

# Investor Presentation

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as of August 10, 2026



Nasdaq: NAMS



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# Obicetrapib and Obicetrapib plus Ezetimibe Could Address Significant Unmet Need in Cardiometabolic Diseases



**Significant unmet** need for oral LDL-C lowering therapy as adjunct to statins



**Simple, once-daily, low-dose** CETP inhibitor with statistically significant LDL-C lowering and effects beyond LDL, observed across three Phase 3 trials and five Phase 2 trials



**Convenient oral format** potentially enables broad adoption and combinations to address unmet need, if approved



Comprehensive IP protection **until mid 2043**

**\$8 billion+**

global peak sales opportunity with 30m+ patients in the US alone not achieving LDL-C lowering goals

**>3,500 pts**

of favorable safety/tolerability data comparable to placebo. Blinded data in >9,500 patients

**35-40%**

LDL-C lowering versus placebo as a monotherapy\*

**~45%**

Lp(a) lowering versus placebo as a monotherapy<sup>+</sup>

**~50%**

LDL-C lowering versus placebo in combination\*\* with ezetimibe

**21%**

observed reduction in MACE favoring obicetrapib at 1-year<sup>^</sup>

**Observed beneficial effects beyond LDL, on ApoB, non-HDL-C, LDL-P, HDL-C and other markers of glycemic control, renal function, and Alzheimer's disease**

# Previous Years Were a Time of Groundwork, Goals and Growth



Completed and announced results for BROOKLYN, BROADWAY, and TANDEM Phase 3 studies



Building world-class commercial and MSL functions



Doubled in size (>100) with new hires and offices in Amsterdam, NL, Miami, FL, and Philadelphia metro area



Secured funding to support US commercial launch, if approved, with cash at end of 2Q 2026 of ~\$678M\*

2024



Composition of matter IP granted



Announced topline results for BROOKLYN Phase 3 trial



Announced topline results for TANDEM Phase 3 trial



Announced topline results for BROADWAY Phase 3 trial



Announced Alzheimer's topline

2025



Full AD data was presented at AAIC on July 30



EMA submission



CHMP Positive Opinion

2026



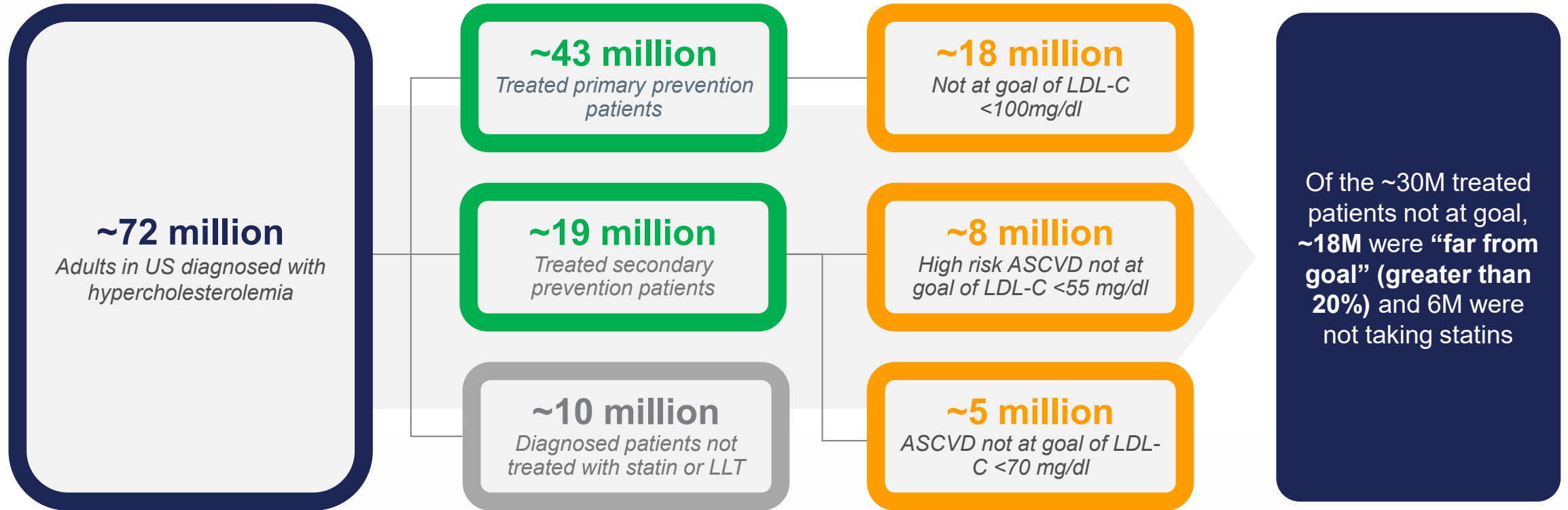
European Approvals



RUBENS Phase 3 Topline

\* Includes cash and cash equivalents and marketable securities

# Obicetrapib Could Address the ~30M Patients in US on Drug but not at Goal

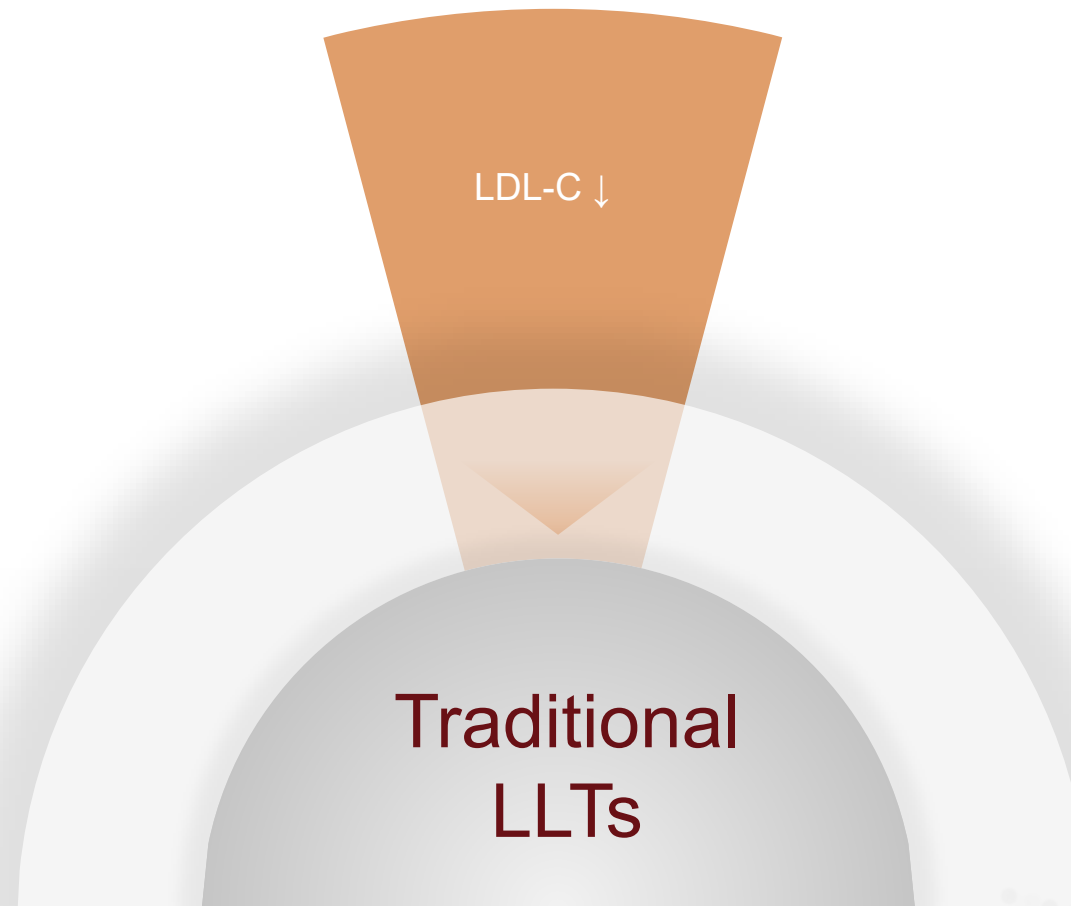


## US Branded Lipid Lowering Market

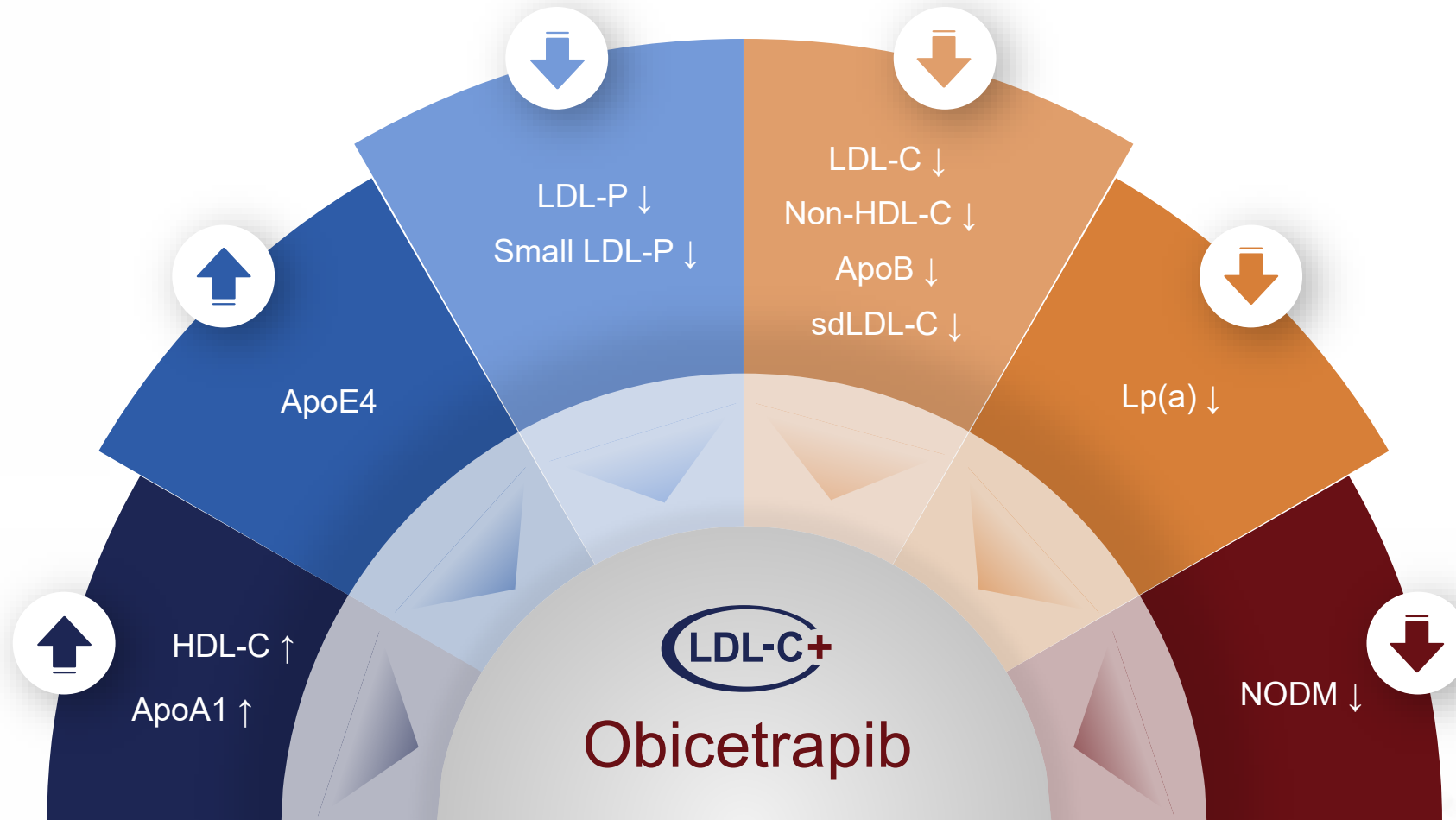
Potential key factors limiting penetration include **product limitations** and **market access** hurdles:  
**Low prescriber enthusiasm for existing TPPs**  
**Payors restrict access**

# Currently, Treatment of Hyperlipidemia is One-Dimensional

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# The Future of Hyperlipidemia Treatment Goes Beyond LDL-C



# The Future of Hyperlipidemia Treatment Goes Beyond LDL-C

Obicetrapib was observed to **impact multiple risk factors**, potentially leading to more comprehensive risk management across **additional patient subpopulations**

## LDL Particles

- Approximately **43 million** patients with hyperlipidemia exhibit elevated levels of **LDL-P of > 1300 nmol/L\***
- Prevalence is **even higher** in individuals **with type 2 diabetes<sup>^</sup>**
- Presence of LDL-P is a **significant independent risk factor for coronary heart disease (CHD)**, even when traditional lipid measures like LDL-C and triglycerides are within normal ranges<sup>^</sup>

## Lp(a)

- In the US, close to **75 million** patients have an elevated Lp(a) score of **> 100-125 nmol/L\*\* ^^**
- An estimated **14.3 million** of these patients also have hypercholesterolemia

## Diabetes

- **27.9 million** hyperlipidemia patients are estimated to have concomitant diabetes\*
- In addition, there are an estimated **27.2 million<sup>^</sup>** individuals with hyperlipidemia that are pre-diabetic

## ApoE4

- ApoE4 is the **strongest genetic risk factor** for late-onset AD and associated with a **22-45%** increased risk of CVD<sup>1-5</sup>
- Approximately **20 million** patients with hyperlipidemia are at increased risk for AD
- Prevalence of ApoE4 in AD is up to **66%<sup>1-5</sup>**

\* Source: Toth, Peter P. et al. *Atherosclerosis*. Volume 235, Issue 2, 585 - 591. ^Hoogeveen RC, Gaubatz JW, Sun W, Dodge RC, Crosby JR, Jiang J, Couper D, Virani SS, Kathiresan S, Boerwinkle E, Ballantyne CM. Small dense low-density lipoprotein-cholesterol concentrations predict risk for coronary heart disease: the Atherosclerosis Risk In Communities (ARIC) study. *Arterioscler Thromb Vasc Biol*. 2014 May;34(5):1069-77. doi: 10.1161/ATVBAHA.114.303284. Epub 2014 Feb 20. PMID: 24558110; PMCID: PMC3999643. \* Ray K, et al. *European Journal of Preventive Cardiology* (2021) 28, 1279–1289. ^ Footnote: Prediabetes was defined as fasting plasma glucose values of 100 to 125 mg/dL or A1C values of 5.7% to 6.4%. Reference: National Diabetes Statistics Report | Diabetes | CDC\*\* Tsimikas et al. *J Am Coll Cardiol*. 2022; 80: 934-946 ^^ Shapiro, M et al. *Journal of Clinical Lipidology*, Volume 19, Issue 1, 28 - 38 Aβ, amyloid β; AD, Alzheimer's disease; Apo, apolipoprotein; CVD, cardiovascular disease; HDL-C, high density lipoprotein cholesterol; LDL-C, low density lipoprotein cholesterol; Lp(a), lipoprotein(a). 1. Abondio P et al. *Genes (Basel)*. 2019;10(3):222. 2. Feringa FM et al. *Front Aging Neurosci*. 2021;13:690372. 3. Hottman 2017. 4. Jeong W et al. *Mol Cells*. 2019;42(11):739-746. 5. Williams et al. *BMC* 2020.c

# Lipid Lowering Market Primed for Commercial Success

| Observations <sup>1-4</sup> |                                                                                     | Key Takeaways                                                                                                                                    |                                                                                                        |
|-----------------------------|-------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------|
| 1                           |    | We have seen LLT market growth each year for the last 5 years                                                                                    | Very large opportunity – even small amounts of penetration can result in significant financial success |
| 2                           |    | Non-statin growth continues in the high double digits                                                                                            | Shift towards more aggressive treatment using alternatives / adding to statins                         |
| 3                           |    | Guidelines have shifted to recommend more intense and earlier intervention not only with statins, but with ezetimibe, PCSK9's and bempedoic acid | Governing bodies and KOL's are prioritizing the importance of treating to goal                         |
| 4                           |    | Access to PCSK9's has been increasing, and utilization controls have been decreasing                                                             | Payers are willing to cover medications in LLT space if priced appropriately                           |
| 5                           |   | PCSK9 growth continues to accelerate 10 years post launch with 45%+ growth in net sales in each of the last 2 years                              | Commercial success is being demonstrated and is accelerating                                           |
| 6                           |  | FDA labeling changes continue to support broad access and earlier treatment                                                                      | Labels are less restrictive and support broad usage                                                    |

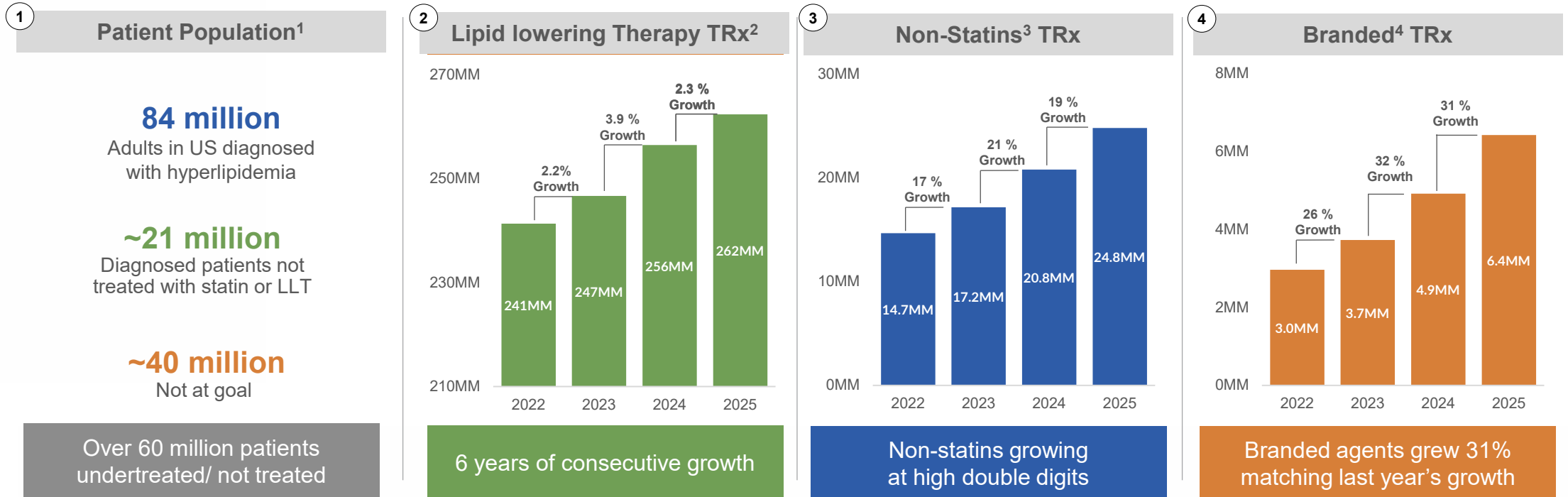
1. Based on third party prescription data on file

2. Grundy SM, et al. J Am Coll Cardiol. 2019;73(24):3168-3209.

3. Lloyd-Jones DM, et al. J Am Coll Cardiol. 2022;80(14):1366-1418

4. 2025 ACC/AHA/ACEP/NAEMSP/SCAI Guideline for the Management of Patients With Acute Coronary Syndromes: A Report of the American College of Cardiology/American Heart Association Joint Committee on Clinical Practice Guidelines | Circulation

# Market Dynamics: Growth of Non-Statin (+19%) and Branded (+31%) Volume Continue to Grow Rapidly Year Over Year



**Market Growth of 2.3% in 2025, adds 6 million additional TRx's, of which 1.5 million are branded TRx's**

Source: Based on third party prescription data on file

1. Forian Healthcare Closed Claims and EHR Data (2022-2024)

2. All Lipid Lowering therapies: Statins, Ezetimibe and combinations; PCSK9 and BPA;

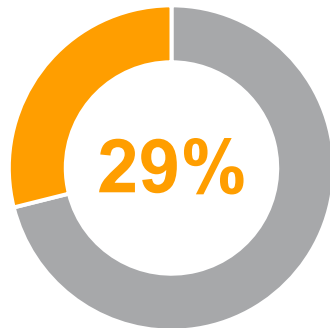
3. Non-Statins : Ezetimibe and combinations; PCSK9 and BPA;

4. Branded: PCSK9 and BPA;

# Majority of ASCVD/HeFH Patients are not Achieving LDL-C Targets

**Primary prevention HeFH**  
patients with  
an LDL-C target <100 mg/dL  
(2011-2017)<sup>1</sup>

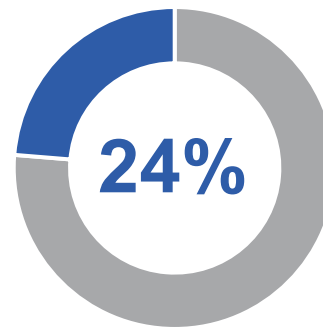
LDL-C < 100 mg/dL



<1/3 achieved  
LDL-C <100 mg/dL

**ASCVD patients** with an  
LDL-C target of LDL<70 or  
<55 mg/dL (2017-2018)<sup>2</sup>

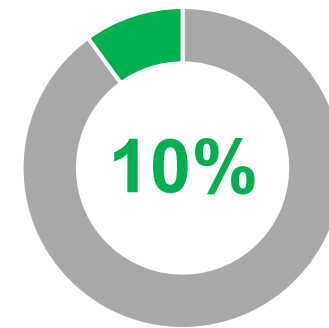
LDL-C < 70 mg/dL



~1/4 achieved  
LDL-C <70 mg/dL

**Very high risk ASCVD**  
**patients** with an LDL-C  
target <55 mg/dL (2020-  
2021)<sup>3</sup>

LDL-C < 55 mg/dL



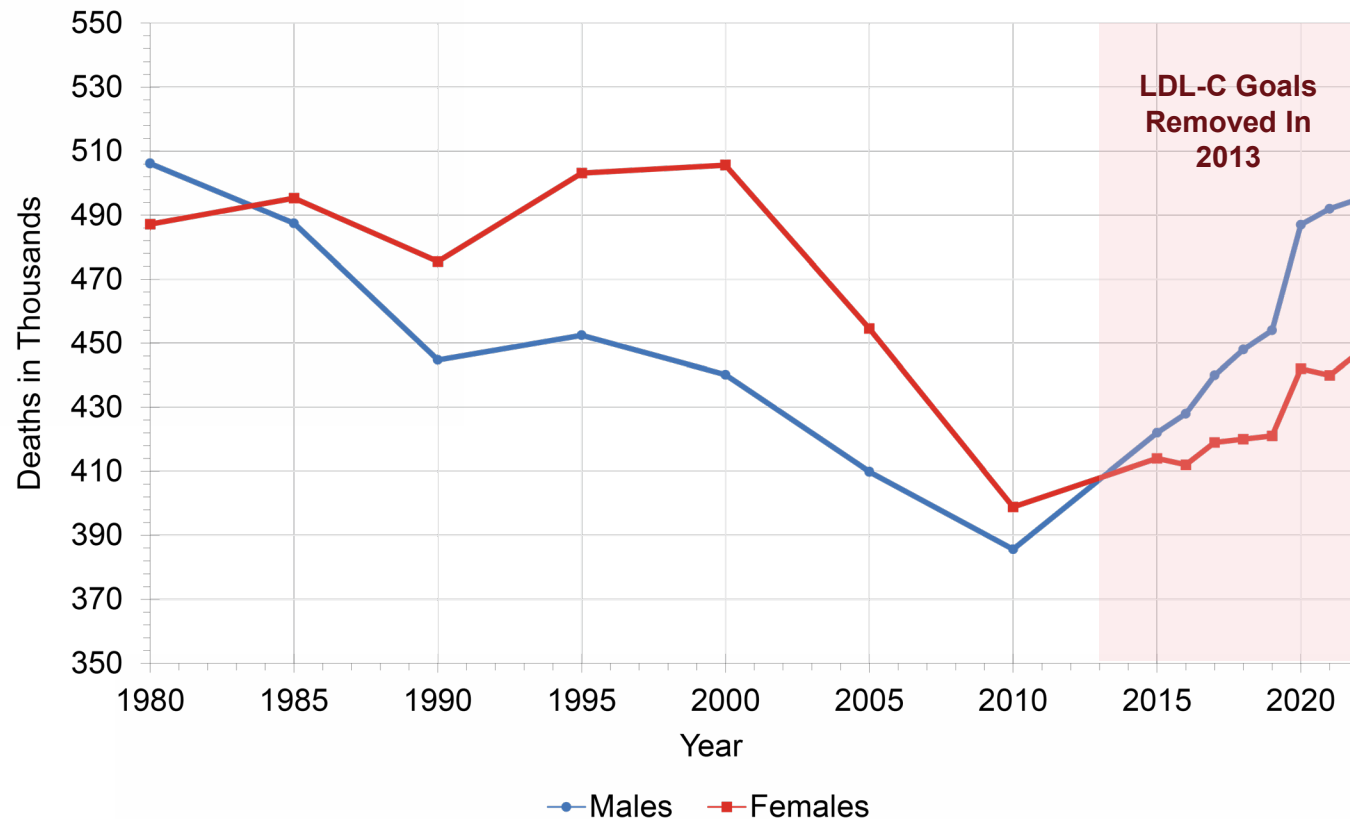
10% achieved  
LDL-C <55 mg/dL

ASCVD=atherosclerotic cardiovascular disease; HeFH=heterozygous familial hypercholesterolemia; LDL-C=low-density lipoprotein-cholesterol.

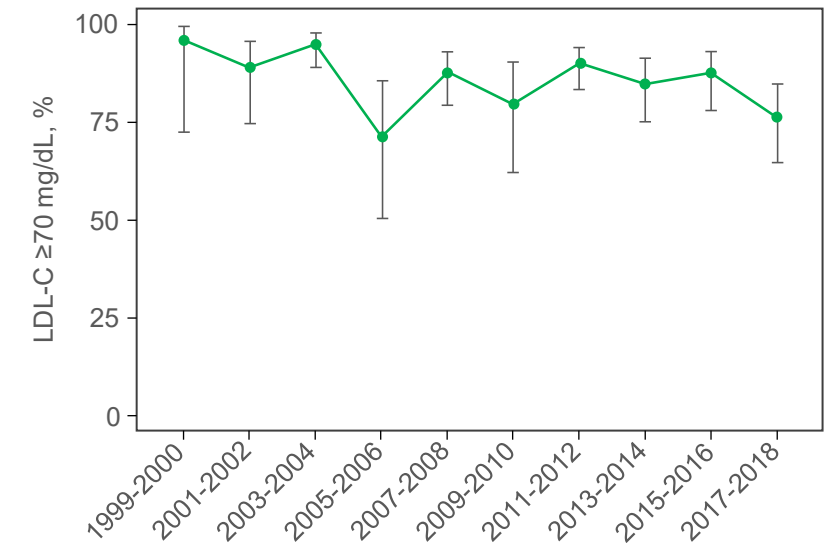
1. Schreuder MM, et al. LDL cholesterol targets rarely achieved in familial hypercholesterolemia patients: A sex and gender-specific analysis. *Atherosclerosis*. 2023 2. Gao Y, Shah LM, Ding J, Martin SS. US trends in cholesterol screening, lipid levels, and lipid-lowering medication use in US adults, 1999 to 2018. *J Am Heart Assoc*. 2023;12(3):e028205; 3. Katzmann JL, et al. Simulation study on LDL cholesterol target attainment, treatment costs, and ASCVD events with bempedoic acid in patients at high and very-high cardiovascular risk. *PLoS One*. 2022;17(10):e0276898;

# CV Events Took an Alarming Turn Following Removal of LDL-C Guidelines in 2013

CVD Mortality Trends for US Males and Females, 1980 to 2022<sup>1</sup>



Trends in Prevalence of High LDL-C in US Adults, NHANES 1999-2018 with History of ASCVD<sup>2</sup>



**~75% of ASCVD patients are NOT at their risk-based LDL-C goal**

# Physicians Have Limited Treatment Options to Meet Patients Needs



Available and investigational treatment options have limitations

|                          | Ezetimibe <sup>(1)</sup>                                                            | Nexletol <sup>(2)</sup>                                                               | PCSK9i <sup>(3)</sup>                                                                 | Obicetrapib <sup>(5)</sup>                                                            | Obi + Eze <sup>(5)</sup>                                                              |
|--------------------------|-------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------|
| Approval                 | Approved                                                                            | Approved                                                                              | Approved                                                                              | LDL-C data 2024                                                                       | LDL-C data 2024                                                                       |
| MACE Benefit             | 7%                                                                                  | 13%                                                                                   | 15%                                                                                   | 21%*                                                                                  | TBD                                                                                   |
| Observed LDL-C Reduction | 25%                                                                                 | 17%                                                                                   | 45-50%                                                                                | 35-40%                                                                                | 49-54%                                                                                |
| Administration           | Oral<br>(small molecule)                                                            | Oral<br>(small molecule)                                                              | Injectable<br>(mAb)                                                                   | Oral<br>(small molecule)                                                              | Oral<br>(small molecule)                                                              |
| Dosing                   | 10mg                                                                                | 180mg                                                                                 | 140-150mg                                                                             | 10mg                                                                                  | 20mg<br>(10mg Obi + 10mg Eze)                                                         |
| Food Effect              | No                                                                                  | No                                                                                    | No                                                                                    | No                                                                                    | No                                                                                    |
| Safety & Tolerability    | Safe, well-tolerated                                                                | Tendon rupture & gout warning on label                                                | Safe, injection site reactions                                                        | Well-tolerated compared to placebo                                                    | Well-tolerated compared to placebo                                                    |
| Lp(a) lowering           | None                                                                                | None                                                                                  | 15-30%                                                                                | 34-56%                                                                                | 63%                                                                                   |
|                          |  |  |  |  |  |

Note: The above data do not represent head-to-head comparisons. Actual results may differ from expectations. Obicetrapib mono and Ezetimibe combo, along with the Oral PCSK9 have not been approved by any regulatory authority. E= estimated dates. Red represents sub-optimal product characteristics.

Sources: 1. PI Zetia table 7. refers to; Gagne, C et al. Am J Cardiol 2002. LDL-C measured only using Friedewald 2. PI Nexletol; study 2. refers to; Goldberg, A et al. JAMA 2019;322(18):1780-1788. LDL-C measured using Friedewald and direct assay for LDL-C <50 mg/dL. 3. multiple studies: Blom, D et al. N Engl J Med 2014; Kereiakes, D et al. Am Heart J 2015.; Ray, K. N Engl J Med 2020. 4. Ballantyne, C et al. JACC 2023;81(16) 5. See slide 12 for LDL-C, \*MACE benefit from exploratory analysis in BROADWAY at 1 year 6. MK0616 was observed to have adverse events comparable to pbo in Phase 2b trials

# Limited New Therapies on the Horizon



Available and investigational treatment options have limitations

## Phase 3 Data Readouts

## Current Phase

## Observed LDL-C Reduction

## Administration

## Dosing

## Food Effect

## Safety & Tolerability

## Lp(a) Lowering

### LIPFENDRA<sup>(1)</sup>

(enlicitide)

LDL-C data 2025  
(CVOT data 2029E)

Approved

50-59%  
(~20% with food)

Oral  
(peptide)

380mg  
(20mg API + 360mg SNAC)

Yes  
(8hr fast & 30min wait)

SNAC technology has  
previously been observed  
to have  
tolerability concerns<sup>(6)</sup>

20-25%



### AZD0780<sup>(2)</sup>

TBD

Phase 3  
Ongoing

35-51%

Oral  
(small molecule)

30mg

No

Well-tolerated  
SAEs and case of AST/ALT  
> 5x ULN

7-20%



### Obicetrapib<sup>(3)</sup>

LDL-C data 2024

CVOT  
ongoing

35-40%

Oral  
(small molecule)

10mg

No

Well-tolerated  
compared to placebo

34-56%



### Obi + Eze<sup>(4)</sup>

LDL-C data 2024

CVOT  
ongoing

49-54%

Oral  
(small molecule)

20mg  
(10mg Obi + 10mg Eze)

No

Well-tolerated  
compared to placebo

63%

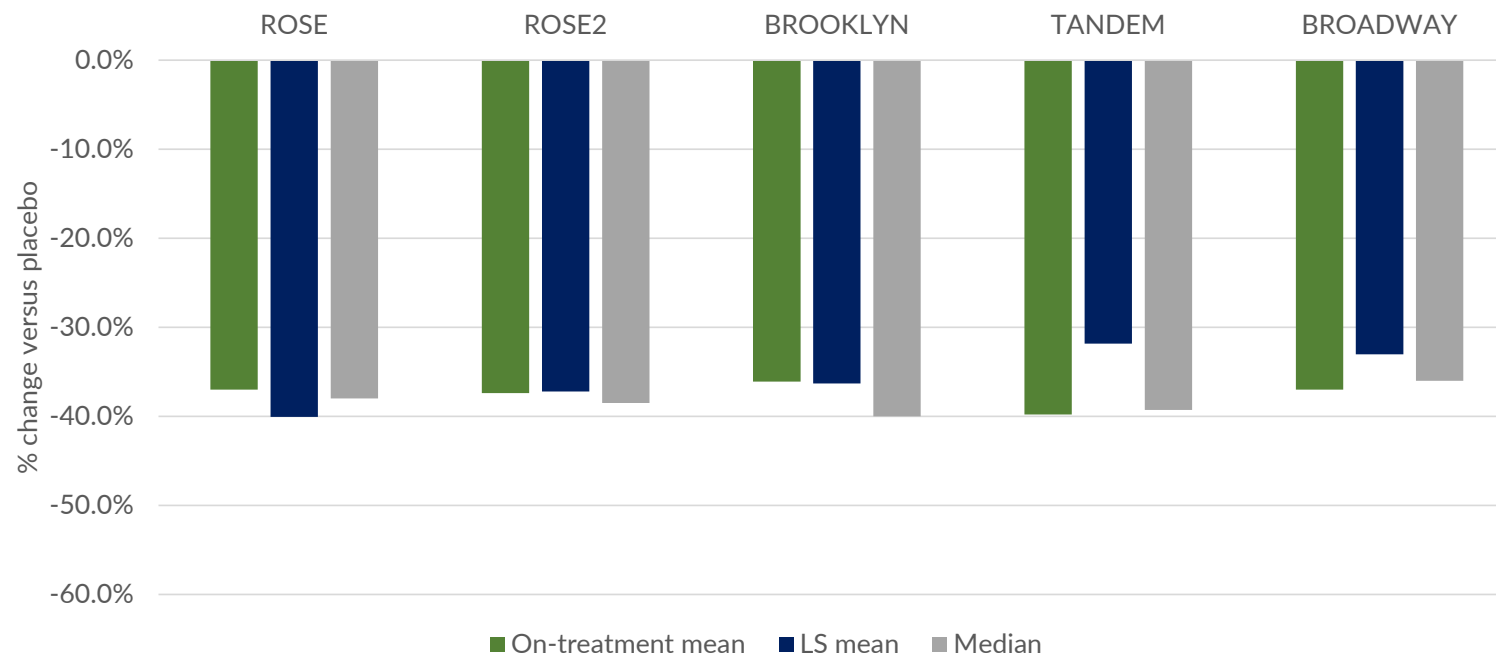


# Obicetrapib Program Designed to Overcome Limitations of Prior CETP Inhibitors

|                                         | Torcetrapib <sup>(1)</sup> | Dalcetrapib <sup>(2)</sup> | Evacetrapib <sup>(3)</sup>                      | Anacetrapib <sup>(4)</sup>     | Obicetrapib <sup>(5)</sup> |
|-----------------------------------------|----------------------------|----------------------------|-------------------------------------------------|--------------------------------|----------------------------|
| Observed LDL-C reduction <sup>(6)</sup> | 25%                        | 7%                         | 11-21%                                          | 17%                            | 35-40%                     |
| CETP inhibition                         | 35%                        | 30%                        | 65%                                             | 80%                            | 97%                        |
| Dosing                                  | 60mg                       | 600mg                      | 100mg                                           | 100mg                          | 10mg                       |
| Blood pressure increase                 | Yes                        | No                         | No                                              | No                             | No                         |
| Aldosterone increase                    | Yes                        | No                         | No                                              | No                             | No                         |
| Lp(a) lowering                          | unknown                    | unknown                    | 20-25%                                          | 20-25%                         | 34-56%                     |
| ApoB lowering                           | 15%                        | None                       | 15-20%                                          | 18%                            | 22-24%                     |
| <b>OUTCOMES STUDIES</b>                 |                            |                            |                                                 |                                |                            |
| Name                                    | ILLUMINATE                 | Dal-OUTCOMES               | ACCELERATE                                      | REVEAL                         | PREVAIL                    |
| Patients                                | 15,067                     | 15,871                     | 12,092                                          | 30,449                         | 9,541                      |
| Baseline LDL-C (mg/dl)                  | 79.7                       | 76.4                       | 81.1                                            | 61                             | 103                        |
| LDL-C reduction (mg/dl)                 | 20                         | NS                         | 25                                              | 11                             | TBD                        |
| Median follow-up                        | 18 mo                      | 31 mo                      | 26 mo                                           | 49 mo                          | 42 mo (expected)           |
| Result (HR)                             | 1.25                       | 1.04                       | 1.01                                            | 0.91                           | TBD                        |
| Explanation                             | Off target tox             | No LDL-C benefit           | Short follow-up but mortality benefit (HR 0.84) | Low baseline and LDL reduction | TBD                        |

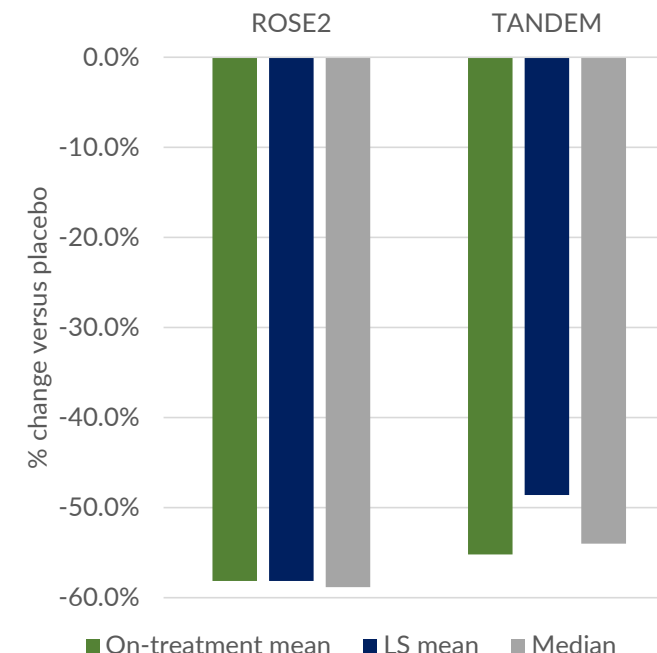
# Consistent LDL-C Reduction Observed Across Our Phase 2 and Phase 3 Trials, Independent of Background Statin and/or Multiple Lipid Lowering Therapy Usage

## Obicetrapib 10 mg monotherapy



Monotherapy reductions of 35-40%

## Obicetrapib plus ezetimibe

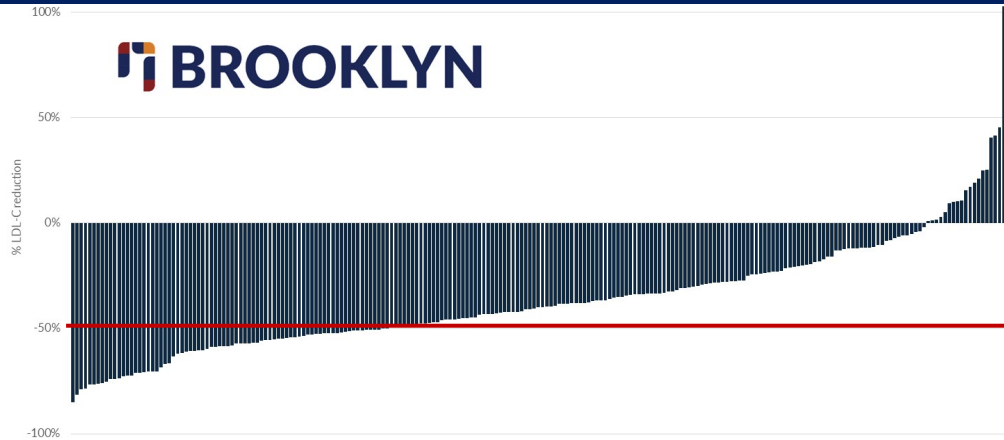


Combo reductions of 50-60%

# LDL-C Responder Analysis at day 84

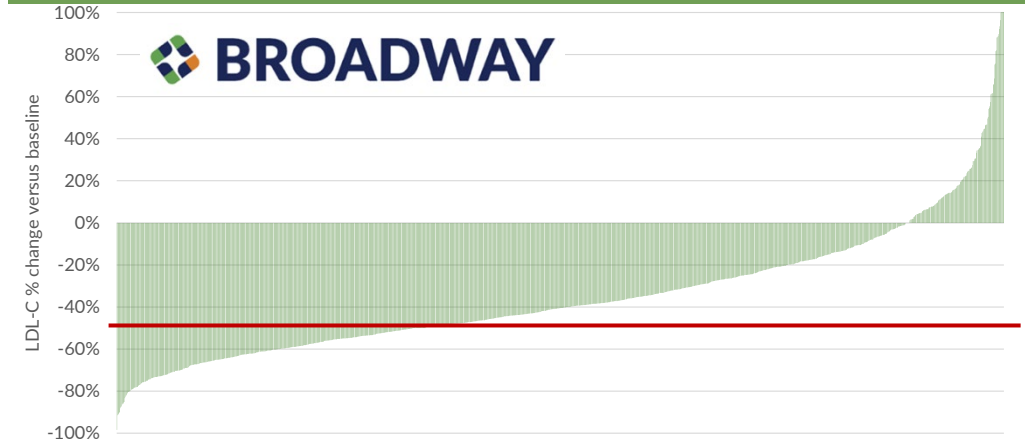
34% of patients with >50% LDL-C reduction

 **BROOKLYN**



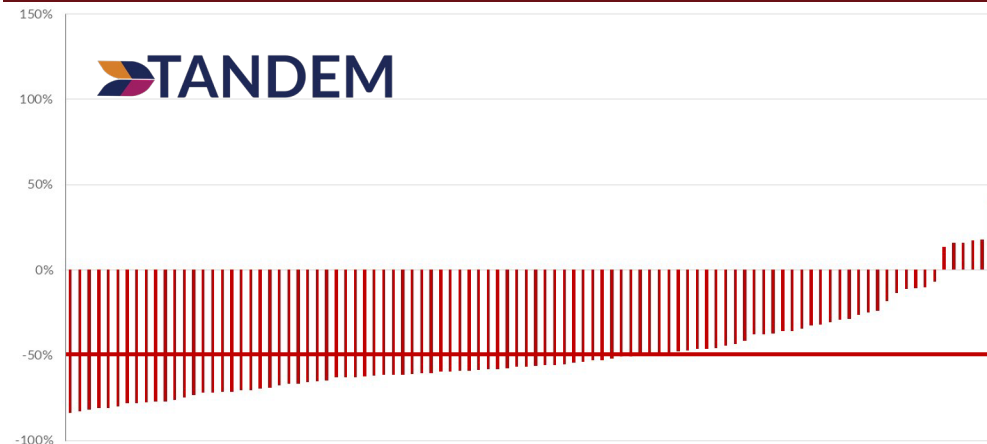
33% of patients with >50% LDL-C reduction

 **BROADWAY**



61% of patients with >50% LDL-C reduction

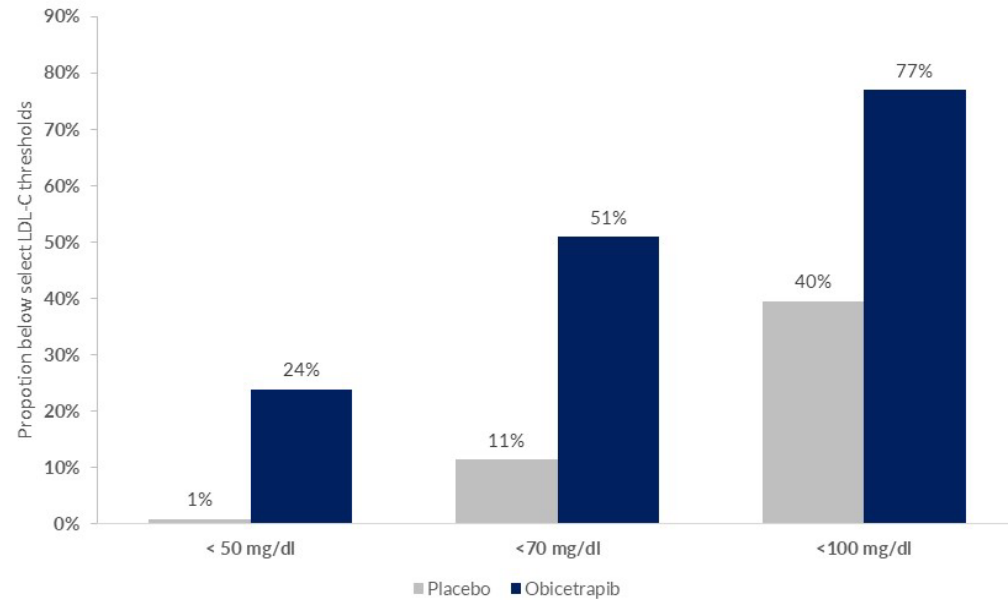
 **TANDEM**



# Greater Proportion of Patients in Obicetrapib Arm Achieved LDL-C Goal

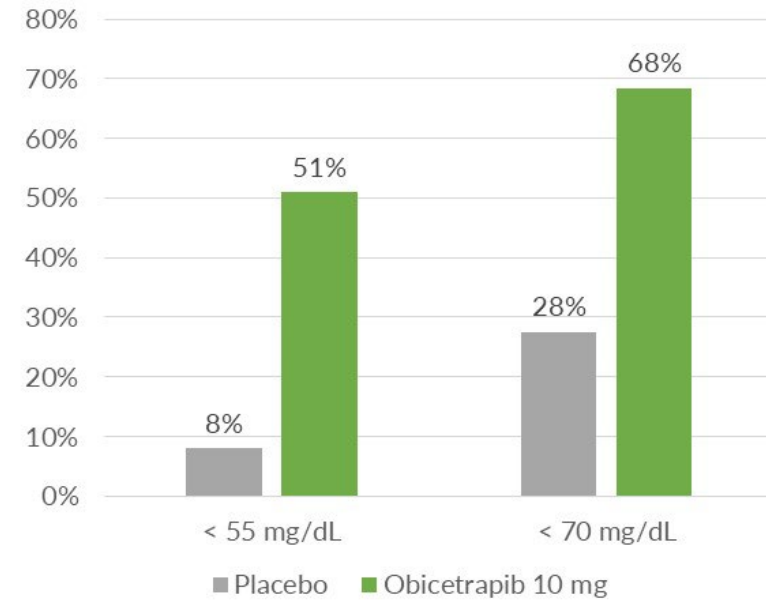
## BROOKLYN

### % of patients achieving LDL-C thresholds at Day 84



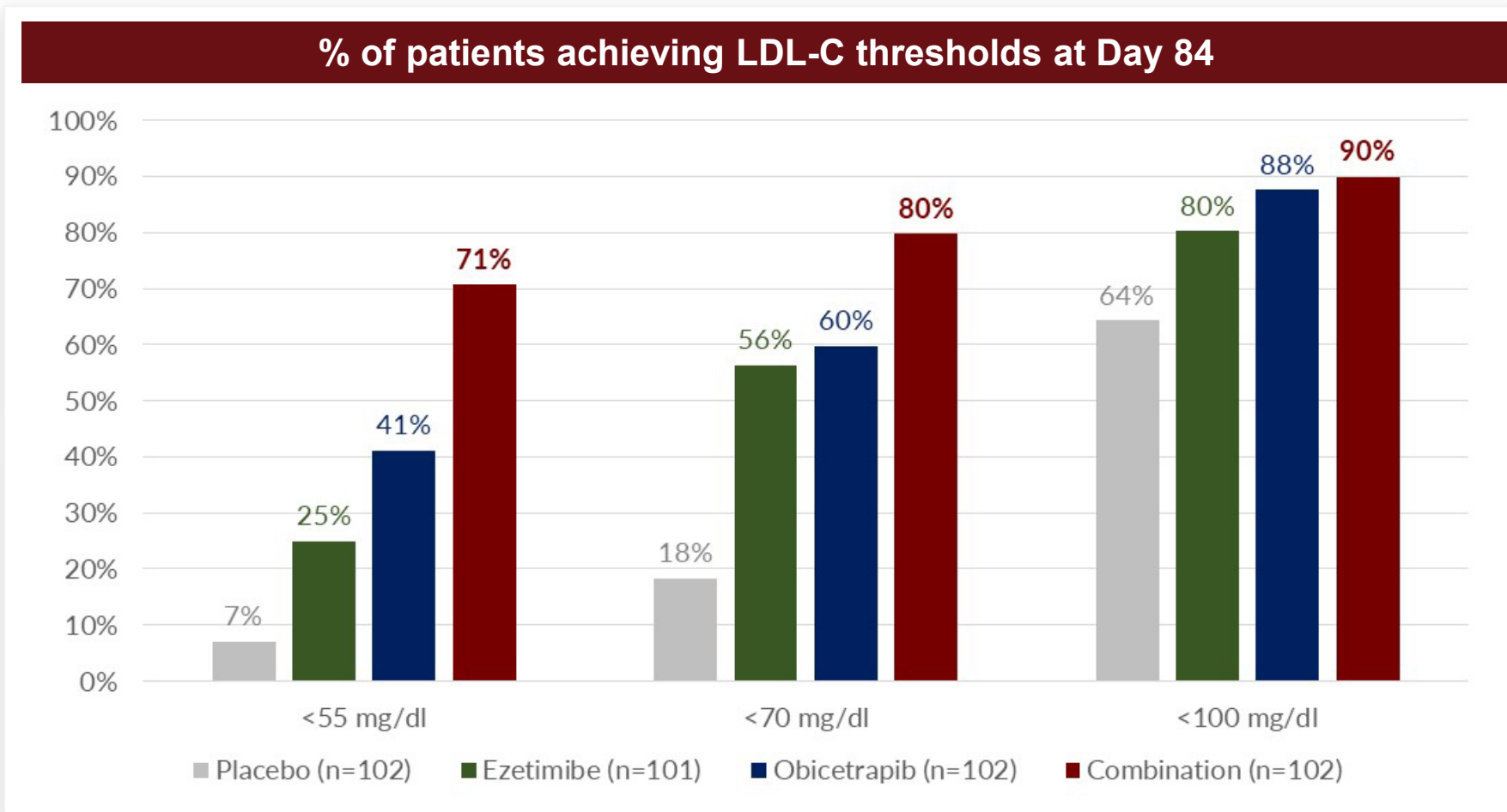
## BROADWAY

### % of patients achieving LDL-C thresholds at Day 84



## Over 70% of Patients on Obicetrapib+Ezetimibe Achieved Less than <55 mg/dL

**TANDEM**



# Exploratory Endpoint: Major Adverse Cardiovascular Events (MACE)

## BROADWAY MACE Data<sup>(1)</sup>

|                                | Placebo<br>N = 844 | Obicetrapib<br>N = 1686 | Hazard Ratio | 95% CI      |
|--------------------------------|--------------------|-------------------------|--------------|-------------|
| All-cause mortality – no. (%)  | 12 (1.4)           | 19 (1.1)                | 0.83         | (0.40-1.71) |
| Coronary heart death – no. (%) | 5 (0.6)            | 8 (0.5)                 | 0.80         | (0.26-2.44) |
| First 4-point MACE – no. (%)   | 44 (5.2)           | 70 (4.2)                | 0.79         | (0.54-1.15) |

## BROADWAY + BROOKLYN Pooled MACE Data <sup>(1)</sup>

|                                | Placebo<br>N = 962 | Obicetrapib<br>N = 1920 | Hazard Ratio | 95% CI      |
|--------------------------------|--------------------|-------------------------|--------------|-------------|
| All-cause mortality – no. (%)  | 14 (1.5)           | 20 (1.0)                | 0.78         | (0.39-1.58) |
| Coronary heart death – no. (%) | 7 (0.7)            | 9 (0.5)                 | 0.63         | (0.24-1.70) |
| First 4-point MACE – no. (%)   | 49 (5.1)           | 75 (3.9)                | 0.75         | (0.53-1.08) |

# MACE Benefit Observed Across Multiple Components After 1 Year of Treatment

## BROADWAY MACE Data<sup>(1)</sup>

|                                      | Placebo<br>N = 844 | Obicetrapib<br>N = 1686 | Hazard Ratio | 95% CI      |
|--------------------------------------|--------------------|-------------------------|--------------|-------------|
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| First 4-point MACE – no. (%)         | 44 (5.2)           | 70 (4.2)                | 0.79         | (0.54-1.15) |
| Coronary heart death – no. (%)       | 5 (0.6)            | 8 (0.5)                 | 0.80         | (0.26-2.44) |
| Nonfatal MI – no. (%)                | 11 (1.3)           | 20 (1.2)                | 0.91         | (0.44-1.90) |
| Stroke – no. (%)                     | 5 (0.6)            | 14 (0.8)                | 1.39         | (0.50-3.86) |
| Coronary revascularization – no. (%) | 34 (4.0)           | 49 (2.9)                | 0.72         | (0.46-1.11) |

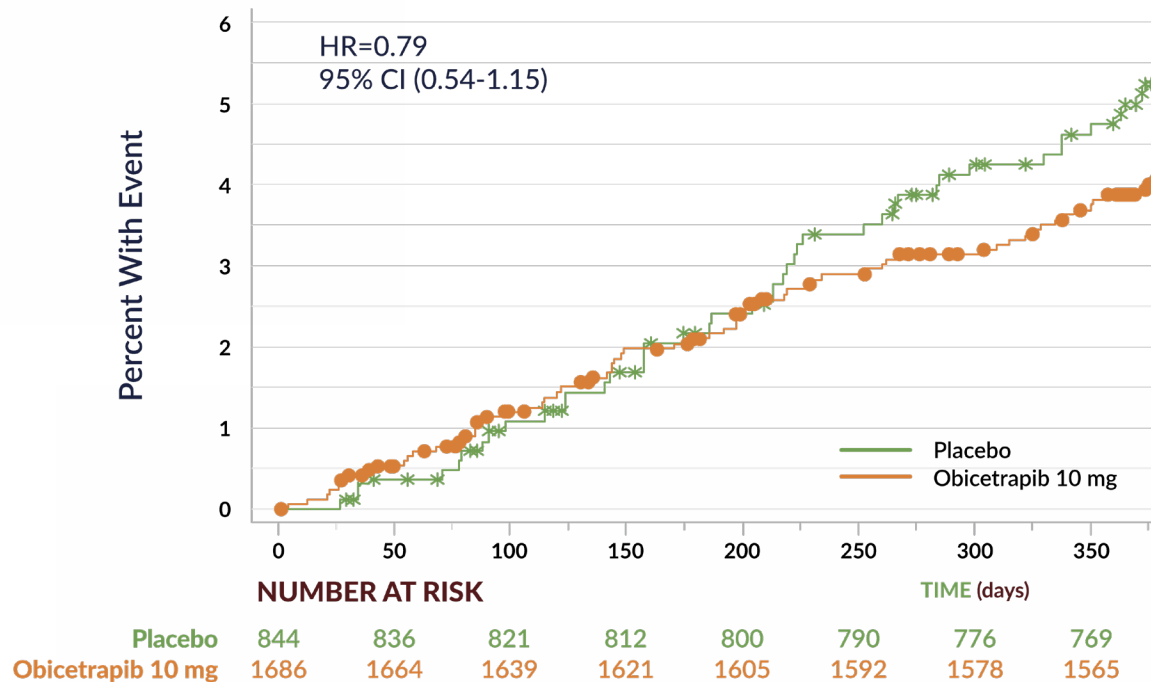
4-point MACE: CHD death, Non-fatal myocardial infarction, non-fatal stroke, coronary revascularization

(1) MACE was evaluated in BROADWAY as an exploratory endpoint. There are limitations on the interpretation of MACE data derived from both BROOKLYN and BROADWAY studies given they were not designed or powered to assess MACE. MACE data presented above may not be predictive of the MACE data we observe in PREVAIL, which has been designed to measure MACE benefit.

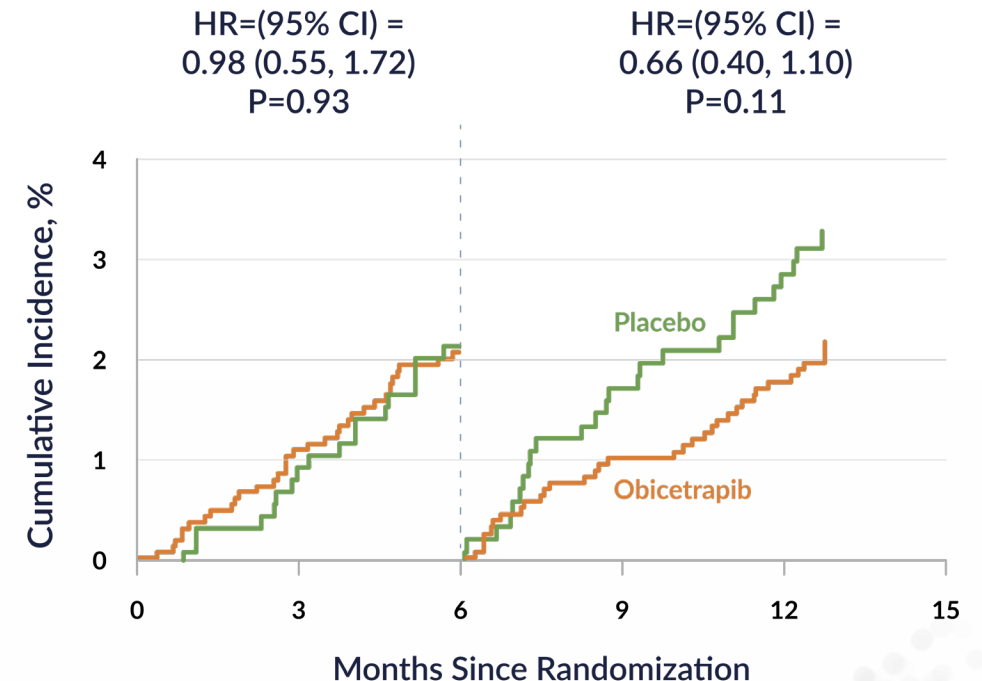
# Kaplan-Meier Curve Separates at Day 200 Driven by Attenuation of MACE-4 Events in the Obicetrapib Arm



Kaplan Meier Curve



Landmark Analysis: Censored at 6 months



4-point MACE: CHD death, Non-fatal myocardial infarction, non-fatal stroke, coronary revascularization

(1) While not powered to detect a MACE benefit, we observed positive MACE data as part of our review of the exploratory endpoint. BROADWAY was not powered to measure MACE benefit and the MACE data presented below may not be predictive of the MACE data we observe in PREVAIL, which has been designed to measure MACE benefit.

# Explaining BROADWAY's Observed 21% MACE Reduction

## Methods of Estimating MACE Reduction



Based on the CTT line, the expectation for a 33% reduction in LDL-C from placebo would be a 9% MACE reduction, after 12 months



LDL-P reduction potentially provides 6-7% additional MACE reduction

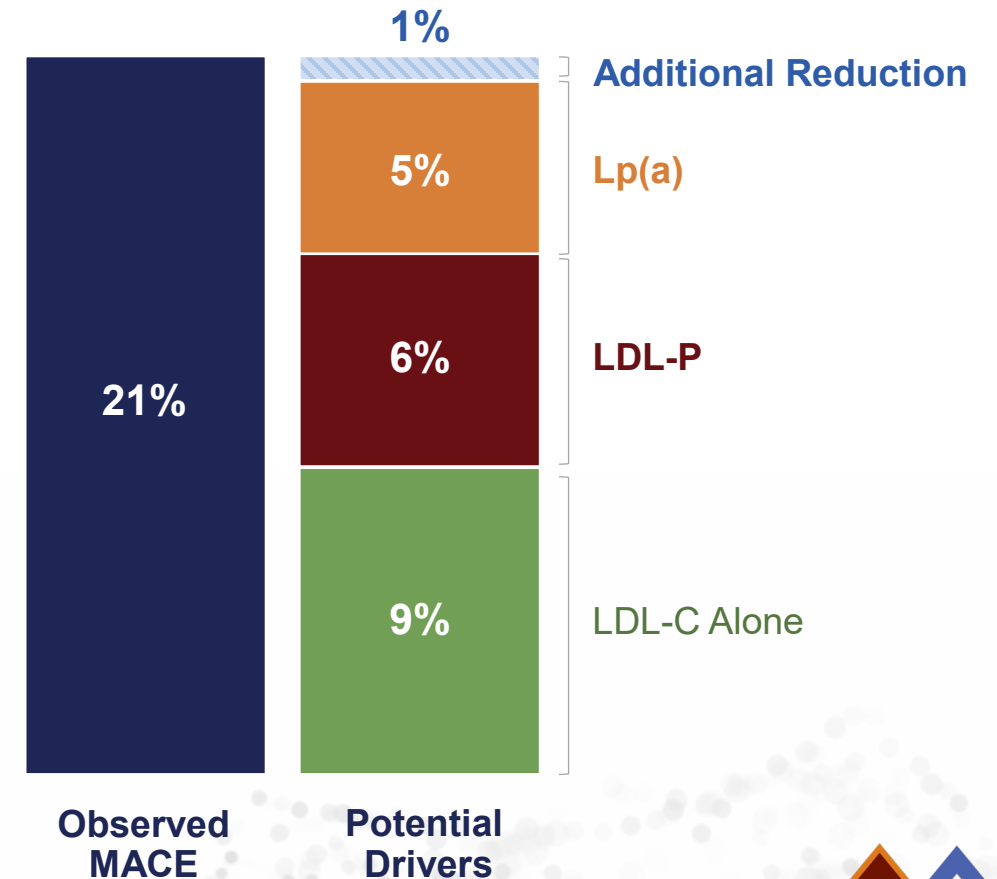


Lp(a) reduction potentially provides an additional 5% MACE reduction



These elements may not be additive as they are correlated to some extent. In this scenario, there is room for additional reduction to be driving incremental gains

## Obicetrapib's Observed MACE Reduction



Note: Actual results may differ from hypothetical calculation.

Source: Cholesterol Treatment Trialists Collaboration; Folse et al. Athero 2014, Quseda et al, Atherosclerosis 35 (2023) 165-177;

Berman, A, Biery, D, Besser, S. et al. Lipoprotein(a) and Major Adverse Cardiovascular Events in Patients With or Without Baseline Atherosclerotic Cardiovascular Disease. JACC. 2024 Mar, 83 (9) 873-886; Data on file

\*\* MACE includes CHD death, myocardial infarction, ischemic stroke and coronary revascularization in adults.

## Overview of Events of Special Interest

|                                                          | Placebo N= 843<br>n (%) | Obicetrapib N= 1685<br>n (%) |
|----------------------------------------------------------|-------------------------|------------------------------|
| AST or ALT > 3 x ULN                                     | 8 (0.9)                 | 10 (0.6)                     |
| Bilirubin > 2 x ULN                                      | 4 (0.5)                 | 2 (0.1)                      |
| CK > 5 x ULN                                             | 3 (0.4)                 | 5 (0.3)                      |
| <b>NODM or worsening of glycemic control</b>             | 338 (40.1)              | 592 (35.1) (p = 0.015)       |
| - AE indicating new/worse type 1 or 2 diabetes           | 30 (3.6)                | 58 (3.4)                     |
| - Initiation of diabetes medication                      | 104 (12.3)              | 186 (11.0)                   |
| - HbA1c ≥ 6.5% (where baseline HbA1c < 6.5%)             | 55 (6.5)                | 84 (5.0)                     |
| - Two consecutive glucose values > 126 mg/dL             | 248 (29.4)              | 459 (27.2)                   |
| - HbA1c increase from baseline > 0.5%                    | 133 (15.8)              | 234 (13.9)                   |
| - Worsening glycemic control                             | 199 (23.6)              | 350 (20.8)                   |
| <b>Renal function worsening</b>                          | 77 (9.1)                | 127 (7.5)                    |
| - eGFR < 30 mL/min/1.73m <sup>2</sup>                    | 13 (1.5)                | 13 (0.8)                     |
| - 25% decrease in eGFR from baseline:                    | 70 (8.3)                | 115 (6.8)                    |
| - Increase of Serum Creatinine ≥ 0.3 mg/dL from baseline | 61 (7.2)                | 91 (5.4)                     |
| Macular degeneration                                     | 0 (0.0)                 | 1 (0.1)                      |

# BROADWAY Designed to be a Predictor of PREVAIL

## BROADWAY

1° endpoint – week 12

N = 2532

Obicetrapib 10 mg (2:1 randomization)

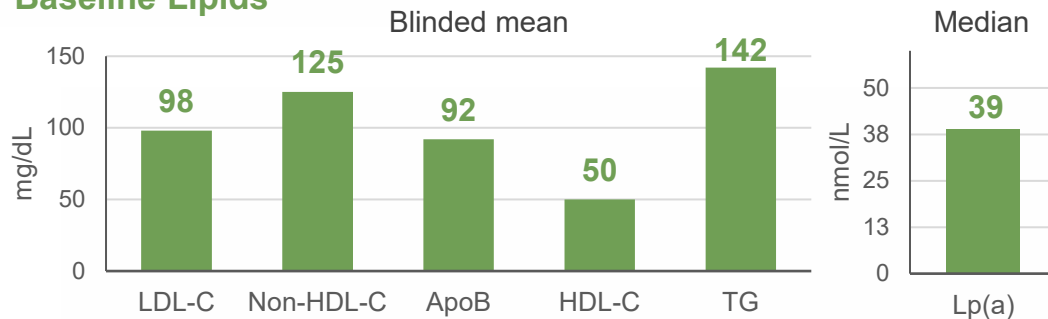
Placebo

13-months

### Key Inclusion Criteria

- ASCVD or HeFH
- LDL-C  $\geq 100$  mg/dL
- LDL-C  $\geq 55$  mg/dL w/risk factors, or
- Maximally tolerated lipid lowering therapy

### Baseline Lipids



### Baseline Lipid Modifying Therapy

- Any statin: 91%
- High intensity statin: 65%
- Ezetimibe: 26%
- PCSK9i: 4%
- SGLT2i: 11%
- GLP-1: 6%

## PREVAIL

LDL-C endpoint

N = 9541

Obicetrapib 10 mg (1:1 randomization)

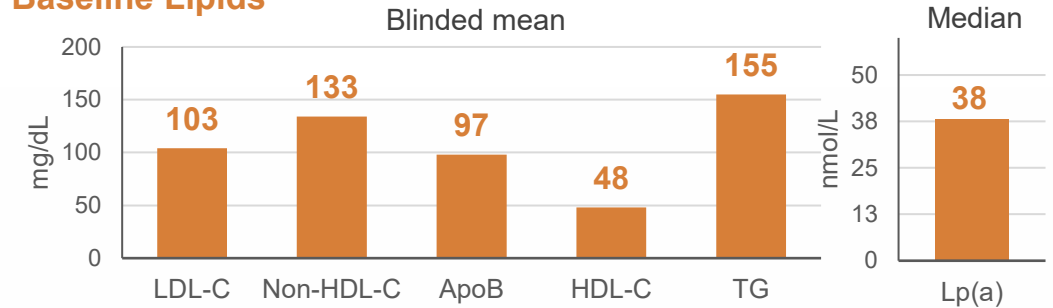
Placebo

54-months

### Key Inclusion Criteria

- ASCVD
- LDL-C  $\geq 100$  mg/dL
- LDL-C  $\geq 55$  mg/dL w/risk factors, or
- Maximally tolerated lipid lowering therapy

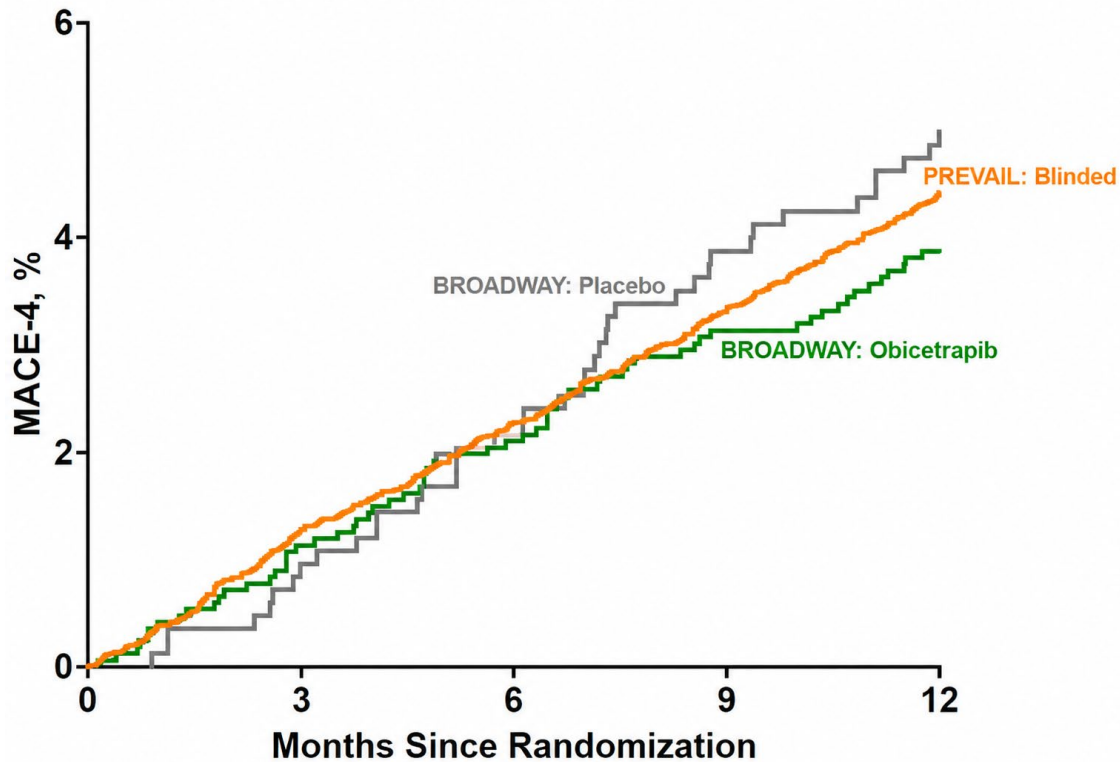
### Baseline Lipids



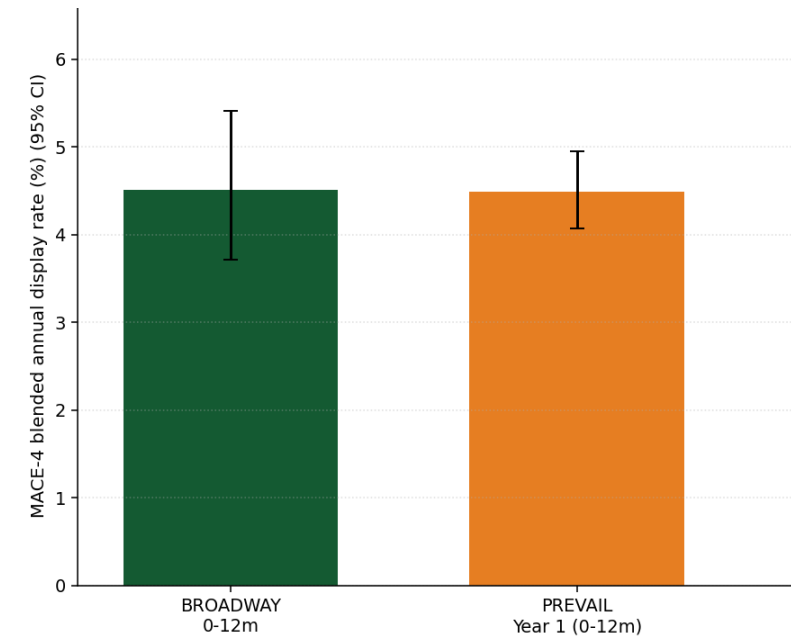
### Baseline Lipid Modifying Therapy

- Any statin: >90%
- High intensity statin: 70%
- Ezetimibe: 23%
- PCSK9i: 2%
- SGLT2i: 14%
- GLP-1: 6%

# Analysis of BROADWAY and PREVAIL Year 1 Event Rates<sup>(1)</sup>



## Blinded/Blended Year 1 Event Rates



4-point MACE: CHD death, Non-fatal myocardial infarction, non-fatal stroke, coronary revascularization

(1) While not powered to detect a MACE benefit, we observed positive MACE-4 data as part of our review of the exploratory endpoint. BROADWAY was not powered to measure MACE benefit and the MACE data presented below may not be predictive of the MACE data we observe in PREVAIL, which has been designed to measure MACE benefit.

PREVAIL data is preliminary and subject to change as MACE events continue to be identified and adjudicated (including with respect to periods presented above). Additionally, Year 1 and Year 2 MACE event trends may not be indicative of future trends.

# PREVAIL Designed to Apply Lessons Learned from Previous CVOTs to Reduce Risk and Demonstrate Obicetrapib's Full Benefit



**Greater LDL-C lowering activity anticipated**  
**Targeting higher baseline LDL-C patients**



**Higher *absolute* LDL-C reduction expected to lead to greater MACE benefit**



**Longer duration of follow up**  
**Targeting higher-risk patient population**



**Maximizes opportunity for MACE reduction**



**Differentiated secondary endpoints**

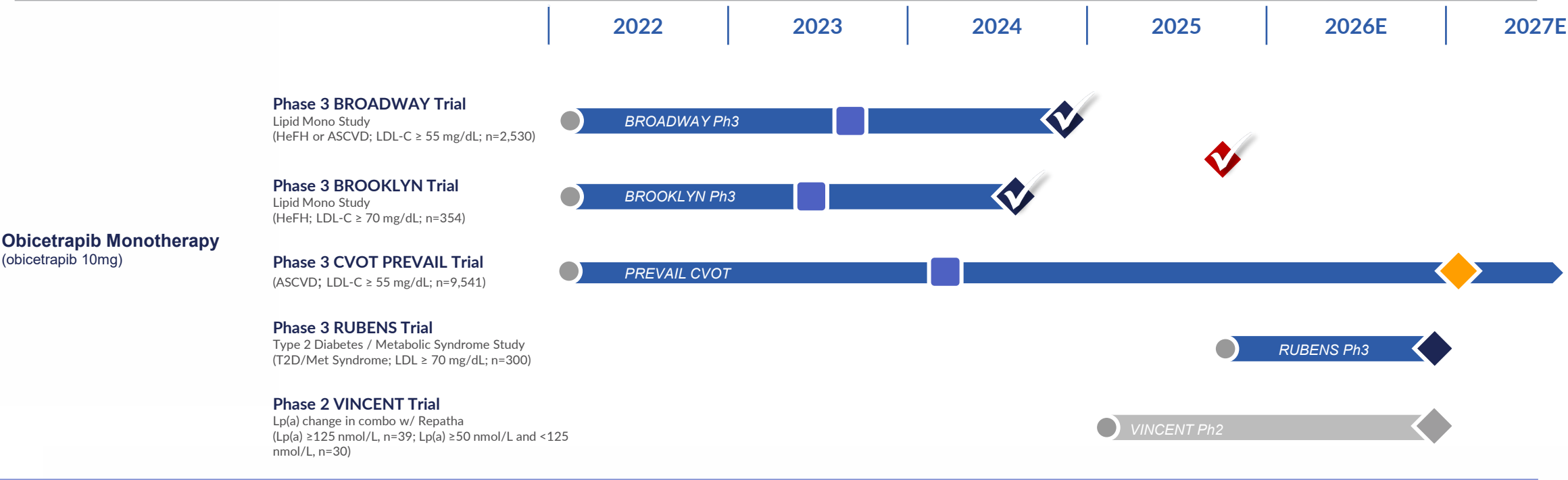


**Potentially enhanced commercial profile vs. other LDL-C lowering agents**

## Primary endpoint selection

Coronary heart disease (CHD) death, non-fatal MI, fatal and non-fatal ischemic stroke, coronary revascularization

# Multiple Data Readouts to Drive Future Momentum



LEGEND

●

Initiation

■

Enrollment Complete

◆

Ph2 (estimated)

◆

Ph3 (estimated)

◆

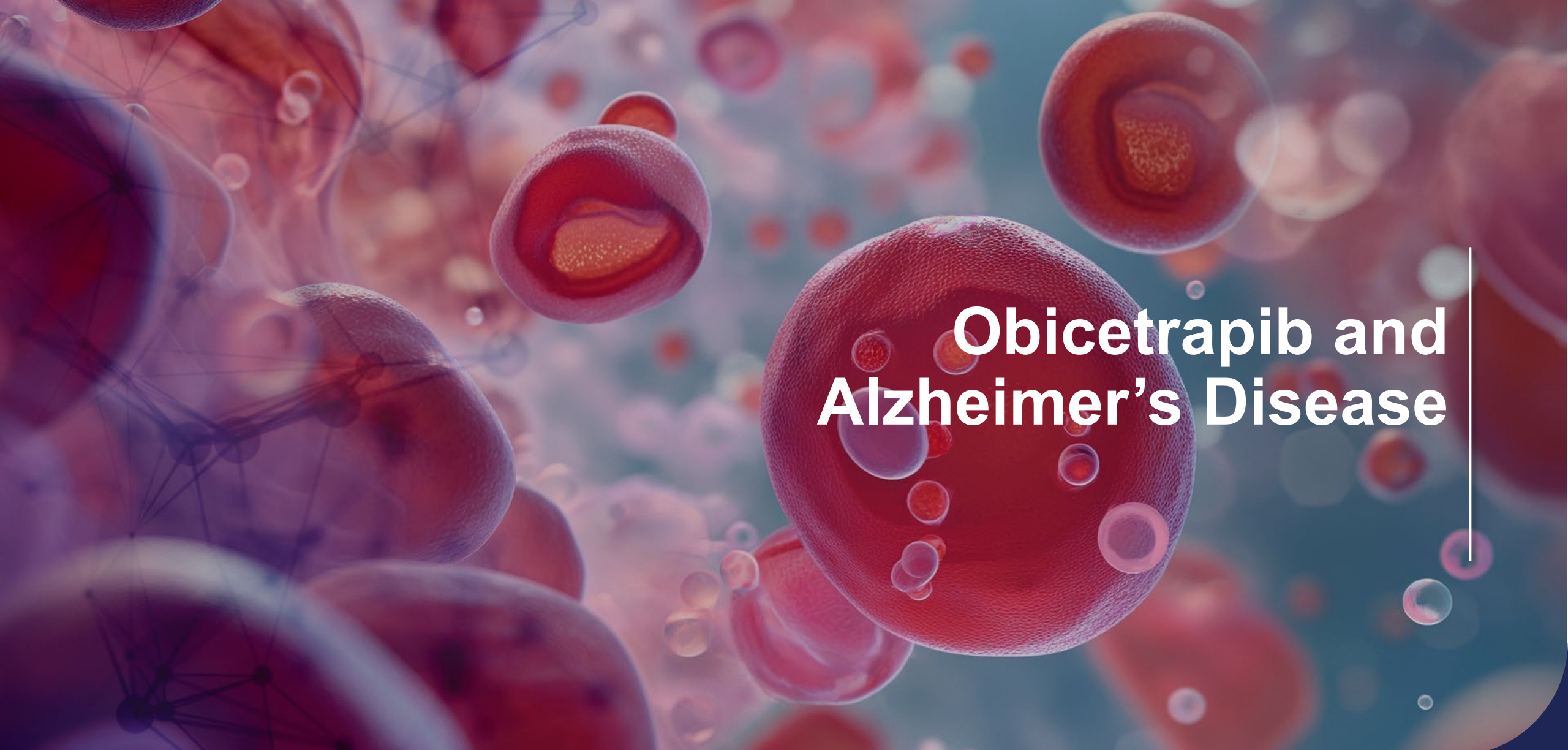
CVOT Interim

◆

European Regulatory Filings\*

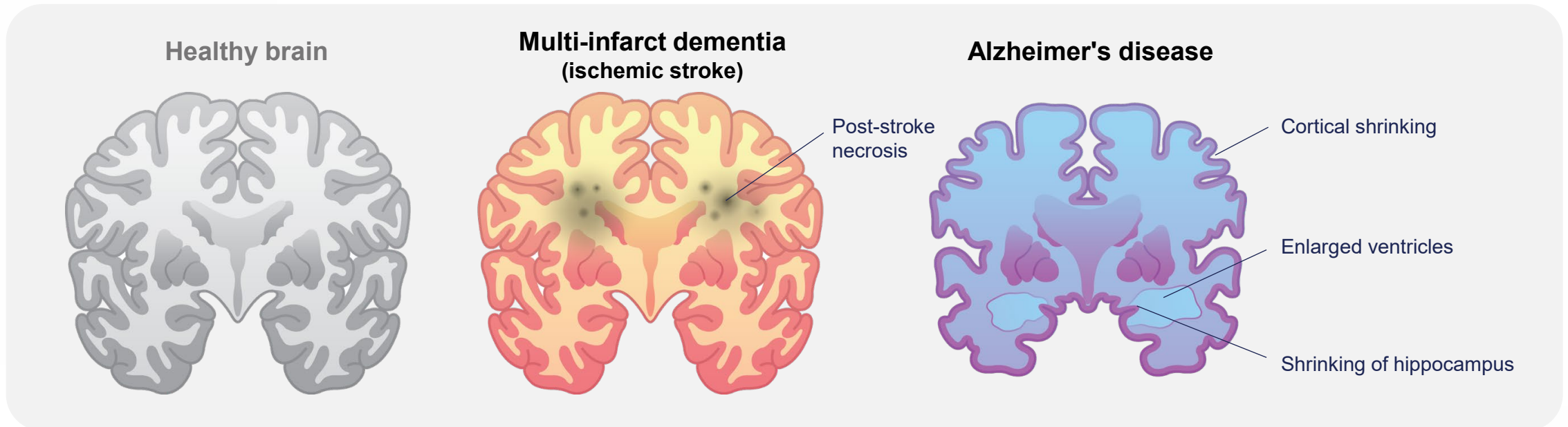
28 \*MAAs were filed and accepted for review by EMA, UK and Switzerland regulatory authorities  
Note: Other than as noted, the pipeline represents trials that are currently ongoing. 2026 and beyond are projections and subject to inherent limitations. Actual results may differ from expectations.





# Obicetrapib and Alzheimer's Disease

# CETPi Bears Therapeutic Potential for Both Multi-infarct Dementia and Alzheimer's Disease

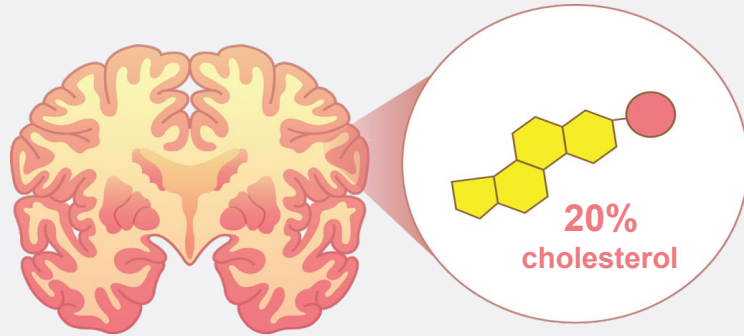


There is emerging evidence of a vascular component in multi-infarct dementia, **implying that LDL-lowering would help reduce atherosclerosis in the brain** <sup>(1)</sup>

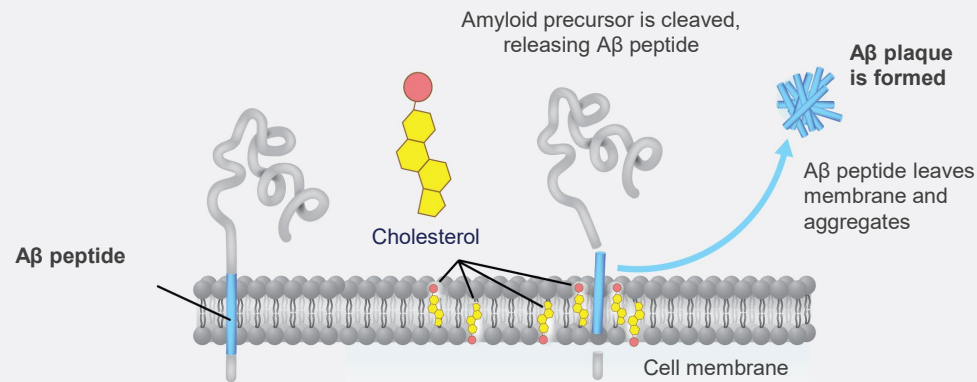
Alzheimer's disease results in the **generalized degeneration of the brain**, with over 50M people affected WW and an economic burden of >\$1 trillion <sup>(2)</sup>

# Elevated Cholesterol Levels in the Brain Precede Formation of Amyloid-Beta (A $\beta$ ) Plaques

The brain is the most cholesterol-rich organ; it contains only 2% body mass but has 20% of the body's cholesterol



## A $\beta$ plaque formation inside AD brain cells



**During early development**, oligodendrocytes synthesize large quantities of cholesterol for myelination

**In adults**, when myelination is complete, glial cells and, to a lesser extent, neurons account for the steady-state production of cholesterol

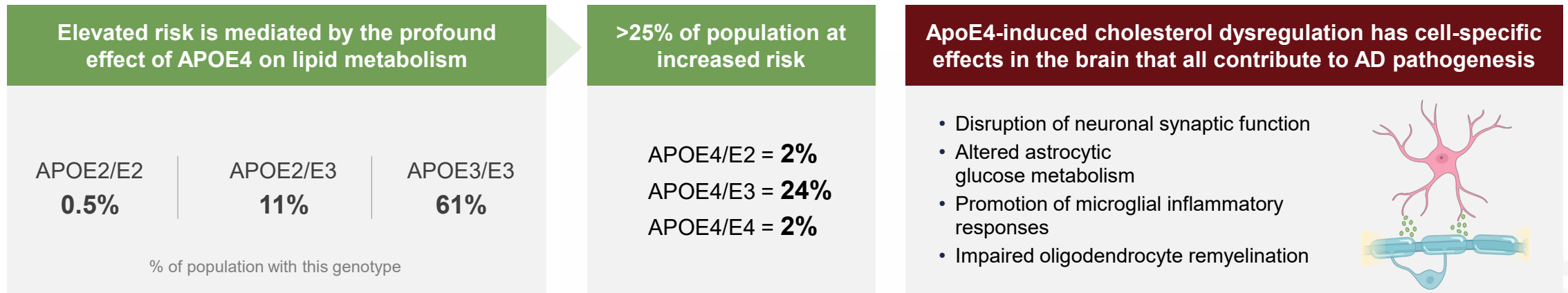
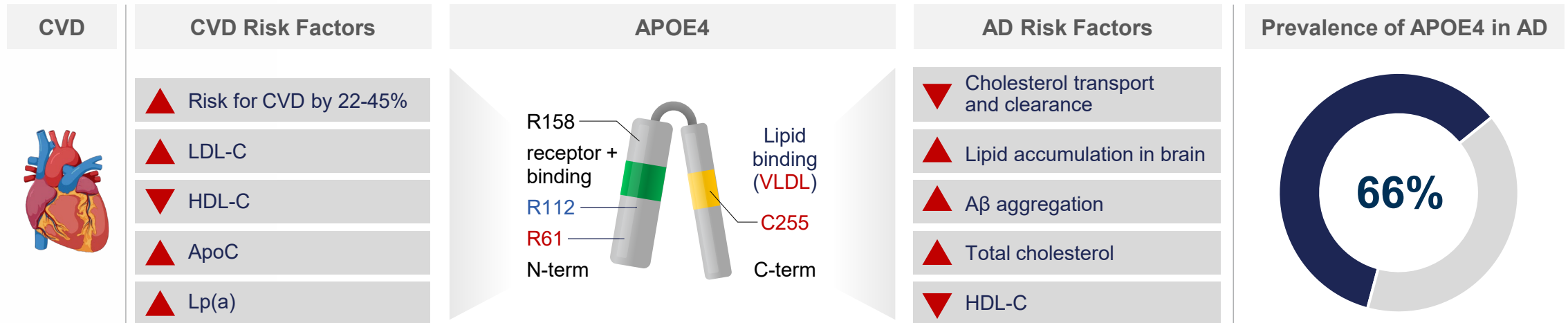
**The brain uses a unique mechanism** for cholesterol recycling and redistribution

- Involves an ApoE-mediated lipoprotein pathway unique to the CNS whereby cholesterol is turned over
- Excess oxysterols are ultimately delivered to the liver for secretion into the bile

**Cholesterol itself does not exit the brain** but instead is converted to a metabolite, 24S-hydroxycholesterol

- 24S-hydroxycholesterol is a proxy for excess cholesterol that will be measured in our phase 2a clinical trial

# ApoE4 is a Risk Factor for Both CVD and Alzheimer's Disease<sup>1-5</sup>



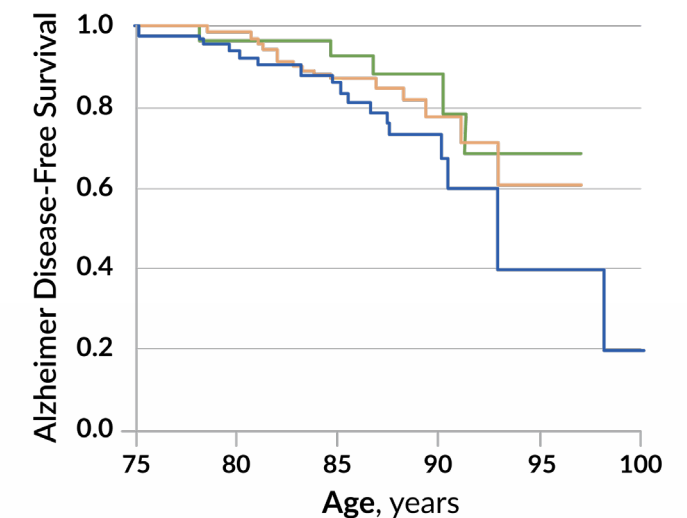
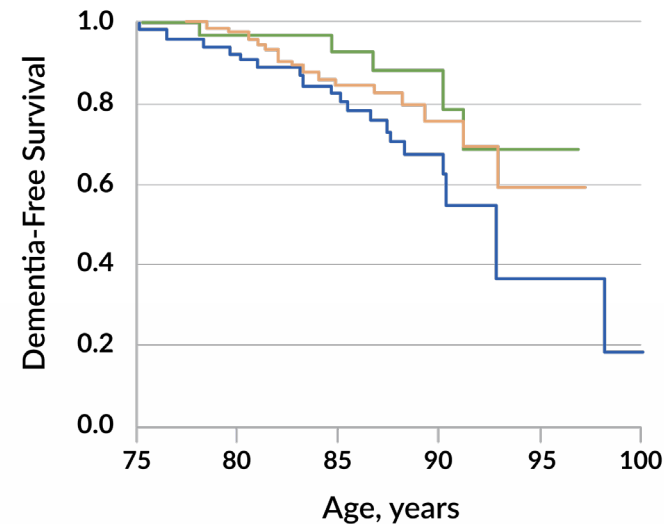
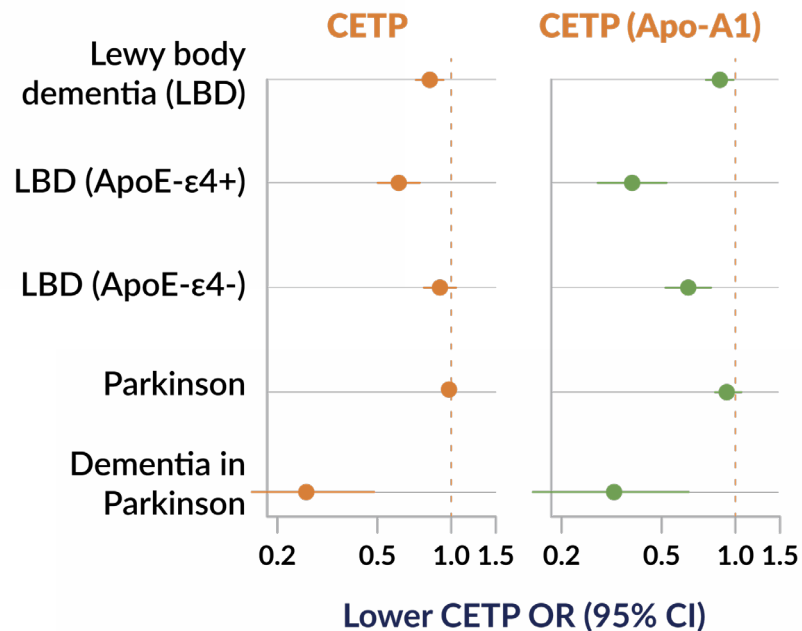
# CETP Loss-of-Function (LoF) Genotype May be Associated with Slower Memory Decline and Lower AD Risk



CETP's potential involvement in CNS cholesterol homeostasis is supported by genomic data



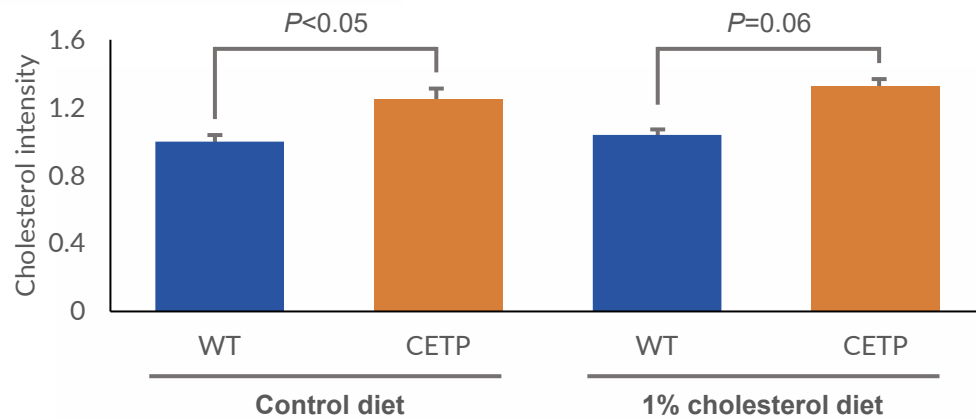
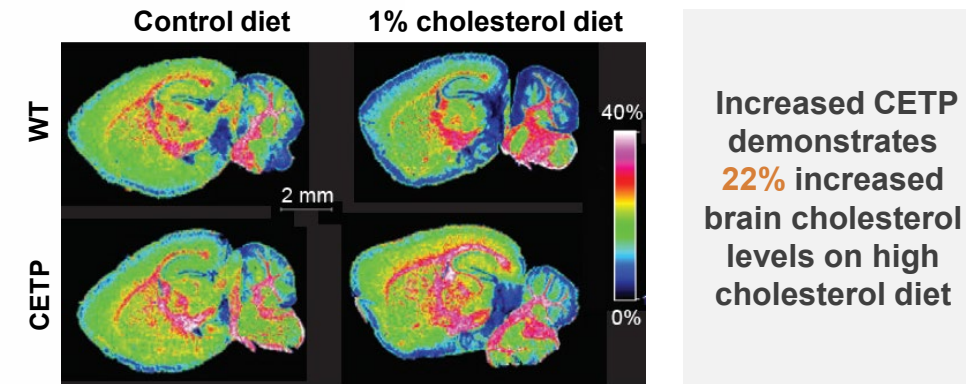
CETP LoF may be associated with lower CETP activity & a corresponding increase in HDL levels



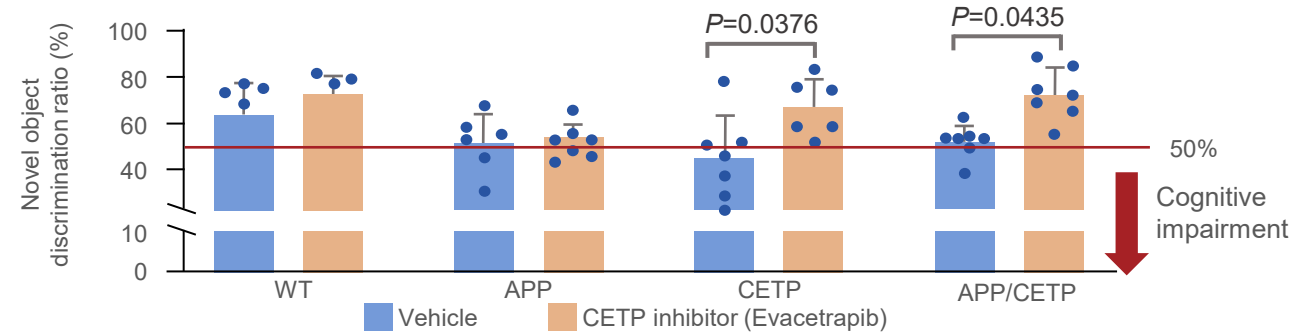
— 2 LoF CETP genes — 1 LoF CETP gene — Normal CETP

# CETP Knock-in Mice Have Increased Brain Cholesterol Levels and CETPi Observed to Rescue Cognition in Preclinical Models of CETP-induced AD

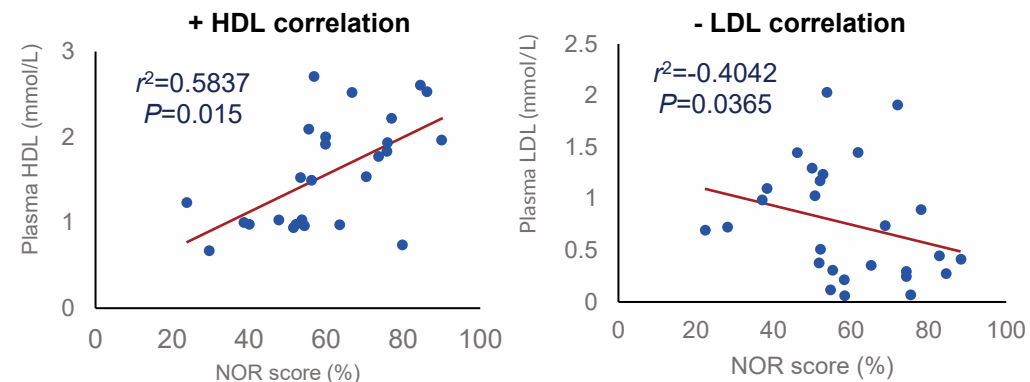
CETP transgenic mice have lower plasma HDL-C levels and increased brain cholesterol levels<sup>1</sup>



CETP inhibition with evacetrapib led to cognitive improvement<sup>2,3</sup>



Plasma Lipoproteins Were Correlated With NOR Score



**Positive correlation** observed between NOR score and HDL quantification in CETP and APP/CETP-expressing female mice

**Negative correlation** observed between NOR score and LDL quantification in CETP and APP/CETP-expressing female mice

A separate study demonstrated that **CETP inhibitors can enter the brain**<sup>4</sup>

AD, Alzheimer's disease; APP, amyloid precursor protein; CETP, cholesterol ester transfer protein; CETPi, CETP inhibitor; HDL, high-density lipoprotein; HDL-C, high-density lipoprotein cholesterol; LDL, low-density lipoprotein; NOR, novel object recognition; WT, wild-type.

1. Oestereich F et al. *J Lipid Res.* 2022;63(9):100260. 2. Phénix J et al. *Alzheimer's Dement.* 2021;17(suppl3):e051134. 3. Phénix J et al. Poster presented at: Alzheimer's Association International Conference; July 26-30, 2021; Denver, USA. 4. Phénix J et al. *Front Pharmacol.* 2023;14:1171937.

# Obicetrapib on Top of Max Tolerated Lipid-Modifying Therapy: Alzheimer's Biomarker Sub-study

**Objective:** To evaluate the effect of obicetrapib on biomarkers related to Alzheimer's Disease compared to placebo over 12-months

**Study design:** Randomized, double-blind, placebo-controlled

## Main Inclusion Criteria:

Have a fasting serum LDL-C at Screening (Visit 1) as follows:

- Have a fasting serum LDL-C  $\geq 55$  mg/dL ( $\geq 1.4$  mmol/L) to  $<100$  mg/dL ( $<2.6$  mmol/L) OR non-HDL-C  $\geq 85$  mg/dL ( $\geq 2.2$  mmol/L) to  $<130$  mg/dL ( $<3.4$  mmol/L) with at least 1 risk enhancer

OR

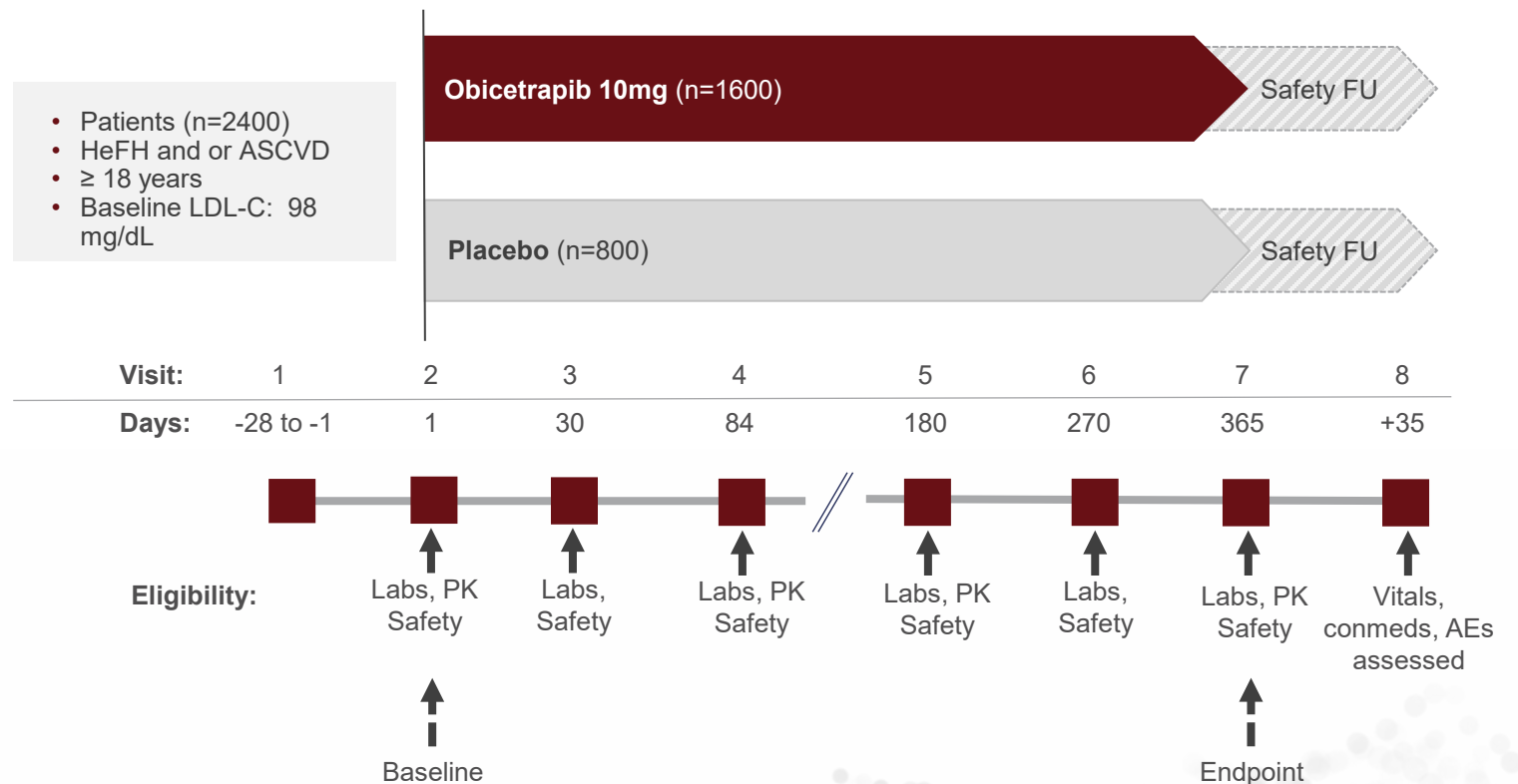
- Have a fasting serum LDL-C  $\geq 100$  mg/dL ( $\geq 2.6$  mmol/L) OR non-HDL-C  $\geq 130$  mg/dL ( $\geq 3.4$  mmol/L).

## Main Exclusion Criteria:

- CV disease  $< 3$  months
- HoFH
- Uncontrolled hypertension

## Biomarker Endpoints:

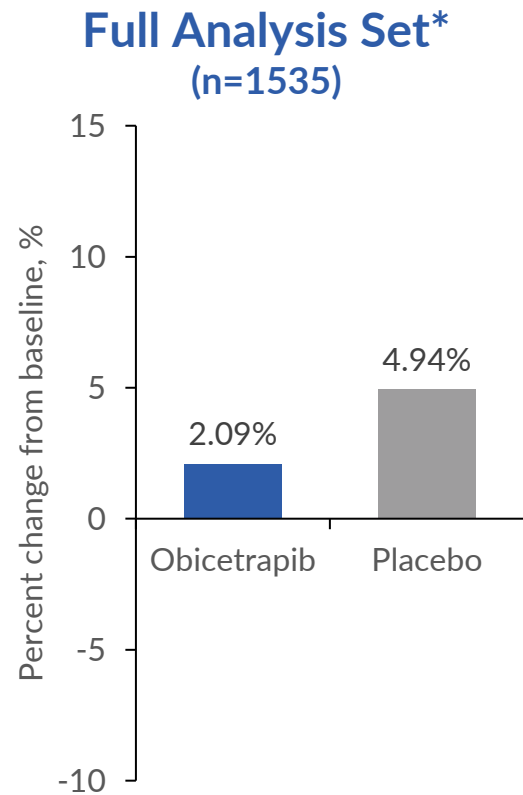
- p-tau217
- Ratio of A $\beta$ 42 to A $\beta$ 40 (A $\beta$ 42/A $\beta$ 40)
- GFAP
- NFL
- p-tau181



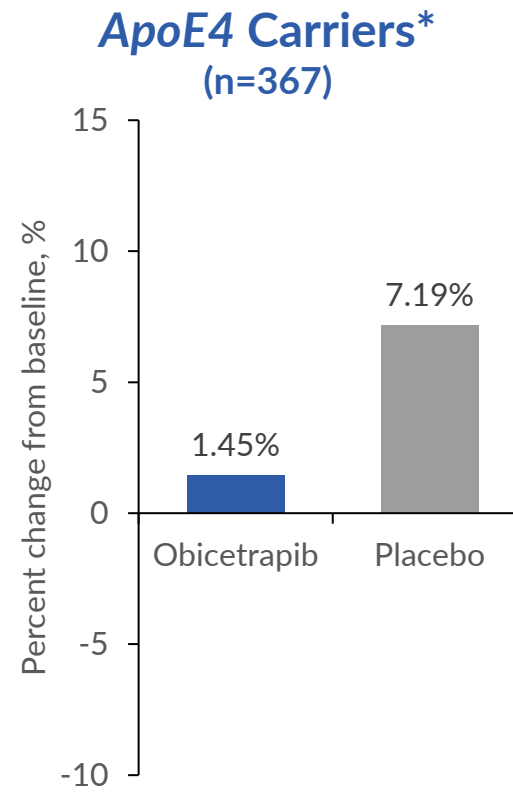
Note: Davidson MH et al. 2025. Manuscript in preparation.

# Significant Reduction in p-tau217 Progression Over 12 Months Observed with Greater Reductions as ApoE4 Risk Factors Increase

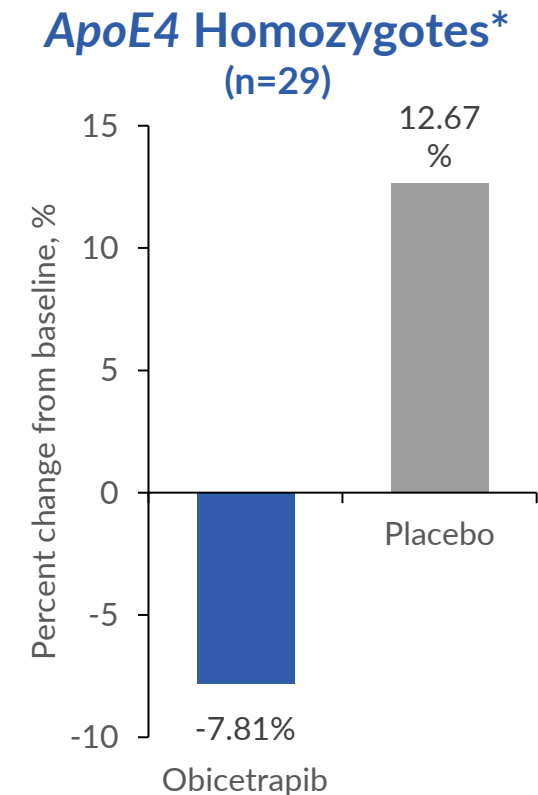
**2.84% benefit between  
obicetrapib and placebo (P=0.025)**



**5.74% treatment benefit  
in ApoE4 carriers (P=0.022)**

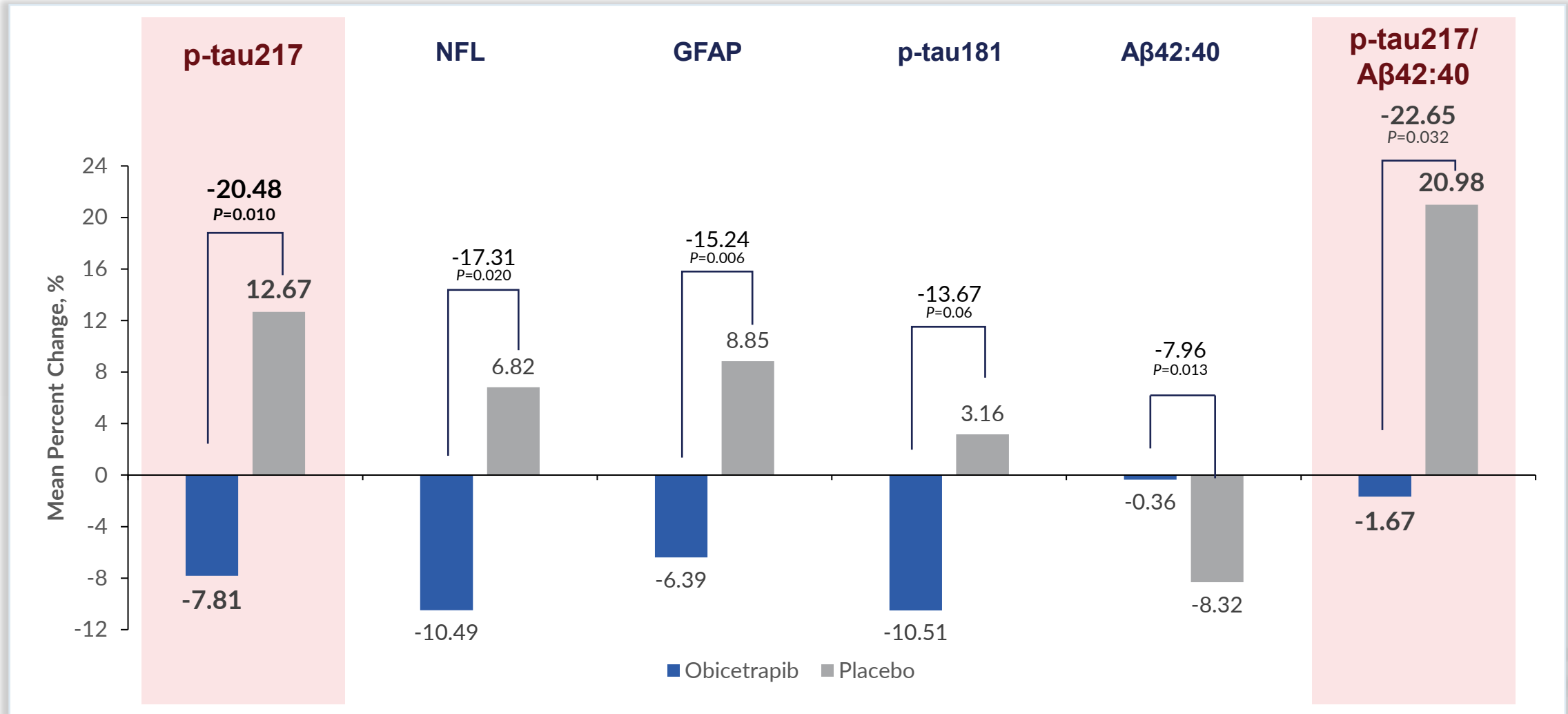


**20.5% treatment benefit in  
ApoE4 homozygotes (P=0.010)**



\*Results reflect adjustment for mean-centered baseline p-tau217 and mean-centered age. Participants with known ApoE status (based on phenotypic analysis), baseline p-tau217 above the lower limit of quantification (LLQ), and available end-of-study p-tau217 were included in the current analyses. ApoE, apolipoprotein E; p-tau, phosphorylated tau. ApoE4 carriers defined as ApoE3/E4 or ApoE4/E4, based on phenotypic analysis Davidson MH et al. 2025. Journal of Prevention of Alzheimer's Disease

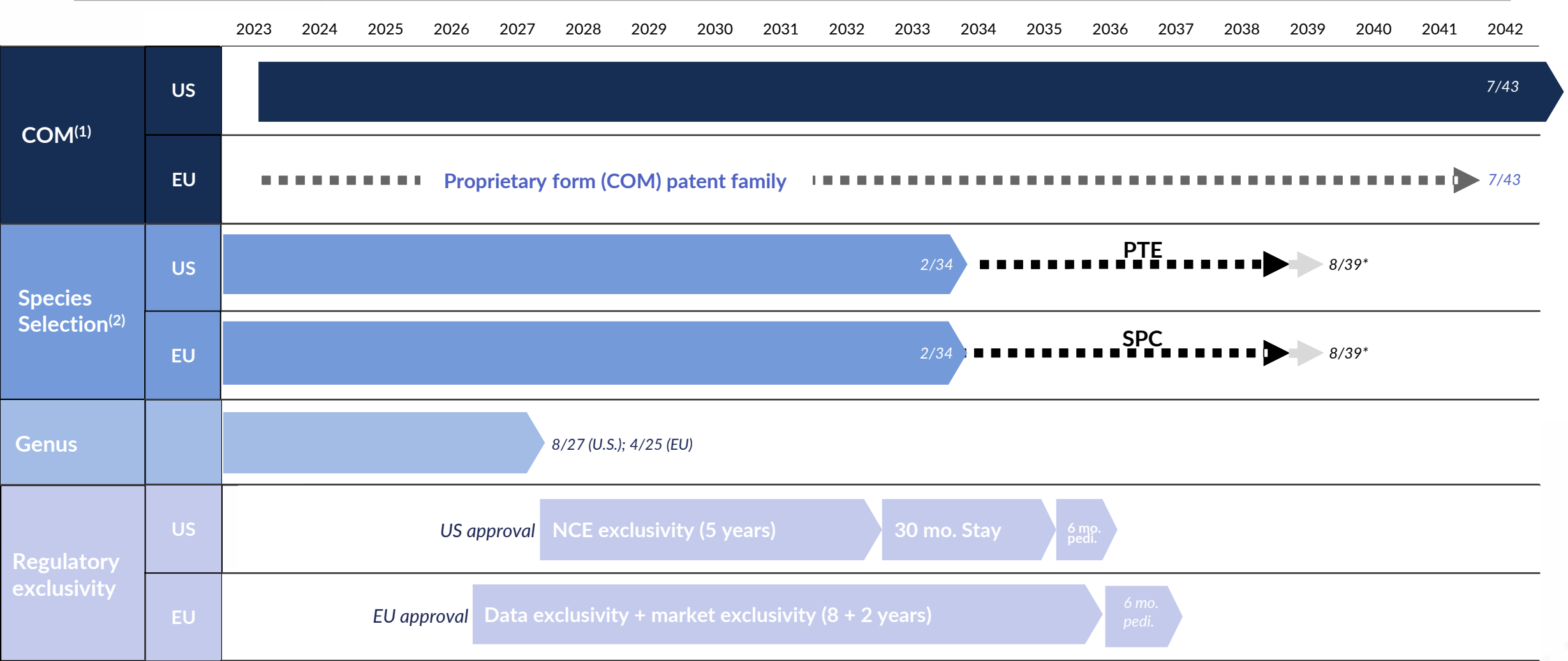
# Obicetrapib's Impact on Additional AD Biomarkers in E4/E4 Carriers



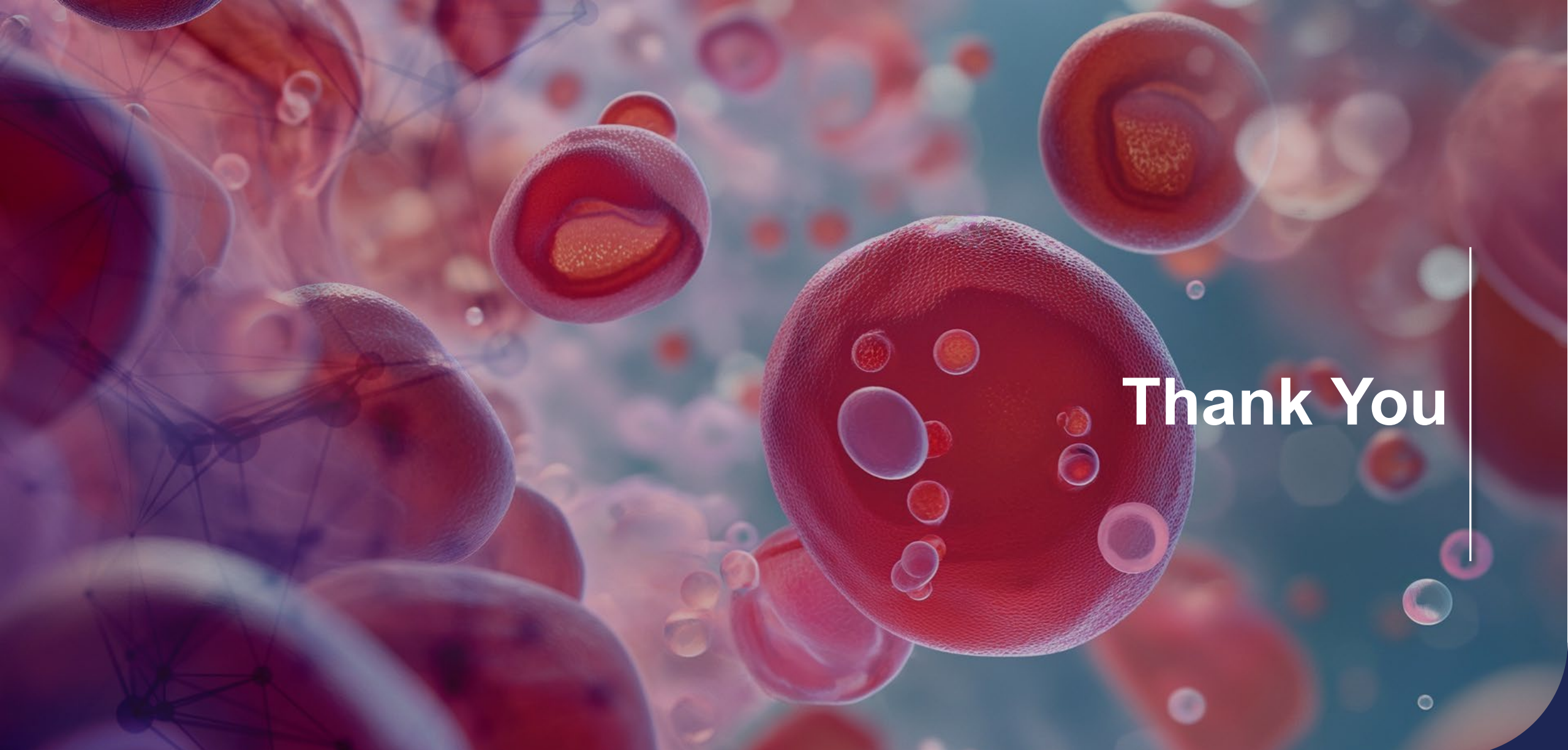


# Intellectual Property

# Comprehensive Patent Portfolio with Composition of Matter IP into 2043



Note: Regulatory approval and exclusivity dates are illustrative and shown for information purposes only; actual results may differ.  
Filled colors = granted patents & dotted lines = pending patents; one patent only to be selected for SPC/PTE; an earlier US approval leads to earlier regulatory expiry & shorter PTE;  
\*including pediatric extension 6m; actual results may differ from expectations. (1) US 12,006,305. (2). Low dose/ species selection patents US 10,653,692, US 11,013,742, US 11,642,344; US 12,186,315 statin combo patent US 10,300,059



Thank You