A photograph of a smiling grandfather, grandmother, and young granddaughter looking at a tablet together in a park. The grandfather is on the left, wearing glasses and a light-colored shirt. The grandmother is on the right, smiling. The young girl is in the center, holding the tablet. The background is a blurred green park.

Developing next-generation
immunotherapies that address
cancer immune resistance

April 2023

KA (Nasdaq)

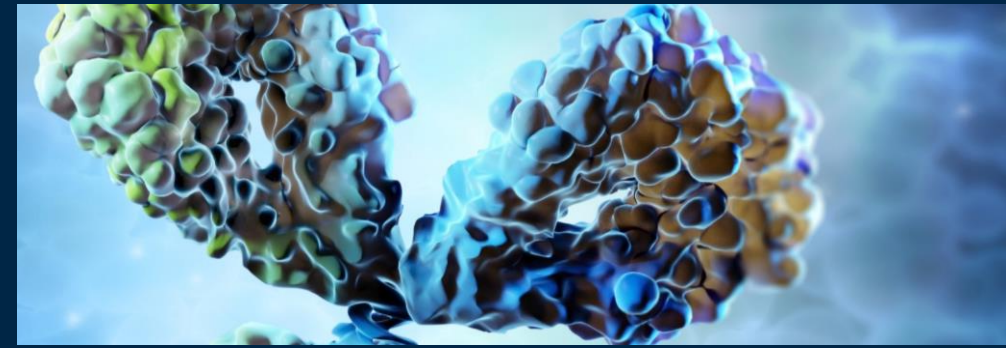
Disclaimers and other information

Cautionary Statements Regarding Forward-Looking Statements

This presentation contains “forward-looking statements” within the meaning of the Private Securities Litigation Reform Act of 1995. The use of words such as, but not limited to, “believe,” “expect,” “estimate,” “project,” “intend,” “future,” “potential,” “continue,” “may,” “might,” “plan,” “will,” “should,” “seek,” “anticipate,” or “could” and other similar words or expressions are intended to identify forward-looking statements. Forward-looking statements are neither historical facts nor assurances of future performance. Instead, they are based on Kineta’s current beliefs, expectations and assumptions regarding the future of Kineta’s business, future plans and strategies, clinical results and other future conditions. New risks and uncertainties may emerge from time to time, and it is not possible to predict all risks and uncertainties. No representations or warranties (expressed or implied) are made about the accuracy of any such forward-looking statements.

Such forward-looking statements are subject to a number of material risks and uncertainties including, but not limited to: the adequacy of Kineta’s capital to support its future operations and its ability to successfully initiate and complete clinical trials; the difficulty in predicting the time and cost of development of Kineta’s product candidates; Kineta’s plans to research, develop and commercialize its current and future product candidates, including, but not limited to, KVA12123; the timing and anticipated results of Kineta’s planned pre-clinical studies and clinical trials and the risk that the results of Kineta’s pre-clinical studies and clinical trials may not be predictive of future results in connection with future studies or clinical trials; the timing of the availability of data from Kineta’s clinical trials; the timing of any planned investigational new drug application or new drug application; the risk of cessation or delay of any ongoing or planned clinical trials of Kineta or its collaborators; the clinical utility, potential benefits and market acceptance of Kineta’s product candidates; Kineta’s commercialization, marketing and manufacturing capabilities and strategy; developments and projections relating to Kineta’s competitors and its industry; the impact of government laws and regulations; the timing and outcome of Kineta’s planned interactions with regulatory authorities; Kineta’s ability to protect its intellectual property position; Kineta’s estimates regarding future revenue, expenses, capital requirements and need for additional financing; and those risks set forth under the caption “Risk Factors” in Kineta’s most recent Annual Report on Form 10-K, as well as discussions of potential risks, uncertainties and other important factors in Kineta’s subsequent filings with the Securities and Exchange Commission. Any forward-looking statement speaks only as of the date on which it was made. Except as required by law, Kineta undertakes no obligation to publicly update or revise any forward-looking statement, whether as result of new information, future events or otherwise.

Kineta is developing next-generation immunotherapies that address cancer immune resistance



Innate Immunity Focused Pipeline

KVA12123

- VISTA blocking mAb to address immunosuppression in the TME
 - Opened Phase 1/2 clinical study evaluating KVA12123 alone and in combination with pembrolizumab in advanced solid tumors
- Preclinical Anti-CD27 agonist mAb to address exhausted T cells

Catalysts

3Q23 | KVA12123 initial clinical safety data
4Q23 | KVA12123 initial clinical efficacy data (ORR)

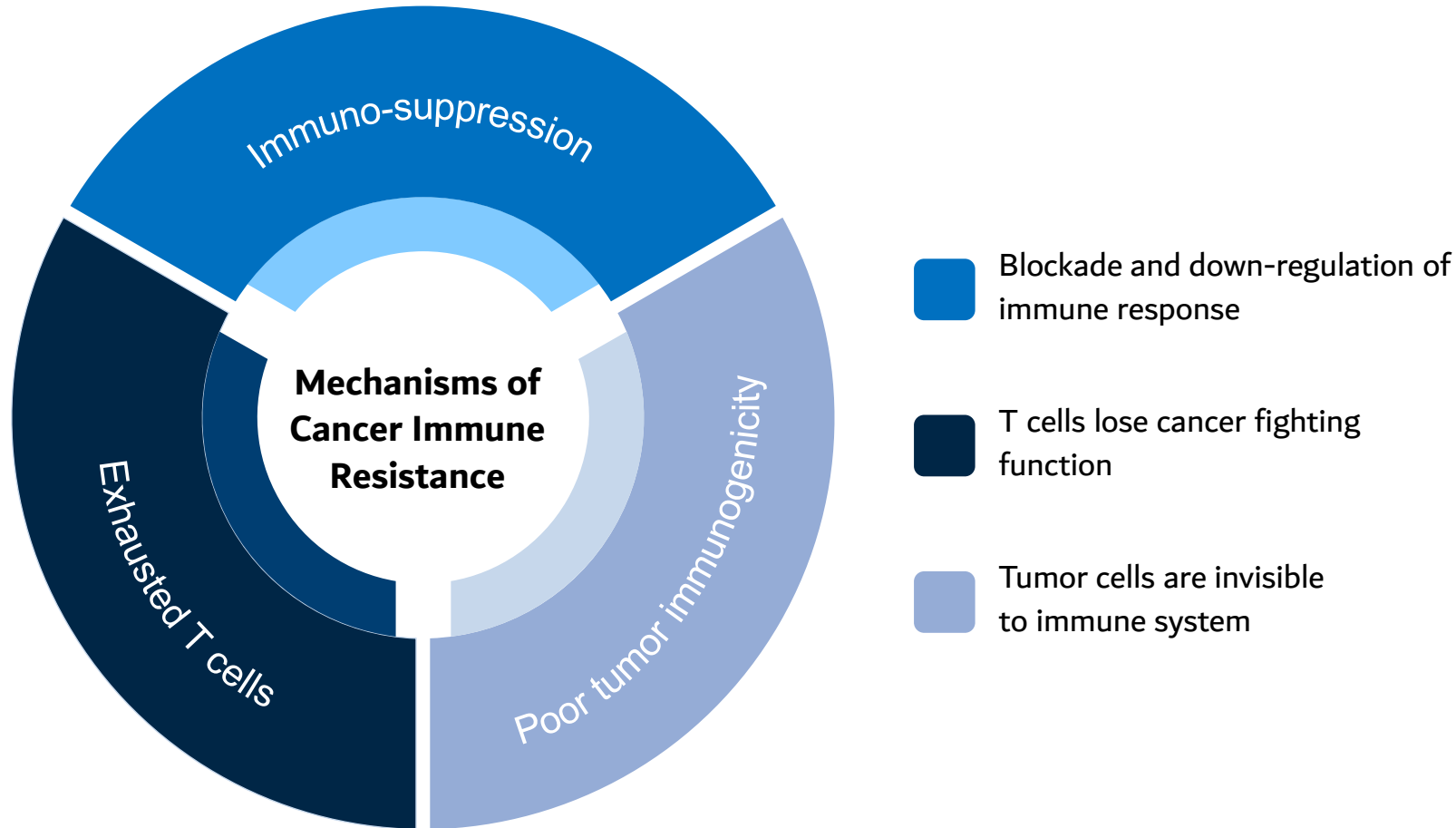
Runway

Through mid-2024

Partnerships



Immune resistance is a major challenge with current cancer therapy



Next-generation cancer treatments require:

Improving survival for checkpoint inhibitor (CPI) non-responders **(70-80%)***

Reprogramming the immune system to attack cancer

Integrating **innate and adaptive immune** responses

Kineta pipeline integrates innate and adaptive immunity to address mechanisms of cancer resistance

Innate immunity

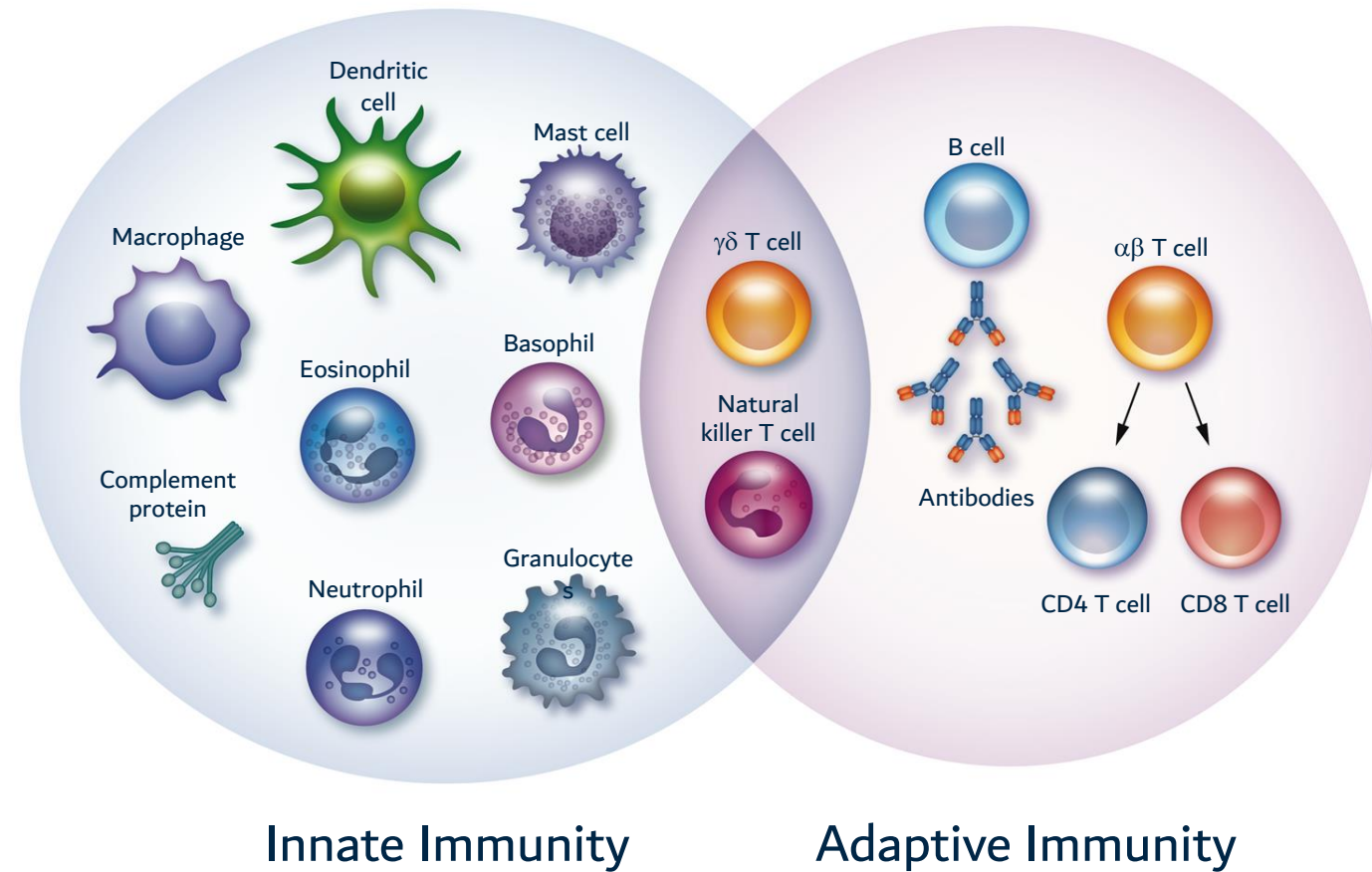
Involved in early response to cancer

Necessary driver for appropriate adaptive immunity

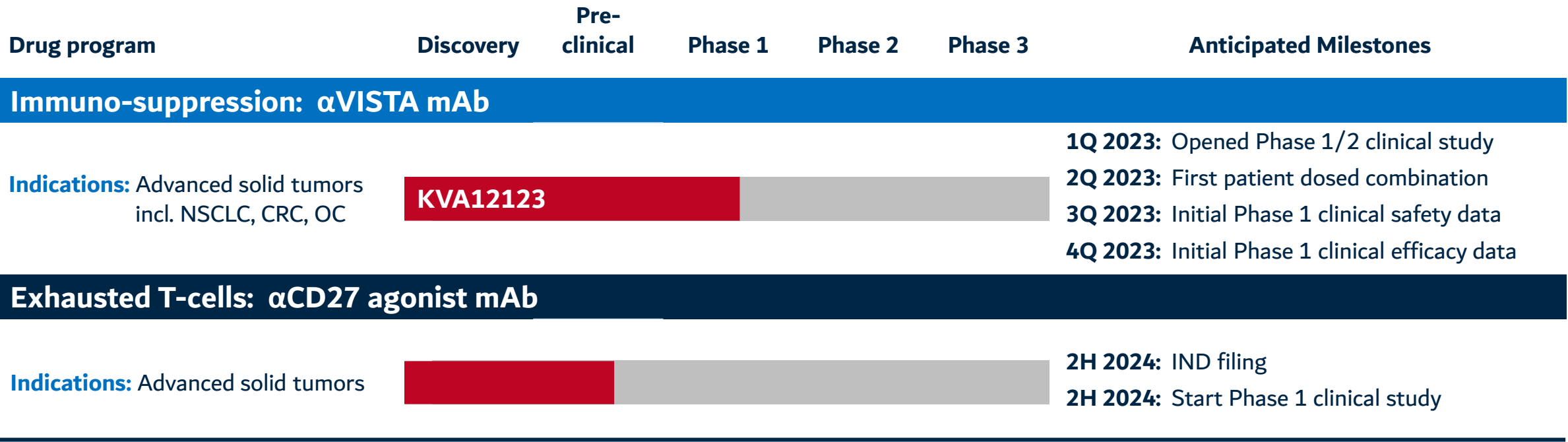
Significant cause of cancer resistance

Adaptive immunity

Most competitor drug development is focused **only** on T cell adaptive immunity



Kineta’s immuno-oncology pipeline aims to address the mechanisms of cancer immune resistance





KVA12123

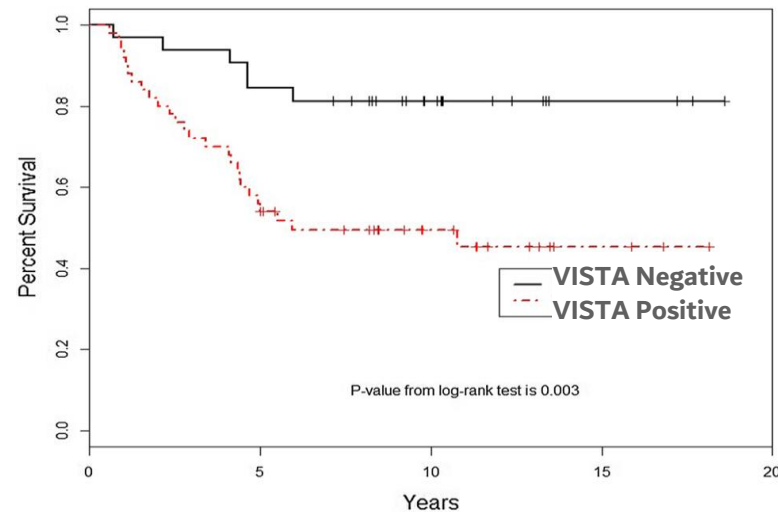
Potentially differentiated
VISTA blocking immunotherapy



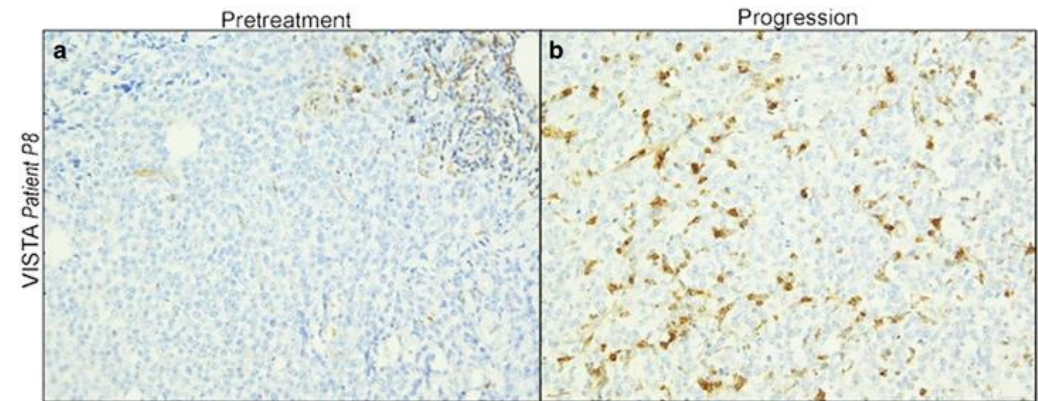
VISTA is a key driver of immunosuppression in the tumor microenvironment

- Immunosuppressive protein expressed on **myeloid cells**
- Highly expressed in **cold tumors** including lung, colon and ovarian cancers
- Correlates with **poor outcomes** in cancer patients
- Up-regulated after CPI therapy and **associated with treatment failure**

Melanoma patient survival by
VISTA expression in tumor-infiltrating immune cells ¹



VISTA expression increases in melanoma patient
during pembrolizumab relapse/progression ²



**Brown staining in human tumors
indicates VISTA expression**

KVA12123: Potentially differentiated VISTA blocking immunotherapy

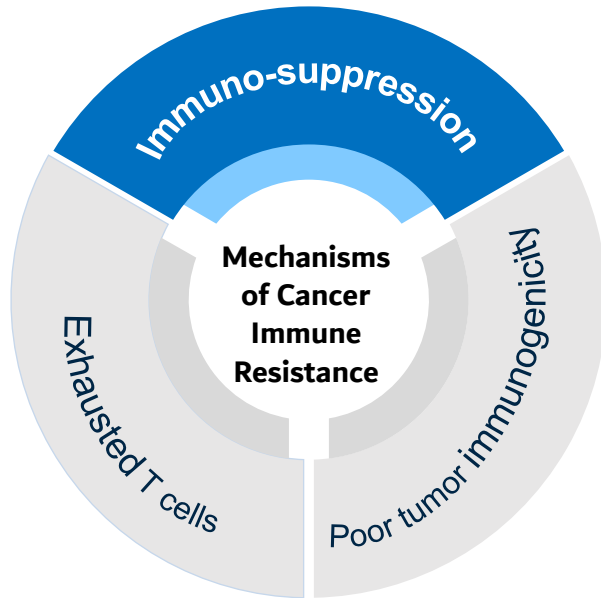
Product	Development stage	Isotype	pH Binding	Single Agent Tumor Model Efficacy	CRS Cytokine Release
Kineta KVA12123	Phase 1	Engineered IgG1 mAb that binds to a unique epitope	Binds at both physiologic pH and acidic pH in the TME	Strong single agent tumor growth inhibition and in combination with PD-1 in preclinical models	No CRS-associated cytokine release or neurotoxicity seen in preclinical models
Hummingbird HMBD002	Phase 1	IgG4	Physiologic	Moderate	IL-6
Pierre Fabre WO180	Phase 1				
Curis* CI-8993	Phase 1 (de-prioritized)	IgG1	Physiologic	Moderate	TNF α , IFN γ , IL2, IL-1 β
Sensei SNS-101	Preclinical	IgG1	Acidic	Weak	TNF α
Pharmabcine PMC309	Preclinical	IgG1	Acidic & Physiologic	Moderate	IFN γ

Other discovery stage programs: Apexigen, Five Prime Therapeutics/BMS, xCella Biosciences

Empty cells indicate no public data available

*Curis announced 11/9/2022 : “Concentrating its resources to focus on and accelerate emavusertib”, the company’s lead asset and “deprioritization of other programs” (CI-8993)

Blocking VISTA can reverse immunosuppression in the TME



Inhibits **MDSC** (myeloid-derived suppressor cells)

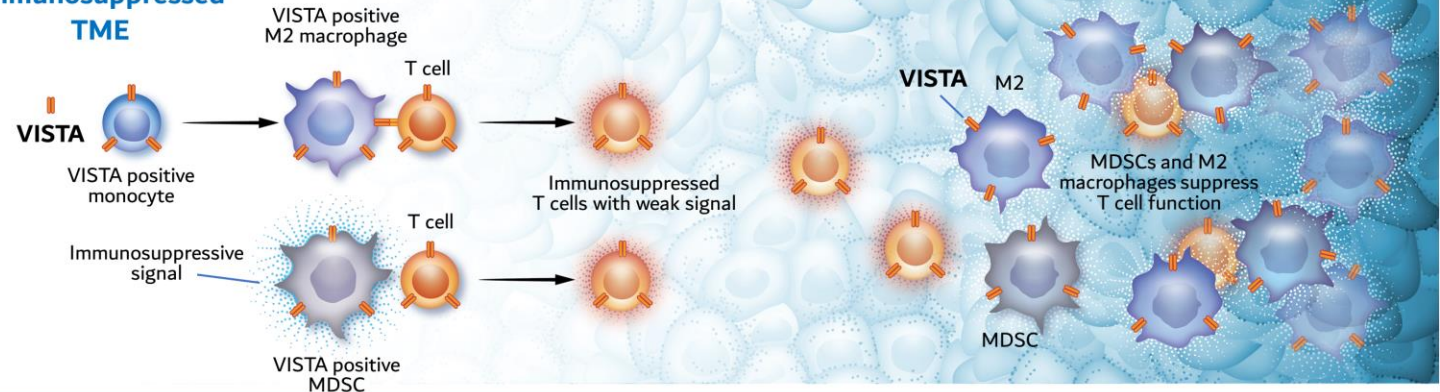
Promotes **T_{eff}** function

Enhances **NK cell** activation

Enhances **monocyte** activation

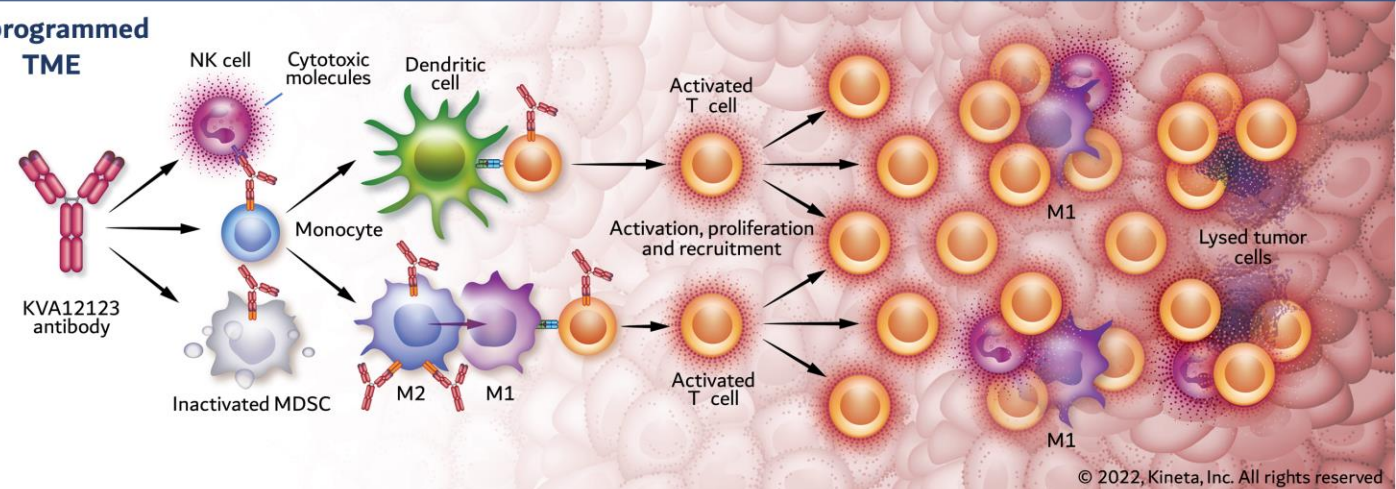
VISTA causes immunosuppression by inactivating T cells

Immunosuppressed TME



KVA12123 targets VISTA with the potential to promote T cell and NK cell anti-tumor function

Reprogrammed TME



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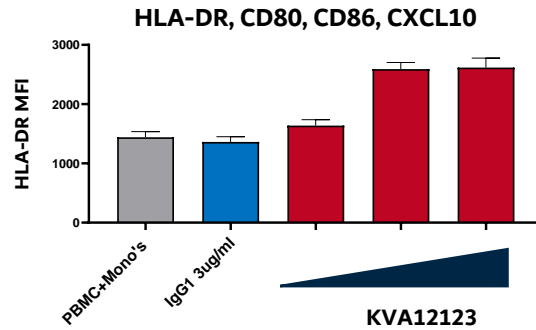
KVA12123 activates both innate and adaptive immune cells *in vitro*



Increases monocyte differentiation and activation



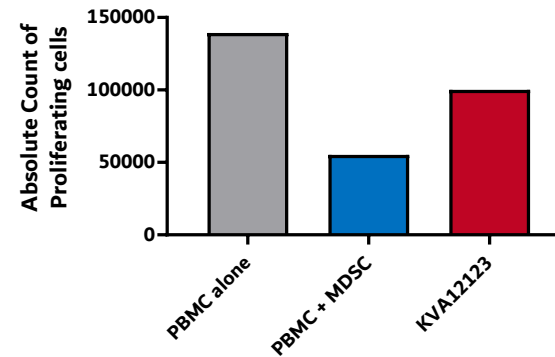
monocyte



Reduces MDSC-mediated T cell suppression



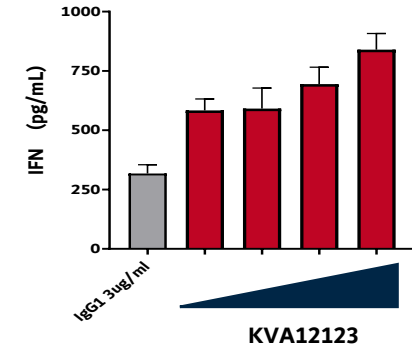
MDSC



Increases HLA-dependent T cell activation



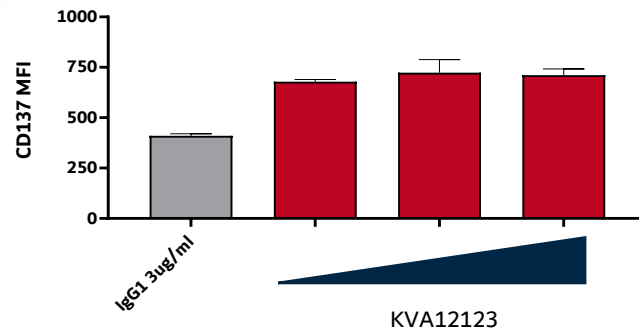
T cell



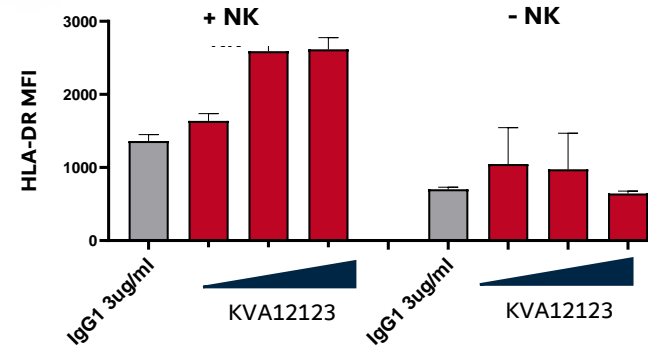
Enhances NK cell activation



NK cell

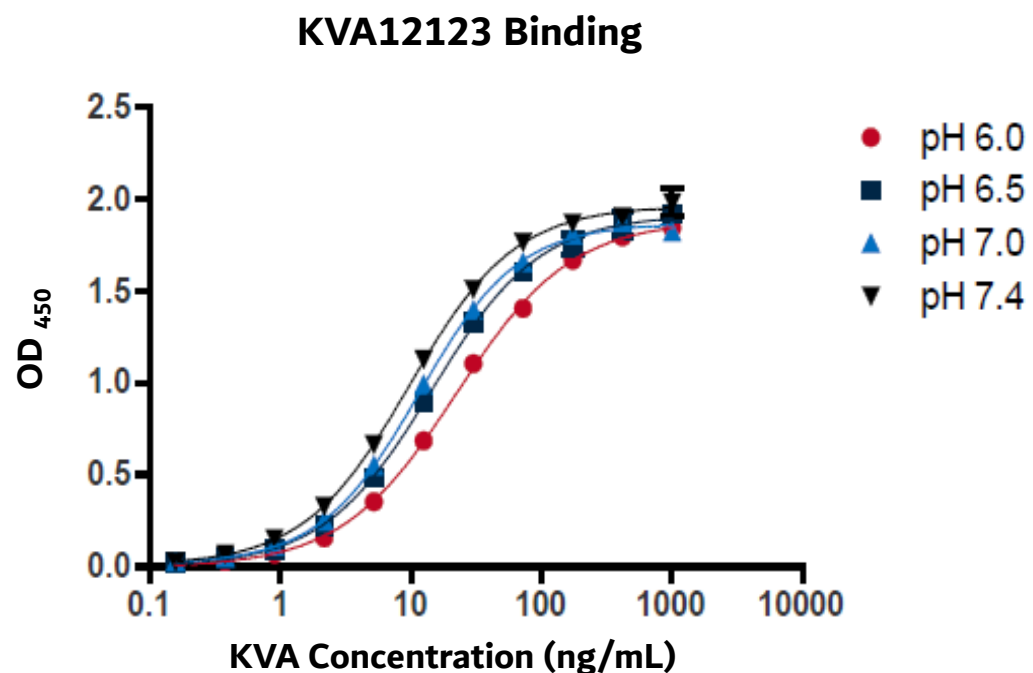


NK dependent mechanism of action

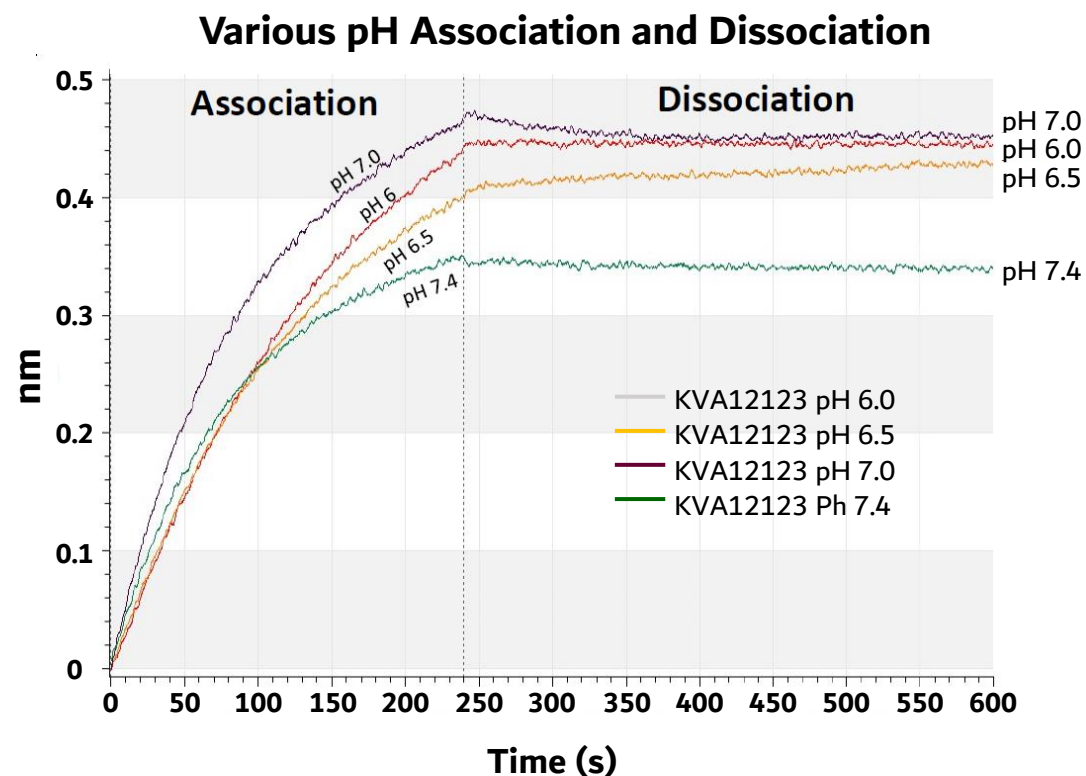


KVA12123 binds at physiologic and acidic pH

ELISA



Octet



Binding studies by ELISA and Octet demonstrate rapid on-rate and slow off-rate from pH 7.4 to pH 6.0

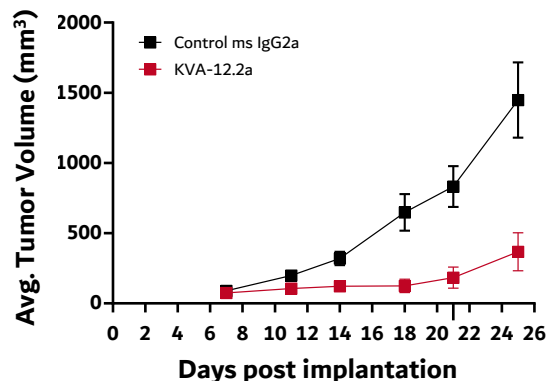
KVA12123 demonstrates single agent tumor growth inhibition and in combination with PD-1 in preclinical models

Monotherapy

Bladder Cancer Model MB49

hVISTA KI mice

Mean Tumor Volume



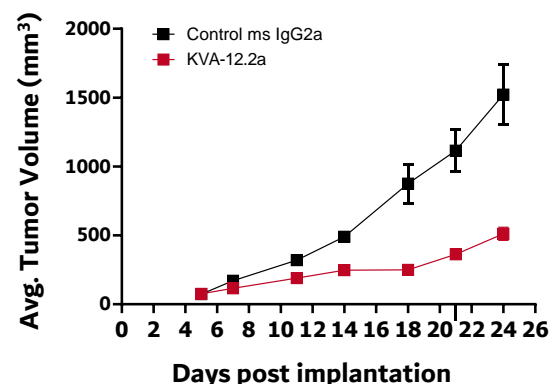
Tumor Growth Inhibition

Anti-VISTA: **75%**

T Cell Lymphoma Model EG7

hVISTA KI mice

Mean Tumor Volume



Tumor Growth Inhibition

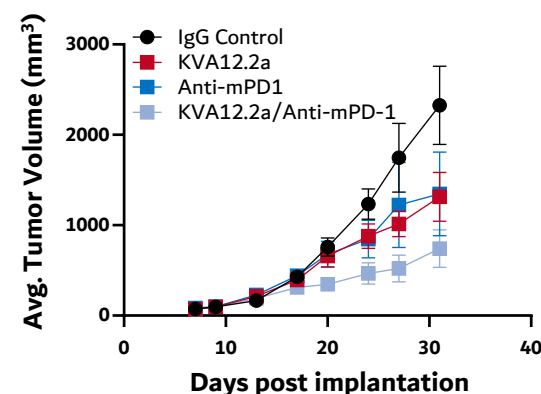
Anti-VISTA: **66%**

Combination therapy

Colon Carcinoma Model MC38*

hVISTA KI mice

Mean Tumor Volume



Tumor Growth Inhibition

Anti-VISTA: **35-42%**

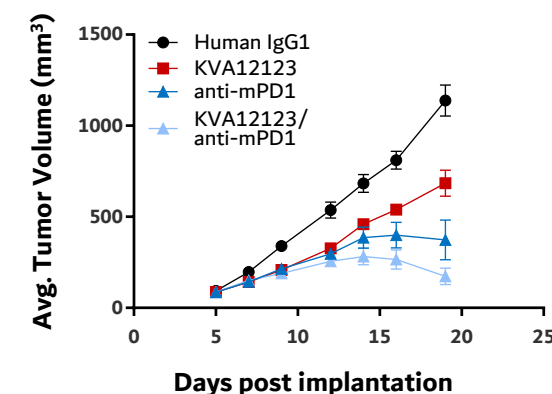
Anti-PD1: **42-60%**

Combination: **68%**

Bladder Cancer Model MB49*

hVISTA KI mice

Mean Tumor Volume



Tumor Growth Inhibition

Anti-VISTA: **40%**

Anti-PD1: **67%**

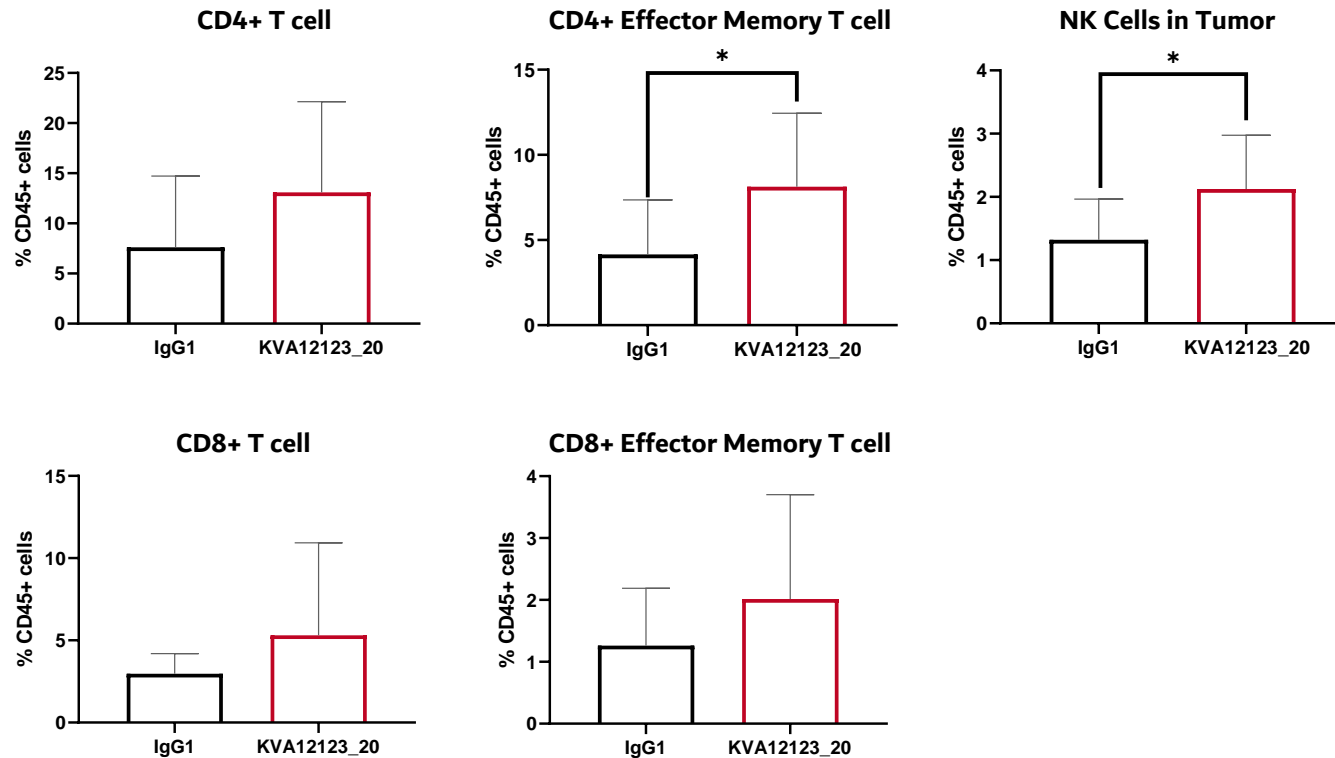
Combination: **85%**

*Combination therapy studies used sub-optimal doses of each agent

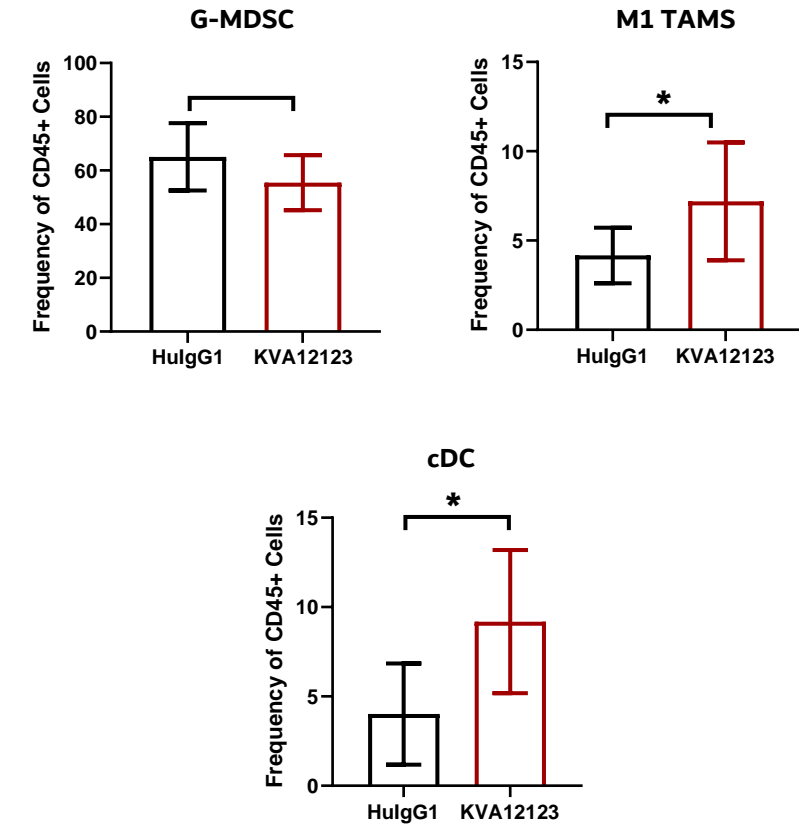
KVA12.2a: mouse isotype equivalent of KVA12123

KVA12123 drives an integrated innate and adaptive anti-tumor immune response in MB49 model (*ex vivo*)

Lymphoid compartment



Myeloid compartment



KVA12123 was observed to be well-tolerated in NHP toxicology studies



No
mortality



No change in CRS cytokine
levels (IL6 or TNF α)



No treatment-related
adverse events



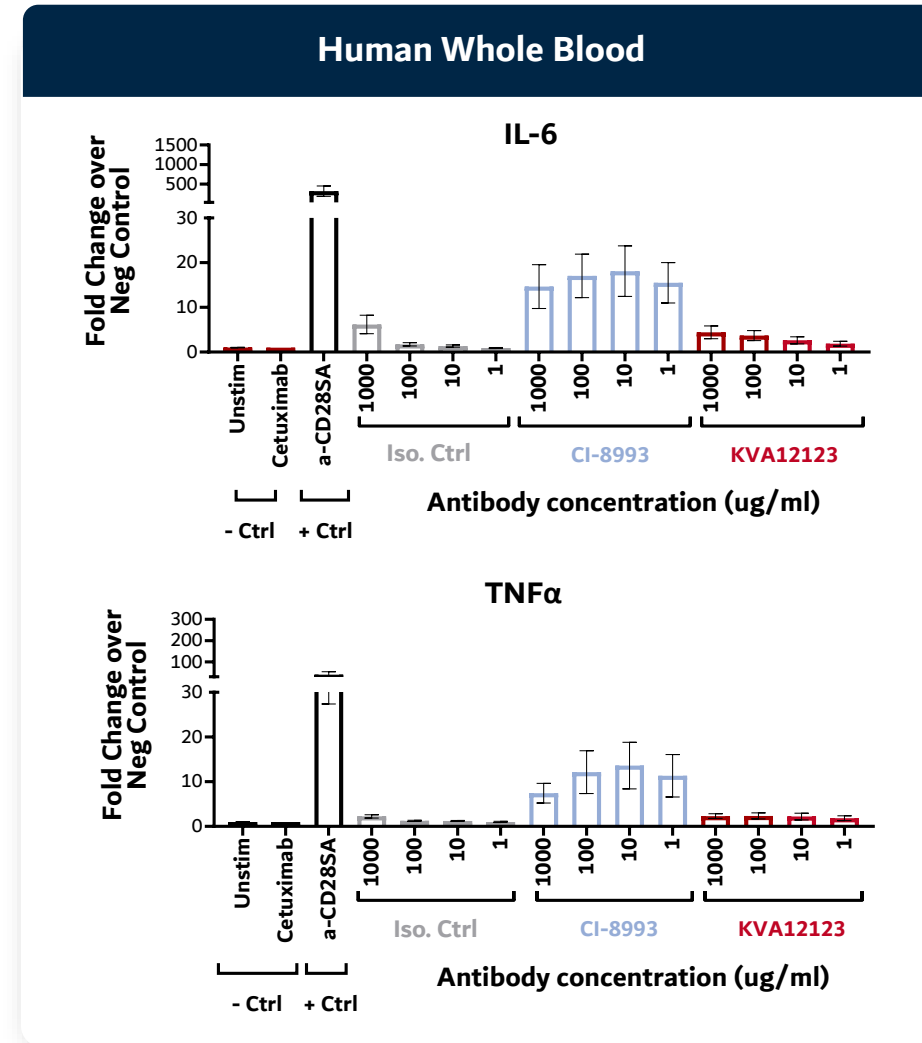
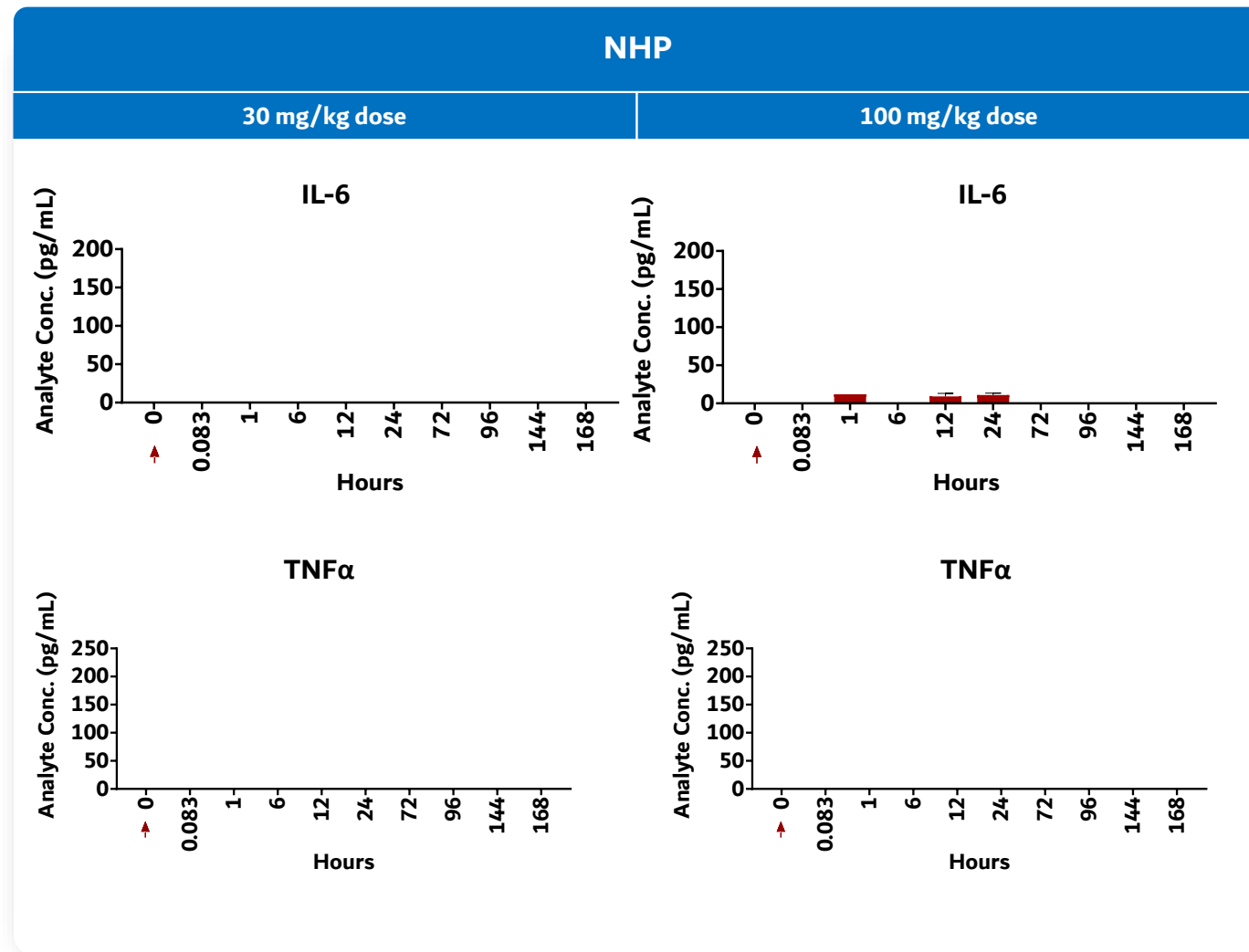
Well
tolerated



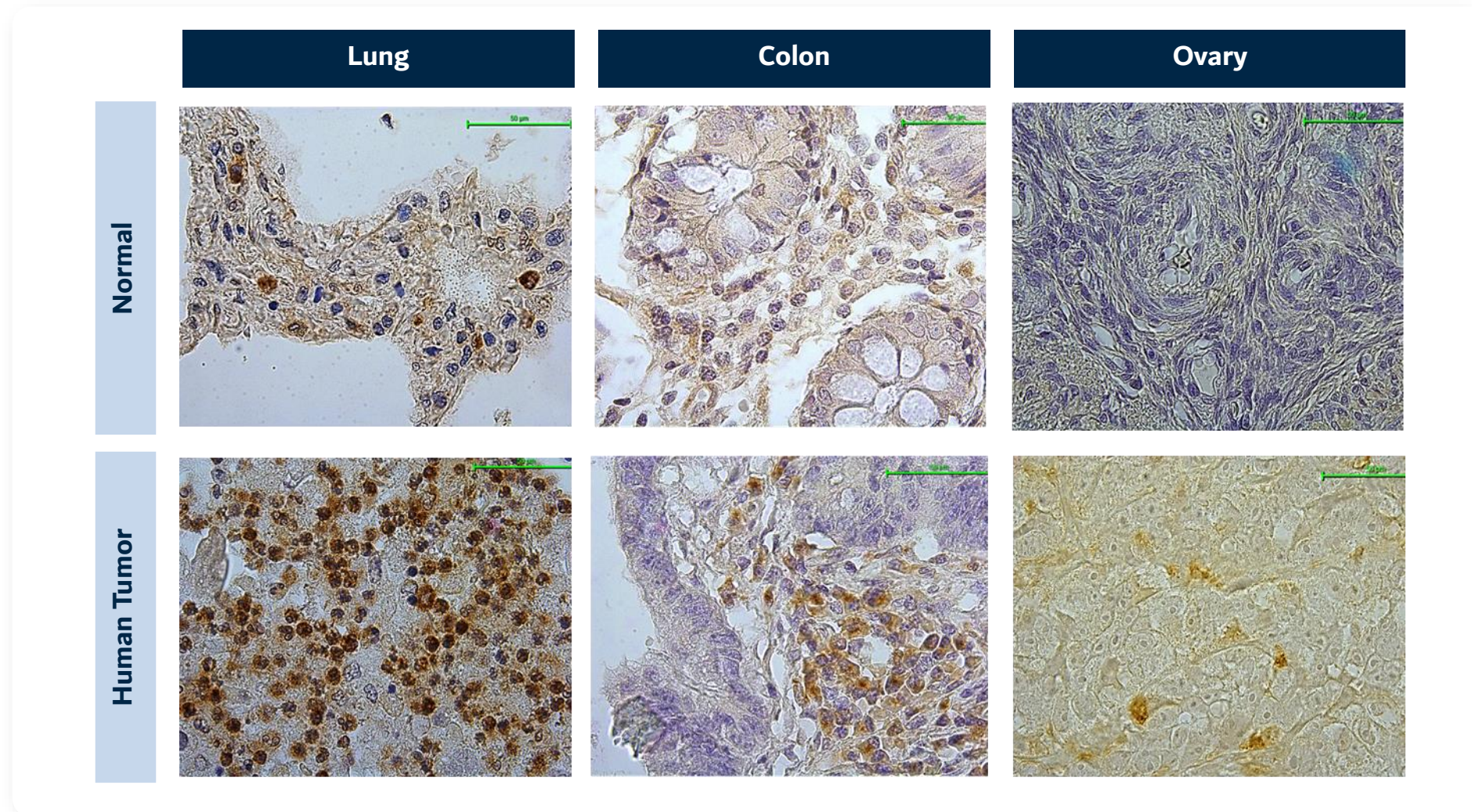
No overt clinical signs
or weight loss

**Kineta has completed multiple, single and repeat-dose toxicology studies in NHP
with doses of KVA12123 up to 100 mg/kg
(>100-fold safety margin over target human exposure)**

KVA12123: No CRS-associated signal in preclinical models in NHP toxicology studies as well as in human whole blood



Clinical applications for KVA12123 are primarily focused on solid tumors with high levels of VISTA expression



Brown staining in human tumors indicates VISTA expression

Phase 1 / 2 open-label clinical trial of KVA12123 alone and in combination with pembrolizumab in patients with advanced solid tumors

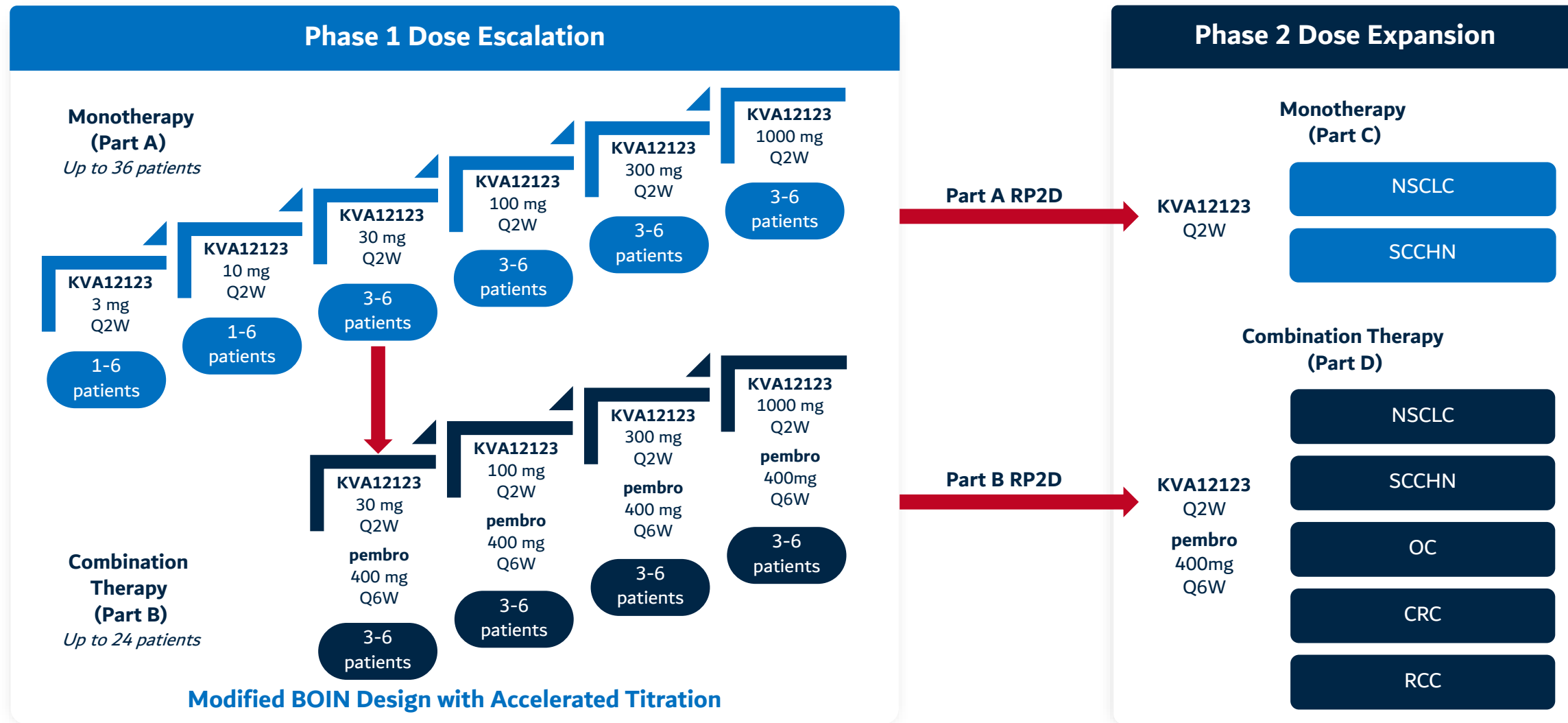
Patient population:

- Phase 1 basket trial in patients with advanced solid tumors (up to 60 patients)
- Phase 2 in NSCLC, HNSCC, OC, CRC, RCC and TBD other patients

Study objectives:

- Primary: Safety and tolerability, recommended Phase 2 dose (RP2D) or maximum tolerated dose (MTD) of KVA12123
- Secondary: Pharmacokinetics, immunogenicity, tumor response in subjects with advanced solid tumors per iRECIST (ORR)
- Exploratory: Biomarker and receptor occupancy

Phase 1 / 2 open-label clinical trial of KVA12123 alone and in combination with pembrolizumab in patients with advanced solid tumors



KVA12123

Clinical trial strategy

Clinical research sites

- Selected to provide diverse advanced solid tumor patients

Merck research collaboration

- Clinical trial collaboration and KEYTRUDA® supply agreement



Exploratory biomarkers:

- Receptor Occupancy (RO)
- Chemokine and cytokine levels in blood
- Immune cell populations in blood
- VISTA expression in tumor pre- and post-treatment

Clinical research sites Parts A & B



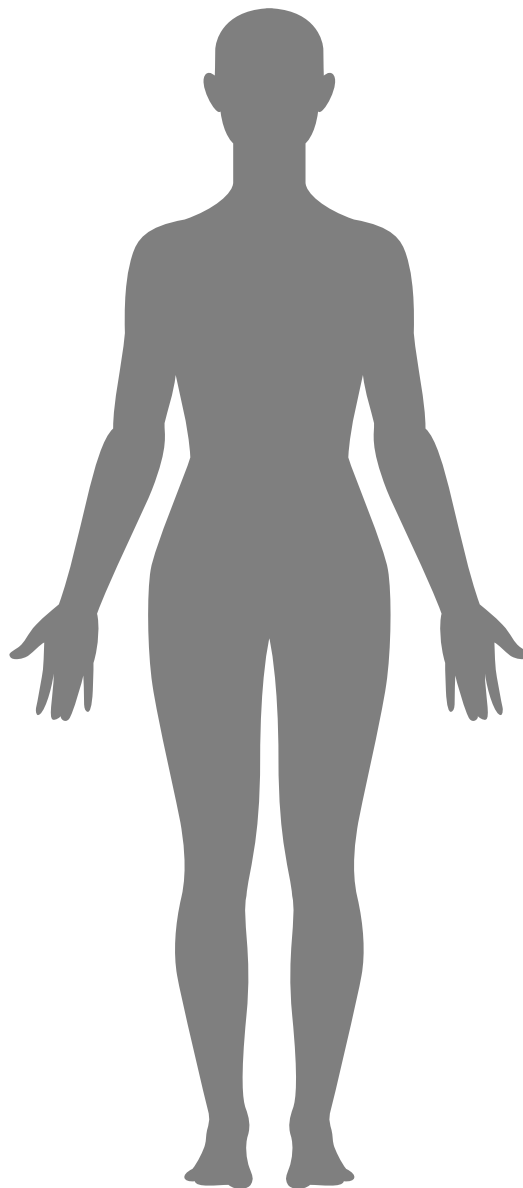
Large commercial market opportunity in initial indications in solid tumors for KVA12123

2.7M

annual new patient population

\$48B

market opportunity



NSCLC ¹

980K annual new patients

\$31.8B market



Colorectal cancer ²

1.1M annual new patients

\$10.3B market



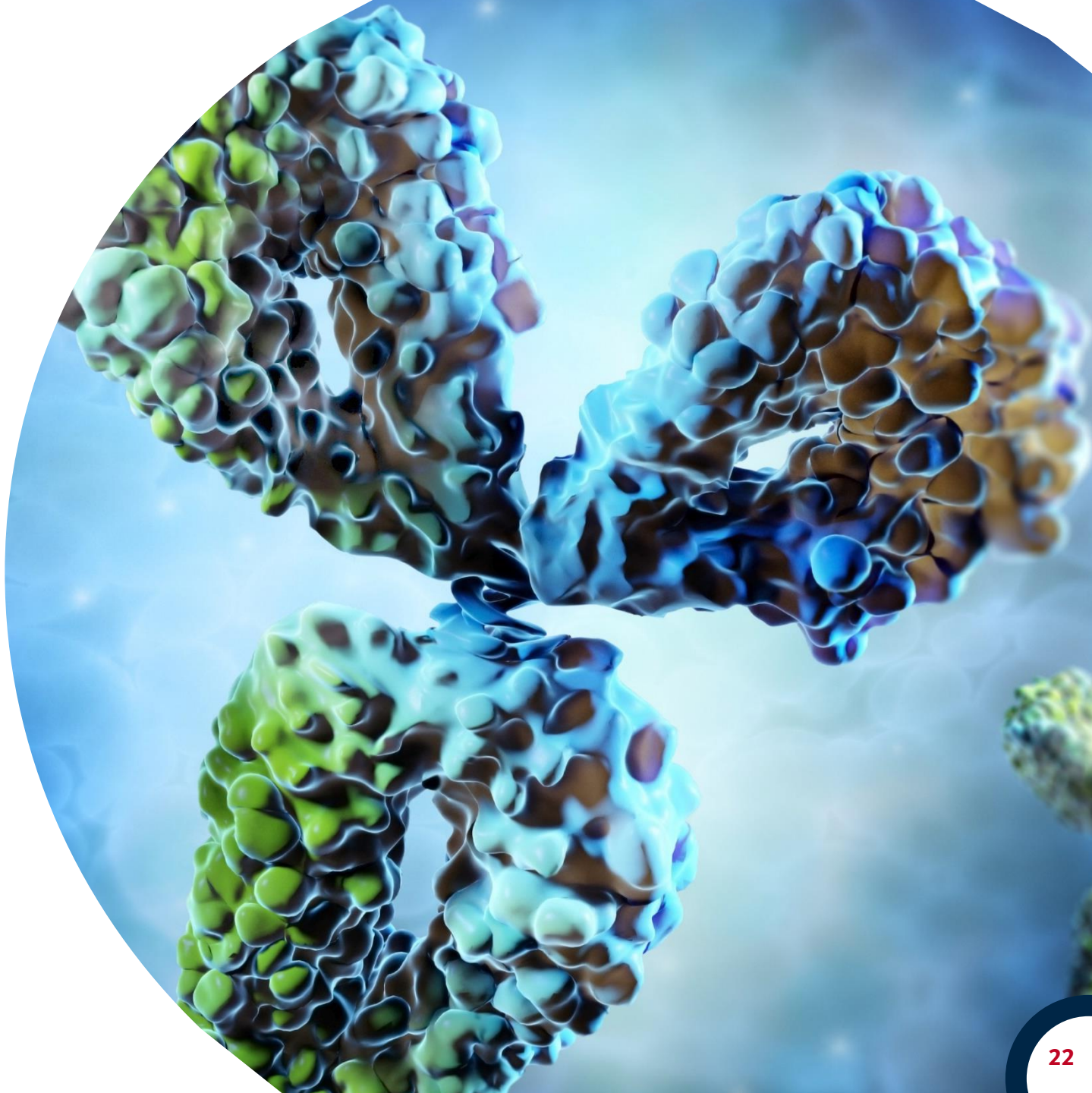
Ovarian cancer ³

660K annual new patients

\$5.9B market

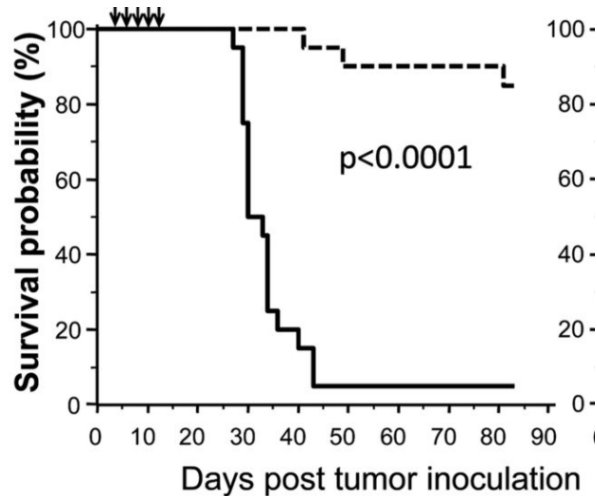


Anti-CD27 agonist
mAb immunotherapy

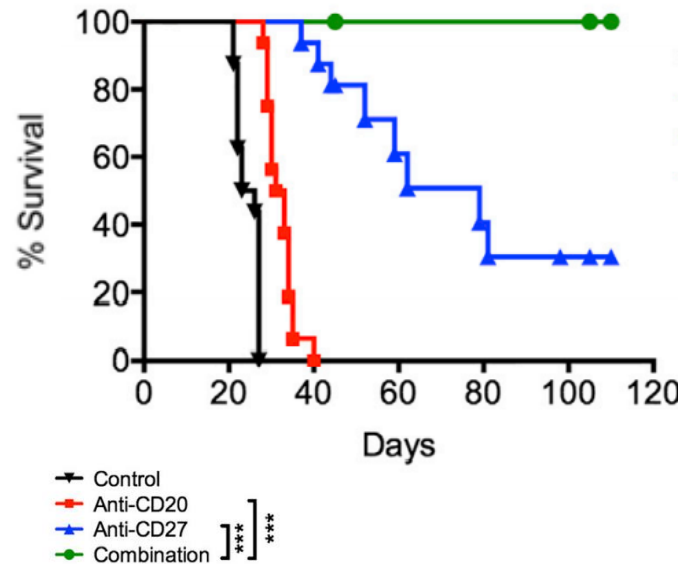


Anti-CD27 agonist antibodies can drive tumor growth inhibition as a monotherapy and in combination with CPIs

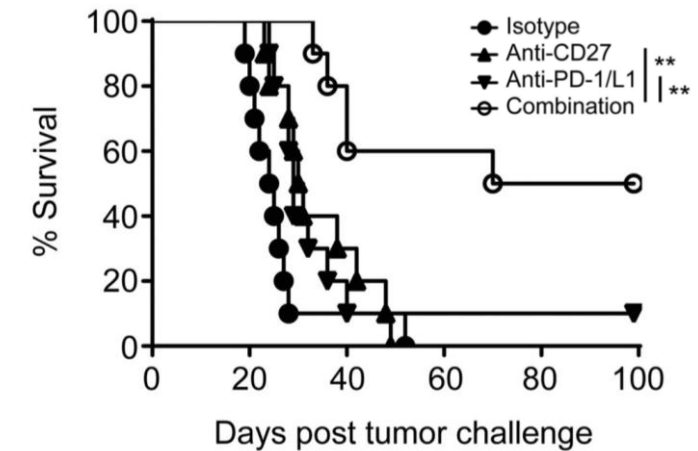
Monotherapy
CT26 Colorectal Cancer ¹



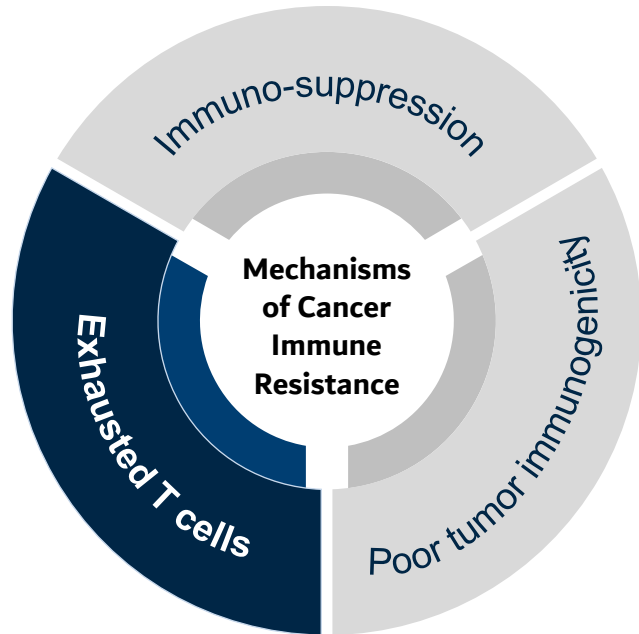
Combination Therapy
BCL-1 B cell lymphoma ²



Combination Therapy
B16-BL6 Melanoma ³



Anti-CD27 agonist to address exhausted T cell mechanism of cancer immune resistance



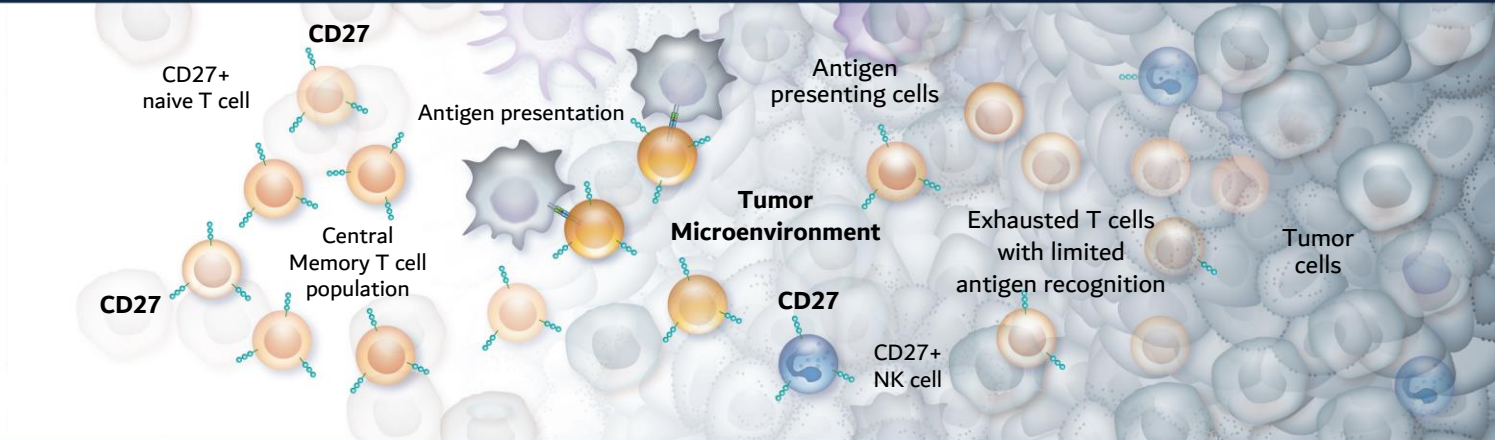
Activates and induces the maturation and migration of naïve **T cells**

Drives the **diversification of the T cell** repertoire

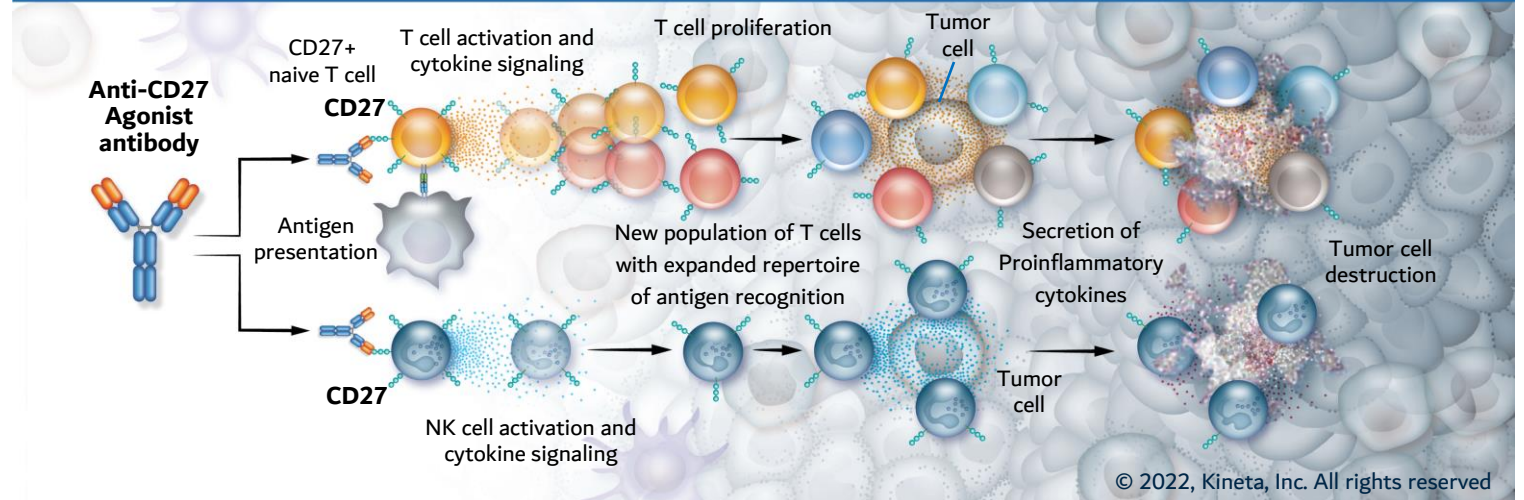
Enhances **NK cell** activation

Activates **low affinity antigens**

Exhausted T cells



CD27 agonist has the potential to generate new populations of functional anti-tumor immune cells



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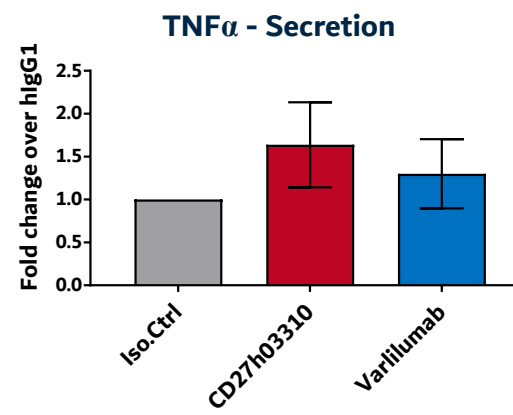
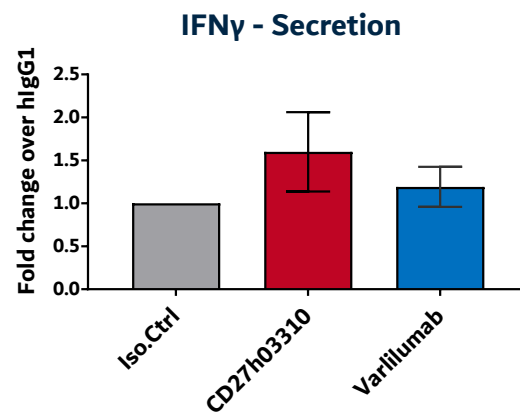
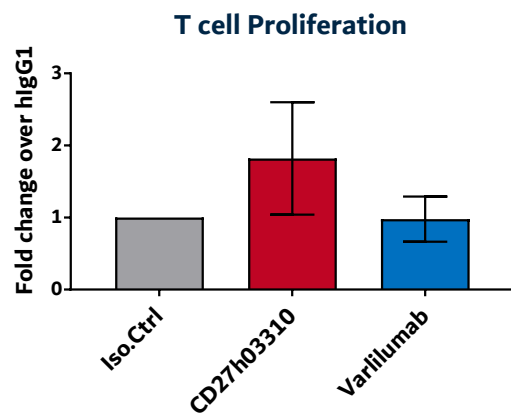
Lead anti-CD27 mAb demonstrates robust agonist activities on T and NK cells in *in vitro* studies



Increases T cell proliferation and activation



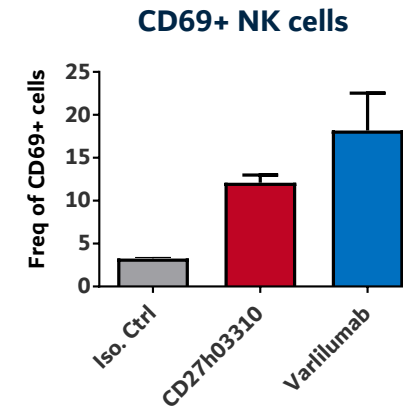
T cell



Increases NK cell activation



NK cell

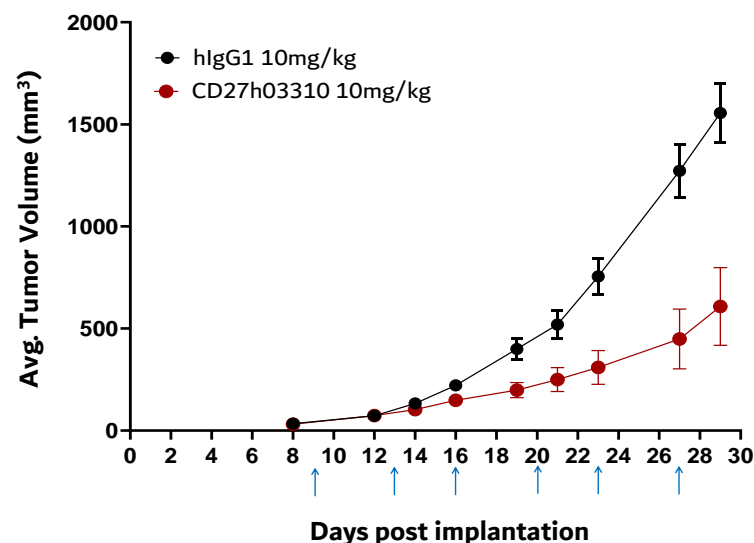


Lead anti-CD27 agonist mAb demonstrates single agent tumor growth inhibition (TGI) in preclinical models

Monotherapy

B Cell Lymphoma Model Raji
SCID mice

Mean Tumor Volume

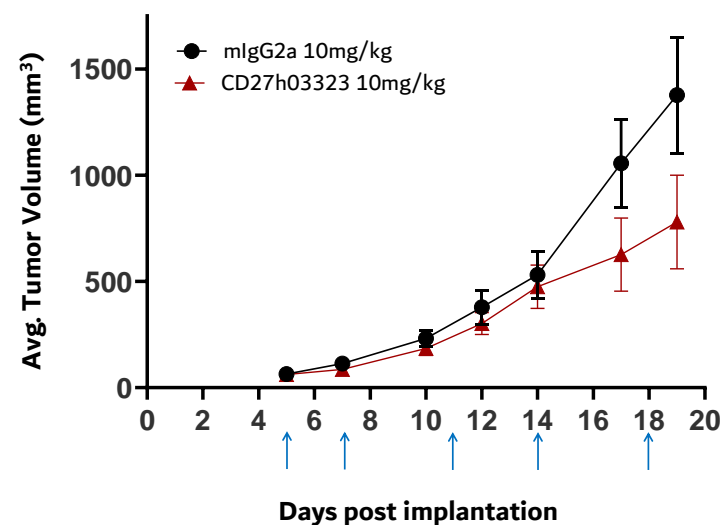


Tumor Growth Inhibition

Lead anti-CD27 mAb: **65%**

T Cell Lymphoma Model EG7
huCD27 KI mice








Mean Tumor Volume



Tumor Growth Inhibition

Lead anti-CD27 mAb: **45%**

Significant catalysts anticipated over the next 24 months

Anticipated Milestones		2023				2024	
		1Q	2Q	3Q	4Q	1H	2H
KVA12123	Opened Phase 1/2 clinical study						
	First patient dosed combination						
	Initial Phase 1 clinical safety data						
	Initial Phase 1 clinical efficacy data (ORR)						
	Initial Phase 2 clinical data						
αCD27 agonist mAb	IND filing						
	Start Phase 1 clinical study						

Experienced leadership team



Shawn Iadonato, PhD

Chief Executive Officer



Craig Philips

President



Thierry Guillaudeau, PhD

Chief Scientific Officer



Keith Baker

Chief Financial Officer



Pauline Kenny

General Counsel



Jacques Bouchy

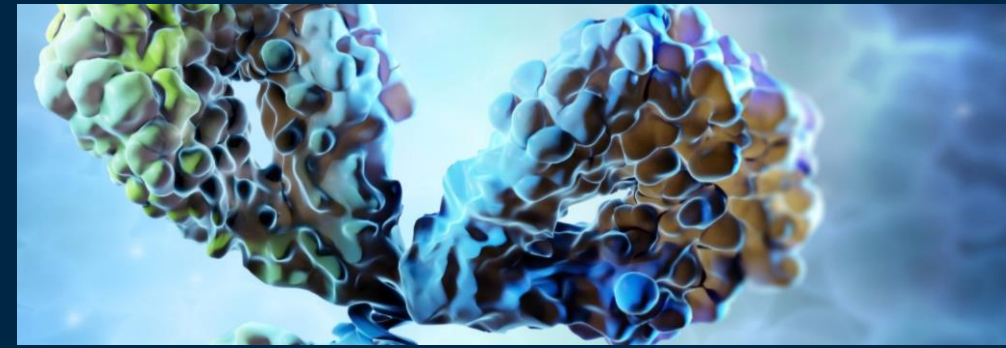
EVP Investor Relations
& Business Development



Schering-Plough



Kineta is developing next-generation immunotherapies that address cancer immune resistance



Innate Immunity Focused Pipeline

KVA12123

- VISTA blocking mAb to address immunosuppression in the TME
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Catalysts

3Q23 | KVA12123 initial clinical safety data
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Runway

Through mid-2024

Partnerships





Developing next generation
immunotherapies for cancer patients




www.kinetabio.com





Appendix

Strategic partnerships provide potential for a significant revenue stream

	Ongoing collaboration	License Agreements with no research obligations by Kineta	
Program	Neuromuscular diseases-ALS	Oncology	Cystic fibrosis
Partner		 <small>A Member of the Roche Group</small>	
Key deal terms	Up to \$530M in milestones	Up to \$96M in milestones	Up to \$965M in commercial only milestones
	Royalties on net sales	Royalties on net sales	Royalties on net sales
	Research collaboration & license agreement		Revenue share on sub-license payments