

# CATALYST BIOSCIENCES

Corporate Overview  
3 March 2020



# Forward looking statements

This presentation includes forward-looking statements that involve substantial risks and uncertainties. All statements included in this presentation, other than statement of historical facts, are forward-looking statements. Examples of such statements include, but are not limited to, potential markets for MarzAA, DalcA and CB 2782-PEG, potential benefits of subcutaneous dosing, potential use of MarzAA as a subcutaneous therapy for patients with hemophilia A or B with inhibitors and other bleeding disorders, potential use of DalcA as a subcutaneous therapy for patients with hemophilia B, potential benefits of CB 2679d-GT as gene therapy, clinical trial results, plans for a registrational trial for MarzAA in second half of 2020, plans for final Phase 2b clinical trial data for DalcA in the second quarter of 2020, plans for non-human primate data for CB 2679d-GT in the second quarter of 2020, and potential milestone and royalty payments from Biogen. Actual results or events could differ materially from the plans, expectations and projections disclosed in these forward-looking statements.

Various important factors could cause actual results or events to differ materially, including, but not limited to, the risk that additional human trials will not replicate the results from earlier trials or animal studies, that potential adverse effects may arise from the testing or use of MarzAA or DalcA, including the generation of antibodies, which has been observed in patients treated with DalcA, that clinical trials will take longer than anticipated to be completed, that costs required to develop or manufacture the Company's products will be higher than anticipated, that Biogen will discontinue development of CB 2782-PEG, competition and other factors that affect our ability to establish collaborations on commercially reasonable terms and other risks described in the "Risk Factors" section of the Company's annual report on Form 10-K filed with the Securities and Exchange Commission on February 20, 2020, and in other filings with the Securities and Exchange Commission. The Company does not assume any obligation to update any forward-looking statements, except as required by law.

## Essential Medicines – Superior Outcomes

### Late-Stage Asset

SQ Marzeptacog alfa  
(activated)  
MarzAA (FVIIa)

**Phase 3 Ready**

### Hemophilia

SQ MarzAA (FVIIa)

SQ Dalcinonacog  
alfa – DalcA (FIX)

Factor IX Gene Therapy

Factor Xa

### Complement

IVT Anti-C3 Dry AMD  
CB 2782-PEG



SQ Systemic  
Complement  
Inhibitors

Protease Engineering Platform

# Pipeline

## Hemostasis

**SQ Marzeptacog alfa (activated) "MarzAA"**  
Hemophilia & bleeding disorders (rFVIIa)

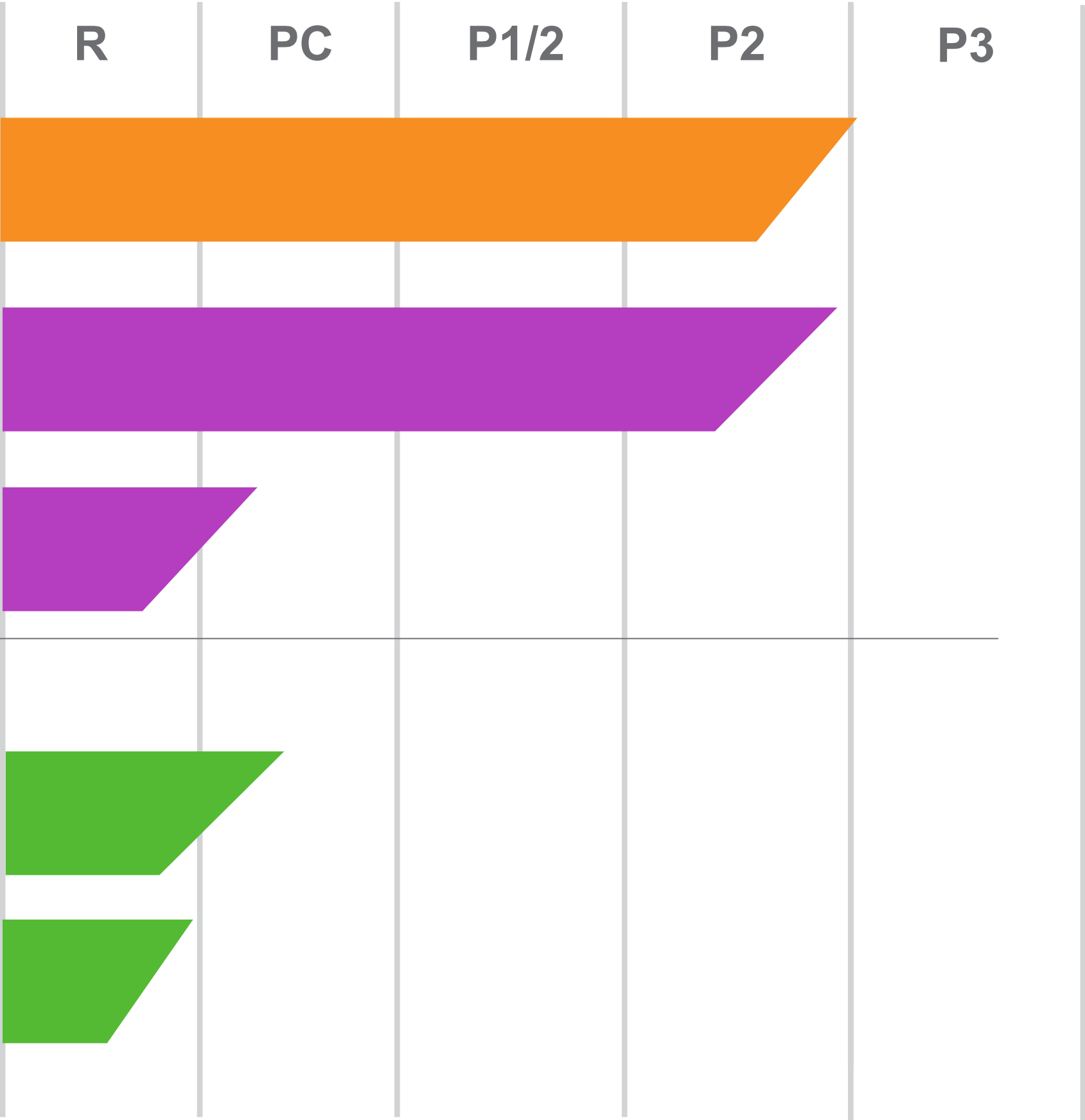
**SQ Dalcinonacog alfa "DalcA"**  
Hemophilia B (rFIX)

**FIX-Gene Therapy**  
Hemophilia B (CB 2679d-GT)

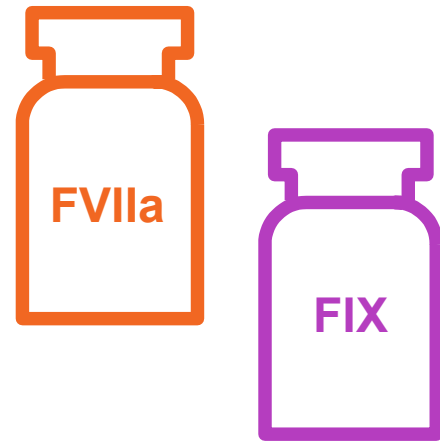
## Complement

**IVT CB 2782-PEG**  
anti-C3 protease for Dry AMD

**SQ Systemic complement inhibitors**



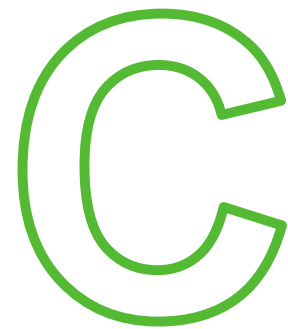
# Investment highlights



Novel subcutaneous factors with orphan drug designation, **MarzAA** & **DalcA** – SQ P2b clinical efficacy demonstrated



Multi-billion-dollar market opportunities



**Anti-C3** collaboration with Biogen

**SQ systemic complement inhibitors** research program



Experienced team



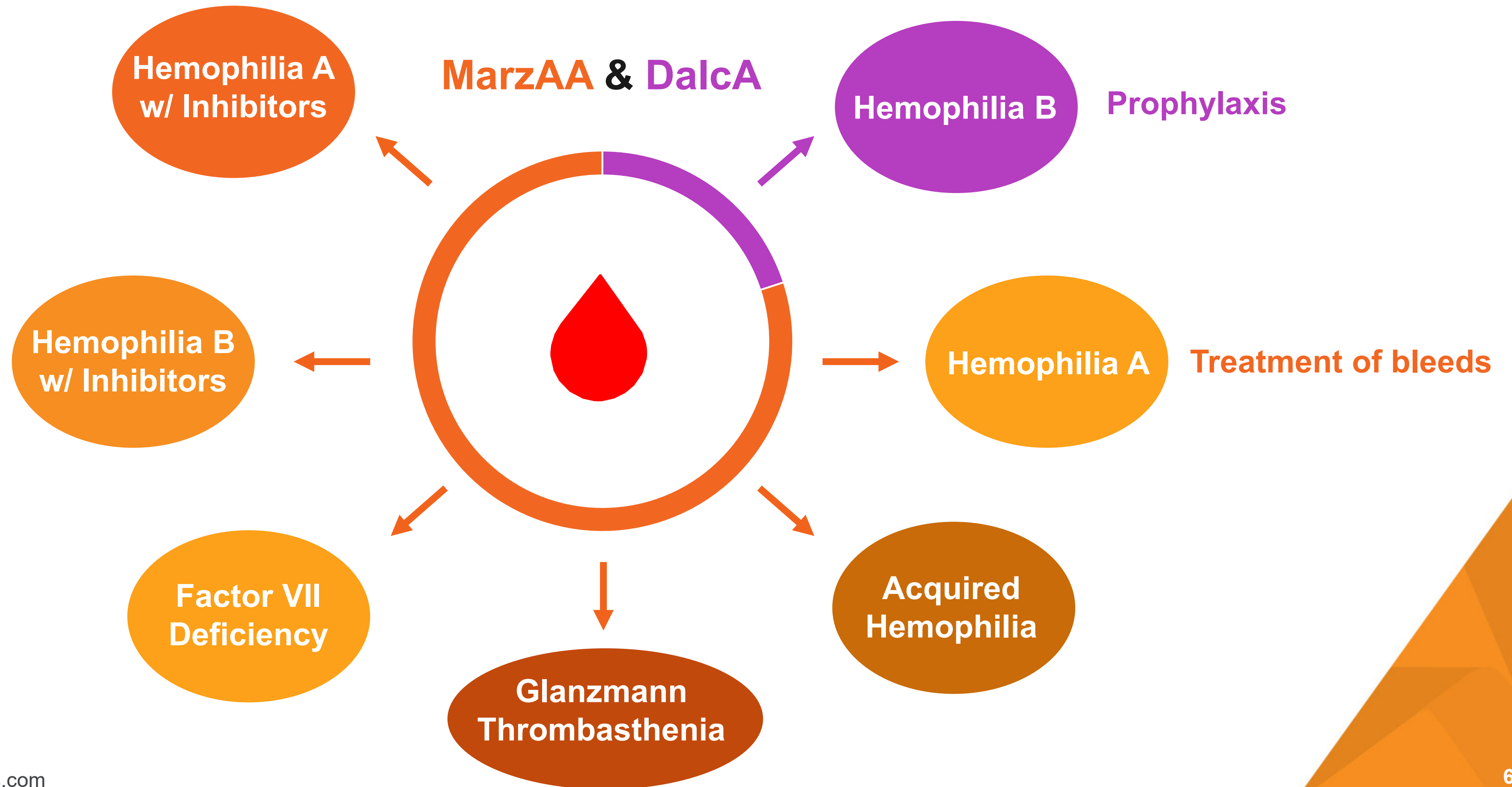
~177 worldwide patents – CBIO retains full ownership of all compounds



Strong balance sheet, ~\$120 M cash

# Addressing unmet needs in orphan bleeding disorders

**SQ treatment of bleeds and prophylaxis – \$4B+ market**





# The Catalyst Biosciences subcutaneous solution

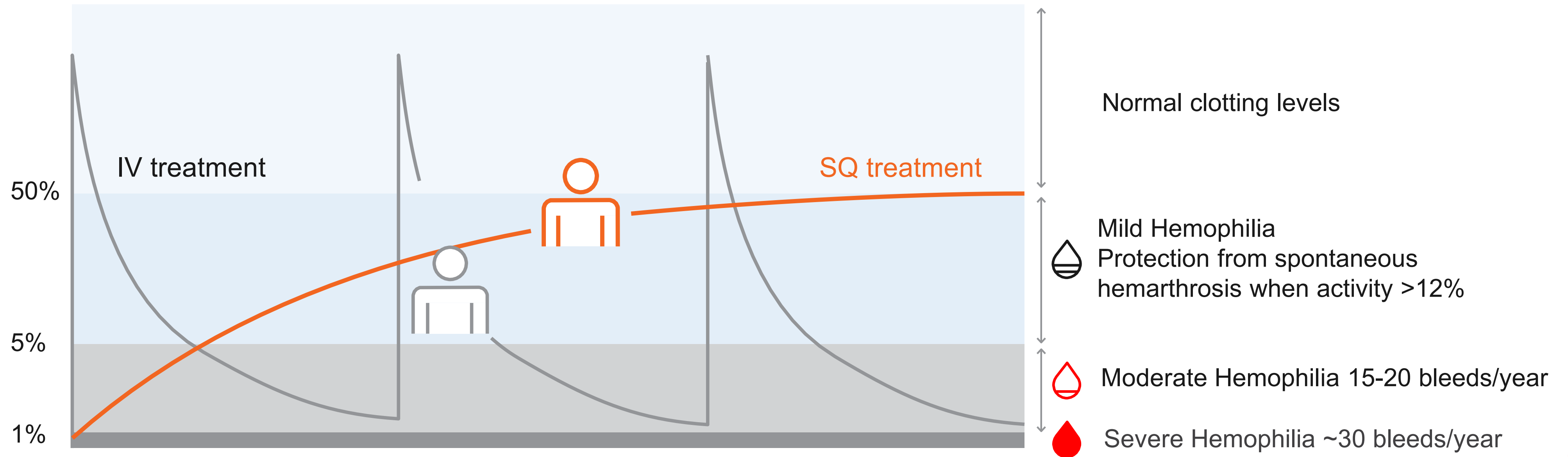


## Our highly potent candidates

- + Quick & simple self-administered SQ injection
- + SQ dosing is the future in hemophilia and other rare hematology indications
- + Ideal for pediatric patients
- + Significantly increases half-life
- + Much higher & more stable factor levels for prophylaxis
- + Enable SQ treatment of bleeding

# The new standard in hemophilia prophylaxis

**Patients in high mild range are protected from spontaneous bleeds**

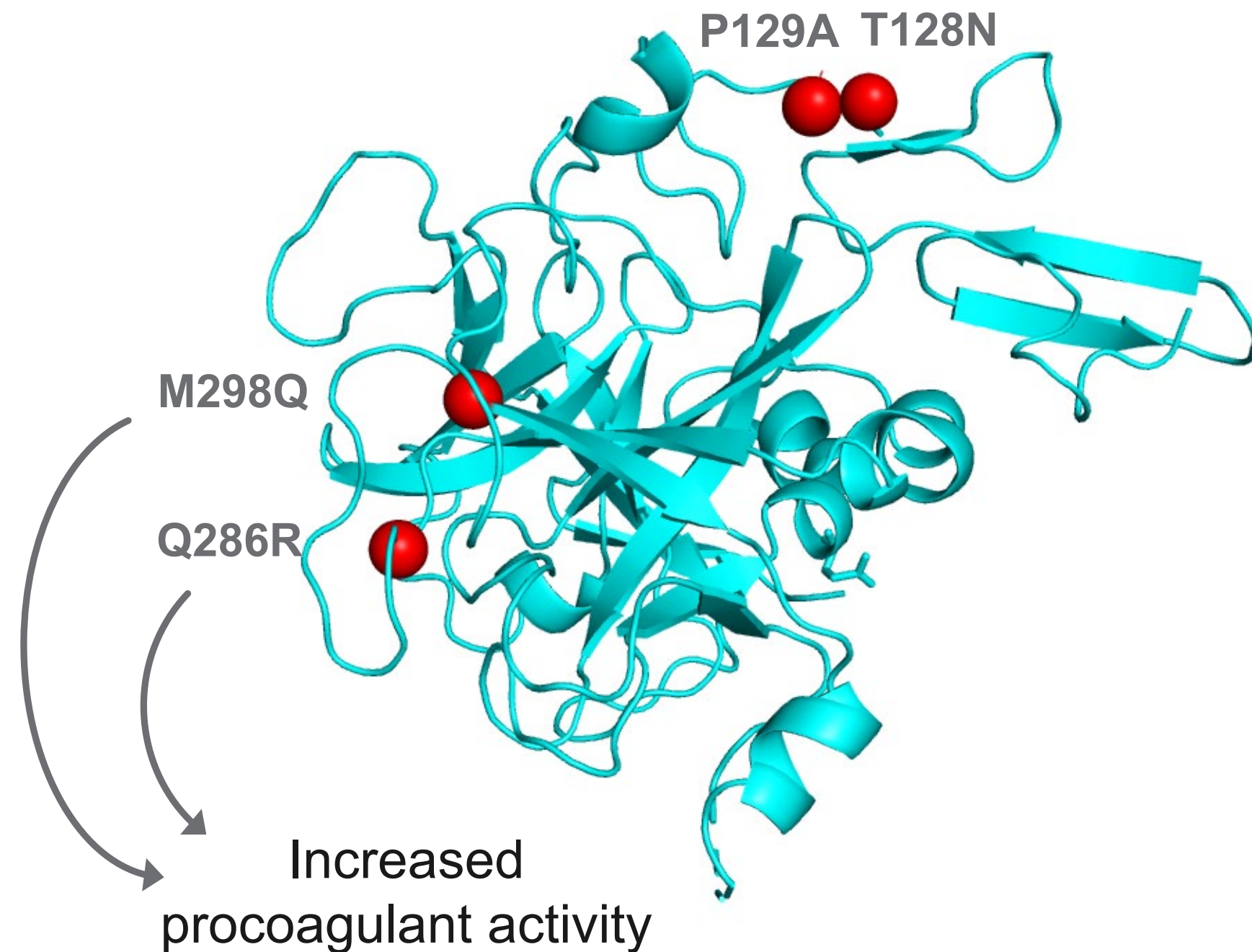


- + Our concept of prophylactic treatment is to keep severe & moderate hemophilia patients in the high mild range
- + Subcutaneous factor treatments build up over time, offering long-term stability in clotting levels



# Marzeptacog alfa (activated): MarzAA rFVIIa

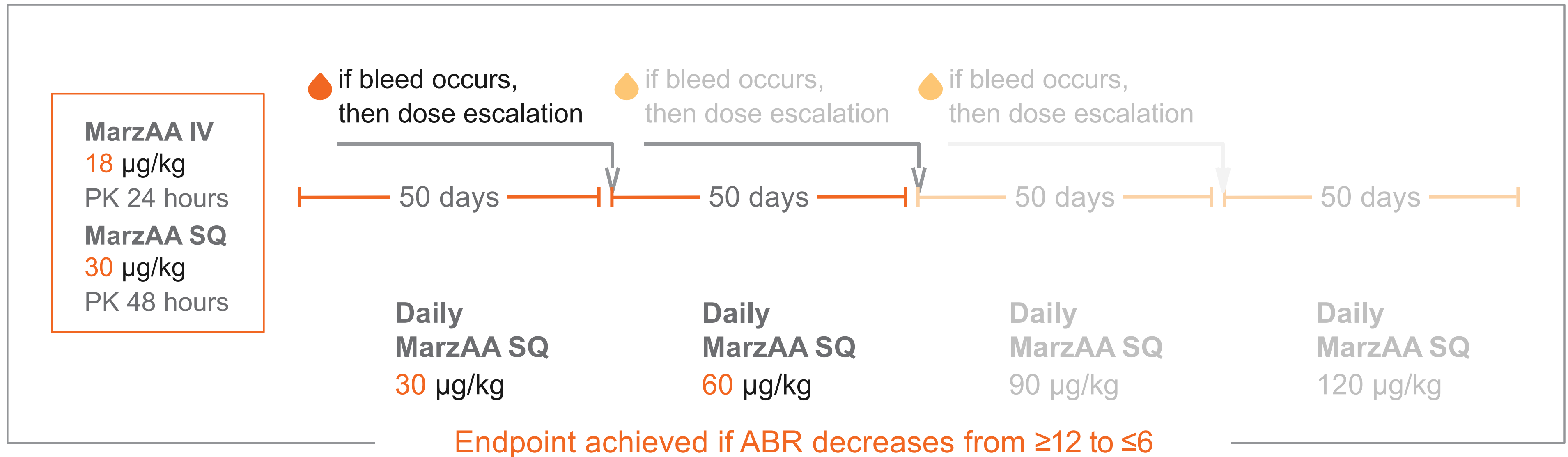
**SQ prophylaxis and SQ treatment of a bleed are clear unmet needs in hemophilia and other bleeding disorders**



- + Four engineered amino acid substitutions within the FVIIa protein
- + 9-fold more potent catalytic activity than NovoSeven RT
- + **Allows subcutaneous dosing**
- + Half-life prolonged when using subcutaneous dosing

**Orphan Drug Designation  
Granted in the US and EU**

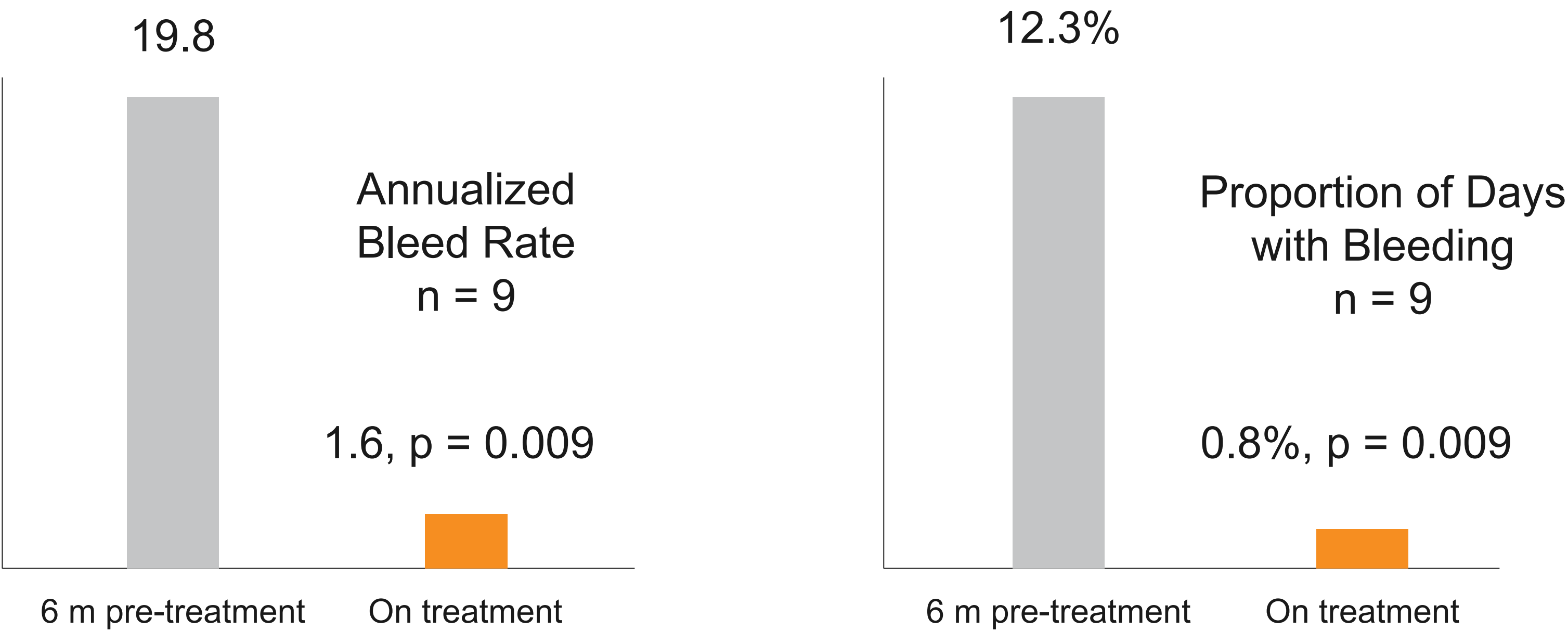
# MarzAA phase 2/3 SQ clinical trial MAA-201 design



- + Patients with documented annual bleeding rate (ABR)  $>12$
- + Open label SQ study with individual dose escalation if needed in Hemophilia A or B with inhibitors
- + Primary endpoint: reduction in annualized bleed rate **at final dose level**
- + Secondary endpoints: safety and tolerability, inhibitor formation

# MarzAA Phase 2 demonstrates clinical efficacy

Greater than 90% reduction in all bleeding; Median ABR zero  
7/9 subjects had no bleeding at final dose level



Safe & well tolerated, ~1% ISRs (6/517 SQ doses) and no ADAs

# In a world of SQ prophylaxis

## Patients & KOLs want SQ treatment of a bleed

Individuals on Hemlibra®  
have breakthrough bleeds

NovoSeven® is safe but is  
administered IV

FEIBA lacks a safety margin  
and is administered IV

## SQ MarzAA meets the profile for an **Ideal Solution**

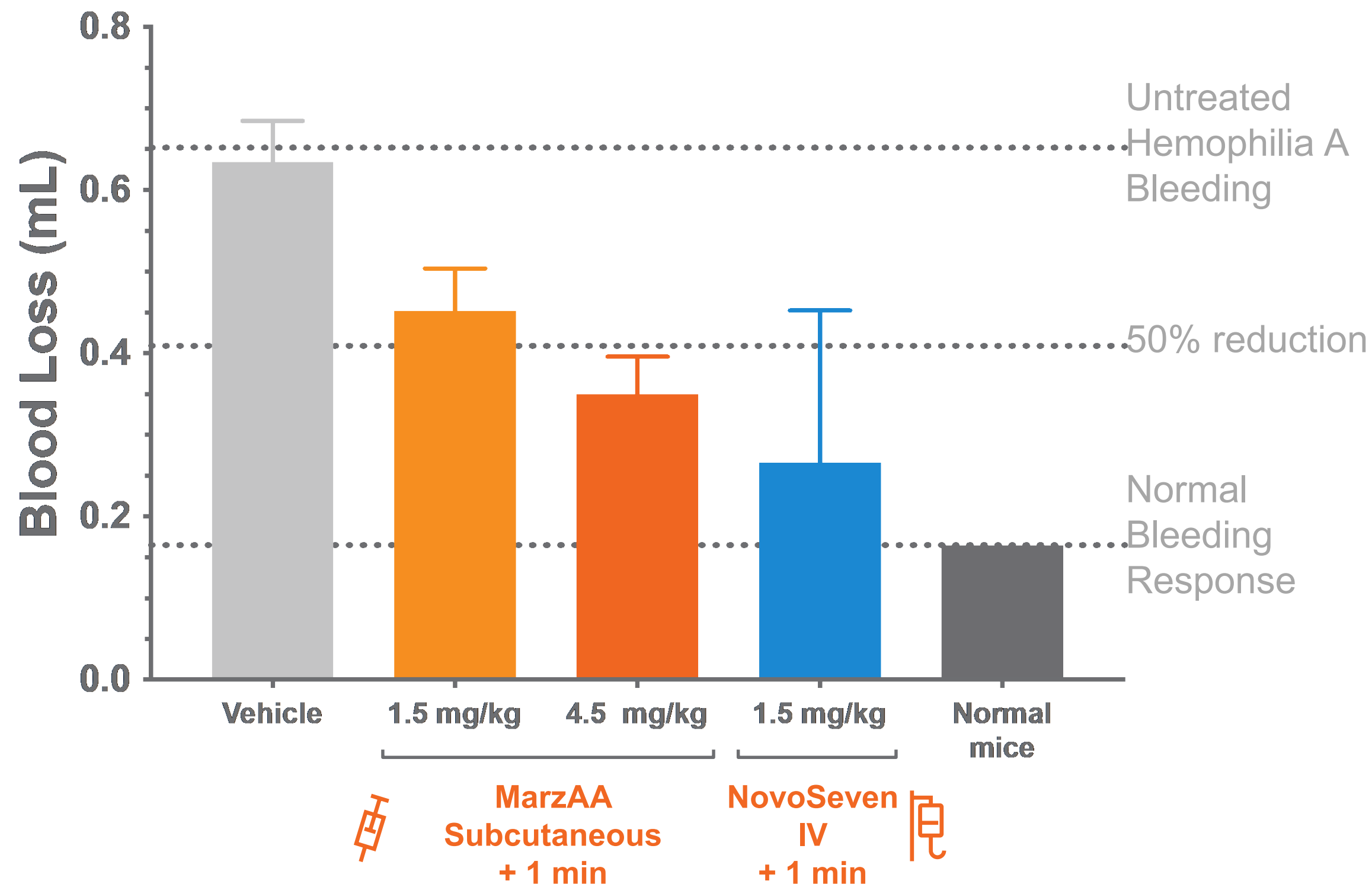
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- ✓ Fast & easy to administer
- ✓ Stops bleeding in a validated preclinical model
- ✓ Can be combined with Hemlibra *in vitro*



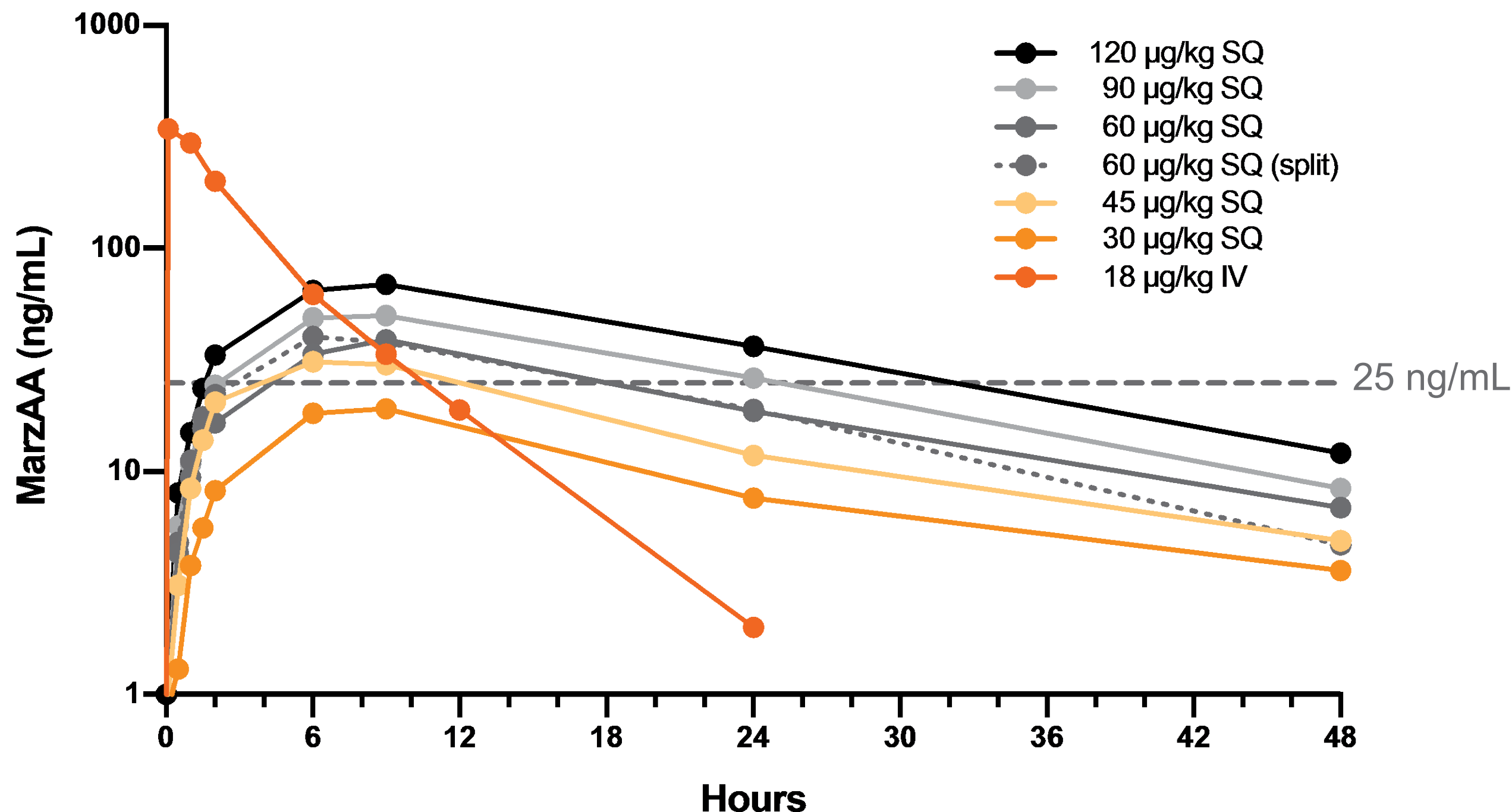
# SQ MarzAA reduces bleeding when dosed **after** injury

## Acute mouse injury model with dosing *after* injury



- + Preclinical tail-cut model in hemophilic rodents is the standard for efficacy
- + SQ MarzAA one minute **after** tail-cut significantly **reduces blood loss**
- + Reduction in blood loss is dose dependent
- + Reduction in blood loss **with SQ MarzAA is similar to IV NovoSeven**

# MAA-102 PK dose levels support SQ treatment of a bleed



- + Dose-proportional increases in Cmax and AUC
- + Rapid increase in levels to target range of 25 ng/mL
- + Target levels can be maintained for 24 hours
- + No ADAs
- + Multiple dose cohorts enrolling
  - 60 µg/kg Q3-hourly twice or thrice

# MarzAA is only bypass agent for **both** SQ prophylaxis and SQ treatment of bleeds

**Attractive commercial profile targeting an existing \$2.2B bypass agent market**

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IV NovoSeven (\$1.2B 2019 sales) validates rFVIIa in multiple rare bleeding disorders

- + Hemophilia A or B with inhibitors
- + Severe Factor VII Deficiency
- + Glanzmann Thrombasthenia
- + Acquired Hemophilia A

**SQ MarzAA has a superior profile to IV NovoSeven – over 100 clinicians surveyed**

- + All physicians surveyed indicated a preference for **SQ MarzAA** over IV N7 in one or more settings
- + **SQ MarzAA** can create & expand multiple prophylaxis markets

# Marzeptacog alfa (activated)

## Phase 3 study to initiate in 2020

Large commercial opportunity across multiple rare bleeding disorders

Demonstrated P2 Clinical efficacy & tolerability for prophylaxis indications

Demonstrated preclinical PoC for SQ treatment of a bleed

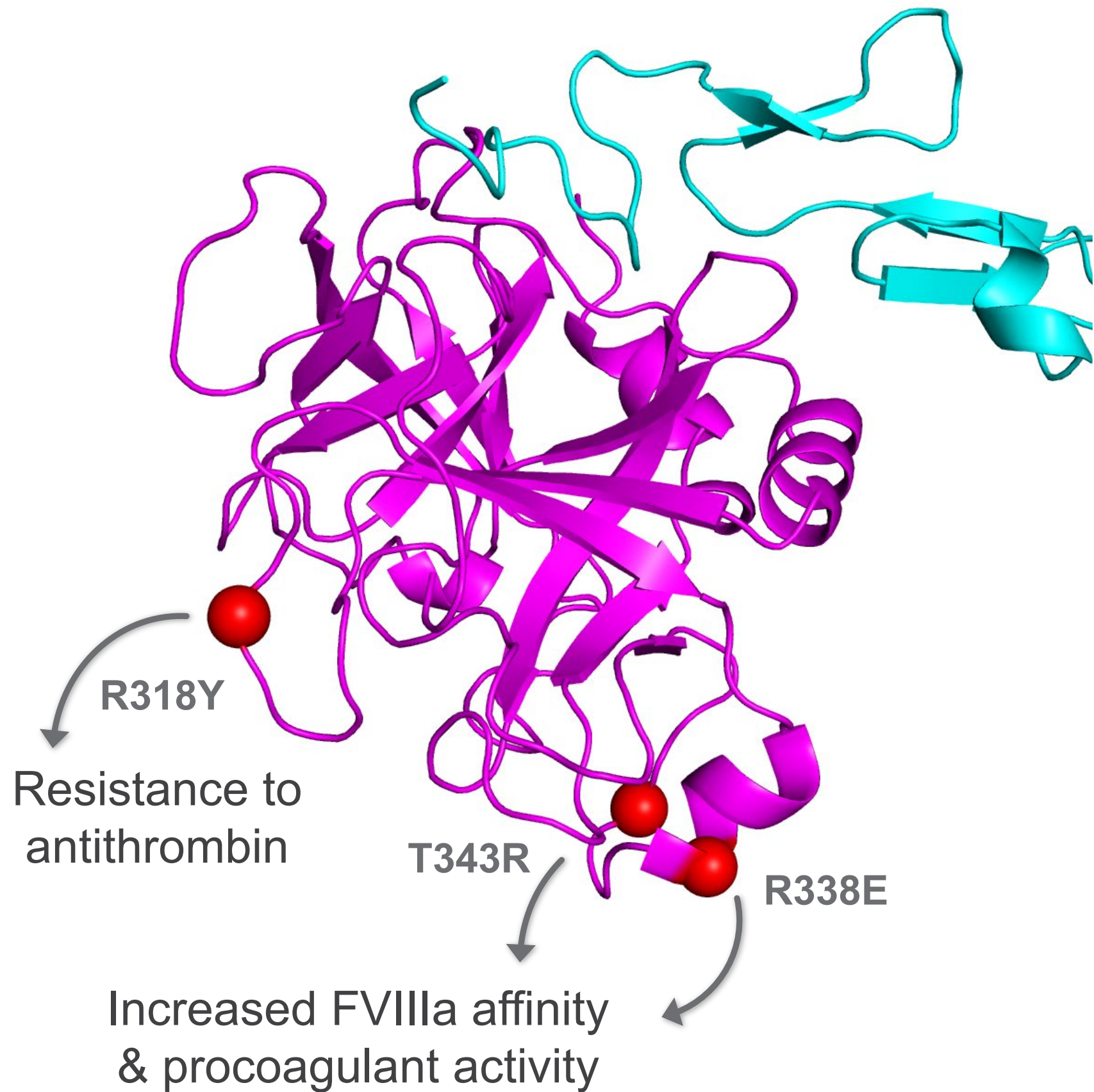
MarzAA can be safely combined with Hemlibra *in vitro*

SQ dose escalation PK/PD study supports treatment of a bleed – final data in Q2 2020

P3 prophylaxis guidance from EMA & MHRA received



# Dalcinonacog alfa: a novel SQ FIX product



## Three substitutions within the FIX protein

- + Increased catalytic activity
- + Higher affinity for FVIIIa
- + Resistance to antithrombin inhibition
- + 22-fold increased potency over BeneFIX

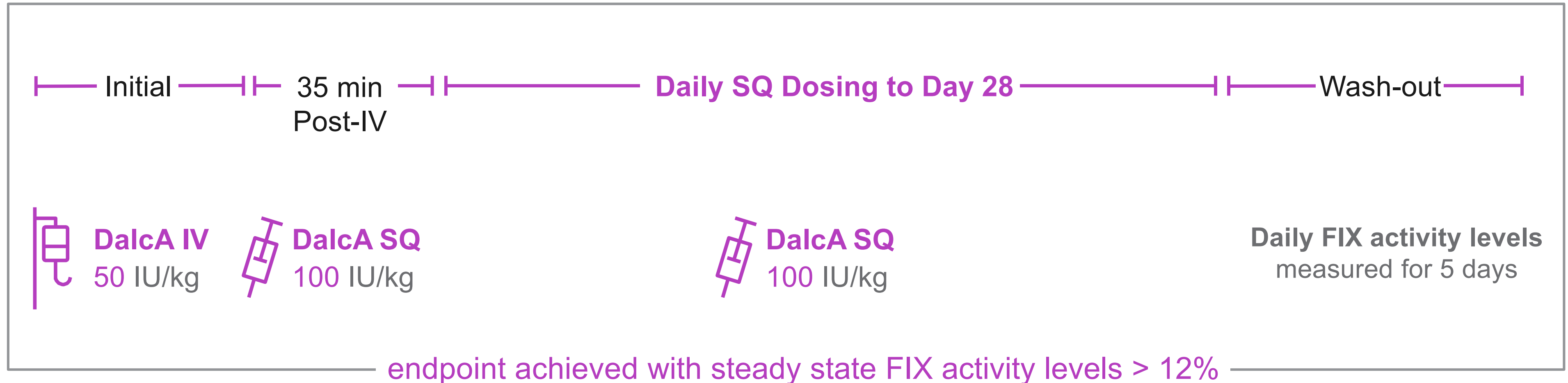
## Differentiated from marketed IV FIXs

- + Simple, small volume SQ administration
- + Enhanced pharmacokinetics with prolonged half-life
- + Excellent extravascular distribution
- + Potential to maintain continuous protective levels

## Orphan Drug Designation in US & EU

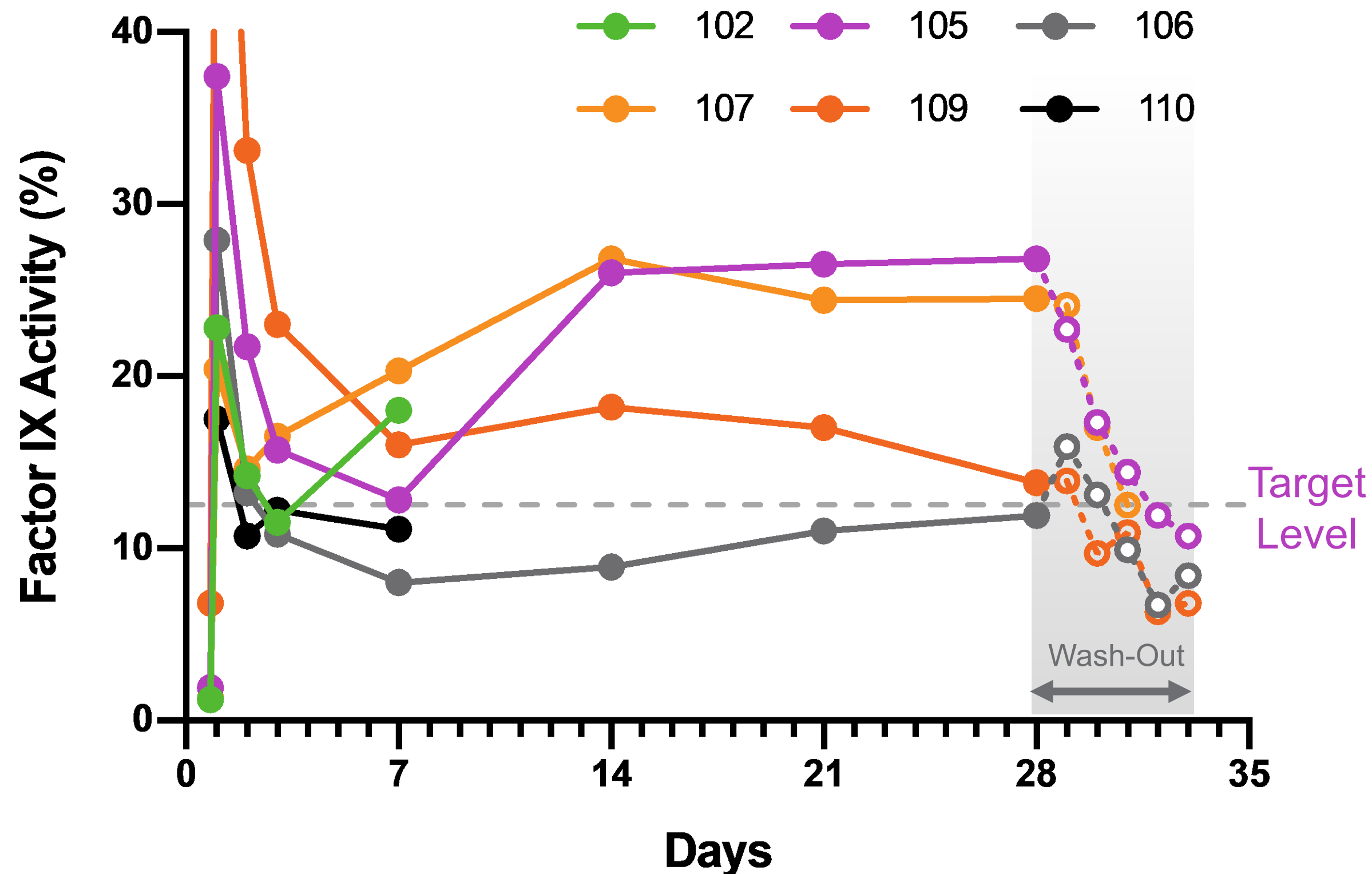
# Dalcinonacog alfa phase 2b SQ clinical trial design

## Enrollment & dosing complete



- + Primary endpoint: **Steady state FIX activity** level above 12% with daily dosing
- + Secondary endpoints: **safety including weekly ADA testing**, pharmacokinetics, pharmacodynamics, bleeding events
- + 10 severe HB patients screened; 6 dosed
- + Rare propeptide mutation excluded
- + HLA profile associated with nAb risk was excluded

# Target levels achieved with 100 IU/kg dosing for 28 Days



## Target FIX >12% Achieved

- + Dosed 6 severe HB subjects
  - 110 continues dosing\*
  - 102 withdrew on Day 7
- + **Steady state FIX levels up to 27%** achieved after 14 days
- + **No breakthrough bleeds**
- + **No ADAs**
- + Consistent PK profiles
- + Terminal half-life is 70-112 h

## Has the potential to be an effective SQ prophylaxis treatment for individuals with Hemophilia B

Trial enrollment & dosing complete

Excellent & consistent therapeutic FIX activity levels attained

No bleeding events during treatment demonstrates effective prophylaxis

No SAEs, systemic hypersensitivity, ADAs or nAb to DalcA or wild-type FIX

Mild to moderate ISR's primarily with initial injections – resolved

Long half-life with SQ administration – potential to lower dose &/or frequency



# FIX gene therapy: CB 2679d-GT for hemophilia B

## CB 2679d-GT in combination with a novel chimeric AAV capsid provides significant improvements

- + Stable high activity levels in a mouse hemophilia B model – **no nAbs**
- + Vector dose reduced 10-fold compared to current constructs
- + Potential for an improved efficacy & safety profile
- + AAV license and sponsored research agreement with Stanford University School of Medicine

## Superior preclinical efficacy of CB 2679d-GT vs Padua

- + 4-5-fold reduction in bleeding time
- + Activity levels elevated throughout the study - **no nAbs**

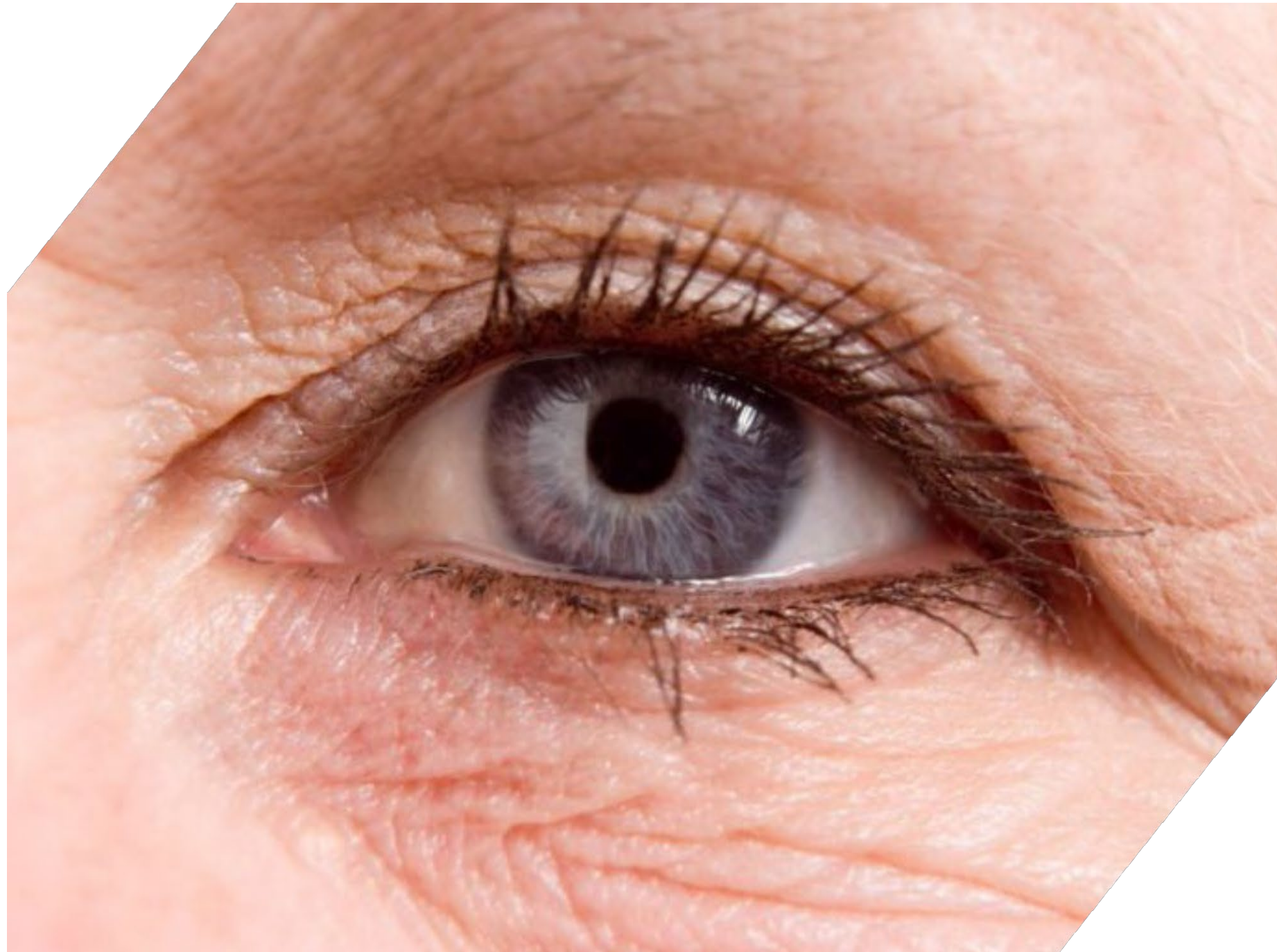
## Wholly-owned & issued patents covering gene therapy

FIX Transgene	AAV Capsid	Study Dose (vg/kg)	FIX Activity (U/mL)
CB 2679d-GT	Novel Chimeric	8.0x10 <sup>10</sup>	20
Padua	TAK-748*	7.4x10 <sup>11</sup>	20
Padua	TAK-748*	7.4x10 <sup>10</sup>	1

\*Weiller *et al.* (2019) *Blood* Vol. 134, Supplement S1 P4633

# CB 2782-PEG anti-complement factor 3 (C3) protease

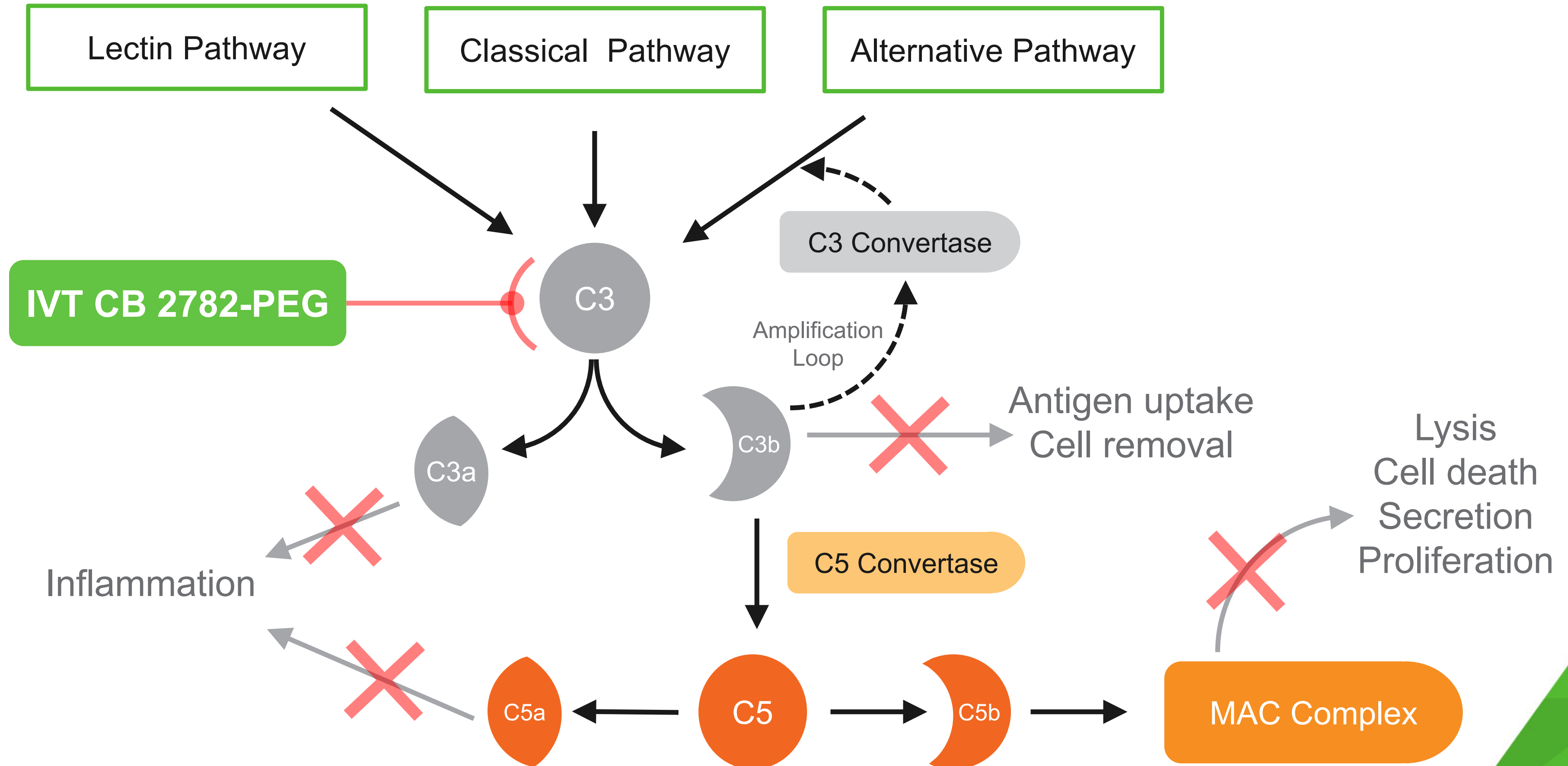
## Geographic Atrophy in Dry AMD



- + Geographic atrophy is an advanced stage of dry age-related macular degeneration that results in the irreversible loss of retina and leads to blindness
- + Dry AMD affects a million people in the United States and over five million people worldwide
- + Global market is estimated at >\$5B with no approved drugs
- + C3 is the only clinically validated target for the treatment of Dry AMD

Sources: National Eye Institute. Facts About Age-Related Macular Degeneration, Tufail 2015, The Eye Diseases Prevalence Research Group 2004, GlobalData

# Targeting C3 blocks the downstream complement cascade





# CB 2782-PEG long acting anti-C3 protease

## Best-in-class anti-C3 profile for dry AMD

- + Generated from Catalyst's proprietary protease engineering platform
- + Potent, selective and long acting anti-C3 protease that degrades C3 into inactive fragments
- + Preclinical NHP PK & PD data\* predict best-in-class human intravitreal dosing three or four times a year









## Biogen Collaboration

- + Announced December 19, 2019
- + \$15M upfront, up to \$340M in milestones and tiered royalties up to low double digits
- + Catalyst to perform fully funded pre-clinical and manufacturing activities
- + Biogen responsible for IND-enabling activities, worldwide clinical development & commercialization





# Milestones

	2019	Q1	Q2	H2
<b>MarzAA</b> (FVIIa)	P2 efficacy 	EoP2	ToB enabling PK/PD 	Registration Trial
<b>DalcA</b> (FIX)	Positive P2b Interim data 	P2b Update 	Final P2b data	
<b>CB 2679d-GT</b> (FIX Gene Therapy)	Preclinical efficacy 	NextGen Vector 	NHP Efficacy	
<b>CB 2782-PEG</b> (dAMD)	Partnership  			

# Financial information

## Selected data

### Financial results

**YE 2019**

Cash & Cash Equivalents .....	\$76.9 M <sup>1</sup>
Operating Expense .....	\$57.3 M
Net Loss (YTD).....	(\$55.2M)
Net Loss per share (YTD).....	(\$4.60)

### Share data

Common Stock Outstanding.....	12,040,835 <sup>2</sup>
Officer & Director ownership .....	7.0%
Fully Diluted Shares.....	14,859,051 <sup>3</sup>

- <sup>1</sup> Excludes \$15M Biogen  
upfront payment  
December 2019 and  
\$34.5M follow-on (gross)
- <sup>2</sup> Excludes ~5.3M shares  
issued in a February 2020  
public offering
- <sup>3</sup> Includes ~1.6M options  
available for issuance

# Team

## President & CEO

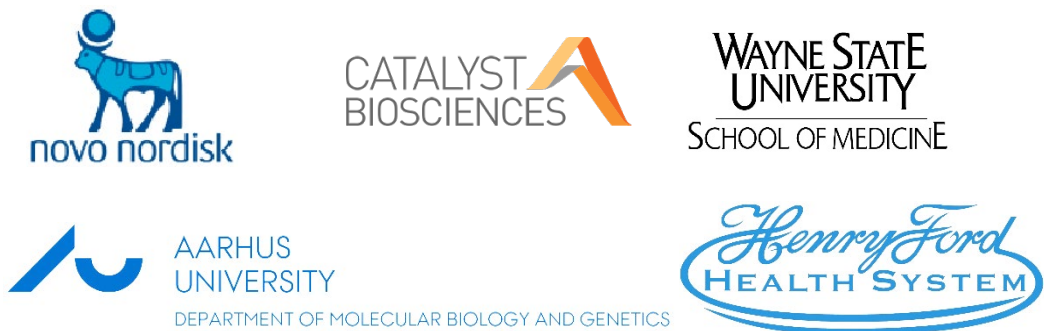
Nassim Usman, Ph.D.



28 years  
in biotech

## SVP, Translational Research

Grant Blouse, Ph.D., M.Sc.



13 years  
in biotech

## Chief Medical Officer

Howard Levy, M.B.B.Ch., Ph.D., M.M.M.



18 years  
in hematology

## SVP, Technical Operations

Andrew Hetherington, M.B.A.



20 years  
in biotech

## SVP, Business Development

Jeffrey Landau, M.B.A.



16 years  
in biotech

# Summary

## Disruptive approach to billion-dollar markets – protease engineering platform

- ✓ **FVIIa: SQ MarzAA ~\$2.2B market**
  - + P2 efficacy & safety demonstrated
  - + P1/2 PK/PD supports ToB
  - + FDA EoP2 in early 2020, P3 expected in 2020
- ✓ **FIX: SQ DalcA >\$1.8B market**
  - + Phase 2b efficacy & safety demonstrated
  - + Final Phase 2b data in 2Q 2020
- ✓ **FIX Gene Therapy: CB 2679d-GT**
  - + Proprietary preclinical gene therapy asset with superior activity vs current clinical constructs
- ✓ **Anti-C3 dAMD: IVT CB 2782-PEG >\$5B market**
  - + Biogen collaboration
  - + \$15M upfront, up to \$340M in milestones and tiered royalties up to low double digits
- ✓ **SQ systemic complement inhibitor program**
  - + Large orphan disease opportunity
  - + Builds complement franchise
- ✓ **Strong financial position**



# THANK YOU

Nasdaq: CBIO

[catalystbiosciences.com](http://catalystbiosciences.com)

