

hope through rigorous science

# **Corporate Presentation**

**November/December 2024** 









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The presentation contains forward-looking statements. Statements made or presented may include statements that are not historical facts and are considered forward-looking within the meaning of Section 27A of the Securities Act of 1933, as amended, and Section 21E of the Securities Exchange Act of 1934, as amended. Words such as "believe," "anticipate," "plan," "expect," "intend," "will," "may," "goal," "potential," "should," "could," "aim," "estimate," "predict," "continue" and similar expressions or the negative of these terms or other comparable terminology are intended to identify forward-looking statements, though not all forward-looking statements necessarily contain these identifying words. We intend these forward-looking statements to be covered by the safe harbor provisions for forward-looking statements contained in Section 27A of the Securities Act and Section 21E of the Exchange Act. These forward-looking statements, including express and implied statements relating to the FDA approval of Attruby (acoramidis), the clinical, therapeutic and market potential of our clinical development programs and our pipeline, our speed of creating new and meaningful drugs and related impact on patients, the efficiency of our engine to rapidly and efficiently deliver medicines, our value creation potential for patients, the potential market sizes and opportunities, the safety, efficacy and mechanisms of our newly FDA-approved Attruby (accoramidis) and other later-stage products including infigratinib and encaleret, our financial position, the potency and safety of our product candidates, the potential benefits of our product candidates, the potential for greater patient access to medications, the affordability and availability of insurance coverage of our medications, and the timing and expectations regarding results of our various clinical trials, reflect our current views about our plans, intentions, expectations and strategies, which are based on the information currently available to us and on assumptions we have made. Such statements reflect the current views of the Company with respect to future events and are subject to known and unknown risks, including business, regulatory, economic and competitive risks, uncertainties, contingencies and assumptions about the Company, including, without limitation, risks inherent in developing therapeutic products, and those risks and uncertainties described under the heading "Risk Factors" in the Company's most recent Annual Report on Form 10-K filed with the U.S. Securities and Exchange Commission ("SEC") and in subsequent filings made by the Company with the SEC, which are available on the SEC's website at www.sec.gov. In light of these risks and uncertainties, many of which are beyond the Company's control, the events or circumstances referred to in the forward-looking statements, express or implied, may not occur. The actual results may vary from the anticipated results and the variations may be material. You are cautioned not to place undue reliance on these forwardlooking statements, which speak to the Company's current beliefs and expectations only as of the date of the presentation. Except as required by law, the Company disclaims any intention or responsibility for updating or revising any forward-looking statements made or presented at the presentation in the event of new information, future developments or otherwise. No representation is made as to the safety or effectiveness of the product candidates for the therapeutic use for which such product candidates are being studied.

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Indicated for the treatment of adult patients with ATTR-CM to reduce cardiovascular death and cardiovascular-related hospitalization



## Our objective function

### Patient impact...

Objective: max 
$$\int_{0}^{t} \sum_{Drugs \ i=1}^{N} \frac{\Delta QALY \ (i)}{patient} * patients \ (i)$$

as many new and meaningful drugs
that have a profound impact on as
many patients as possible

### ...through sustainable value creation

### Each project must be:

- Based on beautiful science with a high probability of technical success (POTS)
- NPV positive (driven by ROIC, g, WACC)

## BridgeBio is a new type of biopharmaceutical company

From: To:

Slow and bureaucratic decision making



Rapid and decentralized decision making

Expensive platforms with long lead times before proof-of-concept data



Assets selected to target genetic diseases at their source

High fixed costs



Variablized and flexible costs

Limited sources of capital



Strategic toolkit of financing options at the levels of the portfolio and affiliate companies

Incentives at the portfolio level



Incentives at the level of each asset to <a href="preserve">preserve</a> focus at the level of biology

## The right approach: decentralized R&D, centralized infrastructure



Build "minimum viable companies" to de-risk programs as quickly and efficiently as possible



Build **central infrastructure** for functions with economies of scale, such as commercial



Leverage hyper-experienced R&D practitioners who are focused on the science of each individual program





Leverage seasoned company builders and centralized capital allocators to take the best possible shots on goal

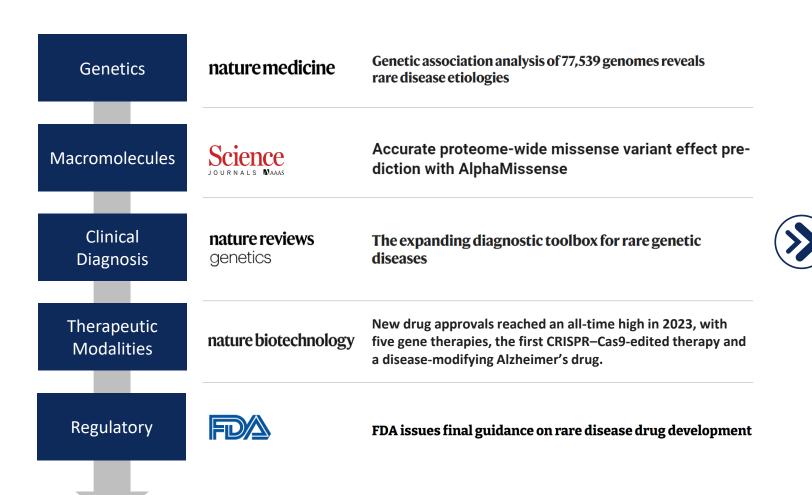


Provide investors with increased choice in where to participate in our ecosystem

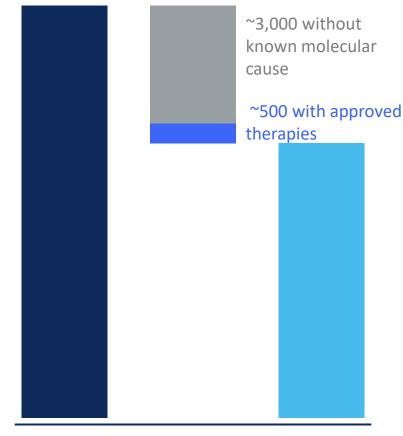


Provide investors with a de-risked portfolio of assets; enable access to low-cost debt

## The right space: capitalizing on a scientific revolution to treat a massive unmet need for genetic diseases



There are **10,000+ counted rare diseases** affecting **450 million+ people globally** 



This leaves **hundreds of millions of people** across **6,500 diseases** with known molecular cause who are **anxiously waiting** for therapies

## Our leadership team has world-renowned drug hunters and operators



Neil Kumar, PhD Founder and Chief Executive Officer







Thomas Trimarchi, PhD President & **Chief Operating Officer** 

Goldman Sachs

REGENERON



Brian Stephenson, PhD, CFA Chief Financial Officer





Uma Sinha, PhD Chief Scientific Officer







Robert Zamboni, PhD Chemistry





Jonathan Fox, MD, PhD Chief Medical Officer, Cardiorenal

MyoKardia AstraZeneca



**Ananth Sridhar** Chief Operating Officer, Cardiorenal

REGENERON Genentech



**Christine Siu** Chief Executive Officer, Muscular Dystrophy







**Justin To** Chief Executive Officer, Skeletal Dysplasias



McKinsey & Company



Eric David, MD, JD Chief Executive Officer, Gene Therapy organovo McKinsey & Company



Charles Homcy, MD Founder and Chairman of **Pharmaceuticals** 







Frank McCormick, PhD Founder and Chairman of Oncology







Richard Scheller, PhD Chairman of R&D





Len Post, PhD Advisor





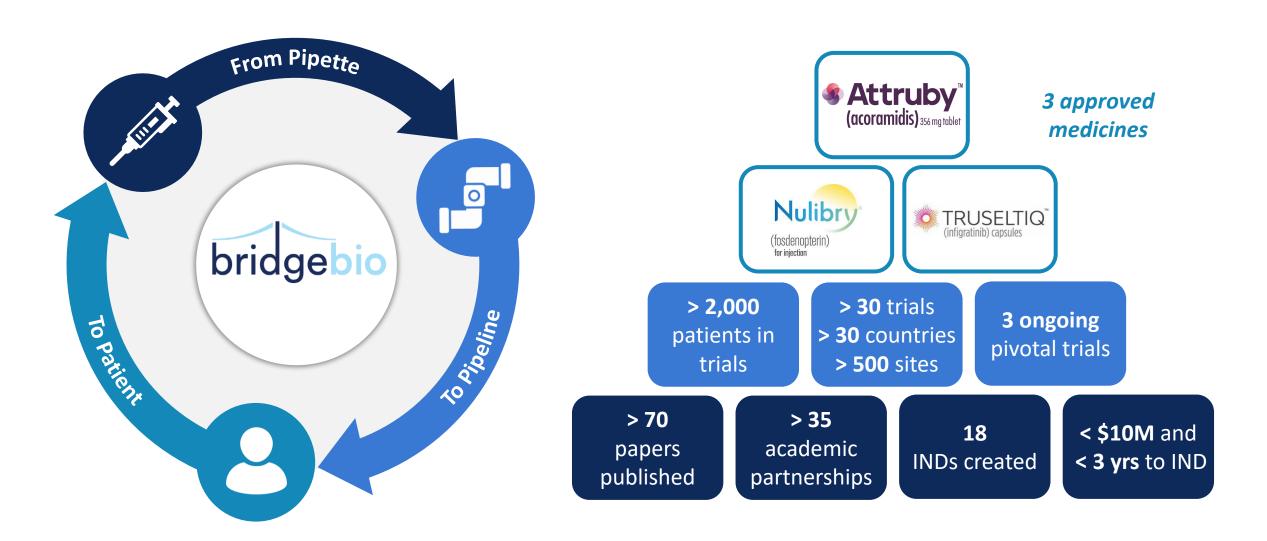


Phil Reilly, MD, JD Advisor





## We have built a sustainable, high velocity engine to deliver medicines



## A pipeline of products that sing across the BridgeBio ecosystem

Program	Indication	Pre-clinical	Phase 1	Phase	e 2 Pl	nase 3	Approved	Patients (US + EU)	Market Opportunity
Attruby (acoramidis)	Transthyretin Amyloidosis (ATTR-CM)						$\checkmark$	500,000+	\$15-\$20B
Nulibry (fosdenopterin)	Molybdenum Cofactor Deficiency (MoCD) Type A						$\checkmark$	<100	Partnered
Truseltiq (infigratinib)	Cholangiocarcinoma						$\checkmark$	37,000	Partnered
Infigratinib	Achondroplasia (ACH)							55,000	\$2B+
(low-dose)	Hypochondroplasia (HCH)							55,000	\$2B+
Encoloret	Autosomal Dominant Hypocalcemia Type 1 (ADH1)							20-25,000	\$1B+
Encaleret	Post-Surgical Hypoparathyroidism (PSH)							200,000	\$1B+
BBP-418	Limb-Girdle Muscular Dystrophy Type 2I/R9 (LGMD2I/R9)							7,000	\$1B+
BBP-812	Canavan Disease				Phase 1/2	Pivotal		1,000	TBD

... along with meaningful equity and collaboration with two clinical-stage sister companies, BridgeBio Oncology Therapeutics and GondolaBio

## Three Phase 3 clinical trial readouts in the next year

#### **Recent Achievements**

2025

Infigratinib



Published Phase 2 study results in the New England Journal of Medicine



Topline data from PROPEL 3 pivotal study and continued execution in hypochondroplasia

**Encaleret** 



Published Phase 2B study results in the New England Journal of Medicine



Topline data from CALIBRATE pivotal study and continued execution in pediatric ADH1 and post-surgical hypoparathyroidism

**BBP-418** 



Completed enrollment of Phase 3 FORTIFY study in patients with LGMD2I/R9



Topline data from FORTIFY study interim analysis, supporting opportunity to pursue accelerated approval in US

## Attruby Now Approved





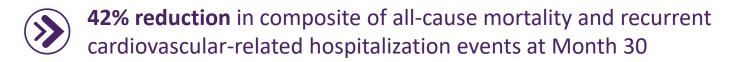


## **Attruby is now FDA Approved – Key Highlights**



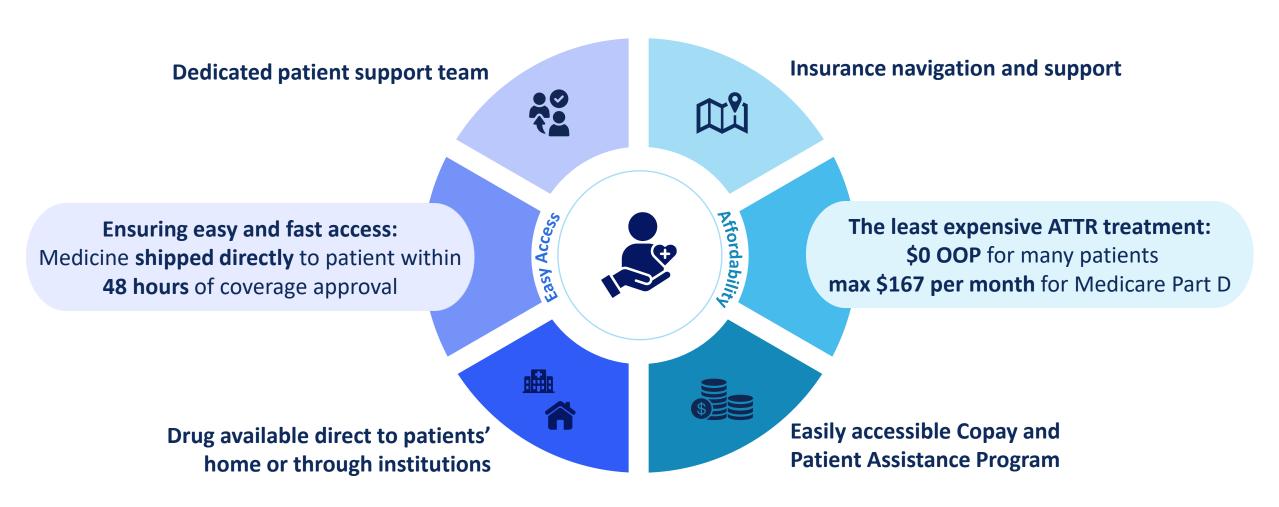




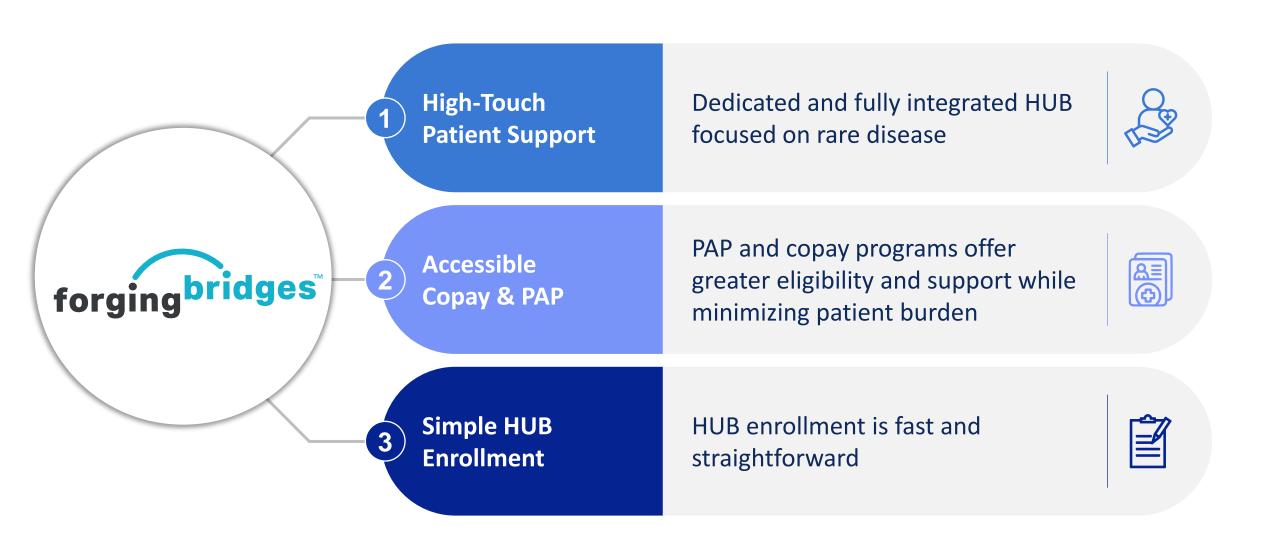


**50% reduction** in the cumulative frequency of cardiovascular-related hospitalization events at Month 30

# Our responsibility to patients spans beyond R&D – we are committed to ensuring the best access and affordability of any ATTR-CM medicine



## Our support programs are laser focused on providing access to patients



# Our commercial infrastructure is deployed across the US to ensure full provider coverage

#### **ATTR-CM Prescriber Market**



8K+ prescribing HCPs, ~100 COEs hold significant influence with prescribers & communicate overall product value

## **BridgeBio Coverage**



Dedicated commercial & US-field-based team with singular focus on serving patients with ATTR-CM



Salesforce is built to adequately cover every HCP and institution prescribing in ATTR-CM

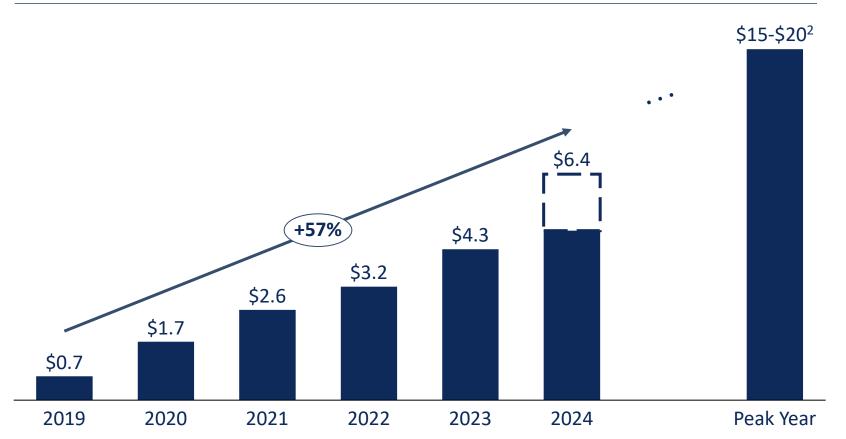


Robust segmentation to maximize effectiveness

## ATTR-CM is a multi-billion-dollar market primed for continued expansion

#### Global annual ATTR market sales<sup>1</sup>

\$B



#### **Drivers of market growth include:**

- Increased global adoption of non-invasive diagnostic tools
- Established international market familiarity with oral TTR stabilizers
- Tailwinds from the Inflation Reduction
   Act anticipated to reduce patient outof-pocket expenditure in US
- Durable market growth with tafamidis polymorph patent protection through 2035<sup>3</sup>

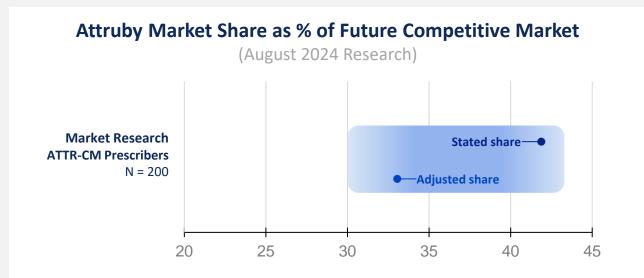
<sup>&</sup>lt;sup>1</sup>ATTR market includes all approved drugs for ATTR-PN and ATTR-CM. 2024 sales annualized as of Q2 2024.

<sup>&</sup>lt;sup>2</sup>Consensus estimates of \$15B-\$20B ATTR market.

<sup>&</sup>lt;sup>3</sup>Orange Book.

## New demand study shows significant Attruby market potential vs. partial stabilizer and partial knockdown

### Surveyed HCPs expect 30 - 40%+ peak market share for Attruby



- Survey of 200 HCPs with a history of ATTR-CM prescribing in Rx data
- Included competitive profiles of stabilizer and knockdown products
- Conducted by third-party consulting firm in August 2024, post competitive data release

### **HCP** sentiment towards Attruby is positive

Acoramidis showed dramatic reduction in cardiovascular hospitalizations, and improvement in patient QoL."

**HCP - Northeast** 

There is a mortality benefit and there are also quality of life benefits. It is an oral medication, so that will be well-liked by patients."

HCP - West

Very impressive treatment effect and best data to date on what happens in a contemporary ATTR-CM population."

**HCP - Central** 

## Our Attruby team has experienced industry leaders who have built and launched blockbuster drugs



**Matt Outten** Chief Commercial Officer

- Broad commercial leadership expertise with success across multiple competitive markets
- Led \$5B+ portfolio, 12 FDA approvals spanning 6 disease states and 7 indications (IMBRUVICA, Pharmacyclics)
- Commercial lead on \$21B pharma M&A deal



**Julie Everett** Chief Business Officer

- Successfully led cross-functional teams through multiple rare disease launches, including VOXZOGO and PALYNZIQ (BioMarin)
- Led commercial strategy and execution across ~\$1B portfolio
- ~Decade of strategy consulting leadership focused on launch excellence and lifecycle maximization (Trinity)



**John Whang** Chief Medical Affairs Officer

- Orchestrated multiple successful launches with pioneering therapies in competitive segments – STELARA (Janssen), REPATHA (Amgen), and CAMZYOS (BMS)
- 8+ launches as strategy consultant (McKinsey)
- Demonstrated strategic innovation (Heartline Study J&J / Apple collab) and consistently built outstanding organizations



**Ana Merz** VP. Sales

Launched IMBRUVICA (\$5B+, 12 FDA approvals, 6 disease states, 7 indications in 10 years) and **EPKINLY (3L+ DLBCL)** 



**Sean Doherty** SVP, Marketing

Broad global sales and marketing launch experience including in rare, infectious, and autoimmune diseases



**Scott Collins** SVP, Market Access

Extensive market access experience with consistent coverage across rare disease and oncology, leading large field-based access teams



**Hudson Boyer** VP, Commercial Analytics & Ops

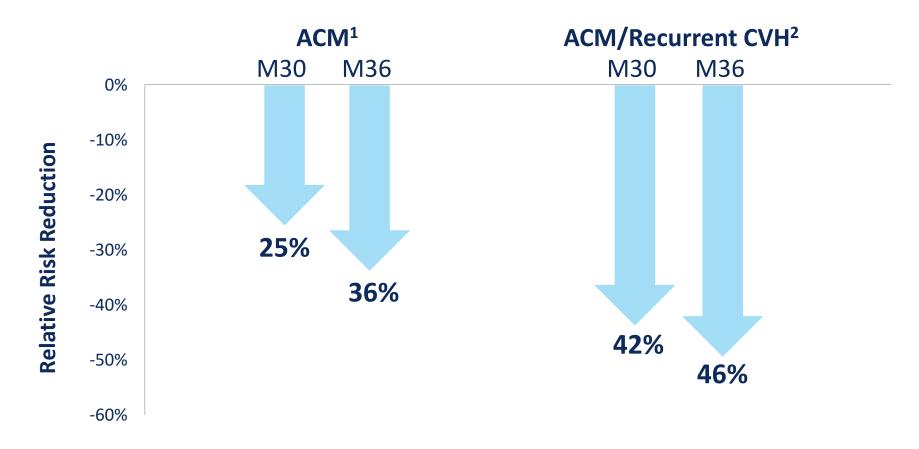
Launches in rare disease, hematology, and immunology; strategy consulting and equity analyst background



Liz Arnold **Head of Commercial Strategy** 

Multiple global launches, expertise in strategy, consulting, and marketing across rare disease, hematology, and OTC

## Recently published data from the OLE further support Attruby's statistically significant benefit on ACM and ACM/Recurrent CVH



Attruby resulted in a **statistically significant** ACM and ACM/Recurrent CVH relative risk reduction at **both Month 36 and Month 42** 

## BridgeBio is committed to advancing Attruby's scientific narrative to ensure as many appropriate patients as possible can benefit



## **ATTR**ibute

**ATTR-CM** WT and hereditary Extended Ph. 3 data disclosures at ACC, ESC-HF, ESC, HFSA, and AHA

ATTRibute-CM Ph. 3 OLE Month 42 safety and efficacy data



## **ATTR**ibute 8

**ATTR-CM** WT and hereditary

Attruby lifecycle management and further analyses of pivotal study data



**ATTR-CM Hereditary** 

Prevention study

Additional real-world evidence generation and publication plan to expand scientific share of voice

Infigratinib



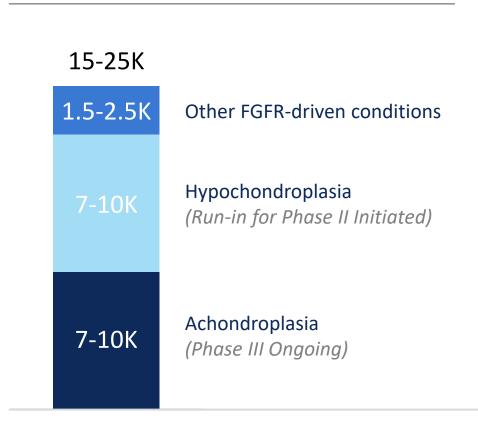
# Achondroplasia is a large, proven commercial market annualizing to ~\$750M today with substantial remaining treatable pool

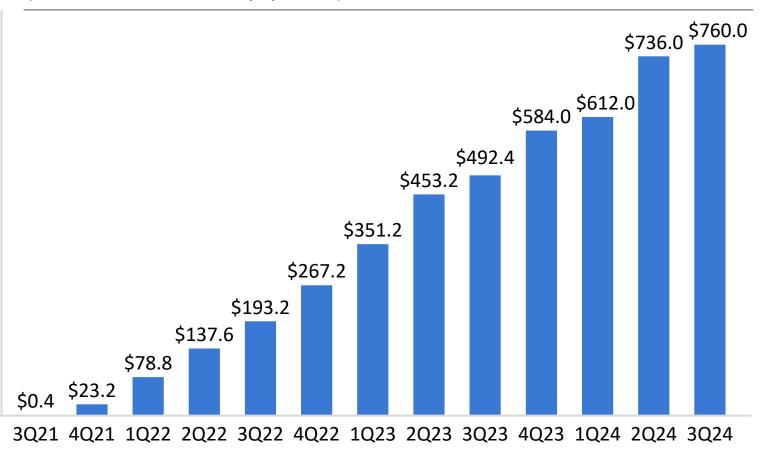
## Addressable patients by indication in US/EU<sup>1</sup>

(current population eligible for treatment)

## Annualized achondroplasia product sales<sup>2</sup>

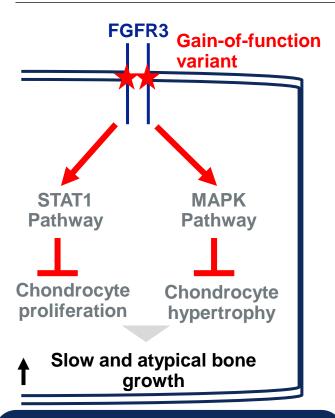
(\$M WW, annualized by quarter)



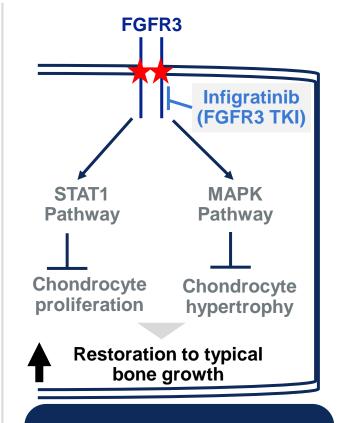


## Infigratinib is a best-in-class FGFR3 inhibitor that targets achondroplasia and hypochondroplasia at their source

#### Mechanism



FGFR3 acts as a "molecular brake" on chondrocyte proliferation and hypertrophy; in ACH or HCH, this brake is **stuck** due to gain-of-function mutations resulting in shortened bones



Infigratinib "releases" the brake, potentially resuming normal chondrocyte function, allowing for restoration of bone growth

#### **Design Principles**



## **Maximize efficacy by targeting** disease at the source

For all the manifestations of ACH and HCH, not just height, which matter for families and physicians



## **Demonstrate safety with low dosing**

Avoiding hypotension & injection site reactions with no hyperphosphatemia, ocular effects or VEGFR3 off-target effects

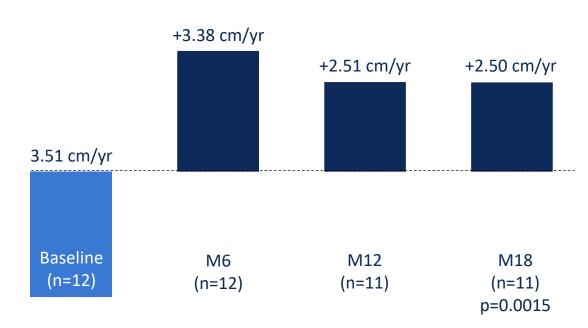


## **Avoid injections and provide** an oral option

For children and families, to reduce burden and pain of treatment

# At 18 months, infigratinib has shown persistent improvement in AHV and body segmentation, along with a favorable safety profile

Mean change from baseline in annualized height velocity (AHV)

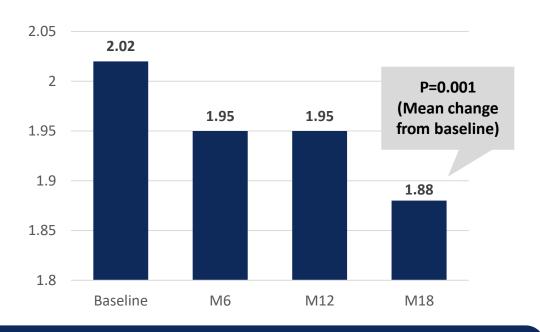


At each timepoint, infigratinib change from baseline AHV is higher than that reported by any other treatment option

At the highest dose, there were no SAEs, most TEAEs were of grade 1 severity, and none were assessed as related to study drug

#### Upper body to lower body segment ratio

(Lower value means improved proportionality)



Infigratinib shows statistically significant proportionality improvements after 18 months

This has potential for a meaningful effect on body proportionality, and if maintained, can be associated with functionality

## Encaleret



## ADH1 is a serious and rare genetic condition for which there are no indicated therapies and diagnosis rates are low

## 12K ADH1 Carriers ~12K carriers of ADH1 causing variants in the US based on 4 population databases<sup>1-2</sup> 9K Treatable 73% of patients with ADH1 are symptomatic<sup>3</sup> **3K – 5K Currently Addressable** 40-60% of symptomatic ADH1 patients are diagnosed today, anticipate increase

#### An analogous ADH1 market is XLH

	XLH	ADH1		
Prevalence (US)	12K <sup>5</sup>	12K		
Disease burden	Hypophosphatemia	Acute hypocalcemia risk, long-term hypercalciuria risk		
Standard of care	Vitamin D, daily phosphate	Vitamin D, daily calcium		
Registrational endpoint	Serum phosphate	Blood and urine calcium		
Projected peak year sales	\$2B+ <sup>6</sup>	\$1B+		

## Population prevalence estimates in literature<sup>1-2</sup>



# Encaleret is a first-in-class, disease-modifying therapy that targets the underlying disease mechanism of ADH1

#### Mechanism

Normal CaSR senses and regulates serum Ca levels to maintain homeostasis

#### **Encaleret** is a **CaSR antagonist**

designed to decrease the sensitivity of CaSRs to extracellular calcium



ADH1 CaSR is overly sensitive to extracellular calcium



<u>restored</u> in the presence of encaleret

Overactive CaSR causes dysregulation of calcium homeostasis, encaleret has the potential to normalize PTH, serum Ca, and urine Ca levels



Restore
PTH
to the normal
range



Restore serum calcium to the normal range



<u>Decrease</u> urine calcium to the normal range

#### **Design Principles**



## First and only investigational treatment directly targeting ADH1 at its source

Potential to restore physiologic mineral homeostasis that is disrupted by CaSR oversensitivity



## **Address common symptomology**

Designed to normalize PTH, serum Ca, and urine Ca levels, potentially resolving key symptoms

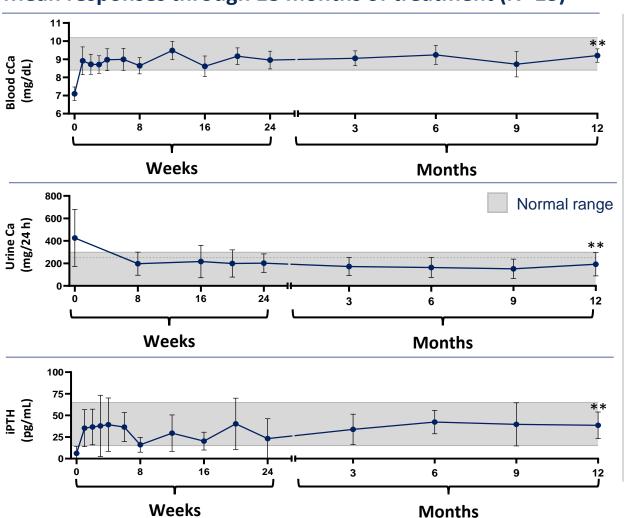


## **Convenient oral dosing**

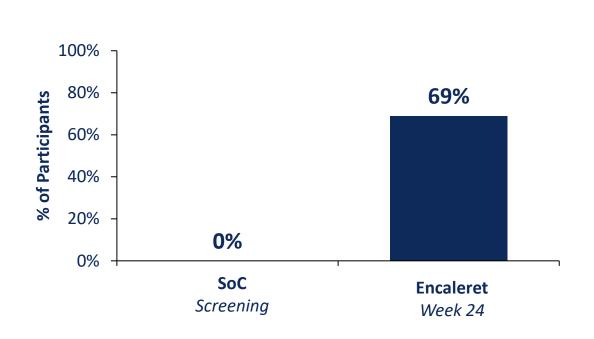
First targeted treatment for ADH1 in a convenient form for patients and families

# Phase 2B results demonstrated rapid and sustained normalization of serum Ca, urine Ca, and serum PTH, without need for dose escalation

Mean responses through 18 months of treatment (N=13)



Participants with blood Ca and urine Ca in the target range



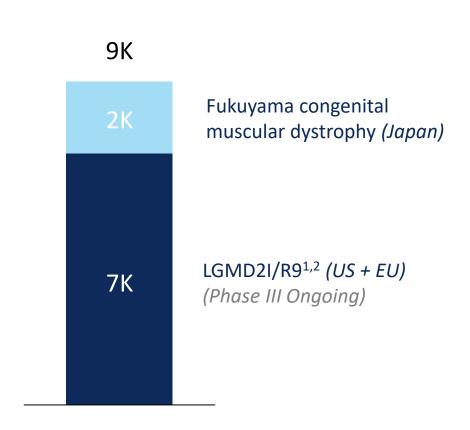
Encaleret normalized mean blood Ca, PTH, and urine Ca in participants with ADH1 over an 18-month period representing a meaningful improvement over SoC

**BBP-418** 



## LGMD2I/R9 is a progressive neuromuscular disease with high unmet need, representing a >\$1B market opportunity in the US and EU

#### Addressable patients by indication



#### **Unmet need**

- LGMD2I/R9 is an inherited neuromuscular disorder characterized by lower-limb weakness and loss of ambulation as well as respiratory decline and cardiac dysfunction
- No approved disease modifying agents for LGMD2I/R9
- Current standard of care is aimed at symptom management and includes physical therapy, steroids, and pain management
- Standard of care does not prevent continuous and progressive decline in LGMD2I/R9 patients

Market opportunity \$1B+

## BBP-418 is a first-in-class, disease-modifying therapy with potential to be the first approved therapy for LGMD2I/R9

#### Mechanism



FKRP glycosylates alpha-dystroglycan (αDG), stabilizing muscle cells by binding extracellular ligands to act as a "shock absorber" for muscle fibers



Partial loss of function in FKRP results in dysfunctional, hypo-glycosylated αDG in muscle cells, increasing cell susceptibility to damage

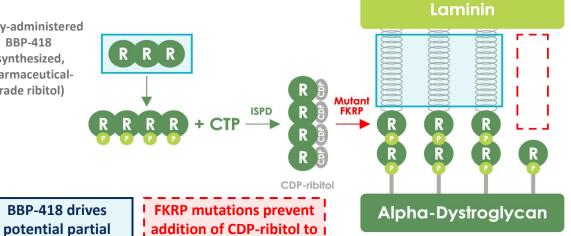


Supply supraphysiological levels of synthesized, pharmaceutical grade ribitol upstream aiming to drive residual activity of mutant FKRP enzyme and increase αDG glycosylation levels

**Orally-administered** BBP-418 (synthesized, pharmaceuticalgrade ribitol)

restoration of

glycosylation of αDG



αDG, limiting function as

a "shock absorber"

#### **Design Principles**



## Provide first disease-modifying therapy

For patients with LGMD2I/R9 and potentially applicable for other  $\alpha$ -dystroglycanopathies



## **Avoid safety concerns with FKRP modulation**

Avoid off-target effects using a synthesized version of an endogenous compound with an encouraging safety profile

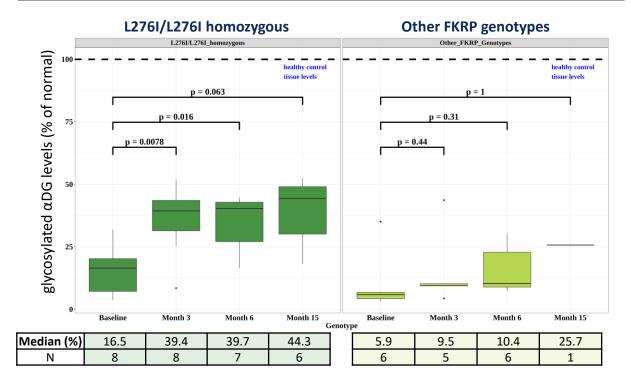


#### Convenient oral medicine

To reduce burden for patients and avoid safety concerns

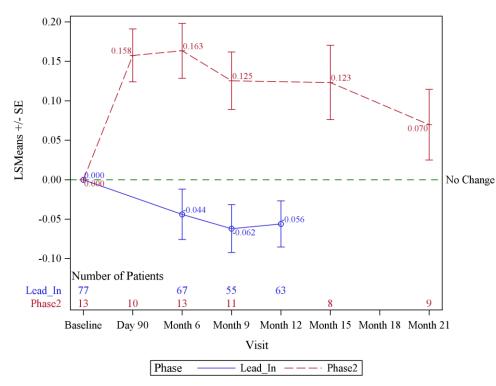
## Significant increases in glycosylated $\alpha DG$ in muscle and stabilization of ambulatory measures were observed in a Phase 2 study of BBP-418

Increase in glycosylated αDG in muscle observed post dosing with BBP-418 (median ± 95% CI)



Patient samples were interpolated to standard curve to determine % of normal glycosylation of αDG + 3 mo = Part 1, 90-day, +6 mo = Part 2, Month 3, + 15 mo = Part 3, Month 9 Median and 10–90% percentile are shown, Wilcoxon test was used to determine significance MLB-01-003 Listing 16.4.1 and 16.1.4.2

#### Change from baseline in 10-meter walk test (m/s)



Blue line denotes natural history; red line denotes on-treatment data from Ph. 2 study. Stabilization of 100-meter timed test and NSAD was also observed at 21 mo. MLB-01-001 Listing 16.2.1 and MLB-01-003 Listing 16.2.1

BBP-812



## Canavan disease is a severe, fatal, and ultra-rare neurodegenerative pediatric disease with no approved therapies

#### **Unmet need**

- Canavan is an ultra-rare neurodegenerative disease with ~1,000 patients across the US and EU
- Canavan is usually fatal within the first two decades of life, and >25% of patients die by the age of 10 years<sup>1</sup>
- Children with Canavan exhibit global and severe cognitive, motor, and language impairment, missing or regressing on most developmental milestones
- They require around the clock care they cannot hold their heads up, sit, crawl, walk, are generally unable to speak, and suffer from seizures and spasticity
- There are no therapies available for Canavan disease



<sup>1</sup>Bley A, et al. Orphanet J Rare Dis. 2021 PMID: 34011350

## BBP-812 is a first-in-class, disease-modifying therapy that targets Canavan disease directly at its source

#### Mechanism

#### **Healthy NAA NAA** metabolic pathway metabolic pathway in Canavan Disease N-acetylaspartic **▲ N-acetylaspartic** acid (NAA) acid (NAA) **Aspartoacylase** (ASPA) Acetate Acetate **Aspartate Aspartate** Healthy Demyelination of neurons Neuron

BBP-812 is an AAV9 gene therapy which directly replaces the mutated ASPA gene that causes Canavan disease

#### **Design Principles**



## **Provide first disease-modifying therapy**

Target the condition directly at the source, utilize single registrational study & biomarker for accelerated approval



## Provide therapy with known safety profile

Leverage safety profile from approved AAV9 gene therapy (Zolgensma)

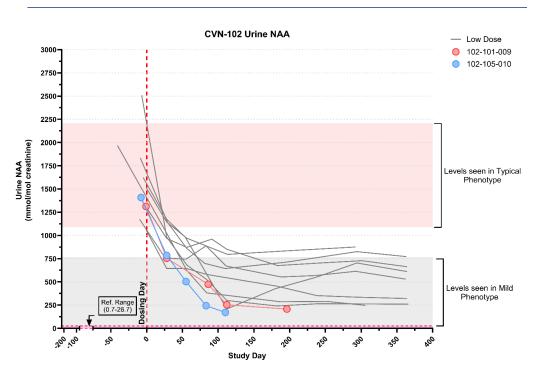


## **Avoid invasive neurosurgery**

Provide a less invasive IV treatment option to minimize burden for patients and their caregivers

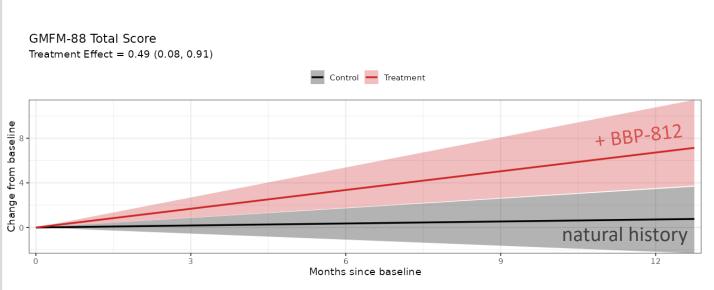
# Significant, sustained reductions in NAA levels and improvement in motor function after 12-months of BBP-812 dosing in Phase 1/2 study

### **Urine N-acetylaspartic acid (NAA) levels**



- BBP-812 reduces urine NAA from levels associated with typical Canavan disease to levels associated with mild disease
- Preliminary high-dose data suggest higher BBP-812 doses further reduce urine NAA levels

### **Gross Motor Function Measure (GMFM-88) Trajectory Analysis**



• Trajectory analysis shows clear separation in GMFM-88 Total Score between individuals dosed with BBP-812 in treatment study (CVN-102, in red) vs. individuals in the natural history study (shown in gray).

## **About Attruby and BridgeBio**

### About Attruby™ (acoramidis)

Attruby is the only near-complete (≥90%) stabilizer of Transthyretin (TTR) approved in the U.S. for the treatment of adult patients with ATTR-CM to reduce cardiovascular death and cardiovascular-related hospitalization. Attruby was generally well-tolerated. The most common side effects were mild and included diarrhea and abdominal pain that were resolved without drug discontinuation. BridgeBio offers an extensive suite of programs to help patients access our medicines. Visit Attruby.com for more information, including full Prescribing Information.

#### About BridgeBio Pharma, Inc.

BridgeBio Pharma, Inc. (BridgeBio) is a new type of biopharmaceutical company founded to discover, create, test, and deliver transformative medicines to treat patients who suffer from genetic diseases. BridgeBio's pipeline of development programs ranges from early science to advanced clinical trials. BridgeBio was founded in 2015 and its team of experienced drug discoverers, developers and innovators are committed to applying advances in genetic medicine to help patients as quickly as possible. For more information visit bridgebio.com.