

Jan 2026

Redefining the design and discovery of therapeutic antibodies

ADAGENE

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Company Highlights

Potential best-in-class anti-CTLA-4 with blockbuster opportunity



ADG126 is a novel anti-CTLA-4 leveraging our SAFEbody® platform
Broad therapeutic Index, Top tier efficacy, and safety profile in late-stage MSS CRC with no liver metastases (NLM)
Global 3L+ CRC opportunity to exceed \$1Bn

SAFEbody® technology validated by strategic collaborations



Technology licensing agreements with: Sanofi, Exelixis, Conjugatebio and Third Arc Bio
Eligible to receive over \$3.3 billion in milestones and royalties

Pipeline candidates showcase platform versatility



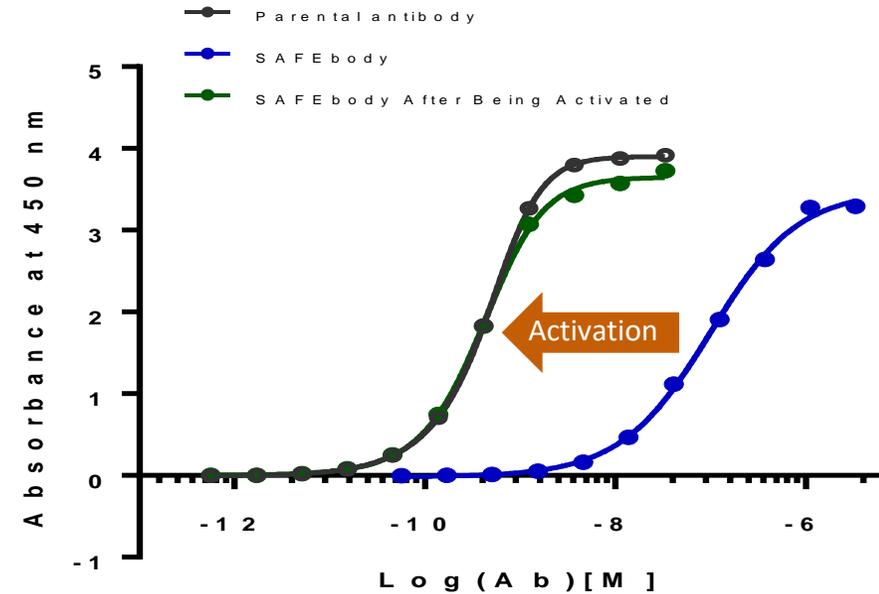
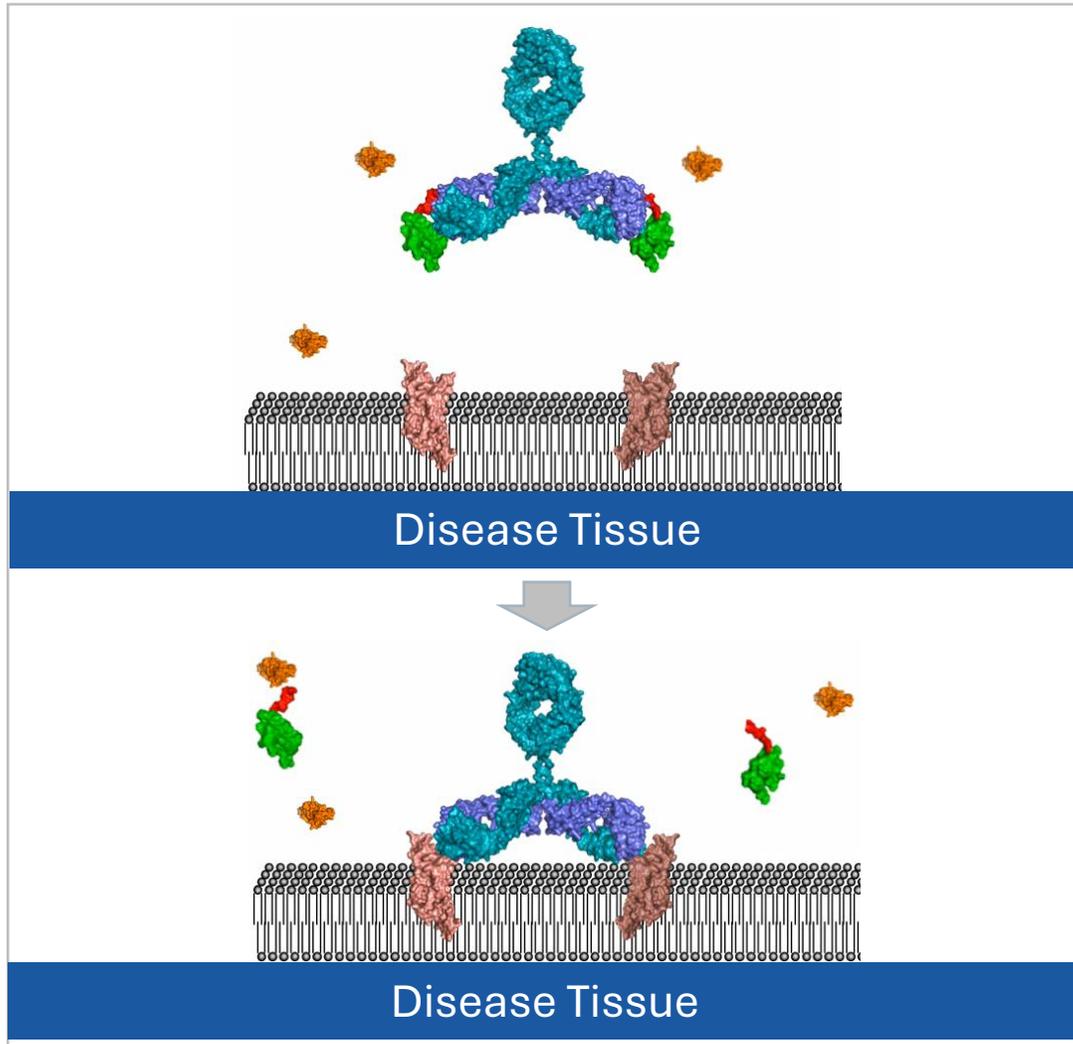
ADG138- Double masked HER2xCD3 T-cell engager
Multiple masked T-cell engagers in discovery

Strong cash balance with runway into 2027



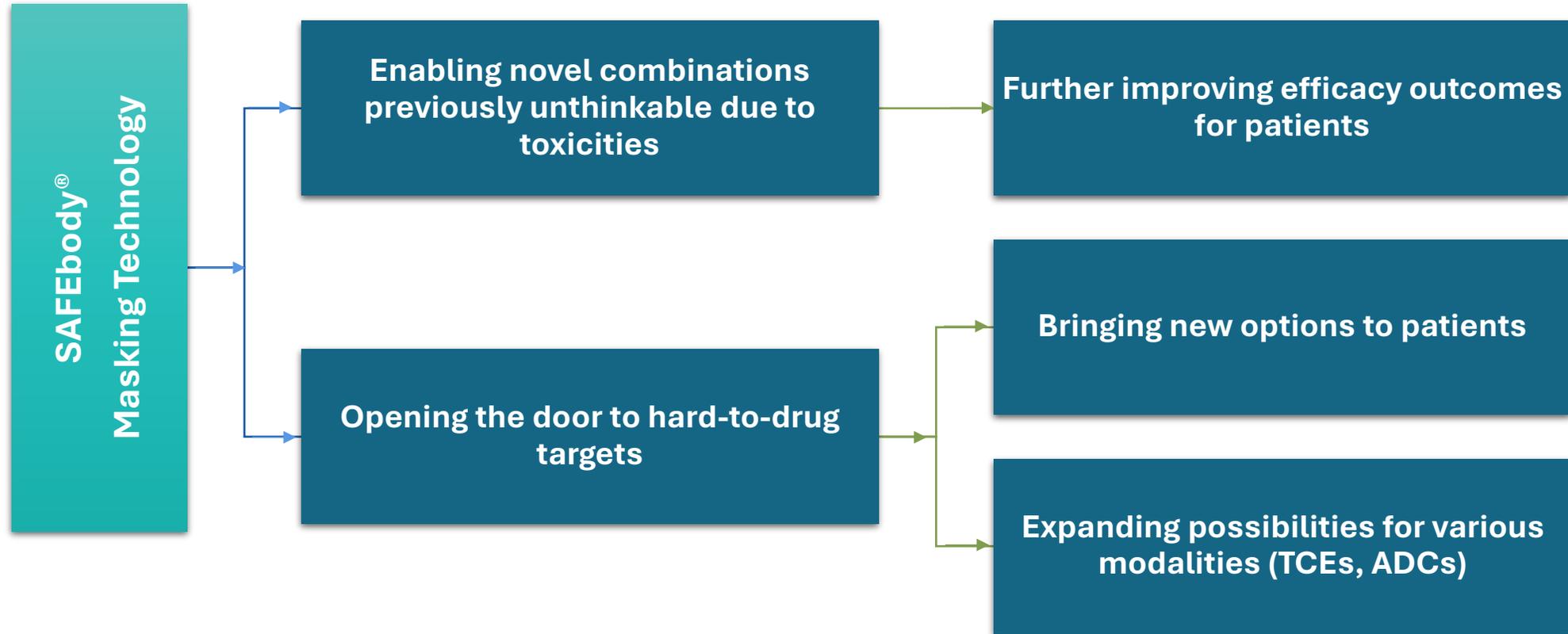
Unaudited cash and cash equivalents : US\$74.5 million as of December 31, 2025
Cash runway into late 2027

SAFEbody® : Masking Technology for Enhanced Therapeutic Index



- **Selective activation** in disease tissues (e.g., tumor microenvironment) to significantly minimize side effects
- **Validated by** robust preclinical and clinical data
- Suitable for applications in **oncology and autoimmune diseases**

SAFEbody® Can Broaden the Range of Possibilities Across Multiple Therapeutic Areas



Adagene Pipeline – Unlocking the Potential of Complex Targets

Program & Technology	Target	Development stage				
		Discovery	IND Enabling	Ph 1	Ph 2	Pivotal
ADG126 SAFEbody	CTLA-4					
ADG206 Fc engineered SAFEbody	CD137					
ADG138 TCE SAFEbody	HER2xCD3					
ADG152 TCE SAFEbody	CD20xCD3					

Strategic Investment from Sanofi in July 2025

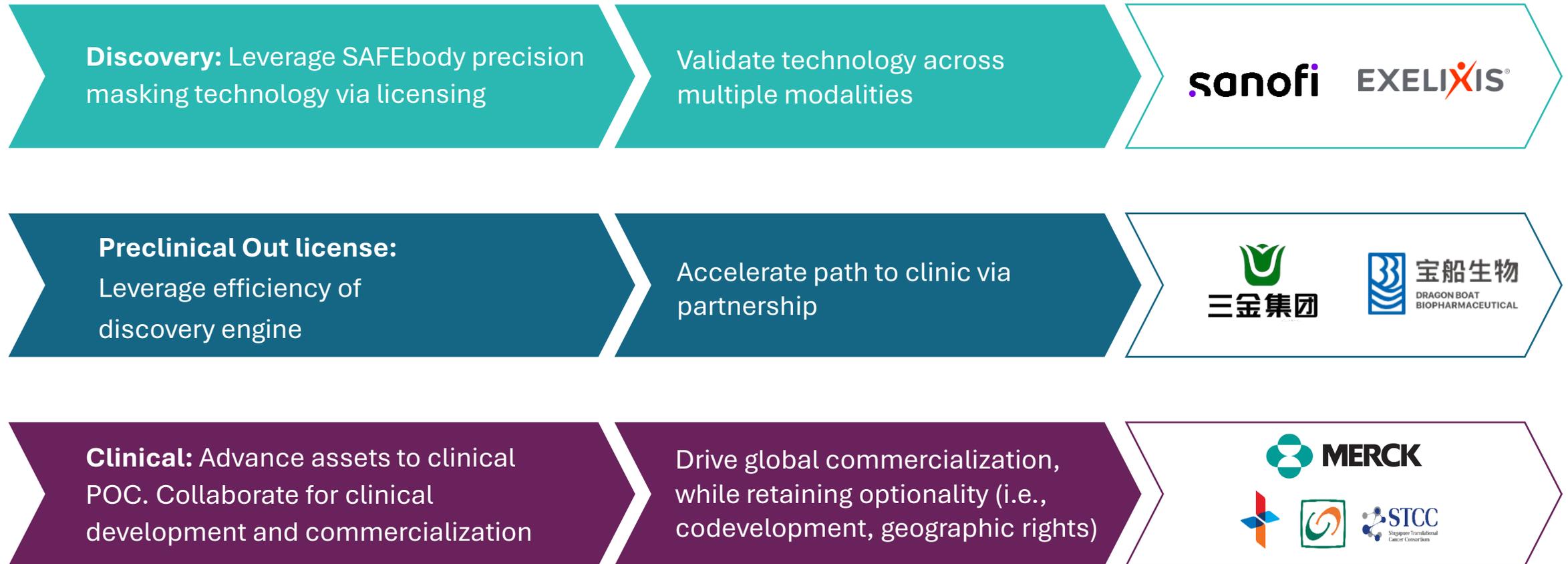
Sanofi to invest up to \$25 million to support Adagene's R&D, including a randomized phase 2 trial of anti-CTLA-4 SAFEbody, ADG126 (muzastotug), in MSS CRC.

Clinical Collaboration: Adagene to supply ADG126 for a Sanofi-sponsored phase 1/2 combo trial in advanced solid tumors, enrolling 100+ patients. Adagene continues to own worldwide commercial rights to ADG126.

Platform Collaboration: Sanofi exercised option on third SAFEbody discovery program, triggering an option exercise fee, as well as potential milestone and royalty payments.

Extended Cash Runway: Funding expected to support operations into 2027

Collaborations Provide Near Term Revenue, and Validate Our Platform



2026 Milestones

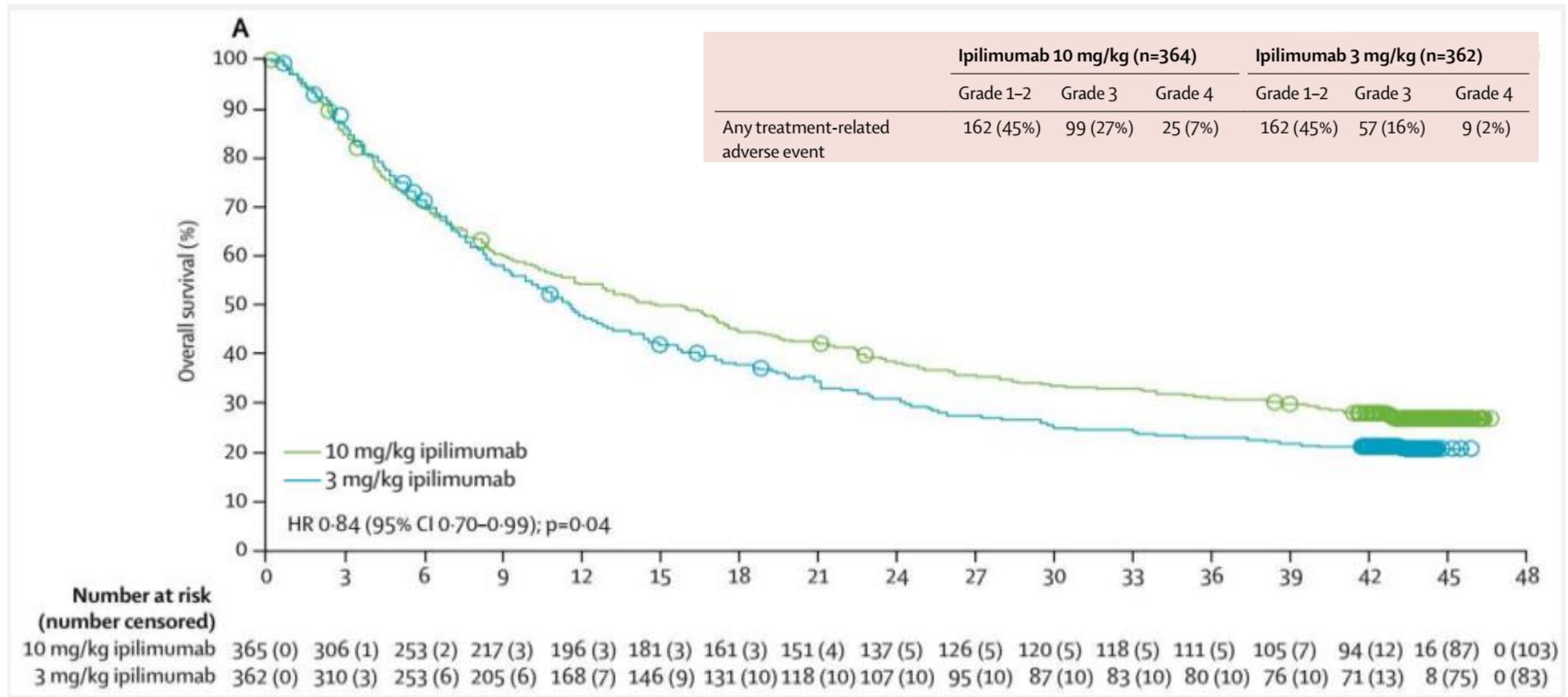
- Q1 2026: Data update from the ongoing Phase 1b/2 study of muzastotug + pembrolizumab in 3L+ MSS CRC, including 41 patients in the 10 mg/kg cohorts and 26 patients in the 20 mg/kg cohorts.
- Complete enrollment of the ongoing randomized Phase 2 dose-optimization study with muzastotug, which is being conducted in alignment with FDA Project Optimus, and designed to allow dose regimen selection for Phase 3.
- Provide preliminary clinical data, including pathological responses, to inform future development from investigator-initiated Phase 2 trial for neoadjuvant muzastotug + pembrolizumab in colorectal cancer.
- Provide initial clinical data from a new cohort of patients in the ongoing Phase 1b/2 study of muzastotug + pembrolizumab in combination with standard of care (fruquintinib) in MSS CRC patients.
- Share results of the clinical trial collaboration with Roche, which evaluates muzastotug in triplet combination with atezolizumab and bevacizumab in first-line treatment of locally advanced or metastatic hepatocellular carcinoma (HCC; liver cancer).
- Establish additional collaboration/licensing agreements.

A circular inset showing a microscopic view of cells with numerous small, clear bubbles or vesicles scattered throughout. The background is a dark, reddish-purple hue. The text is centered over this image.

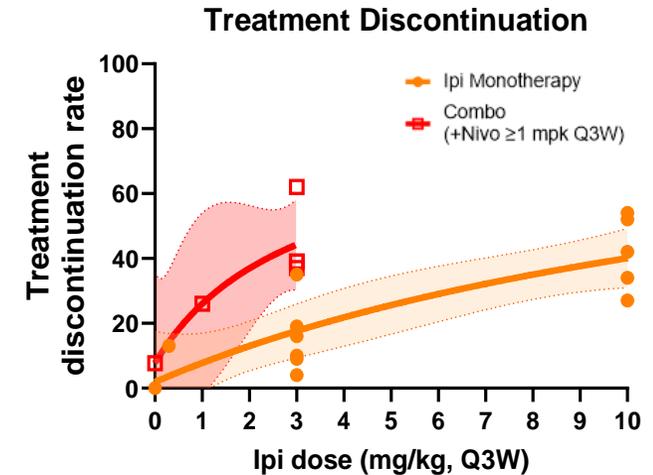
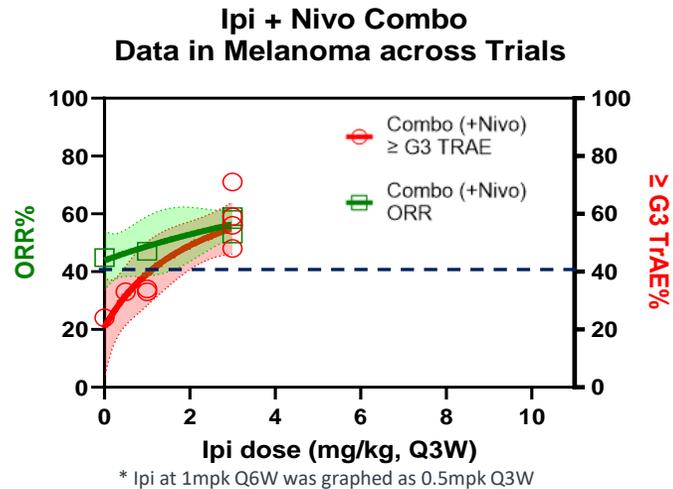
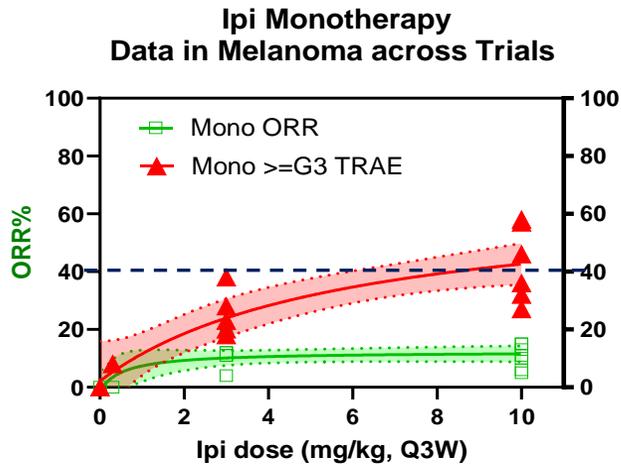
ADG126:

A New Paradigm for CTLA-4

The First Generation of Anti-CTLA-4 Agents Has Demonstrated Dose-Dependent Toxicity and Efficacy



The Increased Ipilimumab Resulted in A Disproportionate Increase in Toxicity With Limited Improvement in Response Rates



- ✓ Efficacy outcomes with ipilimumab were significantly improved when combined with anti-PD-1 (nivolumab).
- ✓ However, the incidence of severe adverse events leading to discontinuations also rose substantially, even with a threefold reduction in the dose of ipilimumab

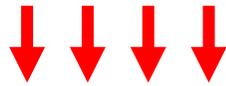
- ✓ Combination tox is similar at >3-fold lower Ipi dose to mono
- ✓ The discontinuation rate remains low for EXP cohorts (**6%**) for **ADG126 + Pembro** in MSS CRC (01/15/2025 data cut across all studied cohorts).

Restoring Anti-Tumor Immunity With Anti-CTLA-4 is A Clinically and Commercially Validated Strategy for Solid Tumors

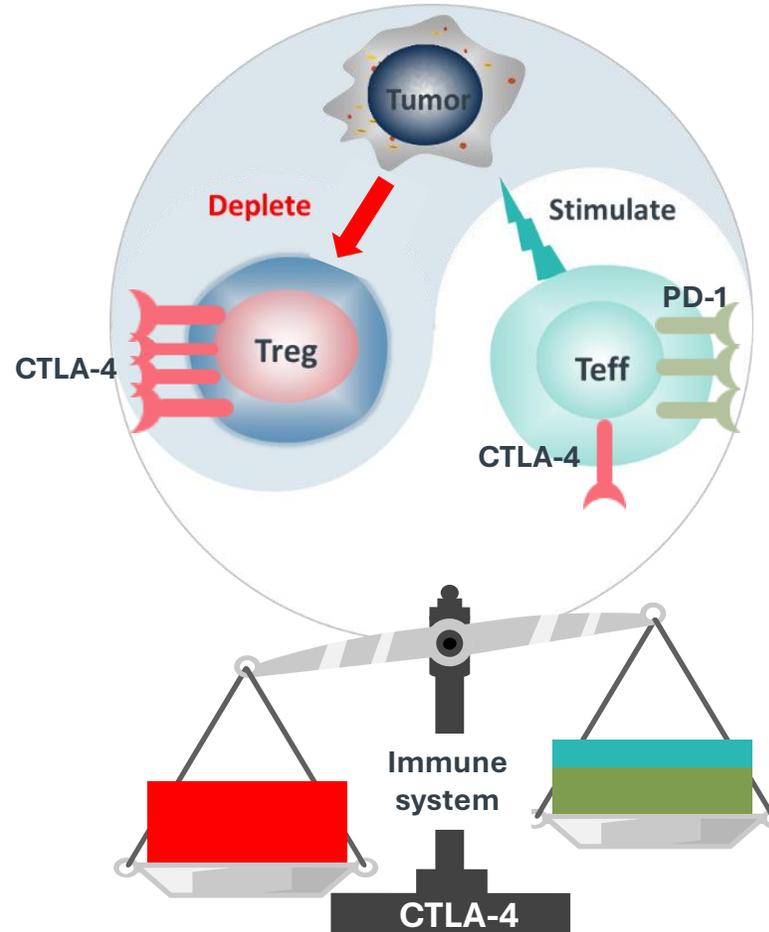
T Regulatory (Treg) Cells

Deplete CTLA-4-mediated Treg cells in tumors

Treg (CTLA-4++++)



- Reducing immune suppression by depleting CTLA-4 mediated Tregs in TME



T Effector (Teff) Cells

Soft prime CTLA-4-mediated T cell
Reinvigorate tumor exhausted CD8+ Cells



Teff (PD-1+++)



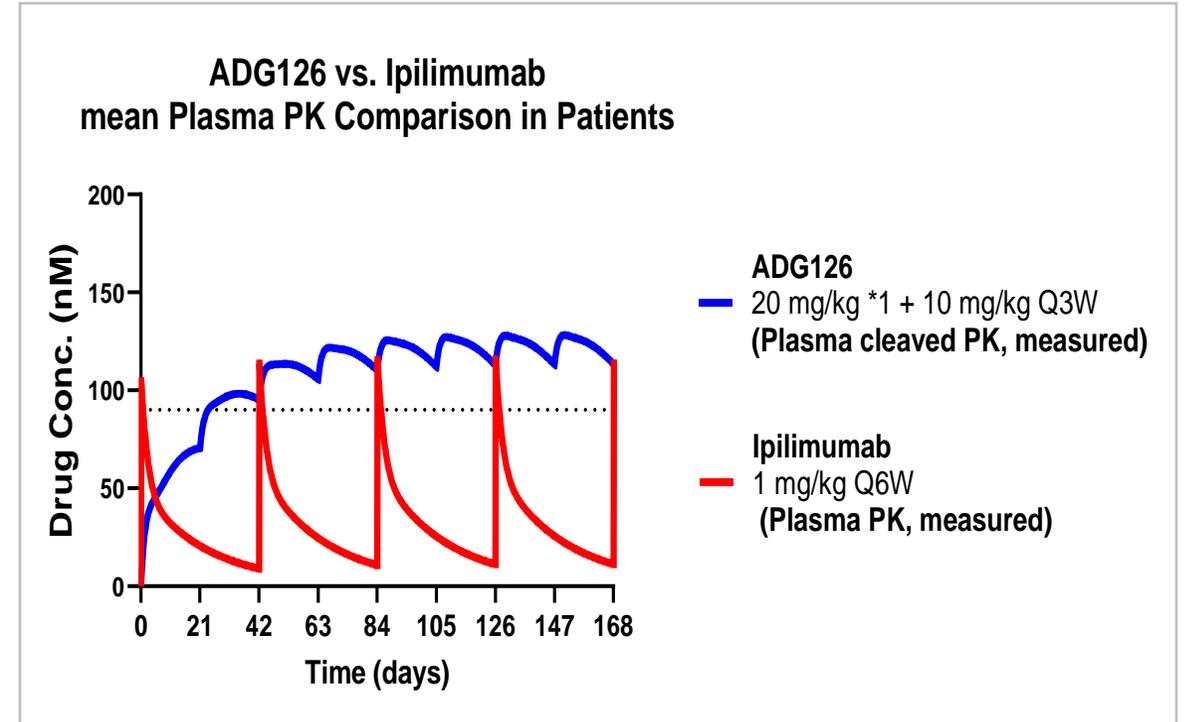
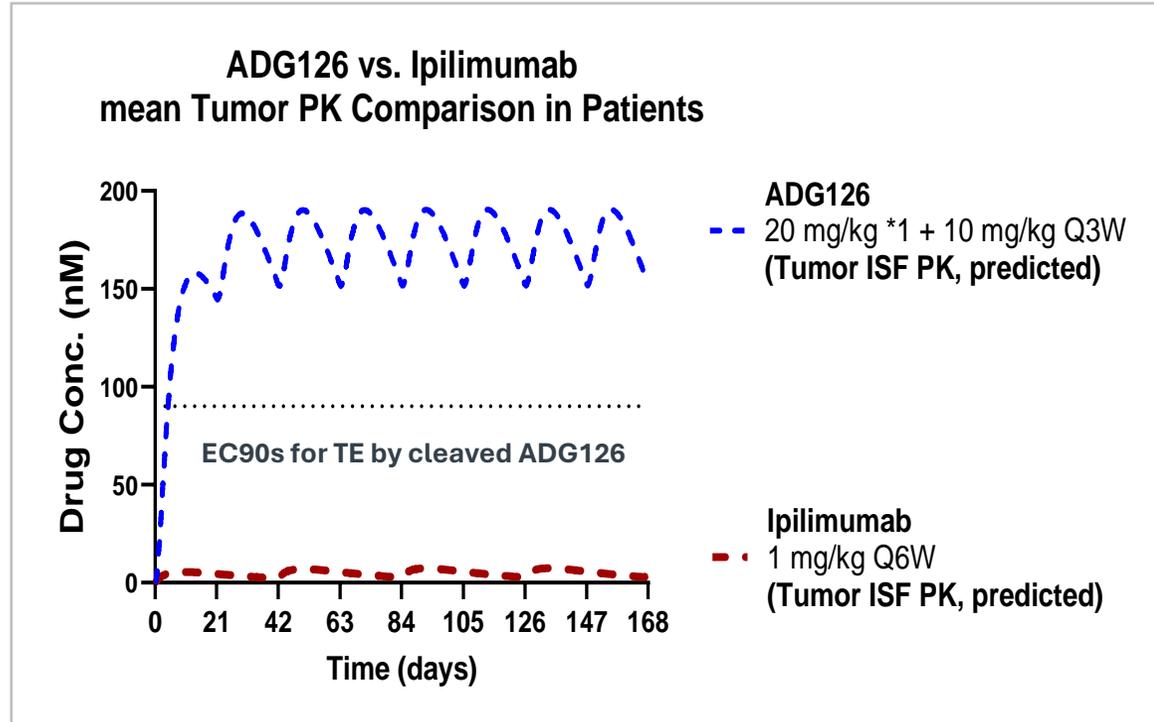
Teff (CTLA-4+)

- Increasing the exposure of activated ADG126 on T effector cells in TME while reducing its exposure in peripheral blood

- Two anti-CTLA-4 drugs have been approved by the FDA in several solid tumor indications: Yervoy (ipilimumab) and Imjudo (tremelimumab)

Enhanced Therapeutic Index Resulting From Increase of Cleaved ADG126 Over Ipilimumab in Tumor vs Blood

At 20mpk single LD + 10mpk Q3W regimen for ADG126, the predicted mean tumor active drug conc. is > 25-fold of Ipi at 1 mpk Q6W (range: 26-70X)



- Adagene will evaluate neoadjuvant high-dose ADG126 + KEYTRUDA® in up to 20 patients (Apr 2025 enrollment) with Major Pathologic Response as primary endpoint. ([NCT06846268](#), Led by Singapore's National University Cancer Institute)
- Matched biopsies to evaluate effects on tumor immune profile and tissue pharmacokinetics.

Anti-CTLA-4 Can Tackle Key Aspects of the Immune-Suppressive Tumor Microenvironment of MSS CRC

Tregs (Regulatory T Cells): High

Tregs exhibit high levels of CTLA-4 and are therefore significantly depleted by Fc active anti-CTLA-4

TILs (Tumor-Infiltrating Lymphocytes): Minimal

Anti-CTLA-4 improves the priming of naive T cells and enhances the activation and proliferation of T effector cells

PD-L1 Expression Levels: Negative or Low

Single-agent PD-1 inhibition is substantially less effective in this setting

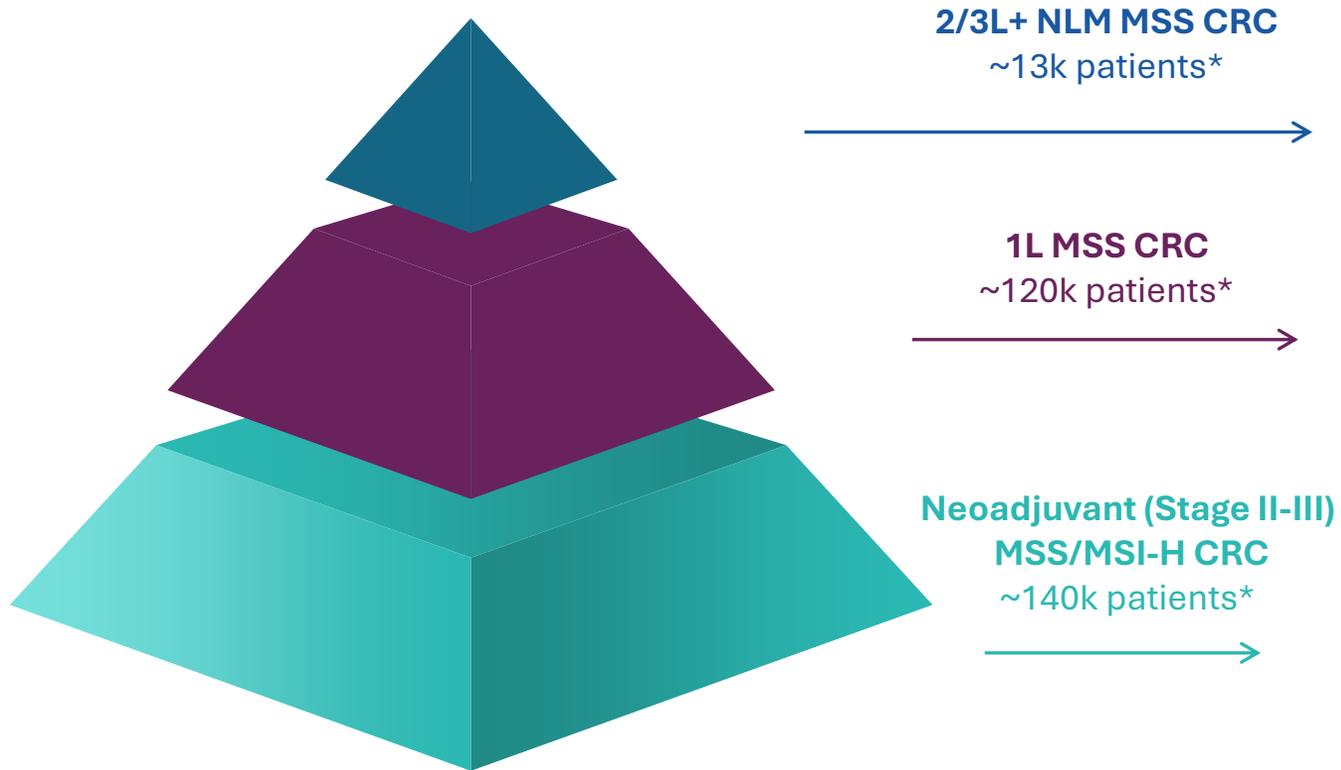
Clinical data suggest that the addition of anti-CTLA-4 agents to PD-1-based regimens can be beneficial in tumors with low PD-L1 expression (see Checkmate-067)

ADG126 + Pembro: Higher Response Rates With Lower Discontinuations Than Standard of Care Treatments

ADG126-P001 study (data from 2025 ASCO)

Compounds	Sunlight TAS102 plus Avastin	Fruquintinib		ADG126 Dose Level + Pembro 200mg Q3W	
		w/o Liver mets	with Liver mets	10 mg/kg Q3W	20 mg/kg Q6W
ORR (%)	6.1	4.3	4.9	17	29
≥G3 TRAEs	72%	36-46%		20%	17%
Discontinuation Rate	13%	15-20%		10	0

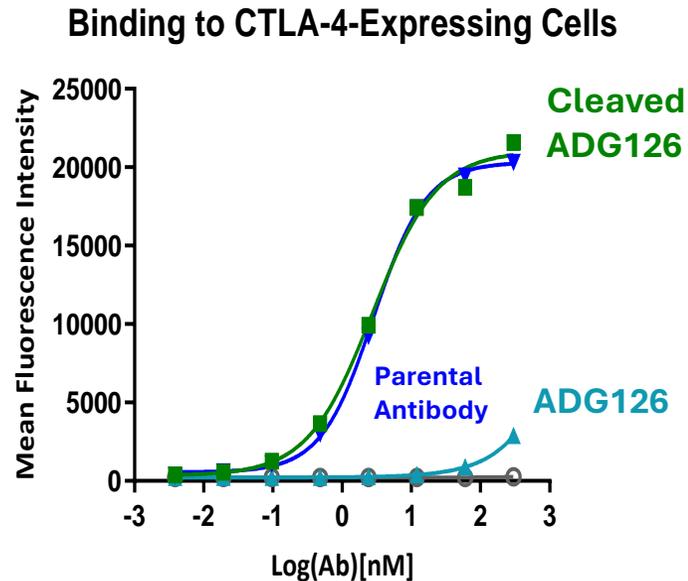
3L+ MSS CRC Without Liver Metastases is a Substantial Initial Opportunity



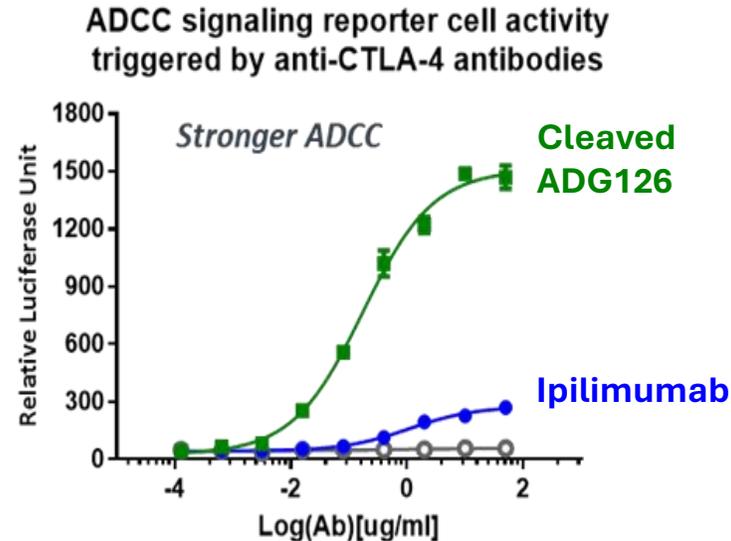
- Patients with no liver metastases represent up to 25-35% of all MSS CRC patients
- Regulatory precedent for Phase 3 trial in MSS CRC with no liver metastases
- Thanks to its safety profile, ADG126/pembrolizumab could be added to the current standard of care (e.g., TAS-102 and bevacizumab)

ADG126: A Novel Anti-CTLA-4 Incorporating a Protease Cleavable Mask, and Offering a Broad Therapeutic Index

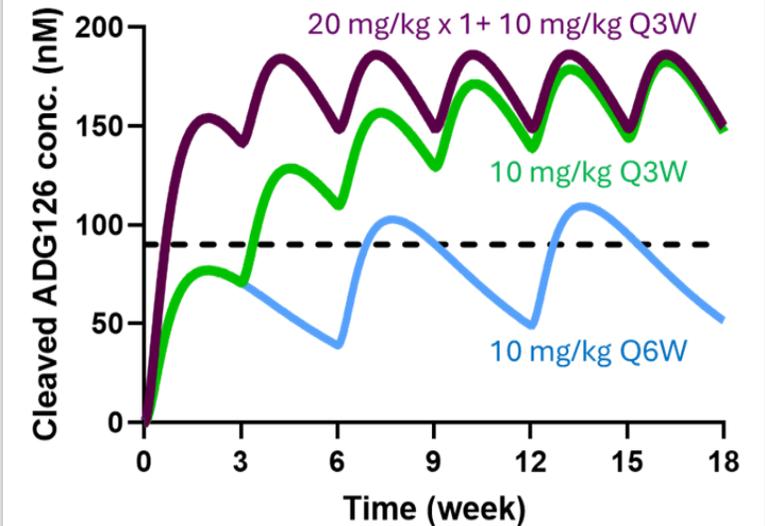
Protease cleavable mask enabling tumor-specific activation



Enhanced ADCC thanks to a differentiated epitope



Dose/Efficacy/Safety



Safety and Efficacy of ADG126 (an Anti-CTLA-4 Masking Antibody) in Combination with Pembrolizumab: Updated Results of Phase 1b/2 Study in Advanced MSS CRC

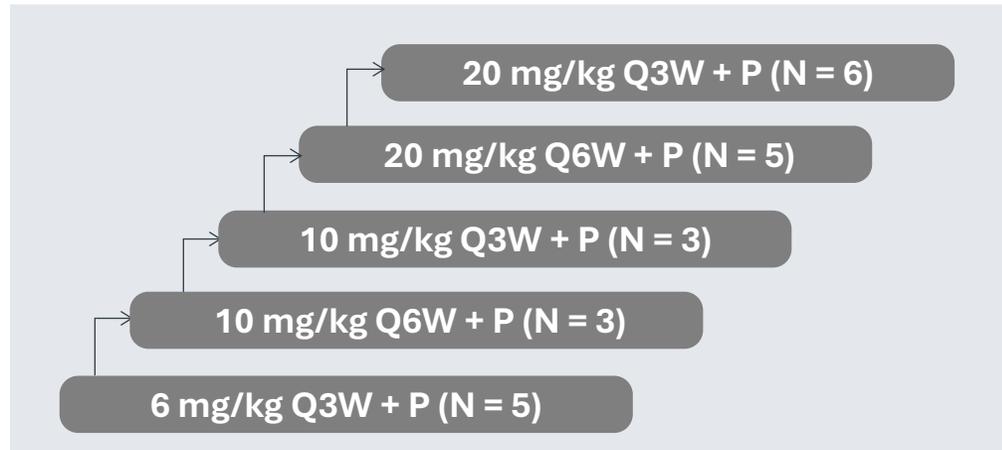
Daneng Li^{1*}, Sun Young Kim^{2*}, Manish R. Patel³, Hee Kyung Kim⁴, Sunil Sharma⁵, Sang Joon Shin⁶, Jeeyun Lee⁷, Sae Won Han⁸, George Lau⁹, Brigitte Ma¹⁰, Yan Li¹¹, Songmao Zheng¹¹, Luke Chung¹¹, Ping Xiao¹¹, Kristine She¹¹, Qinghai Zhao¹¹, Dana Hu-Lowe¹¹, Stanley Frankel¹¹, Michael J. Chisamore¹², Peter Luo¹¹, Jiping Zha¹¹, Marwan Fakih¹

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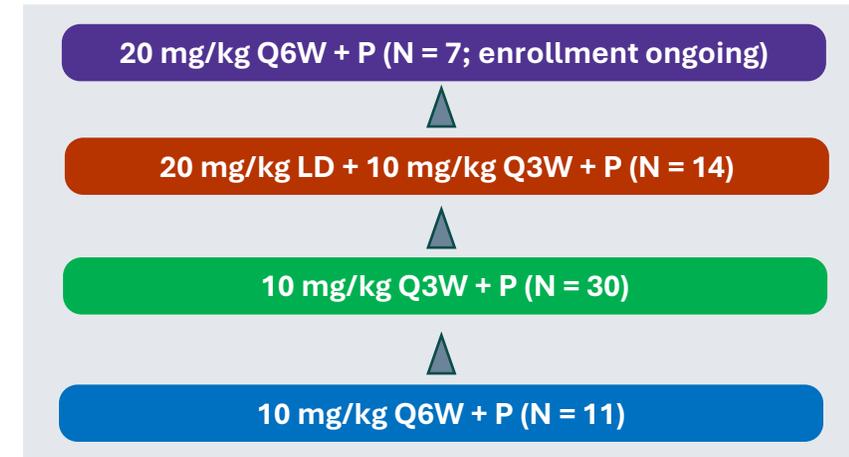
Methods and Study Design Schema: ADG126-P001 Study

- This is a Phase 1b/2, open-label, multicenter dose escalation and expansion combination study of ADG126 + pembrolizumab (200 mg, Q3W) in advanced solid tumors. The study design schema for the dose escalation (DE) and dose expansion (EXP) MSS CRC cohorts is shown below:

Dose Escalation (ADG126 Dose Level)



Dose Expansion (EXP) in MSS CRC*



MSS CRC Patients' Characteristics

- As of April 22, 2025, 97 Pts have been treated in ADG126-P001: 22 Pts in dose escalation (all comers) and 75 Pts in dose expansion (primarily MSS CRC).
- 67 Pts (5 from DE, 62 from EXP) are metastatic MSS CRC.

Baseline Characteristics of MSS CRC Patients

Characteristics	N=67
Median Age (Years), (range)	58 (24-75)
Female, n(%)	35 (52)
Race, n(%)	
Asian, (n%)	46 (69)
White, n(%)	21 (31)
ECOG 0/1, n(%)	24 (36%)/43 (64%)
Prior line of therapy \geq 3, n(%)	23 (34)
Prior immunotherapy, n(%)	0
Demographics	
US, n	24
SK, n	36
CHN/HK, n	7 (CHN=5; HK=2)
Without Liver Metastasis (NLM), n(%)	66 (99)
Peritoneal involvement, n(%)	15 (22)

ADG126 Has Been Tested Across Various Doses and Schedules with pembrolizumab, Showing Broad Safety Margins

Summary of TRAEs from ADG126 + Pembro Treatments in MSS CRC Patients

Dose levels (mg/kg)	N	All G n (%)	G1 n (%)	G2 n (%)	G3 n (%)	Discont. Rate (%)
All	67	55 (82)	20 (30)	22 (33)	13 (19)	3 (4)
10 mg/kg Cohorts	41	34 (83)	10 (24)	18 (44)	6 (15)	3 (7)
10 mg/kg Q6W	11	8 (73)	2 (18)	6 (55)	0	0
10 mg/kg Q3W	30	26 (87)	8 (27)	12 (40)	6 (20)	3 (10%)
20 mg/kg Cohorts	26	21 (81)	10 (38)	4 (15)	7 (27)	0
20 mg/kg Q6W	12	10 (83)	6 (50)	2 (17)	2 (17)	0
20 mg/kg x1 +10 mg/kg Q3W	14	11 (79)	4 (29)	2 (14)	5 (36)	0

- **Overall:** No dose-limiting toxicities (DLT) or G4/5 TRAEs; low discontinuation rate (4%).
- **10 mg/kg Q3W:** Average follow-up time of 13.2 months; manageable safety profile consistent with previous reports.
- **20 mg/kg Q6W:** Lower G3 TRAE% than that from 20 mg/kg x1 +10 mg/kg Q3W cohort.
- **20 mg/kg x1 +10 mg/kg Q3W:** Manageable AE/safety profile. Infrequent use of infliximab.

ADG126 plus Pembrolizumab – Summary of Adverse Events

Summary of $\geq 10\%$ TRAEs from ADG126 + Pembro Treatments in MSS CRC Patients (N=67)

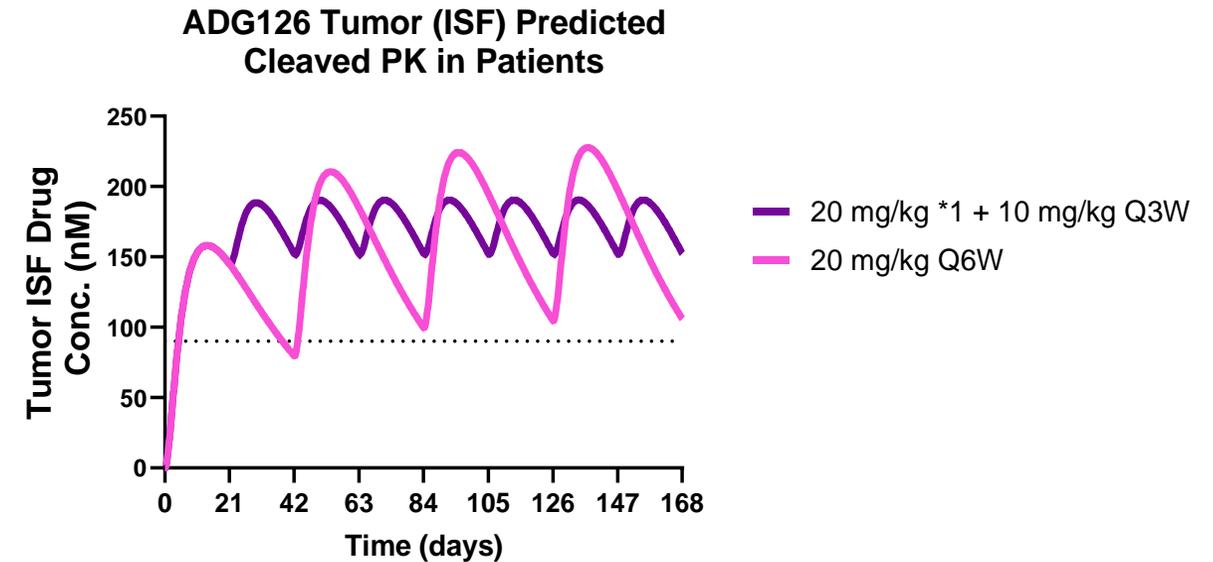
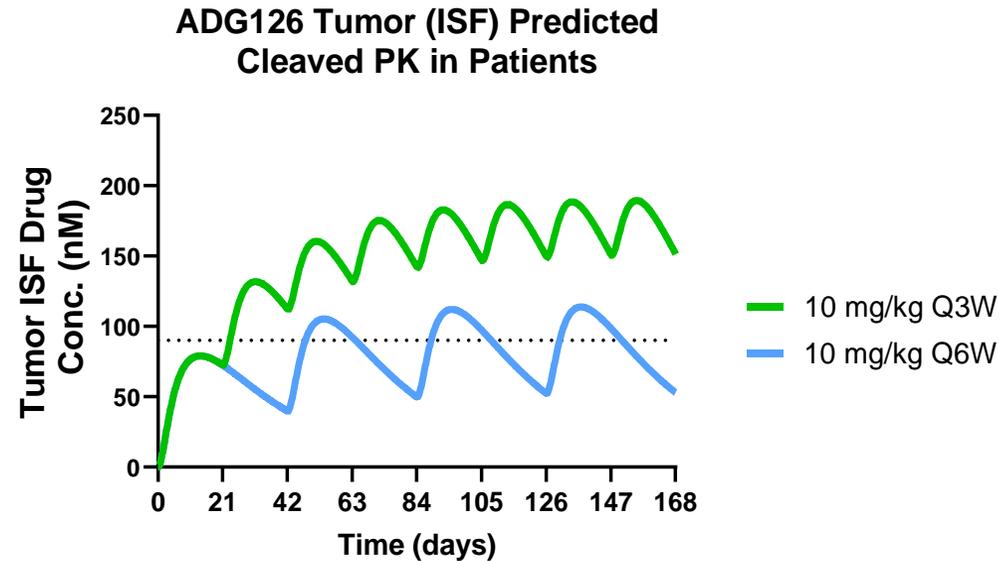
Preferred Term ($\geq 10\%$)	All G n (%)	G1 n (%)	G2 n (%)	G3 n (%)
Any TRAE	55 (82.1)	20 (29.9)	22 (32.8)	13 (19.4)
Pruritus	23 (34.3)	19 (28.4)	4 (6)	0
Diarrhea	9 (13.4)	3 (4.5)	4 (6)	2 (3)
Hypothyroidism	9 (13.4)	3 (4.5)	6 (9)	0
Fatigue	8 (11.9)	6 (9)	2 (3)	0
Adrenal insufficiency	7 (10.4)	2 (3)	5 (7.5)	0

ADG126/Pembrolizumab Has Demonstrated a Robust Dose-Dependent Efficacy

Clinical Activity Parameters of MSS CRC Cohorts (NLM)

ADG126 Dose Level + Pembro 200mg Q3W	10 mg/kg	20 mg/kg
	Q3W (N=29)	Combined (N=21)
ORR% (95% CI)	17 (6-36)	29 (11-52)
BoR, N (%)		
PR	5 ^a (17)	6 ^b (29)
SD	17 (59)	11 (52)
DCR (CR+PR+SD)%, (95% CI)	76 (56-90)	81 (58-95)
6-month CBR%, (95% CI)	38 (21-58)	Data not mature
Median PFS, months (95%CI)	4.8 (2.6-6.7)	NR (2.7-NA)
6-month PFS%, (95% CI)	39.5 (21.8-56.7)	50.4 (20.7-74.2)
Median DoR, months (95%CI)	6.2 (4.2-NA)	NR (6 PRs ongoing)

Our Pharmacokinetic Model Indicated That The 20mg Cohorts Should Reach Optimal Tumor Concentrations of Active ADG126 More Rapidly in the TME

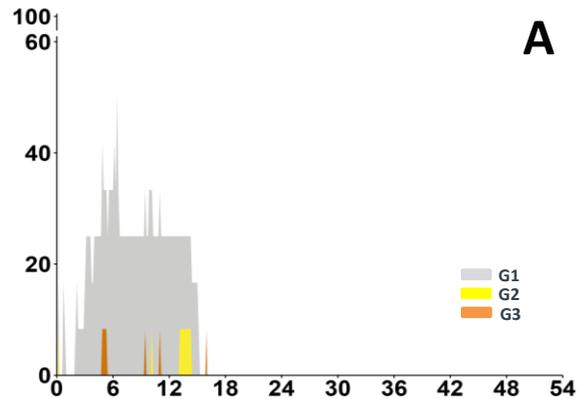


Black dashed line = Upper bound of in vitro human T cell binding EC_{90s} by cleaved ADG126 (e.g., target efficacious concentration within TME)

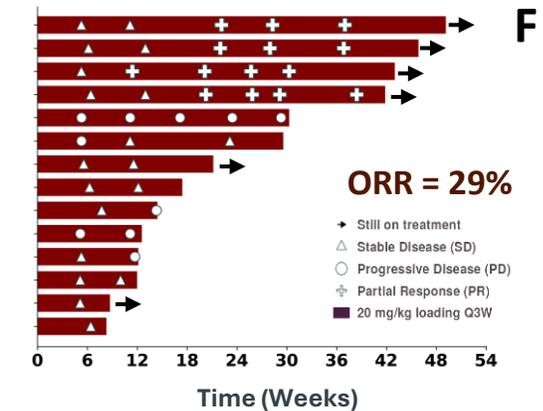
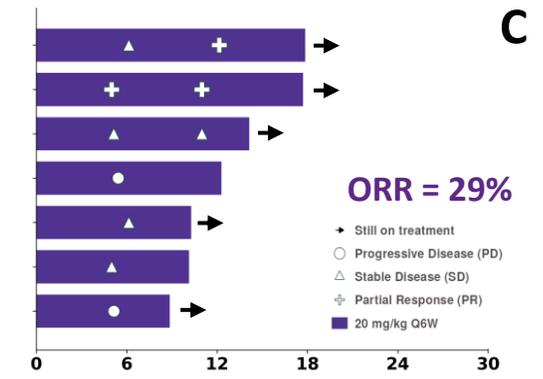
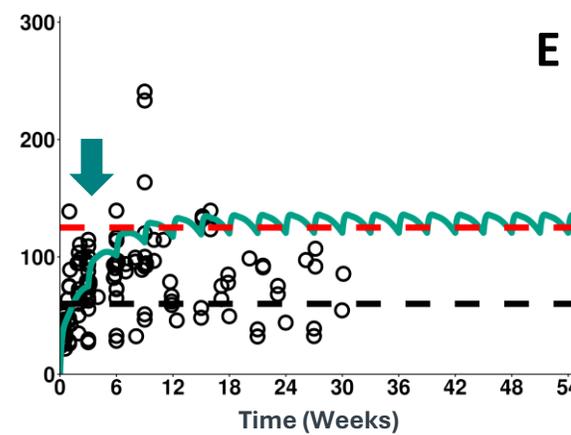
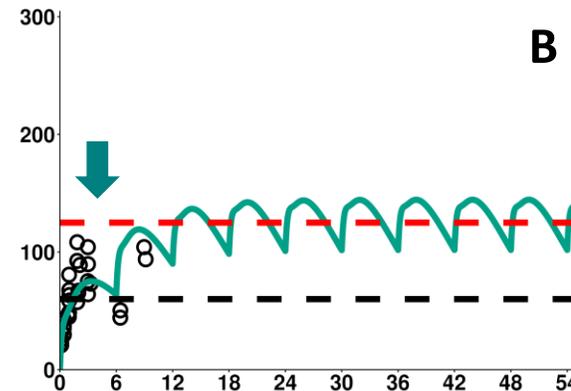
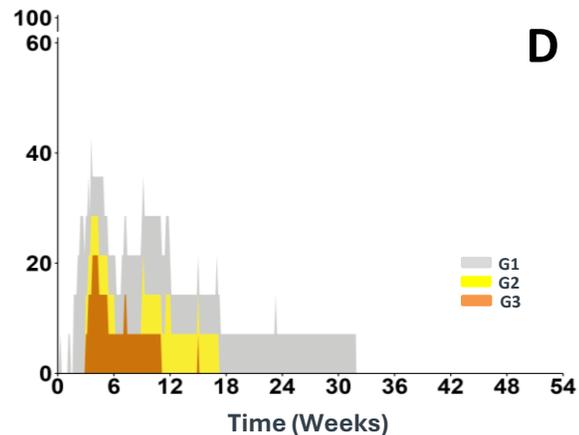
The 20 mg/kg Cohorts Achieved a confirmed ORR of 29% With A Manageable Safety Profile

Safety Kinetics, Plasma Cleaved ADG126 and ORR of 20 mg/kg Cohorts

**ADG126
20 mg/kg Q6W
(N=12; 7 from
EXP)**

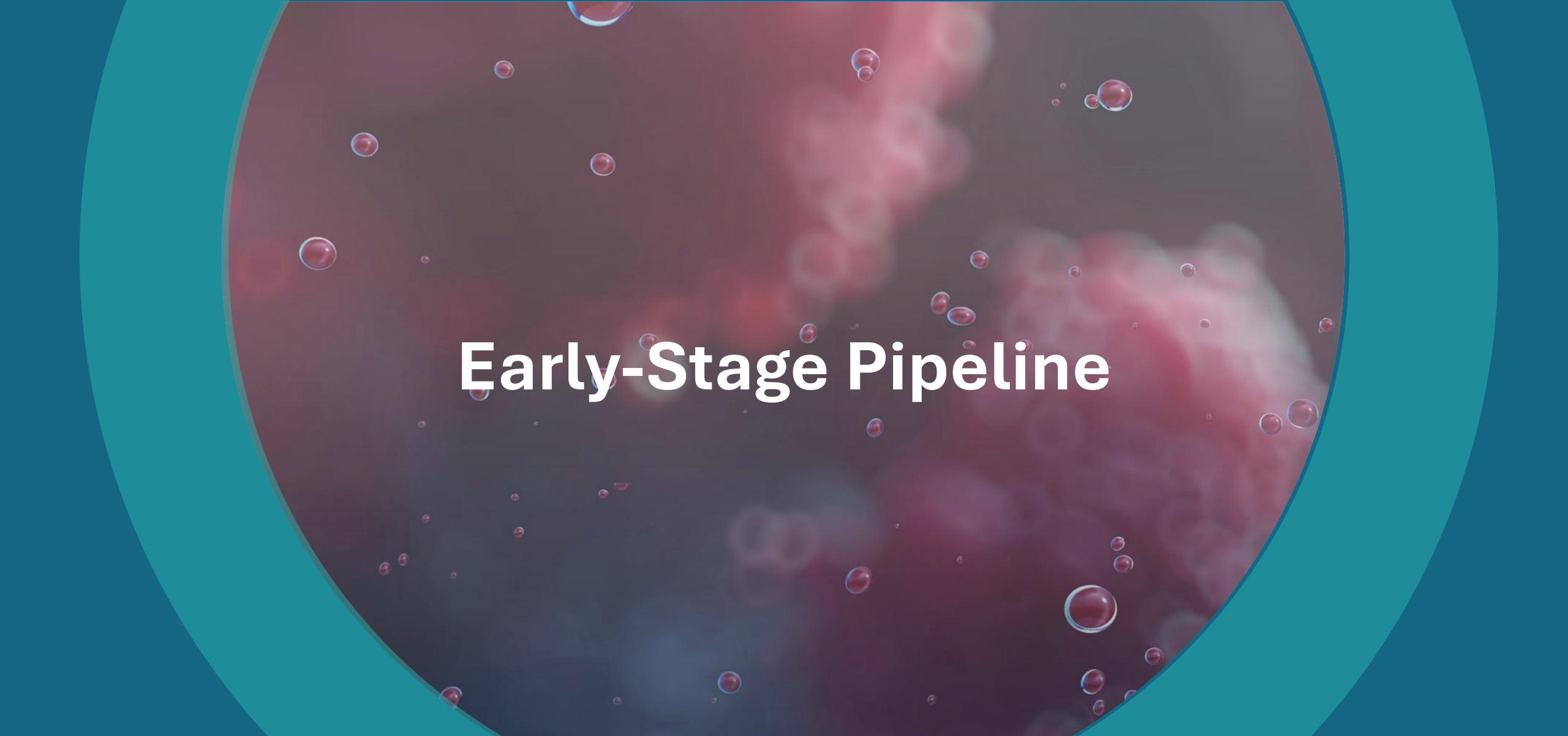


**ADG126
20 mg/kg x1 +
10 mg/kg Q3W
(N=14)**



Vision for the Future: Transforming Cancer Care with ADG126 and Pembrolizumab

New Paradigm	Durable Survival Benefits	Cornerstone Therapy	Transformative Impact
<p>Utilizes precision engineering, tumor-selective activation, and broad combinability.</p>	<p>Promotes long-term immune surveillance.</p>	<p>Seamlessly integrates into SOC treatments and enables future innovations.</p>	<p>Turning advanced cancers into manageable diseases—or achieving cures.</p>

A circular inset showing a microscopic view of cells with numerous small, clear bubbles or vesicles scattered throughout. The background is a dark, reddish-purple color. The text "Early-Stage Pipeline" is centered over this image.

Early-Stage Pipeline

ADG138: Novel, Double Masked HER2xCD3, Bispecific POWERbody™

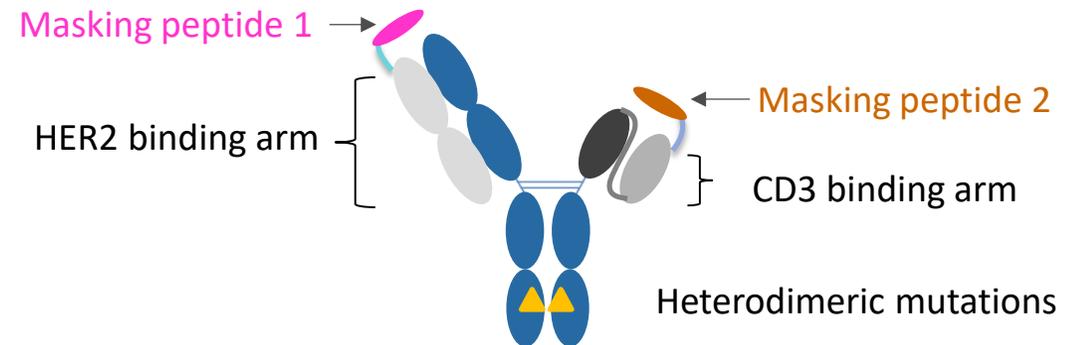
ADG138 integrates bispecific TCE (T cell engager) with precision **masking on both arms** to control cytokine release syndrome and on-target off-tumor toxicity for single agent and combination therapies in **HER2-expressing solid tumors**

Potency: Anti-tumor activity in HER2 high and low expressing tumors, as well as resistant refractory tumors, relative to DS-8201

- ✓ **Safety:** 100-fold greater reduction in cytokine release syndrome compared to its parental TCE
- ✓ **Synergistic anti-tumor activity** when combined with anti-CD137 or anti-PD-1 therapy in HER2 positive tumors

Next step: IND-enabling studies ongoing

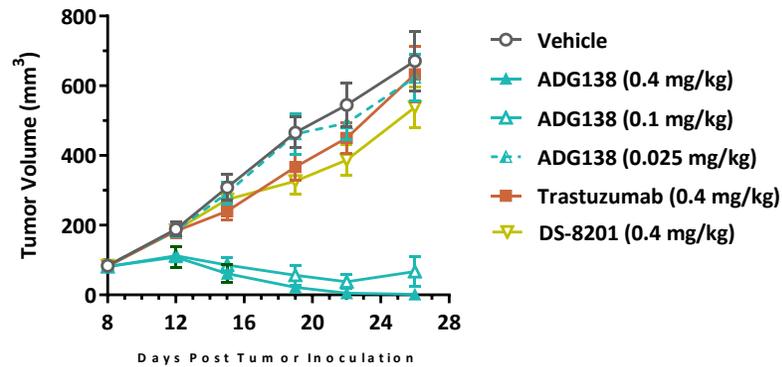
ADG138



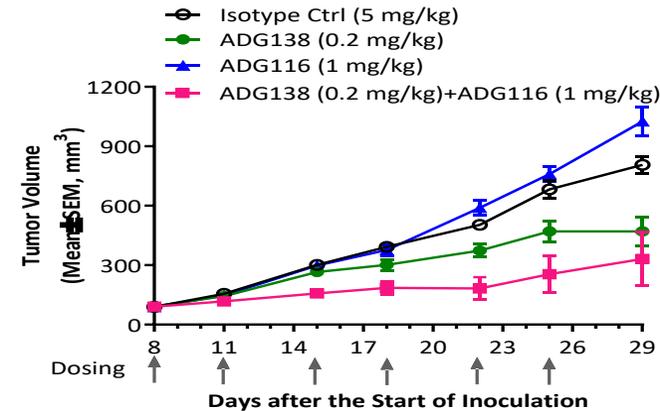
ADG138 Has Potent In vivo Antitumor Activity Both as Single Agent and in Combination with ADG116 (anti-CTLA-4)/106 (CD137 Agonist)/CD28

Combinations of ADG138 with I-O agents exhibited synergistic anti-tumor activities in mouse tumor models

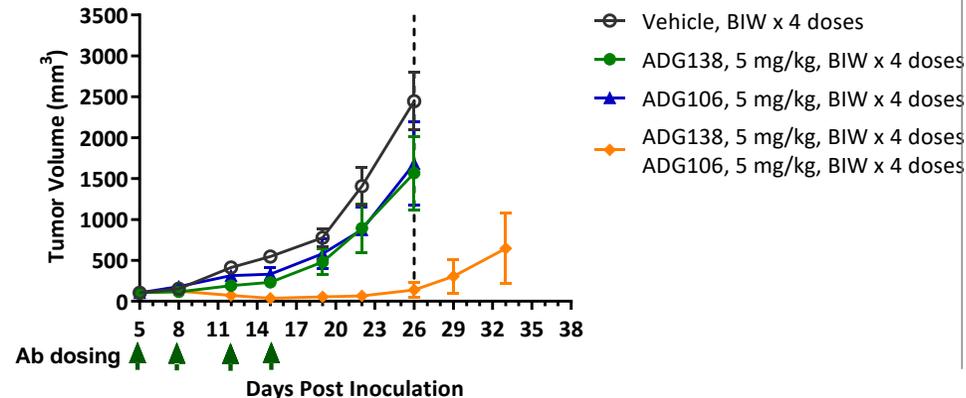
HT55/hPBMC Xenograft Model



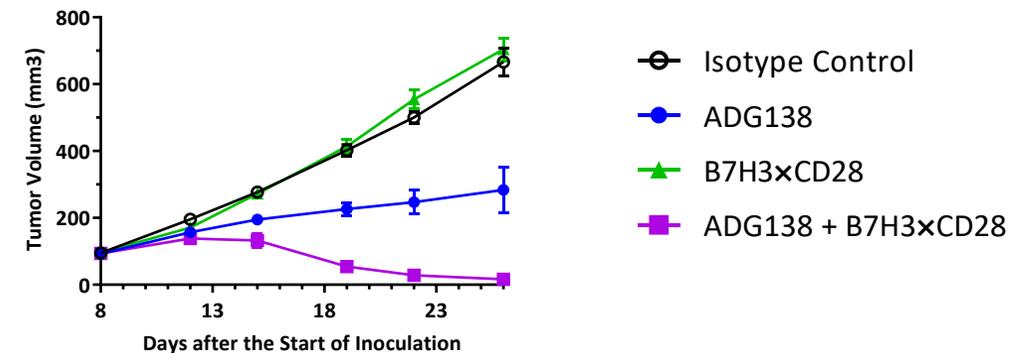
SK-OV3/hPBMC Xenograft Model



MC38-HER2 Cancer Model



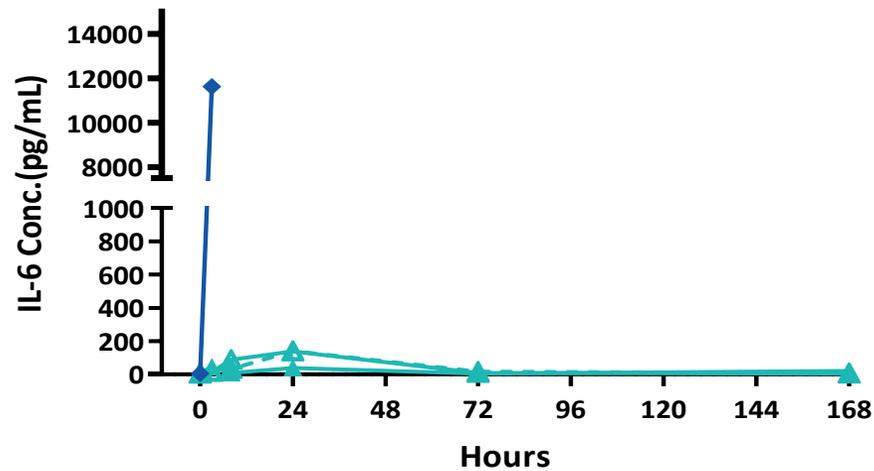
SK-OV3/hPBMC Cancer Model



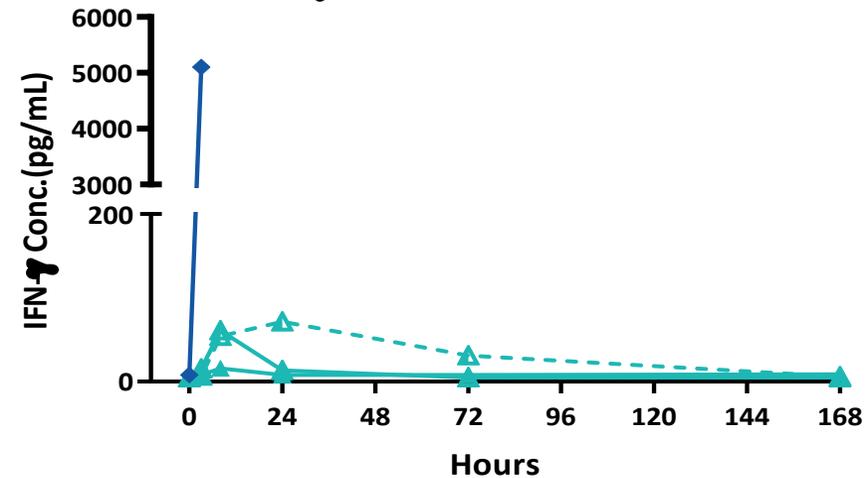
ADG138 Significantly Enhanced Safety Margin Compared to Parental Ab in An Exploratory Monkey Tox Study

	ADG138 Parental 0.2 mg/kg (QWx1)	ADG138 SAFEbody 10 mg/kg (QWx2)	ADG138 SAFEbody 30 mg/kg (QWx2)	ADG138 SAFEbody 60 mg/kg (QWx1)
Tolerability	Not Tolerated	Tolerated	Tolerated	Tolerated

IL-6 levels in monkey serum



IFN- γ levels in monkey serum



◆ ADG138 parental (0.2 mg/kg)
 ▲ ADG138 (10 mg/kg)
 ▲ ADG138 (30 mg/kg)
 ▲ ADG138 (60 mg/kg)

- ADG138 has >500x safety margin vs parental Ab in cynomolgus monkeys
- Project is at the IND enabling stage with an expression of ~5 g/l for CMC scale up

ADG152: Anti-CD20xCD3 is First Bispecific T-cell engager from POWERbody™ Platform

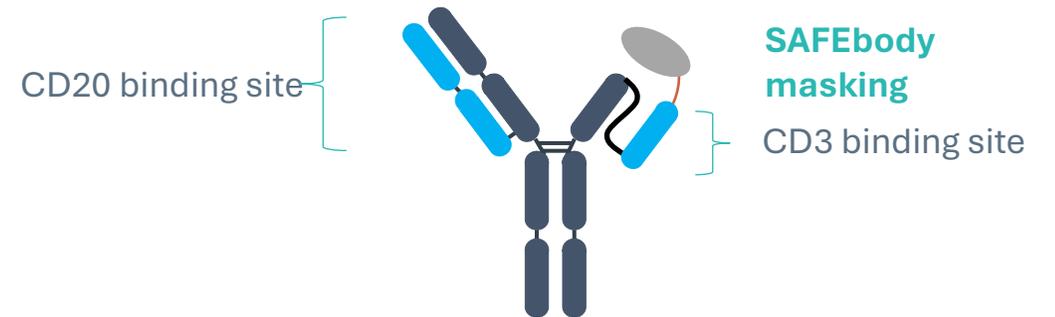
Integrates SAFEbody precision masking technology to minimize cytokine release syndrome (CRS) and on-target/off-tumor toxicities for an increased therapeutic index

Enhanced binding to CD20; anti-CD3 arm has tailor-made affinity for CD3 with SAFEbody technology

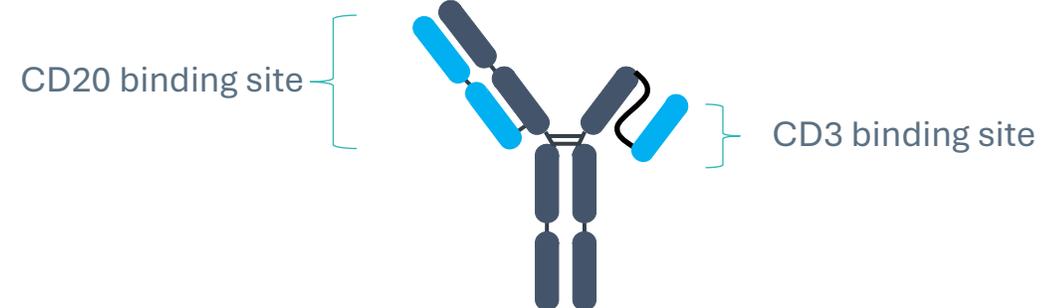
- ✓ **Potency:** Antitumor activity as a single agent in the mouse xenograft tumor model
- ✓ **Safety:** ~100-fold less CRS than a plamotamab analog in monkeys
- ✓ **PK:** Improved half-life and area under the curve than a plamotamab analog in monkeys

Next step: IND-enabling studies ongoing

ADG152 POWERbody



ADG152 Parental Antibody

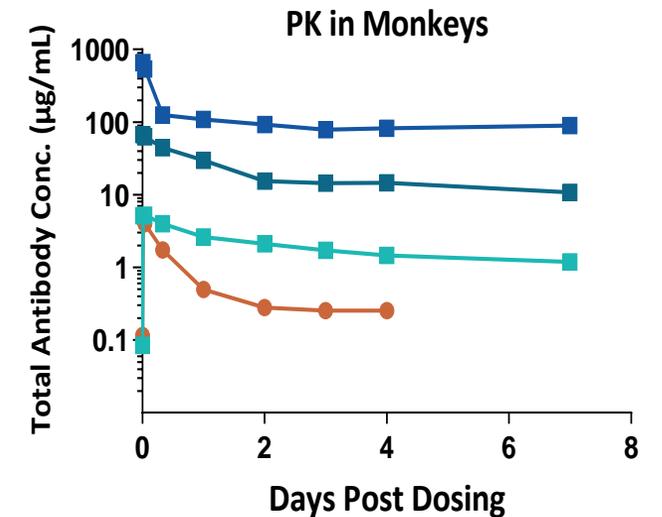
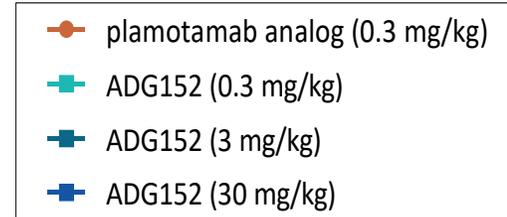
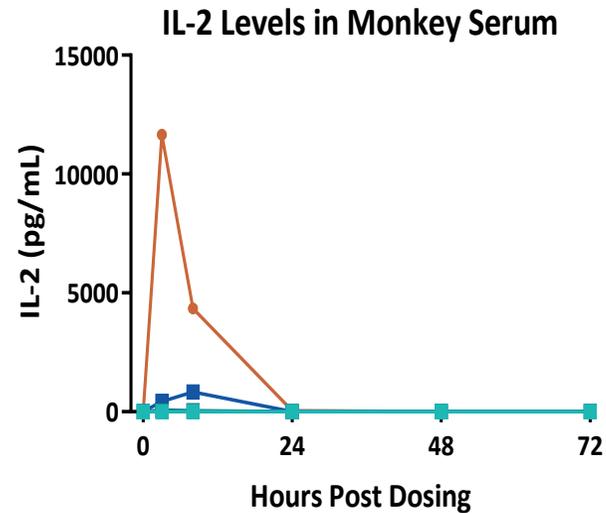
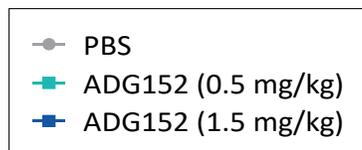
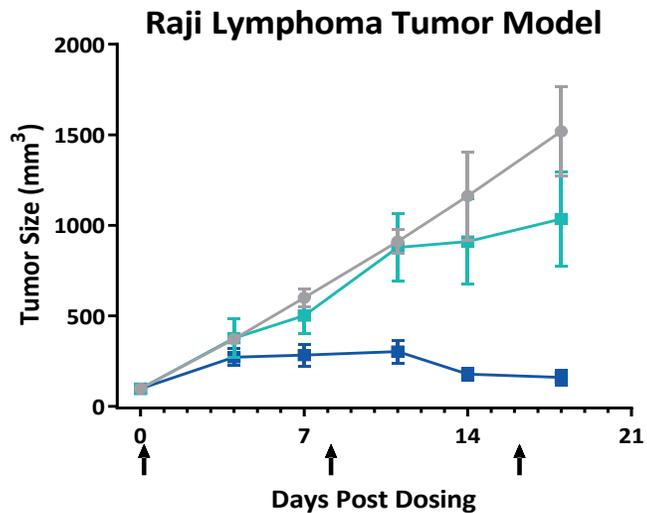


ADG152: Strong Efficacy, Improved Safety and PK Compared to a Plamotamab Analog

Strong anti-tumor activity in the mouse xenograft tumor model

Strong anti-tumor activity in the mouse xenograft tumor model

Strong anti-tumor activity in the mouse xenograft tumor model





Thank You!

ADAGENE