



**Corporate Brief Q2 2024**

Juan Pablo Frías, MD  
Chief Medical Officer and Head of Diabetes  
Biomea Fusion  
Redwood City, CA

## Disclaimer

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Excellent Science - Combining Validated Targets with Breakthrough Chemistry

## We Aim to Cure

➤ Experienced Management Team

➤ Novel FUSION™ System

➤ BMF-219 – Phase II - Lead Asset

➤ BMF-500 – Phase I

➤ Discovery Programs



Biomea Fusion is a clinical-stage biopharmaceutical company focused on the discovery and development of **oral covalent small-molecule drugs** to treat patients with metabolic diseases and genetically defined cancers. We believe that our approach may lead to significant improvement and extension of life for patients. Our team is engaged in all phases of drug discovery and development, including target selection, small molecule design, and preclinical and clinical studies to develop innovative medicines.

Aiming to Develop some of the Most Impactful Medicines of our Time

# A Long History of Developing Successful Drugs - Together



**Thomas Butler**  
Chairman & CEO



**Ramses Erdtmann**  
President & COO



**Juan Frías, M.D.**  
Chief Medical Officer



**Naomi Cretcher**  
Chief of People



**Heow Tan**  
Chief Technical & Quality Officer



**Steve Morris, M.D.**  
Chief Development Officer



**Franco Valle**  
Chief Financial Officer



Co-Founder

The FUSION™ SYSTEM

**BMF-219\***

Co-Inventor



560, 420, 280, 140 mg tablets | 140, 70 mg capsules



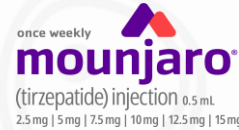
Co-Inventor



Co-Founder



560, 420, 280, 140 mg tablets | 140, 70 mg capsules



once weekly (tirzepatide) injection 0.5 mL  
2.5 mg | 5 mg | 7.5 mg | 10 mg | 12.5 mg | 15 mg



(dapagliflozin) 5mg & 10 tablets



(empagliflozin) tablets 10 mg/25 mg



once weekly (dulaglutide) injection 0.5 mL  
0.75 mg | 1.5 mg | 3.0 mg | 4.5 mg



ONCE - WEEKLY semaglutide injection 2.4 mg



(exenatide) injection



(canagliflozin) tablets



semaglutide injection 0.5mg, 1mg, 2mg



sitagliptin



560, 420, 280, 140 mg tablets | 140, 70 mg capsules



560, 420, 280, 140 mg tablets | 140, 70 mg capsules



EXTENDED-RELEASE CAPSULES



100 mg tablets



100 mg tablets



30 mg TABLETS



150 mg tablets



100 mg tablets



150 mg capsules



30mg TABLETS



560, 420, 280, 140 mg tablets | 140, 70 mg capsules

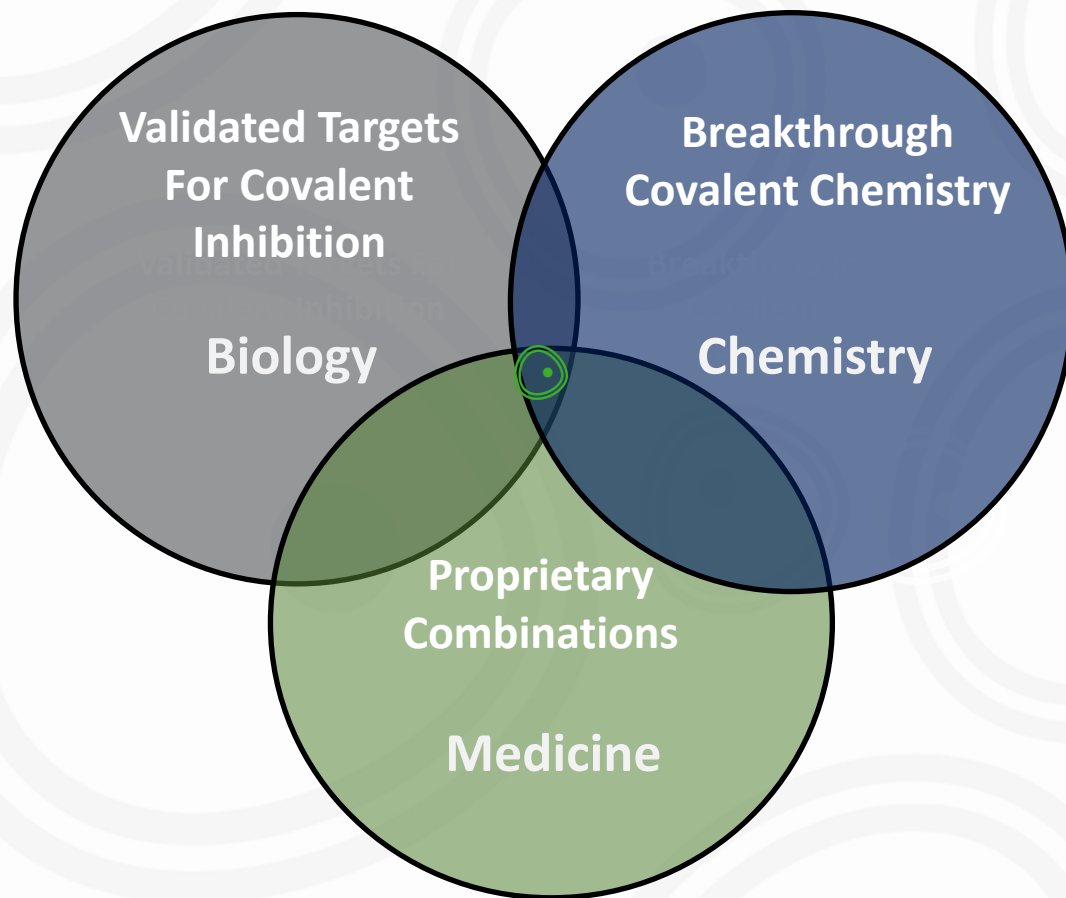


We Aim to Cure™

Confidential

\*Note: BMF-219 is an investigational new drug

## Biomea's Development Principles



Drugs pursuing **Validated Disease Targets** have a ~2x higher likelihood of approval than molecules pursuing a new mechanism of action

Sources: Nelson et al. (2015) Nat Genet.; Thomas et al. (2016) BIO; In a Landscape of 'Me Too' Drug Development, What Spurs Radical Innovation? HBS Weekly Review (Jun 2018)



**Covalent Small Molecule Inhibitors** provide deep target inactivation and a wider therapeutic window, allowing for longer duration on therapy

Sources: Singh et al. (2011) Nature Reviews Drug Discovery; Cheng et al. (2020) Journal of Hematology & Oncology; Strelow (2017) SLAS Discovery; Kalgutkar & Dalvie (2012) Expert Opin. Drug Discov.;



**Combination Therapy** with non-overlapping resistance mechanisms results in more durable responses and better outcomes

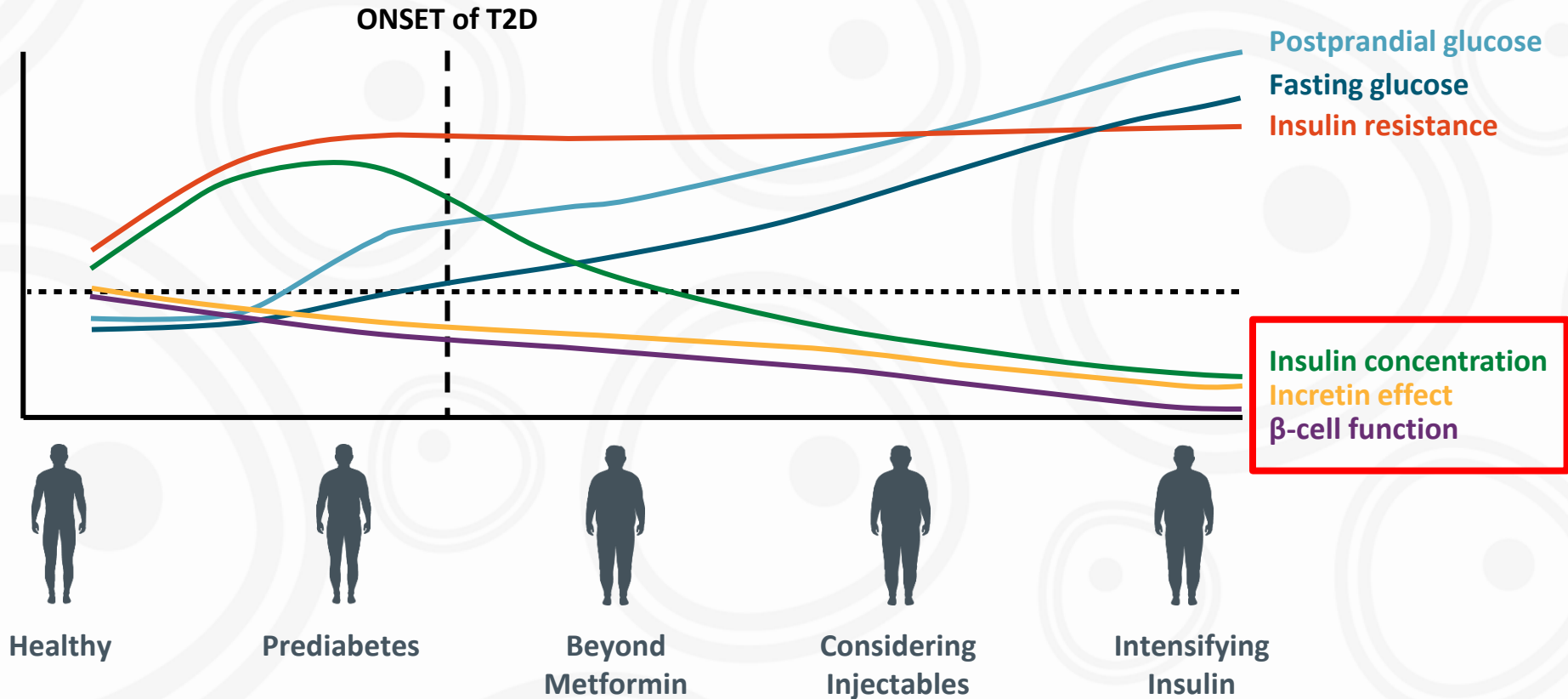
Sources: Palmer et al. (2019) eLife; Mokhtari et al. (2017) Oncotarget

## Biomea's Pipeline Expands into 5 Clinical Trials with 2 Novel Agents

### Multiple Upcoming Milestones in the Near Term

	Study	Indications	Milestones	Expected Timeline
BMF-219 Menin Program	COVALENT-111	Type 2 Diabetes	Phase II - Dose Expansion Initial Data Readout	2024
	COVALENT-112	Type 1 Diabetes	Phase II - Initial Proof of Concept	2024
	COVALENT-101	Liquid Tumors	Phase I - Dose Escalation Completion	2024
	COVALENT-102	Solid Tumors	Phase I - Dose Escalation Completion	2024
BMF-500 FLT3 Program	COVALENT-103	AML/ALL (acute leukemia)	Phase I - Dose Escalation Completion	2024
Additional Program	Target # 3	TBA	Announce IND Candidate	2024

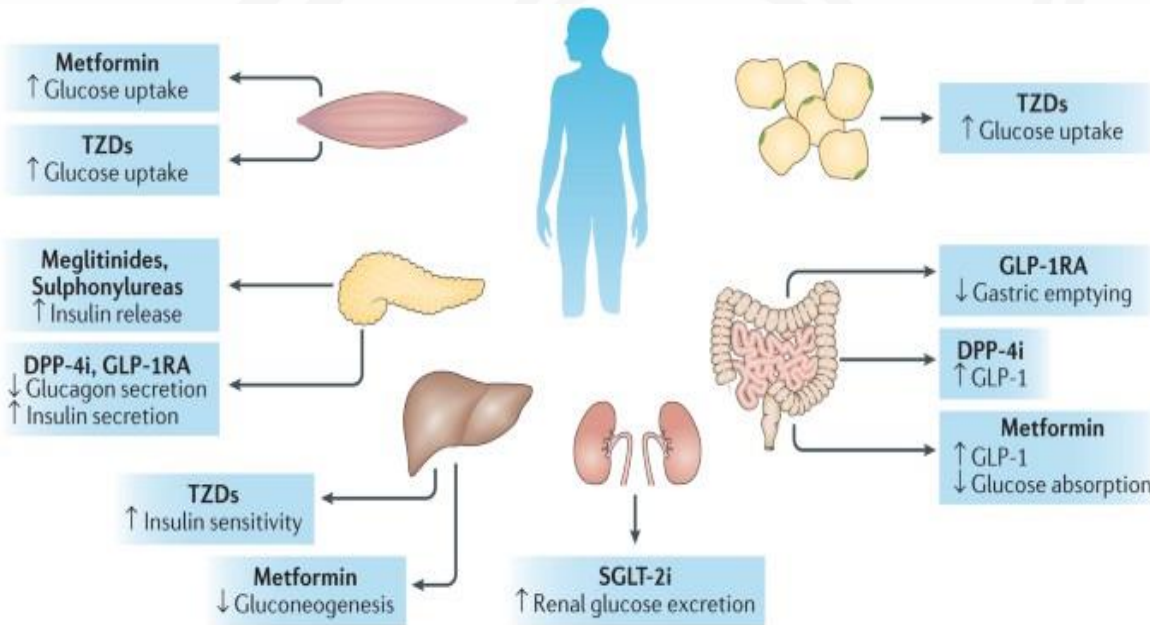
# Natural History of Types 2 Diabetes – A Progressive Decline in Beta-Cell Function



T2D, type 2 diabetes  
Kendall, D. Am J Med. 2009;122:S37.

## BMF-219 Mechanism of Action

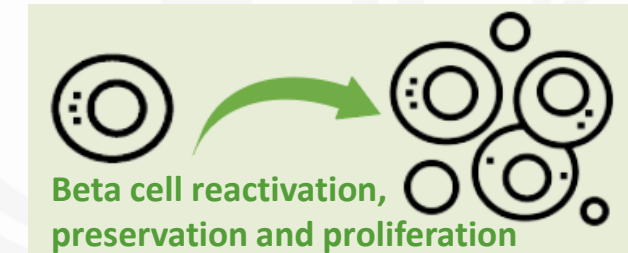
# BMF-219 is a Potential First in Class Diabetic Agent – Addressing the Root Cause of Disease



[Nat Rev Endocrinol 12, 337–346 \(2016\).  
https://doi.org/10.1038/nrendo.2016.51](https://doi.org/10.1038/nrendo.2016.51)

Currently approved therapies are primarily targeting the **Symptoms of Type 2 Diabetes: *Hyperglycemia***

## BMF-219: Menin Inhibition a Potential New Class of Diabetes Agents



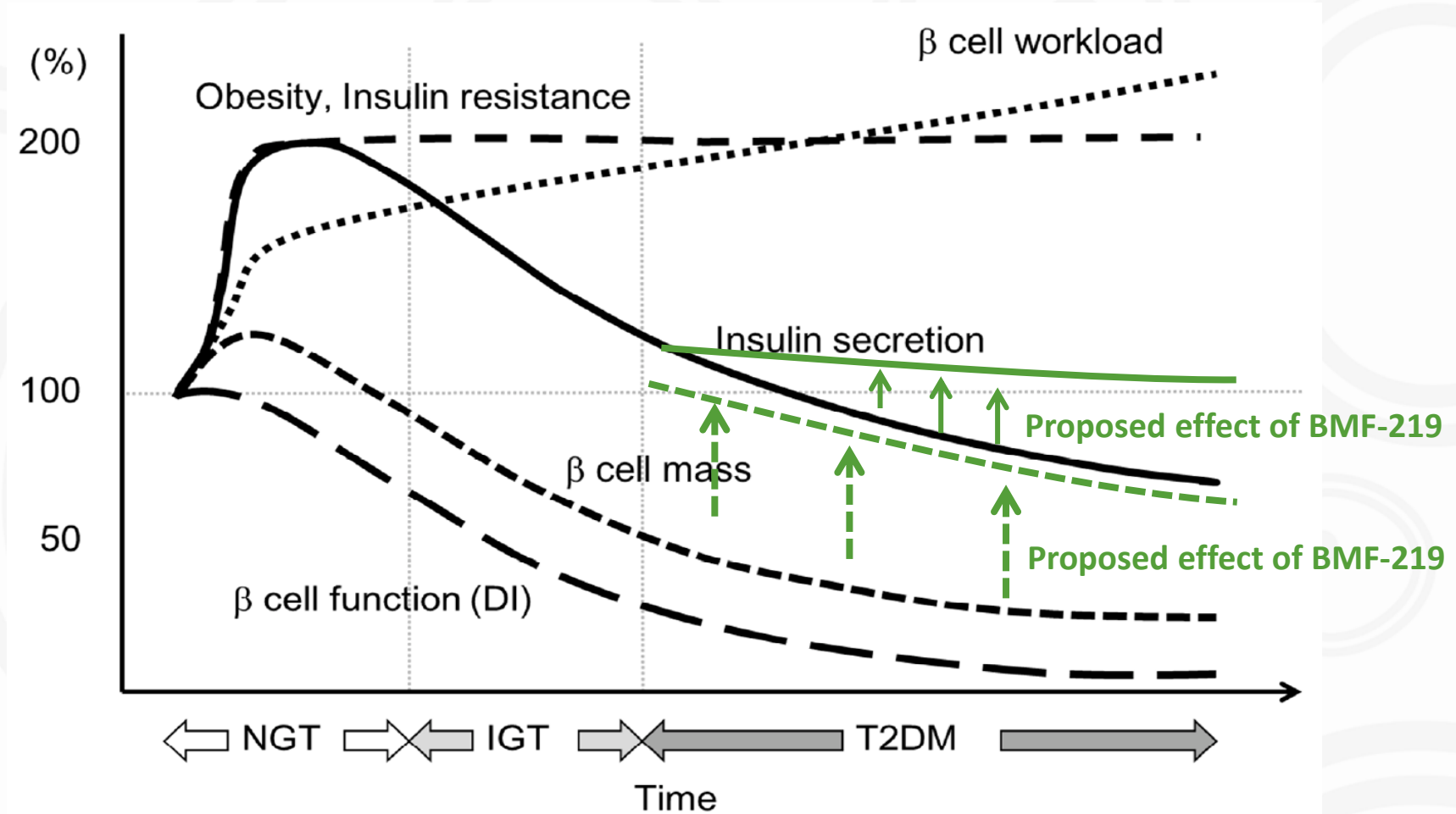
**Beta Cell Mass ↑    Beta Cell Health ↑**

**Control of Glycemia even after Cessation of Dosing**

BMF-219 represents a potential new class of diabetes agents addressing the: **Root Cause of Diabetes**  
- **Loss of Beta Cell Mass and Function** -



# The Goal for BMF-219 is to Improve Glycemic Control without Continuous Medication



BMF-219 is aimed to increase beta-cell mass and function, thereby increase insulin production in order to achieve glycemic control - without the need of continuous medication.

## Investigational BMF-219 Has a Unique Potential Value Proposition in Early Clinical Trials

### How is BMF-219 Differentiated from Currently Available Diabetes Therapies?

Oral Small Molecule

Complementary  
Agent to Available  
Diabetes Therapies

Non-Chronic  
Dosing

Well-Tolerated Safety  
Profile  
after First Read Out

Disease Modifying Potential  
Addressing the Root Cause of Diabetes

Continued Glycemic Control even after  
Cessation of Dosing

Addressable Population may Include All People with Diabetes

## COVALENT-111 Study Design (Type 2 Diabetes)

# Additional Dose Levels and Various Dosing Durations Are Being Explored in the Escalation and Expansion Portion of COVALENT-111

**Part 1 Dose Escalation,**  
4 weeks dosing+ 22 weeks follow up

**Healthy Volunteers**  
n=16

**50 mg QD, n=10**  
x 4 wks

**100 mg QD, n=20**  
x 4 wks

**200 mg QD / 100 mg BID, n=22**  
x 4 wks

**200 mg QD, n=10** → **400 mg QD, n=10**  
x 2 wks, n=10 x 2 wks

**Part 2 Dose Expansion, n=216 – 288 incl.**  
12 weeks dosing + 40 weeks follow-up

Arm A\* **100 mg, n=72**  
x 8 wks

Arm B **100 mg, n=72**  
x 12 wks

Arm C **100 mg, n=72** → **200 mg, n=72**  
x 8 wks x 4 wks

Arm D Anticipated to be added based on data from  
400 mg cohort of escalation portion

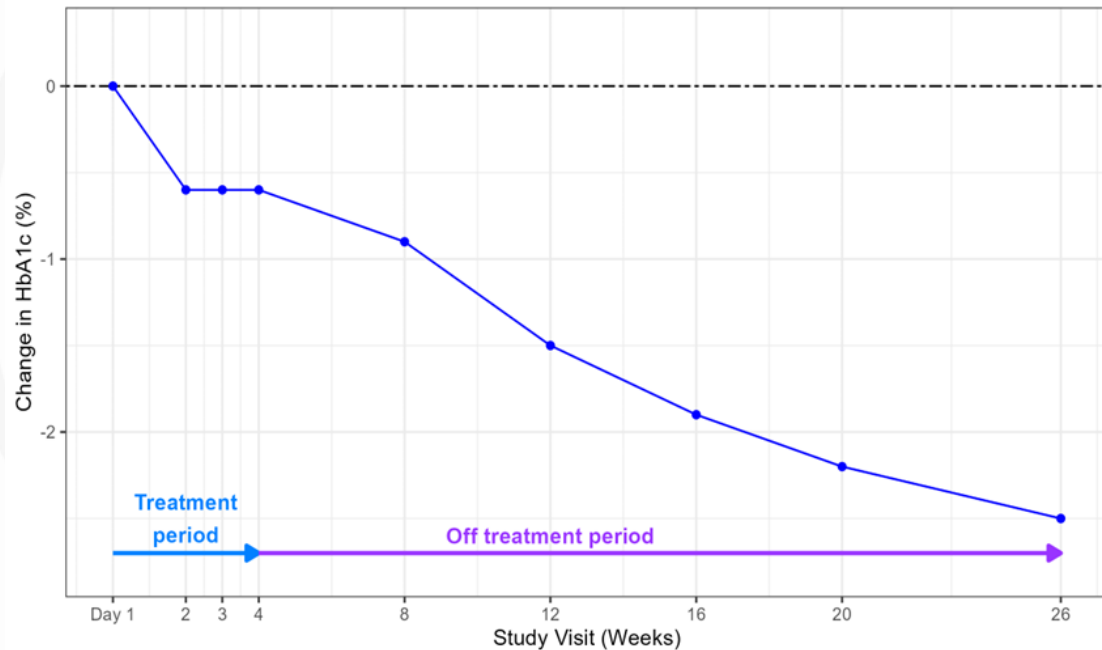
\*Redosing as required at Week 22 for another 4 weeks.

## Case Study 2: 29-year-old man with 4-year history of T2D

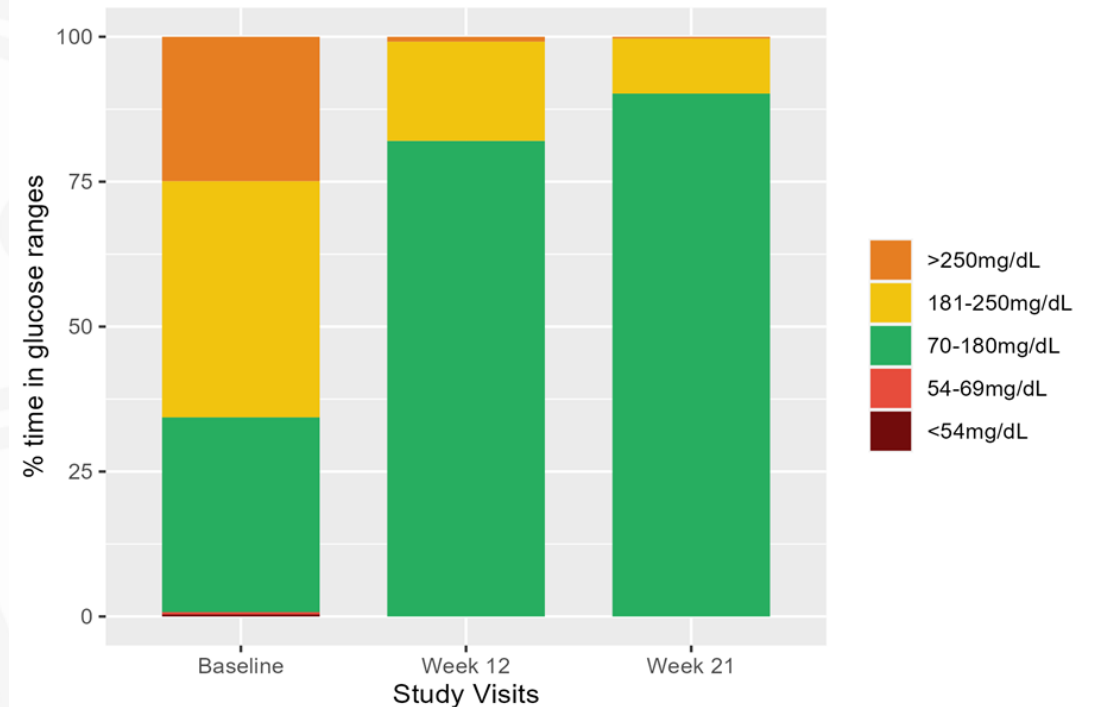
- 29-year-old man with 4-year history of T2D
- Metformin and empagliflozin
- HbA<sub>1c</sub> 9.5%; FPG 146 mg/dL; BMI 25.6 kg/m<sup>2</sup>

- BMF-219 200 mg once daily without food for 4 weeks
- CGM at Week 21 with ~90% TIR<sub>70-180 mg/dL</sub>
- No tolerability issues or related adverse events

### Change in HbA<sub>1c</sub> (%)



### Continuous Glucose Monitoring

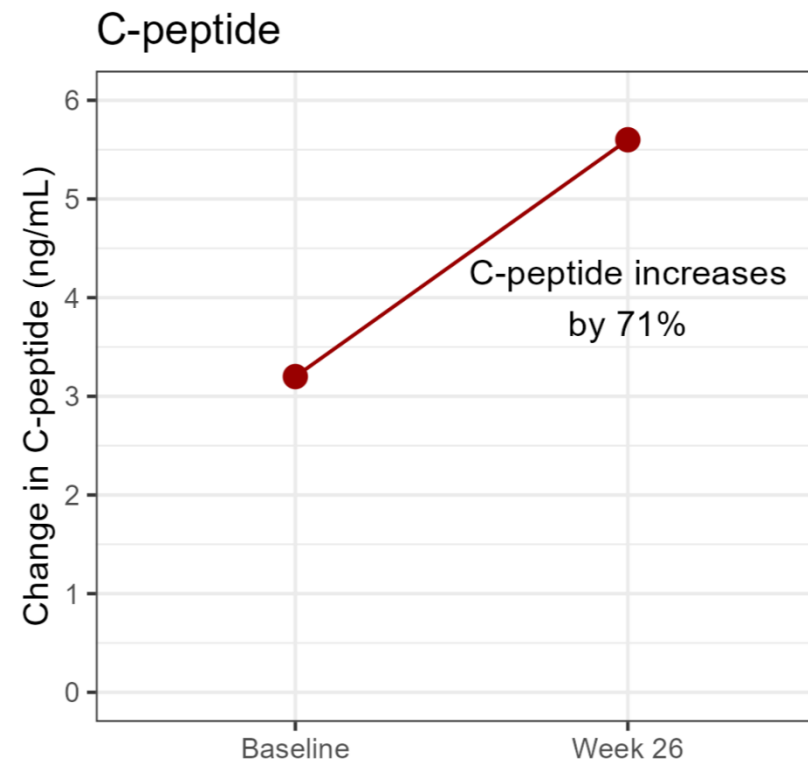
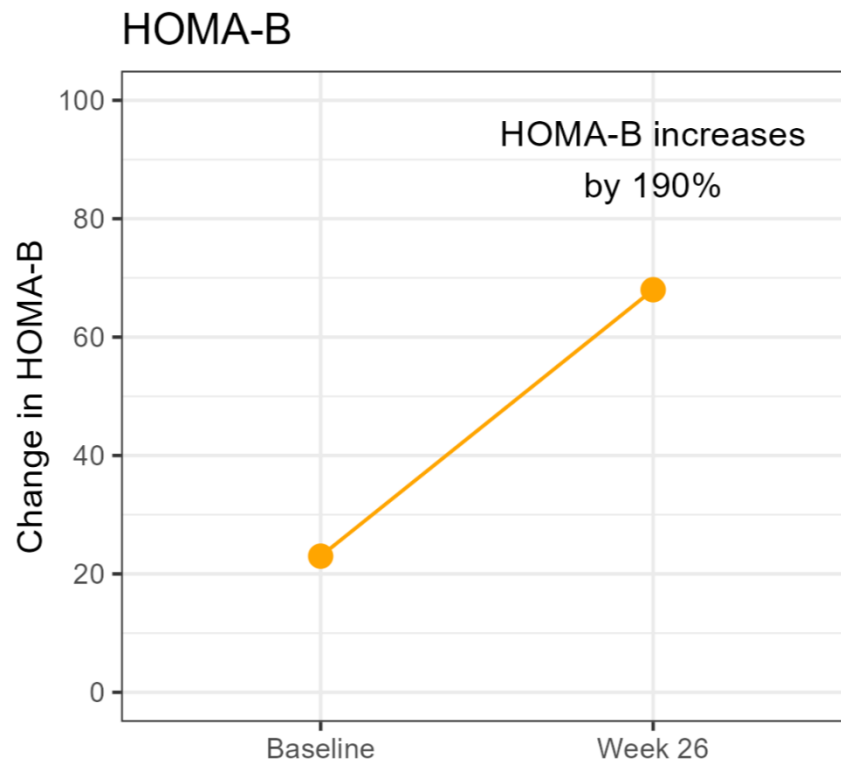


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Change at Week 26



## Summary and Conclusion

- Patients in COVALENT-111 are displaying improved glycemic control while off therapy beyond Week 26 following the 28-day treatment with BMF-219, supporting enhanced beta-cell function as the mechanism of action
- BMF-219 was generally well tolerated with no serious adverse events and no adverse event-related study discontinuations, and no symptomatic or clinically significant hypoglycemia
- 100mg and 200mg dose levels have been selected for the first 3 Arms of the Expansion Phase, which will dose patients up to 12 weeks (compared to 4 weeks in the Escalation Phase) and extended follow-up to Week 52

### Next Steps:

- The Expansion Phase of COVALENT-111 is currently enrolling on schedule.
- Initial 26-week data expected in 2H24

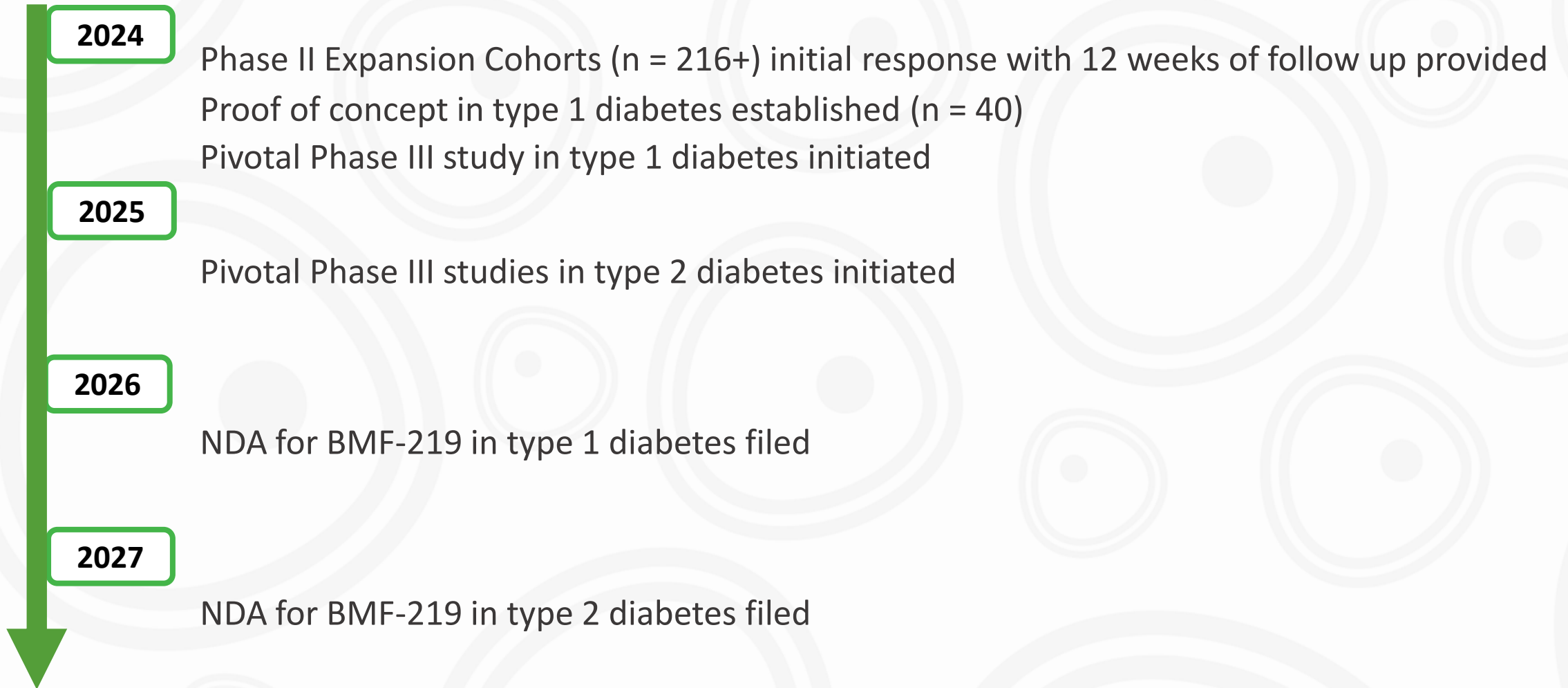
## BMF-219 Induced C-Peptide Increase in the First Two Type 1 Diabetes Patients

Stage 3 Type 1 Diabetes Patients	Patient 1	Patient 2
Baseline Characteristics	58-year-old, diagnosed with type 1 diabetes 3 years ago	24-year-old, diagnosed with type 1 diabetes 7 years ago
Dose Level	BMF-219 200 mg once-daily	BMF-219 100 mg once-daily
Week 4	<ul style="list-style-type: none"> <li>Fasting C-peptide increased by 57% compared to Baseline (study Day 1).</li> <li>During a mixed-meal tolerance test (MMTT) the C-Peptide Index (AUC)<sup>1</sup> increased by 12%.</li> </ul>	<ul style="list-style-type: none"> <li>Fasting C-peptide increased by 16% compared to Baseline.</li> <li>During a MMTT the C-peptide Index (AUC)<sup>1</sup> increased by 30%</li> <li>Near-normal glucose response during the MMTT without receiving any meal-time insulin</li> </ul>
Week 8	<ul style="list-style-type: none"> <li>Fasting C-peptide increased by 80% compared to Baseline.</li> <li>During a MMTT, C-peptide increased up to 200%. The C-Peptide Index (AUC) increased by 40% compared to Baseline</li> </ul>	-
Daily Insulin Usage	Pending	<ul style="list-style-type: none"> <li>Reduction in daily insulin usage during the first four weeks of the study</li> </ul>
Safety	Well Tolerated	Well Tolerated

Data cutoff date: March 7, 2024

<sup>1</sup>C-peptide index is the ratio of serum C-peptide to plasma glucose levels and is used to evaluate beta-cell function

## Next 4 Years BMF-219 in Diabetes





# THANK YOU



Biomea Fusion  
900 Middlefield Road, 4th floor  
Redwood City, CA, 94063  
[biomeafusion.com](http://biomeafusion.com)

