



KalVista
Pharmaceuticals

Corporate Overview

March 2023

Forward-Looking Statements

This presentation and the accompanying oral presentation contain “forward-looking” statements within the meaning of the safe harbor provisions of the U.S. Private Securities Litigation Reform Act of 1995. Forward-looking statements can be identified by words such as: “anticipate,” “intend,” “plan,” “goal,” “seek,” “believe,” “project,” “estimate,” “expect,” “strategy,” “future,” “likely,” “may,” “should,” “will” and similar references to future periods. Examples of forward-looking statements include, among others, statements we make regarding our future financial performance, business plans and objectives, timing and success of our clinical trials, our ability to obtain regulatory approval or the timing of regulatory filings, the potential therapeutic benefits and economic value of our lead product candidates, financing plans, competitive position, industry environment and potential market opportunities.

Forward-looking statements are neither historical facts nor assurances of future performance. Instead, they are based only on our current beliefs, expectations and assumptions regarding the future of our business, future plans and strategies, projections, anticipated events and trends, the economy and other future conditions. Because forward-looking statements relate to the future, they are subject to inherent uncertainties, risks and changes in circumstances that are difficult to predict and many of which are outside of our control. Our actual results and financial condition may differ materially from those indicated in the forward-looking statements. Therefore, you should not rely on any of these forward-looking statements. Important factors that could cause our actual results and financial condition to differ materially from those indicated in the forward-looking statements include, among others, the following: those related to our future financial performance, our ability to raise additional funding when needed, our ability to develop and maintain partnerships, our ability to identify and develop new products in a timely manner, the outcome, cost and timing of our product development activities and clinical trials, market size and acceptance of our products, our ability to maintain, protect and enhance our brand and intellectual property, our ability to continue to stay in compliance with applicable laws and regulations, our ability to scale our business and make key hires and such other factors as discussed under the section titled “Risk Factors” and elsewhere in our Annual Report on Form 10-K, definitive proxy statement and quarterly reports on Form 10-Q that we file with the Securities and Exchange Commission (“SEC”) as well as our other filings and the documents incorporated by reference therein, with the SEC.

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Company Highlights

- Discovery, development, and commercialization of oral, small molecule protease inhibitors
- Lead program sebetralstat for on-demand treatment of orphan disease hereditary angioedema (HAE)
- Sebetralstat phase 3 KONFIDENT trial data anticipated H2 2023, NDA H1 2024
- Sebetralstat would be first oral on-demand HAE therapy and has market transformation potential
- Preclinical oral Factor XIIa program advancing towards IND
 - Initial development in HAE, additional potential indications include thrombosis, inflammation
- All programs internally developed, with full rights and IP protection to at least late-2030s
- Funded into 2025 with \$171.7 million at January 31, 2023

Program Portfolio

Product	Indication	Preclinical	Phase 1	Phase 2	Phase 3	Status
Sebetralstat	On-Demand HAE	KONFIDENT				Trial enrolling, data H2 '23
		KONFIDENT-S (Open-Label Extension)				Trial enrolling
		Orally Disintegrating Tablets				Advancing to sNDA as lifecycle extension
Oral Factor XIIa	HAE Prophylaxis					IND targeted for 2023
Oral Factor XIIa	Thrombosis, inflammation					Future opportunities under evaluation



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Hereditary Angioedema (HAE)

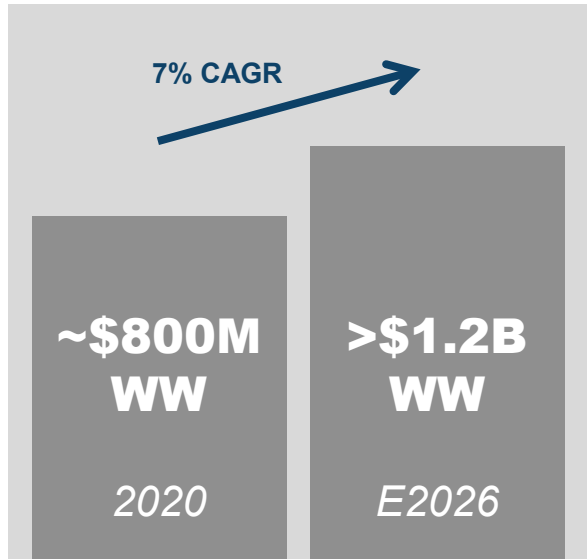
Hereditary Angioedema (HAE)

- Genetic condition causing painful and pronounced swelling in various parts of the body
 - Primarily caused by defect in C1 inhibitor activity, which leads to uncontrolled plasma kallikrein activity and bradykinin release
- Orphan disease: incidence 1 in 10,000 to 1 in 50,000¹
 - Minimum of 6,500 – 8,000 patients in US; similar in EU
 - Incidence consistent worldwide; average patient has ~2 attacks/month
- Approved therapies are primarily injected/infused - high unmet need for efficacious oral administration

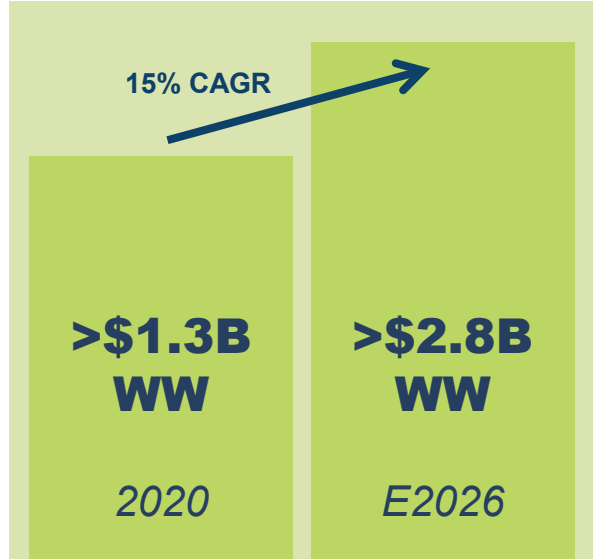
¹www.haei.org

Branded HAE Market Estimated to Be \$4B Revenues by 2026




On-demand/ Breakthrough Treatment



Prophylaxis



Additional Market Growth

- 
Untreated
 Half or more of attacks are not currently treated, including in prophylaxis
- 
Undiagnosed
 30-60% globally
- 
nC1
 Normal C1-INH HAE (currently **no treatments**)

The Sebetralstat Market Opportunity

Leading Share
of Existing
On-Demand
Market:
90k+ scripts/yr at
branded prices



Increased usage
for both on-
demand and
prophylaxis
patients



Patients switching
back from prophy

- Seeking
efficacious and
safe oral option



Future Indications

- Normal C1-INH
HAE
- Short-term
prophylaxis



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Sebetralstat: HAE On-Demand Therapy

Level of Unmet Need in HAE Therapy Is Underappreciated



- All current on-demand products **injected/infused**
- **Pain** and other challenges are **major barriers** to treatment



- Attack treatment is commonly **delayed** and often **too late**
- Some studies show that 50% or more of attacks **aren't treated at all**^{1,2}



- Attacks **still occur** in patients on prophylaxis
- 43% of prophyl patients surveyed **make changes** to try to avoid attacks³

The Sebetralstat Treatment Vision Is Much Better for Patients



- Patients treat at **first symptoms** of attack – **before** swelling begins
- Halt attack at **earliest stages**

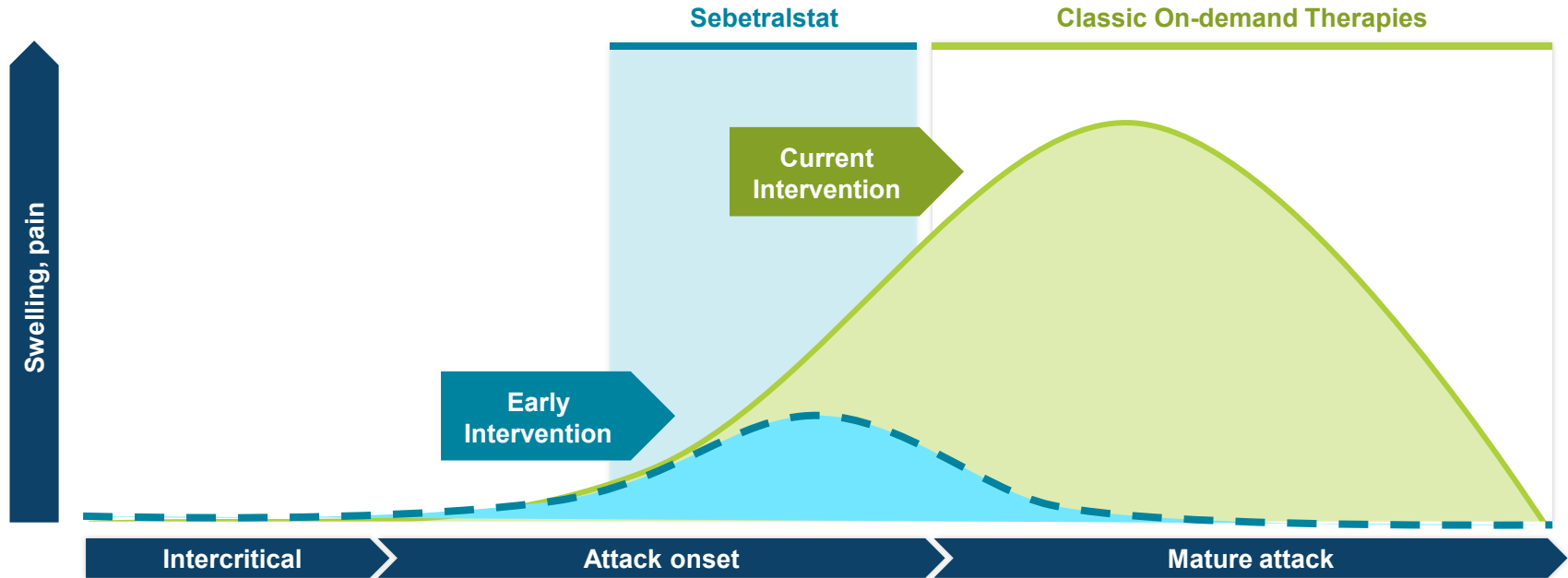


- Patients **treat all attacks**, since it is impossible to predict which ones will **escalate in severity**
- Current **barriers** to treatment are **removed**



- Sebetralstat dosing is **simple** with easy-to-take **tablets**
- Patients can **easily carry, store and access** when needed
- **Oral disintegrating tablet** a further benefit for **pediatrics** and other patients

“Flattening the Curve” of HAE Attack Progression With Early Initiation of On-demand Therapy



Current On-demand Has Many Challenges

Patients say an on-demand oral treatment would simplify the treatment decision and change how and when they treat attacks

95%

*of time patients
would carry
treatment¹*

94%

*of attacks patient
would treat²*

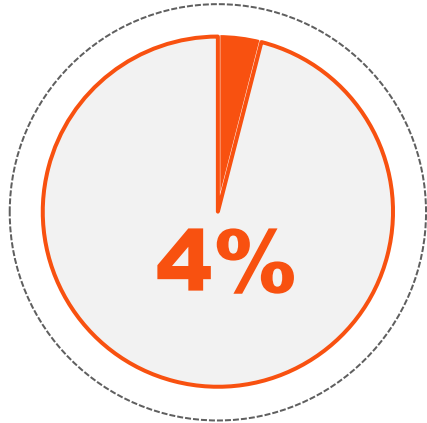
93%

*of patients who
would treat
earlier³*

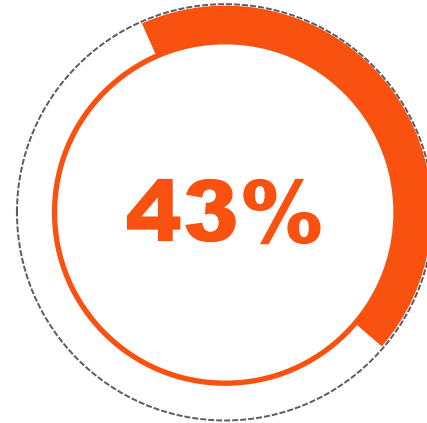
1. Percent of time patients would carry an effective oral on-demand treatment with them outside the home.
2. What percentage of attacks would you treat with an HAE on-demand pill/tablet?
3. Would you treat your attacks faster/earlier with an HAE on-demand pill/tablet?

Even Prophylaxis Patients Have High Unmet Need

While prophylactic treatment reduces attack rates, it does not appear to deliver on the quality of life promised to patients. Prophylaxis patients still do not feel their 100% self and frequently modify their life to avoid triggers.



Only 4% of patients on prophylactic treatment feel their 100% self all of the time

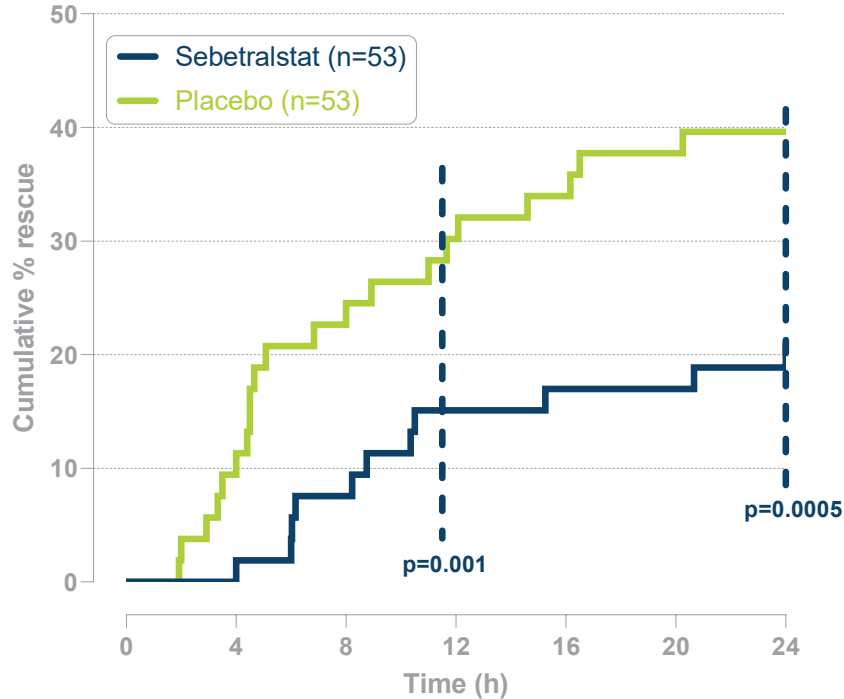


43% of prophylaxis patients adjust their lifestyle to avoid triggers

Positive Sebetralstat Phase 2 Clinical Trial Results

- Sebetralstat met all primary and secondary endpoints
 - Including endpoint being used as primary in ongoing KONFIDENT phase 3 study
- Sebetralstat enables early intervention to maximize treatment success
 - Median time to treatment was 30 minutes
- Significantly improved patient-reported outcomes of treatment effect and attack severity
 - Reduced time to beginning of symptom relief and attack resolution
- Sebetralstat was generally safe and well tolerated
 - No serious adverse events reported, and no patients withdrew due to adverse events

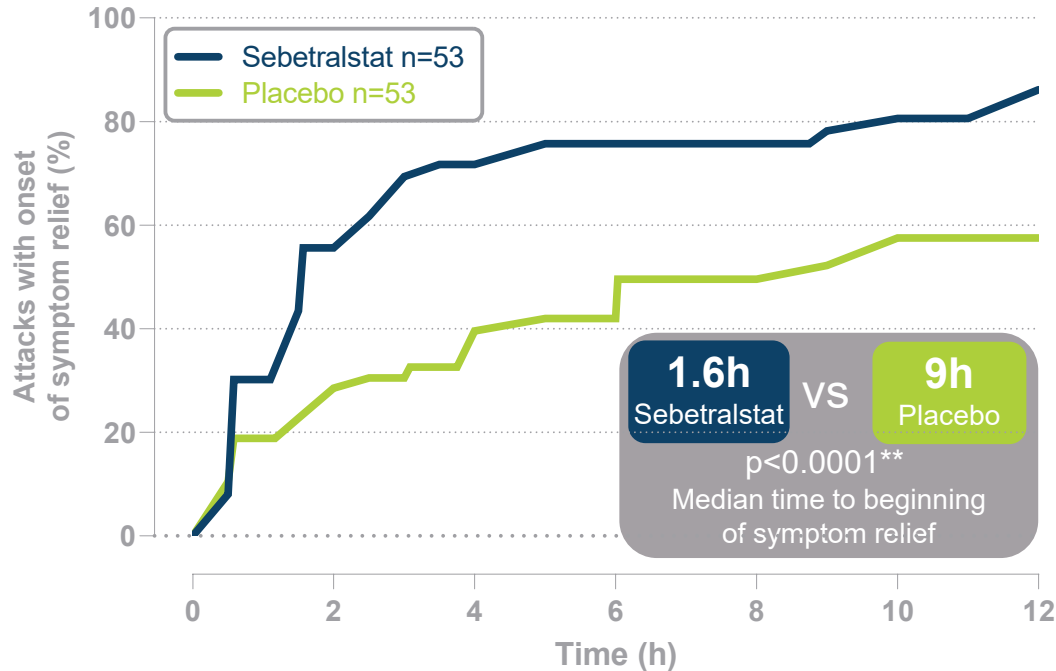
Sebetralstat Met Primary Phase 2 Endpoint



- Sebetralstat significantly increased time to rescue within 12 hours: $p=0.001^*$
 - Placebo 30.2%
 - Sebetralstat 15.1%
- Efficacy maintained at 24 hours
 - $p=0.0005^*$

*Gehan's Generalized Wilcoxon Test

Sebetralstat Reduced Time to Beginning of Symptom Relief



- Assessed using Patient Global Impression of Change (PGI-C)
- Highly significant treatment effect compared to placebo
- **Phase 3 primary endpoint**

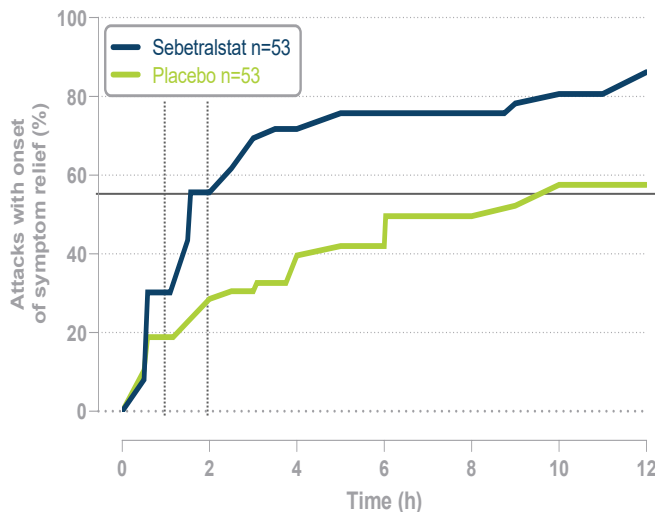
***Gehan's Generalized Wilcoxon Test Full analysis set*

*Censoring occurs where HAE attack was not rated "a little better" or higher or conventional attack treatment was used within 12h
Data on File. KalVista Pharmaceuticals, Inc.*

Sebetralstat Symptom Relief Time Similar to IV-Injected Ruconest (rC1-INH)

Sebetralstat 1.6 hours; Placebo 9 hours
 $p < 0.0001^*$

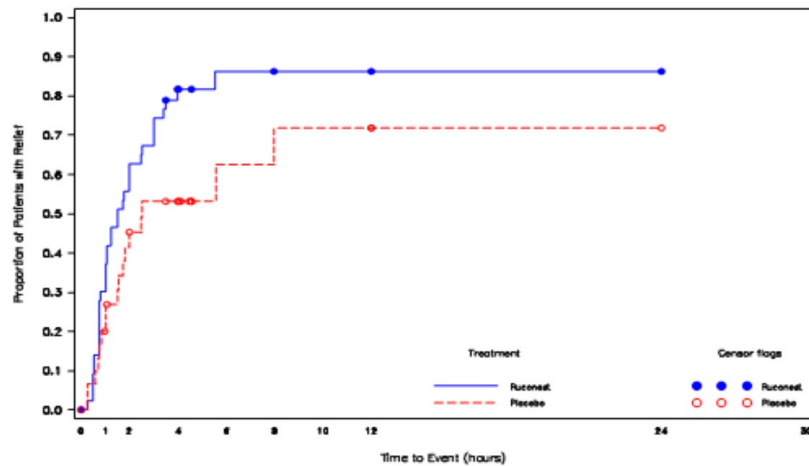
Patient Global Impression of Change (PGI-C)



Ruconest 1.5 hours; placebo 2.5 hours
 $p = 0.031$

Primary endpoint Ruconest Phase 3

Study 1310: Kaplan-Meier Plot of Time to Beginning of Relief of Symptoms with Persistence (Based on Questions 1 and 2 of the TEQ, with Persistence) in the RCT Phase: RCT ITT Analysis Set



*Gehan's Generalized Wilcoxon Test

Ruconest data from Charles M. Maplethorpe, MD, PhD. Clinical Reviewer. Summary Basis of Approval, Recombinant C1 Esterase Inhibitor, STN: 125495/0. The results of the trials for sebetralstat and Ruconest may not be directly comparable, as they are not from a single head-to-head clinical trial

Sebetralstat Phase 3 Trial Design

- Double-blind, crossover trial assessing 300 mg and 600 mg sebetralstat versus placebo
- Each patient treats 3 attacks at home
 - One with each treatment in a randomized, blinded sequence
 - Patients can redose if symptoms warrant, no statistical impact on primary or secondary endpoints



Primary endpoint:

- Time to beginning of symptom relief (PGI-C)

Key secondary endpoints:

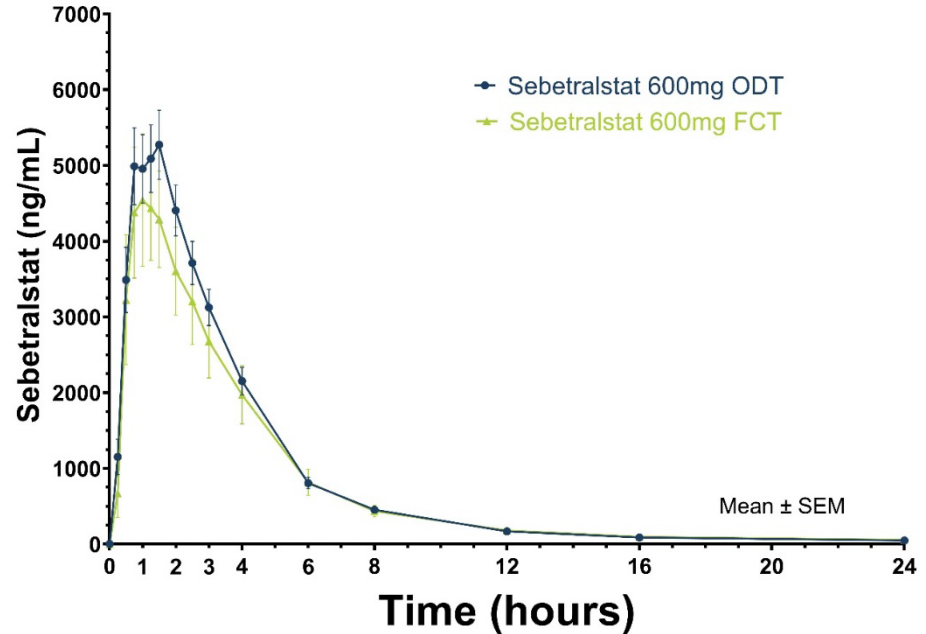
- Time to improvement (PGI-S)
- Time to resolution (PGI-S)

Sebetralstat Phase 3 Trial Design

- The KONFIDENT trial will end when 84 patients have completed three attacks
 - HAE type 1 and 2, including adults, adolescents and patients with ongoing prophylaxis
 - All attack locations eligible, including laryngeal attacks
- FDA-agreed primary endpoint: time to beginning of symptom relief using PGI-C
 - *A priori* secondary endpoint in phase 2 trial
 - Expected to be sufficient to file NDA
 - FDA does not recommend the use of VAS to support the primary endpoint
- At least 90% power to detect the phase 2 treatment effect
- First patient recruited March 2022; Data anticipated H2 2023, NDA filing H1 2024

Orally Disintegrating Tablet (ODT) Further Enhances Options for Patients

- ODT increases ease of dosing for younger patients or those with difficulty swallowing
- Phase 1 data shows similar pharmacokinetics to current film-coated tablets (FCT)
- Expected to be lifecycle extension in US and EU; potentially launch formulation in other geographies



*KalVista data on file

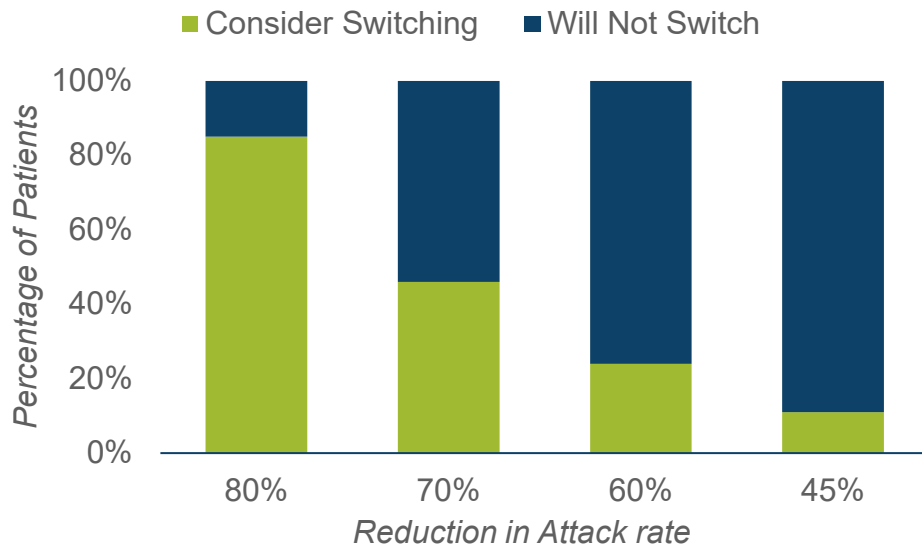


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Factor XIIa

Oral Factor XIIa Designed to Meet Unmet Need in HAE Prophylaxis

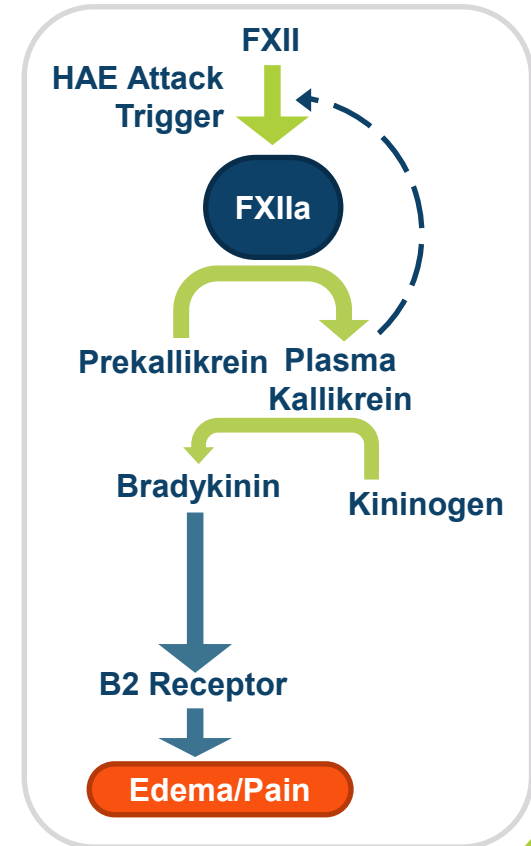
Patients will not trade lower efficacy for oral dosing



➔ Our goal is to achieve an oral treatment with an efficacy profile similar to injectable treatments

Factor XIIa Sits at Top of HAE Attack Cascade

- Factor XIIa (FXIIa) activates the kallikrein kinin system
 - Generates plasma kallikrein, leading to uncontrolled bradykinin release in HAE
 - FXIIa and plasma kallikrein inhibitors selectively block bradykinin generated by the plasma kallikrein kinin system, unlike bradykinin receptor antagonists
- FXIIa inhibitory antibody has been shown clinically to reduce HAE attack frequency
 - At least as efficacious as approved therapies against other targets
 - No known chronic safety implications

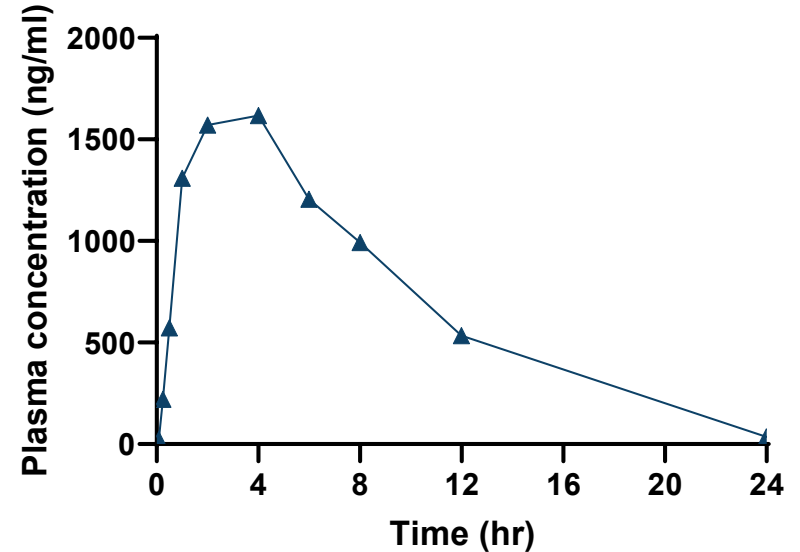


KalVista Is Building a Portfolio of Potent, Oral FXIIa Inhibitors

High potency for FXIIa in multiple series

Compound	FXIIa IC ₅₀ (nM)	Series
1	10	A
2	9.7	A
3	12	B
4	1.9	B
5	7.8	C
6	7.5	C
7	3.5	C
8	2.7	C

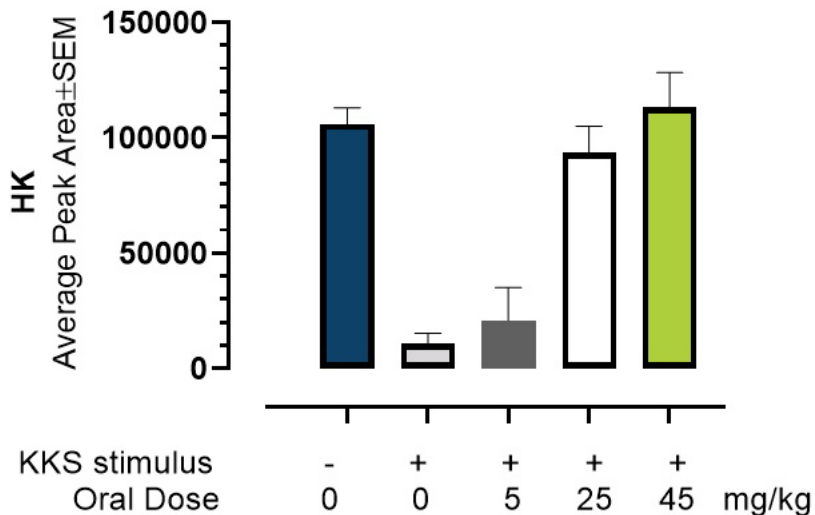
Plasma exposure following oral dosing in rats



- High potency and >1000-fold selectivity against other proteases
- Oral profiles consistent with once-daily dosing in humans
- Multiple distinct series reduce risk, yield broader IP and support development for multiple indications

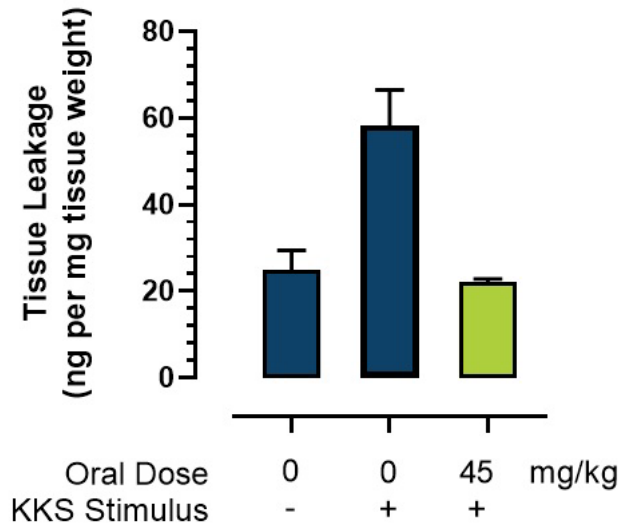
Orally delivered FXIIa Inhibitor Blocks Kallikrein-Kinin System Effects in Mouse Models of Angioedema

Blocks cleavage of kininogen and release of bradykinin



Carrageenan-stimulated paw edema model

Prevents angioedema in HAE-related tissues



Angiotensin-converting enzyme inhibitor angioedema model

Factor XIIa Inhibitor Program Is Advancing Toward IND

- Strong scientific rationale and positive clinical evidence for FXIIa inhibition in HAE prophylaxis
- KalVista is advancing the first oral FXIIa inhibitors towards the clinic – no known competitors in oral therapy
- FXIIa implicated in other indications that may represent large future opportunities, including thrombosis and inflammation
- IND enabling studies underway

KalVista Value Proposition

- The only company developing distinct oral treatments for the full spectrum of HAE disease management
- Sebetralstat on-demand phase 3 KONFIDENT data expected H2 2023, NDA H1 2024
- Sebetralstat will be the first oral on-demand therapy and has market transformation potential
- Oral FXIIa inhibitor program future growth opportunity
 - Initial development in HAE prophylaxis, based on clinical validation of target and high unmet need
 - FXIIa also has potential in large market opportunities in inflammation and thrombosis
- Funded into 2025



NASDAQ: KALV
