



OBESITY INNOVATION, AT SCALE.

Company Presentation

September 2025

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METSERA: A LEADING CLINICAL-STAGE OBESITY COMPANY

Breadth and capabilities to meet the demands of a potential \$170B consumer market¹

Obesity: Among the largest potential markets in the history of biopharma

- Over \$170B in peak sales projected¹, propelled by consumer demand
- Nutrient-stimulated hormone (NuSH) analogs² in early innings – with major unmet need remaining

Metsera: A next-generation injectable and oral obesity franchise, engineered for scale and market leadership

Multiple clinical programs

- Monthly GLP-1 RA (Ph. 2b)
- Monthly amylin RA (Ph. 1)
- Monthly amylin RA + GLP-1 RA co-administration (Ph. 1)
- Oral GLP-1 RA peptide (Ph. 1)

Capabilities to develop, manufacture, and launch at scale

- Peptide engineering platform
- Translational engine
- Development engine
- Scaled manufacturing

Proprietary library built upon ~20,000 NuSH analog peptides

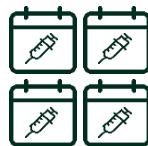
- Injectables and orals
- Miscible and combinable
- Current-gen: GLP-1, GIP
- Next-gen: amylin, GCG, PYY

4 clinical stage programs | 4+ INDs in 2025 | Multiple clinical catalysts over next 6-12 months

A PLATFORM BUILT TO OUTPERFORM FUTURE MARKET LEADERS

Reducing barriers to scale in a potential \$170B+ consumer-driven market¹

FUTURE MARKET LEADERS ARE BEATABLE



Lifelong, weekly injections



Complex, prolonged titration



Tolerability issues
(particularly for orals)



Limited scalability

METSEREA PORTFOLIO: DESIGNED FOR MARKET LEADERSHIP

Monthly dosing
with or without titration

Scalable oral delivery
with injectable-like performance

Multi-NuSH combinations
to optimize weight loss/tolerability

Leading scalability
via potency, half-life, bioA, and dose frequency

METSERA ORIGIN STORY

Three foundational NuSH platforms in a company founded by industry leaders

TWO DEALS PROVIDING THREE FOUNDATIONAL NuSH PLATFORMS



MINT: ~20,000 miscible NuSH analog peptides engineered for differentiated performance

- Current (GLP-1, GIP) and next-generation (amylin, GCG, PYY) targets

HALO™ half-life extension platform

- 2-3x longer than competing peptides in development

MOMENTUM™ platform for scalable oral peptide delivery

- >5x greater oral bioavailability *in vivo* than Rybelsus®
- Versatility to work with multiple NuSH analog peptides



CLIVE MEANWELL
WHIT BERNARD
PETER WIJNGAARD
CHRIS VISIOLI

STEVE MARSO



STEVE BLOOM
JAMES MINNION



BRIAN HUBBARD
MIKE SERRANO-WU

>90,000 patients randomized in Ph. 2 and 3 at <25% the cost of large pharma benchmarks

50,000+ patients randomized in GLP-1 RA outcomes studies

Pioneers in NuSH biology; discovered and engineered ~20,000 peptides

Engineered small molecules targeting APOL1 (Vertex) and PCSK9 (AZ)

OUR PROGRAMS AND CORE DEVELOPMENT STRATEGIES

Four differentiated development strategies

STRATEGY	PROGRAM Target / Mechanism	STAGE OF DEVELOPMENT					ANTICIPATED MILESTONES
		DISCOVERY	IND / CTA- ENABLING	PHASE 1	PHASE 2	PHASE 3	
FULLY BIASED, MONTHLY GLP-1 RA	MET-097i Fully Biased GLP-1 RA			Phase 2b ongoing			VESPER-1 preliminary readout September 2025 ¹ ; VESPER-3 preliminary readout YE 2025 / early 2026; VESPER-2 preliminary read-out early 2026
MONTHLY AMYLIN ANALOG + GLP-1 RA	MET-233i Amylin Analog			Phase 1/2a ongoing			Twelve-week preliminary read-out late 2025
	MET-233i + MET-097i Amylin Analog + Fully Biased GLP-1 RA			Phase 1/2a ongoing			Twelve-week preliminary readout YE 2025 / early 2026
ORAL PEPTIDE PLATFORM (MOMENTUM™)	MET-097_o / MET-224_o Fully Biased GLP-1 RAs		IND-enabling studies ongoing				Four-week preliminary readout of lead oral late 2025, after completion of IND-enabling studies and if successful in initiating study
	MET-002_o GLP-1 RA			Phase 1 ongoing			
NEXT-GENERATION PROGRAMS	MET-034i GIP RA		IND-enabling studies ongoing				Preliminary tolerability readout late 2025, if successful in initiating study
	MET-067i Glucagon Analog		IND-enabling studies ongoing				
	MET-815i MET-097i Prodrug		IND-enabling studies ongoing				

MET-097i

Monthly, fully-biased
GLP-1 RA and
GLP-1 RA prodrug

MET-233i

Monthly amylin analog
and multi-NuSH
combinations

ORAL NuSH PEPTIDE PLATFORM

Engineered for
scalability and
injectable-like
performance

STRATEGIC MANUFACTURING

Scaling to profitably
serve a potential
\$170B+ consumer-
driven market¹

2025 CATALYSTS

Key milestones in next
6-12 months

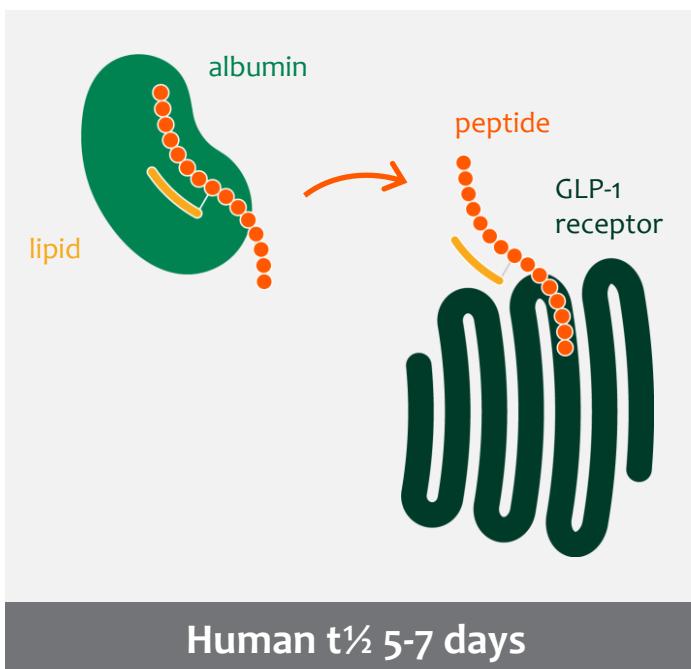
HALO™: ENGINEERING A STEP-CHANGE IN PEPTIDE PERFORMANCE

HALO™ platform generated the two longest-acting clinical-stage therapeutic peptides known

LYS-26-LIPIDATED PEPTIDE

Semaglutide, tirzepatide

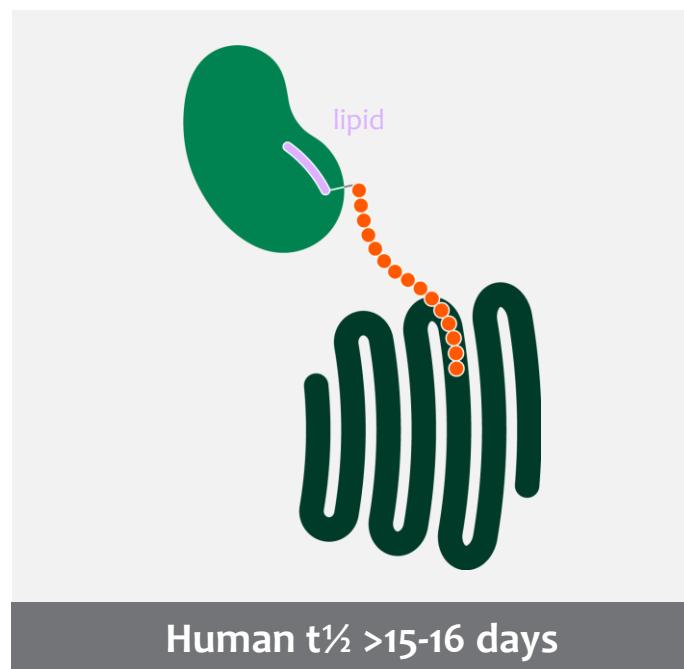
Peptide must dissociate from albumin
to bind GLP-1 receptor



HALO™ LIPIDATED PEPTIDE

MET-097i, MET-233i, Metsera orals

Peptide binds to GLP-1 receptor with or
without dissociation from albumin



BENEFITS OF LONG HALF-LIFE

Potentially afforded by Metsera's
proprietary HALO™ platform

Convenient dosing regimens
QM dosing and titration-free dosing

Improved scalability
More effect with less API

Improved tolerability
Lower peak-to-trough variability

Proprietary iterative engineering
Difficult to mimic

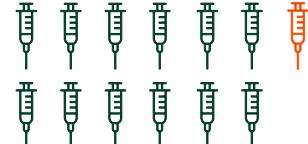
MET-097i: A MONTHLY, FULLY-BIASED GLP-1 RA

Engineered for class-leading durability, potency, and scalability

ATTRIBUTES	POTENTIAL IMPLICATIONS
1 Durability	Titration-free weekly dosing and monthly maintenance dosing
2 Potency	Improved scalability due to lower dosage requirements
3 Full Receptor Bias	Dual-agonist like body weight loss
4 BLA Eligibility	Expanded market exclusivity and longer exclusion from IRA negotiations

MET-097i PHASE 1/2 PROGRAM OVERVIEW

A randomized, placebo-controlled Phase 1/2 trial with three parts

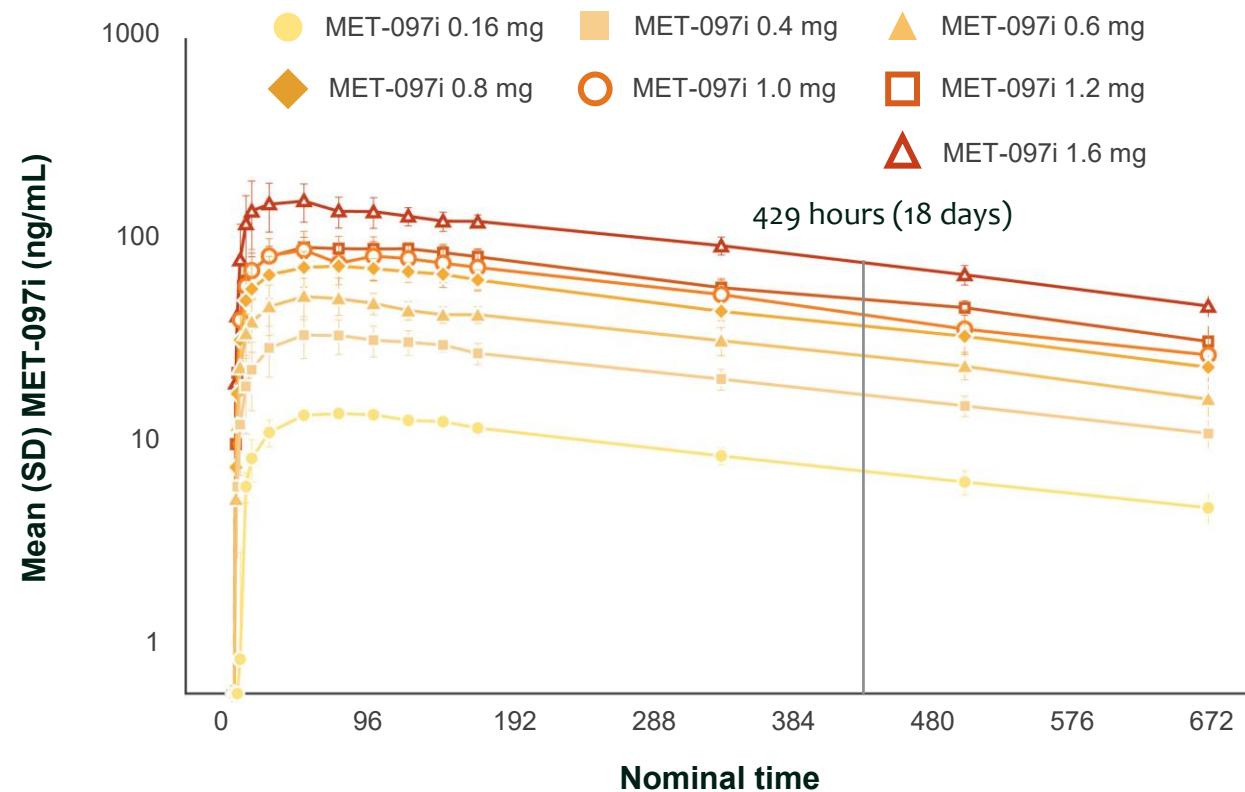
	SAD (Part A)	MAD (Part B)	12-week (Part C)
Objectives	Safety, tolerability, PK, weight loss		<ul style="list-style-type: none"> Weight loss and tolerability through 12 weeks Tolerability of QM switch
Participants		Overweight / obese without T2DM	
Design	N=60	N=62	N=120
	7 cohorts X 	6 cohorts X 	5 cohorts X 
			



MET-097i: THE LONGEST-ACTING NuSH PEPTIDE IN DEVELOPMENT

Half-life potentially enables monthly dosing and titration-free, intermittent dosing

MET-097i OBSERVED HALF-LIFE¹ ~18 DAYS IN SAD



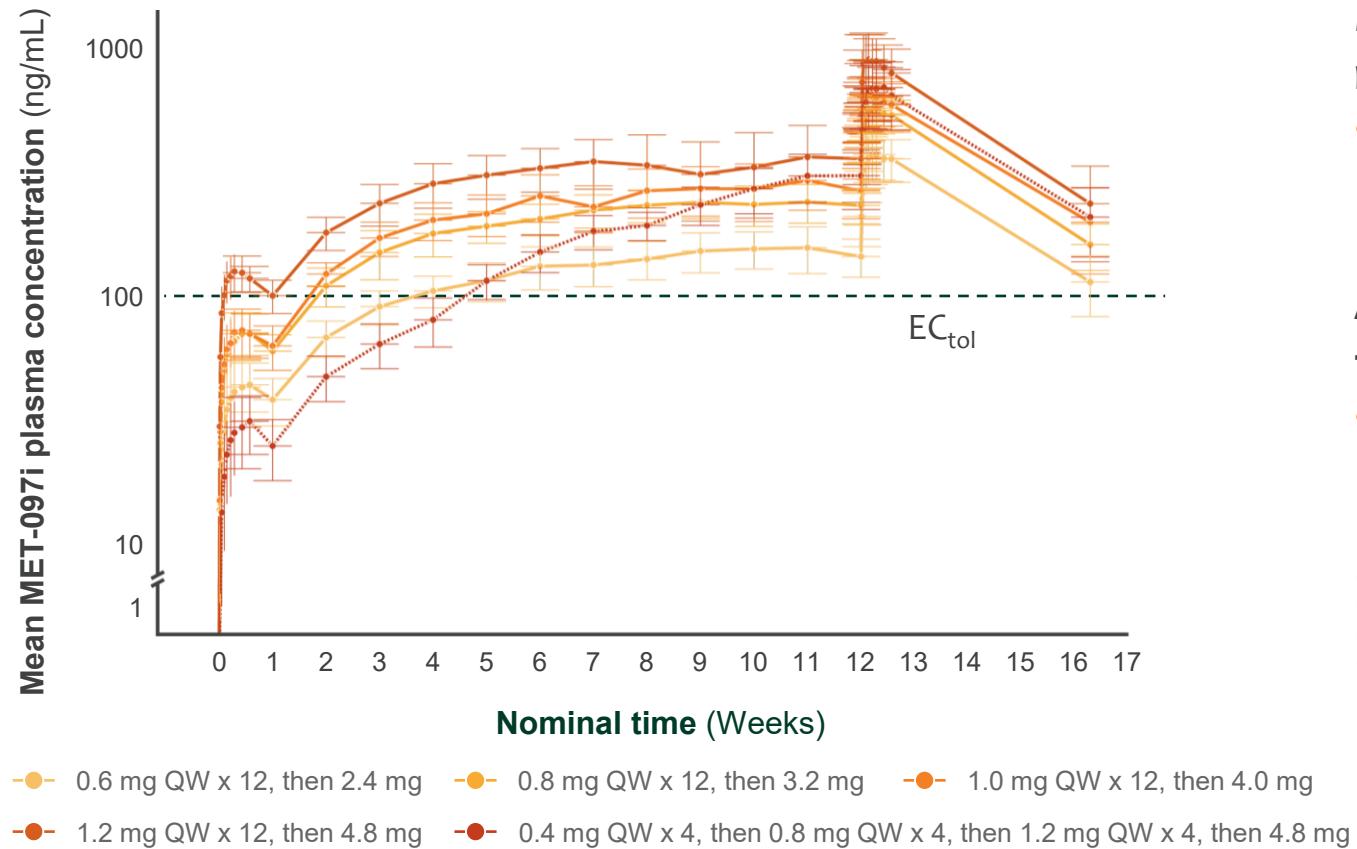
HALF-LIFE IN COMPETITIVE CONTEXT

Asset	Target	Terminal T _{1/2} (hr) ²
Native GLP-1	GLP-1	Minutes
Liraglutide	GLP-1	13
Tirzepatide	GLP-1, GIP	120
Retatrutide	GLP-1, GIP, GCG	144
CT-388	GLP-1, GIP	150
Semaglutide	GLP-1	168
Cagrelinotide	Amylin	159-195
VK-2735	GLP-1, GIP	~170-250
Petrelintide	Amylin	230
ZT-002	GLP-1	260-273
MET-097i	GLP-1	~380
AMG-133 (MariTide)	GLP-1, GIP antag.	343-396 ³

ACCUMULATION WITH MULTIPLE WEEKLY DOSES

Gradual accumulation may enable optimized tolerability and streamlined titration before QM switch

SMOOTH INCREASE IN EXPOSURE, CROSSING TOLERABILITY THRESHOLD



MET-097i accumulated ~4-fold with repeat weekly dosing

- Accumulation escalated exposure beyond tolerability threshold

Accumulation particularly gradual in titrated cohort

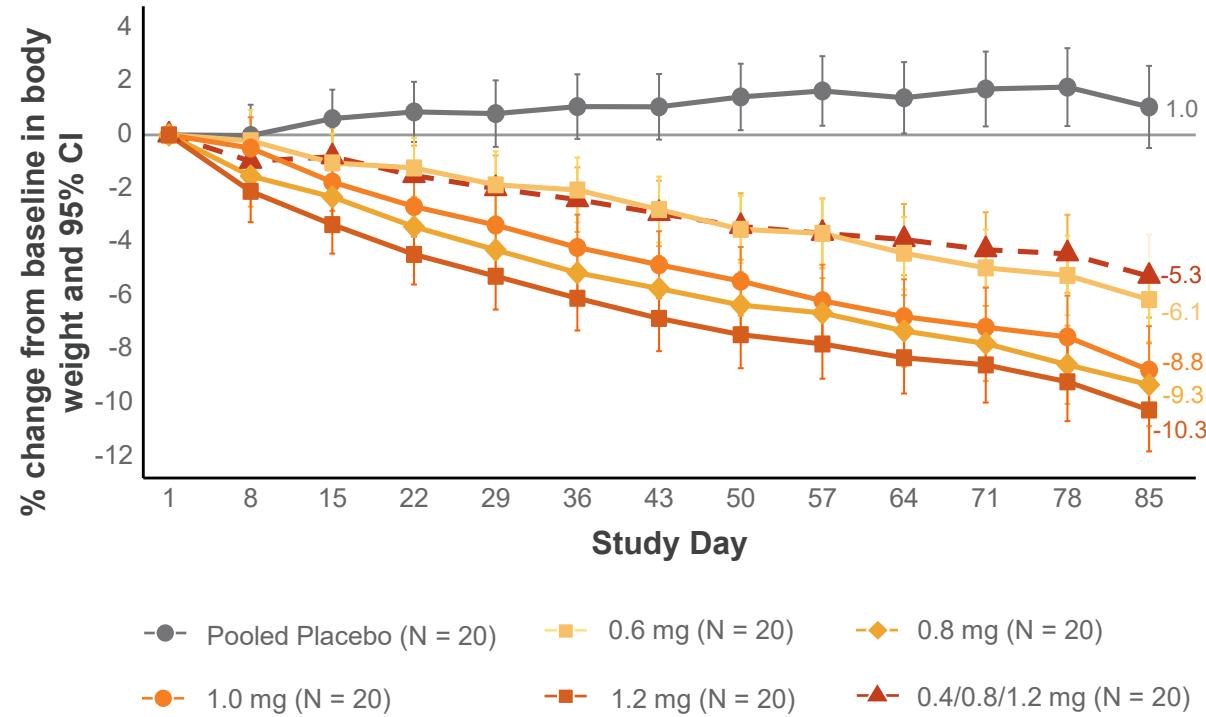
- Potential for differentiated tolerability with simplified titration

Blunted trough-to-peak upon switch to PK-matched QM dose

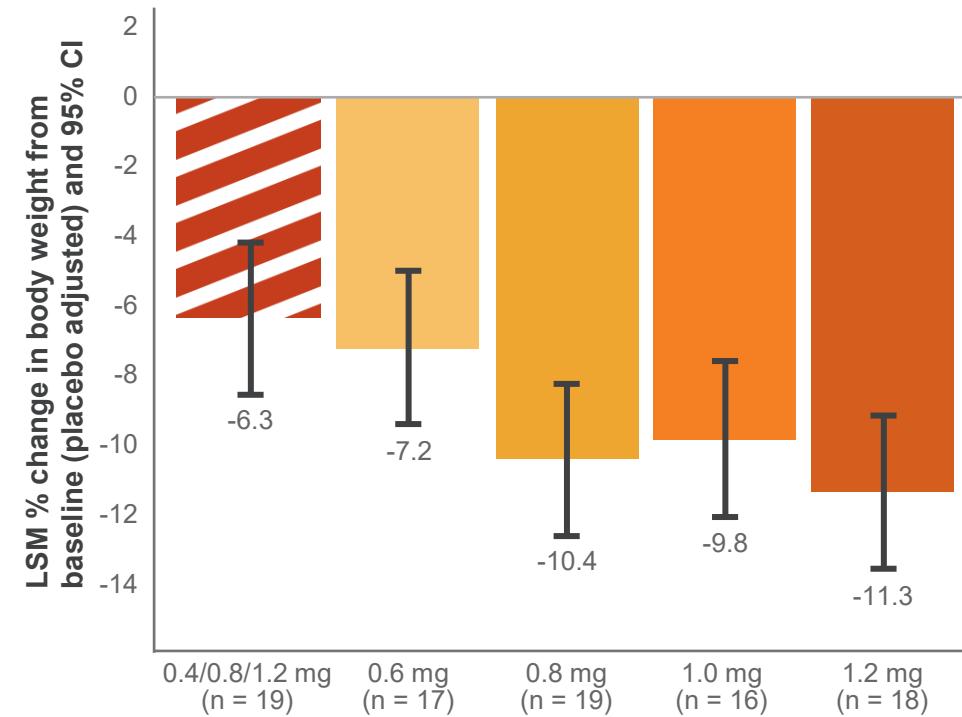
12-WK WEIGHT LOSS AT HIGH END OF COMPETITIVE LANDSCAPE

Consistent with approved and development-stage dual agonists

MEAN % CHANGE FROM BASELINE IN BODY WEIGHT ACROSS COHORTS



MEAN PLACEBO-SUBTRACTED % CHANGE IN BODY WEIGHT AT DAY 85



COMPETITIVE TOLERABILITY AT 12 WEEKS

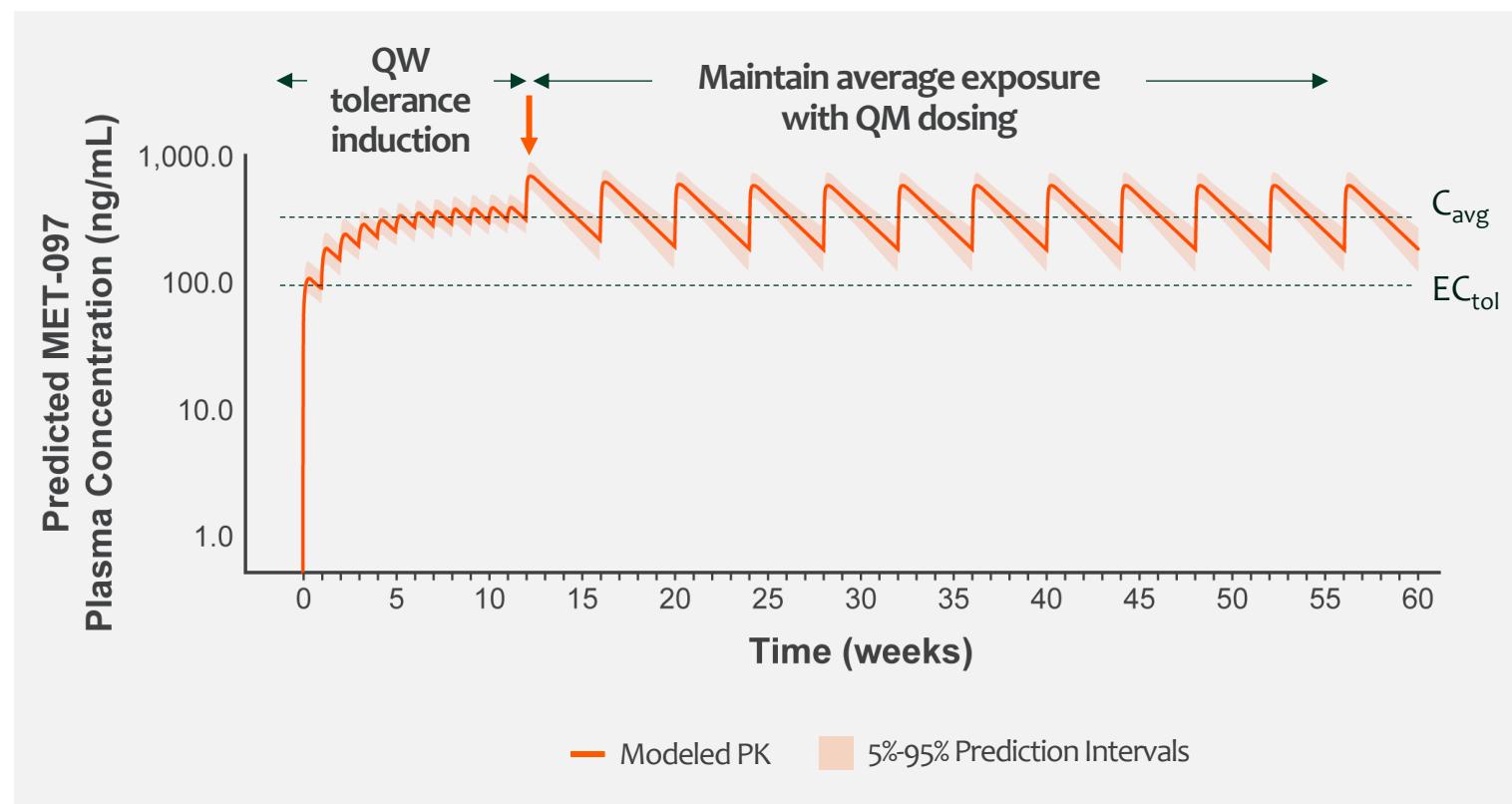
Gastro-intestinal AEs mostly mild – titrated MET-097i particularly well-tolerated

	Pooled Placebo	MET-097i				
		0.4/0.8/1.2 mg N = 20 n(%)	0.6 mg N = 20 n(%)	0.8 mg N = 20 n(%)	1.0 mg N = 20 n(%)	1.2 mg N = 20 n(%)
At Least One GI AE	6 (30.0)	3 (15.0)	12 (60.0)	11 (55.0)	14 (70.0)	17 (85.0)
Nausea	3 (15.0)	1 (5.0)	6 (30.0)	8 (40.0)	10 (50.0)	13 (65.0)
Mild	3 (15.0)	1 (5.0)	6 (30.0)	7 (35.0)	8 (40.0)	10 (50.0)
Moderate	0	0	0	1 (5.0)	2 (10.0)	3 (15.0)
Severe	0	0	0	0	0	0
Vomiting	1 (5.0)	2 (10.0)	6 (30.0)	4 (20.0)	8 (40.0)	12 (60.0)
Mild	0	2 (10.0)	6 (30.0)	3 (15.0)	7 (35.0)	7 (35.0)
Moderate	1 (5.0)	0	0	1 (5.0)	1 (5.0)	5 (25.0)
Severe	0	0	0	0	0	0
Diarrhea	2 (10.0)	0	4 (20.0)	1 (5.0)	4 (20.0)	2 (10.0)

MET-097i PHARMACOKINETICS SUPPORT MONTHLY DOSING

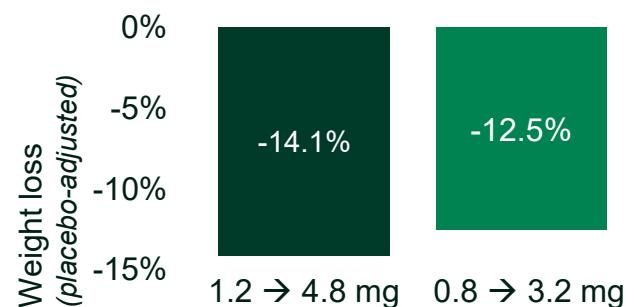
Exposure-matched monthly dose efficacious and well-tolerated at week 16

**OBJECTIVE: INDUCE TOLERANCE WITH WEEKLY DOSE,
MAINTAIN EXPOSURE WITH MATCHED MONTHLY DOSE**



MONTHLY SWITCH DE-RISKED

Continued weight loss 4 weeks after monthly switch (Week 16)¹



Monthly switch well-tolerated²

Week	Vomiting	Nausea
Week 1	16%	34%
Week 13 (2x)	5%	14%
Week 13 (4x)	11%	28%

MET-097i PHASE 2 PROGRAM

Three Phase 2b trials designed to confirm MET-097i's profile and enable rapid Phase 3 initiation

TRIAL	OBJECTIVE	DESIGN	TIMING
Vesper ⁺ 1	Select doses for Phase 3	<ul style="list-style-type: none"> ~225 participants without T2DM Four arms between 0.4 mg and 1.2 mg Dosing without titration 	September 2025 ¹
Vesper ⁺ 2	Evaluate MET-097i in people with overweight / obesity and T2DM	<ul style="list-style-type: none"> ~125 participants with T2DM Four arms between 0.4 mg and 1.6 mg Dosing with and without titration 	Early 2026
Vesper ⁺ 3	Assess weight loss and tolerability after multiple monthly doses	<ul style="list-style-type: none"> ~250 participants without T2DM Four arms receiving 12 weekly doses followed by multiple monthly doses Dosing with and without titration 	Year-end 2025 / early 2026

MET-097i: A HIGHLY DIFFERENTIATED NUSH PROFILE EMERGING

Monthly dosing, tolerability, versatility demonstrated in early trials; significant progress towards BLA

DATA TO DATE CONFIRM COMPETITIVE PROFILE

✓ Monthly dosing de-risked

- Well-tolerated and weight loss 14.1% four weeks after dose¹
- Unlocks exposure-matched monthly dosing

✓ Potentially class-leading tolerability

- Dose escalation with only one case of nausea and two cases of vomiting (n=20)

✓ Weight-loss at high end of competitive landscape

- 11.3% 12-week placebo-adjusted weight loss competitive with leading approved and development-stage dual-agonists²

✓ Titration-free dosing confirmed

- 10.4% placebo-adjusted weight loss with competitive tolerability on a single, fixed dose of 0.8mg/week

UPCOMING MILESTONES

Mid 2025



VESPER-1
28-week Ph. 2b
weekly dosing
(September)

Late 2025



VESPER-3
Ph. 2b monthly dosing
(end 2025 / early 2026)



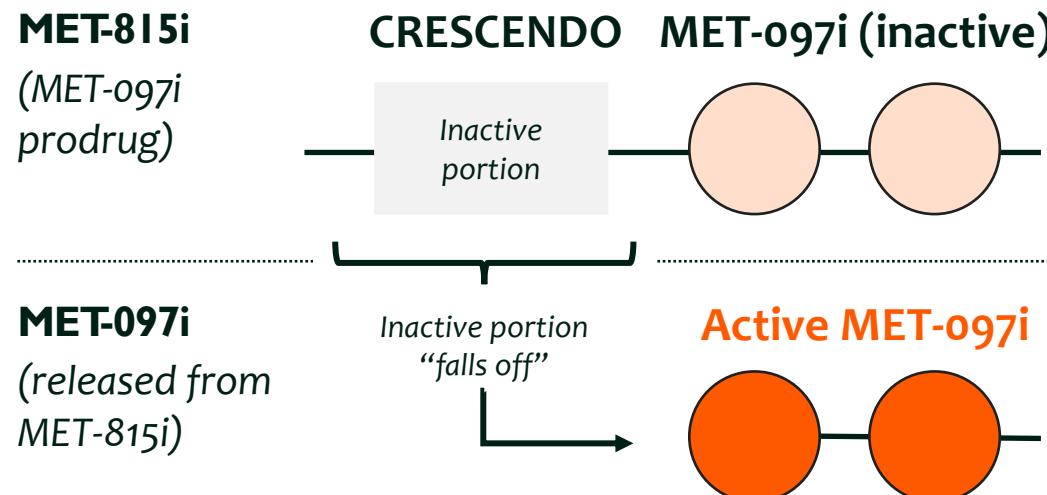
Phase 3 initiation³

CRESCENDO PRODRUG PLATFORM

Potential to reduce dosing frequency to less than monthly

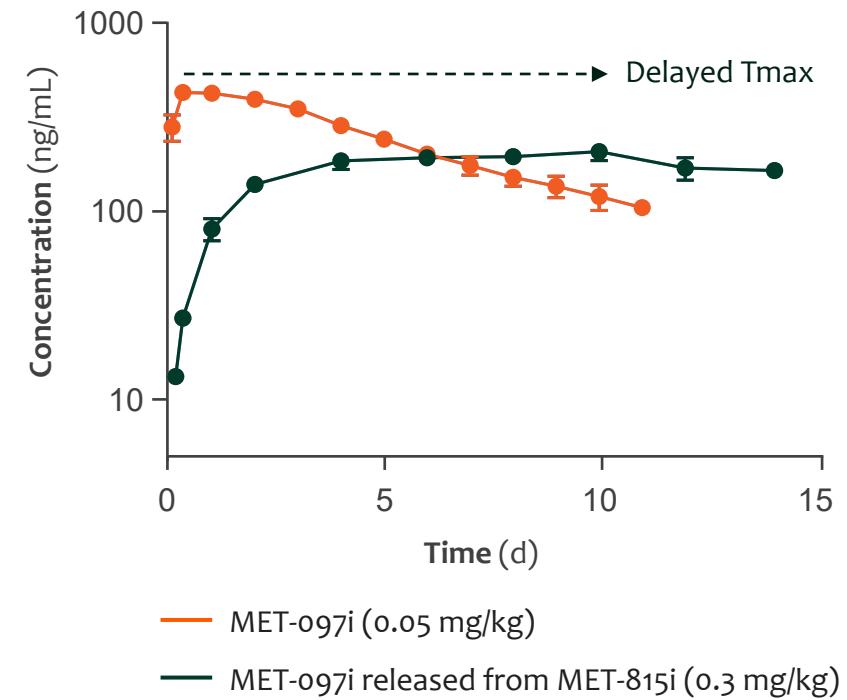
CRESCENDO GRADUAL RELEASE

CRESCENDO enables production of biologically-inactive prodrugs with non-enzymatic release and tunable kinetics



MET-815i GRADUALLY RELEASES MET-097i

Mini-pig pharmacokinetics



MET-097i

Monthly, fully-biased
GLP-1 RA and
GLP-1 RA prodrug

MET-233i

Monthly amylin analog
and multi-NuSH
combinations

ORAL NuSH PEPTIDE PLATFORM

Engineered for
scalability and
injectable-like
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STRATEGIC MANUFACTURING

Scaling to profitably
serve a potential
\$170B+ consumer-
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2025 CATALYSTS

Key milestones in next
6-12 months

MET-233i: A MONTHLY AMYLIN ANALOG

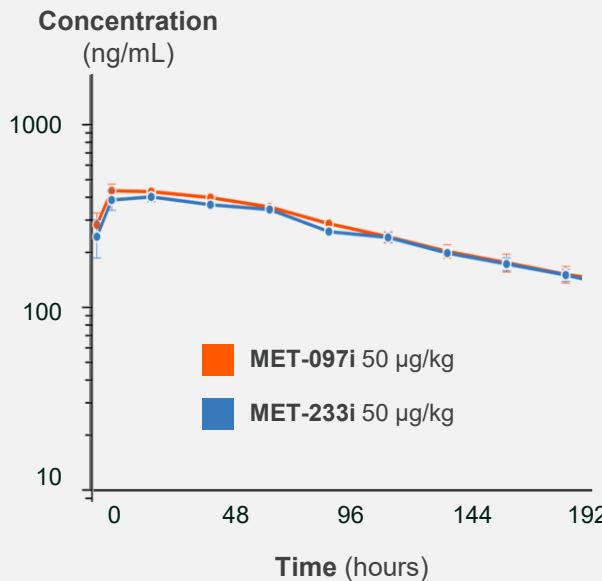
Engineered for durability, potency, and combinability with MET-097i

DURABLE



Supports monthly dosing

Exposure over time after a single dose of Metsera NuSH peptides in pigs

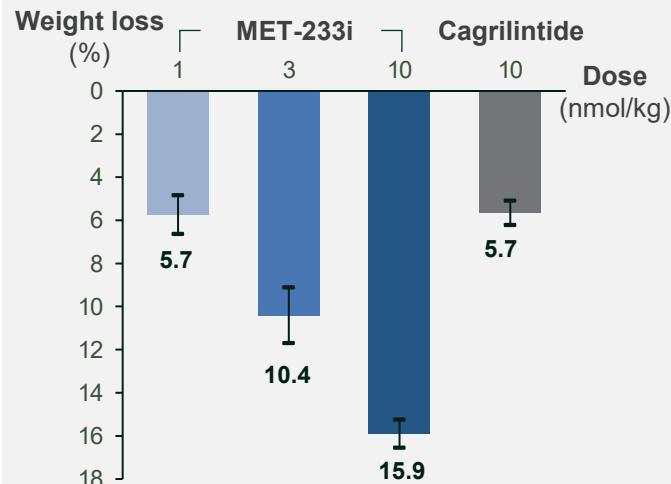


POTENT

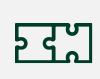


Scalable due to low API needs

Body weight effects at Day 3 after a single dose in rats

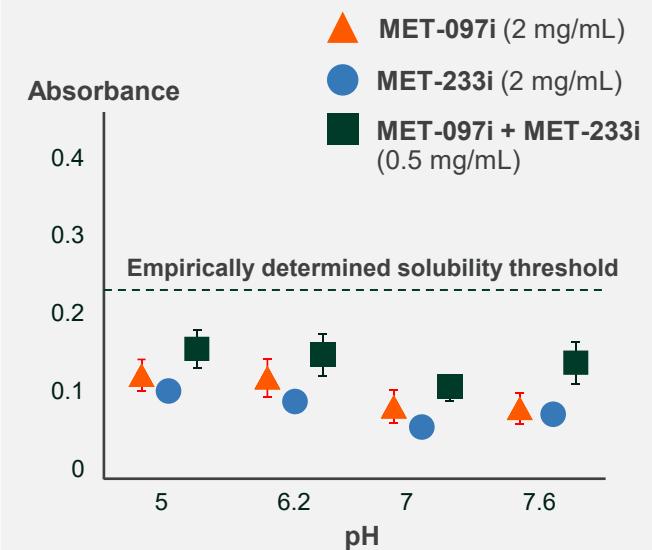


COMBINABLE



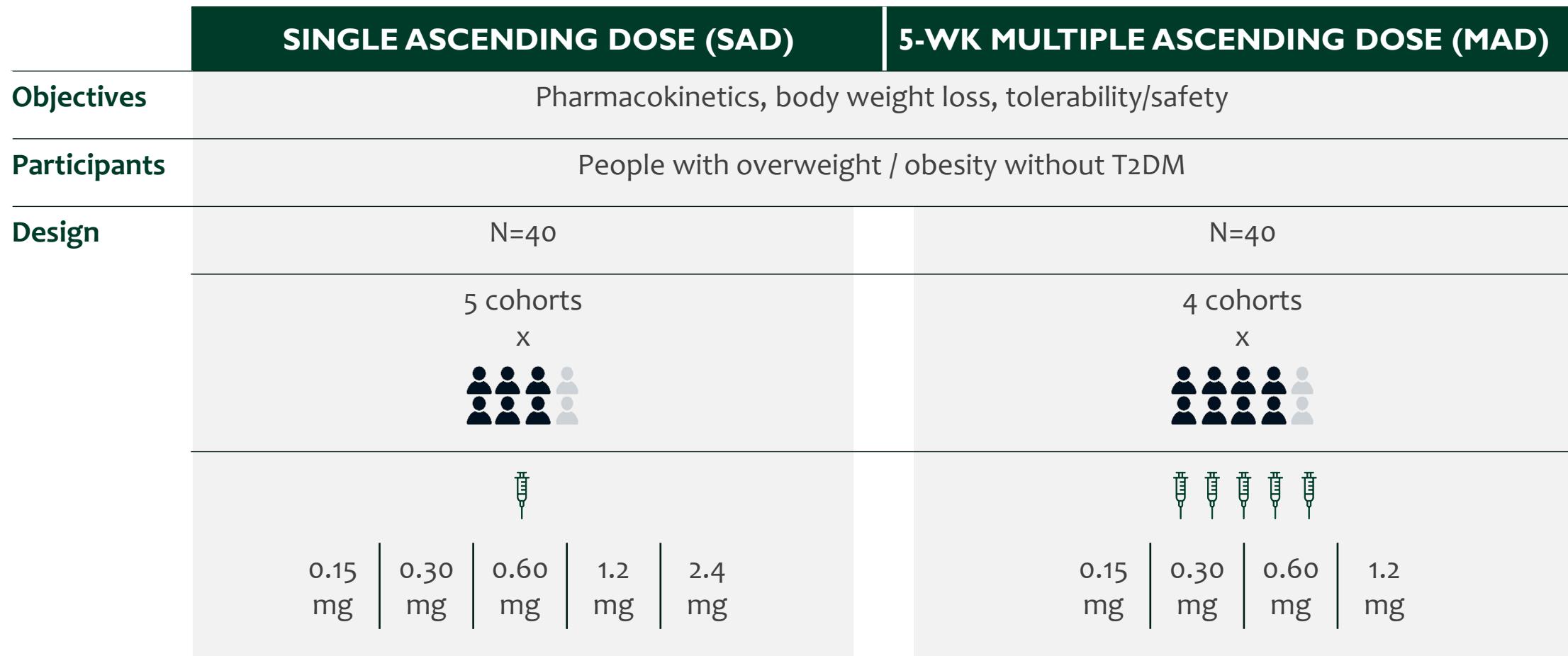
Miscible with MET-097i (GLP-1RA)

Solubility of Metsera NuSH peptides and combination at different pH



MET-233i PHASE I CLINICAL TRIAL

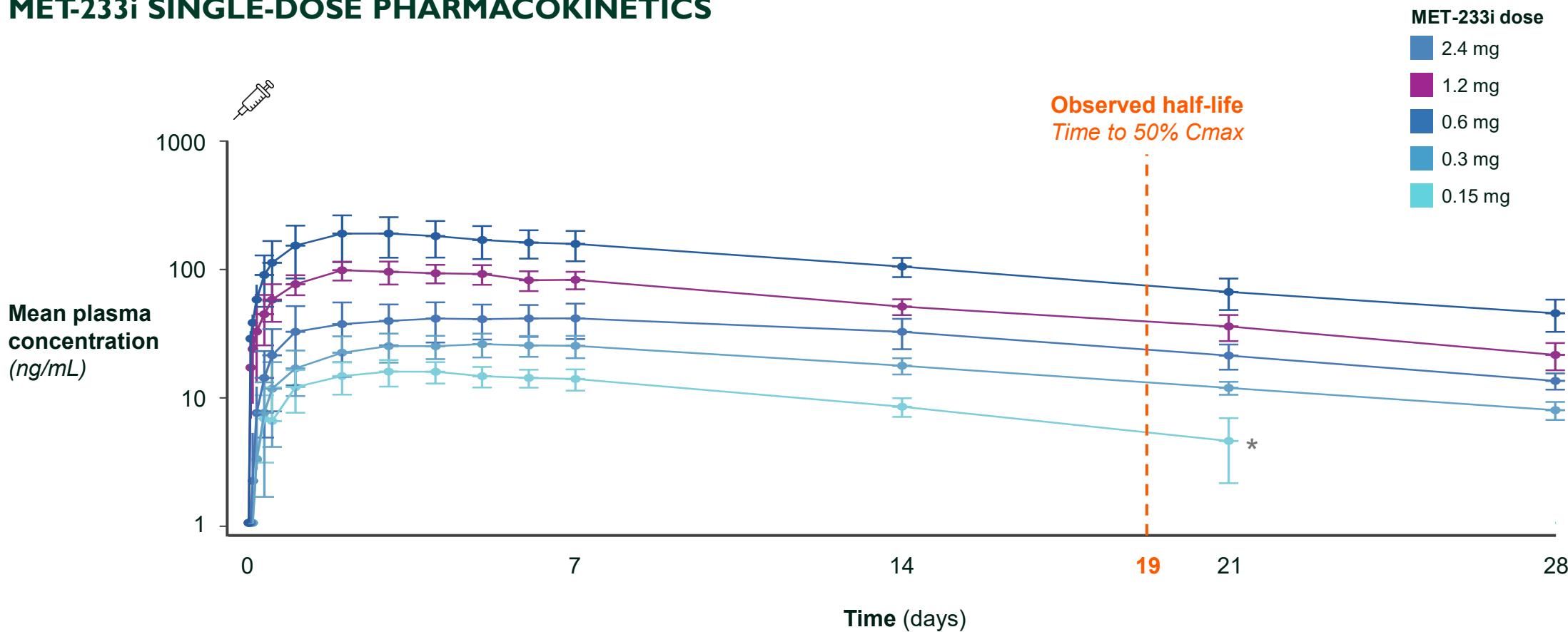
A randomized, placebo-controlled trial in people with overweight / obesity



19-DAY OBSERVED HALF-LIFE SUPPORTS ONCE-MONTHLY DOSING

Dose-linear pharmacokinetics with low variability

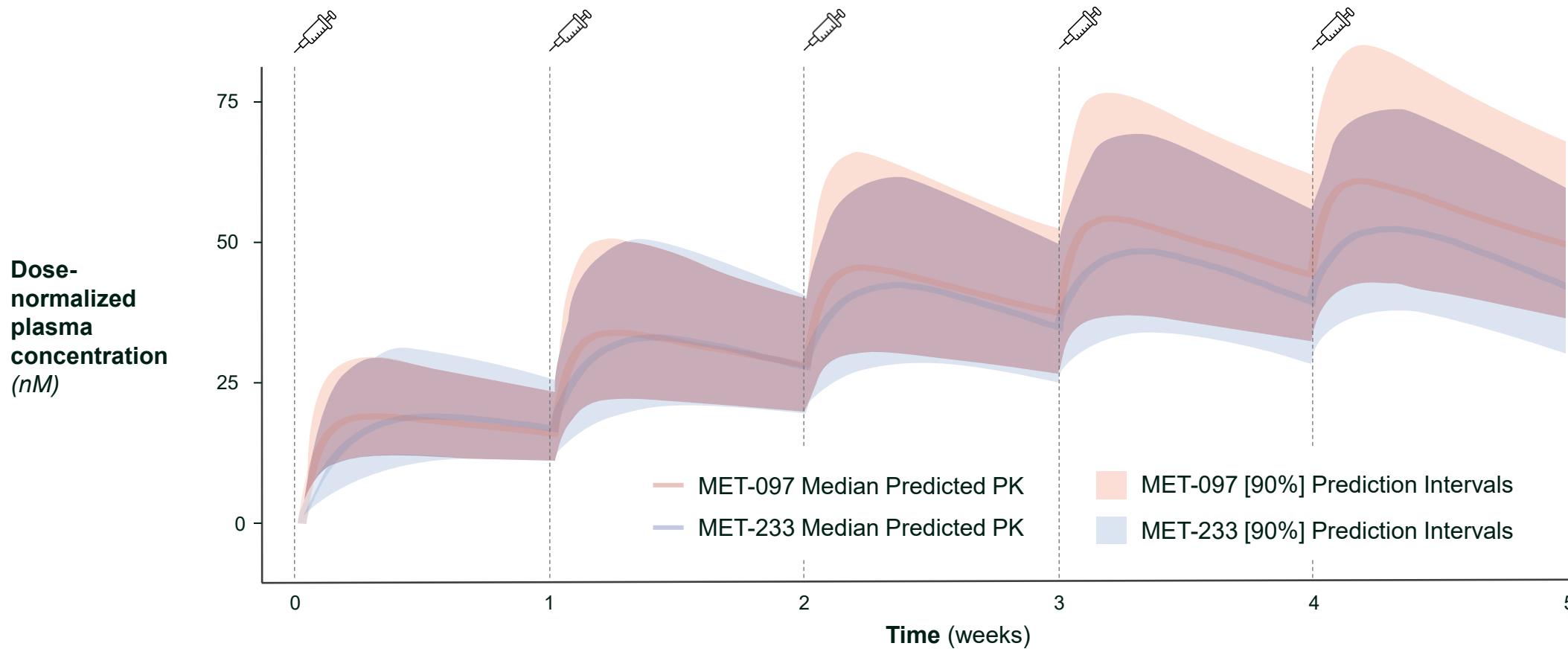
MET-233i SINGLE-DOSE PHARMACOKINETICS



MULTI-DOSE PK ENABLES COMBINABILITY WITH MET-097i

Pharmacokinetics support a potential first-in-class monthly multi-NuSH combination

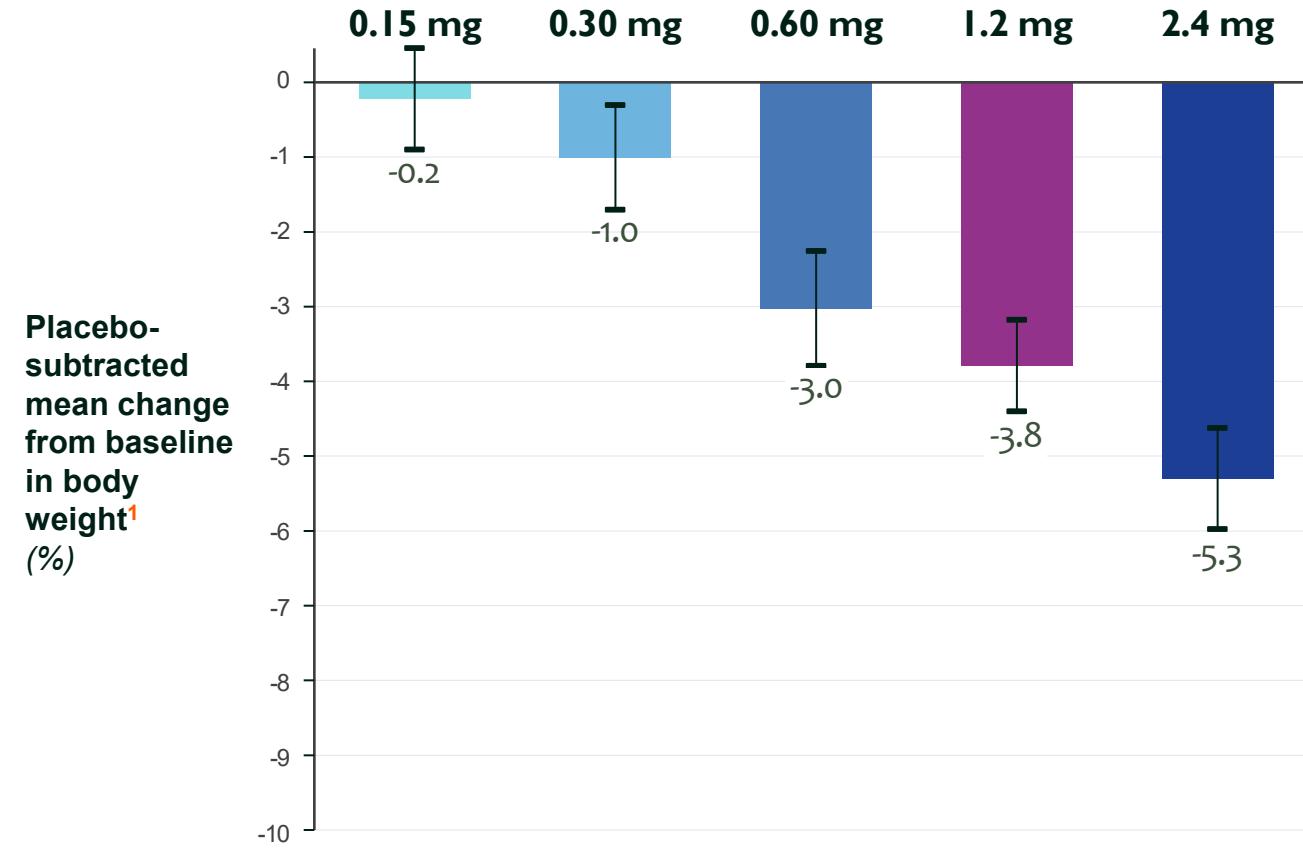
MULTI-DOSE PHARMACOKINETICS OF MET-233i AND MET-097i: 88% OVERLAP



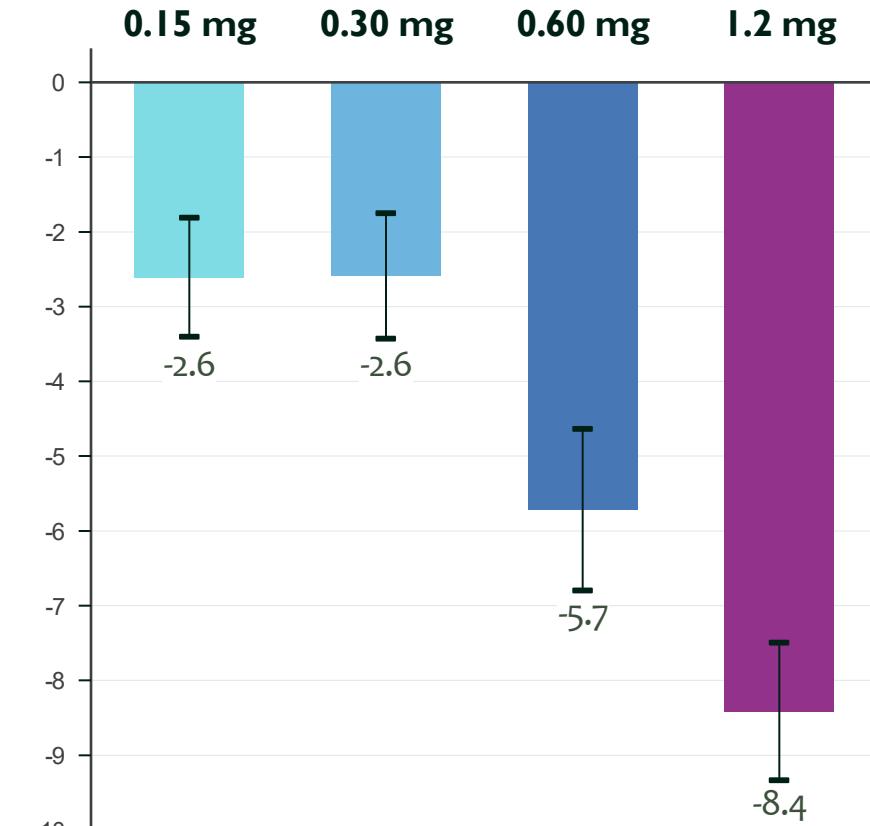
MET-233i SINGLE AND MULTIPLE-DOSE WEIGHT LOSS

Up to 8.4% placebo-subtracted mean weight loss after five doses

SAD: BODY WEIGHT CHANGE AT DAY 8



MAD: BODY WEIGHT CHANGE AT DAY 36



MET-233i WELL-TOLERATED

MAD: All gastrointestinal adverse events mild; starting doses with placebo-like tolerability

ONSET OF GASTROINTESTINAL ADVERSE EVENTS BY WEEK IN MAD

		NAUSEA				VOMITING				EXPOSURE	
		Placebo	MET-233i			Placebo	MET-233i			MET-233i drug exposure level relative to Week 1	
			0.15 mg	0.3 mg	0.6 mg	1.2 mg		0.15 mg	0.3 mg	0.6 mg	1.2 mg
Week	N size	8	8	8	8	8	8	8	8	8	
1		1 (12.5%)	1 (12.5%)	1 (12.5%)	6 (75.0%)	8 (100%)	0	1 (12.5%)	0	3 (37.5%)	3 (37.5%)
2		0	0	0	0	0	0	0	0	0	0
3		0	0	0	0	0	0	0	0	0	0
4		0	0	0	0	1 (14.3%)	0	0	0	0	0
5		1 (12.5%)	0	2 (25.0%)	0	1 (16.7%)	0	0	0	0	0
Total		1 (12.5%)	1 (12.5%)	2 (25.0%)	6 (75.0%)	8 (100%)	0	1 (12.5%)	0	3 (37.5%)	3 (37.5%)

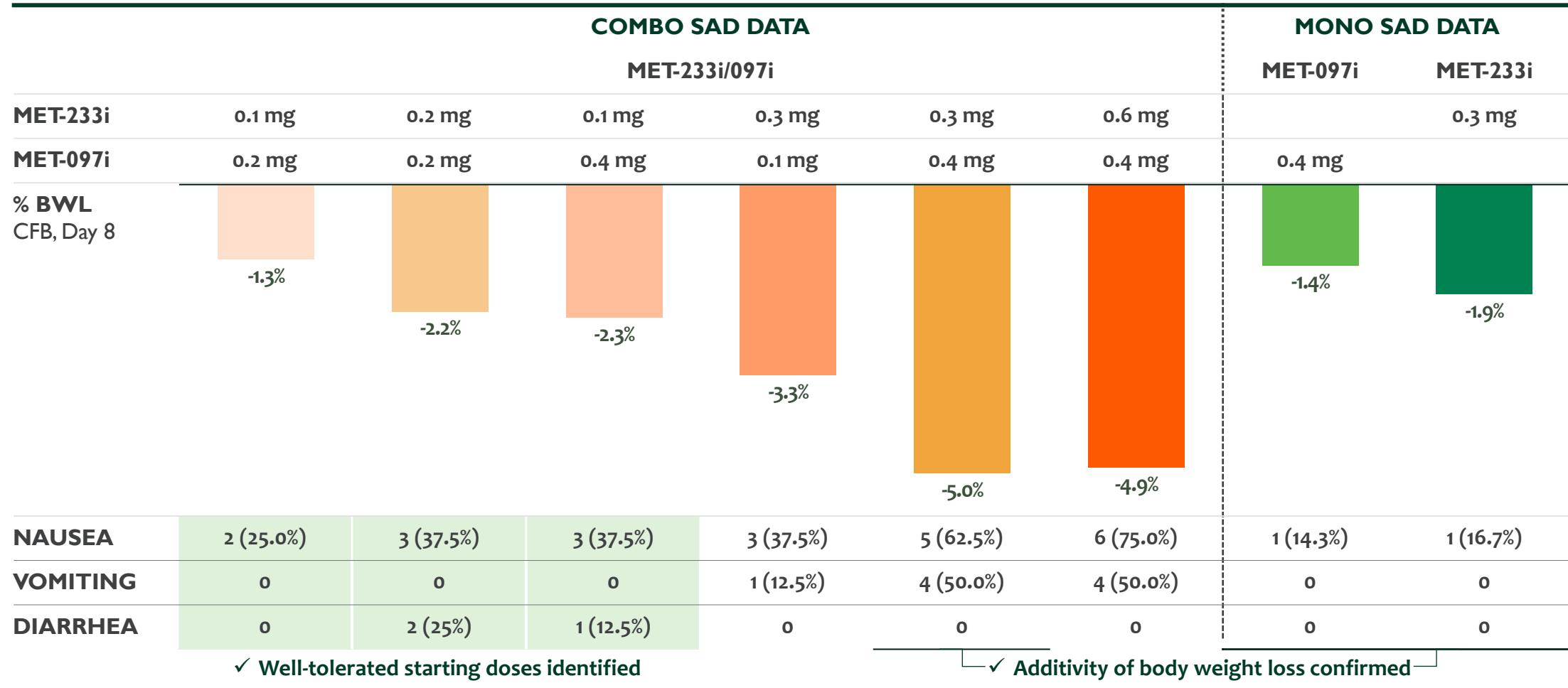
└ Candidate
starting doses ┘

└ Candidate
starting doses ┘

No safety signals.

MET-233/097i SINGLE ASCENDING DOSE (SAD)

Starting doses identified with excellent tolerability and efficacy



MET-233i: A MONTHLY AMYLIN ANALOG

MET-233i mono and MET-233i + MET-097i co-admin studies underway; multiple 2025 datasets expected

AN AMYLIN ANALOG WITH POTENTIAL BEST-IN-CLASS ATTRIBUTES

- ✓ **Pharmacokinetics supporting monthly dosing**
 - 19-day observed half-life
 - Matched to MET-097i
- ✓ **Potentially class-leading potency**
 - 8.4% weight loss after five doses of MET-233i
 - Substantial weight loss after a single MET-233/097i dose
 - Additivity of weight loss established
- ✓ **Excellent tolerability**
 - Placebo-like tolerability at anticipated MET-233i starting doses
 - Excellent tolerability after single MET-233/097i dose
 - All GI AEs in MAD were mild

KEY MILESTONES

Q2 2025

- ✓ **MET-233i monotherapy data**
5-week weight loss and tolerability

Late 2025

- 🚩 **MET-233i monotherapy data**
12-week weight loss and tolerability

- 🚩 **MET-233i / MET-097i co-administration data (End 2025 / Early 2026)**

12-week weight loss and tolerability

MONTHLY TRI- AND QUAD-AGONIST DEVELOPMENT

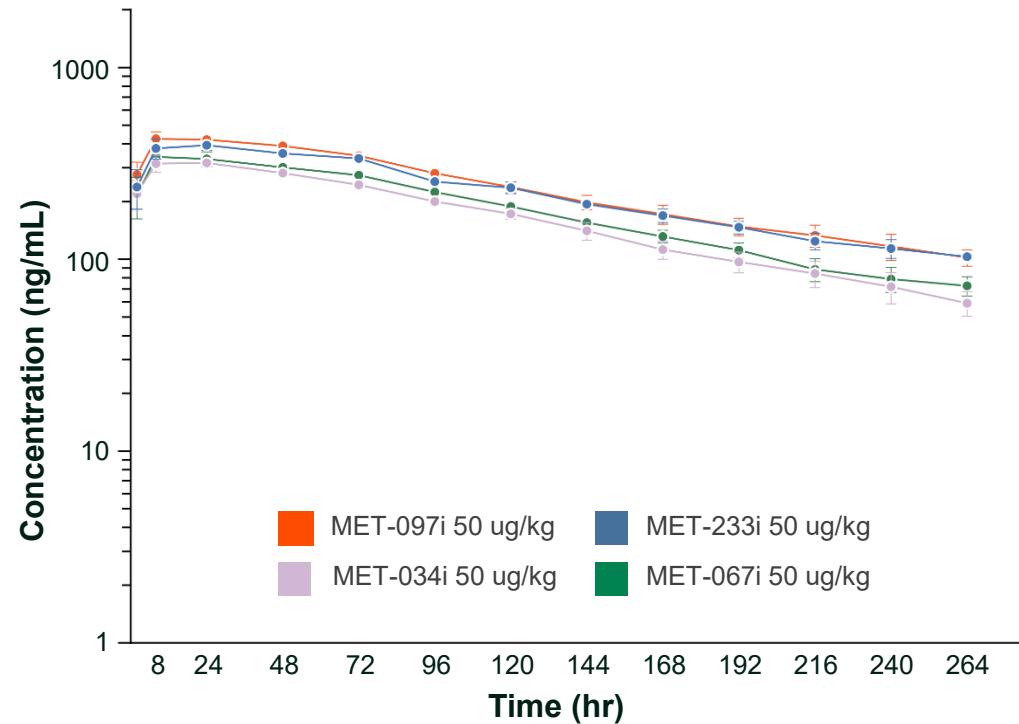
Miscible building blocks to further optimize the relationship between tolerability and efficacy

ULTRA-LONG ACTING QUAD-AGONIST COULD IMPROVE EFFICACY / TOLERABILITY RATIO

Compound	Target	Rationale
MET-034i	GIP	<p>↑↑↑ GI tolerability</p> <p>↑ WL efficacy</p> <p>↑ HbA1C efficacy</p>
MET-067i	Glucagon	<p>↑ WL efficacy</p> <p>↑ MASH efficacy</p> <p>No tolerability cost</p>

PK OBSERVED TO BE SIMILAR FOR THE FOUR COMPONENTS OF THE QUAD-AGONIST

Pharmacokinetic profile (i.v.) in pigs | all compounds dosed at 50 ug/kg



MET-097i

Monthly, fully-biased
GLP-1 RA and
GLP-1 RA prodrug

MET-233i

Monthly amylin analog
and multi-NuSH
combinations

ORAL NuSH PEPTIDE PLATFORM

Engineered for
scalability and
injectable-like
performance

STRATEGIC MANUFACTURING

Scaling to profitably
serve a potential
\$170B+ consumer-
driven market¹

2025 CATALYSTS

Key milestones in next
6-12 months

SCALABLE ORAL PEPTIDES WITH INJECTABLE-LIKE PERFORMANCE

Introducing Metsera's oral platform

CURRENT COMPETITORS

Limitations of oral NuSH pipeline



Oral small molecule NuSH RAs:
Efficacy and tolerability suboptimal

- Limited efficacy
- Tolerability inferior to injectables



Oral peptide NuSH RAs:
Not scalable for a large market

- Bioavailability <1% (Rybelsus®)
- 18+ grams of API per year
- Food and water restrictions

METSERA'S APPROACH

Engineering scalable oral peptide delivery

↑ Bioavailability through **MOMENTUM™** platform

↑ Half-life through **HALO™** platform

↑ Peptide potency through **MINT** engineering principles

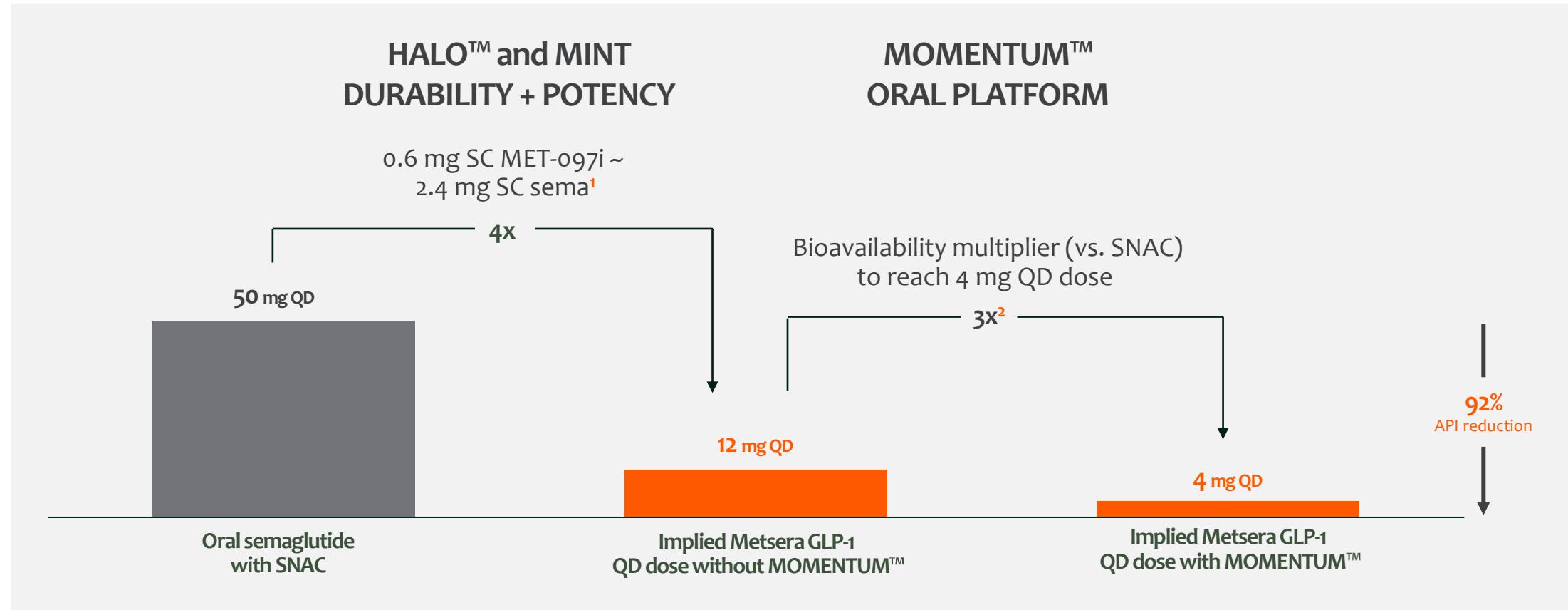
Metsera's oral NuSH pipeline

- MET-002_o
Prototype MOMENTUM peptide
- MET-097_o
FB*, HALO-lipidated GLP-1RA
- MET-224_o
FB*, HALO-lipidated GLP-1RA
- MET-AMY_o
HALO-lipidated oral amylin
- MET-GGG_o
HALO-lipidated triple agonist

MAKING ORAL PEPTIDES SCALABLE

A combination of strategies could deliver a >90% reduction in API versus SNAC

Matching small molecule-like scalability in an oral peptide

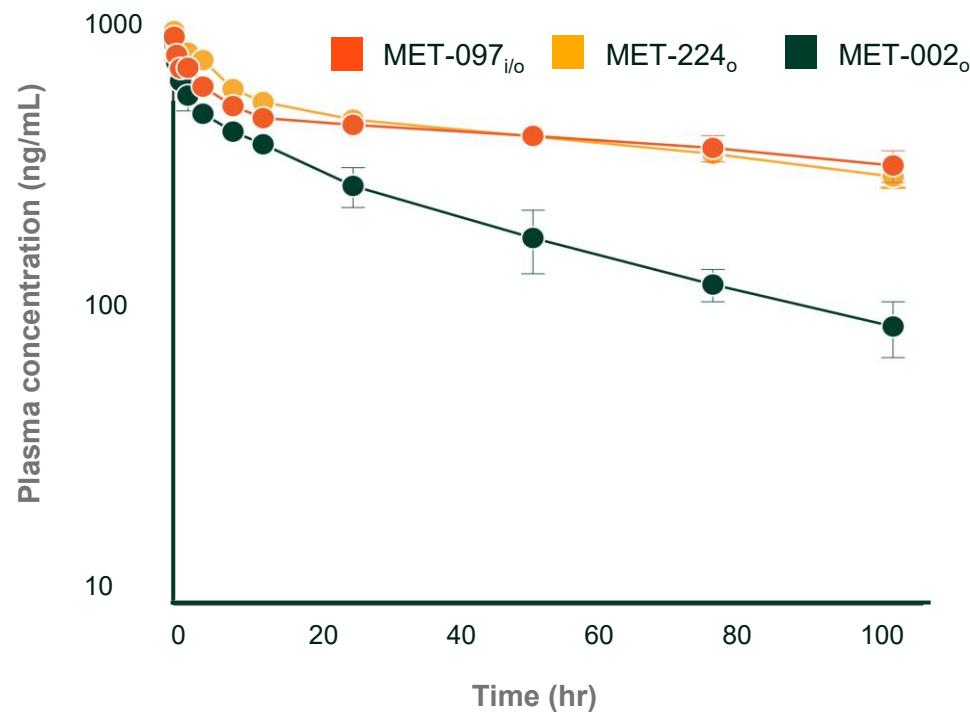


PEPTIDE DURABILITY AND POTENCY

Metsera HALO™ and MINT: Multiplicative effect on chronic efficacy per unit of API

DURABILITY TO REDUCE API NEEDS

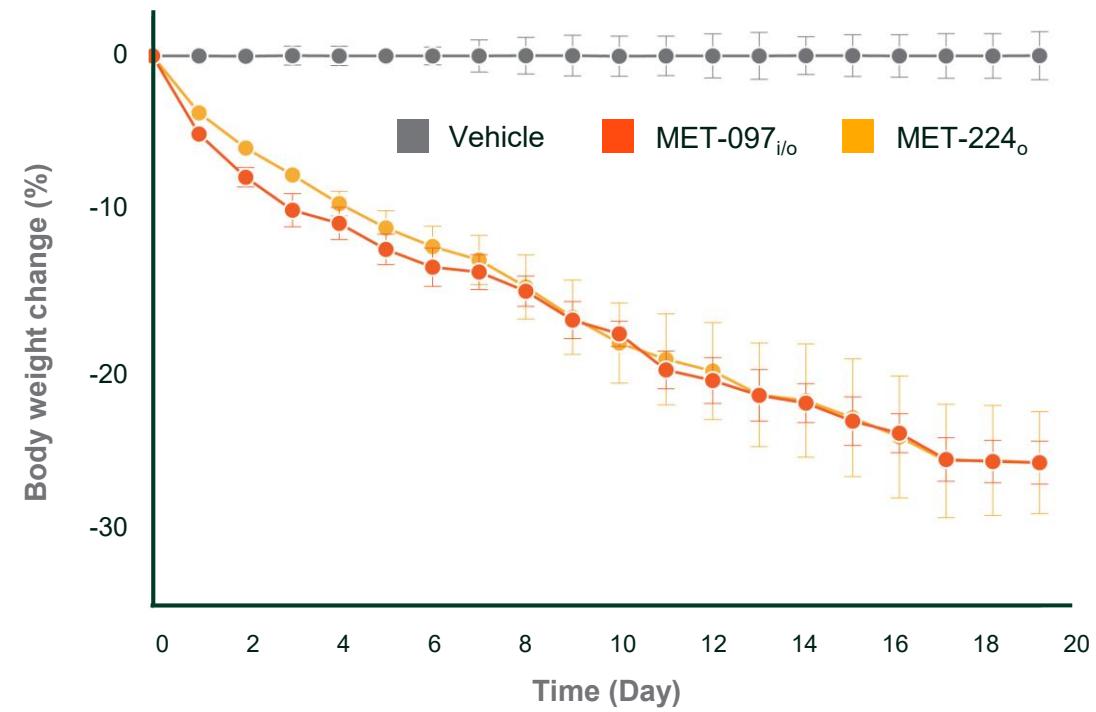
Pharmacokinetic profile (i.v.) in dogs | all doses equivalent



Data presented as mean plasma concentration \pm standard error of the mean (n=3); intravenous dose in dogs

POTENCY TO REDUCE API NEEDS

Chronic dosing (s.c.) in DIO mice | all doses equivalent



Data presented as mean body weight % \pm standard error of the mean (n=5, vehicle; n=9, treatment)

SCALABLE ORAL PEPTIDES WITH INJECTABLE-LIKE PERFORMANCE

First-in-human studies underway; Phase 1/2 data for lead oral GLP-1 RA expected by late 2025

METSERA'S FOCUS: SCALABLE SOLUTIONS

Current approaches to oral delivery self-limiting

- Small molecules: Short-acting pharmacokinetics can constrain efficacy and tolerability
- Oral peptides: Not scalable to-date (18+ grams of peptide per year); food/water restrictions

Metsera approach to address these limitations

- Half-life improvement
- Potency improvement
- Bioavailability improvement
- Substantially lower API requirements
- No food/water restrictions

ONGOING DEVELOPMENT

Phase 1 underway for prototype peptide, MET-002_o

- Objective: Select optimal human formulation

 **Late 2025: 4-week weight loss and tolerability data for selected lead oral¹**

- Lead oral candidates are MET-097_o and MET-224_o

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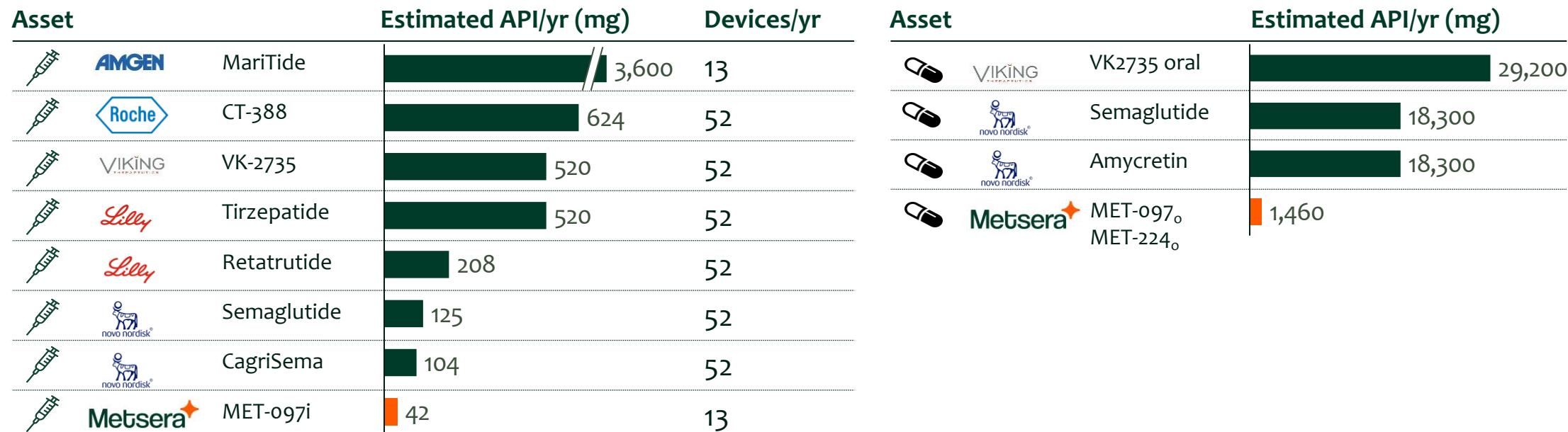
2025 CATALYSTS

Key milestones in next
6-12 months

ENGINEERED TO SCALE TO LARGE POPULATIONS

Metsera has inherent scalability advantages due to potential for QM dosing and low API requirements

Implied per patient, per annum drug substance and device requirements to maintain a 15-20% body weight reduction



SECURING LAUNCH CAPACITY

Building internal manufacturing capability with a trusted partner



AMNEAL+METSEREA

- Joint investment in greenfield plant for API and devices
- Provides launch capacity across injectables and orals
- Leverages Amneal's global network (US and ex-US)

NETWORKED APPROACH FOR COMMERCIAL SUPPLY

- Parallel networked approach for supply chain redundancy
- High-quality partners, e.g., Bachem
- Mitigate geopolitical risk by diversifying commercial suppliers

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2025 CATALYSTS

Key milestones in next
6-12 months

KEY CATALYSTS IN 2025

Execution towards eight potentially inflection events

STRATEGY	MID 2025	LATE 2025
1 MONTHLY GLP-1 RA MET-097i	🚩 VESPER-1¹ (September 2025) 28-week weight loss	🚩 VESPER-3 (end 2025 / early 2026) 28-week weight loss; monthly dosing
2 MONTHLY AMYLIN + GLP-1 RA MET-233i/MET-097i	✓ Amylin monotherapy (Q2 2025) 5-week weight loss & tolerability	🚩 GLP-1 + amylin (end 2025 / early 2026) 12-week weight loss & tolerability
3 ORAL PEPTIDE PLATFORM MET-097o/MET-224o		🚩 Amylin monotherapy 12-week weight loss
4 NEXT-GENERATION COMBINATIONS MET-034i		🚩 Lead oral GLP-1 weight loss & tolerability ³
		🚩 GIP + GLP-1 tolerability ³



THANK YOU

investors@metsera.com

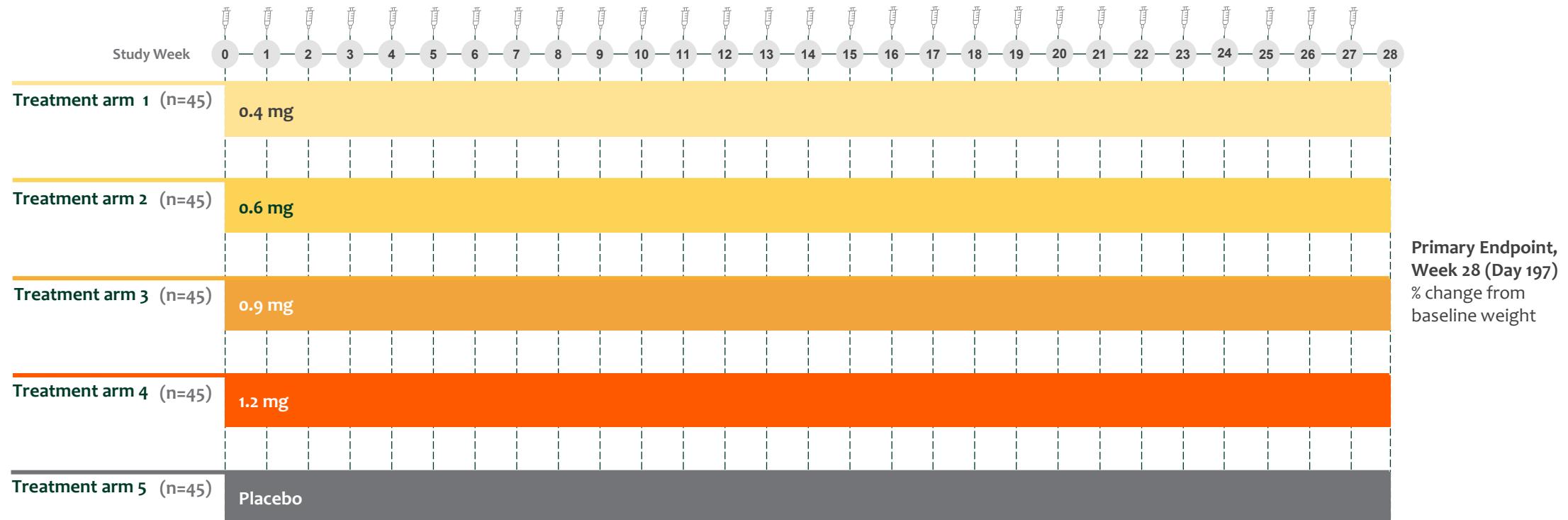


APPENDIX

MET-097i Phase 2b VESPER trials overview

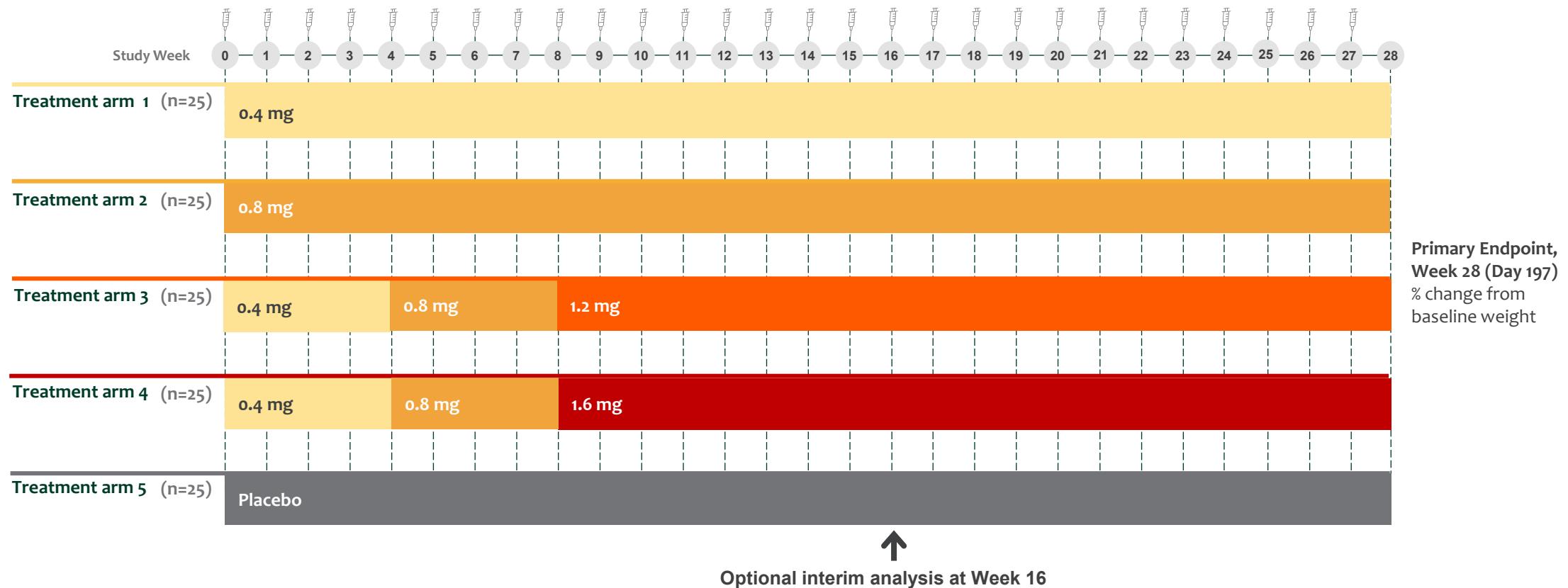
MET-097i VESPER-I TRIAL

28-week, randomized, double-blind, placebo-controlled study of weekly titration-free MET-097i



MET-097i VESPER-2 TRIAL

28-week, randomized, double-blind, placebo-controlled study of weekly MET-097i in T2DM



MET-097i VESPER-3 TRIAL

64-week, randomized, double-blind, placebo-controlled study to evaluate multiple QM MET-097i doses

