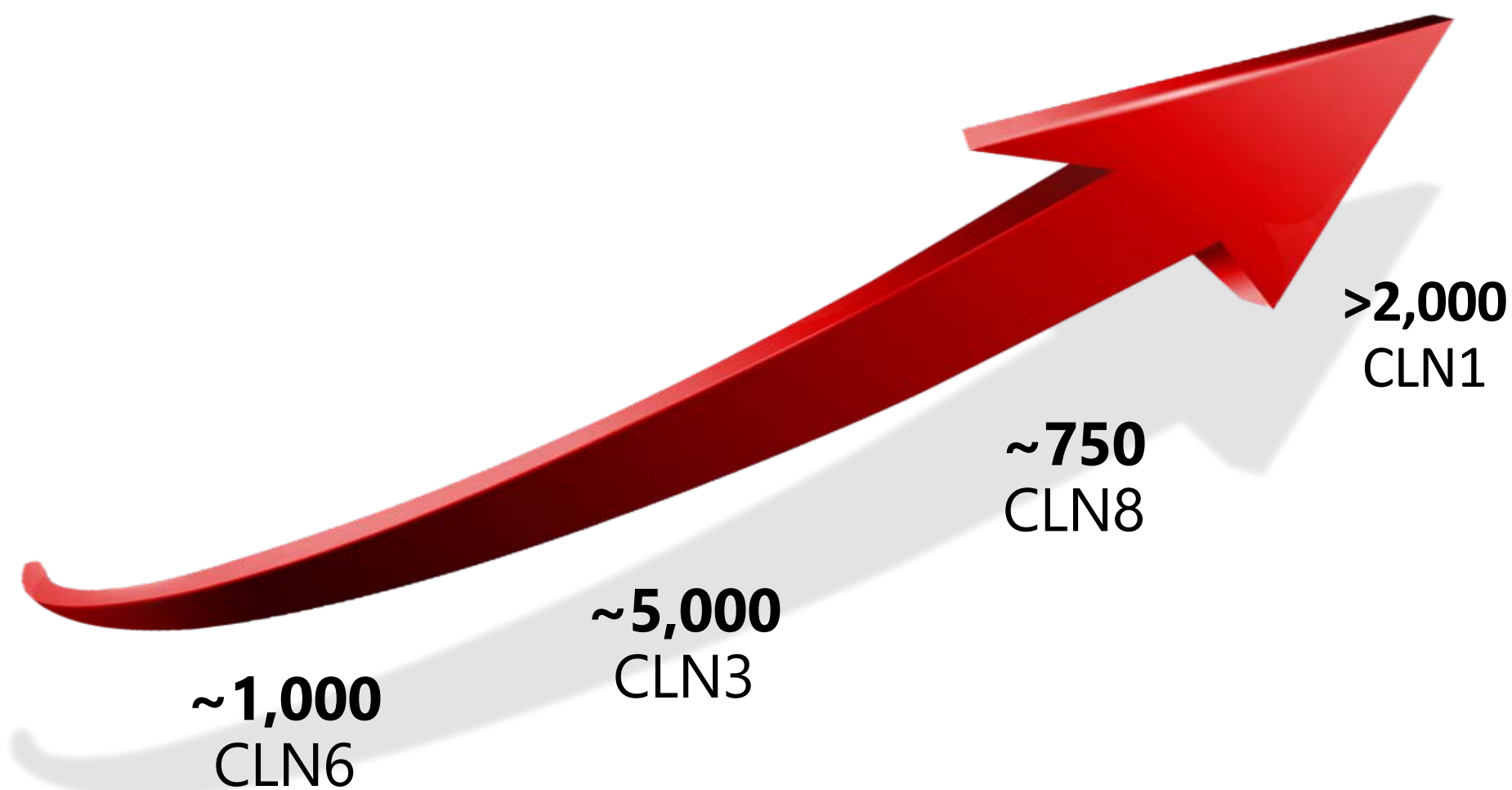
Comparable survival in mouse model



Widespread gene expression in brain of NHPs

Batten Disease Franchise

CLN6 results validate the broad potential of the intrathecal AAV platform to build a valuable and significant franchise to save thousands of children suffering from multiple types of Batten diseases with potential for \$1B+ in recurring peak revenue



Combines Amicus and Penn Expertise Across Lysosomal and Rare Diseases

An R&D platform with rights to 50+ rare diseases



**Protein
Engineering &
Glycobiology
Expertise**

**Clinical and
Regulatory
Expertise**

**Global Commercial
Infrastructure**

**Next-
Generation
Gene Therapy
Platform**

**Team of 200+
scientists bringing
expertise and
experience in:**

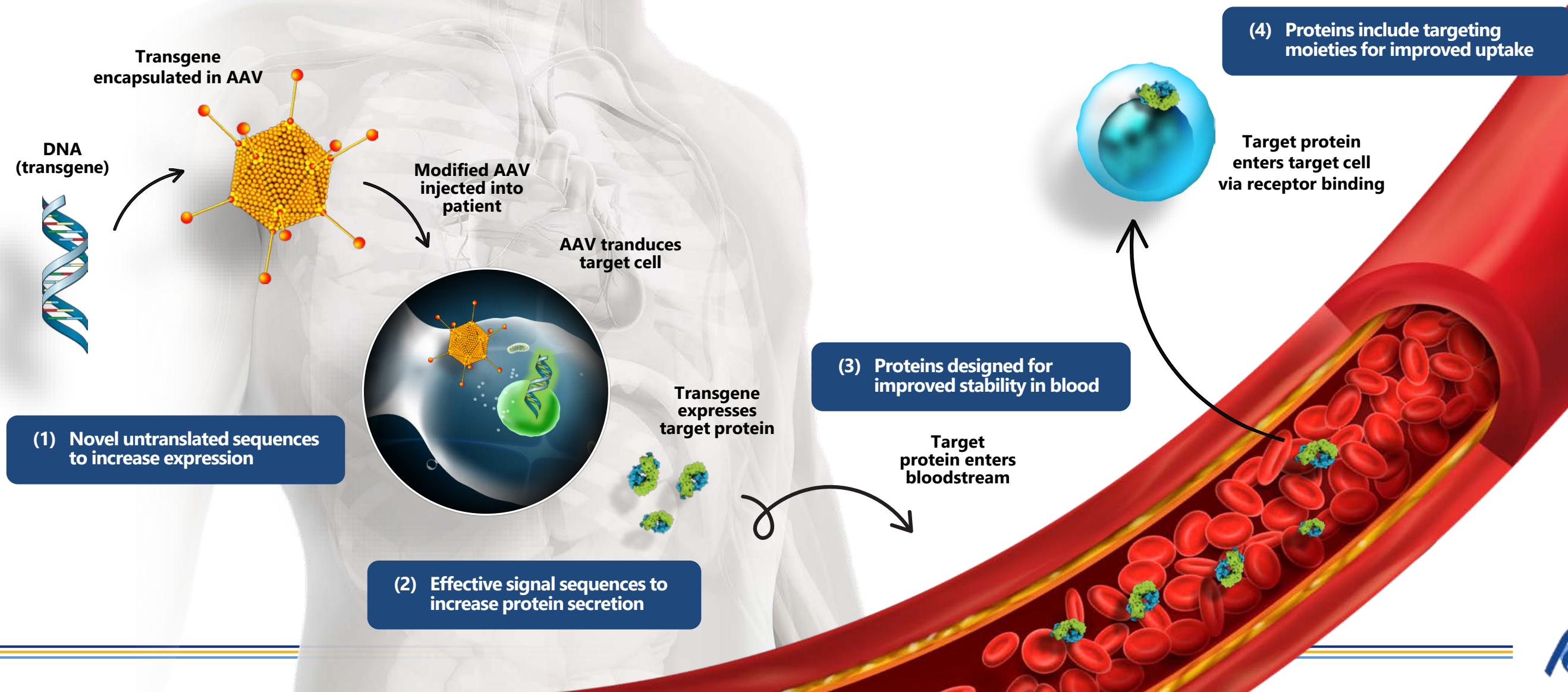
Vectors, Tropisms,
Capsids
Safety
Dosing,
Immunology
Manufacturing,
Scalability



Driving 1-2 new INDs every year starting in 2021

Amicus Approach: Engineered Transgenes for Optimal Cross-Correction

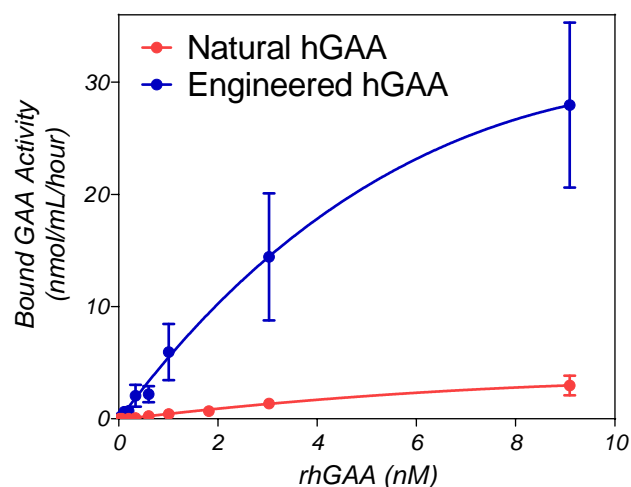
Amicus' unique technologies for protein engineering in Gene Therapy represent a new major platform technology and a groundbreaking advancement in the field



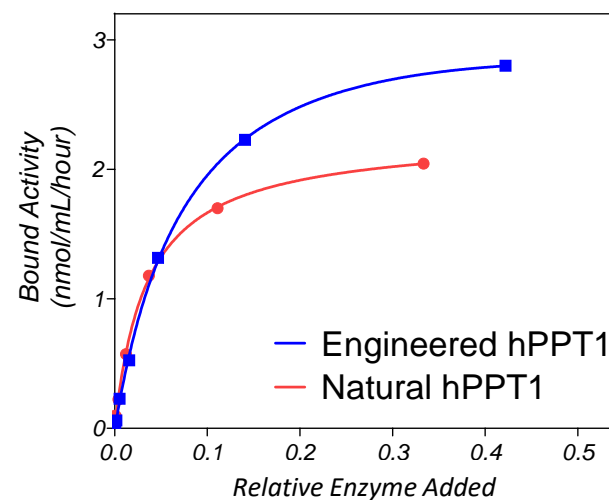
Protein Engineering Platform Has Potential To Be Broadly Applicable to Gene Therapies For Majority of LSDs

Amicus has repeatedly validated the protein engineering platform approach in multiple indications to design transgenes with improved cellular uptake

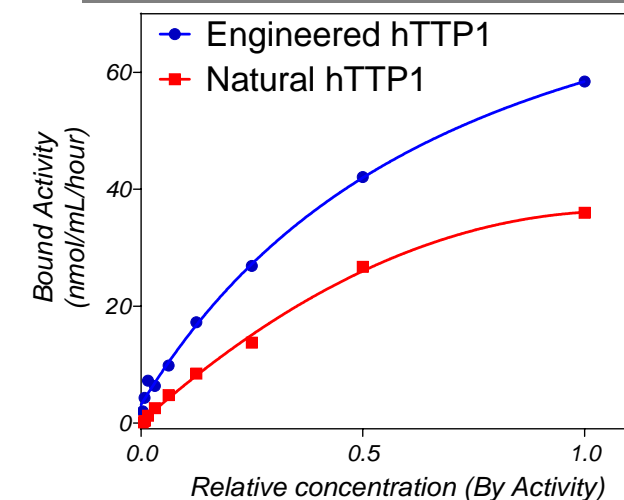
Pompe - GAA



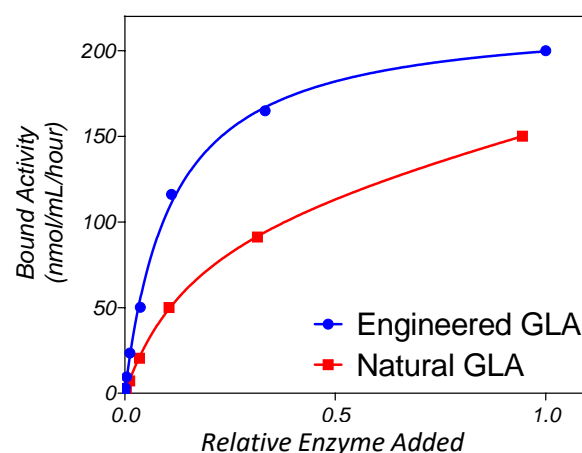
CLN1 – PPT1



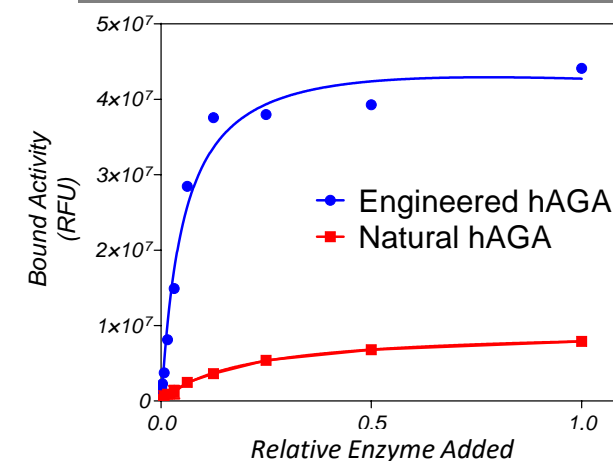
CLN2 – TTP1



Fabry – GLA



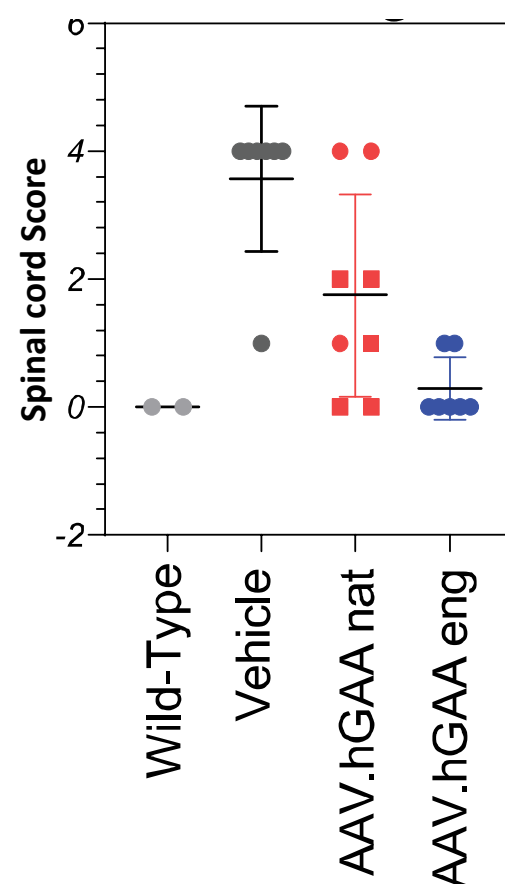
AGU – AGA



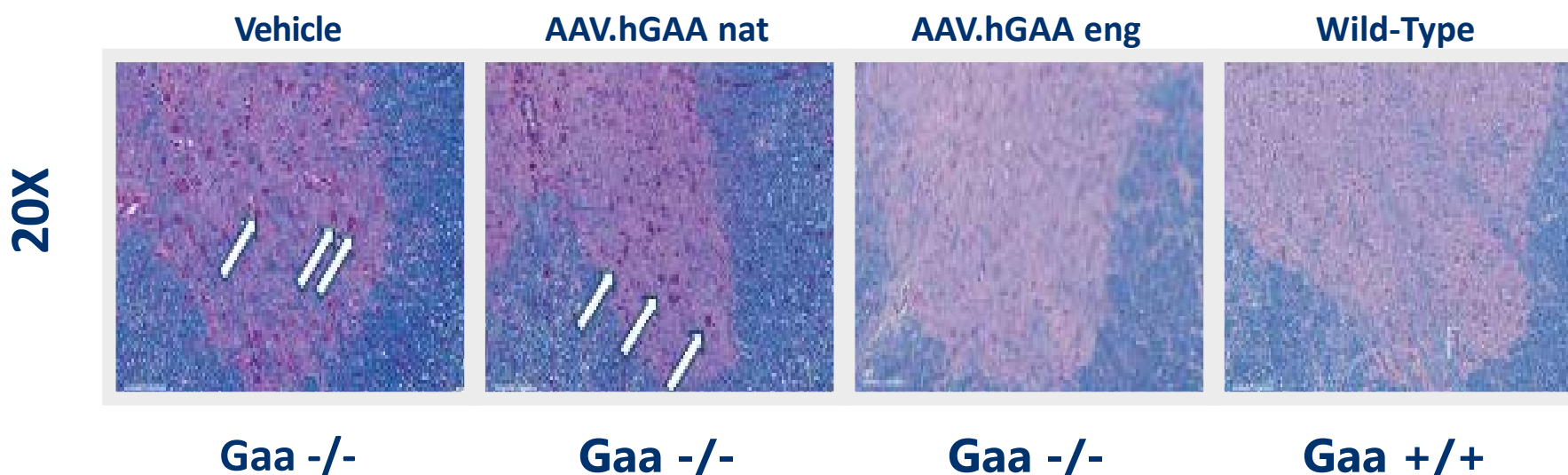
Initial Preclinical Pompe Gene Therapy Results: CNS

Only the AAV with the Amicus engineered hGAA transgene was able to significantly impact cell pathology and glycogen reduction in the CNS

Spinal Cord: Histopath



Spinal Cord: Glycogen PAS

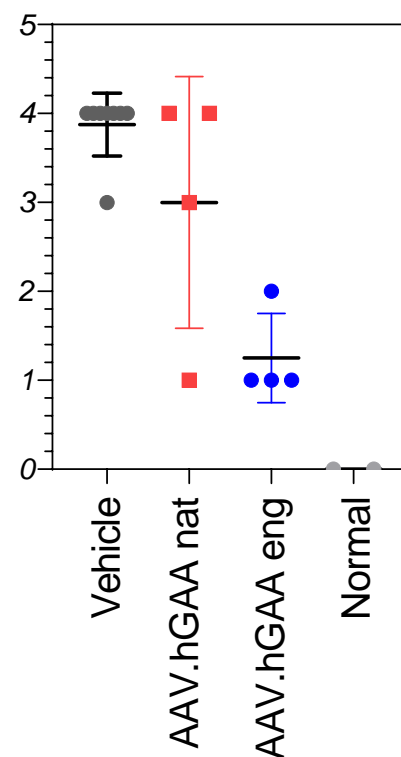


▪ Similar results observed in brain

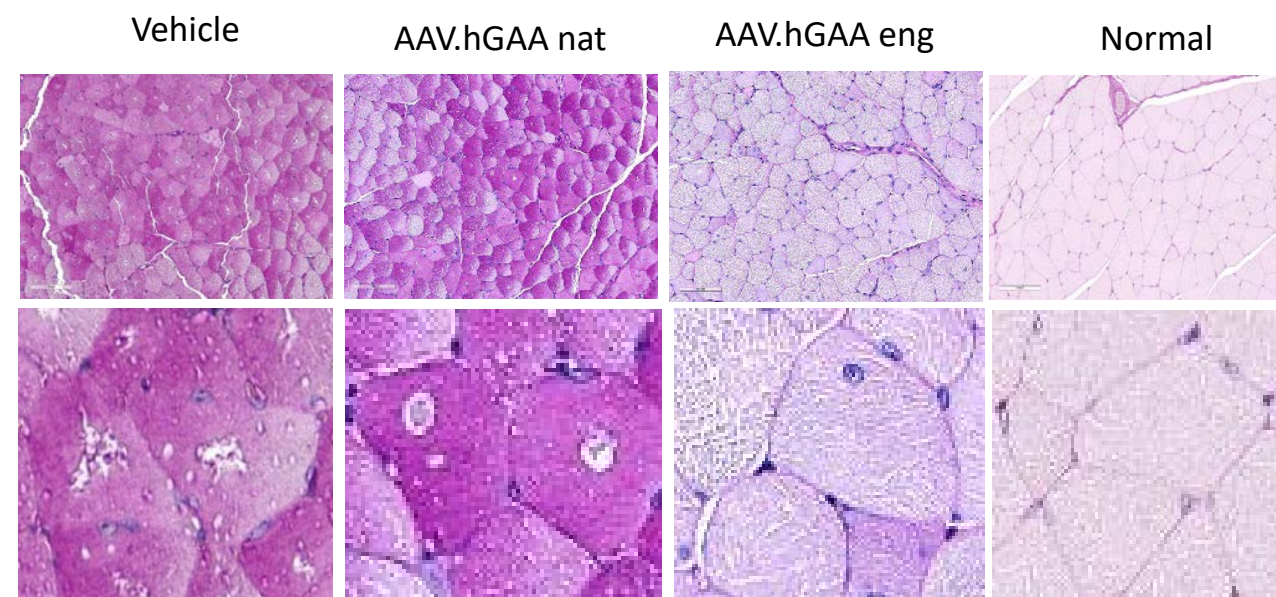
New Pompe Gene Therapy Low Dose Preclinical Data

Results from the low dose (2.5×10^{12} gc/kg) study of engineered AAV-hGAA also showed improved cell pathology and glycogen reduction of the engineered construct versus natural GAA

Quadriceps: Histopath Scoring



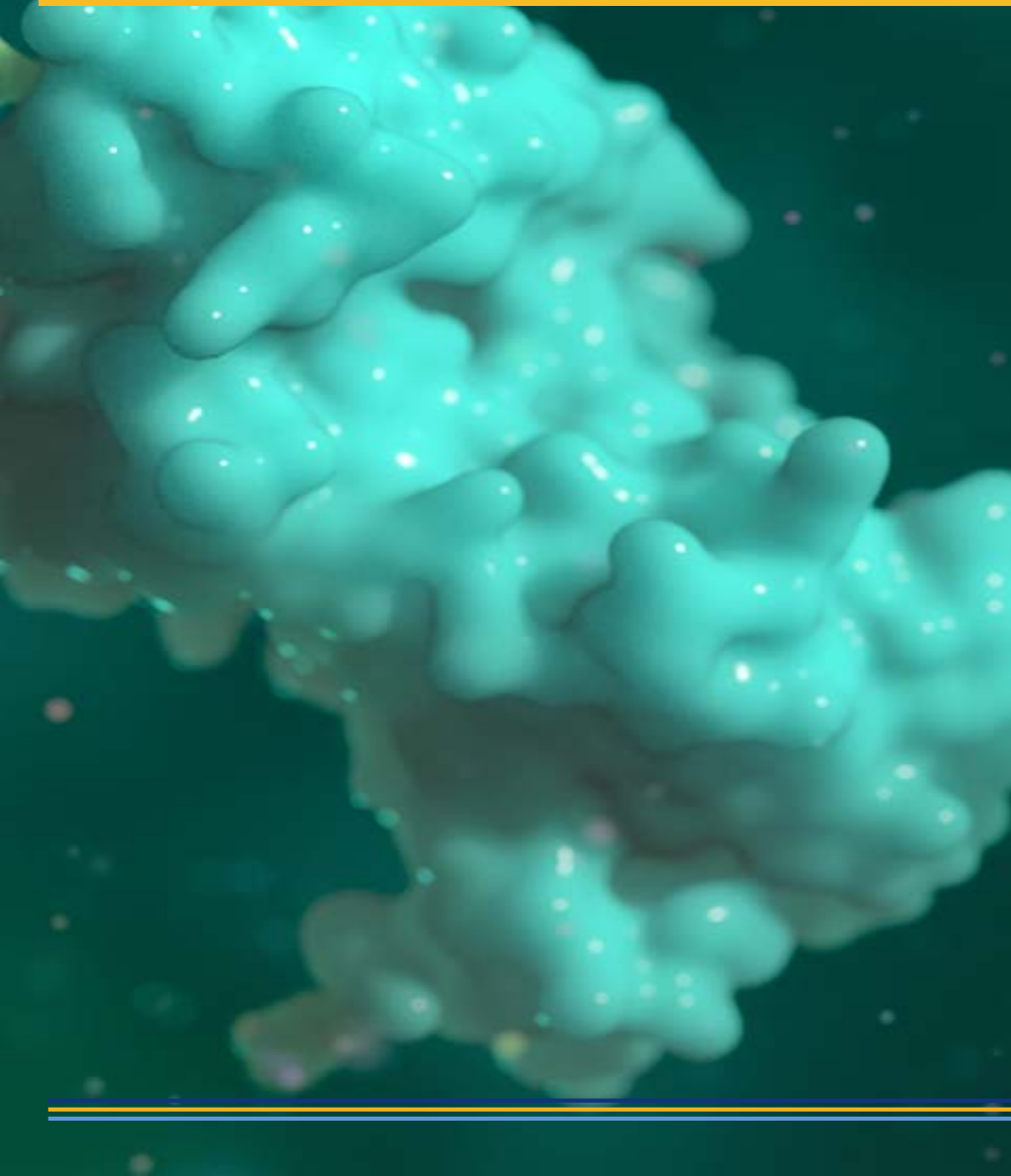
Quadriceps: Glycogen PAS



- Similar design as high dose preclinical study

Pompe Gene Therapy Summary and Next Steps

Initial Pompe preclinical gene therapy data demonstrate differentiated profile and clear pathway toward the clinic



- Clinical candidate selected
- Toxicology batch manufacturing and GMP tech transfer to Paragon underway
- IND enabling toxicology work in progress
- Potential to enter clinic in 1H2021

GTx Manufacturing Strategy for Initial Penn Programs at Catalent

Catalent to leverage Penn's AAV manufacturing expertise and Amicus experience in complex biologics manufacturing and quality control as a competitive advantage.

Catalent
BIOLOGICS

Technologies and capabilities related to Penn collaboration are being transferred to and developed at Catalent

PARAGON
BIOSERVICES

Initial focus on Amicus Pompe AAV gene therapy program

 **Penn**

Penn is supporting technology transfer of existing manufacturing process to Catalent

 **aldevron**

Long-term supply agreements established for research and GMP quality plasmids

 **Amicus**
Therapeutics®
GENE
THERAPY

Gene Therapy: Updates & Key Takeaways



Portfolio of Gene Therapy Programs and Technologies Provides Foundation for Future

- CLN6 Phase 1/2 interim data shows profound impact with potential to become first ever approved gene therapy for fatal brain disease in children
- Plan to report initial data for patients enrolled in CLN3 Phase 1/2 study in 2H'20
- Orphan drug designations granted in U.S. and EU for intrathecal AAV gene therapies for CLN6 and CLN3 Batten disease.
- Pompe gene therapy moving into IND-enabling studies
- Penn Collaboration is R&D engine, with rights to 50+ diseases
- 8 preclinical gene therapies in development



Financial & Operational Strategy

“We are business led and science driven”
- Amicus Belief Statement

Financial Outlook: Key Takeaways

- Company fully funded through major milestones in portfolio and continued global growth
- Cumulative Galafold projected revenue of \$1B+ in 2020-2022 offsets significant majority of company spend/investments
- Extended cash flow runway through OpEx savings, CapEx phasing, program prioritization and increased Galafold revenue projections
- No material business development planned or needed in next several years
- Only modest additional capital required in the outer years to extend runway into profitability with multiple non-equity sources available as/when needed

Financial Summary & Guidance

Strong Balance Sheet with \$450M+ Cash – Cash Runway Well into 2022

FINANCIAL POSITION

Cash	~\$450M+
Cash Runway ¹	Well Into 2022
Debt ^{1,2}	\$152.8M

CAPITALIZATION

Shares Outstanding (as of 12/31/2019)	255,417,869
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FINANCIAL GUIDANCE

FY20 Galafold Revenue Guidance	\$250M-\$260M
FY20 Non-GAAP Operating Expense Guidance	\$410M-\$420M

¹Based on existing operating plan ³Includes \$2.8 million of convertible debt and \$150 million of straight debt

Cash Runway Now to Well into 2022 (~2+ years)

Fully funded through major milestones in portfolio and continued global growth

**\$450M+
Cash
YE2019**

~2+ Years Cash Runway

**Well into
2022**

LEGEND

- ODD - Orphan Drug Designation
- RPD - Rare Pediatric Disease Designation

At Major Inflection Point: Path to Profitability

Clear strategy to build our business, advance our portfolio and achieve profitability with the following key priorities:

- Grow Galafold
- Advance AT-GAA to pivotal data, global approvals and launch
- Progress CLN6, CLN3 and Pompe gene therapies into and through the clinic
- Generate 1-2 gene therapy INDs every year starting in 2021
- Discover and develop next generation protein engineering and gene therapy technologies with Penn

Only modest additional capital required in outer years to extend runway into profitability with multiple non-equity sources available as/when needed

Our Passion for Making a Difference Unites Us

Amicus is now at a major inflection point and positioned to create significant shareholder value ahead while advancing our mission for patients.



Appendix

Non-GAAP Reconciliation

Amicus Therapeutics, Inc.
Reconciliation of Non-GAAP Financial Measures
(in thousands)

	Years Ended December 31,		
	2019	2018	2017
Total operating expenses - as reported GAAP	\$ 464,311	\$ 405,618	\$ 472,679
Research and development:			
Share-based compensation	17,575	11,740	10,328
Research and development asset acquisition expense	—	100,000	—
Selling, general and administrative:			
Share-based compensation	26,855	17,520	12,773
Loss on impairment of assets	—	—	465,427
Changes in fair value of contingent consideration payable	3,297	3,300	(234,322)
Depreciation and amortization	4,775	4,216	3,593
Total operating expense adjustments to reported GAAP	52,502	136,776	257,799
Total operating expenses - as adjusted	\$ 411,809	\$ 268,842	\$ 214,880