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KODIAK

THE OPHTHALMOLOGY MEDICINES COMPANY

APRIL 2019

SPECIAL NOTE REGARDING

FORWARD-LOOKING STATEMENTS

These slides and the accompanying oral presentation 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 initiation, timing, progress and results of our preclinical and clinical studies and our research and development programs, our ability to advance product candidates into, and successfully complete, clinical studies, and the timing or likelihood of regulatory filings and approvals 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 forward-looking 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 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.



OUR MISSION

The Ophthalmology
Medicines Company

1 HIGH SCIENCE INSIDE

Disruptive products and platforms to tackle the biggest challenges in ophthalmology.

2 “GO-TO” MEDICINES

Develop KSI-301 to be the anti-VEGF of choice for all patients with retinal vascular disease.

A pipeline directed to the major retina indications of wet AMD, dry AMD, and diabetic eye disease.

3 GLOBAL FRANCHISE

A singular focus in ophthalmology with a global scope.

From discovery through commercialization.

STRONG

COMPANY LEADERSHIP

Management Team with Experience to Build a Leading Biotech Company

BOARD OF DIRECTORS

Deep Biotech and Governance Experience

Victor Perlroth MD
Chairman

KODIAK

Felix J Baker PhD
Director

Baker Brother
Advisors LP

Bassil I. Dahivat PhD
Director

xencor

Richard S. Levy MD
Director

Robert A. Profusek
Director

**JONES
DAY**

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Founder
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Avidia MPM GUZIK



John Borgeson

Chief Financial Officer

Pfizer venBio LABRYS BIOLOGICS



Jason Ehrlich MD, PhD

Chief Medical Officer
Chief Development Officer

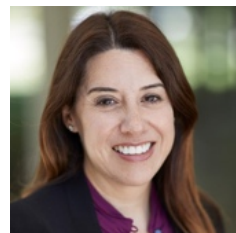
Roche Genentech



Hong Liang PhD

SVP Discovery Medicine

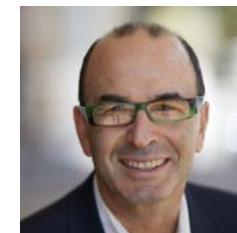
Pfizer RINAT Deltagen



Desiree Beutelspacher

VP, Clinical Operations

OPHTHOTEC eyetech



Joel Naor MD

VP, Clinical Science &
Development Operations

Allergan MACUSIGHT
STEMCELLS QLT Inc.



Almas Qudrat MSc

VP, Quality Operations

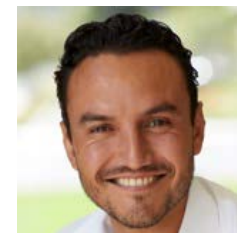
Roche Allergan NEKTAR elan



Stephen Raillard PhD

VP, Chemical Dev. & Mfg

XenoPort AFFYMAX



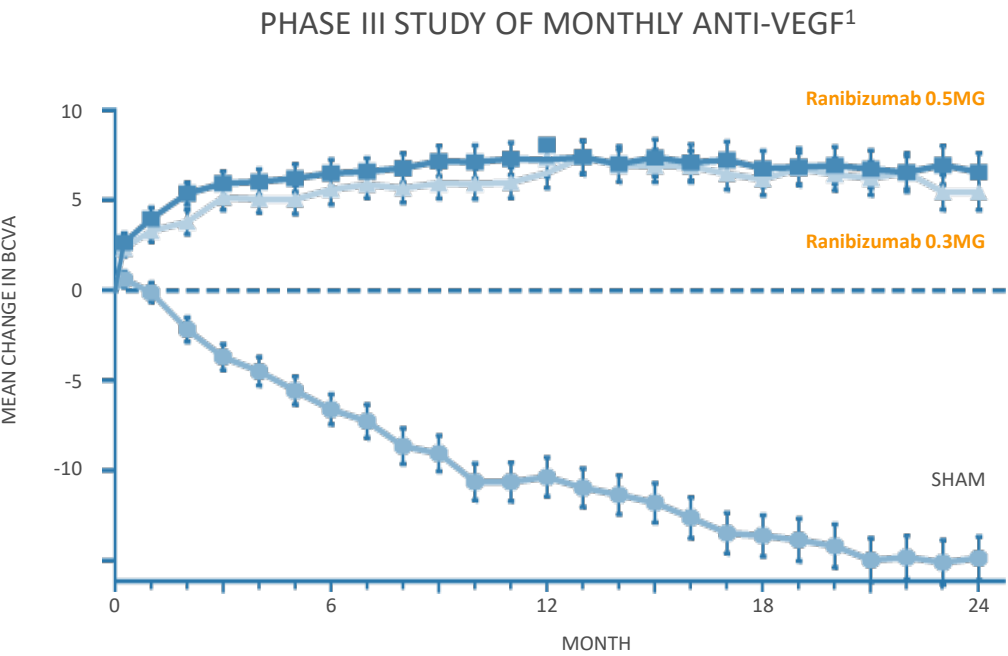
J. Pablo Velazquez-Martin MD

VP, Clinical Research &
Translational Medicine

BAYER

WHAT IS THE PROBLEM TO BE SOLVED?

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



Recommended dosing in first year:

Ranibizumab **12**
(MONTHLY)

Aflibercept **8**
(BI-MONTHLY AFTER 3 MONTHLY
LOADING DOSES)

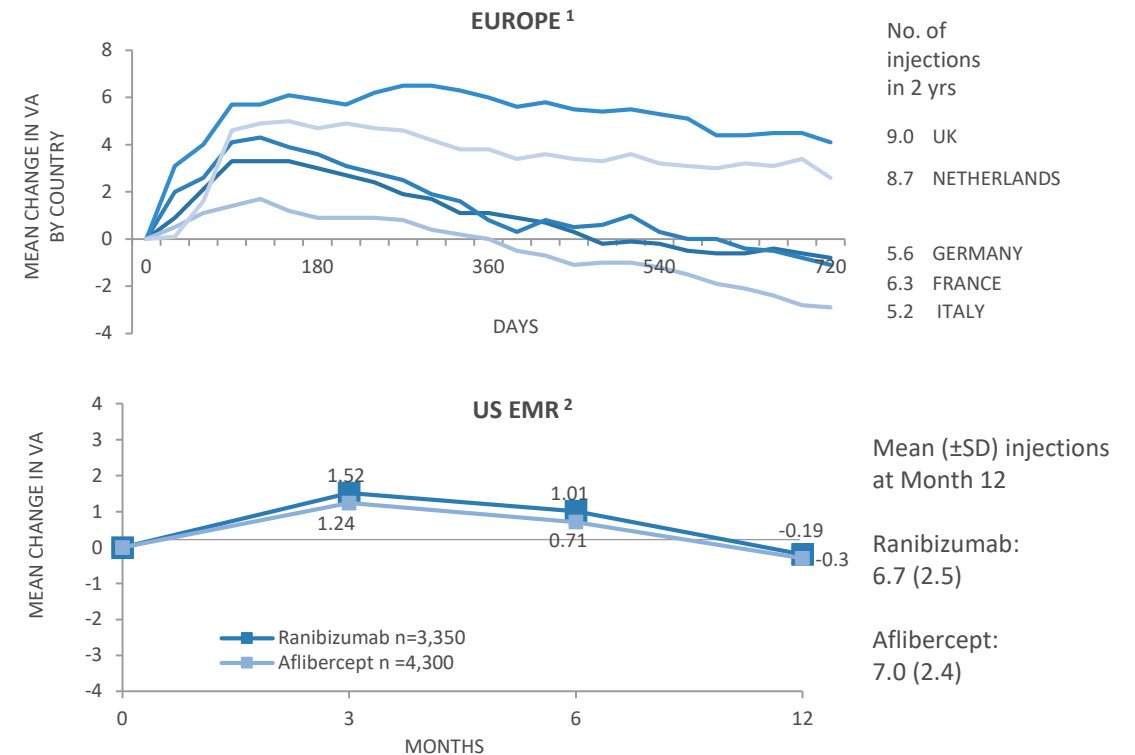
1. Rosenfeld PJ et al; MARINA Study Group. N Engl J Med. 2006;355:1419-14313. 2. Brown DM et al; ANCHOR Study Group. Ophthalmology. 2009;116:57-65

WHAT IS THE PROBLEM TO BE SOLVED?

...yet minimal visual gains are achieved in real-world practice

- 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 agents

1. The AURA Study, adapted from Holz FG et al. Br J Ophthalmol 2015; 99 (2): 220–226.
2. Adapted from Lotery A, et al. Eye (Lond). 2017 Dec;31(12):1697-1706.
EMR= Electronic Medical Records

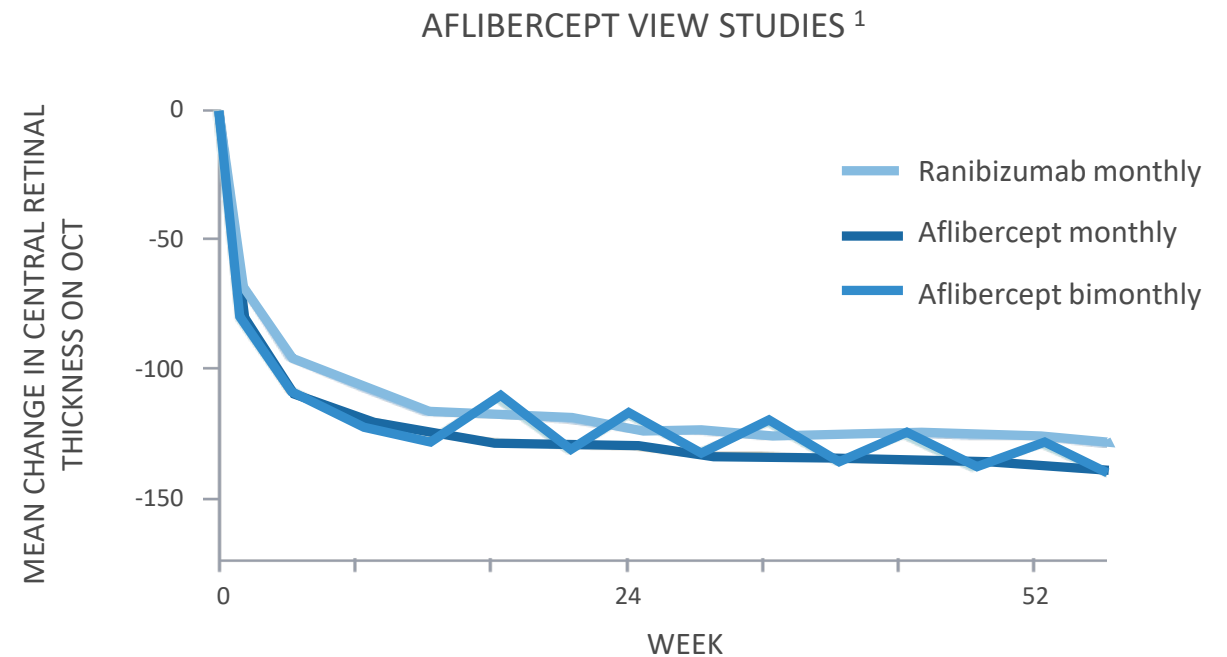


WHY?

Current anti-VEGF drugs do not control disease for long enough between doses.

Undertreatment leads to disease progression and permanent retinal damage.

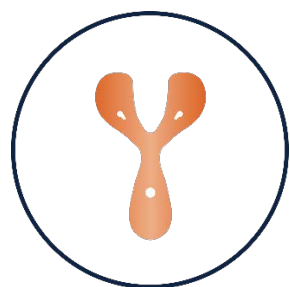
Bimonthly Anti-VEGF Results in Disease Activity between Doses



¹ Heier JS. Ophthalmology. 2012 Dec;119(12):2537-48. 3. Dugel PU. Ophthalmology. 2017 Sep;124(9):1296-1304. CRT= Central Retinal Thickness

ABC PLATFORM™

ANTIBODY BIOPOLYMER CONJUGATE

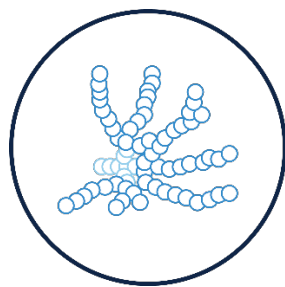


ANTIBODY

IgG1 with inert immune effector function

+

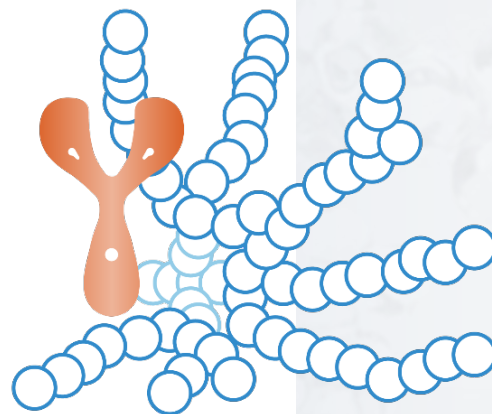
Stable linkage



BIOPOLYMER

Optically clear, high molecular weight phosphorylcholine polymer

=



ANTIBODY BIOPOLYMER CONJUGATE MEDICINES

KSI-301: the “GO-TO ANTI-VEGF” designed to solve the real-world effectiveness problem

SAME WHERE IT MATTERS

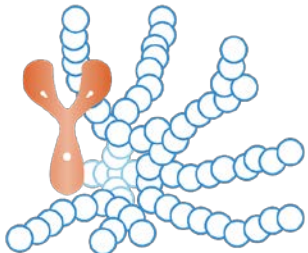
- Clinically proven target: VEGF
- Antibody-based biologic
- Intravitreal injection
- Optically clear solution
- No ocular residues

DIFFERENT WHERE IT COUNTS

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

KSI-301 optimizes size & formulation strength to improve durability

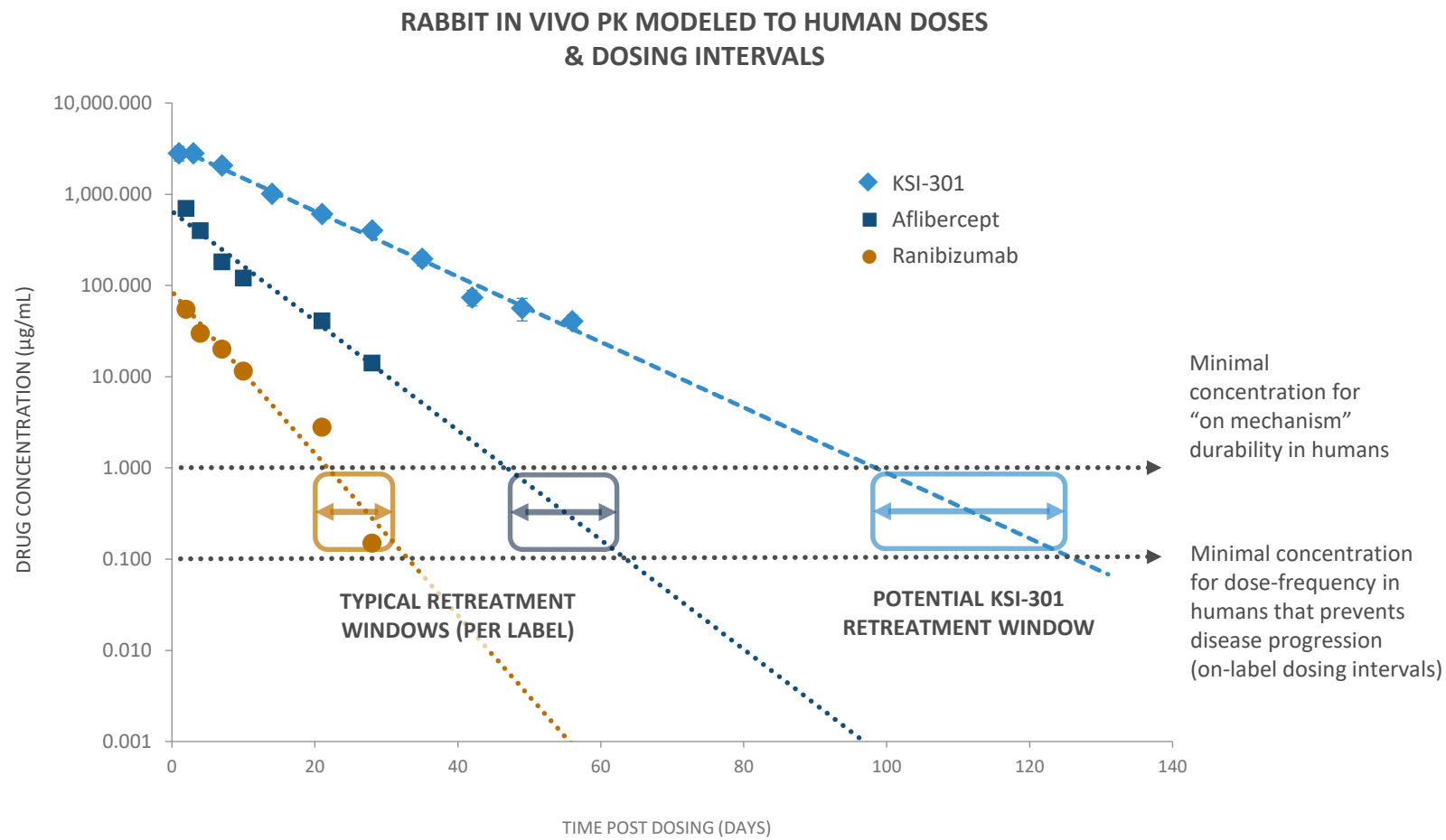
Drug/Candidate:	BROLUCIZUMAB	RANIBIZUMAB	AFLIBERCEPT
Molecule type	Single-chain antibody fragment	Antibody fragment	Recombinant fusion protein
Molecular structure			
Molecular weight	26 kDa	48 kDa	115 kDa
Clinical dose	6 mg	0.3-0.5 mg	2 mg
Equivalent molar dose	22	1	2
Equivalent ocular PK	<1	1	1.5
Equivalent ocular concentration at 3 months	10	1	1,000

KSI-301
Antibody Biopolymer Conjugate (ABC)

950 kDa
5 mg (by weight of antibody)
7
4
1,000,000

KSI-301 bioconjugate has potential for **extended durability** and a **more flexible retreatment window**

- KSI-301 bioconjugate has a flatter (better) ocular PK curve
- An increasing concentration advantage versus other biologics over time

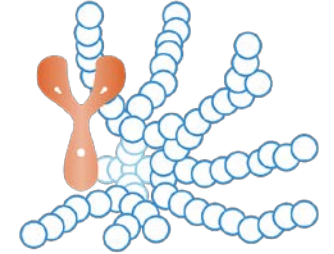
Ranibizumab data: Gaudreault et al (2007) IOVS 46(2) 726 Gaudreault et al (2007) Retina 27(9) 1260
Bakri et al (2007) Ophthalmol 114(12) 2179 || Aflibercept data: EVER Congress Portoroz Slovenia
(2008) Struble (Covance) Koehler-Stec (Regeneron). Aflibercept data adjusted arithmetically to
reflect 2,000µg dose administered (based on rabbit in vivo dosing of 500 µg) || KSI-301 data
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



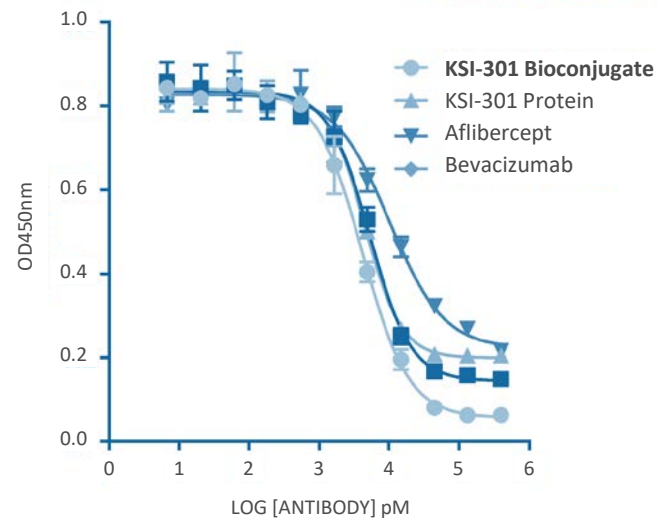
KSI-301 bioconjugate has shown **greater potency in vitro** than unconjugated anti-VEGFs

In vitro assays demonstrate KSI-301 bioconjugate has a **deeper potency** compared to ranibizumab, aflibercept and bevacizumab because of the special nature of its phosphorylcholine biopolymer.

- KSI-301 has high binding affinity to VEGF (K_D 6.75 pM, KinExA 37°C)
- KSI-301 bioconjugate has a deeper potency than other anti-VEGF and even its unconjugated starting protein

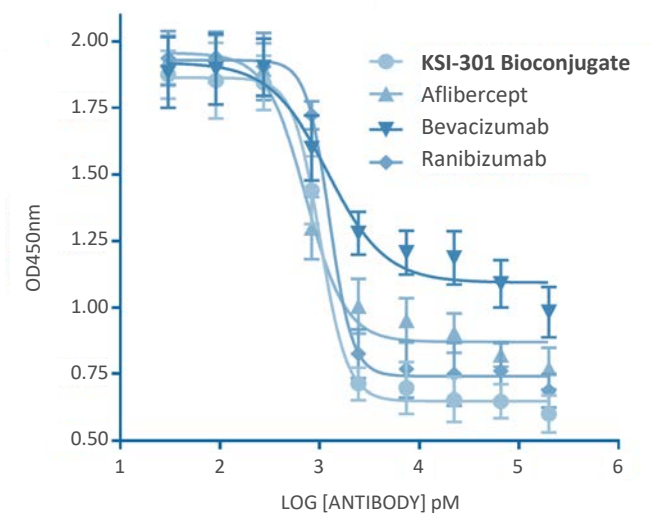


BIOCHEMICAL ASSAY
Anti-VEGF Inhibition of VEGF: VEGFR binding



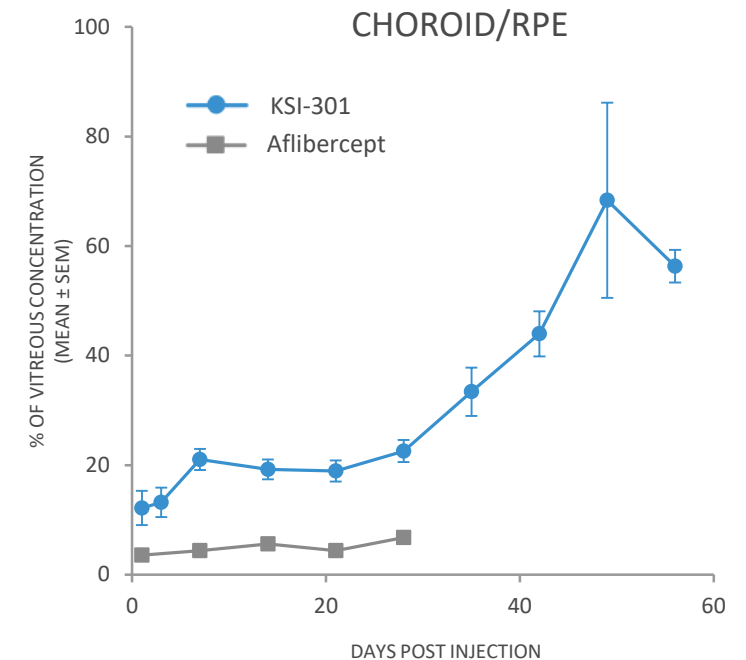
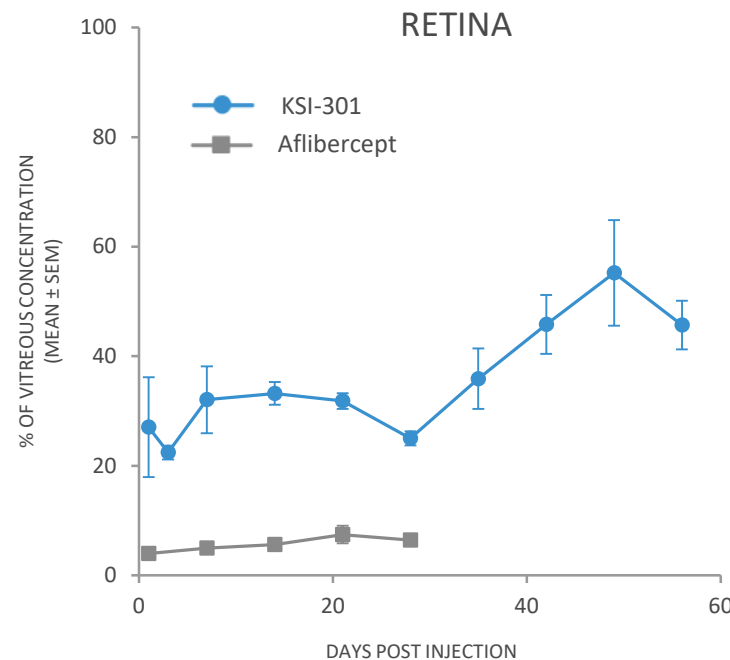
Kodiak data on file

PRIMARY HUMAN RETINAL CELL-BASED ASSAY
Anti-VEGF Inhibition of HRMVEC Proliferation



KSI-301 bioconjugate has **enhanced bioavailability** because of its phosphorylcholine biopolymer

- Ocular tissue bioavailability after single intravitreal injection
- Data from in vivo rabbit models
- Despite 8x larger size, KSI-301 has 8x greater access to retina than aflibercept



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

KSI-301's "high science" design positions it well versus competing efforts

- Using science to solve the durability problem
- Integrates seamlessly - over 26 million intravitreal injections of anti-VEGFs performed globally in 2018
- The single "go-to" anti-VEGF for all patients, encompassing all stages of therapy (induction, maintenance, supplemental)

	GENE THERAPY	PORT DELIVERY SYSTEM	COATED MICROSPHERE DRUG DEPOT	NEW BIOLOGIC TARGETS	KSI-301
Targets Validated Pathway					
Delivery	Surgery (complications)	Surgery (complications)	Intravitreal Injection	Intravitreal Injection	Intravitreal Injection
Anticipated Durability Improvement					
No Supplemental Injections					
No Residual Foreign Material in the Eye					
Use for early disease modification					

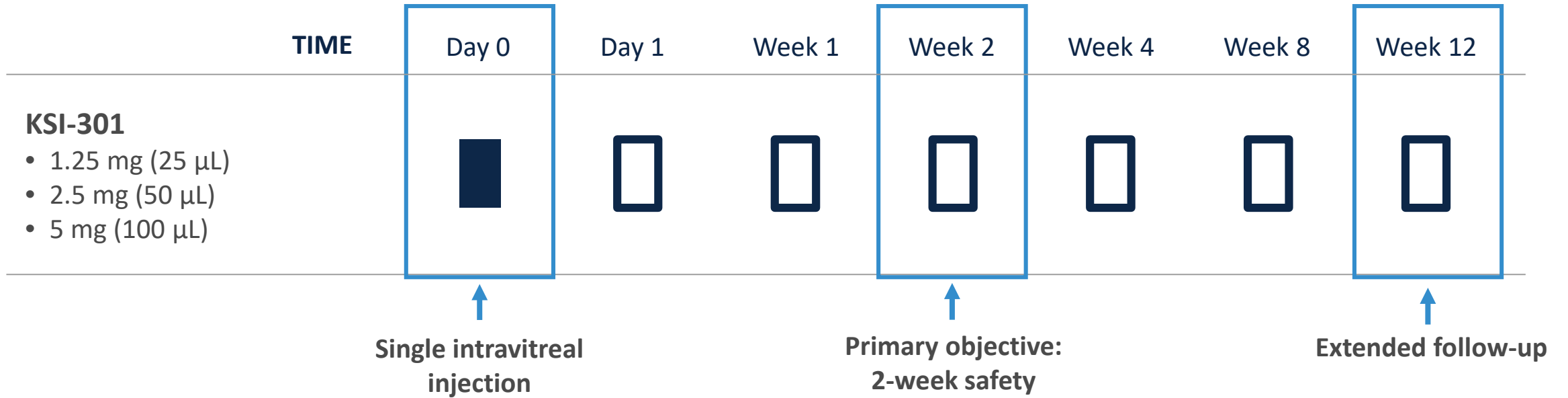
OUR GOAL WITH KSI-301

Develop KSI-301 to be “first off-the-shelf”
by meeting the individual needs of key
stakeholders globally

- ✚ Patient & Patient’s Family
- ✚ Retina Specialist & Care Team
- ✚ Retina Practice Owner
- ✚ Payer
- ✚ Health System

KSI-301
PHASE 1
CLINICAL STUDY

Single Ascending Dose Study in Diabetic Macular Edema Patients



- 9 subjects – 3 per dosing cohort, 7-day safety review between each cohort
- Conducted at 5 US sites
- Single dose with observation to 12 weeks (no retreatment)

PATIENT INFORMATION

Demographic and Ocular Baseline Characteristics

DEMOGRAPHICS	
Age (years, mean)	62
Gender	7M, 2F
OCULAR CHARACTERISTICS Study Eye, n=9	
Previously Received Anti-VEGF	8/9
Number of Anti-VEGF Treatments in Last Year median (range)	3 (0, 7)
Time Since Last Anti-VEGF, Days median (range)	95 (52,>365)
IOP, mmHg mean (SD)	15 (2)
OCT Central Subfield Thickness, Microns mean (SD)	565 (182)
Baseline BCVA, ETDRS Letters mean (SD)	47 (12)
Baseline BCVA, Snellen Equivalent	20/100

SAFETY OUTCOMES

Every dose level was well-tolerated through the 12 week follow-up period

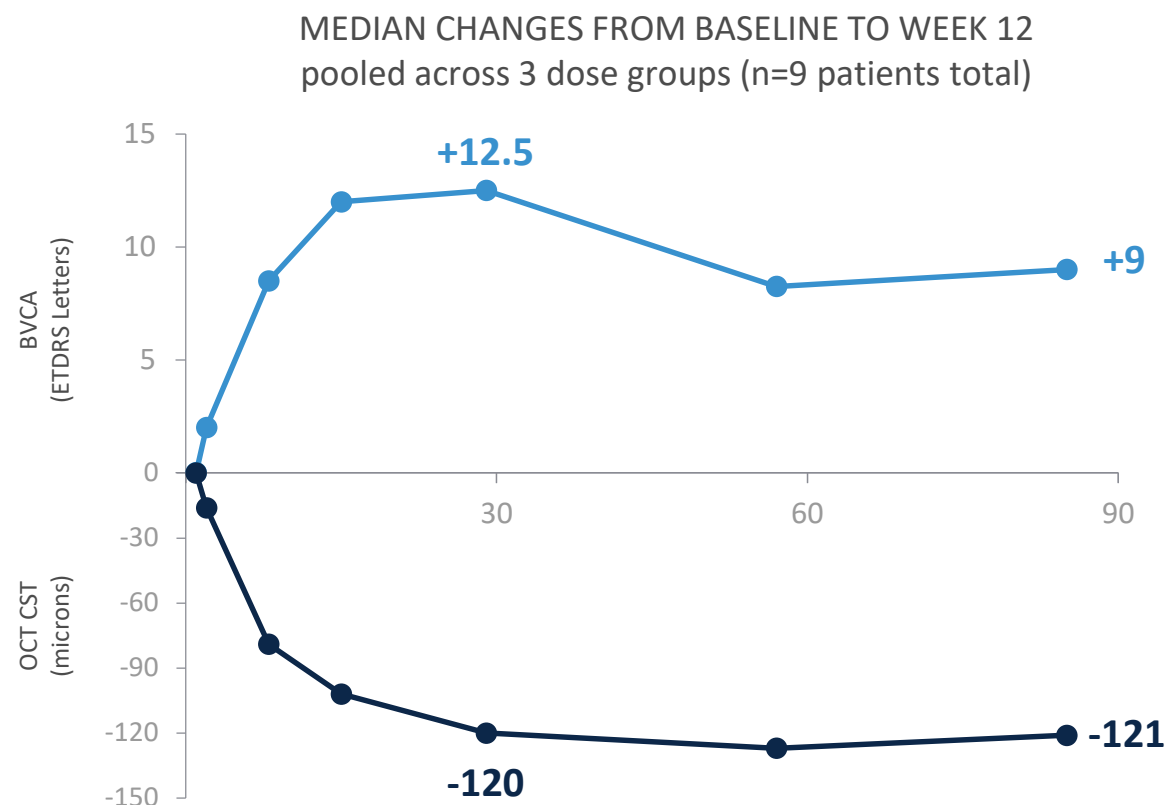
- No dose limiting toxicities
 - No drug-related adverse events or drug-related serious adverse events
 - No intraocular inflammation
- Optically clear media after each injection
 - No anti-drug antibodies detected in any patient
- Systemic levels 1/3 of bevacizumab C_{max} and 1/6 of D28 level (1.25mg dose)¹

NUMBER OF PATIENTS WITH ANY AE=4	N	SERIOUS	RELATED
OCULAR AEs			
Foreign body sensation	1	N	N
Subconjunctival hemorrhage	2	N	N
Floaters (reported in both eyes)	1	N	N
Visual flashes	1	N	N
NON-OCULAR AEs			
Fall	1	N	N
Worsening of coronary artery disease	1	Y	N
Swollen Feet	1	N	N

1. Avery RL et al. Retina. 2017 Oct;37(10):1847-1858

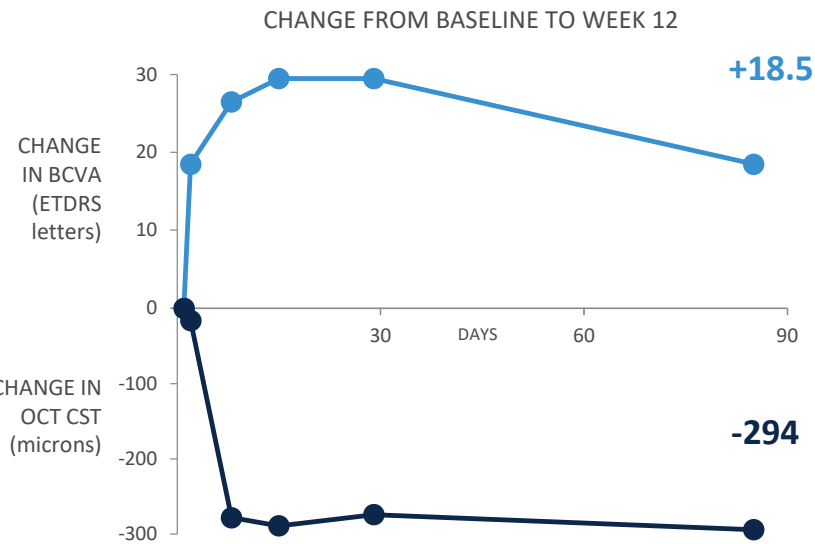
Improvements in vision and retinal thickness after single-dose KSI-301 maintained through 12 weeks

- Rapid, high magnitude responses as early as 1 week after dosing
- Durable improvements out to 12 weeks



KSI-301
CASE
EXAMPLES
—

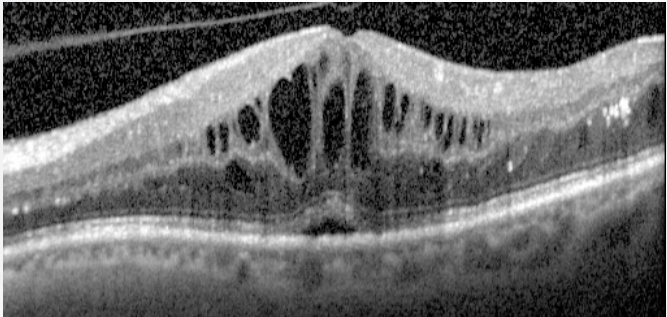
CASE EXAMPLE 1



CLINICAL HISTORY SUMMARY (SITE REPORTED):

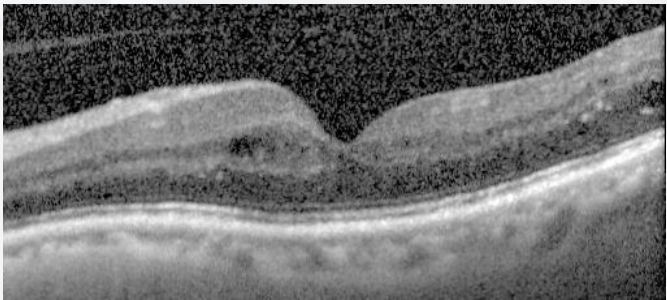
	Date	Treatment	VA Snellen	CST
Retrospective	1/2018		20/40	-
	4/2018	Bevacizumab	20/40	431
	6/2018	Bevacizumab	20/60	655
	8/2018	KSI-301	20/160	636

Resolution of macular edema sustained through 12 weeks in patient with prior suboptimal response

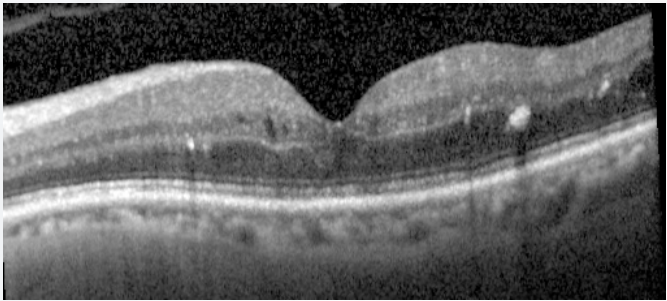


BASELINE

Single dose
KSI-301 (5 mg)

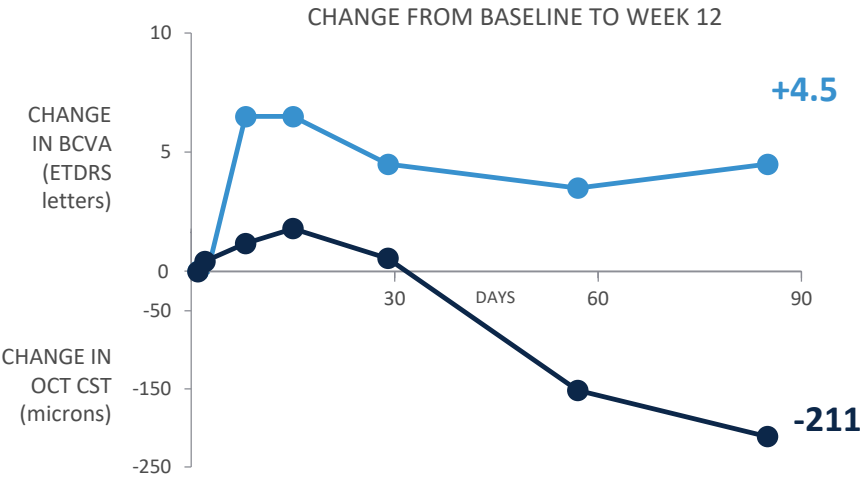


WEEK 4



WEEK 12

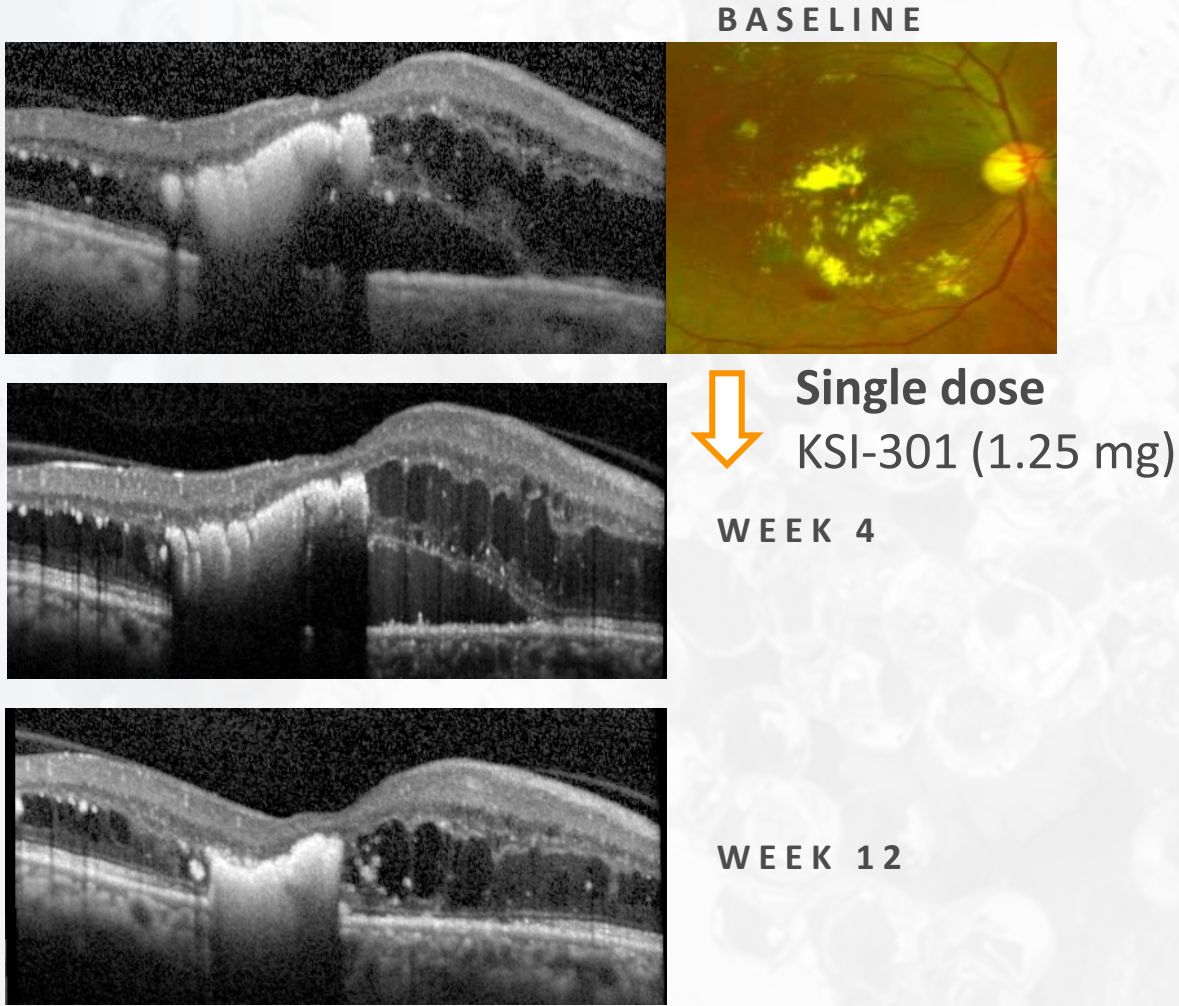
CASE EXAMPLE 2



CLINICAL HISTORY SUMMARY (SITE REPORTED):

	Date	Treatment	VA Snellen
Retrospective	1/2018	Bevacizumab	20/60
	3/2018	Bevacizumab	20/100
	4/2018	Bevacizumab	20/150
	5/2018		20/350
	7/2018	KSI-301	20/80

Resolution of subretinal fluid through 12 weeks in patient with extensive foveal lipid exudates



IMPORTANT EARLY
DEVELOPMENT QUESTIONS

SUCCESSFULLY
ADDRESSED



Manufacturability



Optical Clarity



Target Tissue Access



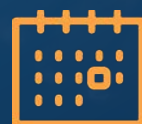
Safety



Speed of Onset



Potency



Clinical Durability

PHASE 1B

OPEN LABEL STUDY

Wet AMD, DME, RVO

- Open-label study to further explore KSI-301 safety, bioactivity, durability
- Actively enrolling (NCT03790852)
- Approximately 50 patients
- Anti-VEGF **treatment naïve** patients only
- **3 loading doses** in every patient
- **7-month follow-up** to explore durability (vision, retinal thickness)
- On track to complete recruitment in 1st half of 2019

Week		0	4	8	12	16	20	24	28	32	36	Primary Endpoint
KSI-301 5 or 2.5 mg	wAMD											
	DME											
	RVO											

 KSI-301 injection

 Dosing as needed (PRN)

 Retreatment criteria assessment

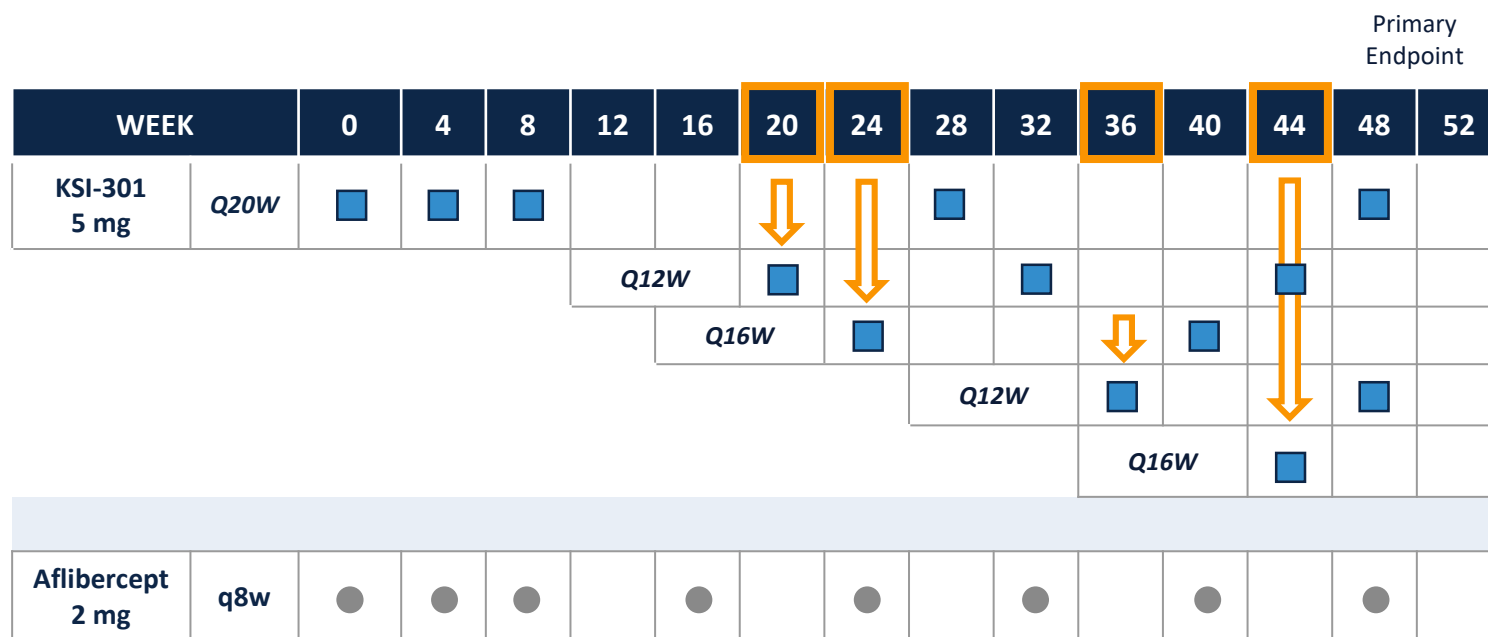
PHASE 2

RANDOMIZED CONTROLLED STUDY

In wet AMD (US/EU)

Pivotal study design, head-to-head
against standard of care aflibercept

- All patients $\geq$ Q12W with KSI-301
- As infrequent as Q20W dosing with KSI-301
- Non-inferiority pivotal design study
- Estimated 400 patients (US/EU)
- On track to begin enrolling in 2Q 2019, with interim (durability) in 2020 and primary (vision) in 2021



■ KSI-301 injection

● Aflibercept injection

↓ Disease Activity Dosing Adjustment

□ Disease Activity Assessment Visit

1ST HALF

2ND HALF

Complete recruitment into phase 1b
multiple-dose study in patients
with wet AMD, DME, and RVO

Start of global pivotal phase 2/3
head-to-head study against
aflibercept in wet AMD

Submit China IND's for phase 2
trials in wet AMD, DME and DR

Continued phase 1b data

Present phase 1b data at key
ophthalmology meetings
(e.g. ASRS, AAO)

Interim phase 2/3 data
from global wet AMD

Initiate China studies
in wAMD and DME

Disclose NPDR strategy

Interim phase 2 data
from China studies

Initiate global pivotal
phase 2/3 in DME

R&D Day

2019

2020

KSI-301 POTENTIAL

CATALYSTS

KEY TAKEAWAYS



KSI-301 is a novel **Antibody Biopolymer Conjugate** that inhibits VEGF

- Same where it matters, different where it counts



Phase 1a single ascending dose study results

- Well-tolerated at all dose levels
- **Rapid-onset, high-magnitude improvements sustained to 12 weeks**



Kodiak executing on a comprehensive and aggressive clinical strategy for KSI-301 in retinal vascular diseases

- Phase 1b data in 2019
- Initiating multiple Phase 2 pivotal studies in 2019



Objective

The “go-to” anti-VEGF, meeting all treatment needs for VEGF-mediated retinal vascular diseases

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