# Merus closing in on cancer

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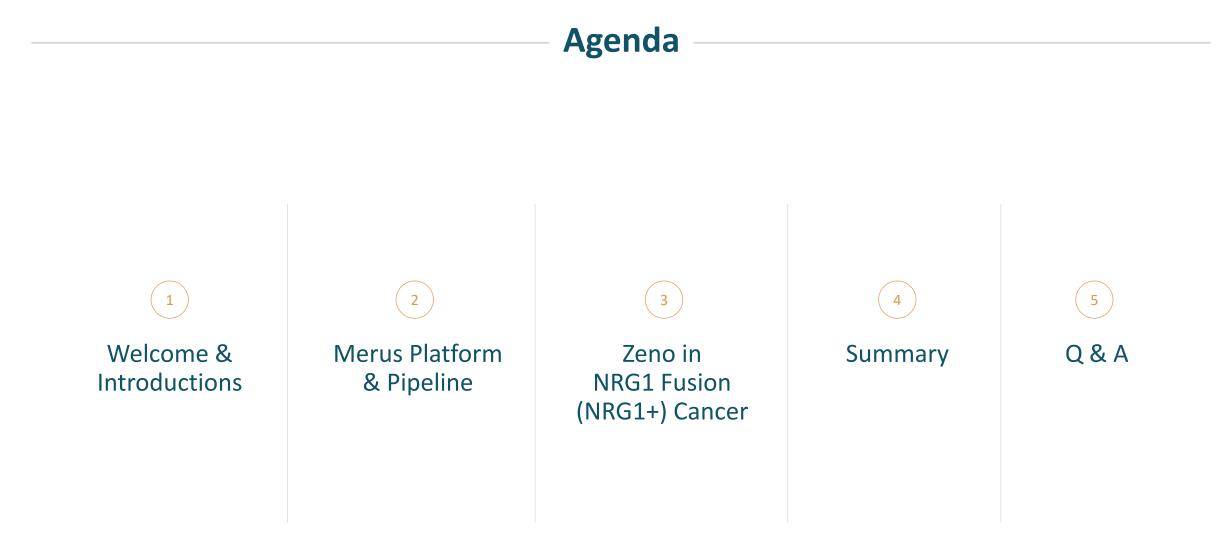
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This presentation (including any oral commentary that accompanies this presentation) contains forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995. All statements contained in this presentation that do not relate to matters of historical fact should be considered forward-looking statements, including without limitation statements regarding the impact our Biclonics<sup>®</sup> and Triclonics<sup>®</sup> platforms can have on cancer, our product candidates' potential to treat certain types of tumors, the timing of regulatory filings and the timing and anticipated data read outs or results from our clinical trials and our collaborations, and anticipated cash runway. These forward-looking statements are based on management's current expectations. These statements are neither promises nor guarantees, but involve known and unknown risks, uncertainties and other important factors that may cause our actual results, performance or achievements to be materially different from any future results. performance or achievements expressed or implied by the forward-looking statements, including, but not limited to, the following: we have incurred significant losses, are not currently profitable and may never become profitable; our need for additional funding, which may not be available and which may require us to restrict out operations or require us to relinquish rights to our technologies or antibody candidates; potential delays in regulatory approval and impacts of the COVID-19 pandemic, which would impact our ability to commercialize our product candidates and affect our ability to generate revenue; the unproven approach to therapeutic intervention of our Biclonics<sup>®</sup>, and Triclonics<sup>®</sup> technology; our limited operating history; economic, political, regulatory and other risks involved with international operations; the lengthy and expensive process of clinical drug development, which has an uncertain outcome; the unpredictable nature of our early stage development efforts for marketable drugs; potential adverse public reaction to the use of cancer therapies; potential delays in enrollment of patients, which could affect the receipt of necessary regulatory approvals; failure to obtain marketing approval internationally; failure to compete successfully against other drug companies; potential competition from other drug companies if we fail to obtain orphan drug designation or maintain orphan drug exclusivity for our products; our reliance on third parties to conduct our clinical trials and perform clinical development and the potential for those third parties to not perform satisfactorily; our reliance on third parties to manufacture our product candidates, which may delay, prevent or impair our development and commercialization efforts; protection of our proprietary technology; our patents being found invalid or unenforceable; potential lawsuits for infringement of third-party intellectual property; our ability to attract and retain key personnel; managing our growth could result in difficulties.

These and other important factors discussed under the caption "Risk Factors" in our in our Annual Report on Form 10-Q for the period ended March 31, 2022 filed on May 9, 2022 with the Securities and Exchange Commission, or SEC, and our other reports filed with the SEC, could cause actual results to differ materially from those indicated by the forward-looking statements made in this presentation. Any such forward-looking statements represent management's estimates as of the date of this presentation. While we may elect to update such forward-looking statements at some point in the future, we disclaim any obligation to do so, even if subsequent events cause our views to change.





# Q&A



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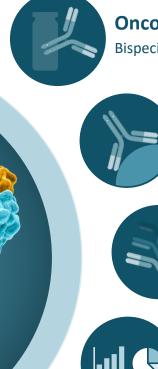
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# **Merus Overview**



#### Oncology-focused Company Developing Multispecific Antibody Therapies

Bispecific and trispecific cancer therapeutic candidates based on the human IgG format



Four clinical stage assets with proof of concept data on Zeno in NRG1-fusion cancer<sup>1</sup>, and early encouraging data on Peto in Head and Neck cancer<sup>2</sup>



#### Leading Multispecific Antibody (Multiclonics®) Platforms

Common light chain format permits broad high throughput evaluation of Biclonics<sup>®</sup> and Triclonics<sup>®</sup>, to develop clinical stage assets with meaningful clinical responses in patients



#### Near Term Trial Updates and Strong Cash Position Beyond 2024\*

Upcoming clinical milestones, and program updates planned over the next 12-18 months: Zeno registrational directed program, Peto clinical update, and MCLA-129 initial clinical data



#### Strategic Collaborations to Unlock Platform Value

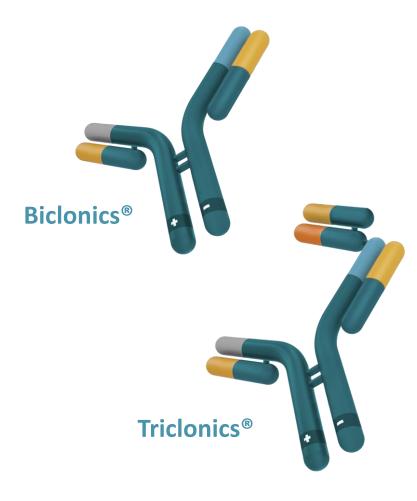
Multiple strategic collaborations and license agreements

<sup>1</sup> Schram et al., ASCO 2022, efficacy data cutoff April 12, 2022, safety cutoff January 12, 2022
 <sup>2</sup> Hollebecque et al., AACR-NCI-EORTC 2021, data cutoff August 9, 2021
 \*See May 9, 2022 10-Q noting our belief that our cash, cash equivalents and marketable securities, will fund our operations beyond 2024.



# **Merus Multiclonics®**

Bispecific and Trispecific therapeutic candidates for cancer with broad application for human disease



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#### Large-scale screening

*To select the best Biclonics® and Triclonics® from up to 1,000s of candidates* 

#### Fully human IgG format

- Ease of manufacturing
- Low immunogenicity risk
- Predictable in vivo behavior
- Durable, consistent half life
- Potential for ADCC enhancement and Fc silencing

#### **Robust IP portfolio**

Patents covering Multiclonics® technology, including

- Common light chain antibody generation and screening
- Dimerization by charge engineering

# **Merus Clinical Pipeline**

PROGRAM	BISPECIFIC TARGETS	INDICATION(S)	PRECLINICAL	PHASE 1	PHASE 1/2	STATUS
Zenocutuzumab (Zeno) (MCLA-128)	HER2 x HER3	NRG1+ cancer				<ul> <li>Phase 1/2 registration- directed trial ongoing</li> <li>Clinical update at ASCO 2022</li> </ul>
Petosemtamab (Peto) (MCLA-158)	LGR5 x EGFR	Solid tumors				<ul> <li>Phase 1 trial ongoing</li> <li>Clinical update planned 2H22</li> </ul>
MCLA-145	CD137 x PD-L1	Solid tumors				<ul> <li>Phase 1 trial ongoing</li> </ul>
MCLA-129	EGFR x c-MET	Solid tumors	(China)			<ul> <li>Phase 1/2 trial ongoing</li> <li>Clinical update planned 2H22</li> </ul>
ONO-4685*	PD-1 x CD3	Relapsed/Refractory T Cell Lymphoma; Psoriasis	ono			• Phase 1 trial ongoing





# Efficacy and safety of zenocutuzumab, a HER2 x HER3 bispecific antibody, in advanced *NRG1* fusion-positive (NRG1+) cancer

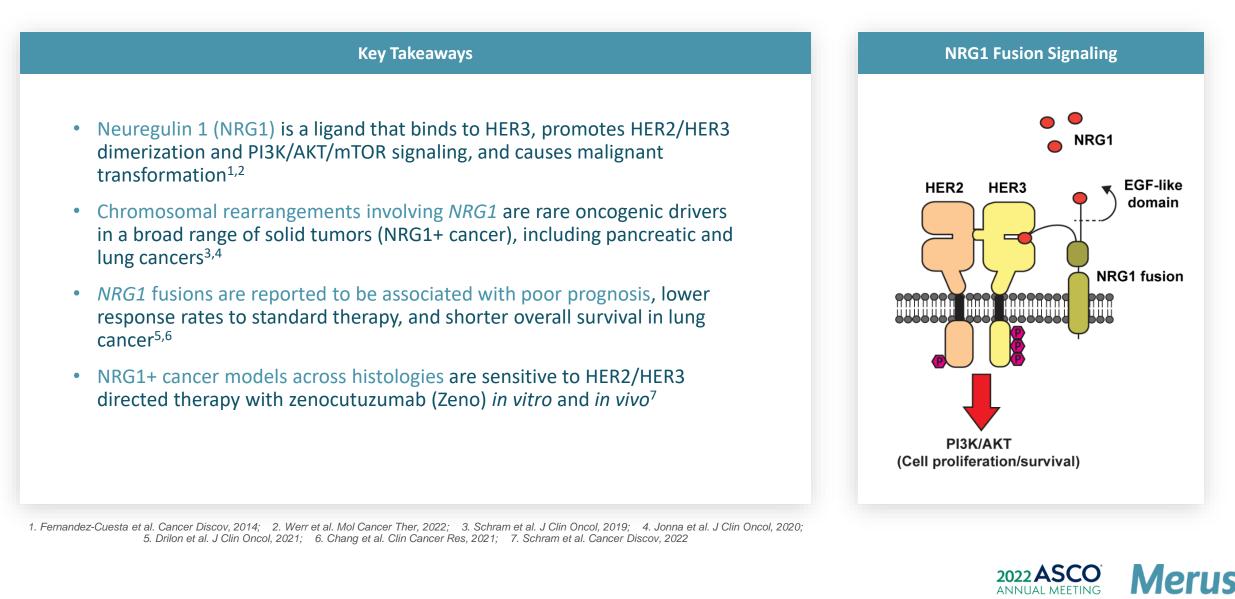
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 Princess Margaret Cancer Centre, Toronto, ON, Canada;
 St. Marianna University School of Medicine, Kawasaki, Japan;
 URCOT, Hôpital Louis Pradel, Hospices Civils de Lyon Cancer Institute, Lyon, France;
 Curie Institute, Saint-Cloud, France;
 Netherlands Cancer Institute, Amsterdam, Netherlands;
 The University of Texas MD Anderson Cancer Center, Houston, TX, USA;
 Karmanos Cancer Institute, Detroit, MI, USA;
 Ruesch Center for the Cure of Gastrointestinal Cancers, Lombardi Comprehensive Cancer Center, Georgetown University Medical Center, Washington, DC, USA;
 Hospital Vall d'Hebron, Barcelona, Spain;
 Merus NV, Utrecht, Netherlands



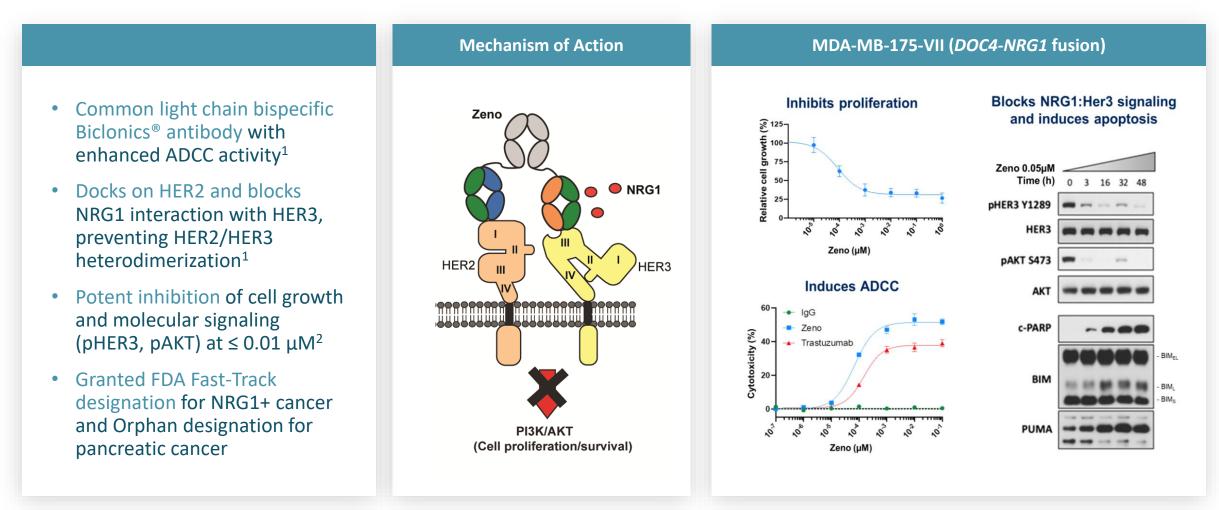
# Background

#### NRG1 Fusions are Clinically Actionable Targets



# Background

#### Zenocutuzumab is a Novel Therapeutic for NRG1+ Cancer

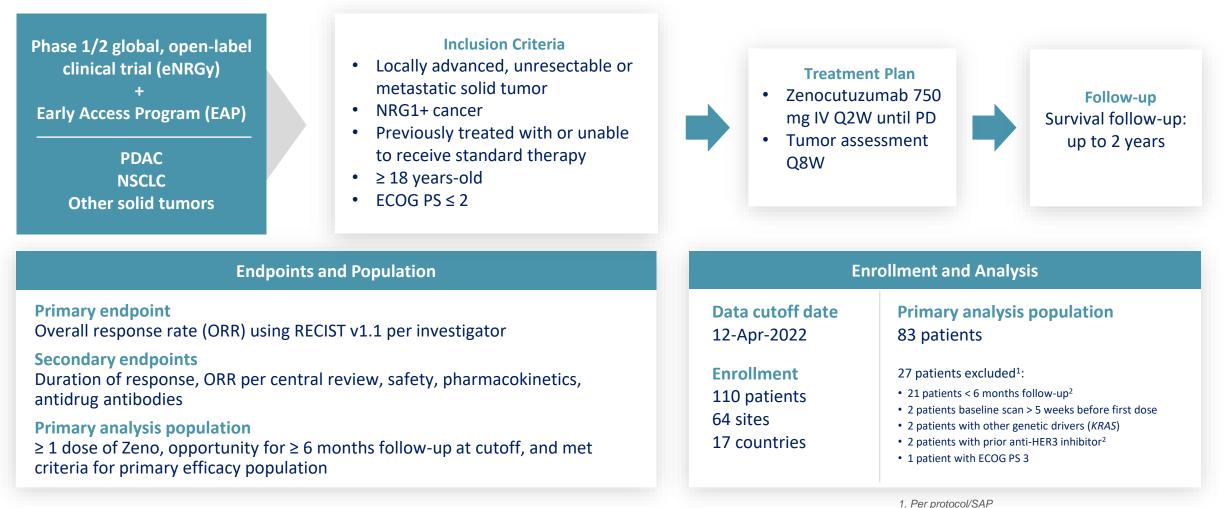


1. Geuijen et al. Cancer Cell, 2018; 2. Schram et al. Cancer Discov, 2022



# Schema

#### Global, Multicenter Zenocutuzumab Development Program



2. One patient had 2 reasons for exclusion



# **Patient Population**

#### Demographics and Disease Features of NRG1+ Cancer

All Patients (N=83)				
Enrolled in eNRGy trial / EAP, n (%)	72 (87) / 11 (13)			
Age in years, median (range)	59 (22 - 84)			
Male / female, n (%)	34 (41) / 49 (59)			
ECOG PS 0 / 1 / 2, n (%)	35 (42) / 47 (57) / 1 (1)			
Race, n (%) <sup>1</sup>				
White	47 (57)			
Asian	27 (33)			
Other	3 (4)			

1. Data not reported for 6 patients

All Patients (N=83)	
Metastatic disease, n (%)	82 (99)
Measurable disease, n (%)	79 (95)
Primary tumor, n (%)	
NSCLC <sup>2</sup>	47 (57)
PDAC	19 (23)
Breast cancer	7 (8)
Cholangiocarcinoma	3 (4)
CRC	3 (4)
Other <sup>3</sup>	4 (5)

2. Adenocarcinoma (N=42), IMA (N=4), mixed adeno-squamous carcinoma (N=1)

3. Endometrial soft tissue sarcoma, pancreatic neuroendocrine carcinoma, renal cell carcinoma, unknown primary



# **Patient Population and Disposition**

#### Prior Therapy, Diagnostics, and Molecular Features of NRG1+ Cancer

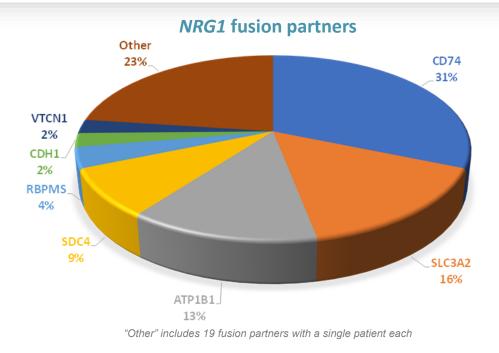
All Patients (N=83)	
Prior systemic therapy	
N lines, median (range) <sup>1</sup>	2 (0 - 8)
Prior afatinib, n (%)	9 (11)
Patient disposition	
Treatment ongoing, n (%)	20 (24)
Reason for discontinuation, n (%)	
Disease progression <sup>2</sup>	61 (73)
Other <sup>3</sup>	2 (2)
Duration of exposure, months	
Median (range)	6.3 (1 - 21)

1. 11 patients were treatment-naïve in the metastatic setting

2. Includes radiological and clinical progression

3. Unrelated AE of dyspnea due to underlying progression, pregnancy

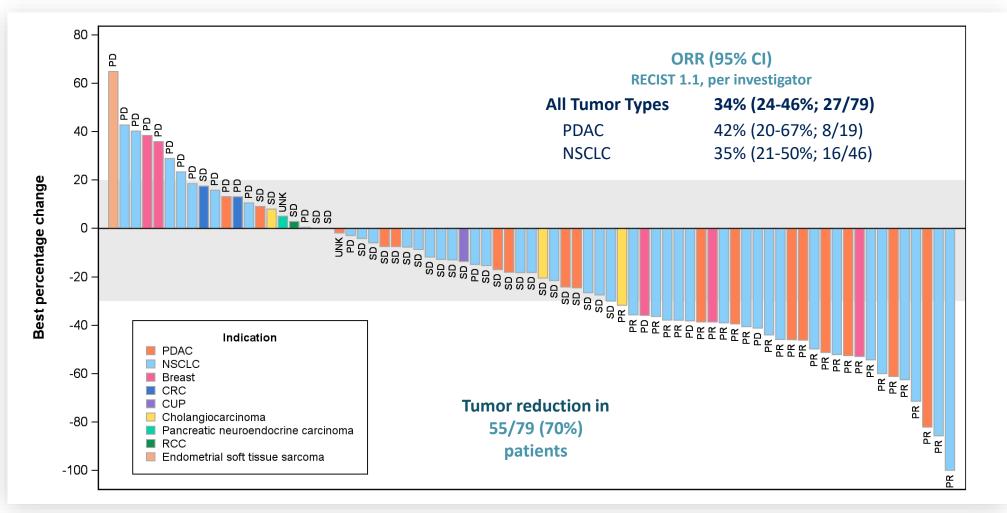
All Patients (N=83)	
NRG1 identification technology, n (%)	
RNAseq	64 (77)
DNAseq	18 (22)
Nanostring	1 (1)





# Zenocutuzumab Activity in NRG1+ Cancer

Best Percent Change in Target Lesions from Baseline

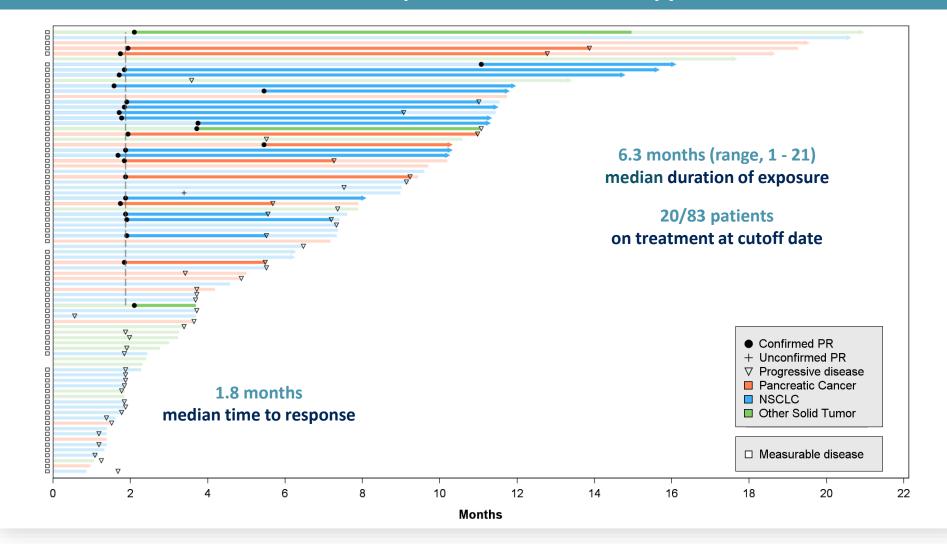


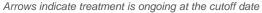
*Note:* The waterfall plot shows data for 75 of 79 patients. Change in tumor size could not be measured for 4 of the 79 patients, 3 due to absence of post baseline assessment (early progression) and one due to incomplete assessment.



# Zenocutuzumab Activity in NRG1+ Cancer

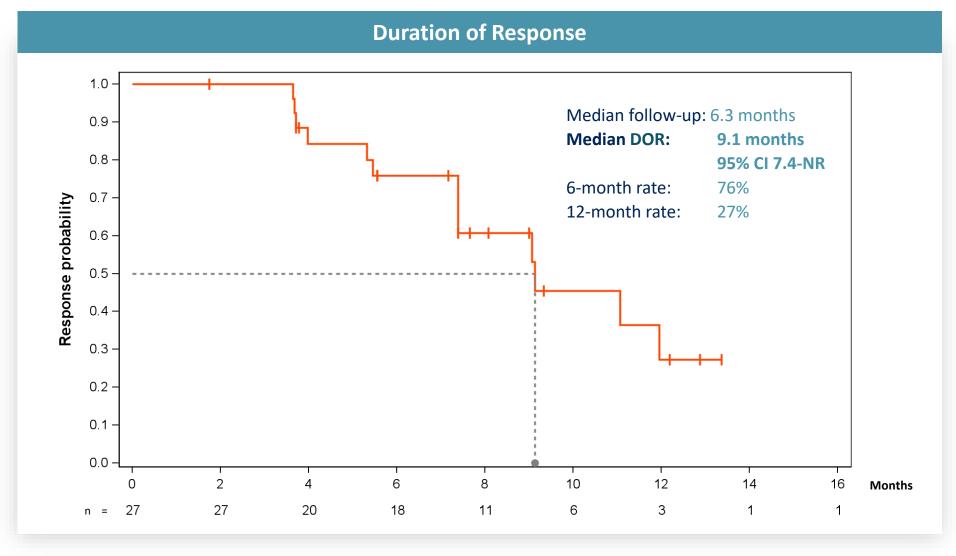
Time to Response and Time on Therapy







# Zenocutuzumab Activity in NRG1+ Cancer

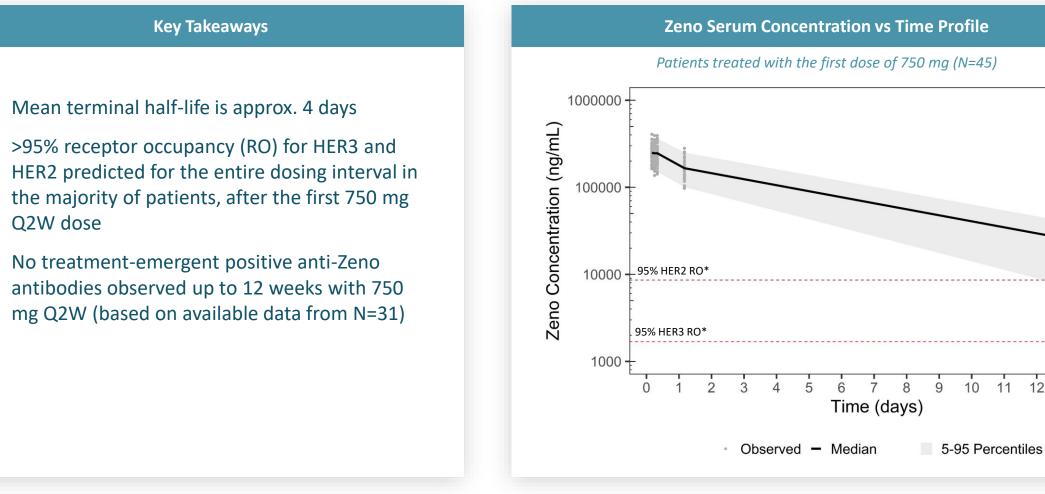


Note: 20/83 patients were on treatment at cutoff date



# **Pharmacokinetics and Immunogenicity**

#### Zenocutuzumab achieved substantial receptor occupancy and was not immunogenic



\* RO based on average of KD values for binding affinities. Geuijen et al. Cancer Cell, 2018.



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Q2W dose

# Safety Profile

#### Zenocutuzumab is well tolerated

Key Takeaways		AEs Irrespective of Causality (>10%)			Treatment-Related AEs (>10% and all Grade 3-5		
		ALL GRADES	GRADE 3-4	GRADE 5	ALL GRADES	GRADE 3-4 <sup>2</sup>	GRADE 5
Safety profile of 208	Patients with ≥1 AE	92%	36%	3%	61%	5%	0.5%
	Diarrhea	32%	2%	-	21%	0.5%	-
atients treated with	Asthenia/fatigue	30%	4%	-	12%	0.5%	-
eno at the RP2D <sup>1</sup> in	Nausea	20%	1%	-	10%	0.5%	-
	Anemia	19%	3%	-	1%	-	-
e single agent	Infusion-related reaction <sup>3,4</sup>	15%	1%	0.5%	15%	1%	0.5% <sup>3</sup>
rogram	Dyspnea	14%	4%	-	2%	0.5%	-
·	Vomiting	13%	0.5%	-	4%	-	-
Low incidence of Grade	Abdominal pain	12%	1%	-	2%	0.5%	-
3 treatment-related	Constipation	11%	-	-	2%	-	-
AEs	Decreased appetite	10%	0.5%	-	4%	-	-
L3	AST increase	9%	3%	-	2%	0.5%	-
Low incidence of severe gastrointestinal and skin toxicity, and no clinical cardiotoxicity	Cough	8%	0.5%	-	1%	0.5%	-
	ALT increase	7%	3%	-	1%	0.5%	-
	Myalgia	4%	0.5%	-	2%	0.5%	-
	Neutropenia	3%	1%	-	2%	0.5%	-
	Hypertension	1%	1%	-	0.5%	0.5%	-
, ,	Platelet count decrease	1%	0.5%		0.5%	0.5%	-
<1% of patients discontinued due to AEs	Hyperuricemia	0.5%	0.5%	-	0.5%	0.5%	-
	Lymphadenitis	0.5%	0.5%	-	0.5%	0.5%	-
	Нурохіа	0.5%	0.5%	-	0.5%	0.5%	-
	Bacteremia	0.5%	0.5%	-	0.5%	0.5%	-

Safety data cut off:12-Jan-2022 1. 101 patients with 750 mg Q3W; 26 patients with QW; 81 patients with 750 mg Q2W

2. No Grade 4 treatment-related AEs reported

One Grade 5 hypersensitivity (previously reported; Alsina et al. ASCO, 2017)
 Composite term covering preferred terms considered by the investigator to be IRRs occurring within 24 hours of infusion start



# Zenocutuzumab

**Conclusions** 



#### Durable responses in previously treated advanced NRG1+ cancer

- ORR 34% overall (n=79; 95% CI: 24-46%)
- Median DOR 9.1 months (95% CI: 7.4-NR)
- Antitumor activity across multiple tumor types



#### Extremely well tolerated safety profile

- Most adverse events were low grade
- Very low rate of discontinuations due to toxicity



#### Offers potential new standard of care for patients with NRG1+ cancer

- Currently no approved targeted therapy for NRG1+ cancer
- Significant unmet medical need



# Launch Plan for Potentially First and Best in Class Treatment

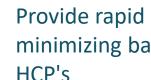
Establish Zeno as the Standard of Care for NRG1+ Cancer





Maximize upfront testing for NRG1 to identify appropriate patients and guide treatment decisions





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Provide rapid access to Zeno by minimizing barriers for patients & HCP's



# **Innovative Merus Technology Platform**

#### Zeno is potentially a new NRG1+ specific & tumor agnostic standard of care





Zeno has demonstrated clinical activity across multiple NRG1+ tumor types and fusion partners Zeno is **welltolerated** and has a patient-friendly every other week dosing regimen



On track for a **BLA filing for Zeno in NRG1+ cancer** for patients who have received prior standard therapy



Potential first and best in class medicine for NRG1+ cancer



# Q&A



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