

# KODIAK

THE OPHTHALMOLOGY MEDICINES COMPANY

**Corporate Presentation** 

August 2024 🛸

#### SPECIAL NOTE REGARDING

### FORWARD-LOOKING STATEMENTS

These slides contain "forward-looking statements." Forward-looking statements are based on our current expectations of future events and are subject to risks and uncertainties that could cause actual results to differ materially and adversely from those in or implied by such forward-looking statements. For a discussion of risks and uncertainties and other important factors, any of which could cause our actual results to differ from those contained in the forward-looking statements, see the section entitled "Risk Factors" in our most recent Form 10-K, as well as discussions of potential risks, uncertainties, and other important factors in our subsequent filings with the SEC. These forward-looking statements speak only as of the date hereof and Kodiak undertakes no obligation to update forward-looking statements, and readers are cautioned not to place undue reliance on such forward-looking statements. Kodiak®, Kodiak Sciences®, ABC™, ABC Platform™, and the Kodiak logo are registered trademarks or trademarks of Kodiak Sciences Inc. in various global jurisdictions.

# THE OPHTHALMOLOGY MEDICINES COMPANY OUR MISSION



### TRAILBLAZING SCIENCE

Our creative and thoughtful foundation



### 2 "GO-TO" MEDICINES

Our challenge to the status quo



### **3** SINGULAR FOCUS IN OPHTHALMOLOGY

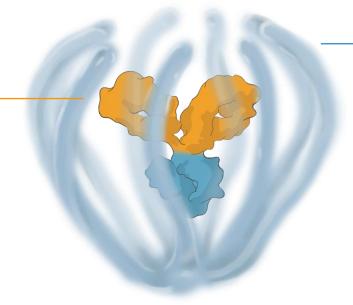
Our 24 / 7 / 365

### ABC Platform: Enabling multi-mechanism therapies empowered for durability

 Our Antibody Biopolymer Conjugate ("ABC") Platform combines the best durability with the right efficacy and is the foundation of tarcocimab tedromer and KSI-501, two "ABC" investigational medicines in late-phase clinical development

### Antibody or Other Biologic

Engineered to exhibit high binding affinity and specificity. Any biologic including antibodies and aptamers can be conjugated to the biopolymer via a stable, site-specific linkage



### **Biopolymer**

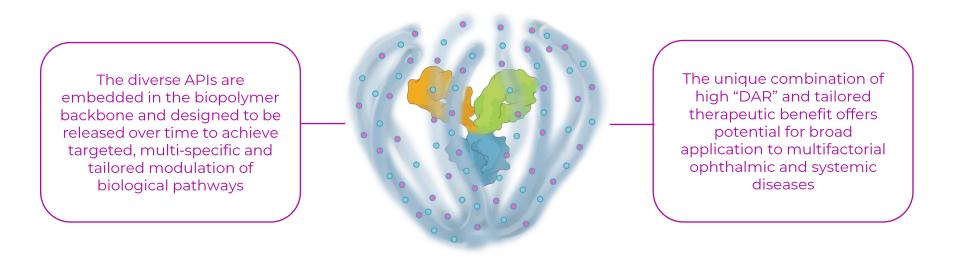
Engineered to make medicines last longer and extend their therapeutic benefit. It is also designed to combine multiple active pharmaceutical ingredients (APIs)

The biopolymer is optically clear and made of phosphorylcholine, the primary hydrophilic component of human cell membranes

Antibody Biopolymer Conjugate ("ABC")

### ABC Platform Evolution: Designed for multi-specific, high "DAR" medicines

 We are expanding our early research pipeline of duets and triplets built from our modular ABC Platform that embeds diverse active pharmaceutical ingredients ("APIs") including small molecules, proteins, peptides, macrocycles and oligonucleotides in the biopolymer backbone to enable high drug-antibody ratio ("DAR") medicines with targeted, multispecific, tailored modulation of biological pathways for ophthalmic and systemic diseases



### Our product candidates: a portfolio of three clinical programs to address key limitations of today's therapies across a broad spectrum of retinal diseases

"ABC" Platform-derived biologics: The best durability with the right efficacy for high-prevalence retinal vascular diseases



Tarcocimab Anti-VEGF ABC

- **Objective:** to have a compelling first-line durability profile without compromising immediacy
- Longest-acting anti-VEGF biologic (6month predominant) while preserving the flexibility to dose monthly
- An intravitreal biologic that can be used in any patient whether they be in the loading (immediacy) phase or in the maintenance (durability) phase
- Enhanced 50 mg/mL formulation



KSI-501 Bispecific Anti-IL-6, VEGF Trap ABC

- **Objective:** to address the opportunity for first-line efficacy with the best durability
- First-in-class bispecific "ABC" designed to address retinal inflammation and vascular permeability simultaneously
- Reflects 10 years of learnings of the "ABC" platform to maximize each patient's efficacy and durability potential
- Enhanced 50 mg/mL formulation

#### Unconjugated biologic: For inflammatory retinal diseases



**KSI-101** Bispecific Anti-IL-6, VEGF Trap Protein

- Objective: to address the underlying disease mechanisms of macular edema secondary to inflammation ("MESI") for which no approved intravitreal biologic therapies exist today
- First-in-class bispecific protein designed to address retinal inflammation and vascular permeability simultaneously
- 100 mg/mL formulation provides
   high strength and potency

#### Science Updates for our "ABC" Platform Biologics (Tarcocimab and KSI-501)

- 1. Supported by a true science of durability (conjugate design, animal ocular  $t_2^{1/2}$  data and human ocular  $t_2^{1/2}$  data) in contrast to current anti-VEGFs
- 2. Enhanced formulation containing both conjugated and unconjugated antibody is intended to balance durability and immediacy

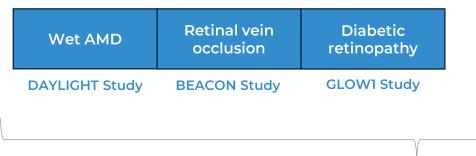
VEGF: vascular endothelial growth factor; IL-6: interleukin 6; mAb: monoclonal antibody

### TARCOCIMAB TEDROMER

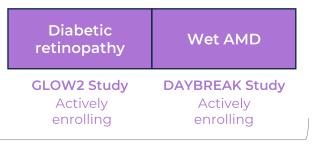
Our objective is for tarcocimab to have a compelling first-line durability profile without compromising immediacy for patients with high-prevalence retinal vascular diseases

- Five Phase 3 studies are planned for inclusion in a Biologic License Application (BLA)
- Three are complete with compelling durability demonstrated and two are in process

#### Completed Phase 3 studies: Primary endpoint met and extended durability demonstrated



#### Two new Phase 3 studies in process: Using the enhanced formulation of tarcocimab



### Planned BLA package

### Three successful Phase 3 studies in diabetic retinopathy (DR), retinal vein occlusion (RVO) and wet AMD with compelling durability demonstrated

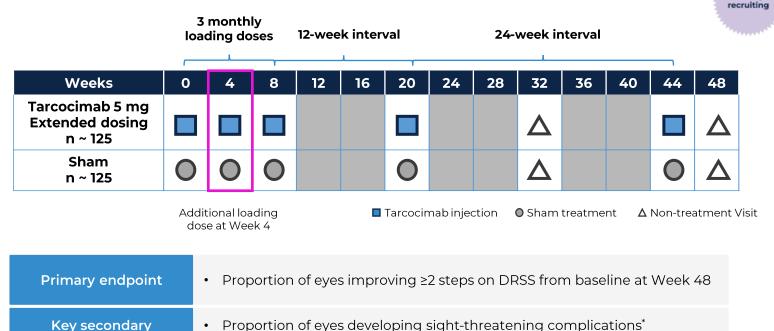
	Study design	Primary endpoint	Extended durability	
Diabetic retinopathy Phase 3 GLOW1 Study	<ul> <li>Superiority study</li> <li>tarcocimab Q24W after 3 initiating doses vs sham</li> </ul>	$\checkmark$	$\checkmark$	Signature durability demonstrated with all patients on 6-month dosing
Retinal vein occlusion Phase 3 BEACON Study	<ul> <li>Tarcocimab Q8W after 2 monthly loading doses vs aflibercept Q4W</li> </ul>	$\checkmark$	~	Doubled treatment interval at primary endpoint (month 6) and ~50% of patients on 6-month dosing at Year 1
Wet AMD Phase 3 DAYLIGHT Study	<ul> <li>Tarcocimab Q4W vs aflibercept Q8W after 3 monthly loading doses</li> </ul>	$\checkmark$	Not Applicable	Monthly dosing of tarcocimab demonstrated favorable safety and non- inferior efficacy at Year 1

 In addition to these studies, tarcocimab was also studied in the Phase 2b/3 DAZZLE study in wet AMD and in the Phase 3 GLEAM and GLIMMER studies in DME. These studies did not meet primary endpoint but did demonstrate strong 5 and 6-month durability in the majority of patients.

### New Phase 3 study: GLOW2 in Diabetic Retinopathy

endpoints

 GLOW2 features a similar study design as the successful GLOW1 study, with the benefit of an additional 3<sup>rd</sup> monthly loading dose (Week 0, 4, 8)



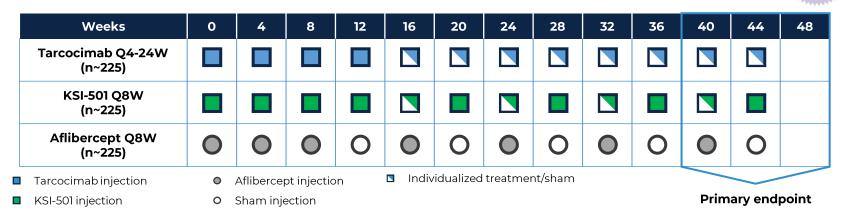
Proportion of eyes improving ≥3 steps on DRSS from baseline

KODIAK 'Sight-threatening complications were defined as: proliferative diabetic retinopathy (PDR), Vitreous hemorrhage or tractional retinal detachment due to PDR, diabetic macular edema, and anterior segment neovascularization. DRSS: diabetic retinopathy severity score; GLOW1 study NCT05066230; GLOW2 study NCT06270836

Actively

### New Phase 3 study: DAYBREAK in Wet AMD

- DAYBREAK includes parallel investigational arms of tarcocimab and KSI-501 vs aflibercept
- The objective is to evaluate the efficacy and safety of tarcocimab and KSI-501 and to support registration in wet AMD for both investigational medicines



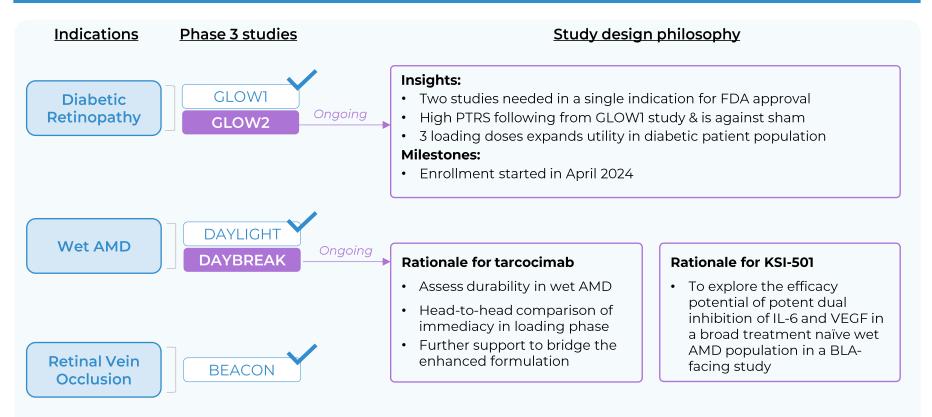
• Mean change in BCVA (ETDRS letters) from baseline to the average of Week 40, 44 and 48

- Tarcocimab: individualized Q4W to Q24W dosing following 4 monthly loading doses to assess 6-month durability potential
- KSI-501: fixed Q8W dosing with additional individualized dosing (up to monthly dosing) following 4 monthly loading
  doses to explore the efficacy potential of potent dual inhibition of IL6 and VEGF in a broad treatment naïve wet AMD
  population

Actively

recruiting

### GLOW2 and DAYBREAK study: philosophy and rationale



"My experience with tarcocimab-treated patients in your trial is you have the durability but you didn't dry as well in the loading phase. But with a formulation of conjugated and unconjugated antibody, then you have fixed that, and you have a drug that primes itself and then takes patients longer. Together with monthly reimbursement where needed, I don't know why you wouldn't be a contender for first-line after step therapy from Avastin."

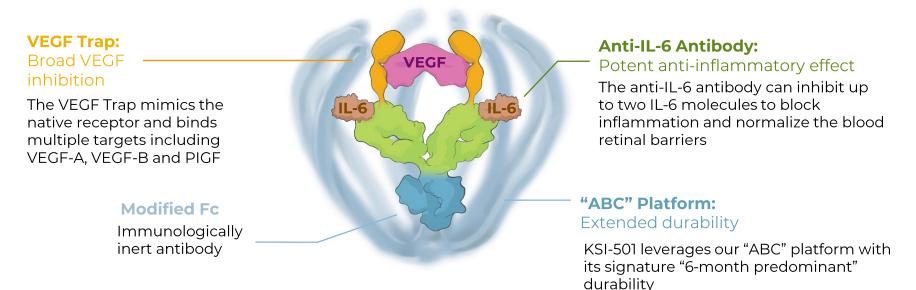
### **KSI-501**

Designed to address the opportunity for improved efficacy with extended durability in high-prevalence retinal vascular diseases by targeting retinal inflammation and vascular permeability simultaneously

- Inflammation has been shown to play a significant role in high-prevalence retinal vascular diseases. However, no treatments exist that concurrently address vascular permeability and inflammation
- KSI-501 is designed to inhibit VEGF and interleukin-6 (IL-6), a pro-inflammatory cytokine and immune growth factor, combining two powerful mechanisms of action to address retinal vascular disease and the underlying inflammatory cascade

# KSI-501 is a first-in-class bispecific designed for highly efficient binding to both IL-6 and VEGF, built on Kodiak's "ABC" platform

• The anti-permeability effect of VEGF inhibition is the primary effector, with the anti-inflammatory effect of IL-6 inhibition offering the potential for additional clinical benefits

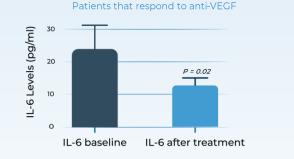


Enhanced 50 mg/mL formulation containing both conjugated and unconjugated forms is intended to balance durability and immediacy

### KODINK

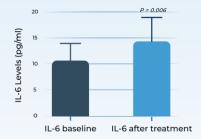
## In addition to VEGF, IL-6 driven inflammation is implicated in retinal vascular disease

- IL-6 is a pro-inflammatory cytokine and immune growth factor implicated in the pathophysiology of multiple retinal diseases
  - Vitreous IL-6 levels are significantly elevated in retinal disease patients vs controls
  - IL-6 stimulates defective angiogenesis independent of VEGF and is implicated in anti-VEGF treatment resistance
  - Increased levels of IL-6 are associated with poor functional outcomes in wet AMD and diabetic macular edema (DME) patients treated with anti-VEGF monotherapy
- Adapted from Chalam et al. (2014). Journal of Ophthalmology, Article ID 502174. Mean with SEM plotted.
- Adapted from Sepah, Y.J., Do, D.V., Mesquida, M. et al. Aqueous humour interleukin-6 and vision outcomes with anti-vascular endothelial growth factor therapy. Eve (2024)

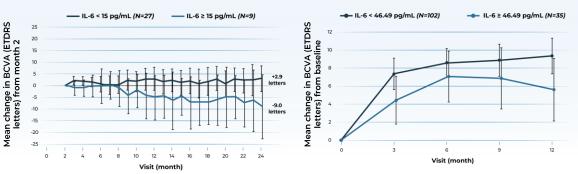


### Aqueous humor IL-6 levels significantly correlate with anti-VEGF treatment response in wAMD<sup>1</sup>

Anti-VEGF treatment resistant patients

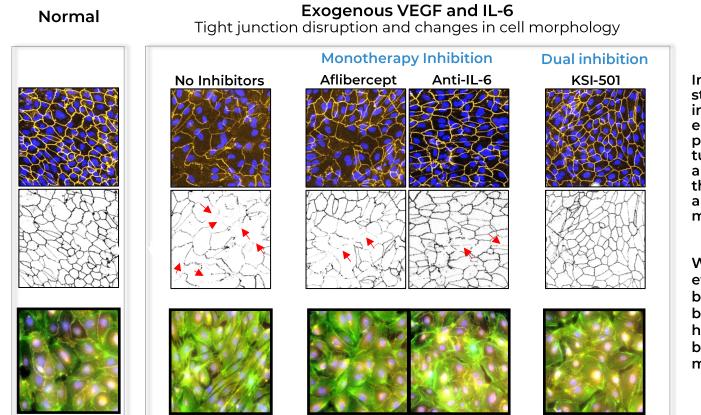


Higher levels of IL-6 in aqueous humor are correlated with poorer BCVA outcomes over time in wet AMD (left) and DME (right)<sup>2</sup>



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# Dual inhibition of VEGF and IL-6 by KSI-501 confers superior normalization of barrier biology vs anti-VEGF or anti-IL-6 monotherapy in preclinical studies



In additional studies, KSI-501 inhibits endothelial cell proliferation and tube formation to a greater extent than anti-VEGF or anti-IL-6 monotherapy

With synergistic effect on the blood retinal barrier, KSI-501 holds potential to be a new diseasemodifying therapy

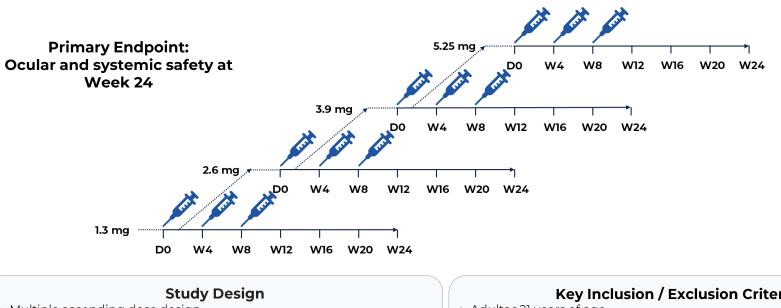
Vascular Cells

**RPE** Cells



RPE cells: nuclei in blue, ZOI (tight junction protein) in yellow. Red arrows indicate gaps in ZO-1 tight junction protein. Vascular cells: nuclei in purple, ZOI (tight junction protein) in yellow, actin in green.

### Phase 1 study of KSI-501 was a multiple ascending dose study in patients with diabetic macular edema



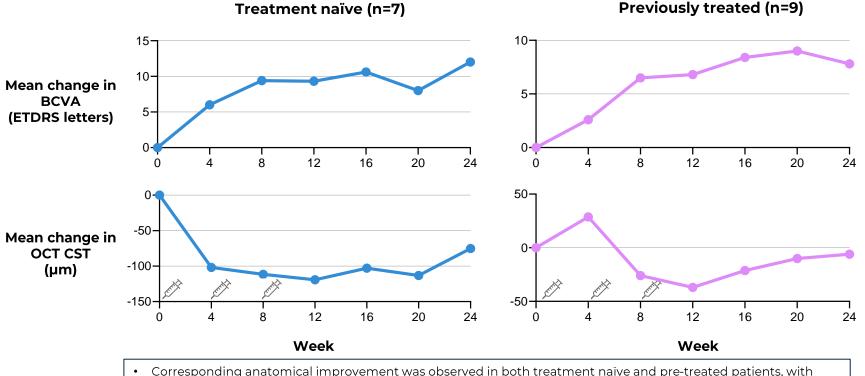
- Multiple ascending dose design
- Conducted at 5 sites in the US
- 3-5 subjects planned to be enrolled for each dosing group, with option for expansion of each group if indicated
- Each subject received 3 monthly doses and was followed for 24 weeks total

#### Key Inclusion / Exclusion Criteria

- Adults ≥21 years of age
- Diabetes mellitus type 1 and 2 (HbA1c ≤12%)
- Vision loss due to DME
- BCVA between 25 and 70 ETDRS letters (20/40 20/320 Snellen)
- DME (CST ≥320 microns)
- Treatment naïve and previously treated with an 8-week washout period

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### KSI-501 demonstrated robust and meaningful visual acuity gains that were sustained over 16 weeks in both treatment naïve and pre-treated patients



- Corresponding anatomical improvement was observed in both treatment naïve and pre-treated patients, with meaningful and sustained improvement in treatment-naïve patients
- Treatment naïve patients are planned to be the target population of Phase 3 studies

n = Number of participants treated;

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BCVA: best-corrected visual acuity; ETDRS: early treatment diabetic retinopathy study; OCT: optical coherence tomography; CST: central subfield thickness

# KSI-501 development plan: Phase 3 DAYBREAK study is actively enrolling patients, with parallel investigational arms of KSI-501 and tarcocimab

Phase 1 study	Diabetic macular edema			
<ul> <li>A multiple ascending dose study of KSI-501 in patients with DME, both treatment naïve and pretreated patients</li> <li>DME is known to have high levels of cytokine-mediated microvascular inflammation in addition to VEGF-mediated vascular permeability</li> <li>Results: repeated monthly dosing of KSI-501 was (1) safe and well tolerated, and (2) achieved clinically meaningful and sustained visual acuity gains and CST reduction</li> </ul>				
Phase 7 DAVRDEAK study				
<ul> <li>Phase 3 DAYBREAK study</li> <li>A 3-arm study: <ul> <li>Tarcocimab dosed Q4W-Q24W after 4 monthly loading doses</li> <li>KSI-501 dosed Q8W with additional individualized Q4W dosing after 4 monthly loading doses</li> <li>Active comparator aflibercept 2mg dosed per label</li> </ul> </li> <li>Enhanced 50 mg/mL formulations of conjugated and unconjugated forms for both tarcocimab and KSI-501 are intended to balance durability and immediacy</li> </ul>		Actively enrolling		

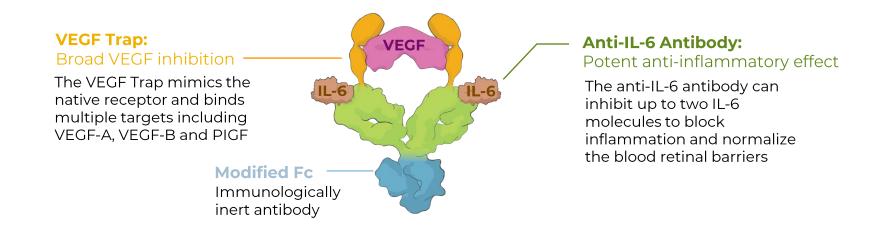
### KSI-101

A potent 100 mg/mL high strength bispecific protein being developed for the treatment of macular edema secondary to inflammation ("MESI") for which no approved intravitreal biologic therapies exist today

- In patients with intraocular inflammation, significant vision loss is most commonly a consequence of macular edema
- Studies show that inflammation and vascular permeability have a synergistic effect on driving disease progression and vision loss due to macular edema, but there are no approved therapies that target both drivers of disease

### KSI-101 is a first-in-class bispecific protein for the powerful treatment of macular edema secondary to inflammation ("MESI")

- KSI-101 is a bispecific protein designed to directly target both IL-6 mediated inflammation and edema, and VEGF-mediated vascular permeability
- The anti-inflammatory effect of IL-6 inhibition is the primary effector, with the anti-permeability effect of VEGF inhibition having an additive and synergistic effect



Currently there are no available intravitreal biologic therapies addressing the spectrum of inflammatory conditions of the retina. Our goal is for KSI-101 to target both underlying disease mechanisms concurrently to prevent vision loss for patients who have macular edema and inflammation

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### Macular edema is the leading cause of vision loss among patients with intraocular inflammation and IL-6 mediated pro-inflammatory signaling is a key disease driver

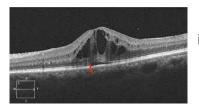
- Macular edema is the leading cause of vision loss among patients with intraocular inflammation
- Signaling mediated by pro-inflammatory cytokines including IL-6 is a key disease driver of macular edema secondary to inflammation ("MESI")
  - Leads to the disruption of the inner and/or outer blood-retina barrier and accumulation of fluid
- Currently there are no approved, targeted therapies
  - Existing treatment is limited in efficacy and has undesirable side effects
  - There is only one other biologic in late-stage clinical development



Intraocular inflammation is the 4<sup>th</sup> leading cause of vision loss in the developed world

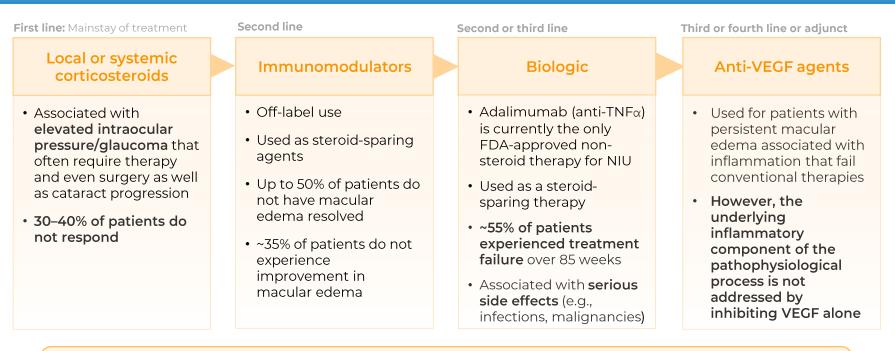
- Up to 50% of patients experience reduced vision
- 10-15% of patients become blind

#### Macular edema is the leading cause of vision loss among patients with intraocular inflammation



1/3 of patients with intraocular inflammation develop macular edema (~110,000 patients in the U.S.)

### Current treatment algorithm for macular edema secondary to inflammation: high unmet need for safer therapies that target the underlying mechanisms of disease



There is an unmet need for potent therapies with a better safety profile. With bispecific IL-6 and VEGF inhibition which confer a synergistic anti-inflammatory and anti-permeability effect, along with the proven safety profile of an intravitreal biologic, KSI-101 can become the first line therapy for all retinal diseases with an inflammatory component

#### NIU: non-infectious uveitis

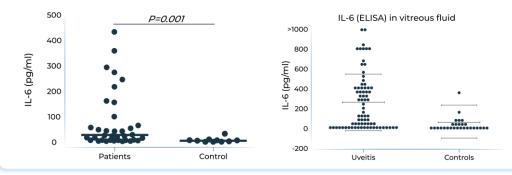
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Sources: 1. Tomkins-Netzer O et al. Ophthalmology. 2015. 122:2351-2359. 2. Jaffe et al. N Engl J Med. 2016. 375:932-43. 3. Rosenbaum et al. Sem Arthrit. 2019. 49: 438-445.

## Studies show that both IL-6 and VEGF play a key role in retinal inflammatory disease

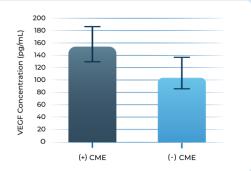
 IL-6 levels are elevated in ocular compartments and in serum in patients with non-infectious uveitis, and further elevated in uveitis patients with macular edema

# Aqueous Humor IL-6 levels are<br/>elevated in patients with<br/>intermediate uveitis1IL-6 levels are elevated in<br/>vitreous fluid of patients with<br/>active uveitis2



• Additionally, persistent inflammation triggers VEGF upregulation. VEGF levels are found to be elevated in aqueous humor of eyes with uveitis and uveitic macular edema, which can lead to angiogenesis, vascular leakage and blood-retinal barrier dysfunction

VEGF levels are elevated in aqueous humor of uveitis patients with macular edema vs without macular edema<sup>3</sup>



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### KSI-101 development plan: Phase 1b APEX study is actively enrolling; FDA alignment on pivotal program: paired studies with 16-week primary endpoint

PHASE 1B			
APEX study	Diabetic macular edema Macular edema secondary to inflammation (MESI)		
<ul> <li>Cohort 1: evaluate 3 dose levels of KSI-101 in DME patients</li> <li>Cohort 2: evaluate 3 dose levels of KSI-101 in patients with MESI</li> <li>Goal is to evaluate safety and tolerability of KSI-101 and to identify 2 dose levels to progress into pivotal studies</li> </ul>		Actively enrolling	

### PHASE 2B/3 (DUAL STUDIES)

PEAK study	Macular adama secondary to inflammation (MESI)	
PINNACLE study	Macular edema secondary to inflammation (MESI)	
<ul> <li>Objective is to evaluate the efficacy and safety of KSI-101 and to support registration in macular edema secondary to inflammation</li> <li>PEAK and PINNACLE are expected to be identically designed studies with 3 arms: a low dose and a high dose investigational arm of KSI-101 with 4 monthly doses followed by PRN (as needed) dosing and a sham arm</li> <li>Primary endpoint will be vision outcomes at week 16 with study completion at week 48</li> </ul>		Target enrollment TBD

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#### Tarcocimab tedromer



An intravitreal biologic that can be used in any patient whether they be in the loading (immediacy) phase or in the maintenance (durability) phase for high-prevalence retinal vascular diseases

- Supported by our science of durability
- Enhanced 50 mg/mL formulation containing both conjugated and unconjugated antibody is intended to balance durability and immediacy
- Three of five Phase 3 studies complete in three indications
- Phase 3 GLOW2 study in DR and Phase 3 DAYBREAK study in wet AMD are actively enrolling



Designed to address the opportunity for improved efficacy with extended durability in high-prevalence retinal vascular diseases by targeting retinal inflammation and vascular permeability

- Supported by our science of durability
- Enhanced 50 mg/mL formulation containing both conjugated and unconjugated antibody is intended to balance durability and immediacy
- Phase 3 DAYBREAK study in wet AMD is actively enrolling, designed to explore the efficacy potential of potent dual inhibition of IL-6 and VEGF in a broad treatment naïve wet AMD population



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A potent 100 mg/mL high strength bispecific protein being developed for the treatment of macular edema secondary to inflammation (MESI) for which no approved intravitreal biologic therapies exist today

- The anti-inflammatory effect of IL-6 inhibition is the primary effector, with the antipermeability effect of VEGF inhibition having an additive and synergistic effect
- Phase 1b APEX study in DME and MESI is actively enrolling, to evaluate the safety and tolerability and to identify two dose levels to progress into dual Phase 2b/3 studies (PEAK and PINNACLE) in MESI

### KODIAK SCIENCES

- \$219 million in cash and cash equivalents as of end of 2Q24
- Planning to achieve meaningful inflection points within current cash runway