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# KODIAK

THE OPHTHALMOLOGY MEDICINES COMPANY

Corporate Presentation December 2020

# FORWARD-LOOKING STATEMENTS

These slides contain forward-looking statements and information. The use of words such as "may," "might," "will," "should," "expect," "plan," "anticipate," "believe," "estimate," "project," "intend," "future," "potential," or "continue," and other similar expressions are intended to identify forward-looking statements. For example, all statements we make regarding the potential licensure of KSI-301 and a single BLA submission in wet AMD, DME, RVO and diabetic retinopathy in 2022; our platform technology and potential therapies; future development plans; clinical and regulatory objectives and the timing thereof; anticipated design of planned clinical trials; expectations regarding the potential efficacy and commercial potential of our product candidates; the anticipated presentation of data; and our ability to advance our product candidates into later stages of development are forward looking. All forward-looking statements are based on estimates and assumptions by our management that, although we believe to be reasonable, are inherently uncertain. All forwardlooking statements are subject to risks and uncertainties that may cause actual results to differ materially from those that we expected. These statements are also subject to a number of material risks and uncertainties that are described in the preliminary prospectus supplement, in our most recent quarterly report on Form 10-Q, as well as our subsequent filings with the Securities and Exchange Commission. Any forward-looking statement speaks only as of the date on which it was made. We undertake no obligation to publicly update or revise any forward-looking statement, whether as a result of new information, future events or otherwise, except as required by law.

#### KODIAK SCIENCES

## WHERE WE ARE TODAY

4 PIVOTAL TRIALS

**3** INDICATIONS

## **SINGLE** BLA FILING EXPECTED IN 2022

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#### KSI-301 CLINICAL EXPERIENCE

Clinical data from 1,500 injections in 400+ patients representing 250+ patient-years of exposure in representative populations in wAMD, DME and RVO

- Safety: Tracking with current standard of care (Lucentis, Eylea)
- Efficacy: Strong and appropriate impact on vision & retinal anatomy in each indication studied
- Durability: Majority of patients going 6-months or longer in wet AMD and DME

#### OPTIMIZED PIVOTAL STUDY PROGRAM

Objective to show disruptive durability with same safety and efficacy as Eylea

DAZZLE wet AMD study enrollment complete; BEACON RVO study and GLEAM / GLIMMER DME now enrolling – Data from all studies expected in 2022

Pivotal studies designed from phase 1b data with tighter criteria for disease activity assessments, shorter durability intervals, high statistical power, maintaining similar (80%+) U.S. treatment naïve population

#### OPERATING WITH CONVICTION

On track for a single BLA filing in the key indications of wAMD, DME, RVO treatment and with NPDR (prevention) indication in a supplemental BLA

Manufacturing investments aligned to clinical opportunity with commercial supply goal of 2.5M+ Prefilled Syringes in Year 1 of launch

Developing bispecific and triplet ABC Medicines for multi-mechanism diseases, including dry AMD and glaucoma

#### POISED COMMERCIAL OPPORTUNITY

Competitive landscape is clearing with competing molecules/technologies demonstrating poor benefit risk profiles

We believe KSI-301 may be able to capture market share from standard of care agents, future biosimilars, and competing late-stage molecules in development

#### 3

#### THE OPHTHALMOLOGY MEDICINES COMPANY

## OUR MISSION



TRAILBLAZING SCIENCE

Our creative and thoughtful foundation



#### 2 GENERATION 2.0 MEDICINES

Our challenge to the status quo



## **3** SINGULAR FOCUS IN OPHTHALMOLOGY

Our 24 / 7 / 365

A PIPELINE OF ABCs FOR RETINA

Kodiak's deepening pipeline of mono-, bi-specific and triplet inhibitors that merge biologics with small molecules to address major causes of vision loss beyond retinal vascular disease

#### MONOSPECIFIC

#### 1 Molecule, **1 Target**

Antibody conjugated to phosphorylcholine biopolymer

**KSI-301** inhibits VEGF— In Phase 3 clinical development

#### BISPECIFIC

#### 1 Molecule, **2 Targets**

Bispecific antibody conjugated to phosphorylcholine biopolymer

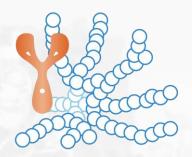
**KSI-501** inhibits VEGF and IL-6 for retinal diseases with inflammatory component - IND planned 2021

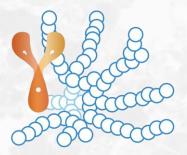
#### TRIPLET

#### 1 Molecule, **3 Targets**

Bispecific antibody conjugated to phosphorylcholine biopolymer embedded with 100's of copies of smallmolecule drug

For high-prevalence multifactorial diseases, such as dry AMD and glaucoma - IND planned 2022







#### THE OPHTHALMOLOGY MEDICINES COMPANY

## FOCUSED ON DEVELOPING ABC MEDICINES<sup>TM</sup> FOR HIGH PREVALENCE RETINAL DISEASES



#### KSI-301 AND KSI-501 FOR RETINAL VASCULAR DISEASES

#### A GROWING \$11B MARKET WITH CLEAR UNMET NEEDS

- · Wet age-related macular degeneration (wet AMD) remains a leading cause of blindness in the elderly
- Diabetes is the leading cause of blindness in working-age adults
- Novel agents like KSI-301 designed to provide long treatment-free durability and/or improve response to therapy are needed
- KSI-501 targets both VEGF & IL-6; supplemental targeting of retinal microvascular inflammation through IL-6 may be of additional clinical benefit

#### **KSI-601 TRIPLETS FOR DRY AMD**

#### DRY AMD IS 10 TIMES MORE PREVALENT THAN WET AMD AND HAS NO AVAILABLE THERAPIES

- Dry AMD frequently leads to irreversible vision loss and substantial functional vision limitations
- There are no available therapies for dry AMD; drugs targeting single pathways have repeatedly yielded no / limited efficacy
- Targeting multiple biological pathways both intracellular and extracellular as enabled by our triplet inhibitor technology may be required for complex, multifactorial diseases like dry AMD
- Durability of a treatment will be key due both to chronic nature of the disease and size of the patient population and will be enabled by ABC Platform based triplets

#### TRIPLETS FOR THE NEURODEGENERATIVE ASPECTS OF GLAUCOMA

#### GLAUCOMA IS A LEADING CAUSE OF IRREVERSIBLE BLINDNESS WORLDWIDE

- Many patients experience worsening of glaucoma and lose vision over time despite maximum medical therapy
- Available therapies today treat intraocular pressure, not the fundamental biology of retinal neural cell loss which is multifactorial in nature
- Our triplets technology is designed to target multiple intra- and extracellular pathways implicated in the neurobiology of glaucoma
- Durability of treatment will be key and will be enabled by ABC Platform based triplets



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## IN THEORY

Intravitreal anti-VEGF agents improve & maintain vision when dosed per label...

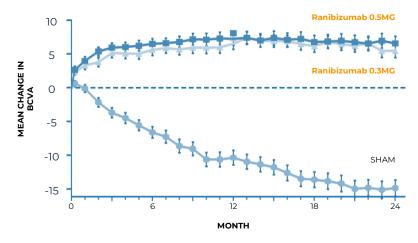
#### Recommended dosing in first year:

Ranibizumab 12 monthly

#### Aflibercept

bi-monthly after 3 monthly loading doses

#### PHASE III STUDY OF MONTHLY ANTI-VEGF 1

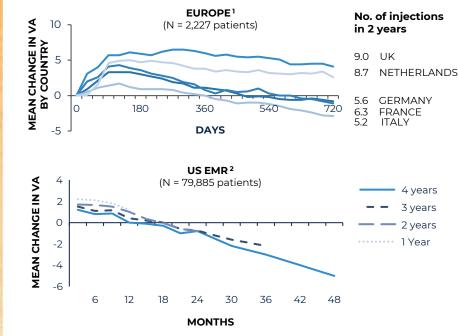


## IN PRACTICE

...yet in the real word, visual gains are minimal and not maintained.

Patients cannot be treated frequently enough and are overextended between doses in the real world. Without continuous high-intensity treatment, vision loss can begin after only 3 months of anti-VEGF therapy.

This pattern is seen globally and with all current medicines.



1. The AURA Study, adapted from Holz FG et al. Br J Ophthalmol 2015; 99 (2): 220-226.

2. Adapted from SIERRA-AMD, Khanani A, et al. Ophthal. Retina 2020 Feb; 4(2):122-123. EMR= Electronic Medical Records

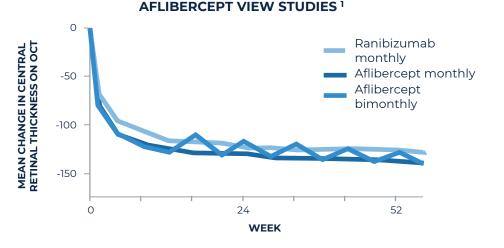
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## WHY?

Current, Generation 1.0 agents do not control disease for long enough between doses.

Undertreatment leads to disease progression and permanent retinal damage. Bimonthly anti-VEGF therapy results in disease activity between doses due to insufficient durability.



# NOW WHAT?

Today's Generation 1.0 anti-VEGF agents are not good enough.



Patients, physicians, and health systems struggle with the limitations of today's Generation 1.0 medicines.



A new class of Generation 2.0 intravitreal therapy is needed.

## What profile may be required to meaningfully change the current paradigm?

	Durability -		1	
Potential Impact	Maintenance Phase	Loading Phase	Efficacy Profile	Safety Profile
	wAMD: >50% reach Q20W		wAMD, DME, and RVO: Non-	Safety profile is in line with aflibercept and ranibizumab
5 to 6 month	<b>DME:</b> >50% reach Q20W	≤ 3 loading doses	inferior to comparator	
predominant	<b>RVO:</b> Non-inferior with Q8W	s s loading doses	<b>NPDR:</b> 2 step change and / or lower event rate	
	<b>NPDR:</b> Compelling efficacy at 2x / year			
	wAMD: >50% reach Q16W or better		wAMD, DME, and RVO: Non-	Safety profile is in line with aflibercept and ranibizumab
4 to 5 month	DME: >50% reach Q16W or better	≤ 3 loading doses	inferior to comparator	
predominant	<b>RVO:</b> Non-inferior with Q8W		<b>NPDR:</b> 2 step change and / or lower event rate	
	<b>NPDR:</b> Compelling efficacy at 3x / year			
	<b>wAMD:</b> 33% Q8W, 33% Q12W, 33% Q16 / 20W			
3 to 4 month predominant	DME: >50% better than Q12W	> 7 loading docos	wAMD, DME, and RVO: Non- inferior to comparator	Safety profile may be worse than aflibercept and ranibizumab
	<b>RVO:</b> Non-inferior with Q8W	≥ 3 loading doses	NPDR: 2 step improvement	
	<b>NPDR:</b> Compelling efficacy at 4x / year			

## KSI-301 Phase 1b data suggest a Generation 2.0 safety, efficacy and durability profile

W	'et	Α	Μ	D

72% have achieved a 6-month treatment-free interval at least once during follow-up<sup>1</sup>

Time to First Retreatment <sup>2</sup>	Percentage
At or before 2 months	8% (4/49)
3 months or longer	92% (45/49)
4 months or longer	82% (40/49)
5 months or longer	66% (27/41)
6 months	49% (20/41)

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#### **Diabetic Macular Edema**

79% have achieved a 6-month treatment-free interval at least once during follow-up<sup>1</sup>

Time to First Retreatment <sup>2</sup>	Percentage
Before 2 months	0% (0/33)
At 2 months	3% (1/33)
3 months or longer	97% (32/33)
4 months or longer	76% (25/33)
5 months or longer	70% (23/33)
6 months or longer	67% (22/33)

#### **Retinal Vein Occlusion**

81% have achieved 4-month or longer treatment-free interval at least once during follow-up<sup>1</sup>

Time to First Retreatment <sup>2</sup>	Percentage
At 1 month	6% (2/34)
2 months or longer	94% (31/33)
3 months or longer	66% (21/32)
4 months or longer	56% (18/32)

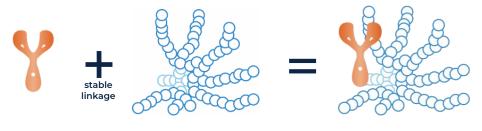
#### Safety and efficacy data in line with today's first-line medicines

1. Data from Phase 1b KSI-301 presentation at AAO 2020 Virtual Annual Meeting , through data cutoff of 15 Sept, 2020

 Time to first retreatment per protocol-specified criteria, after 3 initial monthly doses of 2.5 mg or 5 mg KSI-301. Data from Phase 1b KSI-301 presentation at ASRS 2020 Virtual Annual Meeting, through data cutoff of 09 Jun. 2020

#### ANTIBODY BIOPOLYMER CONJUGATE ABC PLATFORM

Biologics precision-engineered for increased durability and efficacy



#### ANTIBODY

laG1 with inert immune effector function

#### BIOPOLYMER

Optically clear, high molecular weight phosphorylcholine polymer CONJUGATE

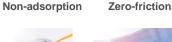
Antibody and biopolymer covalently bound via single site-specific linkage















Stereospecific

#### SAME WHERE IT MATTERS

- Clinically proven targets
- Antibody-based biologic
- Intravitreal: 25M+ injections annually
- Optically clear, no residues
- Fast and potent clinical responses

#### DIFFERENT WHERE IT COUNTS

- Designed-in ocular durability
- Designed-in rapid systemic clearance
- Improved bioavailability
- Improved biocompatibility
- Improved stability

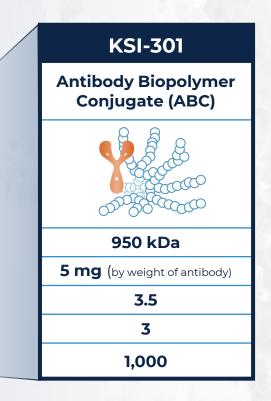
#### KSI-301: AN ANTI-VEGF ABC

## GENERATION 2.0 ANTI-VEGF

KSI-301's high molecular weight & formulation strength can provide an important dosing advantage

Drug:	RANIBIZUMAB (Lucentis)	AFLIBERCEPT (Eylea)	BEVACIZUMAB (Avastin)
Molecule type	Antibody fragment	Recombinant fusion protein	Antibody
Molecular structure	٩	8	Y
Molecular weight	48 kDa	115 kDa	149 kDa
Clinical dose	0.3-0.5 mg	2 mg	1.25 mg
Equivalent molar dose	0.5	1	0.9
Equivalent ocular PK	0.7	1	1
Equivalent ocular concentration at 3 months	0.001	1	NA <sup>1</sup>

Equivalent values are showed as fold changes relative to aflibercept. kDa= kilodalton 1. Lower affinity of bevacizumab precludes a useful comparison



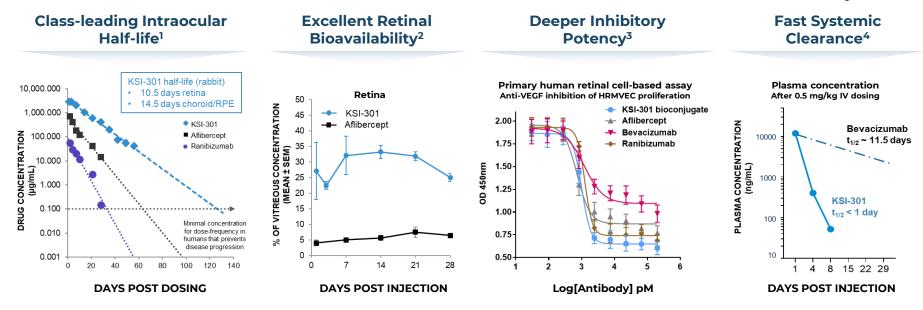
#### **KSI-301 ANTIBODY BIOPOLYMER CONJUGATE** "MORE THAN THE SUM OF ITS PARTS"





Artistic representation of KSI-301

Electron microscope image of KSI-301



1. Data from rabbit model. Ranibizumab data: Gaudreault et al (2007) IOVS 46(2) 726 Gaudreault et al (2007) Retina 27(9) 1260 Bakri et al (2007) Ophthalmol 114(12) 2179 || Aflibercept data: EVER Congress Portoroz Slovenia (2008) Struble (Covance) Koehler-Stec (Regeneron). Aflibercept data adjusted arithmetically to reflect 2,000 µg dose administered (based on rabbit in vivo dosing of 500 µg) || KSI-301 data on file, adjusted arithmetically to reflect 5,000 µg dose administered (based on rabbit in vivo dosing of 725 µg). Error bars reflects standard error of the mean

2. Covance rabbit ADME (absorption, distribution, metabolism, elimination) model: Aflibercept data (2008): EVER Congress Portoroz Slovenia Struble (Covance), Koehler-Stec (Regeneron). KSI-301 data (2017): Covance study, data on file. Error bars reflects standard error of the mean

KSI-301 data: data on file; Bevacizumab data: Yeung et al 2010 Cancer Research.

KODIAK 4. KSI-301 data: Non-human primate toxicology study, data on file; Bevacizumab data: Yeung et al 2010 Cancer Research.

## **OUR GOAL** WITH KSI-301

Develop KSI-301 as a meaningfully differentiated first-line treatment in each retinal vascular disease Better meet the individual needs of key stakeholders globally







#### We are developing KSI-301 to be first line in the 4 major retinal vascular diseases

Target enrollment exceeded Recruitment closed		<b>Now Recruiting</b> First patients randomized in GLEAM / GLIMMER and BEACON		
Wet AMD	Diabetic Macular Edema	<b>Retinal Vein Occlusion</b>	Non-Proliferative Diabetic Retinopathy	
DAZZLE Study (n~550)	GLEAM and GLIMMER Studies (n~450 each)	BEACON Study (n~550)	GLOW Study (n~440)	
KSI-301 <b>once every 3, 4 or 5 months</b> after 3 monthly doses	once every 3, 4 or 5 months once every 2 to 6 months		KSI-301 once every 3, 4 or 6 months	
Comparator	Comparator	Comparator	Comparator	
Aflibercept <b>Once every 2 months</b> after 3 monthly doses	Aflibercept Once every 2 months after 5 monthly doses	Aflibercept Once every month	Sham	

KSI-301 pivotal studies enroll treatment-naïve patients and incorporate key learnings from our Phase 1b study, supporting a high level of confidence in our KSI-301 development program

#### **KSI-301 COMMERCIAL MANUFACTURING**

#### BUILDING CAPACITY TO SUPPLY RAPID MARKET UPTAKE

Expected Year 1 manufacturing capacity to supply 2.5M+ doses with the ability to flex up to 15M+ doses

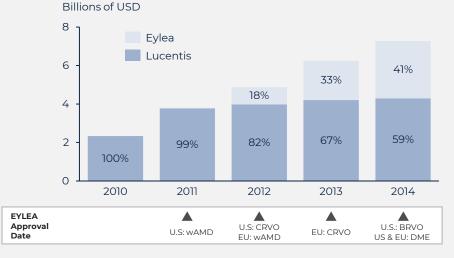
Integrated global pharmaceutical supply chain

Purpose-built Lonza IBEX Dedicate bioconjugation facility to support commercial launch

#### Case study on market adoption

Worldwide anti-VEGF revenue

Can Eylea market share growth educate KSI-301 adoption?



Kodiak aims to submit a single BLA for KSI-301 in wet AMD, DME and RVO in calendar year 2022

Company financial disclosures and product labeling

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## **OUR 2022 VISION**

#### **RETINAL VEIN OCCLUSION**

2022 BEACON Phase 3 top-line data 2022 BLA filing

#### WET AMD

2022 DAZZLE Phase 2b/3 top-line data 2022 BLA filing

#### DIABETIC MACULAR EDEMA

2022 GLEAM / GLIMMER Phase 3 top-line data 2022 BLA filing

2022 THE OPHTHALMOLOGY

MEDICINES COMPANY

#### KSI-501 anti-VEGF/IL-6

2021 IND submitted 2022 Phase la/lb data

#### DIABETIC RETINOPATHY

2023 GLOW Phase 3 top-line data





#### KSI-601 Triplet Inhibitor for dry AMD

2022 IND submitted

Indications submitted in 3 BLA (wAMD, DME and RVO)



**Clinical molecules** 

IND per year beginning 2021

## MILESTONES AND KSI-301 DEVELOPMENT ACCELERATION

#### 2019

#### KSI-301

- Safety, efficacy, durability proof-of-concept established
- Initiation of DAZZLE wAMD pivotal study
- FDA EOP2 meeting
- ✓ \$225MM royalty financing
- ✓ \$317MM equity financing

#### KSI-301

Achieved

 Additional readouts of Phase 1b data

2020

- Maturation of data support pivotal clinical studies
- Manufacturing framework to supply millions of doses in first year of launch
- Initiate two DME Phase 3 trials (GLEAM & GLIMMER)
- Initiate RVO Phase 3 trial (BEACON)
- Complete enrollment in wAMD (DAZZLE)

#### 2021

#### KSI-301

- Initiate NPDR Phase 3 trial (GLOW)
- Presentation of one-year Phase 1b results in wet AMD, DME and RVO
  - Complete enrollment in DME (GLEAM & GLIMMER) and RVO (BEACON) studies
- DAZZLE wet AMD last patient last visit
- KSI-501 (bispecific ABC)
- Submit IND

#### 2022

#### KSI-301

- DAZZLE wAMD pivotal study top-line readout
- RVO pivotal study (BEACON) top-line readout
- DME pivotal studies (GLEAM & GLIMMER) topline readouts
- Submit BLA for wAMD, DME and RVO

#### KSI-501

- Phase 1/2 data in inflammatory retinal diseases
- KSI-601 (triplet ABC) for dry AMD
- Submit IND

#### 2023

#### KSI-301

Potential regulatory approval for wAMD, DME and RVO in US and EU

- Potential commercial launch for wAMD, DME, RVO in US
- DR pivotal study (GLOW) readout
- Submit sBLA for DR pivotal study (GLOW)
- KSI-501
- Additional readouts of Phase 1/2 data
- KSI-601
- Initiate Phase 1/2 study

Potential Milestones 2021 - 23

## **KSI-301 Accelerated Development Strategy**

4 Pivotal Studies to support BLA with All 3 Major Anti-VEGF Indications Run Concurrently

	2019	2020	2021	202	2	2023
Phase 1b Ongoing	121 treatment-naïve wAMD, DMI Safety, efficacy, durability - 18 n					
DAZZLE Pivotal wAMD Target Enrollment Exceeded	012W	treatment naïve patients /-Q20W KSI-301 vs Q8W Eylea	12-month endpoint			
GLEAM DME Phase 3 First Patients Randomized			atment naïve pts. 4W KSI-301 Eylea	12-month endpoint	Single BLA	U.S. commercial launch
GLIMMER DME Phase 3 First Patients Randomized			atment naïve pts. 4W KSI-301 Eylea	12-month endpoint	2022	
BEACON RVO Phase 3 First Patients Randomized		or CRVO	atment naïve BRVO patients -301 vs Q4W Eylea	6-month endpoint		
GLOW DR without DME Phase 3 In Planning			~440 patients Q16W-Q24W KSI-301 v	vs Sham	2-month endpoir	nt <b>BLA 2023</b>

KODIAK BLA: biologics license application; RVO: retinal vein occlusion; BRVO: branch RVO; CRVO: central RVO; wAMD: wet age-related macular degeneration; DME: diabetic macular edema; DR: diabetic retinopathy

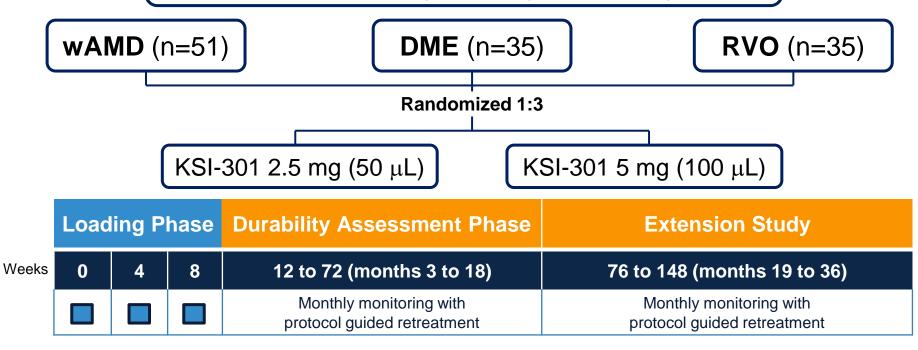
## **KSI-301 CLINICAL DATA**

121 patients dosed in Phase 1b study

## KSI-301 Phase 1b

## insight into durability among treatment naïve subjects

Randomized, open label study to evaluate multidose safety, efficacy & durability



## KSI-301 Phase 1b Retreatment Criteria prespecified by disease state

#### wAMD

- − Increase in CST ≥75  $\mu$ m with a decrease in BCVA of ≥ 5 letters compared to Week 12, OR
- Decrease in BCVA of > 5 letters compared to Day 1, due to worsening wAMD activity, OR
- Decrease in BCVA of ≥ 10 letters compared to the best prior BCVA, due to worsening wAMD activity, OR
- 6 months have elapsed since the last retreatment

#### DME and RVO

- − Increase in CST ≥75 µm with a decrease in BCVA of ≥ 5 letters compared to Week 12 or the prior visit, OR
- Decrease in BCVA of ≥ 10 letters compared to the best prior BCVA, due to worsening DME/RVO disease activity

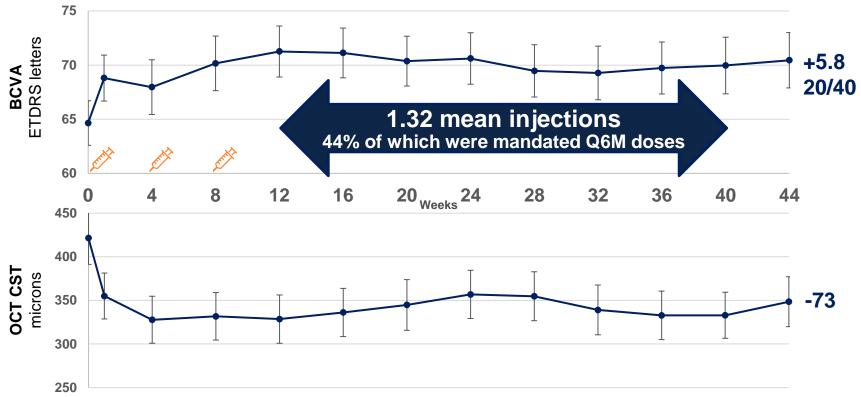
# For all subjects, investigators can retreat at their discretion if significant disease activity is present that does not meet the above criteria

## **KSI-301** Phase 1b Baseline Characteristics

Variable	wAMD Cohort (n=51)	DME Cohort (n=35)	RVO Cohort (n=35)
Age, mean (SD), years	77.9 (10.5)	59.7 (11.7)	63.6 (12.6)
Gender, n (%), female	32 (62.7)	14 (40.0)	13 (37.1)
Race, n (%), White	48 (94.1)	28 (80.0)	31 (88.6)
BCVA, mean (SD), ETDRS letters	63.3 (13.3)	66.8 (10.2)	54.9 (15.4)
Snellen equivalent	~20/50	~20/50	20/80
BCVA, Snellen 20/40 or better, n (%)	20 (39.2)	16 (45.7)	6 (17.1)
OCT CST, mean (SD), microns	430 (162)	453 (110)	675 (237)

## WET AMD

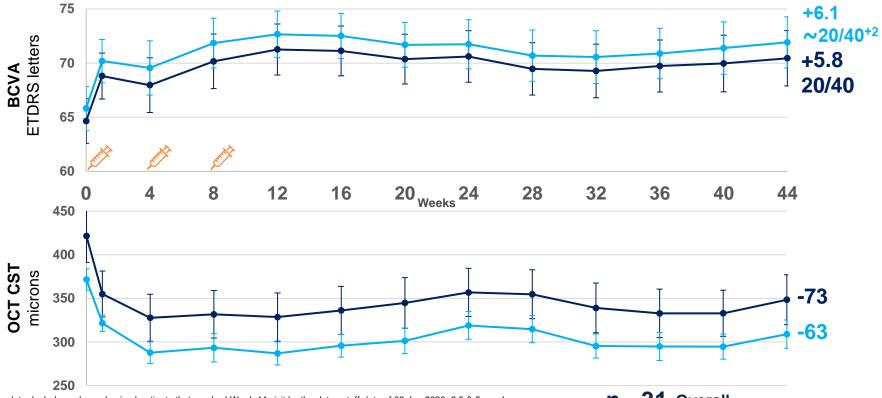
# Efficacy of KSI-301 in Wet AMD change from baseline to week 44 in mean BCVA & OCT



Interim data. Includes only randomized patients that reached Week 44 visit by the data cutoff date of 09 Jun 2020; 2.5 & 5 mg doses pooled. Observed data. Error bars represent standard error of the mean. OCT CST values are site reported and include PED height. BCVA= best corrected visual acuity; OCT= optical coherence tomography; CST= central subfield thickness. Mean injections reflect the average number of injections received per patient between Week 12 and 40 (aflibercept per label mean number of injections 4.0).

**n= 31** Patients reaching Week 44 visit by data cutoff

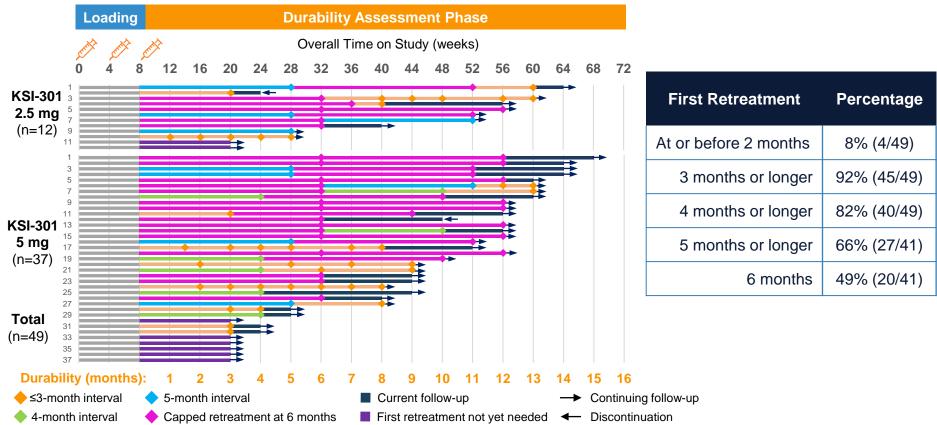
# Efficacy of KSI-301 in Wet AMD in 27/31 subjects without high PEDs



Interim data. Includes only randomized patients that reached Week 44 visit by the data cutoff date of 09 Jun 2020; 2.5 & 5 mg doses pooled. Observed data. Error bars represent standard error of the mean. OCT CST values are site reported and include PED height. High PED defined as presence of a PED with baseline CST ≥500 microns. BCVA= best corrected visual acuity; OCT= optical coherence tomography; CST= central subfield thickness.

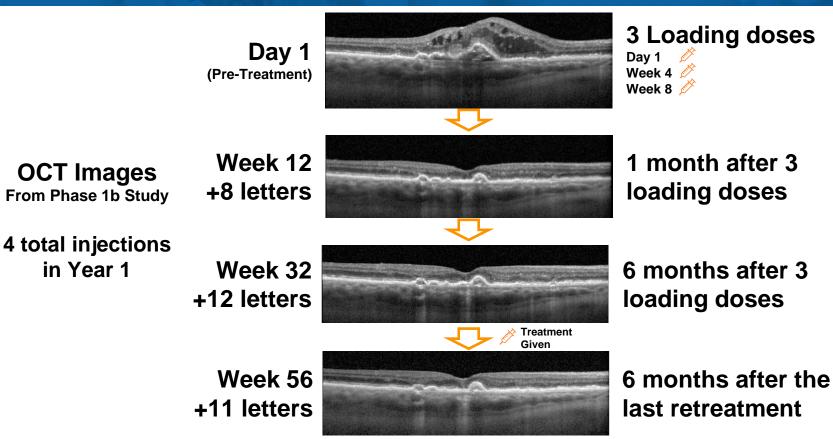
n= 31 Overall n= 27 Without high PEDs

## KSI-301 in wAMD: Durability Assessment Data support 3- to 6-month durability



Interim data. Includes only randomized patients that reached the first retreatment opportunity (Week 12 visit) by the data cutoff date of 09 Jun 2020. Each bar represents an individual patient.

## Case Example: 6-Month Dosing Through 1 Year KSI-301 in wet AMD



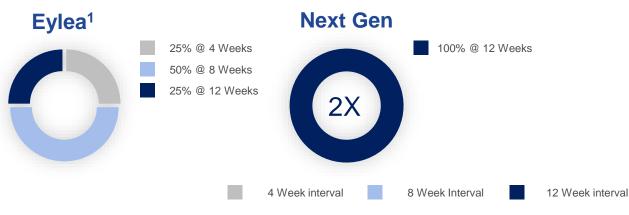
## KSI-301 in wAMD: Maturing dataset is robust and consistent over time

	Angiogenesis Meeting (21 January 2020 cutoff)	ASRS Meeting (09 June 2020 cutoff)
Patient-years Clinical Experience	22.0	41.7
Efficacy Analyses (functional and anatomical)	Week 24 (n=31)	Week 44 (n=31)
Mean change in BCVA	5.9 letters	5.8 letters
Mean change in OCT CST	-58 microns	-73 microns
Mean number injections since week 12	0.16	1.32
Durability Analyses (time to first retreatment)	n=35	n=49
At or before 2 months	<b>9%</b> (3/35)	<b>8%</b> (4/49)
3 months or longer	<b>91%</b> (32/35)	<b>92%</b> (45/49)
4 months or longer	<b>84%</b> (27/32)	<b>82%</b> (40/49)
5 months or longer	<b>72%</b> (21/29)	<b>66%</b> (27/41)
6 months	<b>55%</b> (16/29)	<b>49%</b> (20/41)

31

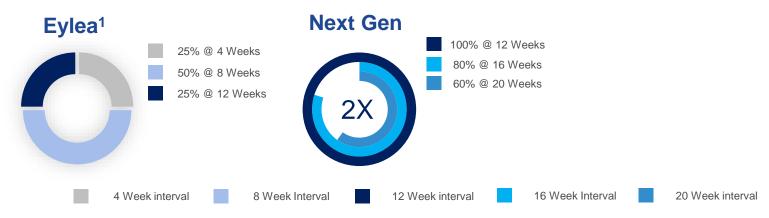
# A next generation biologic should bring nearly all patients to a 12-week interval

	Maintenance Phase					
	4 Weeks 8 Weeks 12 Weeks 16 Weeks 20 Weeks					
Lucentis						
Eylea						
Next Gen						



## A biologic bringing nearly all patients to 12 weeks *and* a majority to 4- and 5- months would be disruptive

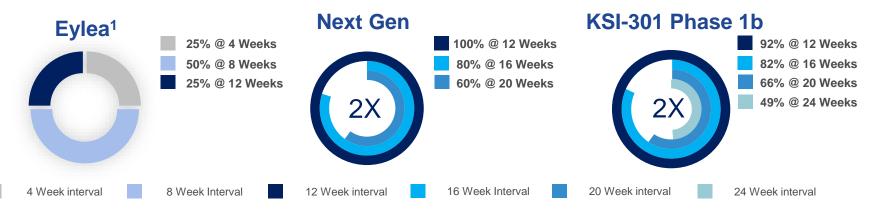
	Maintenance Phase					
	4 Weeks	8 Weeks	12 Weeks	16 Weeks	20 Weeks	
Lucentis						
Eylea			1111111			
Next Gen		.				



1. According to current clinical practice

## Benchmarking: KSI-301 Phase 1b wAMD data KSI-301 time to first retreatment data confirm the potential to be disruptive

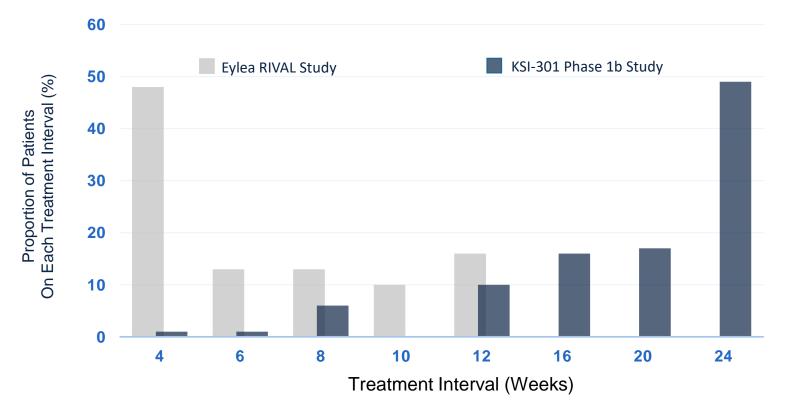
	Maintenance Phase					
	4 Weeks	8 Weeks	12 Weeks	16 Weeks	20 Weeks	24 Weeks
Lucentis						
Eylea						
Next Gen						
KSI-301 <sup>2</sup>		Ň				



1. According to current clinical practice

2. Phase 1b data based on the time to first retreatment

#### Benchmarking in treatment-naïve wAMD: KSI-301 Phase 1b "Generation 2.0" durability compared to Eylea long-interval RCT data

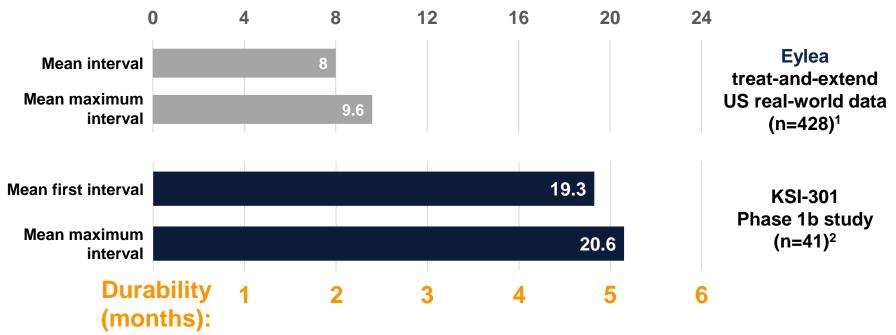


1. Gillies MC, et al. Effect of Ranibizumab and Aflibercept on Best-Corrected Visual Acuity in Treat-and-Extend for Neovascular Age-Related Macular Degeneration: A Randomized Clinical Trial. JAMA Ophthalmol. 2019;137(4):372–379. doi:10.1001/jamaophthalmol.2018.6776

2. For KSI-301: Includes randomized patients that reached the first retreatment opportunity (Week 12 visit) by the data cutoff date of 09 Jun 2020.

## Benchmarking: KSI-301 Phase 1b wAMD data "Generation 2.0" durability compared to Eylea real-world data

#### Mean treatment intervals after the loading phase (weeks)



1. Singer MA, et al. Two-Year Real-World Treat and Extend Patterns and Fluid Outcomes Among Neovascular Age-Related Macular Degeneration Patients Treated With Anti-VEGFs. ASRS 2020 virtual meeting. Available at asrs.org. 2. Includes all randomized patients that received all three loading doses and a first retreatment by the data cutoff date of 09 Jun 2020. For Eylea data set, mean interval is the average interval per patient over two years, and mean maximum interval is the average of the longest interval achieved per patient at any point during follow-up. For KSI-301 data set, first interval refers to the first retreatment, and mean maximum interval is the average of the longest interval per patient at any point during follow-up.

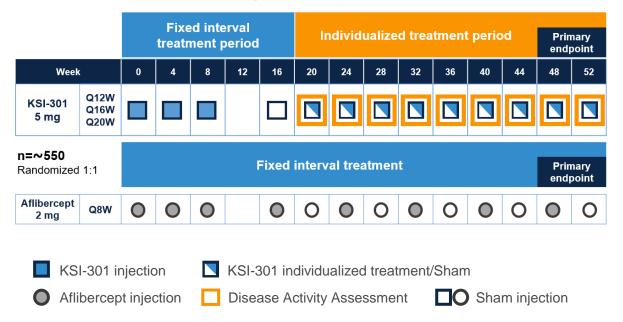
### KSI-301 Phase 2b/3 wAMD DAZZLE Study Dosing with KSI-301 as infrequently as every 20 weeks\*

#### Wet AMD – Phase 1b

First Retreatment	Percentage (n=49)
At or before 2 months	8%
3 months or longer	92%
4 months or longer	82%
5 months or longer	66%
6 months	49%

72% have achieved a 6-month treatment interval at least once during follow-up<sup>1</sup>

## DAZZLE pivotal study evaluates individualized dosing of every 12, 16 or 20 weeks



\*After the loading phase. Clinicaltrials.gov ID NCT04049266, currently in late stages of recruitment 1. As of 15 Sep 2020

## How do DAZZLE Study Disease Activity Assessment Criteria Compare to Phase 1b?

Parameters	Phase 1b Study <sup>1</sup>	DAZZLE study <sup>2</sup>	Change
Visual <i>and</i> anatomical	Increase in CST ≥75 µm with a decrease in BCVA of ≥ 5 letters compared to Week 12, <i>OR</i>	Increase in CST ≥50 µm with a decrease in BCVA of ≥ 5 letters compared to Week 12, <i>OR</i>	Tighter CST control (25 microns)
Visual only	Decrease in BCVA of ≥ 10 letters compared to the best prior BCVA, due to worsening wAMD activity, <i>OR</i>	Decrease in BCVA of ≥ 10 letters compared to the best prior BCVA, due to worsening wAMD activity, <i>OR</i>	No change
	Decrease in BCVA of > 5 letters compared to Day 1, due to worsening wAMD activity	N/A	Eliminated to reduce subjectivity and unnecessary retreatments
Anatomical only	N/A	Increase of ≥ 75 microns compared to Week 12, <i>OR</i>	Added two anatomical-only
	N/A	New Macular Hemorrhage	criteria

wAMD = wet age-related macular degeneration; CST = central subfield retinal thickness; BCVA = best corrected visual acuity.

<sup>1</sup> Clinicaltrials.gov ID: NCT03790852

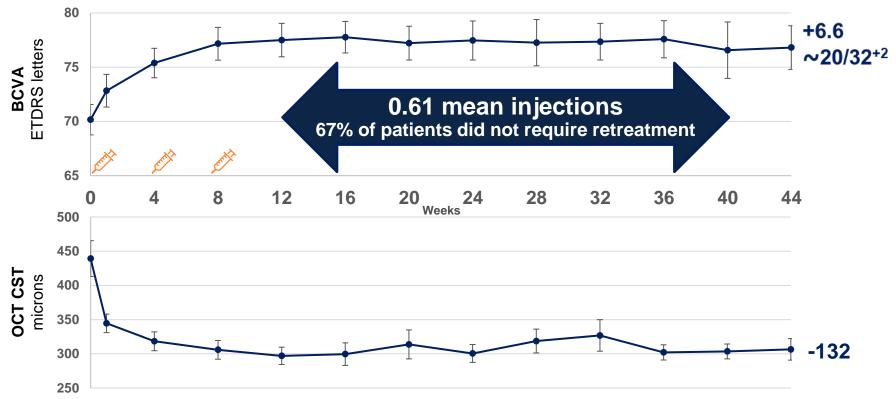
<sup>2</sup> Clinicaltrials.gov ID NCT04049266

## **DAZZLE protocol optimization**

- Building from the exploratory Phase 1b, DAZZLE maintains consistency of key features while further optimizing protocol design
  - 1. Similar patient population treatment naïve wAMD (~80% from USA)
  - 2. Tighter dosing interval ranging from Q4W-Q24W to Q12W-Q20W
  - Tighter disease control tighter disease activity assessments to determine patients' dosing intervals
  - 4. Decreased subjectivity no physician discretion treatment (IRT driven)
  - 5. High statistical power for non-inferiority (>90%)

## DIABETIC EYE DISEASE DME & DR

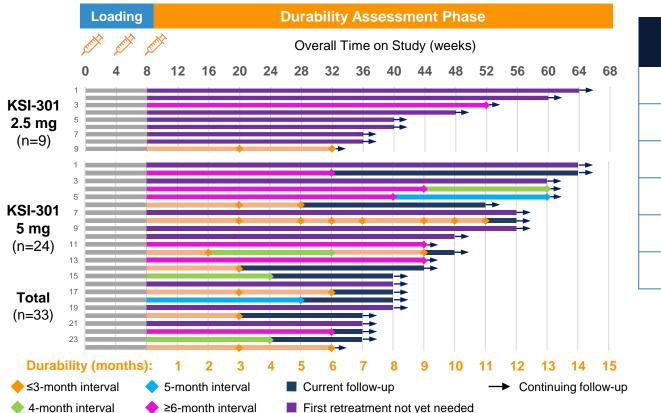
# Efficacy of KSI-301 in DME change from baseline to week 44 in mean BCVA & OCT



Interim data. Includes only randomized patients that reached Week 44 visit by the data cutoff date of 09 Jun 2020; 2.5 & 5 mg doses pooled. Observed data. Error bars represent standard error of the mean. OCT CST values are site reported. BCVA= best corrected visual acuity; OCT= optical coherence tomography; CST= central subfield thickness. Mean injections reflect the average number of injections received per patient between Week 12 and 40 (aflibercept per label mean number of injections 5.0).

**1= 18** Patients reaching Week 44 visit by data cutoff

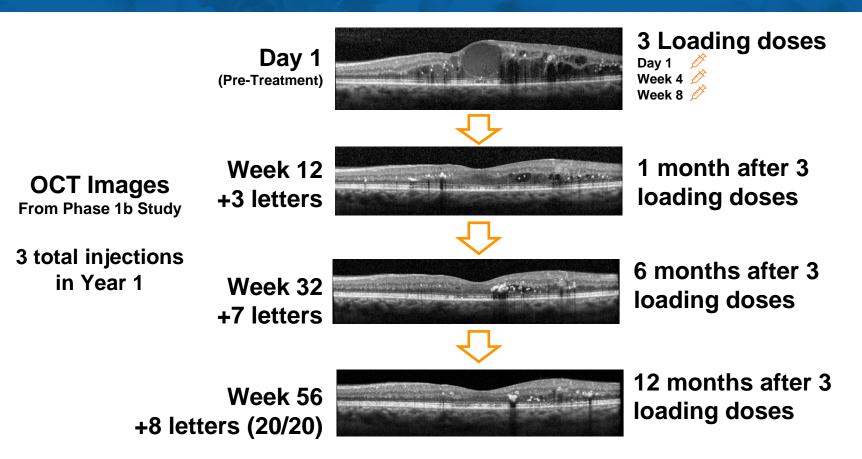
# KSI-301 in DME: 3 loading doses can provide sustained disease control of 3 to 6+ months



Interim data. Includes only randomized patients that reached the first retreatment opportunity (Week 12 visit) by the data cutoff date of 09 Jun 2020. Each bar represents an individual patient. All depicted patients continue to be followed (no discontinuations)

First Retreatment	Percentage
Before 2 months	0% (0/33)
At 2 months	3% (1/33)
3 months or longer	97% (32/33)
4 months or longer	76% (25/33)
5 months or longer	70% (23/33)
6 months or longer	67% (22/33)

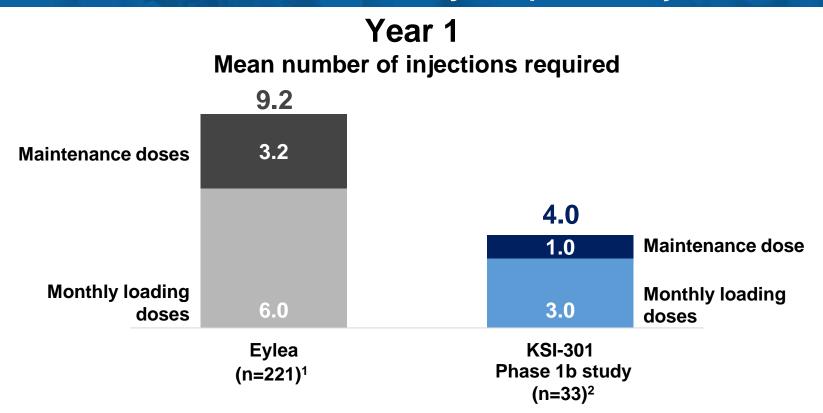
## Case Example: No Retreatments for 12 Months After Loading Phase KSI-301 in DME



# KSI-301 in DME: Maturing dataset is robust and consistent over time

	Angiogenesis Meeting (21 January 2020 cutoff)	ASRS Meeting (09 June 2020 cutoff)
Patient-years Clinical Experience	16.8	29.8
Efficacy Analyses (functional and anatomical)	Week 24 (n=19)	Week 44 (n=18)
Mean change in BCVA	6.8 letters	6.6 letters
Mean change in OCT CST	-133 microns	-132 microns
Mean number injections since week 12	0.21	0.61
Durability Analyses (time to first retreatment)	n=33	n=33
At 2 months	<b>3%</b> (1/32)	<b>3%</b> (1/33)
3 months or longer	<b>97%</b> (31/32)	<b>97%</b> (32/33)
4 months or longer	<b>76%</b> (16/21)	<b>76%</b> (25/33)
5 months or longer	<b>68%</b> (11/16)	<b>70%</b> (23/33)
6 months or longer	<b>64%</b> (9/14)	<b>67%</b> (22/33)

#### Benchmarking: KSI-301 Phase 1b DME data "Generation 2.0" durability compared to Eylea



1. Wells JA. Aflibercept, bevacizumab, or ranibizumab for diabetic macular edema (DRCR Protocol T). N Engl J Med. 2015 Mar 26;372(13):1193-203 (supplemental data).

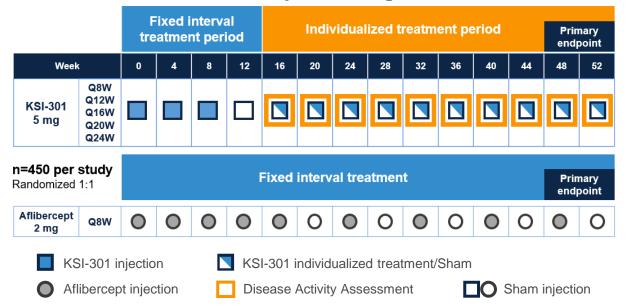
2. Interim data. Annualized injections based on the current monthly injection rate of all DME patients as of the 09 Jun 2020 data cutoff.

#### KSI-301 Phase 3 DME GLEAM and GLIMMER Studies Dosing with KSI-301 as infrequently as every 24 weeks\*

#### DME – Phase 1b

First Retreatment	Percentage (n= 33)
At 2 months	3%
3 months or longer	97%
4 months or longer	76%
5 months or longer	70%
6 months or longer	67%

79% have achieved a ≥6month treatment interval at least once during follow-up<sup>1</sup> GLEAM-GLIMMER pivotal studies evaluate individualized dosing of every 8, 12, 16, 20 or 24 weeks, after only 3 loading doses



### How do GLEAM/GLIMMER Studies Disease Activity Assessment Criteria Compare to Phase 1b?

Parameters	Phase 1b Study <sup>1</sup>	GLEAM/GLIMMER Studies	Change	
Visual <i>and</i> anatomical	Increase in CST $\geq$ 75 µm with a decrease in BCVA of $\geq$ 5 letters compared to Week 12 or the prior visit, <i>OR</i>	Increase in OCT CST $\geq$ 50 µm <u>compared to</u> <u>lowest previous measurement</u> and a decrease in BCVA of $\geq$ 5 letters <u>compared to</u> <u>the average of the 2 best previous BCVA</u> <u>assessments</u> , due to worsening of DME disease activity, <i>or</i>	Tighter and dynamic control of both vision and anatomy	
Visual only	Decrease in BCVA of ≥ 10 letters compared to the best prior BCVA, due to worsening DME activity	N/A	Eliminated to reduce subjectivity and unnecessary retreatments	
Anatomical only	N/A	Increase in OCT CST ≥ 75 µm compared to lowest previous measurement due to worsening of DME disease activity; <i>or</i>	Added two anatomical-	
Only	N/A	New or worsening proliferative DR (PDR)	only criteria	

DME = diabetic macular edema; OCT = optical coherence tomography; CST = central subfield retinal thickness; BCVA = best corrected visual acuity.

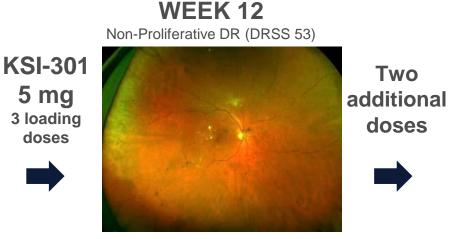
<sup>1</sup> Clinicaltrials.gov ID: NCT03790852

## **GLEAM/GLIMMER** Phase 3 protocol optimization

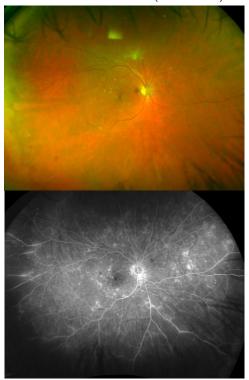
- Building from the exploratory Phase 1b, GLEAM/GLIMMER maintain consistency of key features while further optimizing protocol designs
  - 1. Similar patient population treatment naïve DME (~80% from USA)
  - 2. Tighter dosing interval ranging from open to Q8W-Q24W
  - 3. Tighter disease control tighter disease activity assessments to patients' determine dosing intervals
  - 4. Decreased subjectivity no physician discretion treatment (IRT driven)
  - 5. High statistical power for non-inferiority (>90%)

# The sustained disease control of only 3 loading doses of KSI-301 is also seen in proliferative diabetic retinopathy

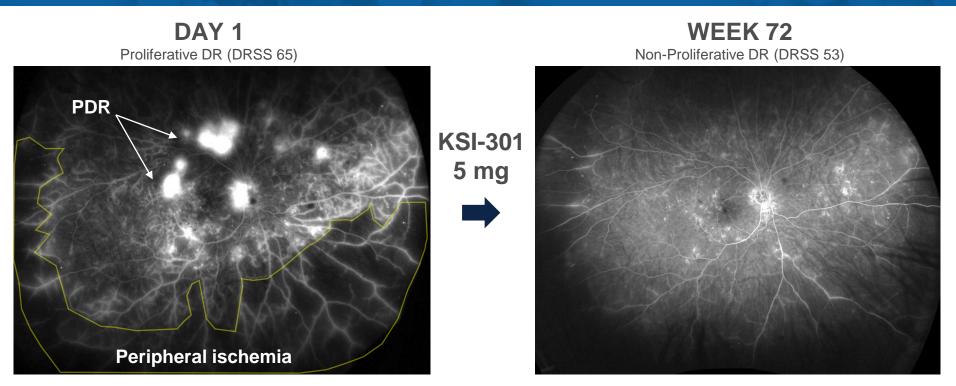
# DAY 1 Proliferative DR (DRSS 65)



Regression from PDR to NPDR Fast and substantial (2-step) improvement, sustained for 18 months with only 2 additional doses (26-week mean retreatment interval) WEEK 72 Non-Proliferative DR (DRSS 53)



In addition to the regression from PDR to NPDR, this patient exhibits signs of peripheral vascular reperfusion

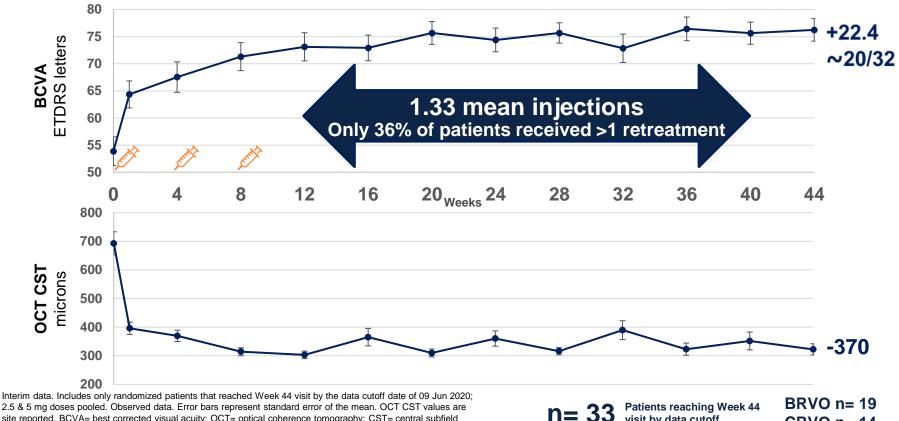


Sustained signs of disease modification for 18 months with only 2 additional doses (26-week mean treatment interval)

PDR= Proliferative DR; NPDR= Non-Proliferative DR; DRSS = DR Severity Scale; DRSS 53 = Severe NPDR; DRSS 65 = Moderate PDR

## RETINAL VEIN OCCLUSION

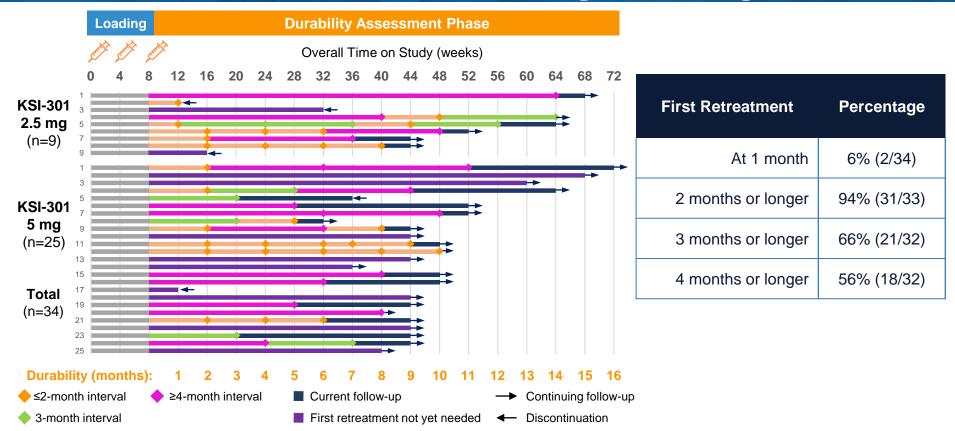
### Efficacy of KSI-301 in RVO change from baseline to week 44 in mean BCVA & OCT



2.5 & 5 mg doses pooled. Observed data. Error bars represent standard error of the mean. OCT CST values are site reported. BCVA= best corrected visual acuity; OCT= optical coherence tomography; CST= central subfield thickness. Mean injections reflect the average number of injections received per patient between Week 12 and 40 (aflibercept per label mean number of injections 8.0).

Patients reaching Week 44 visit by data cutoff **CRVO n= 14** 

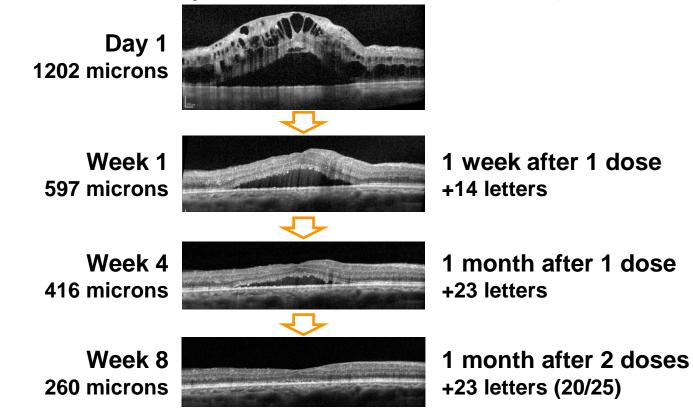
# KSI-301 in RVO: 3 loading doses show potential for 2 to 4 month or longer dosing



Interim data. Includes only randomized patients that reached the first retreatment opportunity (Week 12 visit) by the data cutoff date of 09 Jun 2020. Each bar represents an individual patient.

# Is it possible to control the most severe CRVO cases with only 2 loading doses?

Case Example of KSI-301 in the Phase 1b Study

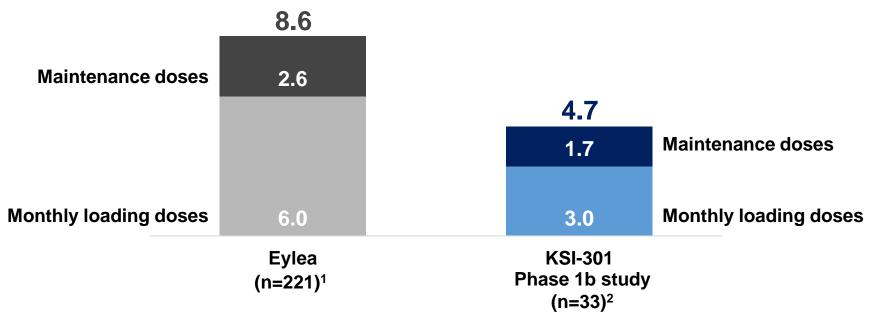


### KSI-301 in RVO: Maturing dataset is more robust and consistent over time

	Angiogenesis Meeting (21 January 2020 cutoff)	ASRS Meeting (09 June 2020 cutoff)
Patient-years Clinical Experience	18.8	29.8
Efficacy Analyses (functional and anatomical)	Week 24 (n=30)	Week 44 (n=33)
Mean change in BCVA	22.2 letters	22.4 letters
Mean change in OCT CST	-350 microns	-370 microns
Mean number of injections since week 12	0.46	1.33
Durability Analyses (first retreatment)	n=33	n=34
At 1 month	<b>6%</b> (2/33)	<b>6%</b> (2/34)
2 months or longer	<b>94%</b> (30/32)	<b>94%</b> (31/33)
3 months or longer	<b>64%</b> (20/31)	<b>66%</b> (21/32)
4 months or longer	<b>53%</b> (16/30)	<b>56%</b> (18/32)

#### Benchmarking: KSI-301 Phase 1b RVO data "Generation 2.0" durability compared to Eylea

#### Year 1 Mean number of injections required



 Injections averaged between the two pivotal aflibercept trials; n represents the total randomized in the aflibercept groups in both studies. Brown DM. Intravitreal Aflibercept Injection for Macular Edema Secondary to Central Retinal Vein Occlusion: 1-Year Results From the Phase 3 COPERNICUS Study. Am J Ophthalmol 2013;155:429–437.Korobelnik JF, et al. Intravitreal Aflibercept Injection for Macular Edema Resulting from Central Retinal Vein Occlusion. Ophthalmology 2014;121:202-208

2. Interim data. Annualized injections based on the current monthly injection rate of all RVO patients as of the 09 Jun 2020.

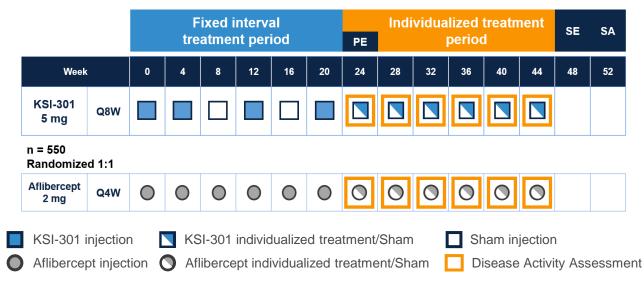
#### KSI-301 Phase 3 RVO BEACON Study Two loading doses with KSI-301 + every 8 weeks

First Retreatment	Percentage (n= 34)	
At 1 month	6%	
2 months or longer	94%	
3 months or longer	66%	
4 months or longer	56%	

RVO - Phase 1h

81% have achieved a 4-month or longer treatment interval at least once during follow-up<sup>1</sup>

## BEACON pivotal study evaluates two loading doses and every 8-week dosing, followed by individualized dosing



#### How do BEACON Study Disease Activity Assessment Criteria Compare to Phase 1b?

Parameters	Phase 1b Study <sup>1</sup>	BEACON Study <sup>2</sup>	Change
Visual <i>and</i> anatomical	Increase in CST $\geq$ 75 µm with a decrease in BCVA of $\geq$ 5 letters compared to Week 12 or the prior visit, <i>OR</i>	Increase in OCT CST $\geq$ 50 µm <u>compared to</u> <u>lowest previous measurement</u> and a decrease in BCVA of $\geq$ 5 letters <u>compared to</u> <u>the average of the 2 best previous BCVA</u> <u>assessments</u> , due to worsening of RVO disease activity, <i>or</i>	Tighter and dynamic control of both vision and anatomy
Visual only	Decrease in BCVA of ≥ 10 letters compared to the best prior BCVA, due to worsening RVO activity	N/A	Eliminated to reduce subjectivity and unnecessary retreatments
Anatomical only	N/A	Increase in OCT CST ≥ 75 µm compared to lowest previous measurement due to worsening of RVO disease activity; <i>or</i>	Added one anatomical- only criteria

RVO = retinal vein occlusion; OCT = optical coherence tomography; CST = central subfield retinal thickness; BCVA = best corrected visual acuity.

<sup>1</sup> Clinicaltrials.gov ID: NCT03790852

<sup>2</sup> Clinicaltrials.gov ID: NCT04592419

#### **BEACON Phase 3 protocol optimization**

- Building from the exploratory Phase 1b, BEACON maintains consistency of key features while further optimizing study protocol
  - 1. Similar patient population treatment naïve RVO (~80% from USA)
  - 2. Tighter dosing from open to fixed q2-month dosing, through 6-month primary endpoint
  - 3. Tighter disease control tighter disease activity assessments to determine dosing interval, in second 6 months of study
  - 4. Decreased subjectivity no physician discretion treatment (IRT driven)
  - 5. High statistical power (>90%)

## SAFETY

#### Multiple-dose safety of KSI-301 Phase 1a/1b program



- Most AEs were assessed as mild and are consistent with profile of intravitreal anti-VEGFs
- To date, 38 SAEs have been reported in 20 subjects none drug related
- Two ocular SAEs in the study eye, not drug related, both resolved
  - Worsening DME secondary to systemic fluid overload
  - Worsening cataract in a diabetic patient
- Only two AEs of intraocular inflammation, both trace to 1+ vitreous cells, with complete resolution
  - Rate of 0.30% (2/657 injections)
  - No vasculitis or retinitis in either patient

## Conclusion

#### KODIAK SCIENCES

#### WHERE WE ARE TODAY

4 PIVOTAL TRIALS

**3** INDICATIONS

### **SINGLE** BLA FILING EXPECTED IN 2022

#### KODIAK

#### KSI-301 CLINICAL EXPERIENCE

Clinical data from 1,500 injections in 400+ patients representing 250+ patient-years of exposure in representative populations in wAMD, DME and RVO

- Safety: Tracking with current standard of care (Lucentis, Eylea)
- Efficacy: Strong and appropriate impact on vision & retinal anatomy in each indication studied
- Durability: Majority of patients going 6-months or longer in wet AMD and DME

#### OPTIMIZED PIVOTAL STUDY PROGRAM

Objective to show disruptive durability with same safety and efficacy as Eylea

DAZZLE wet AMD study enrollment complete; BEACON RVO study and GLEAM / GLIMMER DME now enrolling – Data from all studies expected in 2022

Pivotal studies designed from phase 1b data with tighter criteria for disease activity assessments, shorter durability intervals, high statistical power, maintaining similar (80%+) U.S. treatment naïve population

#### OPERATING WITH CONVICTION

On track for a single BLA filing in the key indications of wAMD, DME, RVO treatment and with NPDR (prevention) indication in a supplemental BLA

Manufacturing investments aligned to clinical opportunity with commercial supply goal of 2.5M+ Prefilled Syringes in Year 1 of launch

Developing bispecific and triplet ABC Medicines for multi-mechanism diseases, including dry AMD and glaucoma

#### POISED COMMERCIAL OPPORTUNITY

Competitive landscape is clearing with competing molecules/technologies demonstrating poor benefit risk profiles

We believe KSI-301 may be able to capture market share from standard of care agents, future biosimilars, and competing late-stage molecules in development



# KODIAK

THE OPHTHALMOLOGY MEDICINES COMPANY