# CATALYST BIOSCIENCES

Piper Sandler Virtual Healthcare Conference 23 November 2020

CatalystBiosciences.com



## 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 forwardlooking statements. Forward-looking statements include statements about the potential benefits of products based on Catalyst's engineered protease platform; potential markets for and advantages of MarzAA and DalcA; plans in Q4 2020 to enroll a pivotal Phase 3 registration study of MarzAA, initiate a Phase 1/2 trial in FVII Deficiency, Glanzmann Thrombasthenia, and patients treated with Hemlibra; the potential for MarzAA and DalcA to effectively and therapeutically treat hemophilia subcutaneously; potential markets for our anticomplement and gene therapy programs; potential payments from Biogen; plans to declare a development candidates in our systemic complement program in Q4 2020; the superiority of CB 2679d-GT over other gene therapy candidates; and the Company's collaboration with Biogen for the development and commercialization of pegylated CB 2782 for the potential treatment of geographic atrophy-associated dry age-related macular degeneration (AMD). Actual results or events could differ materially from the plans, intentions, expectations and projections disclosed in the forwardlooking statements.

Various important factors could cause actual results or events to differ materially, including, but not limited to, the risk that trials and studies may be delayed as a result of the novel coronavirus (COVID-19) outbreak and other factors, that trials may not have satisfactory outcomes, that additional human trials will not replicate the results from earlier trials, that potential adverse effects may arise from the testing or use of DalcA or MarzAA, including the generation of neutralizing antibodies, which has been observed in patients treated with DalcA, the risk that costs required to develop or manufacture the Company's products will be higher than anticipated, including as a result of delays in development and manufacturing resulting from COVID-19 and other factors, the risk that Biogen will terminate Catalyst's agreement, competition and other risks described in the "Risk Factors" section of the Company's quarterly report filed with the Securities and Exchange Commission on November 5, 2020, and in other filings with the Securities and Exchange Commission. The Company does not assume any obligation to update any forwardlooking statements, except as required by law.

## Catalyst Biosciences – Protease medicines



## Protease engineering platform

## Late-stage asset

SQ Marzeptacog alfa (activated)
MarzAA (FVIIa)

Phase 3 in 2020

## Hemophilia

SQ MarzAA (FVIIa)

SQ Dalcinonacog alfa – DalcA (FIX)

Factor IX Gene Therapy

Factor Xa

## Complement

IVT Anti-C3 Dry AMD CB 2782-PEG

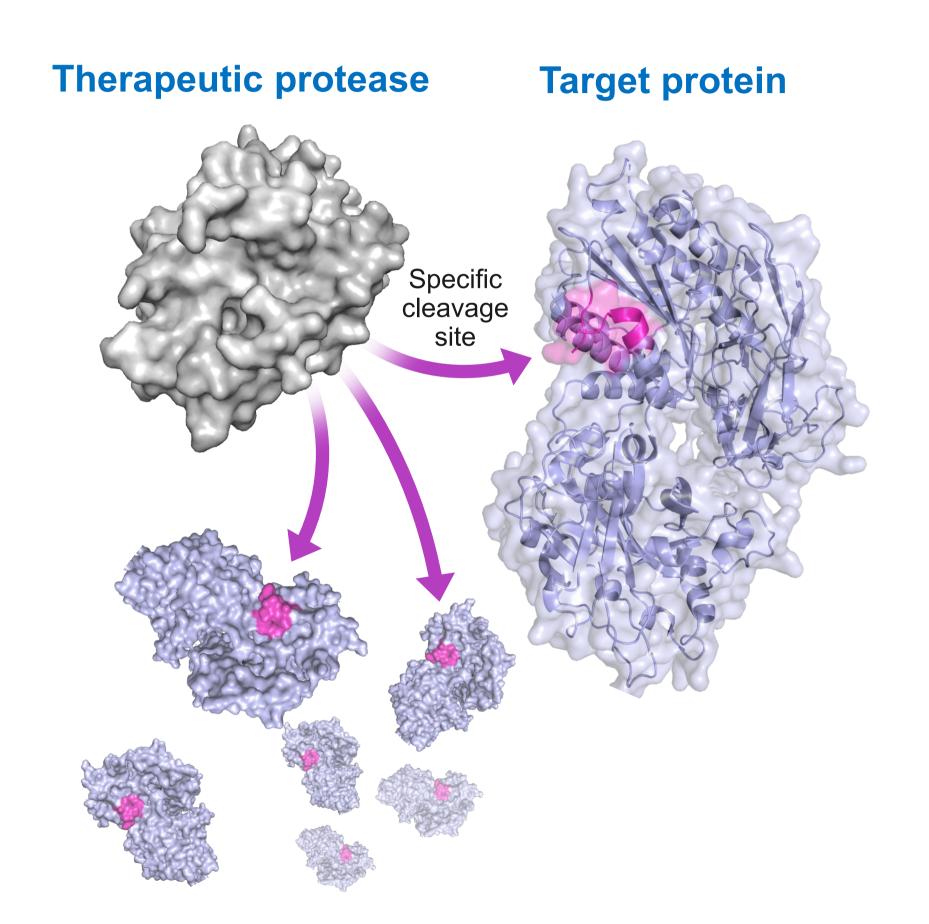


SQ Systemic Complement Inhibitors

## Harnessing the catalytic power of proteases



### One protease molecule regulates 1000s of target molecules



### An adaptable protease platform

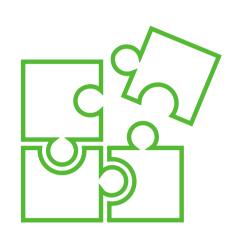
- **Obligation** Demonstrated efficacy of clinical stage assets
- Functionally enhanced natural proteases (FVIIa, FIX)
- Engineered novel protein degraders (Anti-C3)
- Ideal for high concentration drug targets or controlling amplification cascades
- Potential to address novel targets
- Increased potency and extended half-life variants

## **Investment highlights**





Novel subcutaneous factors with orphan drug designation; MarzAA & DalcA – P2 efficacy in prophylaxis studies complete



Anti-C3 Dry AMD with Biogen SQ systemic complement regulator research program



Multibillion-dollar market opportunities



Experienced team



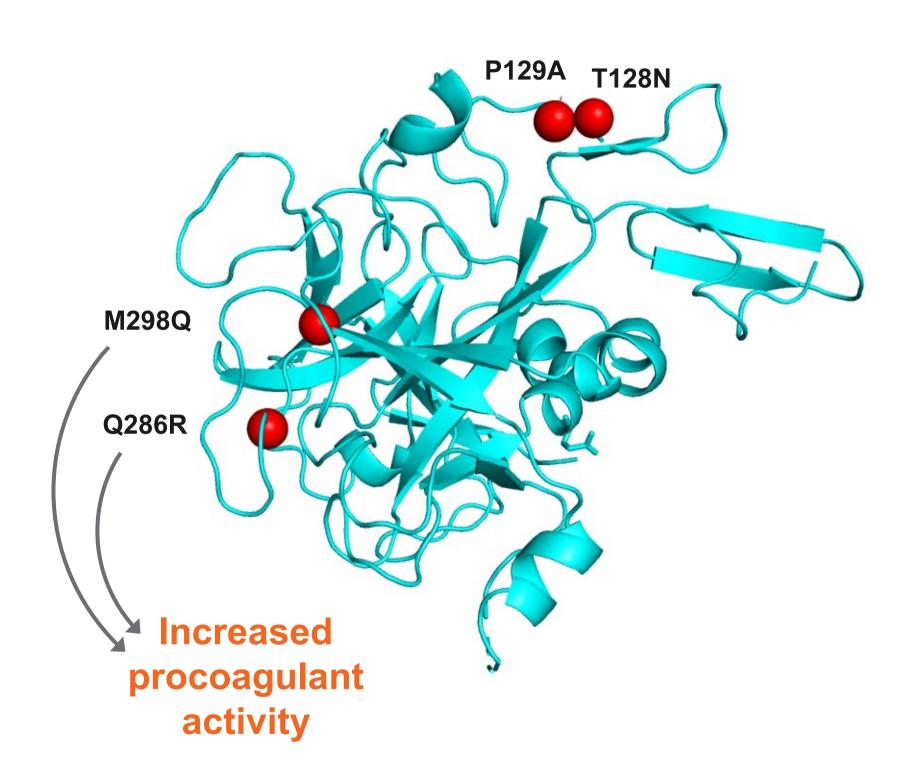
Strong balance sheet, \$104M cash – Q3



178 worldwide patents
CBIO retains full ownership
of all compounds

## Marzeptacog alfa (activated): MarzAA rFVIIa

#### Addresses a clear unmet need in hemophilia & other bleeding disorders



#### 9-fold higher activity vs NovoSeven RT

- + Potency allows for SQ dosing that prolongs half-life
- + Simple, small volume SQ administration

#### Preclinical efficacy of SQ on-demand treatment

+ HA mouse after tail cut; HA dog; HA rat

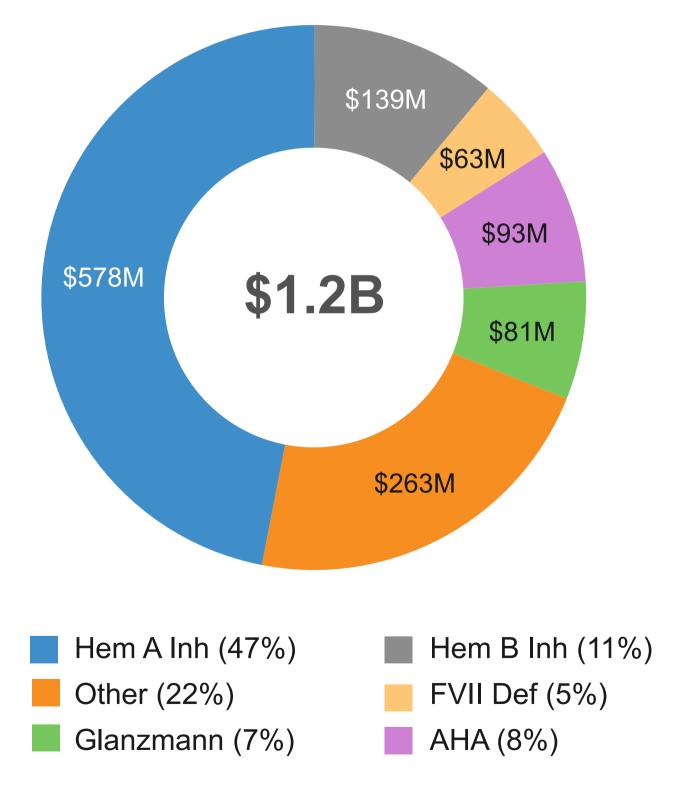
## P2/3 prophylaxis efficacy & safety in HA or HB with inhibitors

+ 46 patients treated including: single dose IV, up to 3 SQ doses/day, & daily SQ up to 97 days

## SQ MarzAA is a large commercial opportunity



## Global NovoSeven sales breakdown by indication (2019)



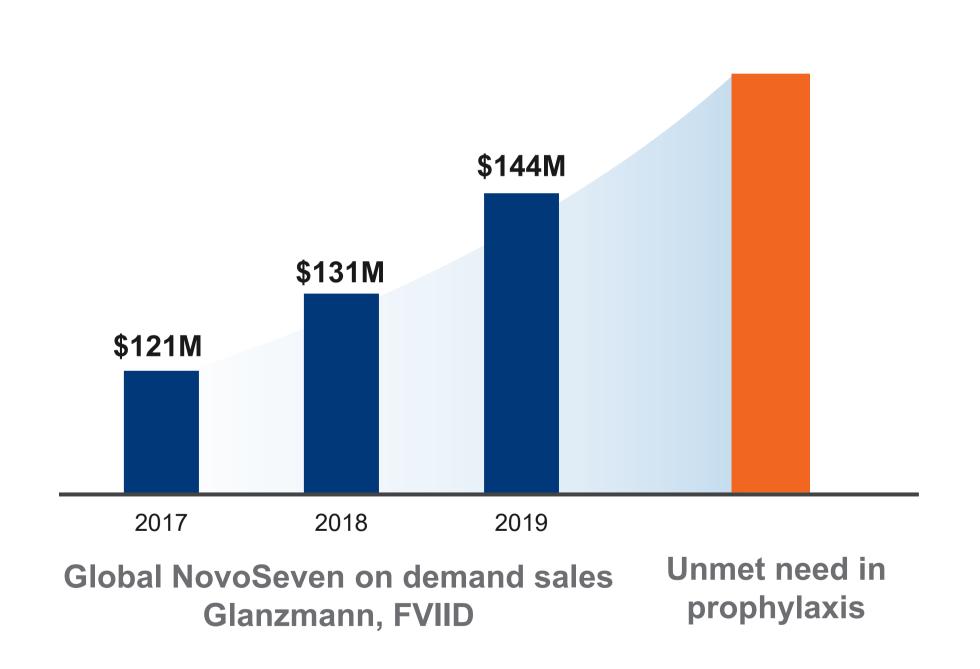
#### SQ MarzAA has a superior profile

- Faster & easier to administer *vs* N7 dosed every 2 hours IV until hemostasis
- MarzAA SQ half-life ~8x longer than N7
- 9-fold higher activity vs N7
- Open Potential to reduce rebleeding
- Stops bleeding in multiple preclinical models
- Can be combined with Hemlibra in vitro without increased thrombogenicity
- Ideal for pediatrics and patients with venous access issues
- Prophylaxis efficacy demonstrated in P2

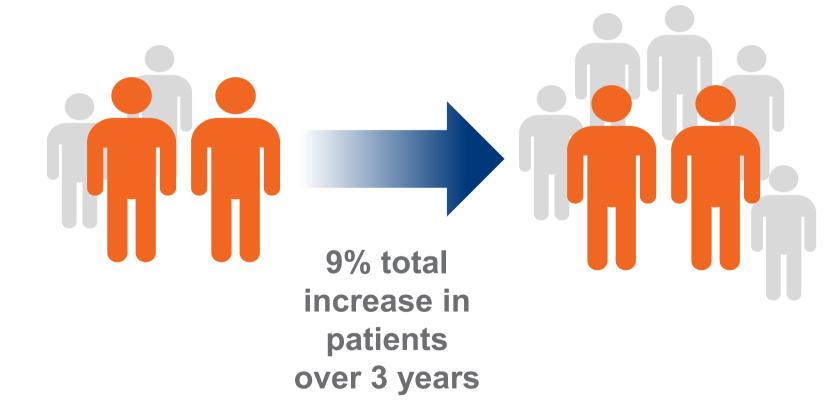
Source: Adivo Associates market research; Catalyst Biosciences market research. Data on file

## MarzAA could be the first prophylaxis for Glanzmann & FVIID





Growing number of Glanzmann and FVIID patients treated with NovoSeven

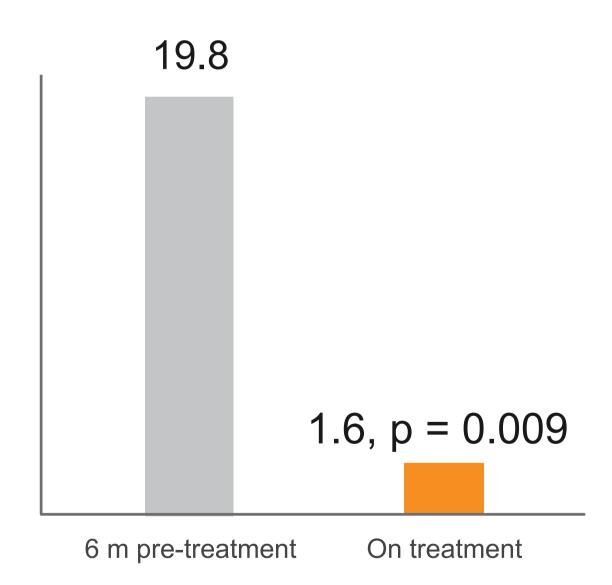


Source: Catalyst Biosciences, Adivo Associates Market Research, Data on file. \*Note: Treated patients may be counted multiple times as patients may have multiple bleeding events per year needing factor treatment

## MarzAA is efficacious with daily prophylaxis

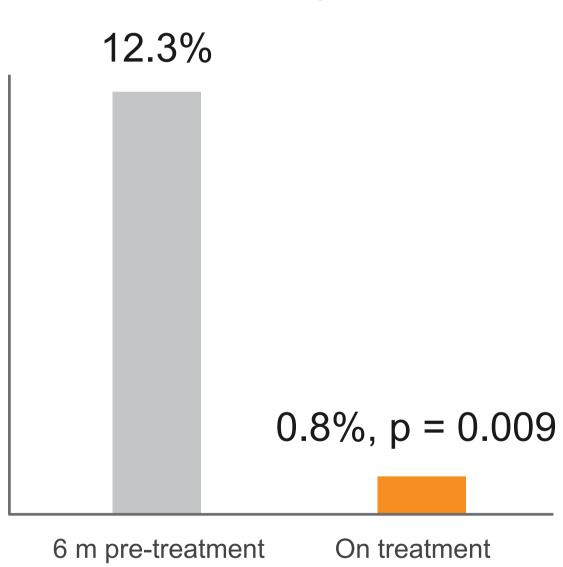
### Phase 2: Daily SQ dosing for 44 – 97 days

#### Annualized bleed rate n = 9



## Proportion of days with bleeding

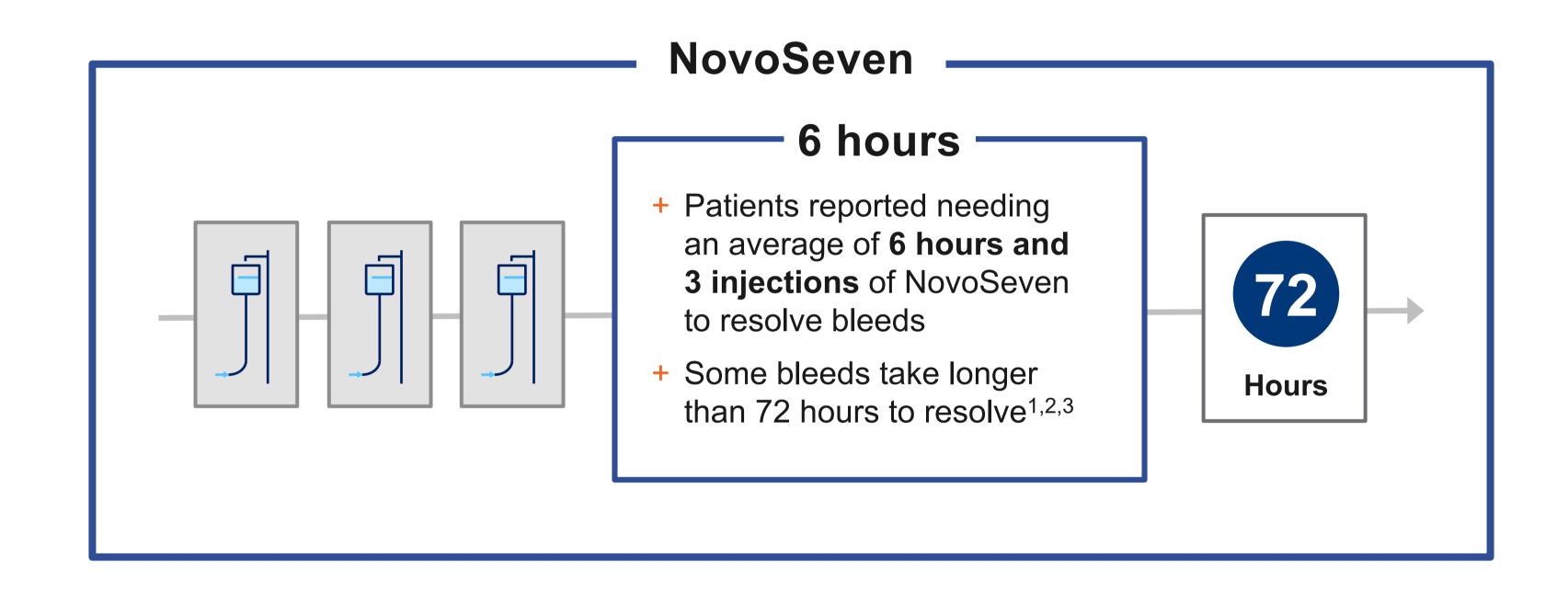
$$n = 9$$



- + Greater than 90% reduction in all bleeding Median ABR = 0
- + 2 subjects had dose escalation from 30 to 60 μg/kg
- + Safe & well tolerated, ~1% ISR (6/517 doses) & no ADA

## Current bypass agents require multiple IVs over the course of hours

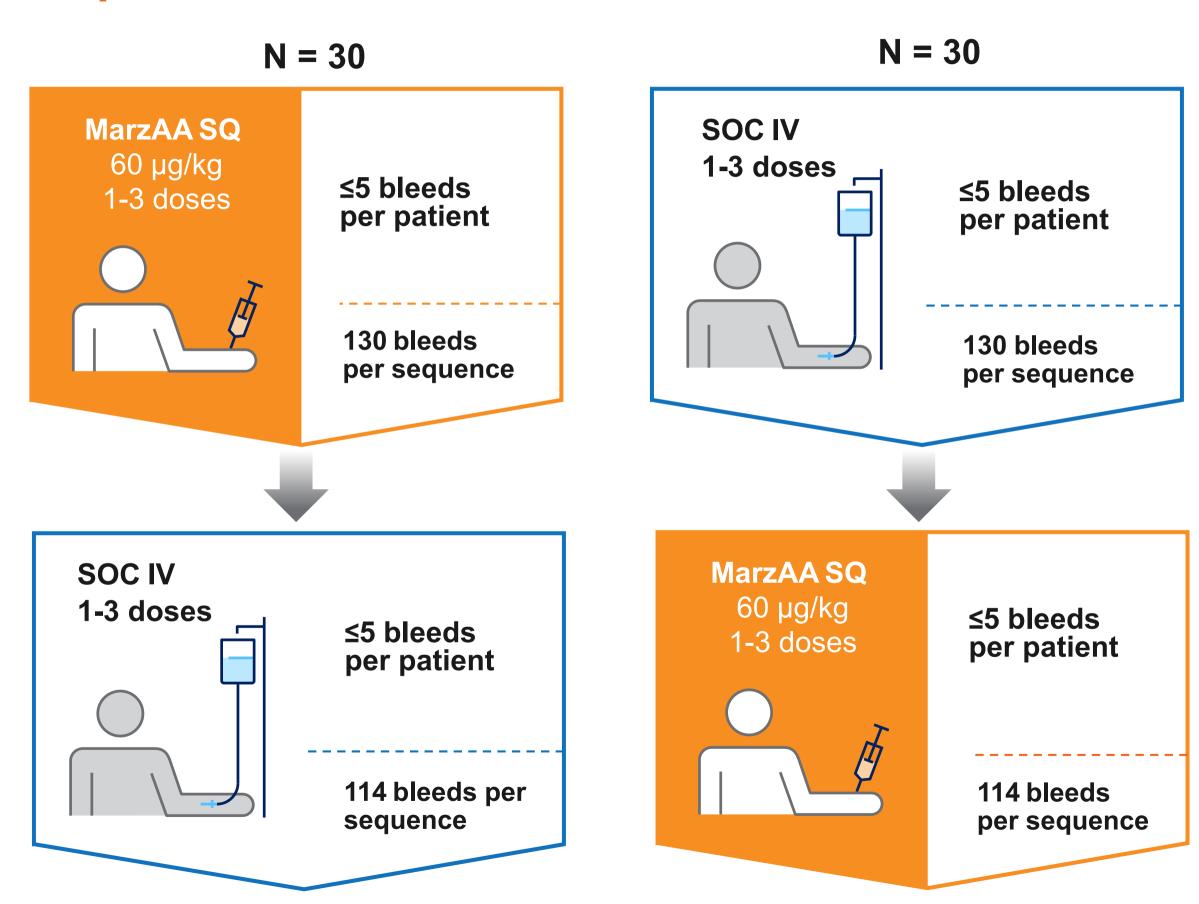




## Crimson 1 Phase 3 study: Treatment of episodic bleeding



### Hemophilia A or B with inhibitors, ABR ≥ 8



#### Primary endpoint

Non-inferior hemostatic efficacy: standard 4-point scale at 24 h

#### Secondary endpoints

Time to bleed resolution; number of doses; rescue meds

#### Safety

Adverse events, anti-drug antibodies (ADA); thrombosis

#### 

- + SOC estimate 85%

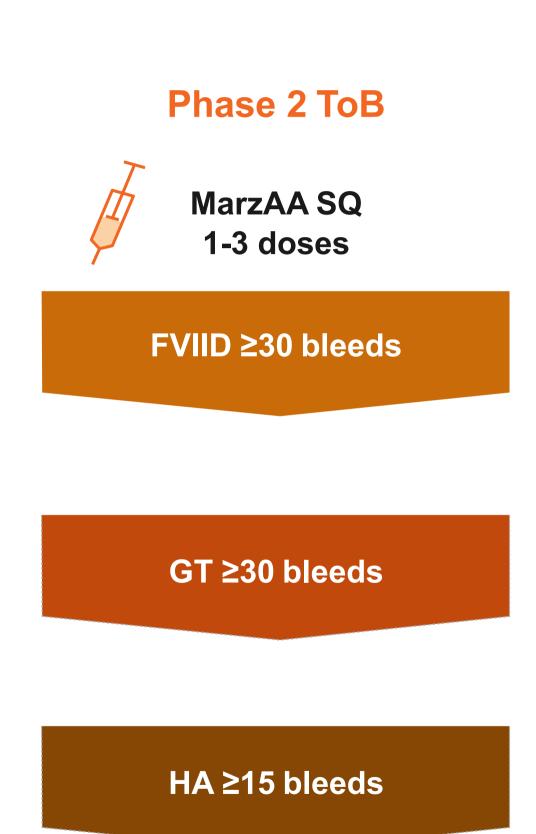
  Excellent/good treatment of bleeds
- + Non-inferiority margin of 12%
- + 2.5% significance, one-sided
- + 90% power

## MAA-202 Phase 1/2 study design



### FVII deficiency, Glanzmann thrombasthenia and HA on Hemlibra: N = 8 each

## Phase 1 PK MarzAA IV each cohort Single dose MarzAA SQ Single dose escalation Multiple dose Q3H

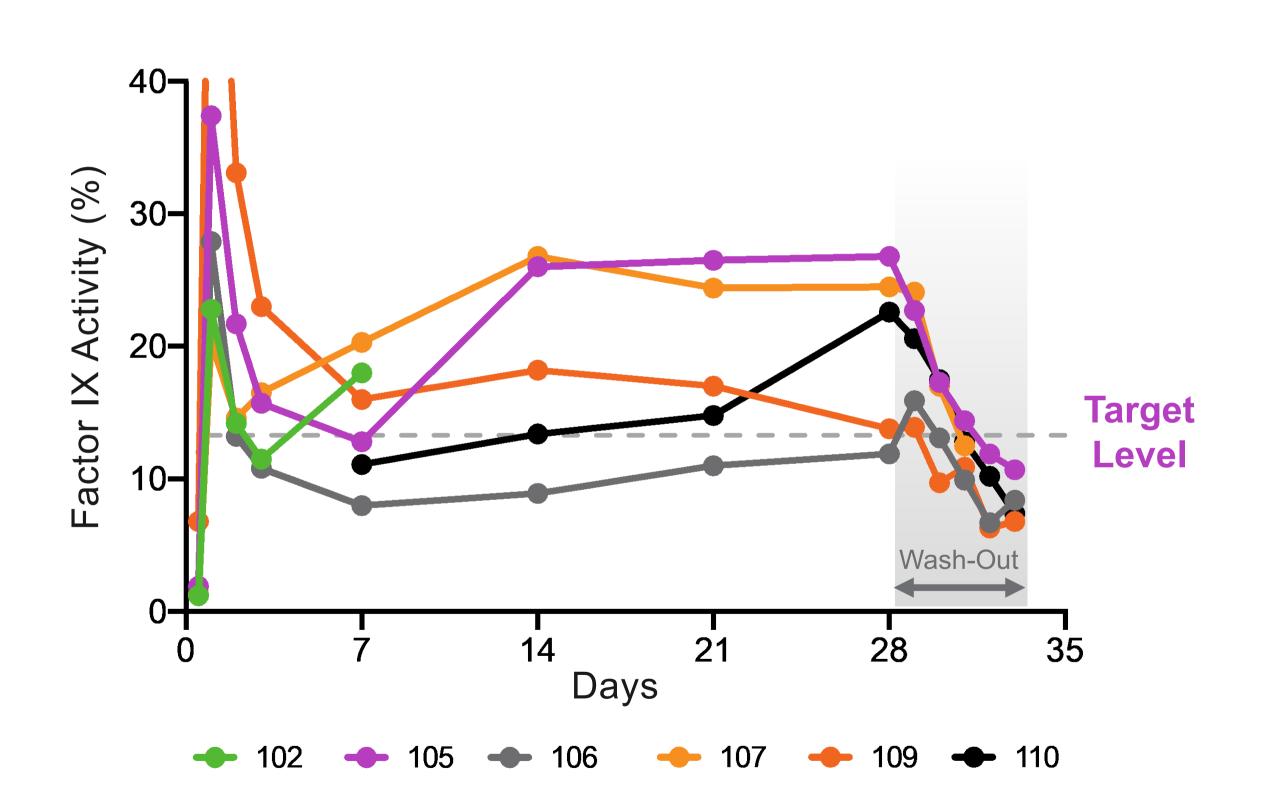


## Phase 1 **Primary endpoint: Pharmacokinetics Secondary endpoint:** Pharmacodynamics Phase 2 ToB **Primary endpoint:** Hemostatic efficacy at 24 hours **Secondary endpoints:** Effective hemostasis at successive timepoints; doses needed; rescue meds Safety: Adverse events and ADA

## DalcA P2b demonstrated efficacy & safety



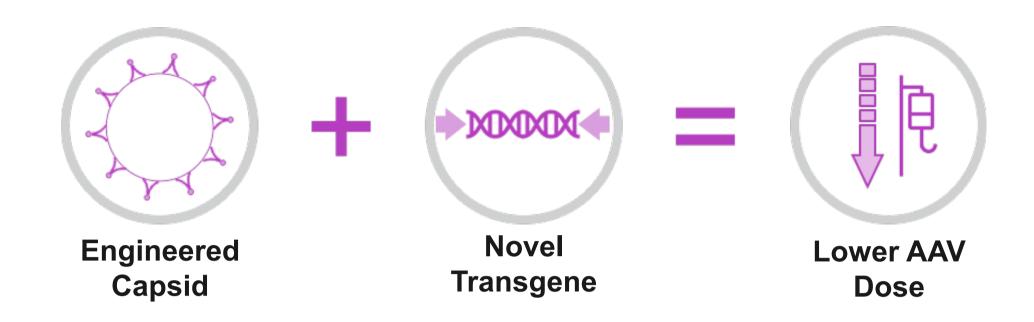
Target levels >12% achieved with daily SQ 100 IU/kg dosing for 28 days



- + Dosed 6 severe HB subjects
  - Subject 102 withdrew on Day 7
- + Steady state FIX levels up to27% achieved after 14 days
- + No breakthrough bleeds
- + No neutralizing ADA
- + Mild to moderate ISR transient & self-limiting
- + Terminal half-life is 2.5 5.1 days

## Catalyst's CB 2679d gene therapy for hemophilia B





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

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



License & sponsored research agreement

## **⊘** CB 2679d-GT has a superior profile *vs* Padua in preclinical studies

- + Stable high activity levels with 1/10th vector dose in mouse model
- + 4 to 5-fold reduction in bleeding time when compared to the Padua
- + Potential for improved efficacy & safety at 1-2 log reduced dose

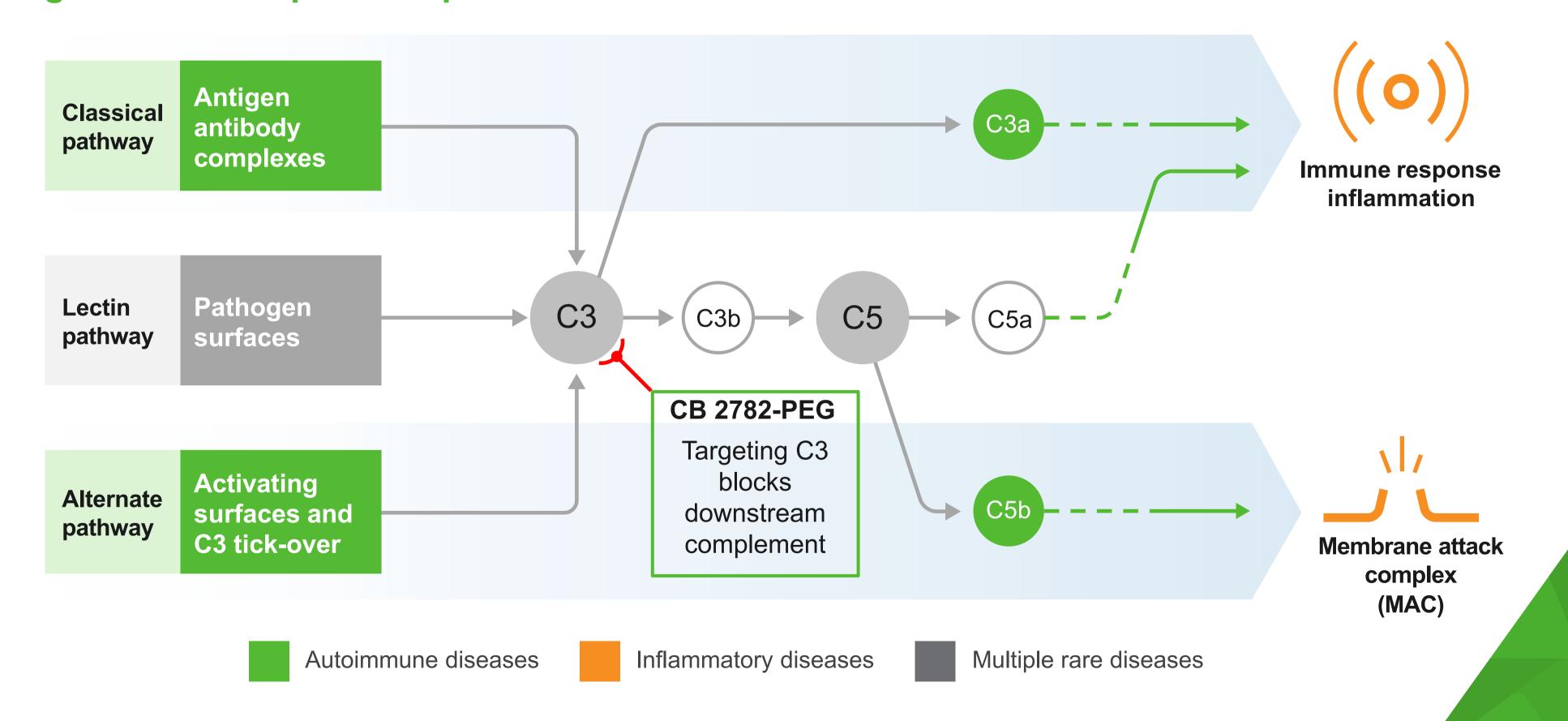
#### 

- + Presented at World Federation of Hemophilia Virtual Summit 2020
- + Additional vector optimization & dose ranging studies ongoing
- **⊘** Wholly-owned & issued patents covering gene therapy

## Targeting complement – a pathway regulated by proteases



Dysregulated complement activity is associated with a broad range of disorders and a logical fit for our protease platform



## 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

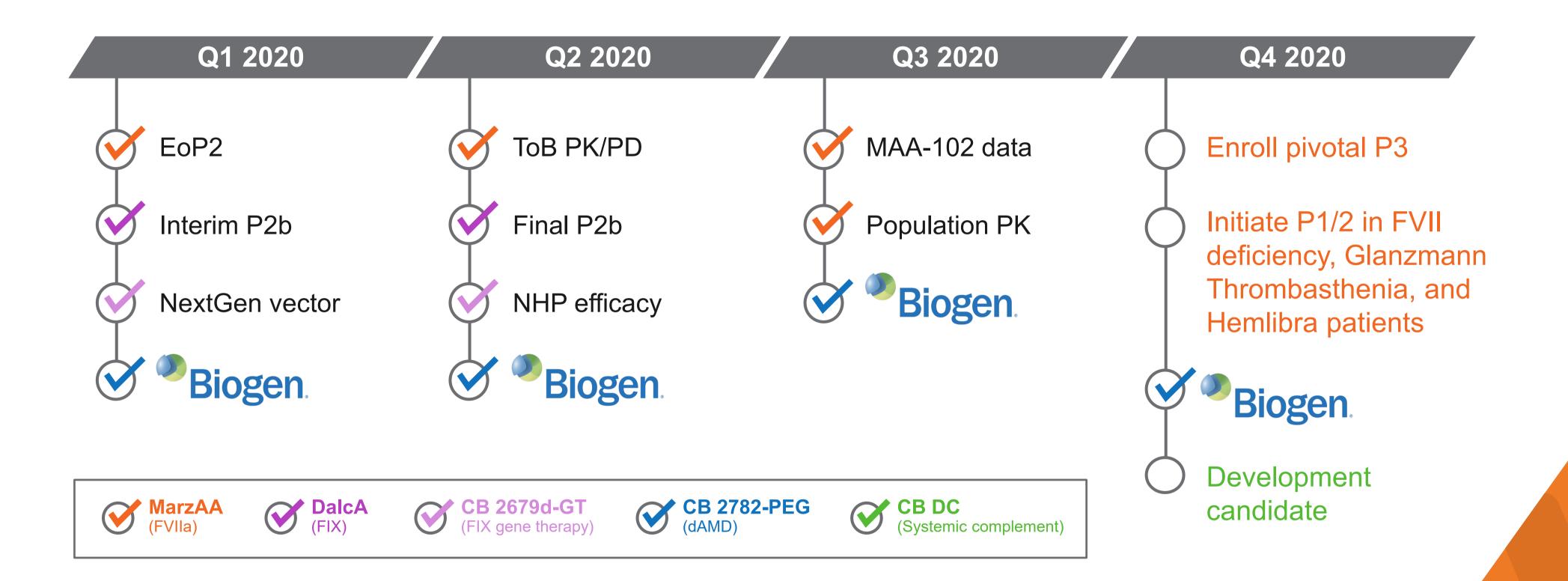
#### **Biogen collaboration**



- + Announced December 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 – 2020





#### **Team**



#### Nassim Usman, Ph.D.

President & CEO









28 years in biotech

#### Grant Blouse, Ph.D.

**SVP Translational Research** 











13 years in biotech

#### **Clinton Musil**, M.B.A

**Chief Financial Officer** 







16 years in biotech & investing/banking

#### Jeffrey Landau, M.B.A.

**SVP Business Development** 









18 years in biotech

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

**Chief Medical Officer** 











20 years in hematology

#### Anju Chatterji, Ph.D.

**SVP Biologics Development & Manufacturing** 







19 years in biotech

## Summary



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

- FVIIa: SQ MarzAA ~\$2.2B market
  - + P1 PK/PD & preclinical data supports ToB
  - + P2 efficacy & safety demonstrated
  - + P3 patient enrollment in Q4 2020
- FIX: SQ DalcA >\$1.8B market
  - + Phase 2b efficacy & safety demonstrated
  - + Potential for less frequent dosing
- FIX Gene Therapy: CB 2679d-GT
  - + Proprietary preclinical gene therapy asset with superior activity and lower dose *vs* current clinical constructs

- Anti-C3 dAMD: IVT CB 2782-PEG >\$5B market
  - + Biogen collaboration
  - + \$15M upfront, up to \$340M in milestones, up to low double digits tiered royalties
- SQ systemic complement inhibitor program
  - + Large \$B+ rare-disease opportunity
  - + Multiple indications & applications
  - + 1<sup>st</sup> development candidate in Q4 2020
- Well capitalized
  - + Cash runway into 2022

## THANK YOU

Nasdaq: CBIO

CatalystBiosciences.com

