



Developing  
**Breakthrough Biologics,**  
Life-changing Medicines™

## Corporate Presentation

May 29, 2020



# Legal Notices

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## **Investigational Agents**



















All Company product candidates described or mentioned herein are investigational and have not yet been approved for marketing by any regulatory authority.

# Building a Leadership Position in Immuno-Oncology

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<b>Late-stage immuno-oncology company</b>	<ul style="list-style-type: none"><li>• December 2020 PDUFA goal date for most advanced product candidate</li><li>• Three additional ongoing or anticipated registration-directed studies</li></ul>
<b>Proprietary platform technologies</b>	<ul style="list-style-type: none"><li>• Bispecific DART® platform technology that exploits multiple mechanisms</li><li>• Fc-engineering to enhance innate and adaptive immunity</li></ul>
<b>Deep and differentiated pipeline</b>	<ul style="list-style-type: none"><li>• Unique immune-based mechanisms</li><li>• Retain major market rights for 6 of 7 clinical assets</li></ul>
<b>Funded to execute on plan</b>	<ul style="list-style-type: none"><li>• \$171M cash, cash equivalents and marketable securities at 3/31/20</li><li>• Multiple 2020 inflection points</li><li>• Cash runway into 2022 via anticipated and potential collaboration payments</li></ul>

# Deep and Differentiated Immuno-Oncology Pipeline

Program (Target)	Potential Indication(s)	First-in-Human (Phase 1)	Proof-of-Concept (Phase 2)	Pivotal	Major Market Rights	
<b>Margetuximab</b> (HER2)	HER2+ Breast					Greater China 
	HER2+ Gastric/GEJ (+retifanlimab/MGD013)					
<b>Flotetuzumab</b> (CD123 × CD3)	AML					
<b>Retifanlimab</b> (PD-1)	Solid Tumors					 <sup>(b)</sup>
<b>Enoblituzumab</b> (B7-H3)	SCCHN (+retifanlimab/MGD013)					Greater China 
<b>MGD013</b> (PD-1 × LAG-3)	Solid Tumors & Heme Malignancies					Greater China 
<b>MGD019</b> (PD-1 × CTLA-4)	Solid Tumors					
<b>MGC018</b> (B7-H3) <sup>(a)</sup>	Solid Tumors					

MGD = DART

MGA = Antibody

MGC = ADC

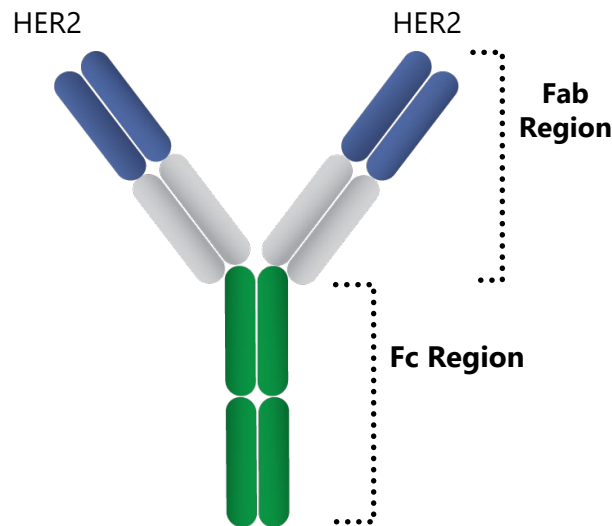
(a) MGC018 is an antibody-drug conjugate (ADC) based on a duocarmycin payload with cleavable peptide linker that was licensed from Byondis (formerly Synthon Biopharmaceuticals).

(b) MacroGenics retains rights to develop its pipeline assets in combination w/retifanlimab (MGA012) and to manufacture a portion of global clinical and commercial supply needs of retifanlimab.

**All Company product candidates described or mentioned herein are investigational and have not yet been approved for marketing by any regulatory authority.**

# Margetuximab: Anti-HER2 mAb Engineered to Enhance Activity of Immune System

December 2020 PDUFA goal date for BLA for HER2+ metastatic breast cancer (mBC)

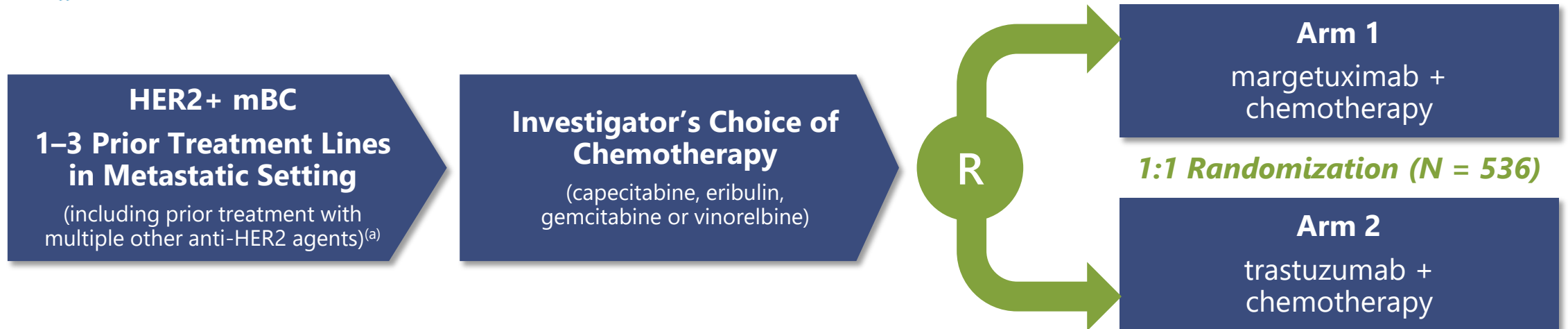


<b>Function/ MoA</b>	<ul style="list-style-type: none"> <li>• Inhibits HER2 signaling (similar to trastuzumab)</li> <li>• Fc region engineered to engage innate and adaptive immunity as mediators of anti-tumor activity</li> </ul>
<b>Clinical Studies</b>	<ul style="list-style-type: none"> <li>• Ph. 3 SOPHIA study versus trastuzumab in HER2+ mBC</li> <li>• Ph. 2/3 MAHOGANY study w/checkpoint inhibitor in HER2+ gastric cancer</li> <li>• Ph. 1/2 expansion cohort w/checkpoint inhibitor in HER2+ solid tumors</li> </ul>
<b>2020 Anticipated Milestones</b>	<ul style="list-style-type: none"> <li>• BLA for HER2+ mBC: PDUFA goal date (12/18/2020)</li> <li>• Final SOPHIA OS analysis (N=385) (2H)</li> <li>• Obtain initial data from MAHOGANY Module A</li> </ul>

*Margetuximab is investigational and has not yet been approved for marketing by any regulatory authority*

# Phase 3 SOPHIA Study Comparing Margetuximab to Trastuzumab

Designed to support registration in 3rd/4th line HER2+ metastatic breast cancer



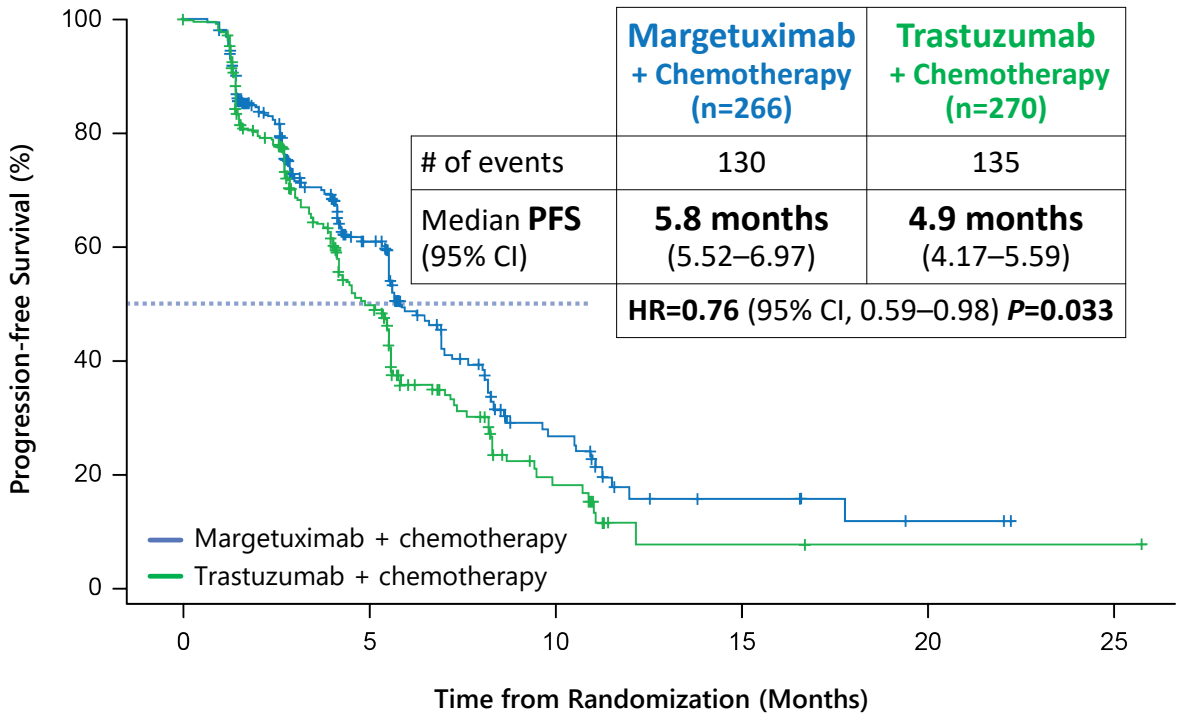
- **Sequential primary endpoints: PFS and OS**
- Patients carrying CD16A (FcγRIIIa) 158F allele were pre-specified exploratory subpopulation

PFS (N=257, HR=0.67,  $\alpha=0.05$ , power=90%)  
OS (N=385, HR=0.75,  $\alpha=0.05$ , power=80%)

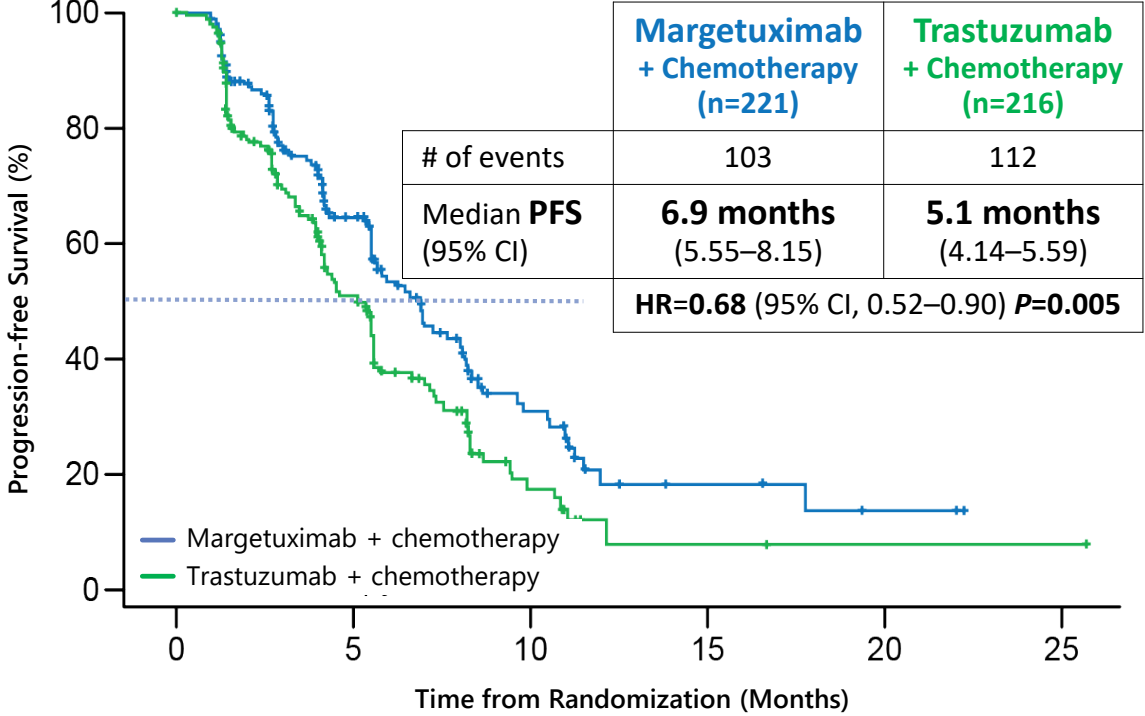
(a) All study patients had previously received trastuzumab and pertuzumab, and approximately 90% had previously received ado-trastuzumab emtansine.

# Primary PFS Endpoint: Margetuximab Demonstrated Superiority to Trastuzumab

**PFS Primary Endpoint (ITT Population):**  
24% Risk Reduction of Disease Progression



**Pre-specified Exploratory Subpopulation (CD16A-158F Carriers):**  
32% Risk Reduction of Disease Progression



Margetuximab	266	174	94	45	21	8	6	4	2	0
Trastuzumab	270	158	74	33	13	2	2	1	1	1

Margetuximab	221	157	84	42	21	8	6	4	2	0
Trastuzumab	216	129	62	30	11	2	2	1	1	1

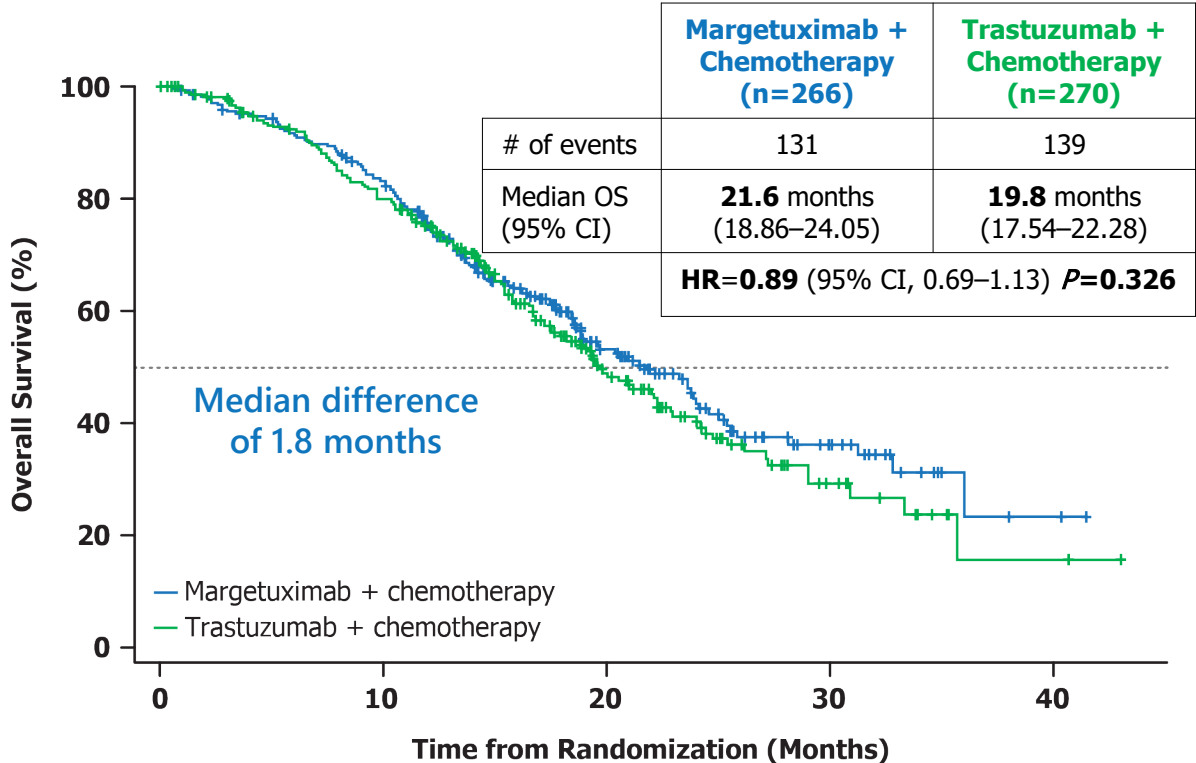
October 2018 data cut-off after 265 PFS events in ITT population.  
CI=confidence interval. ITT=Intent to Treat population: N=536. CD16A 158F Carriers=FF or FV Genotype.  
HR=Hazard Ratio (ITT by stratified Cox model; F Carriers by unstratified Cox model).

Rugo, et al., ASCO 2019

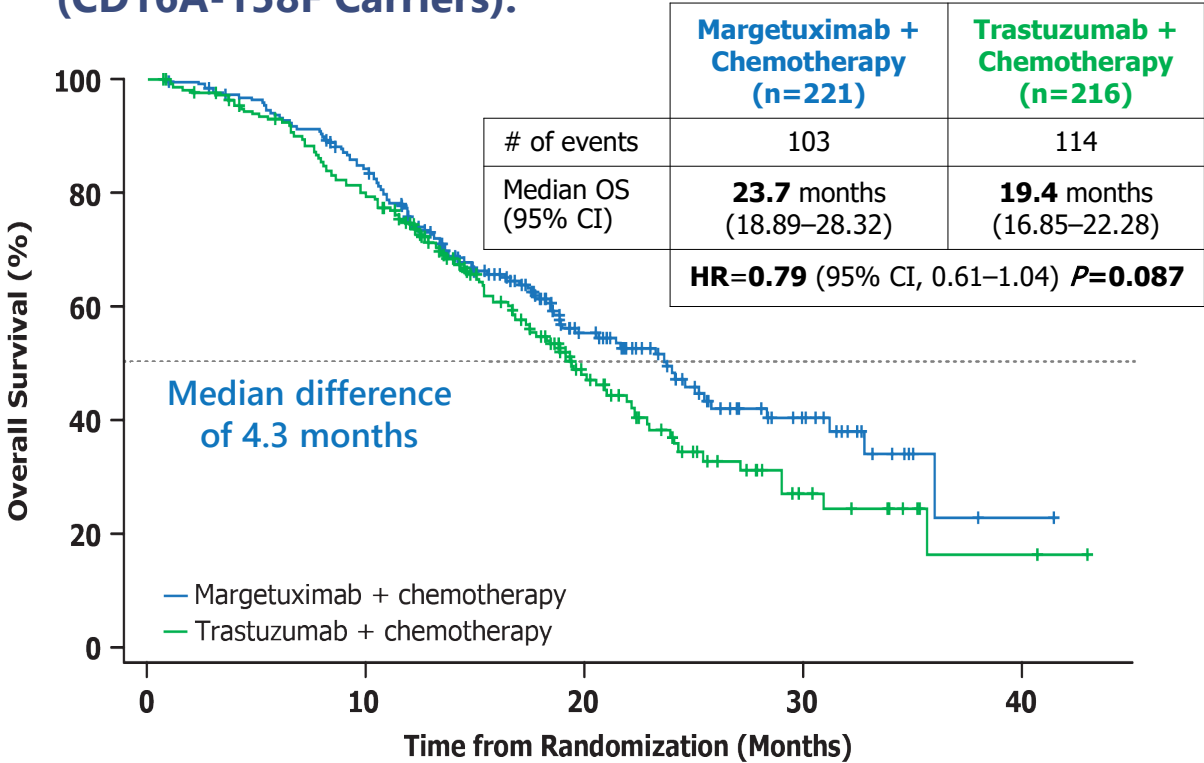
# Second Interim Overall Survival Analysis: Trend Favored Margetuximab

Final analysis expected 2H20

## ITT Population



## Pre-specified Exploratory Subpopulation (CD16A-158F Carriers):



Margetuximab 266 259 249 239 230 214 188 159 131 107 80 64 47 35 31 22 14 9 3 2 2 0  
 Trastuzumab 270 260 246 236 218 205 183 160 126 102 74 57 43 30 22 16 10 6 2 2 2 1 0

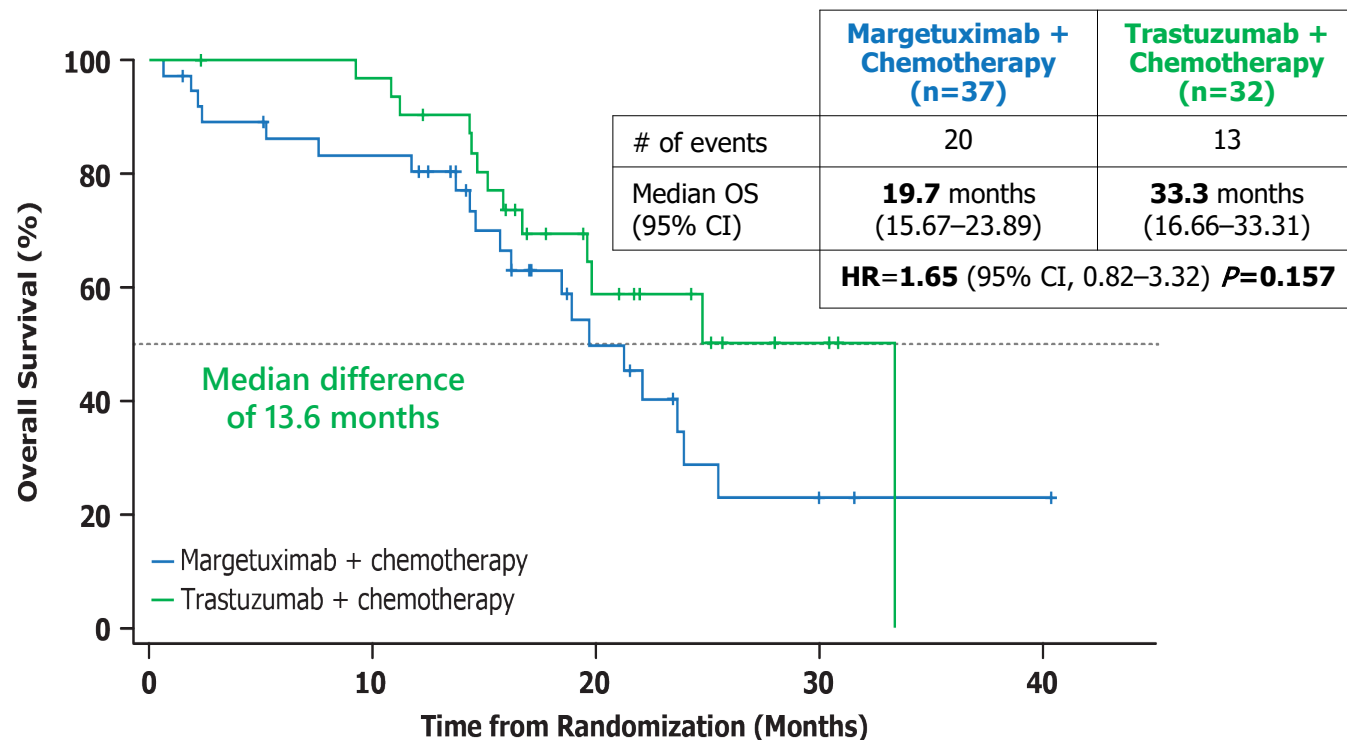
Margetuximab 221 219 212 204 196 181 157 135 111 91 68 55 42 31 27 19 13 8 2 1 1 0  
 Trastuzumab 216 210 201 192 176 165 145 123 98 81 57 43 30 21 16 11 9 6 2 2 2 1 0

September 2019 data cut-off after 270 events in ITT population. Median follow-up: 15.6 months.  
 ITT=Intent to Treat population: N=536. CD16A 158F Carriers=FF or FV Genotype.  
 CI=confidence interval. HR=Hazard Ratio (ITT by stratified Cox model; F Carriers by unstratified Cox model).



# Pre-specified Exploratory OS in CD16A-158 VV Homozygotes

VV subpopulation represents 33 events (270 events in ITT population)



Margetuximab	37	34	32	30	29	29	27	23	19	15	11	9	5	4	4	3	1	1	1	1	1	0	
Trastuzumab	32	32	31	31	31	30	28	27	20	14	11	8	8	4	3	3	1	0					

## Unbalanced patient characteristics

Baseline Characteristic	Margetuximab + Chemotherapy (n=37)	Trastuzumab + Chemotherapy (n=32)
Cancer disease history		
Brain, n (%)	8 (22%)	3 (9%)
Breast, n (%)	10 (27%)	5 (16%)
Liver, n (%)	16 (43%)	10 (31%)
Lung, n (%)	11 (30%)	13 (41%)
Lymph node, n (%)	21 (57%)	16 (50%)
HER2 IHC 3+, n (%)	19 (51%)	18 (56%)
Hormone receptor +, n (%)	23 (62%)	18 (56%)
ECOG PS 1, n (%)	14 (38%)	16 (50%)
>60 years of age, n (%)	16 (43%)	5 (16%)
>2 prior metastatic lines of therapy, n (%)	15 (41%)	9 (28%)

Less favorable

September 2019 data cut-off after 270 events in ITT population. Median follow-up: 15.6 months. CI=confidence interval. HR=Hazard Ratio (by unstratified Cox model).

Rugo, et al., SABCS 2019

# Overall Safety Profiles Similar

## Adverse Events (AE)

	Margetuximab + Chemotherapy (n=264)		Trastuzumab + Chemotherapy (n=266)	
<b>Any grade AE, n (%)</b>	260 (98.5)		261 (98.1)	
<b>Any margetuximab or trastuzumab-related AE, n (%)</b>	160 (60.6)		132 (49.6)	
<b>Grade ≥3 AE, n (%)</b>	142 (53.8)		140 (52.6)	
<b>Grade ≥3 margetuximab or trastuzumab-related AE, n (%)</b>	34 (12.9)		22 (8.3)	
<b>Any SAE, n (%)</b>	43 (16.3)		49 (18.4)	
<b>Any margetuximab or trastuzumab-related SAE, n (%)</b>	5 (1.9)		4 (1.5)	
<b>AE leading to treatment<sup>a</sup> discontinuation, n (%)</b>	8 (3.0)		7 (2.6)	
<b>AEs resulting in death,<sup>b</sup> n (%)</b>	3 (1.1) <sup>c</sup>		2 (0.8) <sup>d</sup>	
<b>AEs of special interest, n (%)</b>	<b>All Grade</b>	<b>Grade ≥3</b>	<b>All Grade</b>	<b>Grade ≥3</b>
Infusion-related reaction (IRR)	35 (13.3)	4 (1.5)	9 (3.4)	0
Discontinuation due to IRRs, n (%)	2 (0.6)	0	0	0
LV dysfunction leading to dose delay or discontinuation, n (%)	4 (1.5)	0	6 (2.3)	0

Safety Population (randomized patients who received any study treatment): N=530. April 2019 cut-off.

(a) Including both anti-HER2 study therapy and chemotherapy. (b) No AEs resulting in death were considered related to anti-HER2 study therapy.

(c) Pneumonia (n=2), pneumonia aspiration (n=1). (d) Pneumonia (n=1), acute kidney injury (n=1). LV=left ventricular; SAE=serious AE.

Rugo, et al., SABCS 2019

# Margetuximab's Potential Role in Treatment of HER2+ mBC

**Need remains for additional therapies in later lines**  
Patients will progress on other HER2-directed therapies

**PFS improvement  
vs. trastuzumab  
in clinical study**

Superiority in  
head-to-head trial

**Flexibility**

Ability to tailor  
treatment by  
selecting among  
four different  
chemotherapies

**Familiarity**

Side effect profile  
is well known  
and manageable

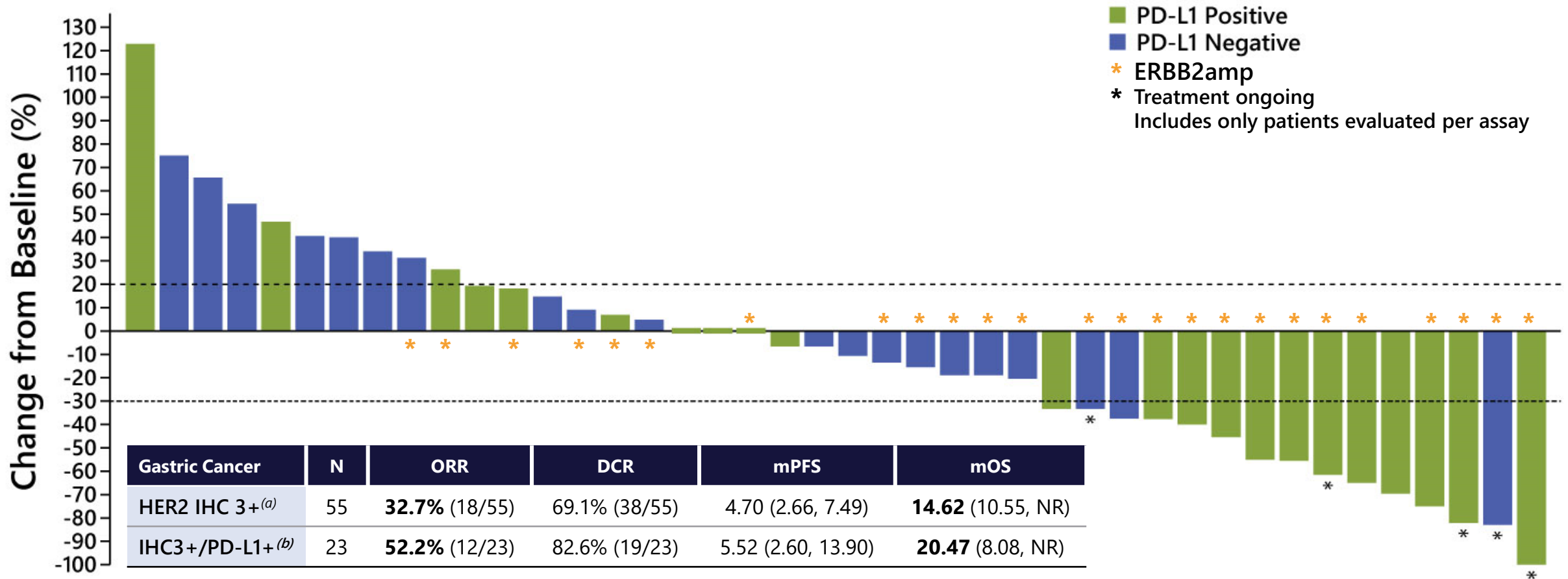
**CD16A  
exploratory  
analysis**

85% of population  
are F carriers

*Margetuximab is investigational and has not yet been approved for marketing by any regulatory authority*

# Promising Activity in Advanced Gastric Cancer Patients in Phase 2 Study

33% ORR in HER2 3+ gastric cancer previously treated with chemotherapy and trastuzumab



Data cut-off July 10, 2019. Includes patients who received ≥1 margetuximab and pembrolizumab dose in expansion phase, and had baseline measurable disease and ≥1 post-baseline disease assessment.

(a) Immunohistochemistry (IHC) test gives score of 0 to 3+ that measures amount of HER2 receptor protein on surface of cells in cancer tissue sample. Score of 0 to 1+ is called "HER2 negative", score of 2+ is called "borderline", score of 3+ is called "HER2 positive."

(b) "PD-L1 Positive" reflects Combined Positive Score (per standard FDA approved assay) ≥1% (PD-L1 tested on archival tissue by IHC; clone 22C3 pharmDx).

Catenacci, et al., ESMO 2019

# Gastric Cancer as Follow-on Indication

Data from 2L margetuximab + anti-PD-1 mAb presents opportunity to advance to 1L

## HER2+ gastric cancer benchmarks

	1st Line	2nd Line			3rd Line
	SOC	SOC	Ongoing Phase 2 Study		Ongoing Study
<b>Agent (Study)</b>	Trastuzumab + Chemo <sup>(a)</sup> (TOGA)	Ramucirumab + Paclitaxel <sup>(b)</sup> (RAINBOW)	<b>Margetuximab + Pembrolizumab<sup>(c)</sup></b>		Pembrolizumab <sup>(d)</sup> (KEYNOTE-61) PD-L1+
			IHC 3+	IHC 3+/PD-L1+	
<b>ORR</b>	47%	28%	33%	52%	15.8%
<b>Median PFS</b>	6.7 mos.	4.4 mos.	4.7 mos.	5.5 mos.	1.5 mos.
<b>Median OS</b>	<b>13.1 mos.</b>	9.6 mos.	<b>14.6 mos.</b>	<b>20.5 mos.</b>	9.1 mos.
<b>≥ Grade 3 TRAEs</b>	<b>68%</b>	Overall: N/A 41% Neutropenia 15% Hypertension 12% Fatigue	20%		14%
<b>Gastric/GEJ Patient Mix</b>	80/20%	80/20%	100%/0%		70%/30%
					80%/20%

SOC = Standard of Care

(a) Data from Herceptin package insert; Bang, et al., Lancet, 2010;

(b) Data from Cyramza package insert; Wilkes, et al., Lancet Oncology, 2014;

(c) Data presented at ESMO 2019; Grade 3 TRAE includes all GC and GEJ patients.

(c) Data presented at ESMO 2019; Grade 3 TRAE includes all GC and GEJ patients.

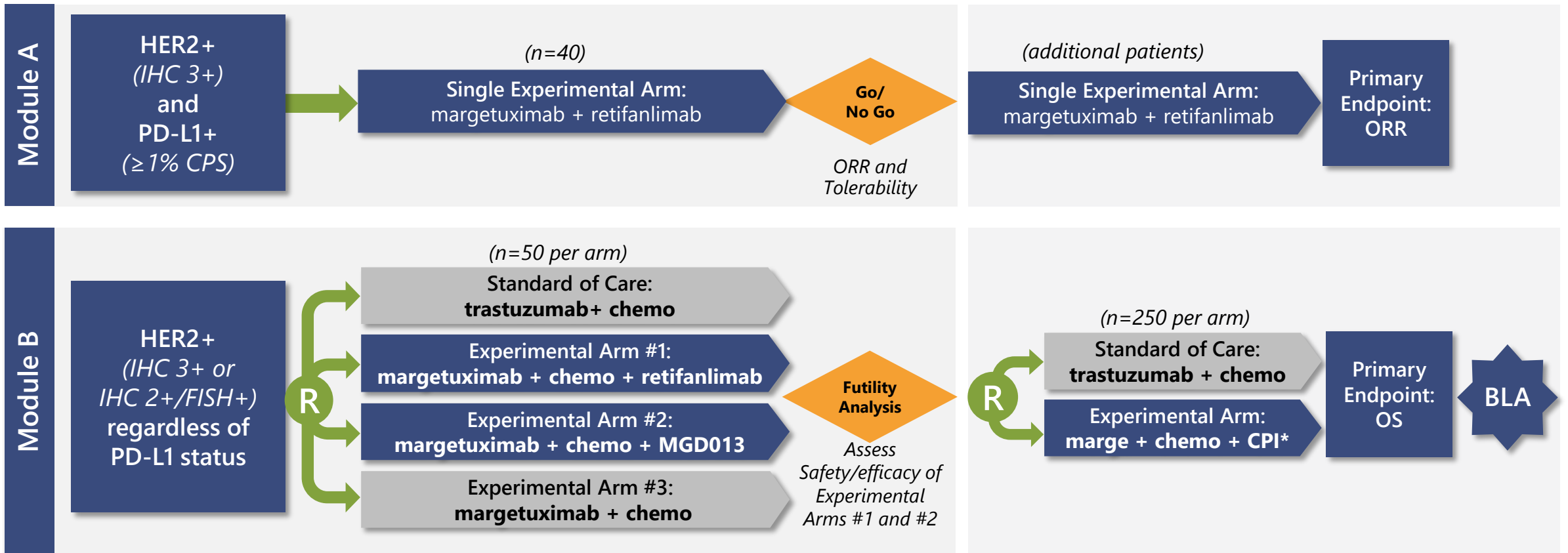
(d) Shitara, et al., 2018, Lancet;

(e) Shitara, et al., 2019, Lancet Oncol.

# MAHOGANY Phase 2/3 Study: Registration Path in 1L Gastric & GEJ Cancer

Module A has potential for U.S. Accelerated Approval of chemotherapy-free regimen

## MAHOGANY



MAHOGANY (Margetuximab in HER2-positive Gastric Cancer

\* Pending chronic tox study (if regimen with MGD013 is selected).

# Flotetuzumab: CD123 × CD3 DART Molecule

*Establishing leadership position among CD123-targeting bispecifics*

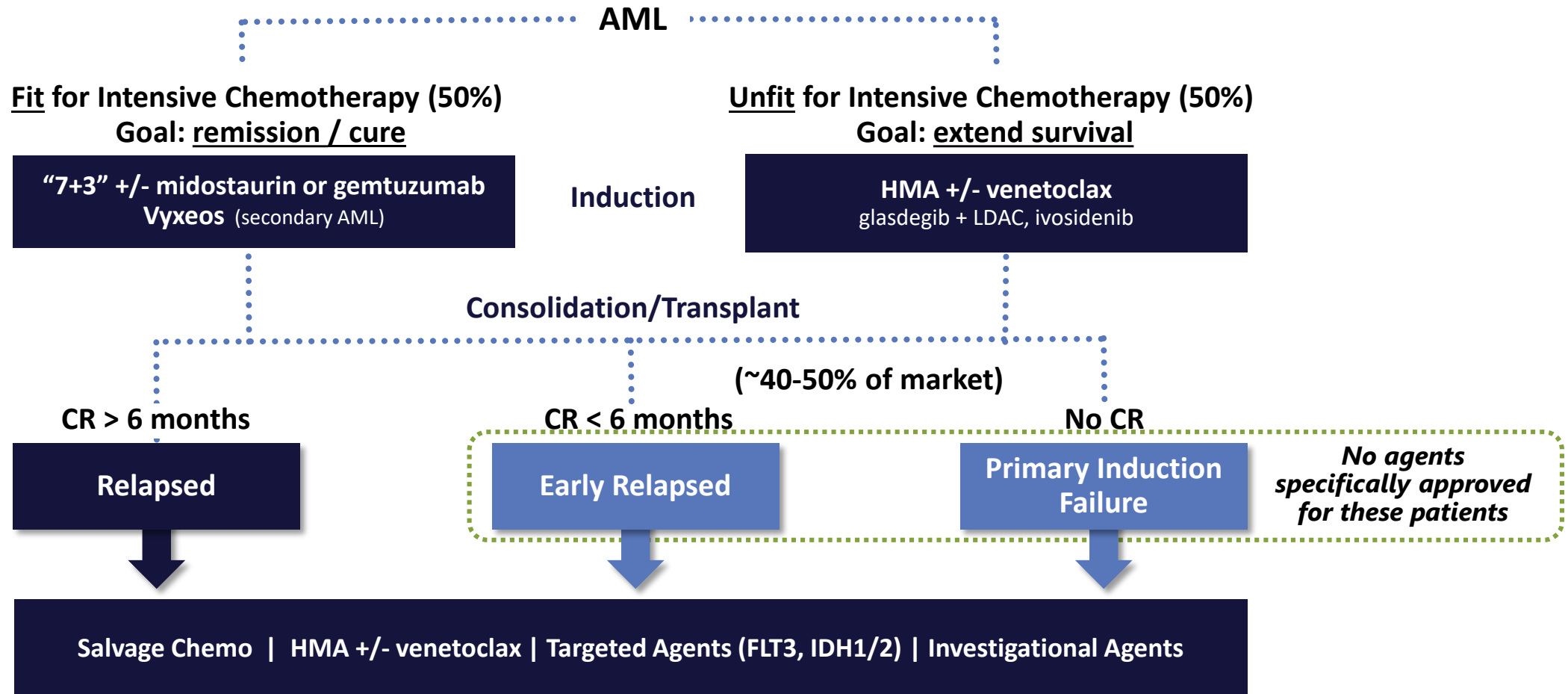


<b>Function/ MoA</b>	<ul style="list-style-type: none"> <li>• Redirected T-cell killing against leukemia cells               <ul style="list-style-type: none"> <li>– Eliminates leukemic stem cells; spares normal hematopoietic stem cells</li> <li>– Engages any T-cell without HLA-restriction</li> </ul> </li> </ul>
<b>Clinical Studies</b>	<ul style="list-style-type: none"> <li>• Single arm study to support registration in primary induction failure (PIF) and early relapse (ER) AML (continuation of ongoing Phase 1/2)</li> <li>• Phase 1 combination with retifanlimab planned in R/R AML</li> </ul>
<b>2020 Anticipated Milestones</b>	<ul style="list-style-type: none"> <li>• Updated data from Phase 1/2 study in 2H 2020</li> </ul>

*Flotetuzumab is investigational and has not yet been approved for marketing by any regulatory authority*

# Primary Induction Failure & Early Relapsed AML: Significant Unmet Need

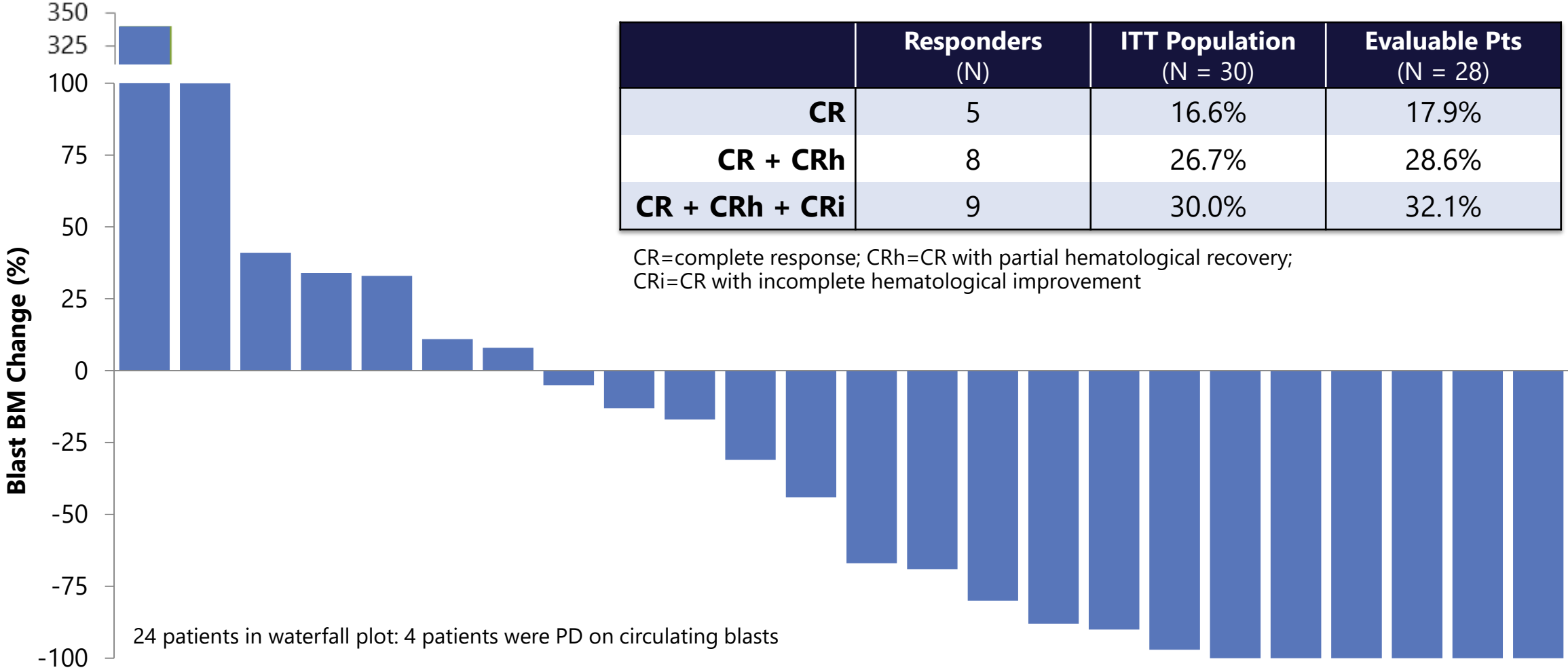
50% of patients have no known targetable mutation; flotetuzumab is mutation-agnostic





# Flotetuzumab is Active in Primary Induction Failure & Early Relapsed AML Patients

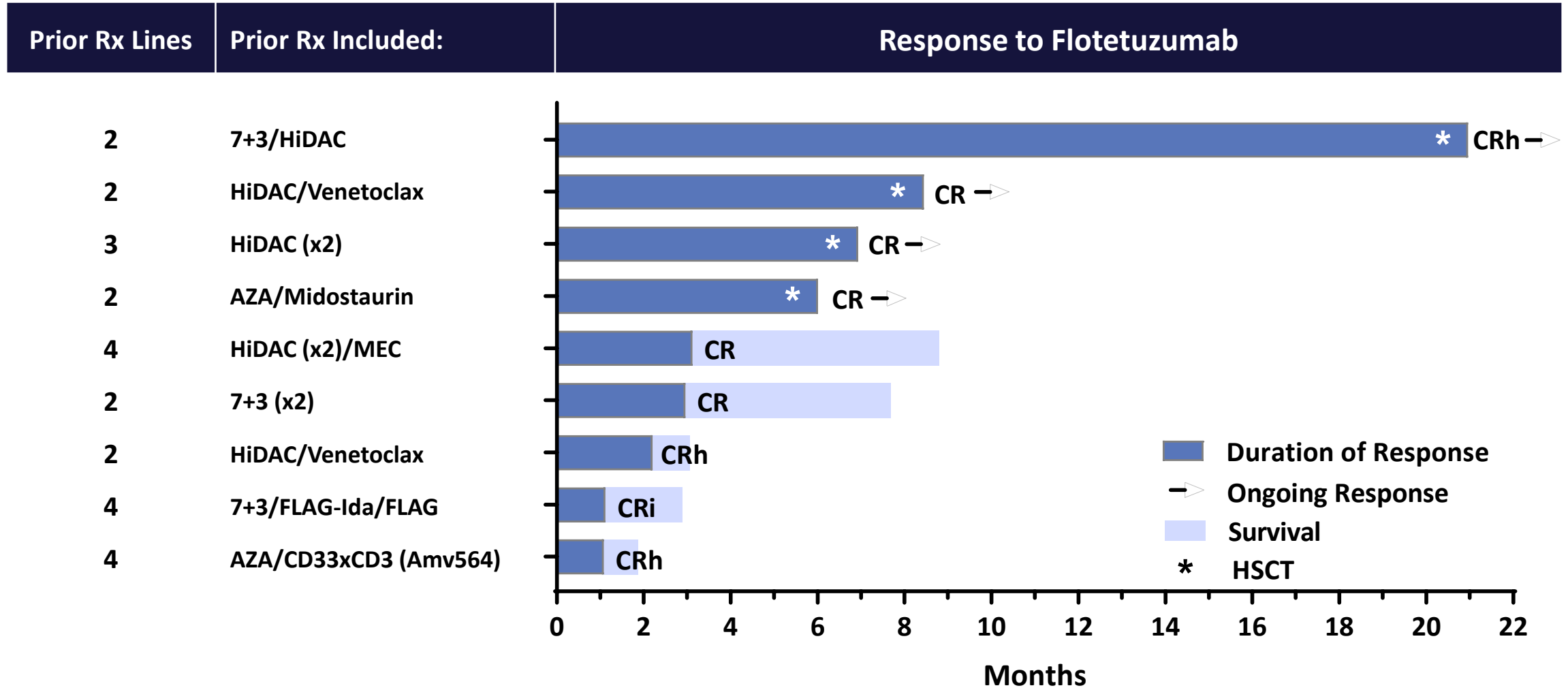
Benchmark analysis suggests historical CR+CRh rates in this setting of ~12.5%<sup>(a)</sup>



(a) Unpublished analysis of the CLASSIC I, VALOR, ADMIRAL trials and additional trials that included venetoclax, gemtuzumab-ozogamicin, and IDH1/2 inhibitors; (n=1328): CR/CRh = 12.5% [95% CI = 7.7%, 19.6%]

Uy, et al., ASH 2019

# Flotetuzumab: Duration of Response in PIF & Early Relapsed AML Patients



\* Four responders (3 CR, 1 CRh) received allo-HSCT consolidation

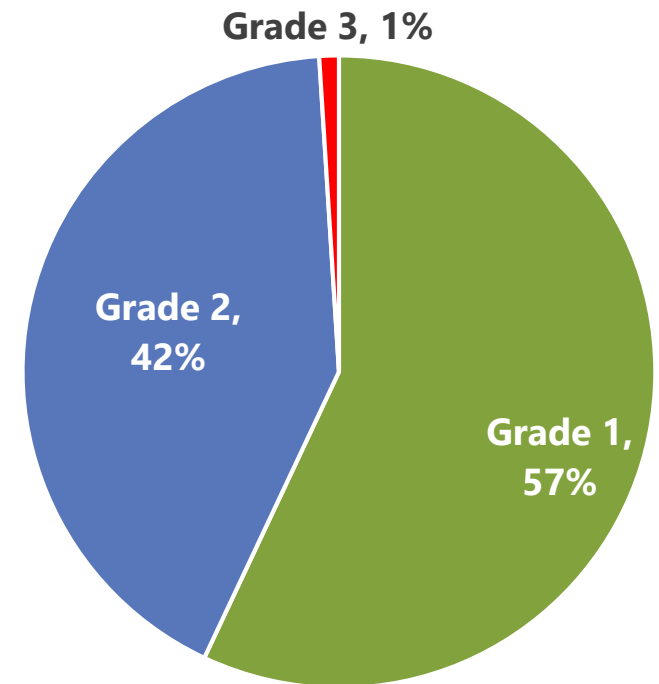
Uy, et al., ASH 2019

# Mitigating Cytokine Release Syndrome Associated With T Cell Engagers

*Decreased CRS severity and increased total flotetuzumab dose intensity*

- Infusion-related reaction /cytokine release syndrome (IRR/CRS) occurred in all (30/30) patients:
  - Mild to moderate (grade 1 or 2) in severity; only one grade 3 event reported in one patient
  - Most events observed were of short duration (Median: Grade 1=1 day; Grade 2=2 days; Grade 3=3 days)
- CRS mitigation strategies:
  - Lead-in dosing schedule for flotetuzumab
  - Early use of tocilizumab as supportive care
  - Short half-life molecule can be “switched-off” (Continuous infusion advantageous for managing exposure)

**Distribution of CRS Events by Grade**

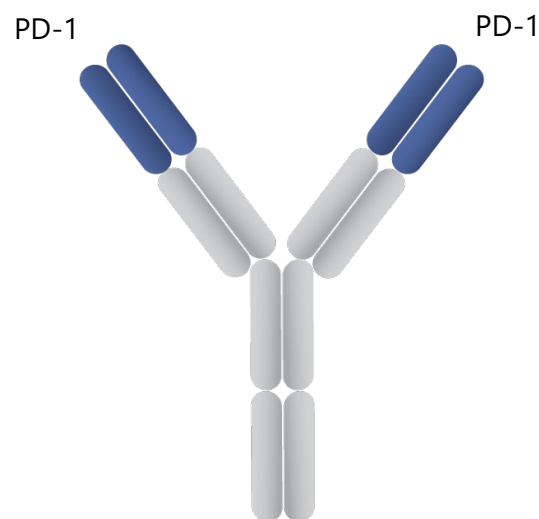



There were no grade 4 events

Uy, et al., ASH 2019

# Retifanlimab (MGA012): Anti-PD-1 antibody

## Global collaboration with Incyte



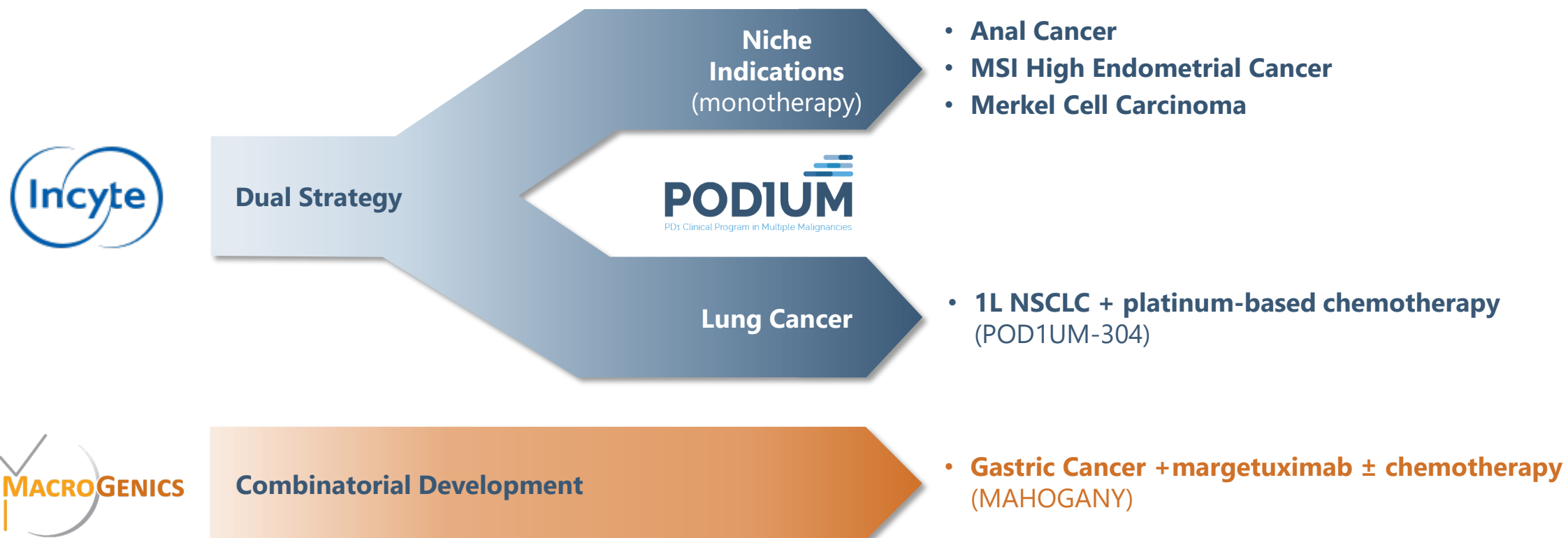
Function/ MoA	<ul style="list-style-type: none"> <li>• Humanized, hinge-stabilized IgG4 mAb</li> <li>• Inhibits PD-1</li> </ul>
Clinical Studies	<ul style="list-style-type: none"> <li>• Five registration-directed studies ongoing or planned in 2020 across a broad range of tumor types<sup>(a)</sup></li> </ul>
Global Incyte Transaction 	<ul style="list-style-type: none"> <li>• Up to \$750M in milestones (\$15M received to date)</li> <li>• Tiered royalties of 15-24% on future retifanlimab sales</li> <li>• Rights to develop pipeline assets with retifanlimab</li> </ul>
2020 Anticipated Milestones	<ul style="list-style-type: none"> <li>• Monotherapy data in anal cancer</li> <li>• Initiation of Ph. 3 randomized study in NSCLC by Incyte</li> </ul>

(a) ClinicalTrials.gov referenced May 28, 2020

**Retifanlimab is investigational and has not yet been approved for marketing by any regulatory authority**

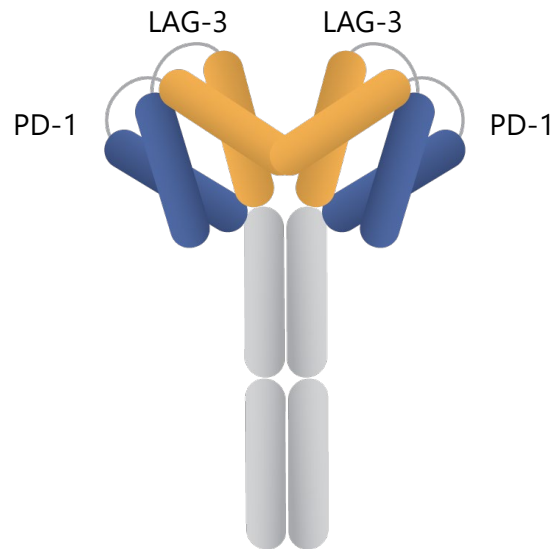
# Comprehensive Development Plans for Retifanlimab

*Multiple potentially registration-enabling clinical studies*



ClinicalTrials.gov referenced May 28, 2020

# MGD013 (PD-1 × LAG-3): First Bispecific Checkpoint Molecule in Clinical Trials



## Function/ MoA

- Simultaneous and/or independent blockade of two checkpoint molecules
- Reactivation of exhausted T cells

## Clinical Studies

- Ph. 1 dose expansion in:
  - Nine tumor types (solid and liquid); checkpoint-naïve and checkpoint-experienced
  - Multiple combination studies ongoing (including expansion cohort with margetuximab in HER2+ tumors)

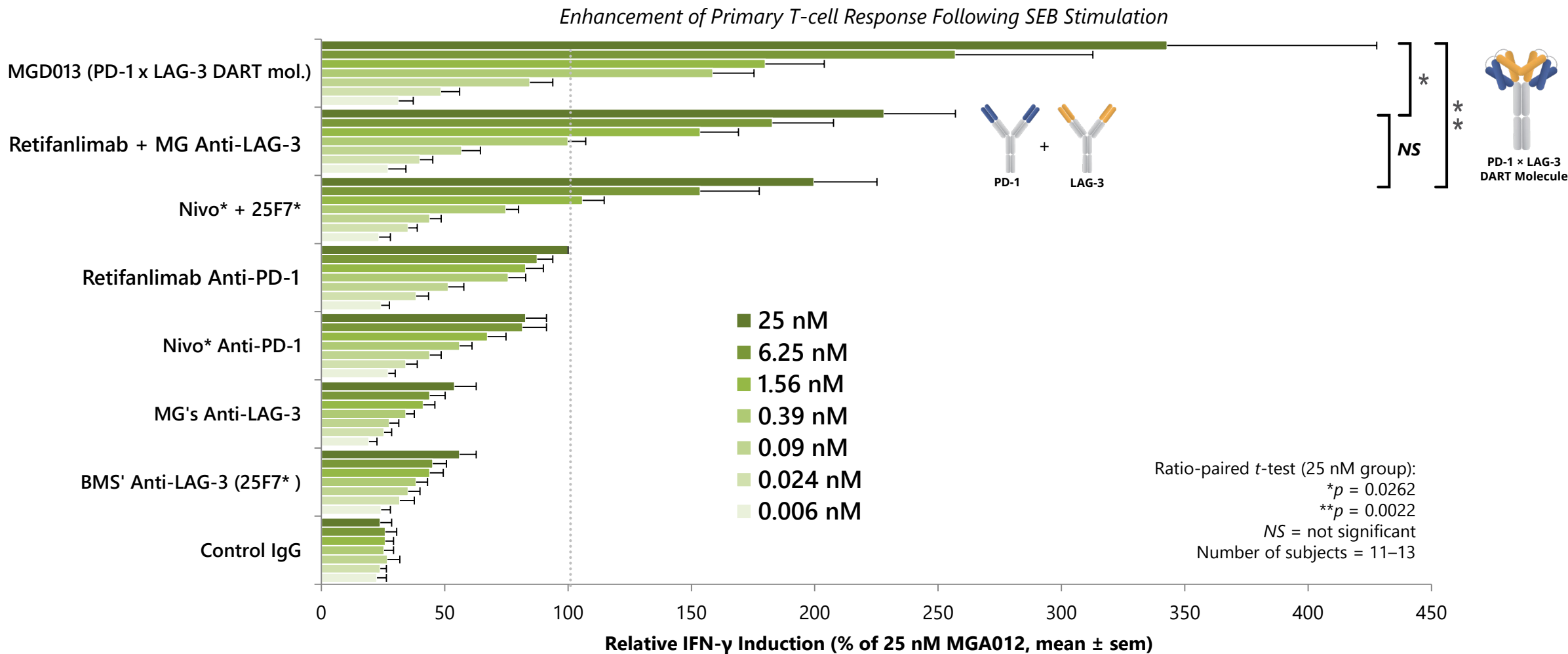
## 2020 Anticipated Milestones

- Select indications for further monotherapy and combination development

*MGD013 is investigational and has not yet been approved for marketing by any regulatory authority*

# MGD013: Synergistic T-cell Activation

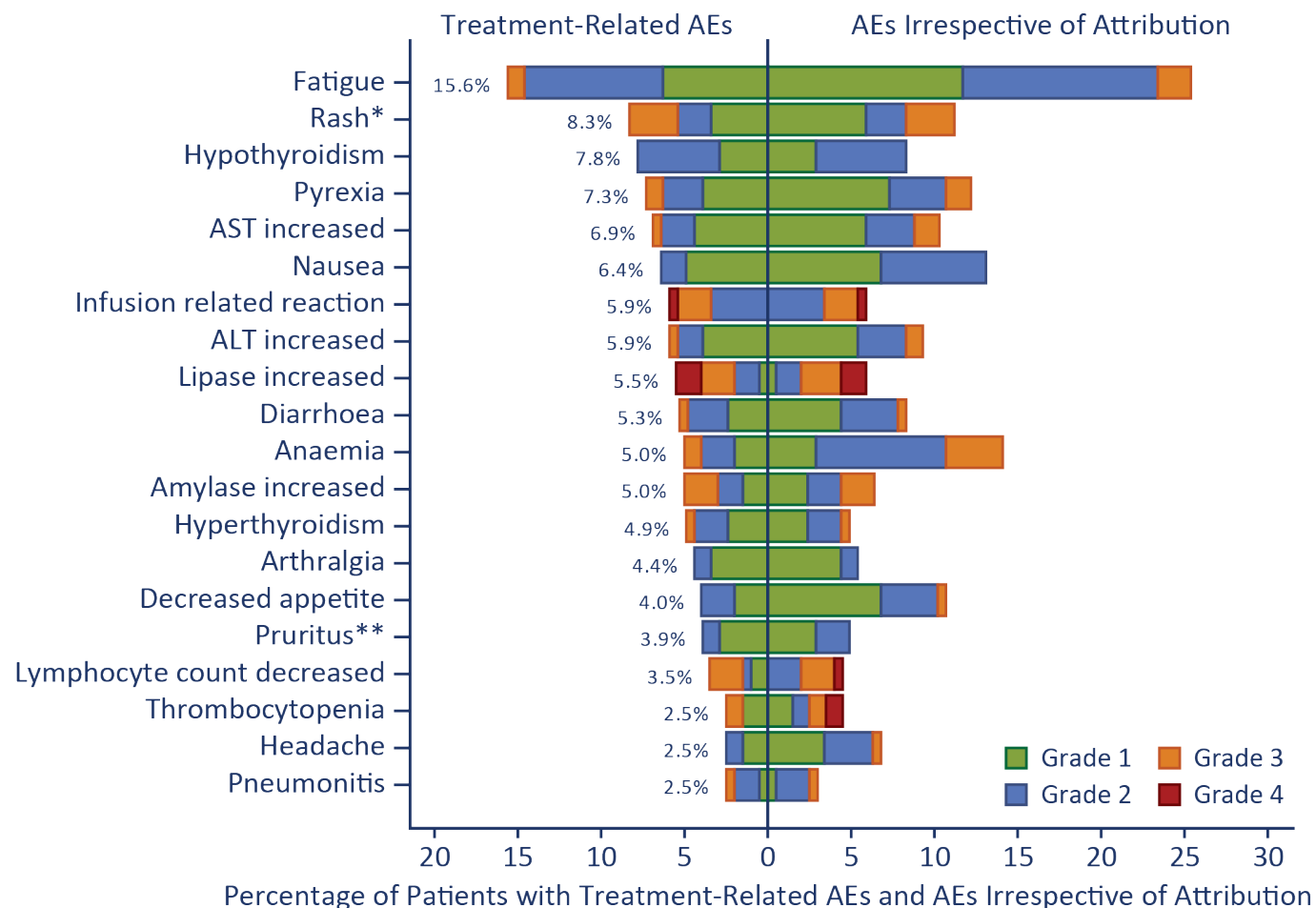
*DART molecule construct enhances T-cell activation vs. anti-PD-1 + anti-LAG-3 mAbs in vitro*



\*IFN $\gamma$  release by 25 nM MGA012 = 3276 $\pm$ 744 pg/ml.

# MGD013 Safety Profile Consistent with PD-1 Antibody Monotherapy

Overall AE Totals	No. (%) of Patients	
	All Grades (N=205)	≥ Grade 3 (N=205)
AE (irrespective of causality)	178 (86.8)	86 (42.0)
Treatment-related AE	118 (57.6)	37 (18.0) <sup>a</sup>
SAE (irrespective of causality)	63 (30.7)	47 (22.9)
Treatment-related SAE	18 (8.8)	11 (5.4)
AE leading to discontinuation	18 (8.8)	16 (7.8)
AESIs in ≥ 2 Patients		
Rash	17 (8.3)	6 (2.9)
Hypothyroidism	16 (7.8)	0 (0.0)
IRR or CRS	13 (6.3)	5 (2.4)
Diarrhoea	11 (5.4)	1 (0.5)
Lipase increased	11 (5.4)	7 (3.4)
Hyperthyroidism	10 (4.9)	1 (0.5)
Arthralgia	9 (4.4)	0 (0.0)
Pneumonitis	4 (2.0)	1 (0.5)
Myalgia	4 (2.0)	0 (0.0)
Peripheral neuropathy	3 (1.5)	1 (0.5)
Hepatitis	3 (1.5)	2 (1.0)
Adrenal insufficiency	2 (1.0)	0 (0.0)



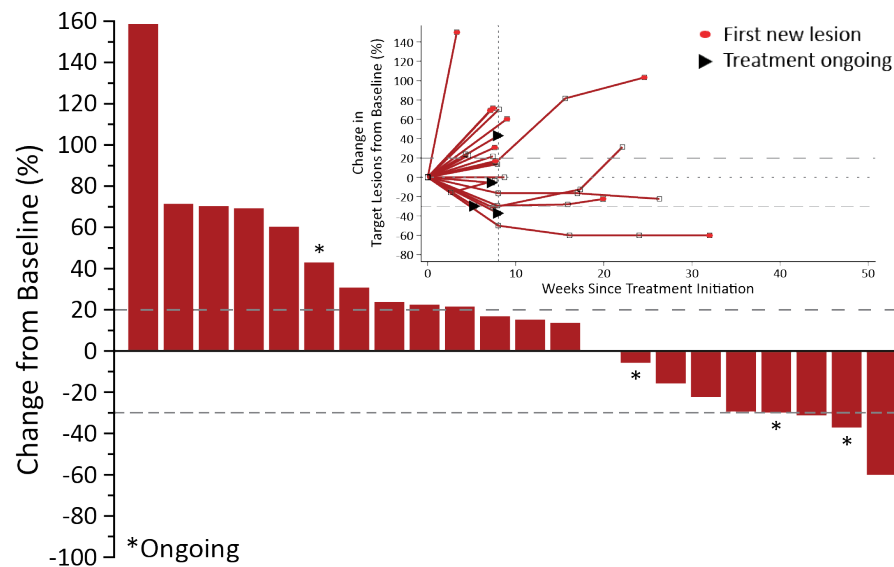
\* Includes MedDRA Preferred Terms of Rash and Maculopapular Rash. \*\* Includes MedDRA Preferred Terms of Pruritus and Generalized Pruritus. Grade 4 drug-related AEs include: lipase increased (n=3), neutrophil count decreased, and IRR (n=1, each). No Grade 5 TRAEs have been reported. AESI = adverse events of special interest. Data cutoff: April, 25, 2020.

Luke, et al., ASCO 2020

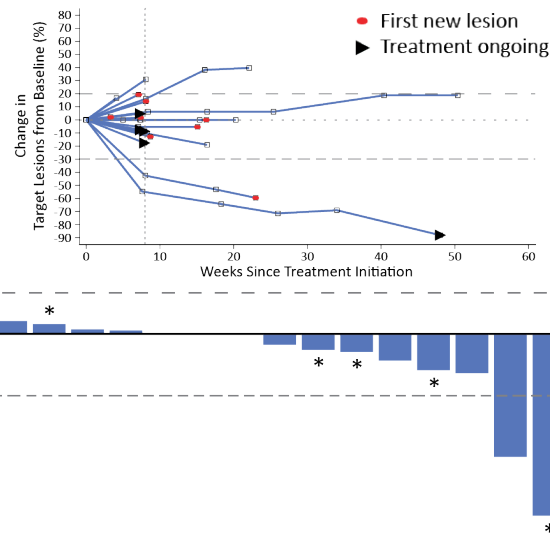


# MGD013 Monotherapy: Anti-tumor Activity Observed in Multiple Tumor Types

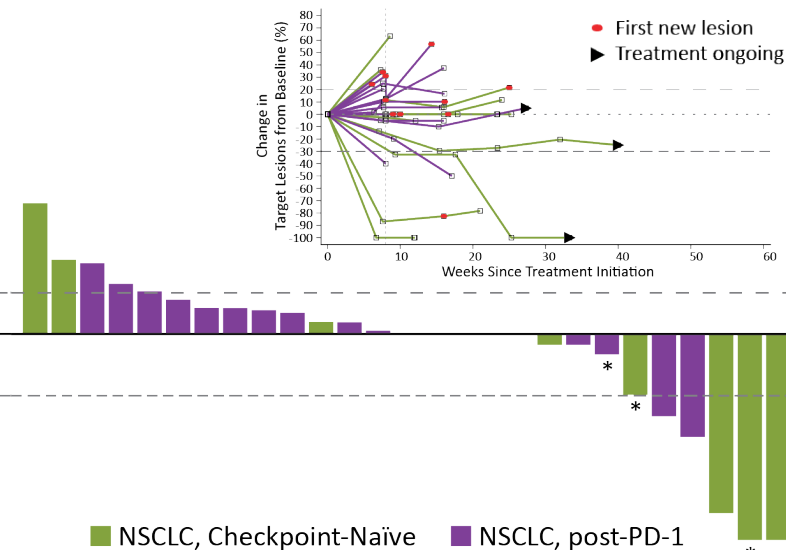
### Triple-negative Breast Cancer



### Epithelial Ovarian Cancer



### Non-small Cell Lung Cancer



	TNBC	EOC	NSCLC, CPI-Naïve	NSCLC, post-PD-1
Evaluable Patients	23	23	14	15
ORR (Confirmed)	4.3% (1/23)	8.7% (2/23)	14.3% (2/14)	0% (0/15)
ORR (Confirmed + Unconfirmed)	17.4% (4/23)	8.7% (2/23)	21.4% (3/14)	13.3% (2/15)
SD	34.8% (8/23)	43.5% (10/23)	50.0% (7/14)	53.3% (8/15)
DCR	39.1% (9/23)	52.2% (12/23)	64.3% (9/14)	53.3% (8/15)

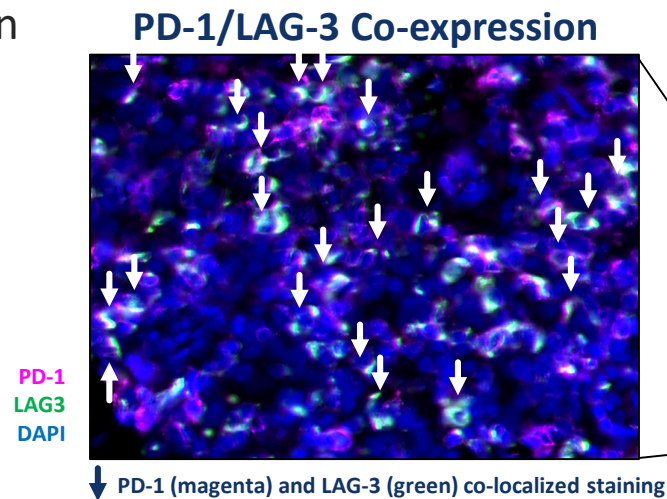
Data cutoff: April, 25, 2020

Luke, et al., ASCO 2020

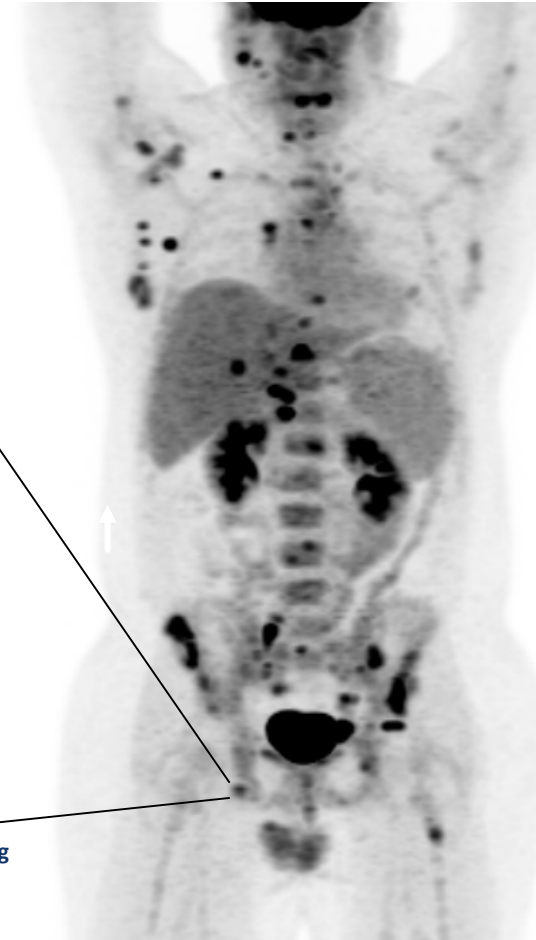
# Complete Response after Single MGD013 Administration

*27-year-old male with DLBCL progressive disease after CAR-T cell therapy*

- Relapsed subsequent to DA-R-EPOCH and JCAR017
- Pre-treatment biopsy: High levels of LAG-3 & PD-L1
- Received MGD013, 600 mg x 1
- Admitted on Day 11 for management of Grade 2 CRS
- CR on Day 24 (per Lugano classification)
- No evidence of CAR-T in circulation
- Allogeneic SCT performed
- Currently in remission:
  - 11 months post-MGD013
  - 9 months post-transplant



Screening



**MGD013**  
Complete Response - Day 24

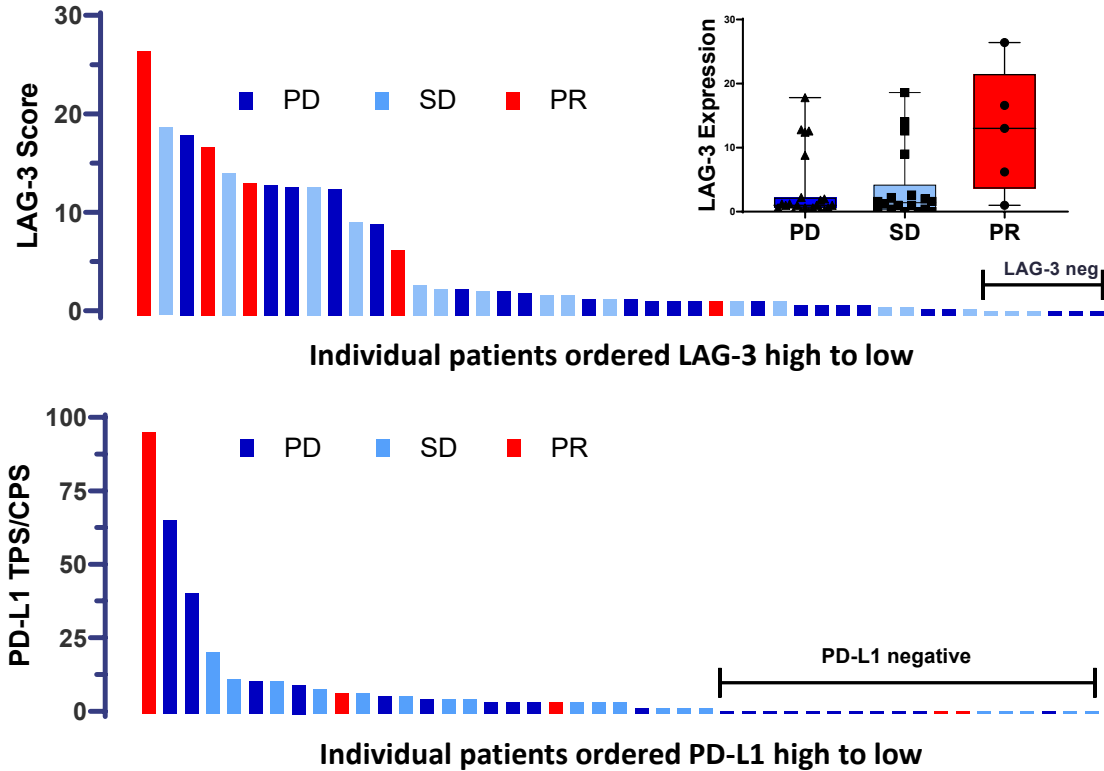


Luke, et al., ASCO 2020

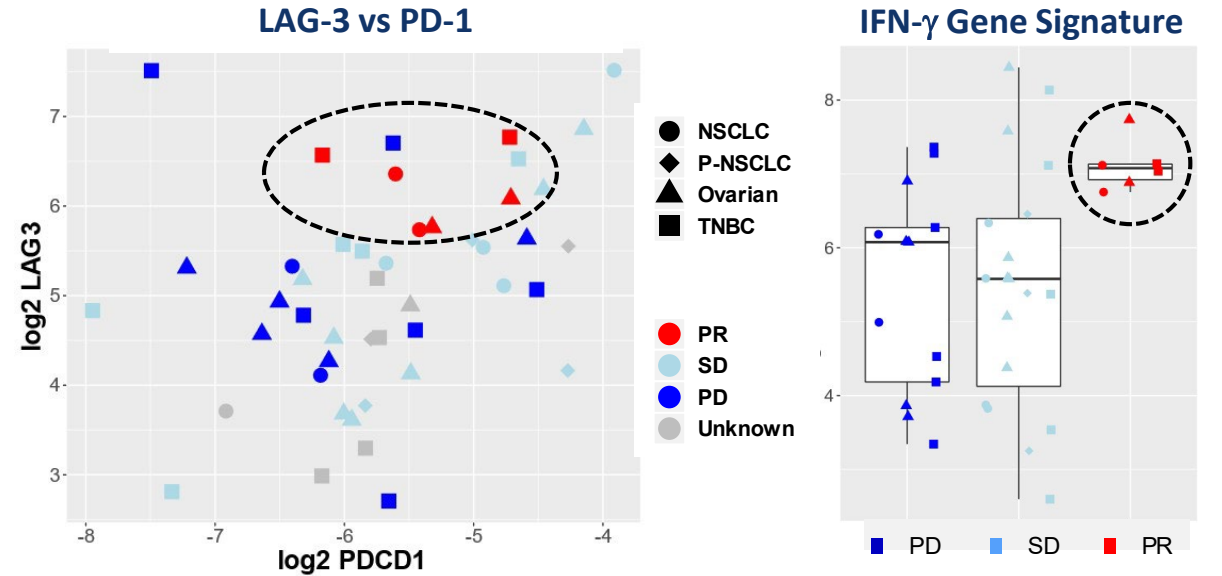
# Monotherapy Objective Responses Associated with LAG-3 Expression

*Inflammatory interferon-γ signature elevated in patients with clinical response*

## Retrospective IHC Analyses



## Transcript Profiling (Baseline Tumor Biopsy)



Objective responses associated with high baseline LAG-3/PD-1 expression and IFN-g gene signature (CXCL9, CXCL10, CXC11, STAT1)

Archival biopsies from TNBC, EOC, and NSCLC expansion cohorts analyzed for LAG-3 (N=46) or PD-L1 (N=45) by IHC. LAG-3 score was determined as per Chen et al., e15086 ASCO 2020. PD-L1 expression was determined per Agilent PD-L1 (22C3) pharmDx kit

The NanoString PanCancer IO 360™ assay was used to interrogate gene expression, including the abundance of 14 immune cell types and 32 immuno-oncology signatures from archival biopsies from EOC (N=14) NSCLC (N=25) and TNBC (N=13) expansion cohorts.

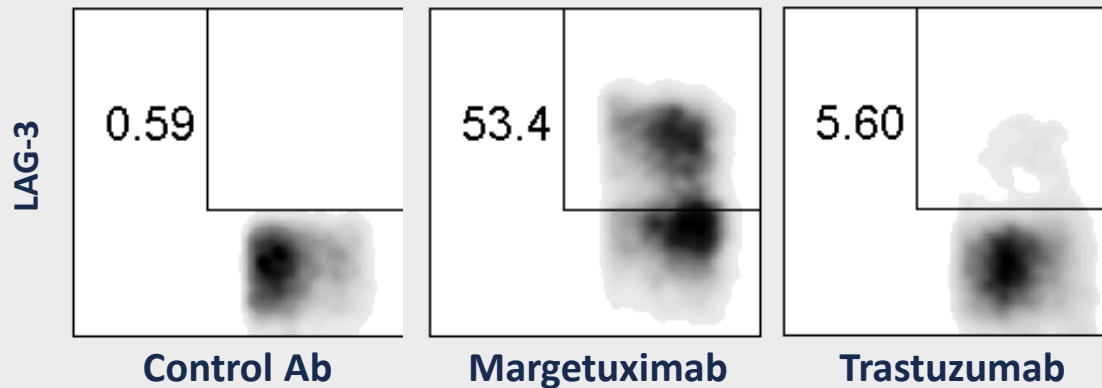
Luke, et al., ASCO 2020

# Can Tumors Be Made More Responsive to PD-1 × LAG-3 Intervention?

*Enhancing effector-cell activation via Fc-engineered mAb*

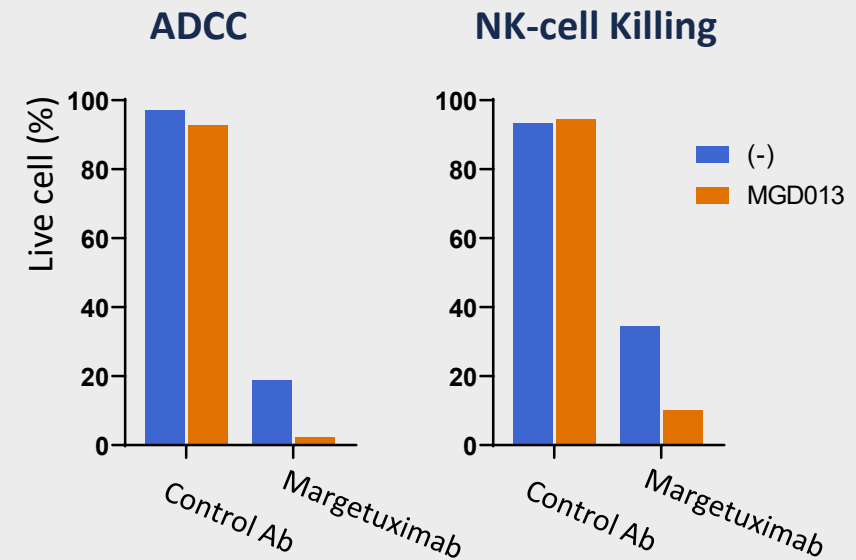
Fc-engineered margetuximab up-regulates LAG-3 and PD-L1 on NK, monocytes and T cells

## Margetuximab Enhances LAG-3 Expression by NK Cells



Human PBMC + N87 (HER2+) gastric cancer cells; E:T=10:1; (IL-2, 20 U/mL)  
Control Ab 50ng/mL, margetuximab/trastuzumab, 5ng/mL; FACS analyses (72h) on CD3<sup>+</sup>CD56<sup>+</sup>-gated NK cells.

PD-1 × LAG-3 (MGD013) enhances lytic activity of immune cells primed by Fc-engineered mAb (margetuximab)

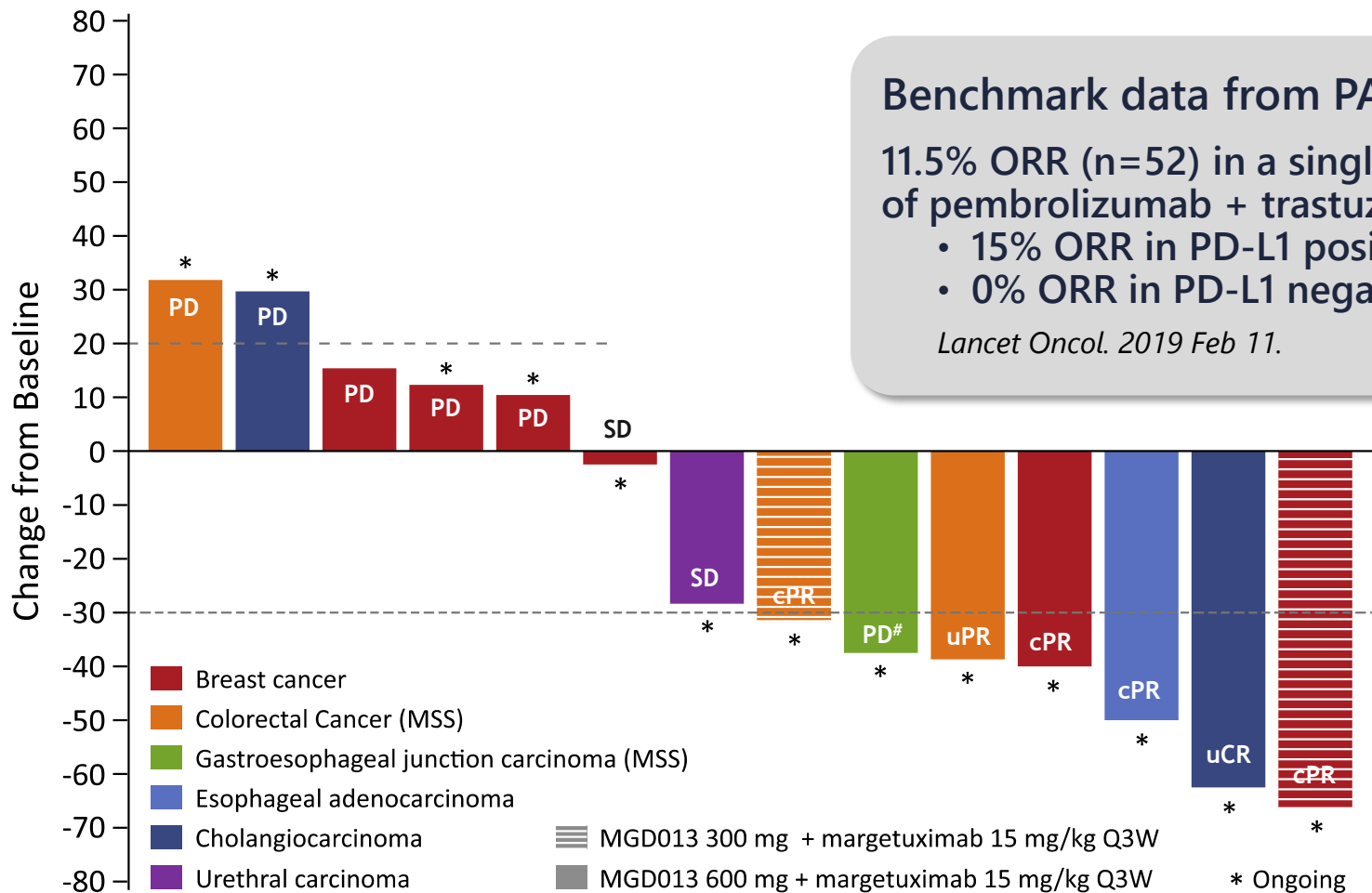


ADCC (target: margetuximab opsonized N87, E:T=10) and NK-cell killing (target: K562, E:T=10) mediated by immune cells activated for 6 days by margetuximab +/- MGD013 in the presence of N87 tumor cells.

Luke, et al., ASCO 2020

# Margetuximab plus MGD013 in Patients with Relapsed/Refractory HER2+ Solid Tumors

Preliminary ORR = 42.9% based on 6/14 evaluable pts (includes unconfirmed objective responses)



**Benchmark data from PANACEA study:**  
 11.5% ORR (n=52) in a single arm, multicenter Ph. 1b/2 trial of pembrolizumab + trastuzumab in HER2+ mBC  
 • 15% ORR in PD-L1 positive (n=6/40)  
 • 0% ORR in PD-L1 negative (n=0/12)  
*Lancet Oncol. 2019 Feb 11.*

**Baseline PD-L1 & LAG-3 in # of Responding Patients (N = 6)**

PD-L1 CPS:	< 1	1	TBD
<b>N</b>	4	1	1

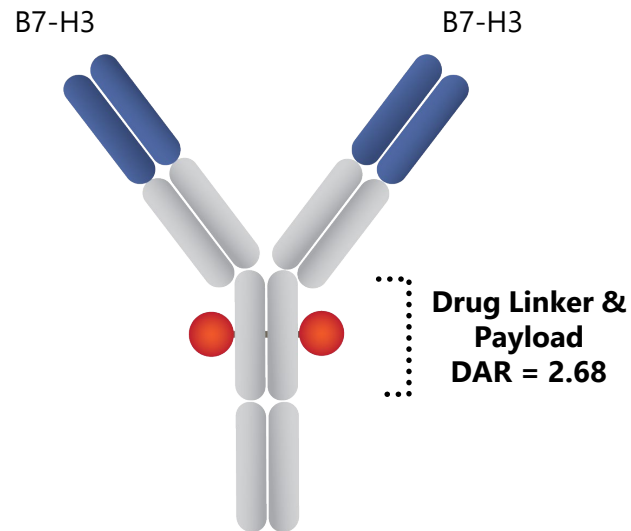
LAG-3 Score:	< 5	5-15	TBD/NE
<b>N</b>	3	1	2

# GEJ pt with apparent pseudo-progression (PD per RECIST), now with 37.5% reduction in target lesions (iPR per iRECIST).

Luke, et al., ASCO 2020

# MGC018: Antibody-Drug Conjugate with Duocarmycin-based Linker Payload

*Leveraging high B7-H3 expression in solid tumors*



## Function/ MoA

- ADC that delivers a potent duocarmycin payload to dividing and non-dividing B7-H3-expressing cells
- Cleavable peptide linker facilitates bystander effect
- Not subject to multi-drug resistance (MDR)

## Clinical Study

- Phase 1 dose escalation in advanced solid tumors (ongoing)

## 2020 Anticipated Milestones

- Initiate Phase 1 dose expansion in metastatic castration-resistant prostate cancer (mCRPC)

*Duocarmycin payload and cleavable peptide linker technology was licensed from Byondis (formerly Synthon Biopharmaceuticals).*

***MGC018 is investigational and has not yet been approved for marketing by any regulatory authority***

# Confirmed High Penetrance in Broad Set of Solid Tumors

Majority of B7-H3 positive tumors express high levels of B7-H3 ( $\geq 2+$ )

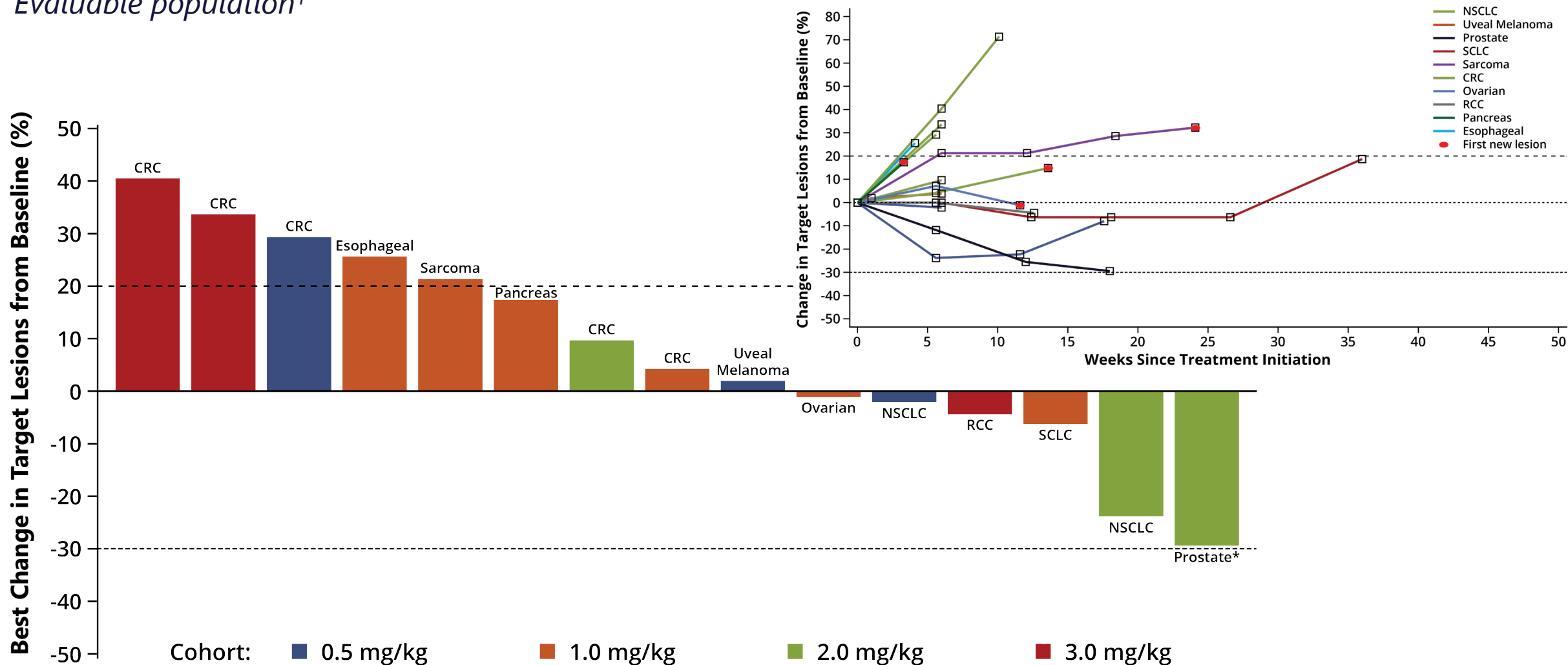
Potential Indications	IHC Summary of >1,400 Tumor Tissue Samples Screened			
	B7-H3 Positive <sup>(a)</sup>		2+ or Above	
Head and Neck	19/19	100%	19/19	100%
Kidney Cancer	77/78	99%	75/78	96%
Glioblastoma	65/66	98%	63/66	95%
Thyroid Cancer	34/35	97%	33/35	94%
Mesothelioma	41/44	93%	39/44	89%
Melanoma	132/146	90%	94/146	64%
<b>Prostate Cancer</b>	<b>88/99</b>	<b>89%</b>	<b>51/99</b>	<b>52%</b>
Pancreas Cancer	69/78	88%	45/78	58%
Bladder	134/156	86%	123/156	79%
Lung Cancer	324/379	85%	300/379	79%
Breast Cancer	189/249	76%	156/249	63%
Ovarian Cancer	59/79	75%	36/79	46%

Limited expression in normal tissue → favorable profile for targeting B7-H3

(a) B7-H3 positivity reflects any grade staining via fixed tumor microarray; B7-H3 is expressed on tumor as well as tumor vasculature.

# Preliminary Evidence of Activity in Multiple Tumor Types

Evaluable population<sup>1</sup>



<sup>1</sup>Patients who received at least one dose and had at least one post-baseline tumor evaluation. \*mCRPC Pt #1. Data were extracted on 06MAY2020.

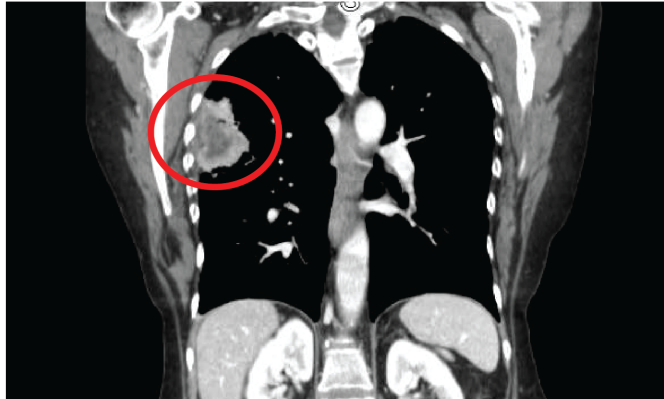
Powderly, et al., ASCO 2020



# Reduction of Pleural-Based Tumor in NSCLC Patient

*MGC018 following progression after five lines of prior therapy*

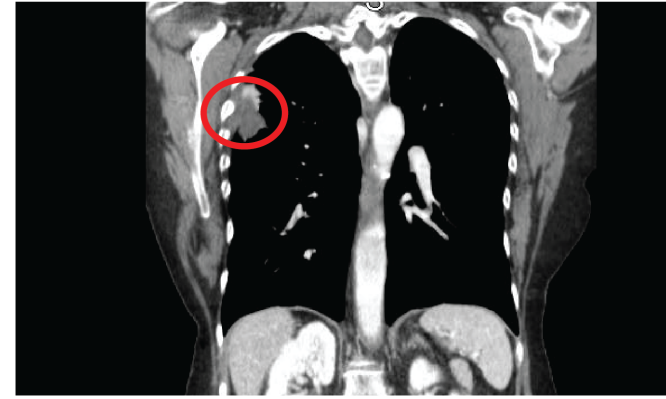
**Baseline (May 23, 2019)**



**2 Doses of MGC018  
(2.0 mg/kg)**  
Decrease in pleural lesion  
read by Investigator



**Week 6 (July 26, 2019)**



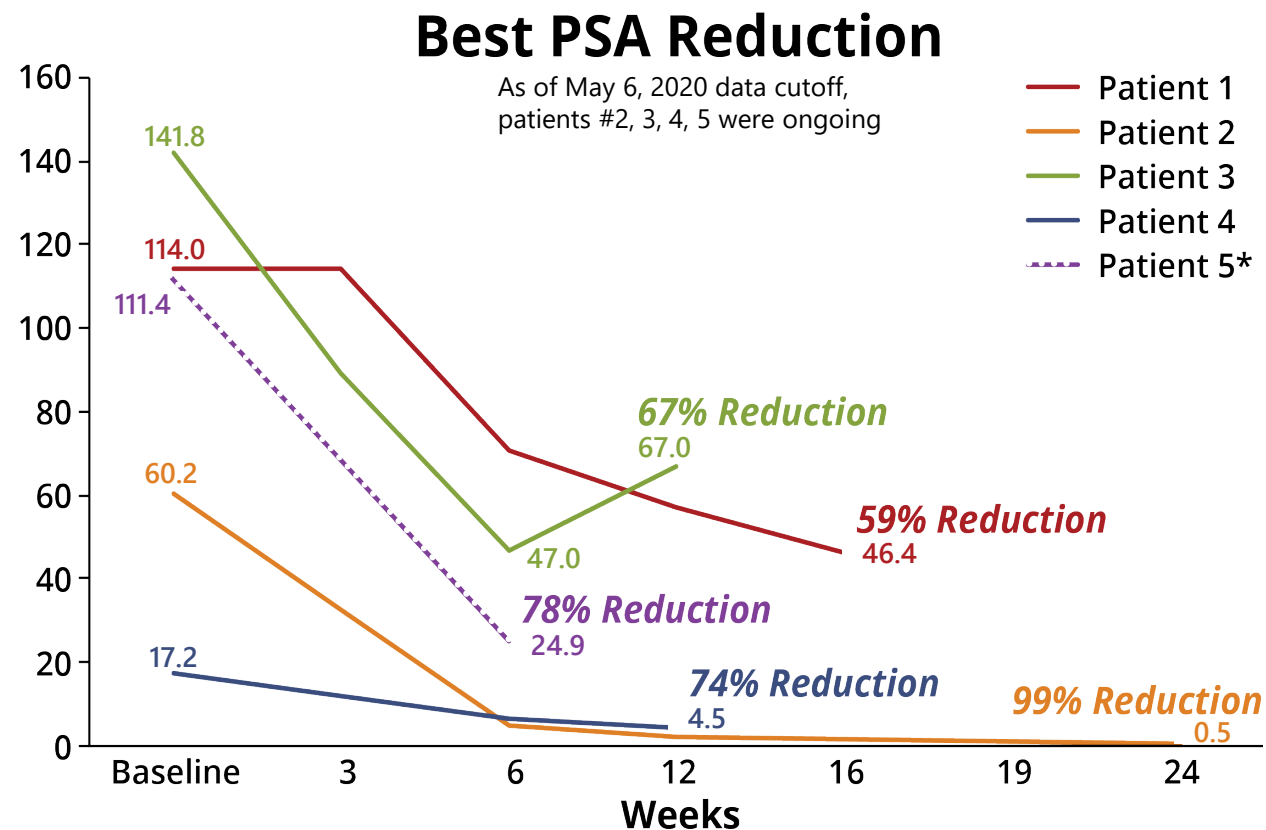
Note image not exact Anterior-Posterior slice as May 23, 2019

Line of Tx	Treatment	Cycles	Duration of Therapy (Months)	Best Response
1	Carboplatin+Paclitaxel+Bevacizumab	4	2	SD
2	Nivolumab	40	16	SD
3	MK-7162 (IDO1 inhibitor)	3	2	SD
4	APG-1252 (Bcl-2 inhibitor)	2	1	PD
5	Pembrolizumab (MK-3475)	2	1	PD
<b>6</b>	<b>MGC018</b>	<b>2</b>	<b>2</b>	<b>SD (~24%)</b>

Powderly, et al., ASCO 2020

# Greater than 50% PSA Decline Following MGC018 in Heavily Pre-treated mCRPC

Patient (Dose)	Line of Therapy	Treatment	Duration of Therapy (months)	MGC018 Response
<b>Patient #1</b> 2 mg/kg One target lesion (lymph node), abdominal adenopathy & bone lesions	1	Docetaxel	4	<b>SD (-29%); 59% PSA Decline</b>
	2	Enzalutamide	24	
	3	Prostvac	5	
	4	Abiraterone	6	
	5	Nivolumab	6	
	6	<b>MGC018</b>	<b>4</b>	
<b>Patient #2</b> 3 mg/kg Bone only disease	1	Docetaxel	6	<b>SD (Ongoing); 99% PSA Decline</b>
	2	Abiraterone	4	
	3	Enzalutamide	12	
	4	Radium 223	6	
	5	<b>MGC018</b>	<b>3+</b>	
<b>Patient #3</b> 3mg/kg Bone only disease	1	Docetaxel	8	<b>SD (Ongoing); 67% PSA Decline</b>
	2	Provenge	2	
	3	Enzalutamide	6	
	4	Abiraterone	9	
	5	<b>MGC018</b>	<b>1.5+</b>	
<b>Patient #4</b> 3 mg/kg Bone only disease	1	Abiraterone	Unknown	<b>SD (Ongoing); 74% PSA Decline</b>
	2	Nivo + Rucaparib	Unknown	
	3	<b>MGC018</b>	<b>1.5+</b>	
<b>Patient #5</b> 3mg/kg Bone only disease	1	Docetaxel	4	<b>SD (Ongoing); 78% PSA Decline</b>
	2	Provenge	12	
	3	Enzalutamide	7	
	4	Abiraterone	7	
	5	Docetaxel	4	
	6	<b>MGC018</b>	<b>1.5+</b>	



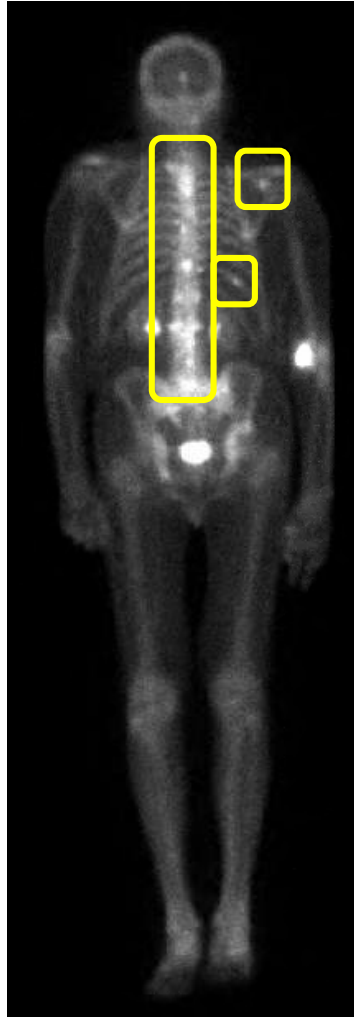
\*Patient #5 data scaled for charting purposes:

Note: Baseline PSA 1,114 ng/mL dropped to 249 ng/mL at Week 6, with a further decline to 95 ng/mL at Week 12 (May 18, 2020; after the May 6, 2020 data cut-off).

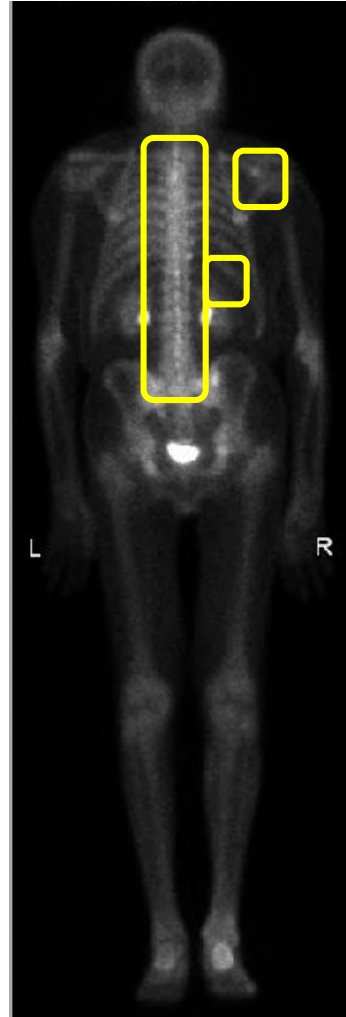
Powderly, et al., ASCO 2020

# 99% PSA Reduction with Substantial Improvement in Metastatic Bone Lesions

*mCRPC Patient #2: Bone lesions of thoracic/lumbar spine, ribs, sternum, and pelvis*



**November 13, 2019**



**February 7, 2020**



**May 1, 2020**

Powderly, et al., ASCO 2020

# Manageable Safety Profile Across Dose Cohorts

*Cytopenias and skin disorders were most common*

## Grade ≥3 Related Adverse Events

System Organ Class Preferred Term	0.5 mg/kg (N=3)	1.0 mg/kg (N=6)	2.0 mg/kg (N=7)	3.0 mg/kg (N=7)	All (N=23)
<b>AT LEAST ONE EVENT</b>	2 (66.7)	2 (33.3)	7 (100)	3 (42.9)	14 (60.9)
<b>Blood and lymphatic system disorders</b>	0	0	2 (28.6)	2 (28.6)	4 (17.4)
Neutropenia	0	0	2 (28.6)	2 (28.6)	4 (17.4)
Lymphopenia	0	0	1 (14.3)	1 (14.3)	2 (8.7)
<b>Gastrointestinal disorders</b>	0	1 (16.7)	0	0	1 (4.3)
Gastrointestinal inflammation	0	1 (16.7)	0	0	1 (4.3)
<b>Investigations</b>	1 (33.3)	2 (33.3)	4 (57.1)	2 (28.6)	9 (39.1)
Lymphocyte count decreased	0	1 (16.7)	2 (28.6)	1 (14.3)	4 (17.4)
Blood alkaline phosphatase increased	0	0	1 (14.3)	1 (14.3)	2 (8.7)
Neutrophil count decreased	0	1 (16.7)	1 (14.3)	0	2 (8.7)
Platelet count decreased	0	0	1 (14.3)	1 (14.3)	2 (8.7)
Lipase increased	1 (33.3)	0	0	0	1 (4.3)
White blood cell count decreased	0	1 (16.7)	0	0	1 (4.3)
<b>Respiratory, thoracic and mediastinal disorders</b>	1 (33.3)	0	0	0	1 (4.3)
Pneumonitis	1 (33.3)	0	0	0	1 (4.3)
<b>Skin and subcutaneous tissue disorders</b>	0	0	3 (42.9)	1 (14.3)	4 (17.4)
Palmar-plantar erythrodysesthesia syndrome	0	0	1 (14.3)	1 (14.3)	2 (8.7)
Rash maculo-papular	0	0	2 (28.6)	0	2 (8.7)
Stasis dermatitis	0	0	1 (14.3)	0	1 (4.3)

Powderly, et al., ASCO 2020

## No Discontinuations Due to Adverse Events at 3.0mg/kg

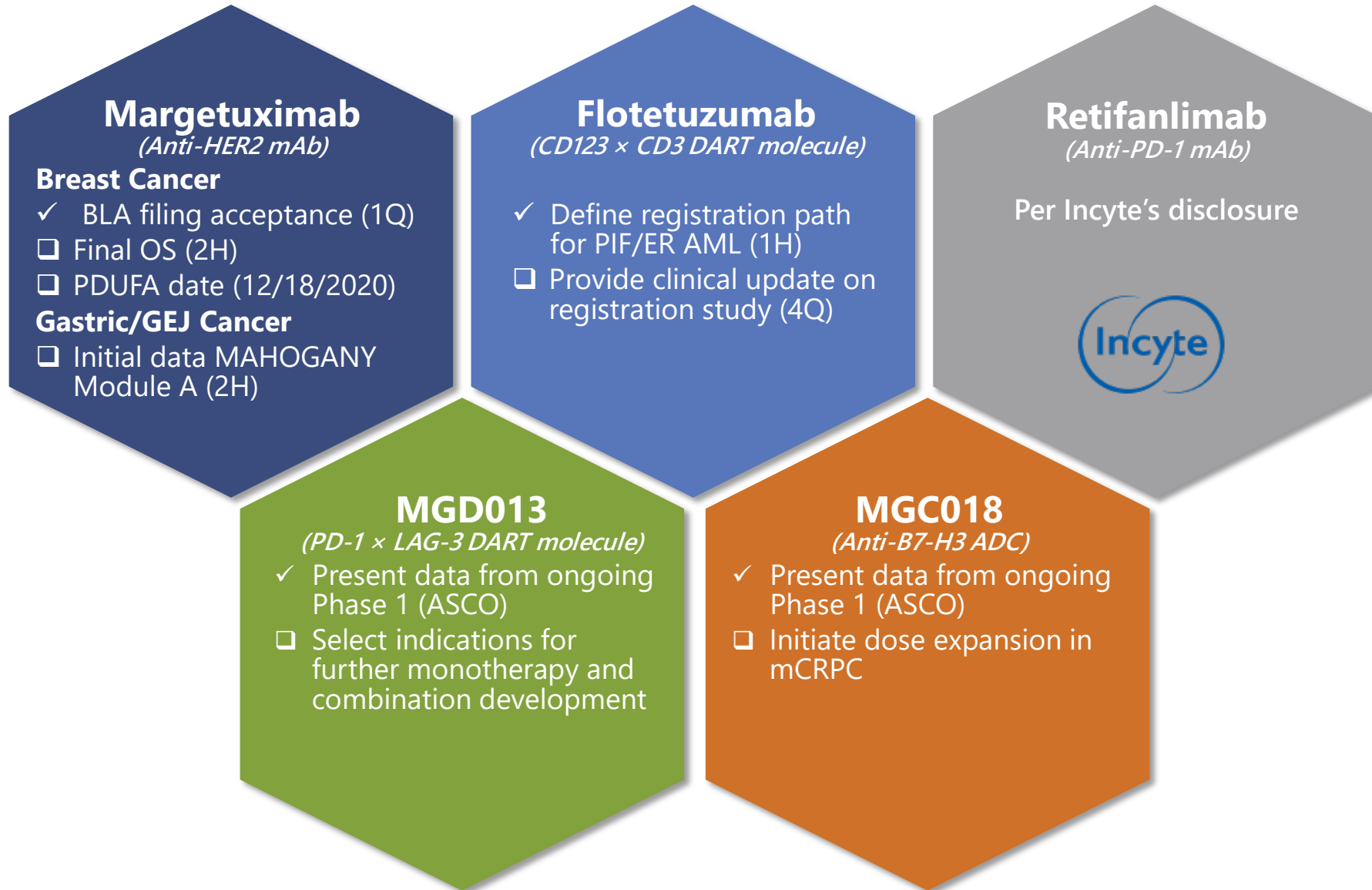
Patients Reporting at Least One Adverse Event	0.5 mg/kg (N=3)	1.0 mg/kg (N=6)	2.0 mg/kg (N=7)	3.0 mg/kg* (N=7)	All (N=23)
Adverse Event	3 (100)	6 (100)	7 (100)	7 (100)	23 (100)
Treatment-Related Adverse Event <sup>1</sup>	3 (100)	4 (66.7)	7 (100)	7 (100)	21 (91.3)
Adverse Event ≥ Grade 3 <sup>2</sup>	3 (100)	4 (66.7)	7 (100)	4 (57.1)	18 (78.3)
Treatment-Related Adverse Event ≥ Grade 3 <sup>2</sup>	2 (66.7)	2 (33.3)	7 (100)	3 (42.9)	14 (60.9)
Serious Adverse Event	1 (33.3)	1 (16.7)	3 (42.9)	0	5 (21.7)
Event that Resulted in Study Discontinuation	1 (33.3)	1 (16.7)	3 (42.9)	0	5 (21.7)
Event that Resulted in Drug MGC018 Withdrawal	1 (33.3)	1 (16.7)	3 (42.9)	1 (14.3)	6 (26.1)
Event that Resulted in Drug MGC018 Dose Reduction	0	0	1 (14.3)	2 (28.6)	3 (13.0)
Event that Resulted in Drug MGC018 Interrupted	1 (33.3)	0	2 (28.6)	5 (71.4)	8 (34.8)
Fatal Adverse Event (pneumonitis)	1 (33.3)	0	0	0	1 (4.3)
Adverse Event of Special Interest (AESI) – Infusion Reaction	0	0	2 (28.6)	5 (71.4)	7 (30.4)

\*Amendment applied to allow dose modification.

- Three treatment-related serious adverse events occurred in three patients:
  - pneumonitis in a patient with concurrent bacterial pneumonia; non-infectious gastroenteritis; and stasis dermatitis in a patient with chronic venous insufficiency
- One dose-limiting toxicity (Grade 4 neutropenia resolved to baseline); no febrile neutropenia observed

<sup>1</sup>Includes events with causality assessments of 'Possible', 'Probable' or 'Definite'. <sup>2</sup>Based on CTCAE criteria version 4.0.3.

# Key Milestones Anticipated in 2020

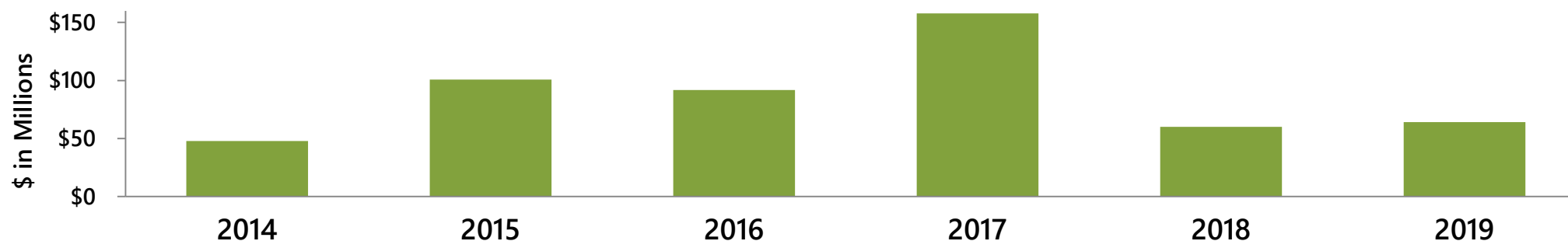


# Financial Overview

- \$171M Cash, cash equivalents and marketable securities as of March 31, 2020
  - Cash runway into 2022 via anticipated and potential collaboration payments
- Historical financial details:

\$ in Millions	2014	2015	2016	2017	2018	2019	1Q Ended March 31,	
							2020	2019
Total Revenues	\$48	\$101	\$92	\$158	\$60	\$64	\$14	\$10
R&D Expense	70	98	122	147	191	195	49	47
Total Operating Expenses	86	121	152	180	231	241	59	57
Cash & Investments	158	339	285	305	233	216	171	320

- Revenues from collaborative and government agreements (> \$525M since 2013 IPO):



# Thank You!

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