# 89bio

# Powerful Science Meaningful Medicines Changing Lives

Nasdaq: ETNB

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#### **Cautionary Note Regarding Forward-Looking Statements**

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We obtained the industry, market and competitive position data used throughout this presentation from our own internal estimates and research, as well as from industry and general publications, and research, surveys and studies conducted by third parties. Internal estimates are derived from publicly available information released by industry analysts and third-party sources, our internal research and our industry experience, and are based on assumptions made by us based on such data and our knowledge of the industry and market, which we believe to be reasonable. In addition, while we believe the industry, market and competitive position data included in this presentation is reliable and based on reasonable assumptions, we have not independently verified any third-party information, and all such data involve risks and uncertainties and are subject to change based on various factors.

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## **Corporate Highlights**



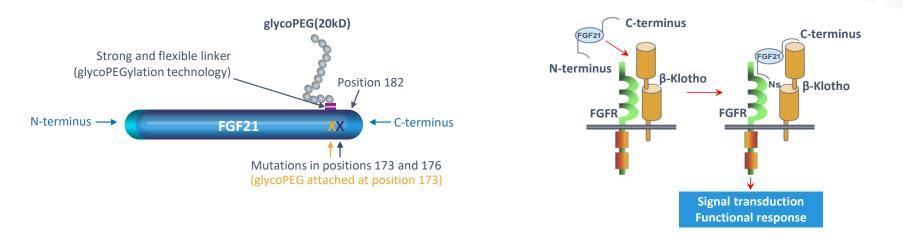
Late-stage clinical development company entering Phase 3 in NASH and SHTG

Pegozafermin (FGF21 analog) has demonstrated
 differentiation within class and category and has the potential to be transformational for patients

Strong balance sheet (\$480.9M\* in cash, cash equivalents and short-term investments)

Highly experienced team with track record of execution

# Pegozafermin is an FGF21 Analog Optimally Engineered to Balance Efficacy and Long Dosing Interval



- Proprietary glycoPEGylation technology commercially validated with approved products
- Increases half-life of native FGF21 (<2 hours) to 55-100 hours based on single ascending dose study
- Low nanomolar potency against FGF receptors 1c, 2c, 3c, similar to native FGF21
- Composition of Matter patent expiring in 2038

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# **Opportunity in NASH**

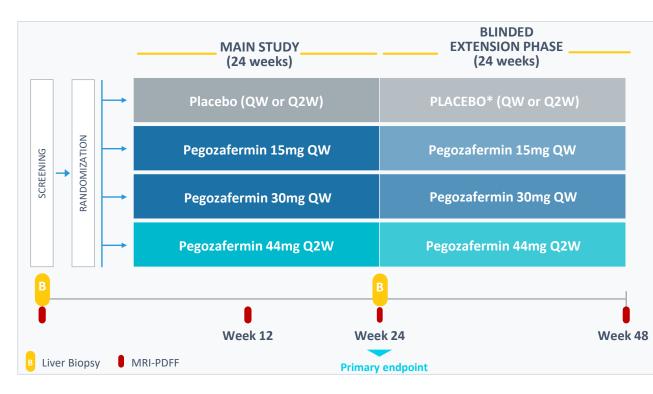




Pegozafermin – Potential to Address an Unmet Need for Effective NASH Treatments as Demonstrated in the Phase 2b ENLIVEN Trial



## ENLIVEN Trial Design Weekly (QW) and Every-Two-Week (Q2W) Dosing



#### **PRIMARY ANALYSIS POPULATION**

• F2-F3 NASH; NAS ≥4

#### **PRIMARY ENDPOINTS**

- ≥1-stage fibrosis improvement with no worsening of NASH<sup>1</sup>
- NASH resolution with no worsening of fibrosis<sup>2</sup>

#### KEY SECONDARY EFFICACY ENDPOINTS

- ≥2-point change in NAS with no worsening of fibrosis
- Non-invasive liver markers (liver fat, liver injury, fibrosis markers)

<sup>1</sup>Improvement in liver fibrosis by ≥1 stage and no worsening of steatohepatitis defined as no increase in NAS for ballooning, inflammation, or steatosis (FDA draft guidance). <sup>2</sup>Resolution of steatohepatitis is defined as absent fatty liver disease or isolated or simple steatosis without steatohepatitis and a NAS score of 0-1 for inflammation, 0 for ballooning and any value for steatosis (FDA draft guidance).

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\*Some placebo patients were re-randomized in the extension phase to receive pegozafermin.

NAS, NAFLD Activity Score; MRI-PDFF, Magnetic resonance imaging-estimated proton density fat fraction; QW: Every week; Q2W: Every 2 weeks

# Potential Challenges with Common Biopsy Reading Approaches

## Single pathologist

- Wholly dependent on idiosyncrasies of that reader
- FDA strongly encourages sponsors to use multiple readers and consensus

## Two pathologists with an adjudicator

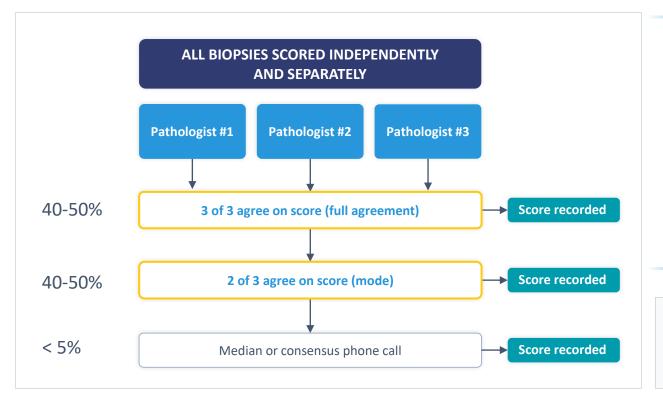
- >50% of scores will be different between the two pathologists
- The adjudicator then acts like a single pathologist to break the tie

### Two pathologists with an alignment call

- >50% of scores will be different between the two pathologists
- Alignment call introduces social pressures to reach agreement and often the senior voice prevails
  - Confirmed by pathologists involved in NASH alignment calls
- Typically the alignment call defaults to the single pathologist

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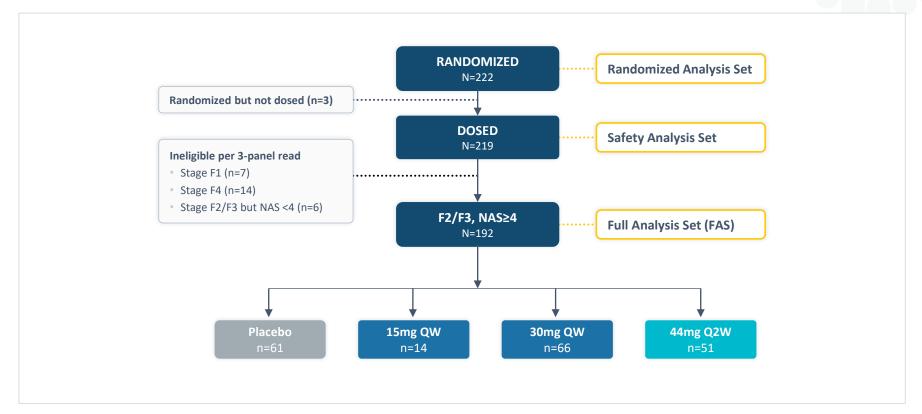
# Objective Biopsy Reading Method Designed to Identify Drug Effect



- Pathologists underwent protocol-specific harmonization training before and during trial
- Pathologists were blinded to subject, treatment and sequence
- >99% of final scores determined by algorithm, which was established a priori, rather than resolving disagreements by inter-reader discussion

Designed to reduce impact of individual reader bias and interreader variability

## Patient Disposition and Analysis Sets



Analysis Sets were prospectively defined

Completer Analysis Set = FAS subjects with biopsies at both baseline and Week 24 (n=164).

O MRI-PDFF Analysis Set = all subjects in FAS with baseline and at least one post-baseline MRI-PDFF assessment (n=181).

# Baseline Characteristics Well Balanced Across Dose Groups

Parameter Mean or %	Placebo (n=71)	<b>15mg QW</b> (n=21)	<b>30mg QW</b> (n=73)	<b>44mg Q2W</b> (n=57)	Total (n=222)
Age (years)	56	55	55	55	56
Female	55%	43%	69%	65%	61%
BMI (kg/m²)	38	38	35	36	37
Type 2 Diabetes	69%	86%	62%	61%	66%
Fibrosis Stage (% F3)	66%	43%	64%	53%	60%
NAFLD Activity Score	5.0	4.8	5.3	5.2	5.1
Liver Fat Content (MRI-PDFF)	16.7%	15.8%	16.7%	15.8%	16.4%
Liver Stiffness (VCTE, kPa)	14.1	11.2	12.5	13.2	13.0
PRO-C3 (ng/mL)	50	62	54	52	53
ALT (U/L)	50	61	60	56	56
AST (U/L)	41	48	47	42	44
HbA1c, overall population (%)	6.6	7.0	6.6	6.7	6.7
Triglycerides (mg/dL)	170	186	175	165	172

Baseline characteristics were consistent in full analysis set (n=192) and the safety set (n=222)

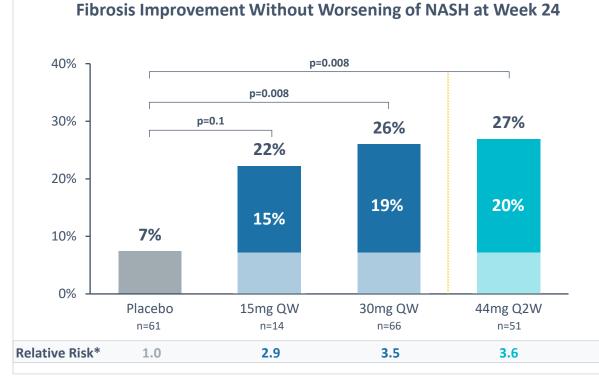
Source: Randomized Analysis Set.



ALT, alanine aminotransferase; AST, aspartate aminotransferase; BMI, body mass index; NAFLD, nonalcoholic fatty liver disease; PRO-

C3, N-terminal type III collagen propeptide; VCTE, Vibration-controlled transient elastography.

# Pegozafermin Demonstrated Statistical Significance on Fibrosis Improvement with Weekly and Every-Two-Week Dosing



 Results were statistically significant for both doses using an ITT analysis (imputes patients with missing biopsies as non-responders) and completers analysis

Relative risk is calculated by dividing drug response by placebo response

\*Relative risk presented is calculated by dividing the drug response by placebo response. Relative risk calculated using statistical methods show similar results. Source: Full Analysis Set; multiple imputation analysis via Cochran-Mantel-Haenszel (CMH) test stratified by type 2 diabetes mellitus (T2DM) status (yes vs. no) and fibrosis stage (F2 vs. F3).

## **Clinical Data in Pre-Cirrhotic Patients**

≥1 Stage Fibrosis Improvement with No Worsening of NASH (placebo-adjusted)

	Pegozafermin Phase 2b   24 weeks Multiple Imputation <sup>1</sup>		Intercept 🚺	Pharmace	igal	inver	ntiva	novo nordisk	ak	ero
			Ocaliva Phase 3   72 weeks	<b>Resmetirom<sup>2</sup></b> Phase 3   52 weeks		Lanifibranor Phase 2b   24 weeks		Semaglutide Phase 2   72 weeks	<b>Efruxifermin</b> Phase 2b   24 weeks Completers Analysis	
Drug response as multiple of placebo response*	3.5	3.6	2.3	1.7	1.9	1.1	1.6	1.3	2.0	2.0
	19%	20%					17%		19%	21%
			13%	10%	12%		1770	10%		
	<b>p=0.008</b> 30mg QW	<b>p=0.008</b> 44mg Q2W	p<0.001	p=0.0002 80mg	p<0.0001 100mg	3% p=ns 800mg	p=0.04 1200mg	p=ns0.4mg QD	p<0.05 28mg	p<0.05 50mg
	(n=66)	(n=51)	(n=308)	(n=316)	(n=321)	(n=63)	(n=69)	(n=56)	(n=38)	(n=34)

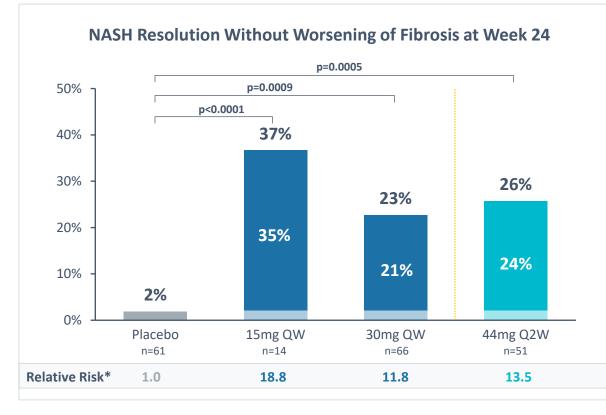
\*Drug response as multiple of placebo response is calculated by dividing drug response by placebo response

<sup>1</sup> Results same for Completer Analysis Set; <sup>2</sup>≥1 stage fibrosis improvement with no worsening of NAS

900 ns= not significant

Note: These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.

## Pegozafermin Demonstrated Statistical Significance on NASH Resolution at All Doses



 Results were statistically significant for both doses using a ITT analysis (imputes patients with missing biopsies as nonresponders) and completers analysis

Relative risk is calculated by dividing drug response by placebo response

\* Relative risk presented is calculated by dividing the drug response by placebo response. Relative risk calculated using statistical methods show similar results. Source: Full Analysis Set; multiple imputation analysis via Cochran-Mantel-Haenszel (CMH) test stratified by T2DM status (yes vs. no) and fibrosis stage (F2 vs. F3).

## Clinical Data in Pre-Cirrhotic Patients NASH Resolution with No Worsening of Fibrosis



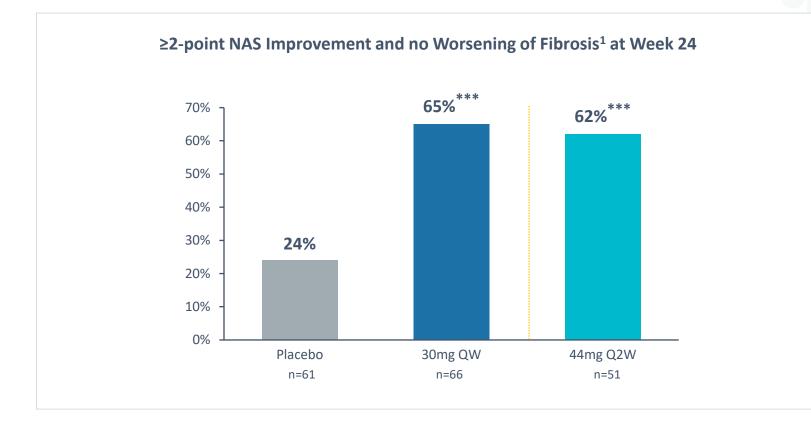
\* Drug response as multiple of placebo response is calculated by dividing drug response by placebo response

 $^{1}$  Results same for Completer Analysis Set;  $^{2}$  NASH resolution with  $\geq$ 2 point reduction in NAS and no worsening of fibrosis

Note: These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons

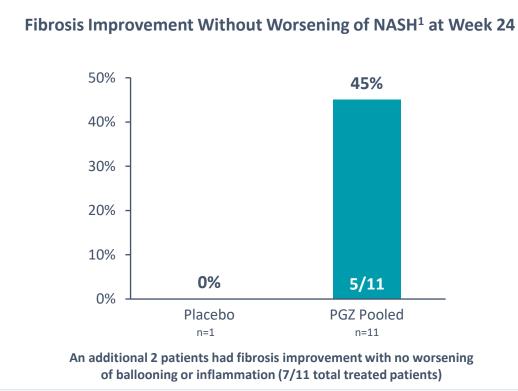
cannot be made, and no head-to-head clinical trials have been conducted.

## Pegozafermin Demonstrated Statistical Significance on ≥2-point NAS Improvement



<sup>1</sup> Full Analysis Set. Analysis via Cochran-Mantel-Haenszel (CMH) test stratified by T2DM status (yes vs. no) and fibrosis stage (F2 vs. F3) \*\*\*p<0.001 versus placebo.

## Descriptive Analysis Data of Cirrhotic (F4) Patients from ENLIVEN



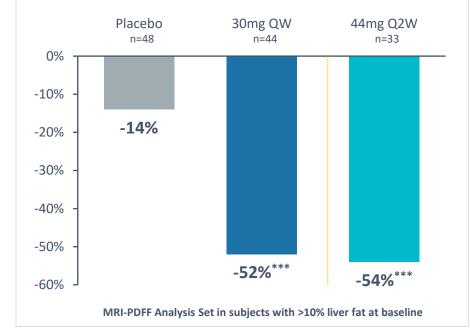
#### **EXPANSION COHORT OF THE PHASE 1B/2A TRIAL<sup>2</sup>**

 In putative F4 subjects (n=6/19), fibrosis improvement ≥1 stage without worsening of NASH ranged from 17% to 57%



<sup>1</sup>Improvement in liver fibrosis by ≥1 stage and no worsening of steatohepatitis defined as no increase in NAS for ballooning, inflammation, or steatosis 12/14 cirrhotic patients enrolled in ENLIVEN had follow-up biopsies at week 24 (11 treated patients and 1 placebo patient). <sup>2</sup>American Association for the Study of Liver Diseases (AASLD) 2022 (abstract #36901). Pegozafermin Demonstrated Robust Liver Fat Reduction with High Responder Rates by MRI-PDFF at Week 24

#### Mean Relative Reduction in Liver Fat vs Baseline<sup>1</sup> at Week 24



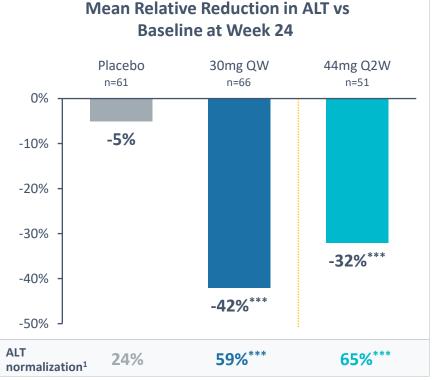
Proportion of Patients Achieving ≥50% Reductions in Liver Fat<sup>2</sup> at Week 24

<b>Placebo</b>	<b>30mg QW</b>	<b>44mg Q2W</b>
(n=48)	(n=44)	(n=33)
13%	66%***	67%***

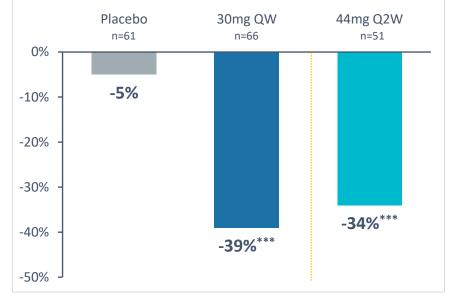
Results for the 15mg QW dose: -33% (n=12; p=ns)

<sup>1</sup>Analysis via mixed model repeated measure (MMRM). <sup>2</sup>Analysis via Cochran-Mantel-Haenszel (CMH) test stratified by T2DM status (yes vs. no) and fibrosis stage (F2 vs. F3). \*\*\*p<0.001 versus placebo

# Pegozafermin Demonstrated Significant Improvements in Markers of Liver Injury (ALT and AST)



#### Mean Relative Reduction in AST vs Baseline at Week 24

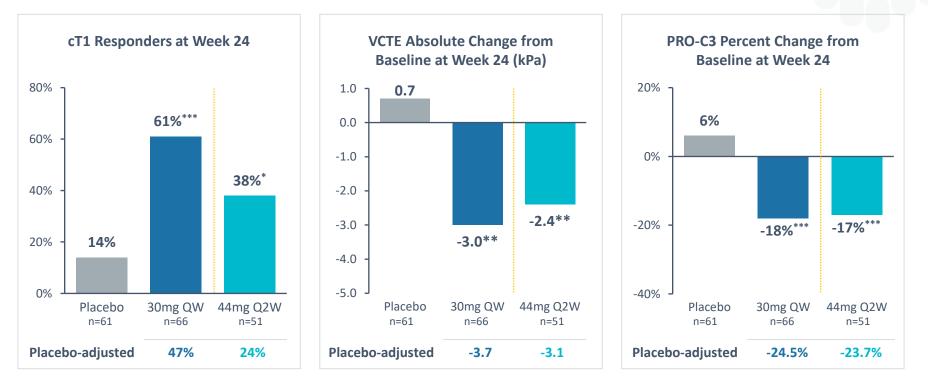


ALT Results for the 15mg QW dose: -38% (n=14; p<0.01)

 $^{1}$ ALT normalization defined as patients with ALT  $\geq$  30 U/L at baseline (n=133) with end-of-study ALT <30 U/L.

Source: Full Analysis Set: Analysis via mixed model with repeated measure (MMRM). Data presented as LS Means. \*\*\*p<0.001 versus placebo.

# Pegozafermin Demonstrated Significant Reductions in Non-Invasive Markers (NITs) of Hepatic Inflammation and Fibrosis



Results for the 15mg QW dose: cT1 40% (n=10; p=ns); VCTE -1.6 kPa (n=14; p=ns); PRO-C3 -5% (n=14; p=ns).

Source: Full Analysis Set for FibroScan and PRO-C3 assessments and MRI-PDFF analysis set for cT1, Analysis via MMRM for cT1 and PRO-C3, ANCOVA for VCTE. A patient is designated a cT1 responder with ≥80 msec reduction as compared to baseline. cT1 analysis was performed at sites where available.

\*p<0.05, \*\*p<0.01, \*\*\*p<0.001 versus placebo.

## Pegozafermin Was Well Tolerated Across Doses Low incidence of treatment-related TEAEs

#### Drug-related TEAEs in ≥10% of patients

Preferred Term	Placebo (n=69)	15mg QW (n=21)	30mg QW (n=72)	44mg Q2W (n=57)
Diarrhea	3%	24%	17%	9%
Nausea	1%	14%	21%	18%
Injection site erythema	3%	14%	14%	5%
Injection site rash	1%	0	10%	4%
Increased appetite	0	10%	13%	5%

Most TEAEs were grade 1 or 2. No tremor reported.

	Placebo	15mg QW	30mg QW	44mg Q2W
Drug-related AEs leading to discontinuation	0	5%ª	6% <sup>b</sup>	2% <sup>c</sup>
Drug-related Serious Adverse Event (SAE)	0	0	0	2% <sup>c</sup>

Related discontinuations: <sup>a</sup> Diarrhea [15 mg QW]; <sup>b</sup> Diarrhea [30 mg QW]; Nausea [30 mg QW]; Diarrhea [30 mg QW]; ISR erythema [30 mg QW]; <sup>c</sup> Pancreatitis [44 mg Q2W]. Unrelated discontinuations: Angina [placebo]; Colon CA [30 mg QW]; COVID-19 [30 mg QW].

# Poised for Phase 3 Advancement in NASH

## **Proposed regulatory and clinical development plans\***

- Meeting with FDA planned in 2H23; pursue EU scientific advice in parallel
- Phase 3 trial in F2/F3 patients with histology endpoint for accelerated approval
- Phase 3 trial in F4 patients with outcomes endpoint for full approval (study in parallel)
- Planned SHTG Phase 3 trials are expected to satisfy safety database requirements and for approval in SHTG

# Pegozafermin Has Potential to Address a Large Commercial Opportunity

## NASH represents a large patient population with significant health risks

• 13M F2-F4 patients eligible in the United States with equivalent number in EU

## Significant market opportunity for pegozafermin as injectable therapy

- Pegozafermin positioned for medium-high risk (F2/F3) non-cirrhotic patients and compensated cirrhotic patients
- Large markets in other disease areas have supported multiple competing orals and injectables, with potent injectables often dominating
  - Injectable GLP-1s account for >\$20B of \$38B in worldwide sales of branded type 2 diabetes drugs despite competing against oral SGLT2 and DPP-4 inhibitors
  - Subcutaneous injectable immunomodulators for rheumatoid arthritis and plaque psoriasis dominate sales (70%) and market share (80%) vs oral competitors
- Best-in-category mechanisms often have multiple successful drug with the same MOA
  - 4 different GLP-1s for T2DM each had sales > \$1B in 2022 (two were the 4th and 5th entrants)
  - 4 different SC anti-IL23 and anti-IL17 drugs for plaque psoriasis each had sales > \$1B in 2022

# Pegozafermin Has Potential to Address a Large Commercial Opportunity

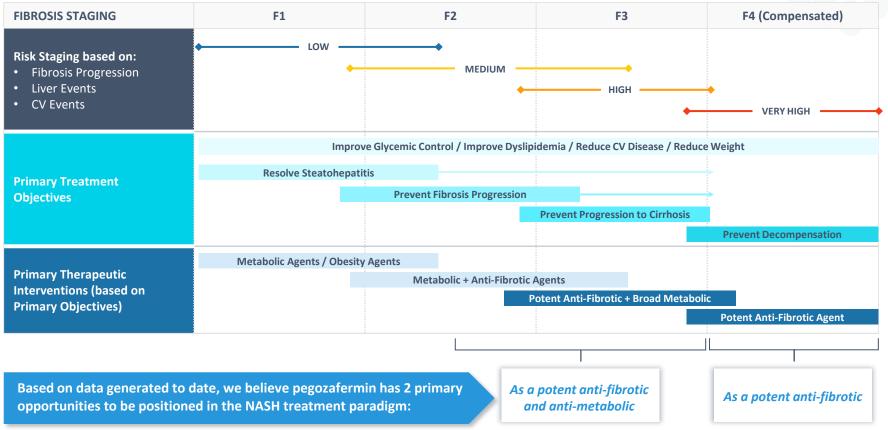
# Pegozafermin has demonstrated a differentiated profile across key attributes in a chronic asymptomatic condition: Efficacy, Tolerability, Convenience

- Pegozafermin has demonstrated potential best-in-class potency with superior tolerability
  - GI adverse events impact patient lives; #1 non-efficacy reason for GLP-1 discontinuations
- Less frequent dosing is highly meaningful to patients and offers physicians optionality
  - 66% of patients would strongly prefer or prefer every-two-week to weekly dosing
  - Competitive markets have shown a preference for extended dosing; Humira (\$21B) and Dupixient (\$9B) are dosed Q2W; Fasenra (anti-IL5) for eosinophilic asthma with similar efficacy profile but less frequent dosing vs. its competitor reached sales parity despite launching 2 years later

## Payers recognize the value proposition in NASH

- Payers expecting premium pricing based on medical necessity
- Payers are unlikely to require diagnostic biopsies to approve treatment

## NASH Treatment Paradigm & Pegozafermin Positioning



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# Opportunity in Severe Hypertriglyceridemia (SHTG)





# Pegozafermin Could Offer an Important New Treatment Option for SHTG

### Large growing patient population with significant health risks; overlap with NASH patient population

- Increasing TG levels increase risk of acute pancreatitis, cardiovascular disease and all-cause mortality
- Emerging evidence of the cardiovascular (CV) benefit associated with TG reduction in patients with CV risk factors

### Significant market opportunity for agent with broad metabolic benefits

- Pegozafermin has a unique selling proposition that is meaningful to prescribers more effective triglyceride reduction with improvements in liver fat and other metabolic measures
- Highly differentiated from approved therapies based on superior broad efficacy and/or safety
- Analyst consensus peak year sales estimated to be greater than \$1 billion (US only)

## Clinical program substantially de-risked

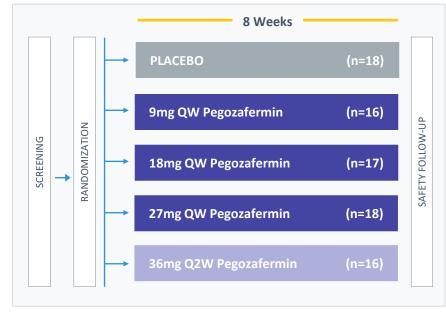
- Phase 3 design similar to positive Phase 2 (ENTRIGUE) design with same primary endpoint
- Agency provided feedback to company on key elements of regulatory path to approval

### SHTG program is synergistic with the NASH program

- Development: Leverages safety database across the two programs to minimize spend across total program
- Commercial: Leverage sales force and infrastructure costs

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# ENtrigue – Phase 2 SHTG Trial Design



Magnetic Resonance Imaging – Proton Density Fat Fraction

QW, once-weekly; Q2W, once every two weeks.

Safety analysis set, n=85 (patients who received at least 1 dose)

Full analysis set, n=82 (patients with at least 1 post-baseline TG assessment) MRI analysis set n=23 (patients with baseline and end of treatment MRIs)

#### **KEY INCLUSION CRITERIA**

- TG  $\geq$ 500mg/dL and  $\leq$ 2,000mg/dL
- Background therapy: statins and/or prescription fish oil and/or fibrates OR none

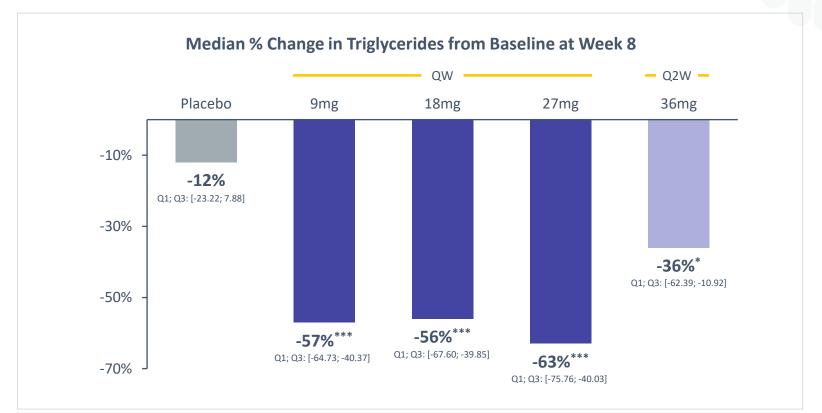
#### **PRIMARY ENDPOINT**

• Primary endpoint: % Change in TGs from baseline

#### **KEY SECONDARY ENDPOINTS**

- Lipids: non-HDL-C, HDL-C, Apo-B
- Liver fat (MRI-PDFF)
- Glycemic control

## Pegozafermin Significantly Reduces Triglycerides Across All Dose Groups Primary Endpoint

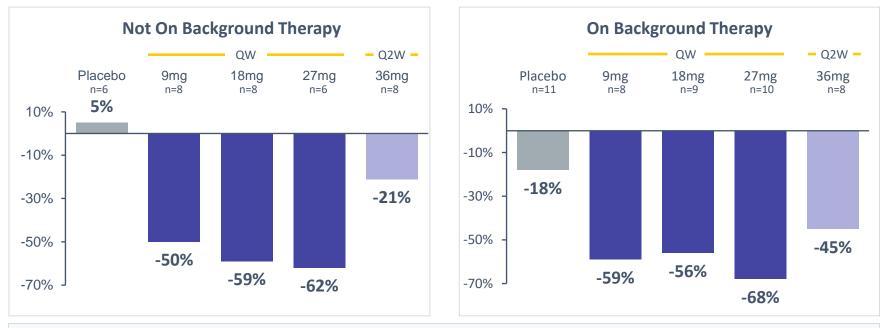


QW: Every week; Q2W: Every 2 weeks



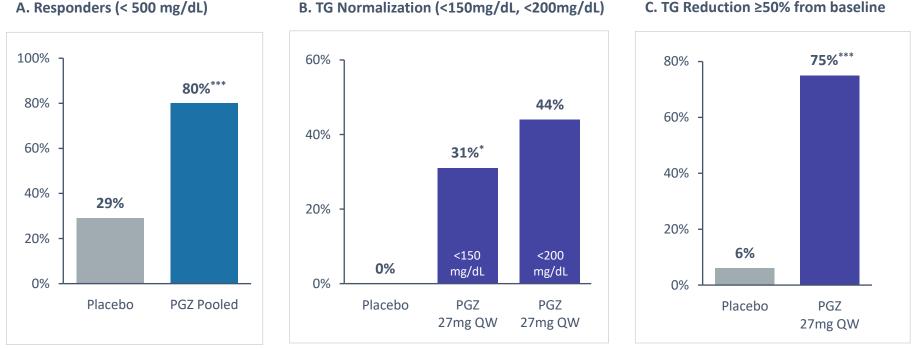
# Pegozafermin Shows Significant Decrease in Triglycerides on Top of Background Therapy

## Median % Change in Triglycerides from Baseline at Week 8



Results are consistent with data from patients on background therapy of statins or statin combos, prescription fish oils, and fibrates

# Pegozafermin Shows Significant Decrease in Triglycerides at Different **Threshold Levels**



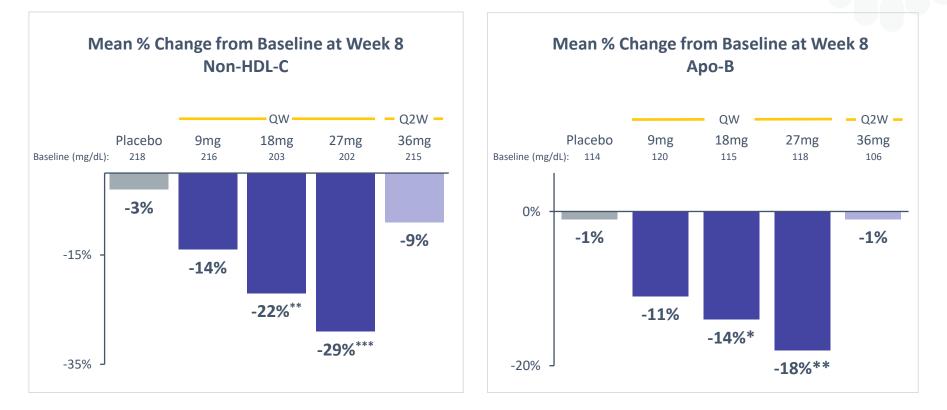
B. TG Normalization (<150mg/dL, <200mg/dL)

C. TG Reduction ≥50% from baseline

Analysis via unstratified Chi-square Test comparing the individual PGZ groups vs placebo. \* p<0.05; \*\* p<0.01; \*\*\* p<0.001 vs. placebo TG Responders defined as patients who achieve TG <500 mg/dL

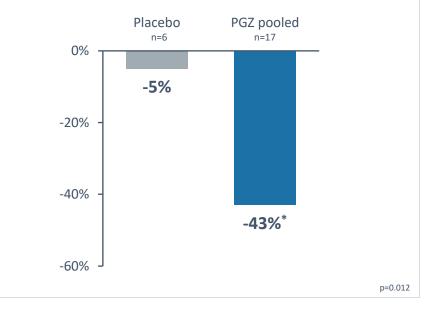
Full Analysis Set

## Pegozafermin Demonstrated Clinically Meaningful Improvements in Non-HDL-C and Apo-B – Key Marker of CV Risk



## Pegozafermin Demonstrated Significant Reduction in Liver Fat Liver Fat Is an Important Potentiator of CV Risk

#### Mean Relative Reduction in Liver Fat vs Baseline at Week 8



#### **HIGH RESPONDER RATES**

- ≥30% Reduction in liver fat: 88% vs 0% in placebo
- ≥50% Reduction in liver fat: 41% vs 0% in placebo
- Normalized liver fat: 24% vs 0% in placebo

Pegozafermin (n=17) and Placebo (n=6)

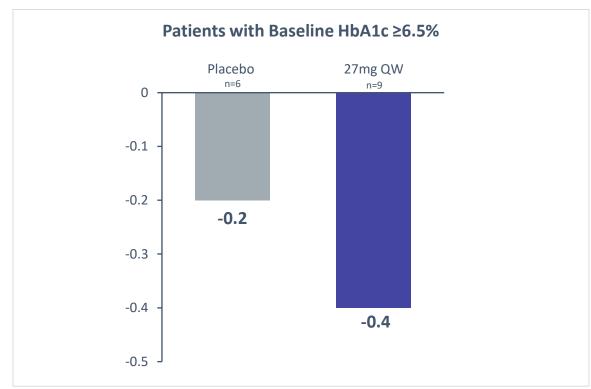
Post-hoc analysis of patients with follow-up MRI-PDFF  $\leq$ 21 days from date of last dose (n=14) resulted in 29% of patient with normalized liver fat and 100% and 50% of patients with mean relative reductions of  $\geq$ 30% and  $\geq$ 50% from baseline, respectively.

Post-hoc analysis of patients with follow-up MRI-PDFF <21 days from date of last dose in 27mg QW cohort (n=5) demonstrated a 63% mean relative reduction from baseline.

\*p <0.05 vs. placebo MRI Analysis Set; p value vs placebo based on ANCOVA analysis

## Pegozafermin Demonstrated Improvement on HbA1c that May Increase With Longer Treatment

## Absolute Change in HbA1c at Week 8



# Pegozafermin Demonstrated Favorable Safety/Tolerability in Phase 2 Study

- Pooled pegozafermin treatment related Adverse Events (AEs) observed in ≥7.5% of patients were:
  - Nausea (10%), diarrhea (9%) and injection site reaction (9%) vs placebo (0%)
  - All Gastrointestinal (GI) AEs were Grade 1 or 2
- No Grade 3 or higher AEs
- No treatment-related SAEs; 2 treatment-related discontinuations at 27mg QW (both Grade 2)
- No tremor or hypersensitivity AEs reported
- No adverse effects on blood pressure or heart rate

# Phase 3 Program Initiation Planned in First Half of 2023

#### **Regulatory**

- FDA feedback supports advancement into Phase 3
- FDA feedback confirms key elements of the overall Phase 3 development program<sup>\*</sup>
  - Primary endpoint: TG reduction from baseline (anticipated to be assessed at the 26-week timepoint)
  - Proposed doses
  - Two well-controlled Phase 3 trials in SHTG patients of one year duration will contribute to the efficacy and safety database required to support the registration package

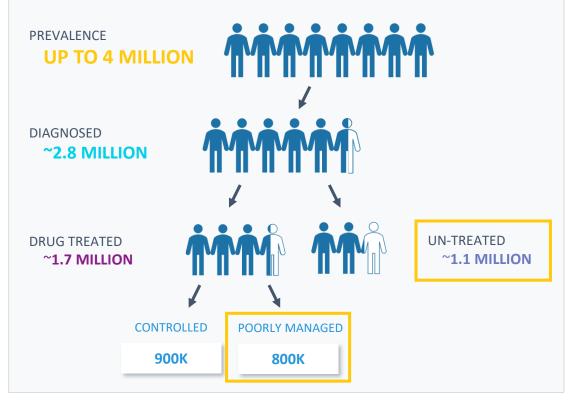
#### **Clinical development**

• Trial start-up activities underway – plan to initiate the first SHTG Phase 3 trial in 2Q23

### **Technical Operations**

• Developed new pre-filled syringe using liquid formulation for use in planned Phase 3 SHTG trial in 2Q23

### SHTG Represents a Large Population with High Unmet Need



Co-morbidity	Prevalence in SHTG population
Fatty Liver Disease (NAFLD)	Up to <b>100%</b>
Type 2 diabetes/Prediabetes	Up to <b>70%</b>
Dyslipidemia	Up to <b>65%</b>

Pegozafermin profile is unique and compelling to physicians because of potential for metabolic benefits



### Pegozafermin Delivers on Key Attributes for Successful SHTG Therapy

MINOR INFLUENCE		MODEST INFLUENCE		MAJOR INFLUENCE
	Hiera	rchy of Attributes for SHTG	Therapy	
RoA/Dosing	Clinical Outcomes	Safety/Tolerability	Metabolic Endpoints	TG Endpoints
<ul> <li>RoA and dosing were seen as the least influential given familiarity with</li> </ul>	<ul> <li>Physicians noted that clinical outcomes are not required to drive utilization in SHTG</li> </ul>	<ul> <li>Safety and tolerability have a lesser impact on treatment decisions compared to efficacy</li> </ul>	<ul> <li>Metabolic endpoints were viewed as additive benefits</li> <li>Fatty liver, HbA1c, and</li> </ul>	<ul> <li>TG lowering is the most influential endpoint to drive utilization</li> <li>Significant efficacy</li> </ul>
injectables in T2D	Physicians were receptive to using TG as a surrogate endpoint		weight loss serve as differentiators	improvement over SoC will drive utilization
		<ul> <li>Generally well- tolerated</li> </ul>	<ul> <li>43% mean relative reduction in liver fat<sup>1</sup></li> </ul>	<ul> <li>63% reduction in TG from baseline<sup>2</sup></li> </ul>
			<ul> <li>0.4% absolute reduction in HbA1c<sup>2</sup></li> </ul>	<ul> <li>80% of subjects achieved</li> </ul>
		Physician Enthusiasm for Metabolic Endpoints	TG<500mg/dL <sup>1</sup>	
		LOW HIGH		
<sup>1</sup> Pooled pegozafermin data at <sup>2</sup> 27mg pegozafermin data at RoA: Route of Administration Source: Physician Interviews:	week 8		Liver fat reduction Decrease in HbA1 PEGOZAFERMIN ATTRIBUTES	

### Pegozafermin Profile Supports Utilization Over Current SoC and Future Competitive Agents

	IN DEVELOPMENT		APPROVED					
	Pegozafermin	APOC3 Potential Fibrates	Prescription Fish Oils		- Statins			
	Potential		Fibrates	Fibiates	Fibrates	Potential Vascepa	Vascepa	Lovaza
Triglyceride reduction	111	<b>V</b> V	<b>VV</b>	✓	~~	$\checkmark\checkmark$		
Liver fat/Fibrosis reduction	J/J	√/—	Worsens liver fat	_		-		
Insulin sensitizing	✓	—	_	_	-	_		
Apo-B lowering	✓	✓	-	$\checkmark$		$\checkmark$		
ALT lowering	✓	Transaminase elevations observed	Monitor ALT	-	May require ALT monitoring	Monitor ALT		

For triglyceride reduction:  $\checkmark \checkmark \checkmark = 260\%$ ,  $\checkmark \checkmark = 31\%-59\%$ ,  $\checkmark = \le 30\%$  — No effect/Not reported



Sources: Feingold KR. Triglyceride Lowering Drugs. [Updated 2021 Apr 1]. In: Feingold KR, Anawalt B, Boyce A, et al., editors. Prescribing information. Corporate presentations. Note: All data regarding third-party molecules on this slide are based on third-party studies, which are in different stages of development, and not our own. Conclusions on this slide are company estimates and are not based on head-to-head results.

### **Corporate Highlights**



Late-stage clinical development company entering Phase 3 in NASH and SHTG

Pegozafermin (FGF21 analog) has demonstrated
 differentiation within class and category and has
 the potential to be transformational for patients

Strong balance sheet (\$480.9M\* in cash, cash equivalents and short-term investments)

Highly experienced team with track record of execution

# 89bio

## Appendix

### **Experienced Management Team Positions 89bio for Success**









Rohan Palekar CEO

CEO, CCO experience

Commercial, strategy, and R&D experience

### Hank Mansbach, MD CMO

20+ years biopharma and R&D leadership in clinical development and medical affairs Ryan Martins CFO

CFO, Strategy/IR, finance, sell-side experience

### Quoc Le-Nguyen CTO & Head of Quality

20+ years biopharma and leadership in technical operations, product supply, and quality





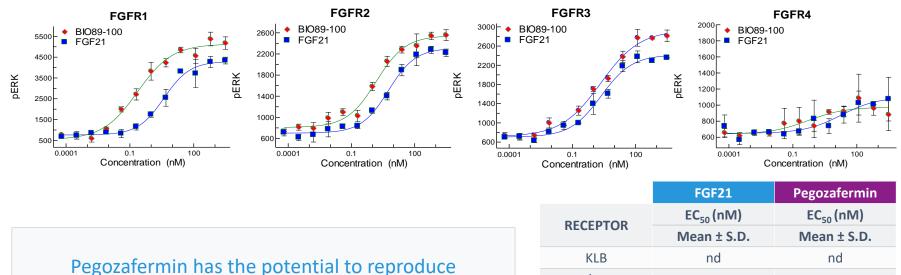




Aduro
 Novartis
 BIOMARIN



### Pegozafermin Exhibits Highly Potent FGF Receptor Agonism



the beneficial metabolic effects of native FGF21

RECEPTOR			
RECEPTOR	Mean ± S.D.	Mean ± S.D.	
KLB	nd	nd	
KLB/FGFR1	4.5 ± 1.0	0.3 ± 0.07	
KLB/FGFR2	4.5 ± 0.9	$1.1 \pm 0.4$	
KLB/FGFR3	$1.8 \pm 0.3$	$1.2 \pm 0.4$	
KLB/FGFR4	nd	nd	
and wet determined, ab ECE4			

nd – not determined; rhFGF19 EC<sub>50</sub> at FGFR4 =  $1.7 \pm 0.4$ 

\* Receptor agonism measured in L6 cells expressing  $\beta$ -klotho and either FGF Receptor 1c, 2c, 3c, or 4 via pERK functional assay \*\* Figures represent data from a single experiment; Table represents mean data from multiple experiments

### Learnings from the Obeticholic Acid NASH Phase 3 Program: **Comparison of Single Central Reader vs. 3-Panel Consensus**

#### **Original Analysis** Consensus Method Read p=0.0002 p<0.0001 25.0-25-Percent Responders NS 23.1 NS 22.4 20.0-20-17.6

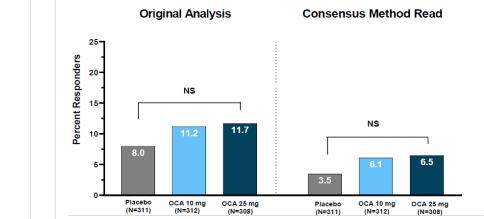
14.1

Improvement of Fibrosis by ≥ 1 Stage without Worsening NASH

OCA 25 mg

(N=308)

#### **Resolution of NASH with No Worsening of Liver Fibrosis**



#### **OBSERVATIONS:**

5.0-

0.0

11.9

Placebo

(N=311)

Placebo response for NASH resolution is >2 fold higher with single reader vs 3-panel consensus

OCA 10 mg

(N=312)

Placebo response similar to ENLIVEN study for both fibrosis improvement and for NASH resolution

OCA 25 mg

(N=308)

#### IMPLICATIONS:

3-panel consensus highlights treatment delta but dampens absolute response

9.6

Placebo

(N=311)

3-panel consensus methodology can reproduce low placebo response in phase 3 trial

OCA 10 mg

(N=312)

### Pre-Specified ITT Analysis Confirms Robustness of Primary Efficacy Results

#### ITT (missing data = non-responder); (n=192) at Week 24

	30mg QW	44mg Q2W		
Fibrosis improvement without worsening of NASH				
Effect Size (placebo-adjusted)	15%	16%		
p-value	0.019	0.015		
NASH resolution without worsening of fibrosis				
Effect Size (placebo-adjusted)	17%	20%		
p-value	0.0019	0.0009		



### Pegozafermin Demonstrated Meaningful Reductions in HbA1c (ENLIVEN)

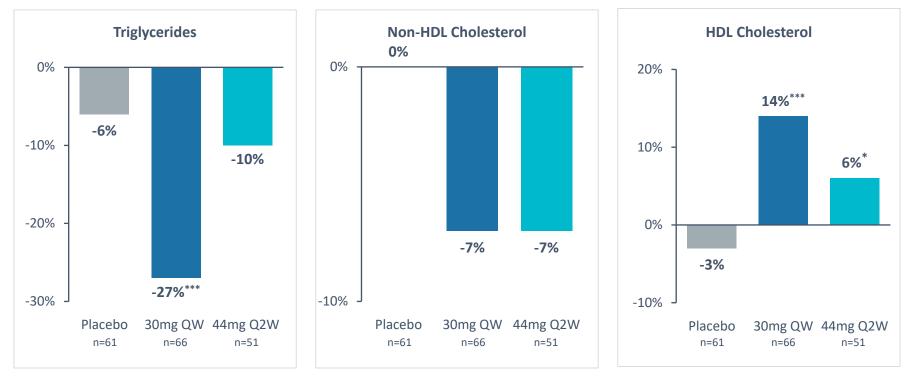


Change in HbA1c from Baseline at Week 24



Source: Full Analysis Set for either overall population or FAS with baseline HbA1c ≥6.5%. Analysis via MMRM. \*p<0.05, \*\*p<0.01 versus placebo.

### Pegozafermin Demonstrated Meaningful Changes in Serum Lipids (ENLIVEN)



#### Percent Change in Serum Lipids from Baseline at Week 24

Source: Full Analysis Set. Analysis via van Eltren Test for triglycerides (reported as median) and mixed model with repeated measure (MMRM). Subjects with missing week 24 triglycerides are

excluded from the non-parametric analysis.

Non-HDL-cholesterol and HDL Cholesterol (reported as LS means) with changes from baseline (absolute or %) as dependent variables.

\*p<0.05, \*\*p<0.01, \*\*\*p<0.001 versus placebo.

### Comparative Profile of FGF21 Analogs - Histology and Liver Markers

	Pegozafe	Pegozafermin (PGZ)		Efruxifermin (EFX)		
	30mg QW	44mg Q2W	28mg	50mg		
Structure (molecular weight)		<ul> <li>GlycoPEGylated FGF21 (40 kDa)</li> <li>Structurally different from other monovalent analogs</li> </ul>		Fc-fusion FGF21 (92 kDa)		
Potency against FGF receptors (EC50) <sup>1</sup>	PGZ (r	FGF21)	EFX (rFGF21)			
1c/2c/3c	0.3 (4.5)/1.1	(4.5)/1.2 (1.8)	0.3 (0.08)/0.7 (0.4)/1.9 (1.0)			
Efficacy (histology)						
Methods	<ul> <li>Rigorous biopsy reading 3 separate independent reader (no bias); algorithm derived score</li> <li>95% + alignment based on 2+ reader agreement</li> </ul>		2 independent readers with consensus based on discussion (no adjudication required)			
Fibrosis 1-point improvement						
Placebo adjusted percent	19%	20%	19%	20%		
Relative risk	3.5	3.6	2.0	2.0		
NASH Resolution						
Placebo adjusted percent	21%	26%	32%	61%		
Relative risk	11.8	13.5	3.1	5.1		
Key liver non-invasive markers						
Liver fat change baseline <sup>2</sup>	-52%	-54%	-52%	-64%		
VCTE kPa (chg baseline, placebo adjusted)	-3.7	-3.1	-1.9	-3.6		
ProC3 (chg baseline, placebo adjusted)	-25%	-24%	-28%*	-33%*		
ALT	-42%	-32%	-38%	-47%		



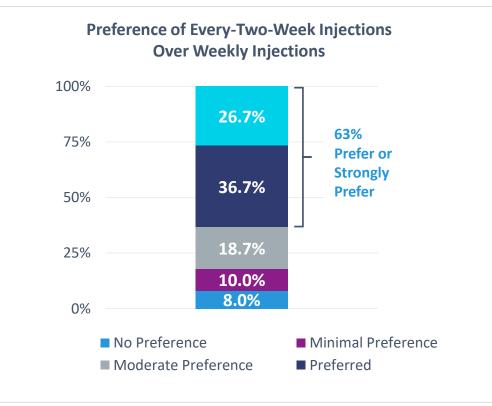
<sup>1</sup>LG cells overexpressing β-klotho and either FGF receptor 1c, 2c, 3c via PERK functional assay. PGZ data is based on MRI-PDFF Analysis Set in subjects with >10% liver fat at baseline Note: All data regarding third-party molecules on this slide are based on third-party studies and not our own. Conclusions on this slide are not based on head-to-head results \*Calculated values

### Comparative Profile of FGF21 Analogs in NASH – Safety/Tolerability

	Pegozafe	Pegozafermin (PGZ)		min (EFX)
	30mg QW	44mg Q2W	28mg	50mg
Tolerability and safety (key terms)				
Diarrhea	17%	9%	35%	33%
Nausea	21%	18%	25%	33%
Frequent bowel	-	-	20%	-
Increased appetite	13%	5%	18%	23%
Injection site erythema	14%	5%	15%	16%
Injection site bruising	-	-	15%	7%

89bio Note: All data regarding third-party molecules on this slide are based on third-party studies and not our own. Conclusions on this slide are not based on head-to-head results

### Over 60% of T2D Patients Prefer or Strongly Prefer Every-Two-Week Injections



- Every-two-week dosing provides opportunity for physicians to optimize therapy to patient preference
- Compliance is important in treatment for chronic, asymptomatic diseases