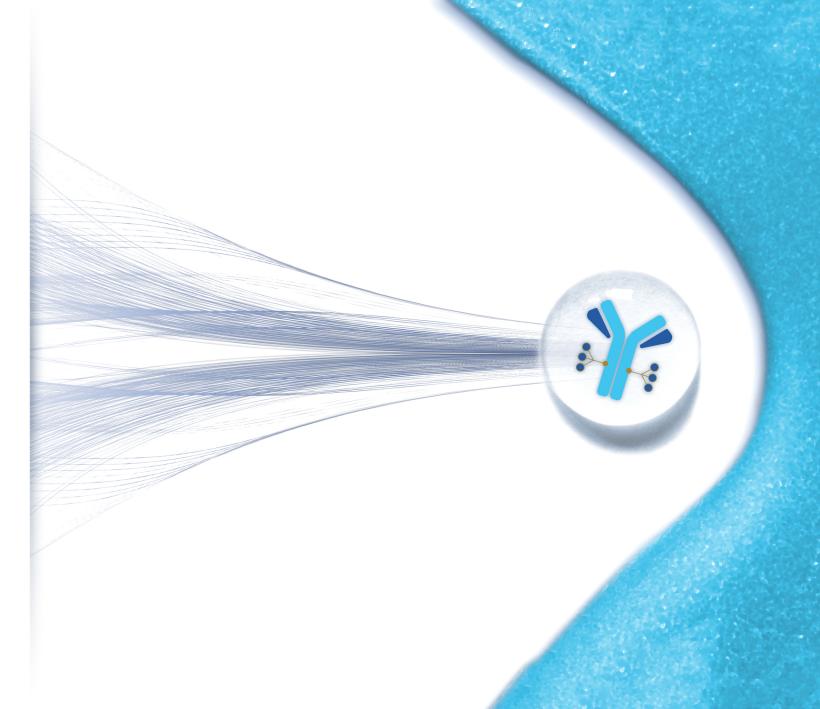


# Accelerating ADC Innovation

...because patients are waiting



March 2020

### **Legal Disclaimer**



This presentation contains "forward-looking" statements within the meaning of federal securities laws. These forward-looking statements are not statements of historical facts and are based on management's beliefs and assumptions and on information currently available to management. Forward-looking statements include information concerning the Company's business strategy and the design, progression and timing of its clinical trials.

Forward-looking statements generally can be identified by terms such as "expects," "anticipates," "believes," "could," "seeks," "estimates," "intends," "may," "plans," "potential," "predicts," "projects," "should," "will," "would" or similar expressions and the negatives of those terms. The Company's operations involve risks and uncertainties, many of which are outside its control, and any one of which, or combination of which, could materially affect its results of operations and whether the forward-looking statements ultimately prove to be correct. Factors that may materially affect the Company's results of operations and whether these forward-looking statements prove to be correct include, among other things, that preclinical testing may not be predictive of the results or success of ongoing or later preclinical or clinical trials, that the development of the Company's product candidates and new platforms will take longer and/or cost more than planned and that the identification of new product candidates will take longer than planned, as well as those listed in our Annual Report on Form 10-K filed on February 28, 2020, with the Securities and Exchange Commission ("SEC") and subsequent SEC filings. Except as required by law, the Company assumes no obligation to update these forward-looking statements publicly, or to update the reasons actual results could differ materially from those anticipated in the forward-looking statements, even if new information becomes available in the future.

Copies of the Company's Annual Report on Form 10-K and our other SEC filings are available by visiting EDGAR on the SEC website at http://www.sec.gov.

### Mersana is Poised for a Transformational 2020



#### **XMT-1536**

## On Track for Near-Term Proof of Concept

- First-in-class asset
- Clinically-Validated
- Wholly-Owned<sup>1</sup>
- Fast-to-market strategy

# First-In-Class Pipeline

1 IND and 2 Development Candidates in 2020

- Addressing unmet patient needs
- Fast-to-market strategies

### **Innovative Platforms**

DolaLock (Dolaflexin, Dolasynthen) and Immunosynthen

- Multiple partnering opportunities
- Efficient product engines

### **Strong Foundation**

~\$100M in Cash<sup>2</sup> +\$15M Credit Facility

- Experienced team
- Operating plan expected to fund important milestones into mid-2021<sup>3</sup>

<sup>&</sup>lt;sup>1</sup> Excluding Brazil

<sup>&</sup>lt;sup>2</sup>Cash, Cash Equivalents, and Marketable Securities as of December 31, 2019

<sup>&</sup>lt;sup>3</sup> Milestones include XMT-1536 Phase 1 clinical study and planned XMT-1592 dose escalation study

### 2020 Will Be a Data Rich Year



#### 2019 ACCOMPLISHMENTS

#### **2020 PRIORITIES**

**IND Candidate** 

**XMT-1536** 

- Established proof of activity & tolerability
- Establish proof of concept

(XMT-1592)

- Established preclinical proof of concept
- Rapid dose escalation

DolaLock **Development** Candidate

Advanced through discovery

Progress into IND-enabling studies

**Immunosynthen** Development Candidate

Advanced through discovery

Select first development candidate

# We are Leveraging our Novel ADC Platforms to Generate Differentiated Product Candidates



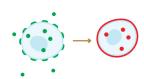
ADC Program	Target	Indication	Platform	Discovery	Preclinical	P1 Dose Escalation	P1 Proof of Concept	P2/Pivotal Study
XMT-1536	NaPi2b	Ovarian Cancer NSCLC Adenocarcinoma	Dolaflexin					
XMT-1592	NaPi2b	NSCLC Adenocarcinoma Ovarian Cancer	Dolasynthen					
To Be Named	B7-H4	Multiple Solid Tumors	Dolaflexin or Dolasynthen					
To Be Named	Multiple	Multiple Solid Tumors	Immunosynthen					
To Be Named	Multiple	Undisclosed	Dolasynthen					
To Be Named	Multiple	Undisclosed	Dolaflexin					
Platform Collabora	tors							
Multiple <b>Serono</b>	Multiple	Undisclosed	Dolaflexin					
ASN004 ASANA BIOSCIENCES	5T4	Undisclosed	Dolaflexin					

# Innovative and Highly Differentiated ADC Technologies and Platforms



#### **DolaLock**

Efficacy without severe neutropenia, neuropathy, or ocular toxicity



- Controlled bystander effect
- Selectively toxic to rapidly dividing cells
- Not a PgP substrate
- Induces immunogenic cell death

#### **Dolaflexin**

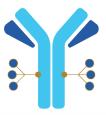
Improved therapeutic index vs. other platforms



- DolaLock payload
- Polymer scaffold
- DAR ~10-12
- Excellent drug like properties

#### Dolasynthen

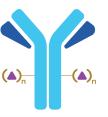
Homogenous & Customizable Platform



- DolaLock payload
- Synthetic scaffold
- Site-specific
- Precise DAR (2-24)

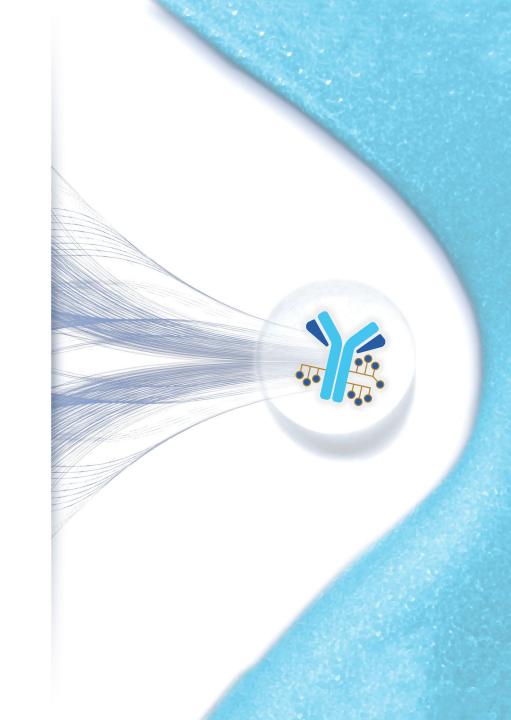
#### **Immunosynthen**

Systemic administration with targeted immunostimulatory effect



- Novel STING agonist
- Complete regression with one dose in multiple models
- Limited effect on systemic cytokines

XMT-1536: First-in-Class Dolaflexin ADC Targeting NaPi2b

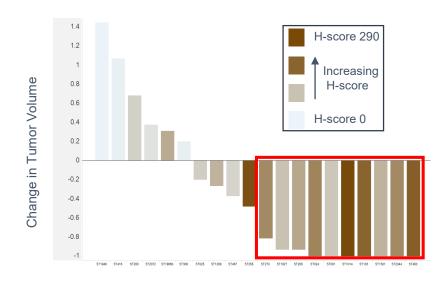


### Leader in Targeting NaPi2b, an Ideal and Validated ADC Target



- Broadly expressed in ovarian cancer and NSCLC adenocarcinoma
  - No detectable expression in squamous NSCLC
  - Limited expression in healthy tissues
- NaPi2b is a lineage marker (not an oncogene) that transports phosphate into the cell
  - The presence of EGFR and KRAS mutations in NSCLC adenocarcinoma correlates with more frequent expression of NaPi2b
- Proprietary biomarker assay can distinguish across low, medium, and high expression
  - Correlation between biomarker expression and response in preclinical and clinical settings
  - Developing companion diagnostic for use in registration enabling study

In Ovarian PDX Models, only tumors with an H-score above cutoff had a tumor response >50%



# XMT-1536: Advancing Through Proof-of-Concept Studies in Ovarian Cancer and NSCLC Adenocarcinoma



#### First-in-Class

- Clinically-validated target
- Fast-to-market strategy
- Wholly-owned<sup>1</sup>

## **Encouraging Clinical Activity**

- Confirmed responses and prolonged stable disease in heavily pretreated and biomarker unselected patients reported at ASCO 2019
- Expansion cohorts ongoing in ovarian cancer and NSCLC adenocarcinoma

#### **Well-Tolerated**

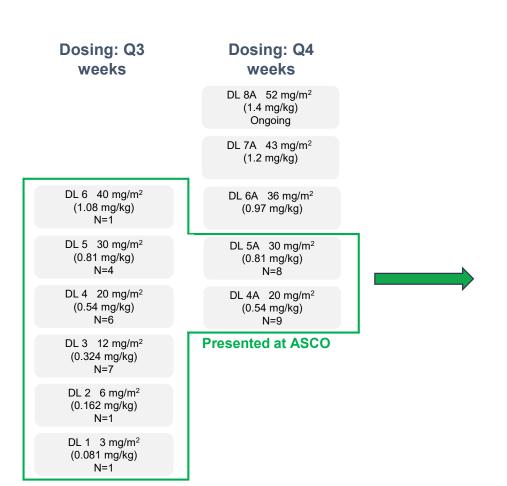
- MTD not yet reached
- Dose escalating to 52 mg/m²
- No severe toxicities commonly seen with other ADCs: neutropenia, ocular toxicities, or peripheral neuropathy
- Transient AST elevation without associated changes in bilirubin

#### **Multiple Data Read Outs Expected in 2020**

### XMT-1536 was Well-Tolerated with Most AE's Grade 1-2



#### Data Presented at ASCO with a Data Cutoff of May 10, 2019



#### Treatment Related Adverse Events in ≥10% of Patients

N = 37	N (%)					
Preferred Term	Grade 1	Grade 2	Grade 3	Total		
Nausea	12 (32)	2 (5)	0	14 (38)		
Fatigue	4 (11)	7 (19)	0	11 (30)		
Headache	5 (14)	5 (14)	0	10 (27)		
Aspartate aminotransferase (AST) increased	3 (8)	2 (5)	4 (11)	9 (24)		
Decreased appetite	1 (3)	6 (16)	0	7 (19)		
Blood alkaline phosphatase increased	6 (16)	0	0	6 (16)		
Vomiting	4 (11)	1 (3)	0	5 (14)		
Gamma-glutamyltransferase (GGT) increased	3 (8)	1 (3)	0	4 (11)		
Myalgia	3 (8)	0	1(3)	4 (11)		
Pyrexia	3 (8)	1 (3)	0	4 (11)		

#### Safety:

• No Grade 4 or 5 treatment-related adverse events (TRAEs)

No Severe Toxicities Associated with Other ADC Platforms such as Neutropenia, Ocular Toxicities, or Peripheral Neuropathy

### XMT-1536 Showed Activity in Heavily Pretreated Patients, Unselected for NaPi2b



#### Data Presented at ASCO with a Data Cutoff of May 10, 2019

#### Clinical Activity at Doses of 20mg/m<sup>2</sup> and Above\*

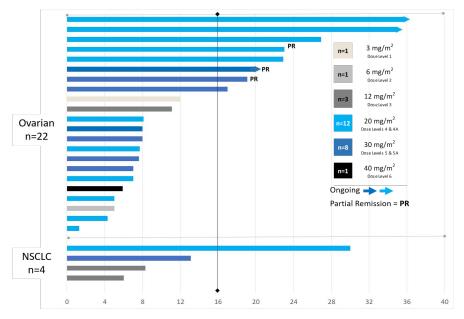
Outcomes in Ovarian Cancer (OC) & Non-small Cell Lung Cancer (NSCLC)	All OC	AII NSCLC	OC ≥20 mg/m²	NSCLC ≥20 mg/m²	OC ≥30 mg/m²
N	19	3	16	2	7
PR*	3 (16%)	0 (0%)	3 (19%)	0 (0%)	2 (28%)
SD*	8 (42%)	2 (67%)	6 (38%)	2 (100%)	3 (43%)
DCR (PR + SD)	11 (58%)	2 (67%)	9 (57%)	2 (100%)	5 (71%)
PD*	8 (42%)	1 (33%)	7 (43%)	0 (0%)	2 (28%)

Response evaluable

Based on objective responses and duration of treatment

\*As measured by RECIST, version 1.1

#### **Clinically Meaningful Treatment Duration > 16 weeks**



Treatment Duration (weeks)

Response evaluable

All Completed Dose Levels (OC and NSCLC Patients), N=26

# XMT-1536: Significant Progress Since ASCO 2019 in Maximizing Patient Benefit and Charting Path to Registration



#### **Higher Dose**

- Cleared 36 and 43 mg/m<sup>2</sup> with primarily Grade 1 and Grade 2 AEs with no severe neuropathy, neutropenia or ocular toxicities
- Increased expansion cohort dose to 43 mg/m<sup>2</sup>
- Currently evaluating 52 mg/m<sup>2</sup>

### Earlier, More Homogeneous Patient Population

 Initiated expansion cohorts in earlier line, more homogeneous populations

#### **Opportunity for Biomarker Selection**

- Correlating H-score and efficacy in anticipation of patient selection
- Collecting archival and fresh tissue to confirm concordance

**FAST-TO-MARKET REGISTRATION STRATEGY** 

### XMT-1536: Path to Pivotal Study in High Unmet Need Indications



## Dose Escalation Data in 1H 2020

#### **Population**

- Late stage platinum-resistant ovarian cancer
- Late stage recurrent NSCLC adenocarcinoma

## Ovarian Cancer Expansion Data in 2Q & 2H 2020

- 1-3 prior lines in platinum resistant
- 4 prior lines regardless of platinum status
- High grade serous histology

## NSCLC Adeno Expansion Data in 1H and 2H 2020

- Prior treatment with a platinum doublet and PD-1/L1 inhibitor
- Prior TKIs if targetable mutation
- Up to 2 prior lines of cytotoxic therapy
- Adenocarcinoma histology

#### Dose

• Evaluating 52 mg/m<sup>2</sup>

- 36 mg/m<sup>2</sup> dose initiated in Aug 2019
- 43 mg/m<sup>2</sup> dose initiated in Dec 2019
- 36 mg/m<sup>2</sup> dose initiated in Aug 2019
- 43 mg/m<sup>2</sup> dose initiated in Dec 2019

# **Current Standard of Care**

**Investigational Agent** 

ORR: 4-12% mPFS: 3-4 mos

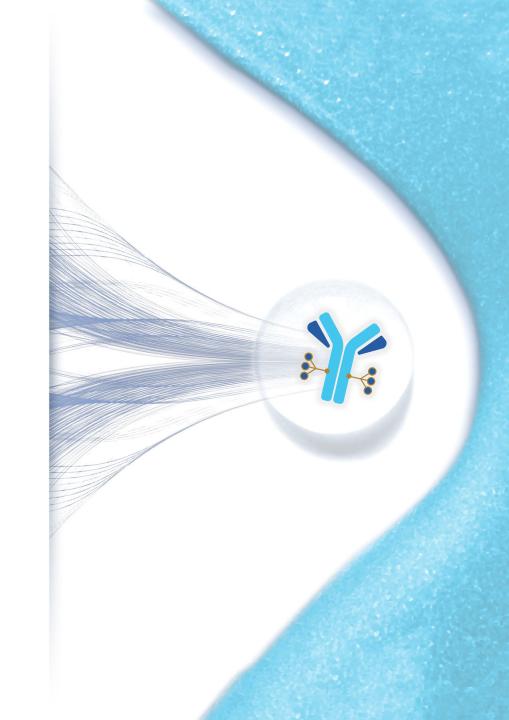
mOS: 9-12 mos

ORR: 14-23%

mPFS: 3-4 mos

mOS: 9-12 mos

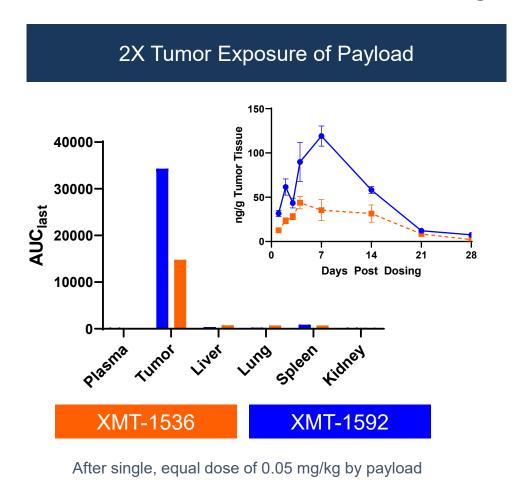
# XMT-1592 is a Dolasynthen ADC Targeting NaPi2b



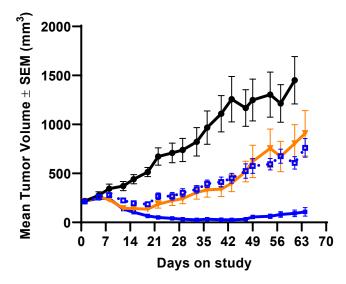
### XMT-1592 Shows Four-Fold Greater Efficacy in Lung Tumor Model



Our Success with NaPi2b Makes it an Ideal Target for Evaluation of the Clinical Differentiation of Dolasynthen



#### 4X Greater Activity in Lung PDX



- → Vehicle
- **XMT-1536** @ 0.1 mg/kg [2.1 mg/kg mAb]
- XMT-1592 @ 0.1 mg/kg [3.0 mg/kg mAb]
- XMT-1592 @ 0.025 mg/kg [0.75 mg/kg mAb]

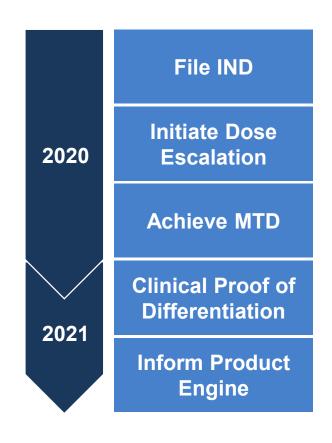
# Leveraging NaPi2b Experience for Rapid Dose Escalation of XMT-1592



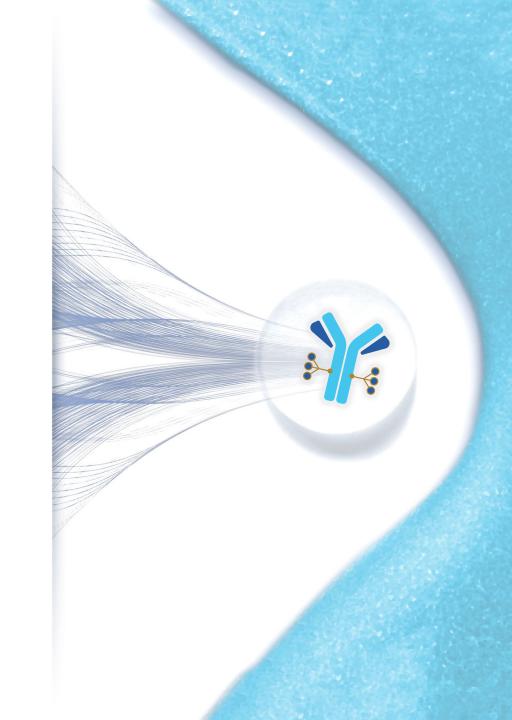
**Leveraging insights from XMT-1536** 

**Target Antibody Payload** Biomarker **Patient Populations Investigators** 

#### **Solidify NaPi2b Leadership**

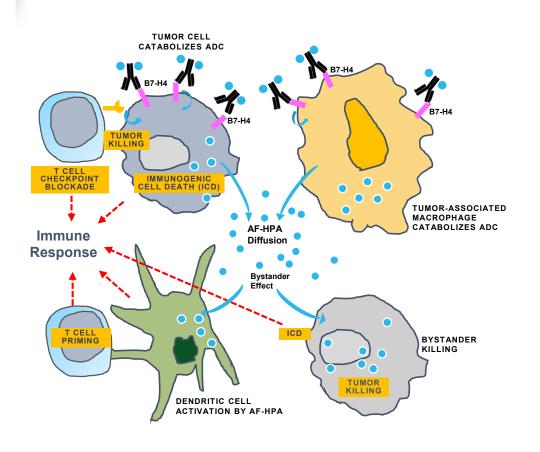


First-in-Class B7-H4 ADC Progressing into IND-Enabling Studies



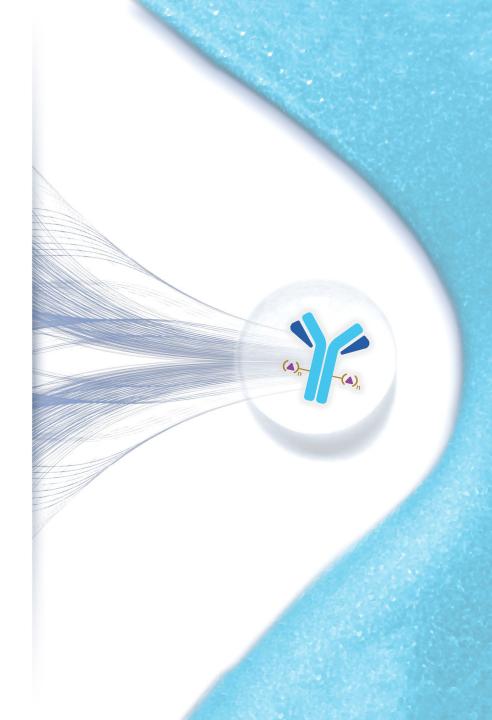
### **B7-H4 Expression Ideally Suited for a DolaLock ADC**





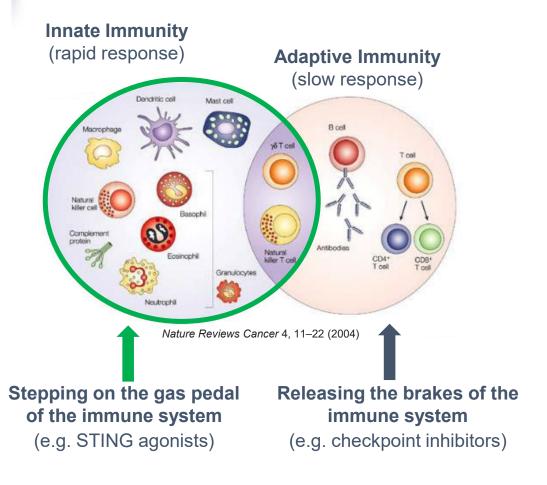
- B7-H4 is expressed on both tumor cells and immunosuppressive tumor-associated macrophages (TAMs)
  - Limited expression in normal tissues
- A DolaLock ADC targeting B7-H4 can exert its effect through multiple mechanisms of action:
  - Uptake by tumor cells and direct cytotoxicity
  - Uptake by TAMs to release payload in the tumor microenvironment
  - Free payload can activate dendritic cells and a secondary immune response
- Expression in PD-L1 negative tumors, provides a potential fast to market opportunities (e.g., triple negative breast cancer)

Immunosynthen Development Candidate in 2020



# Immunosynthen: Strong Rationale for a STING Agonist ADC Approach



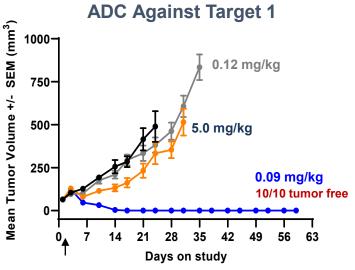


# ADCs are suited to overcome limitations of free agonist:

- Targeted delivery reduces toxicity liabilities
- Improved pharmacokinetics
- Accessibility to metastatic sites
- No restriction on tumor type, location or size

# Immunosynthen ADCs Show *In Vivo* Activity Against Multiple Targets and Immune Memory





21

**Days on Study** 

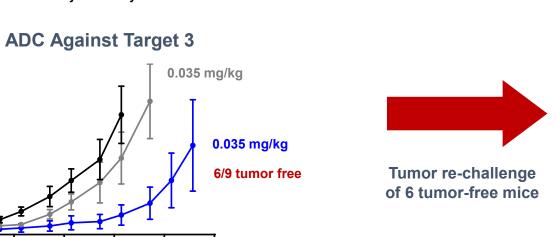
28

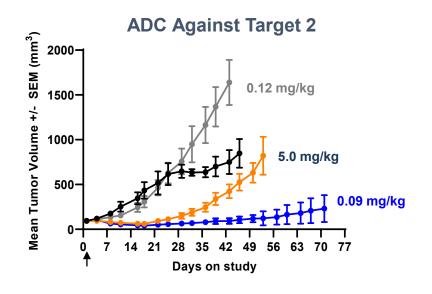
35

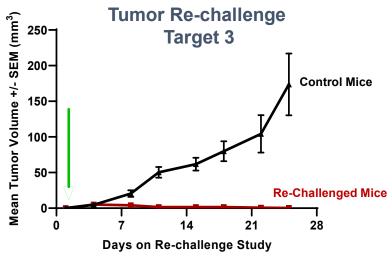
SEM (mm<sup>3</sup>) -008

Mean Tumor Volume +/-





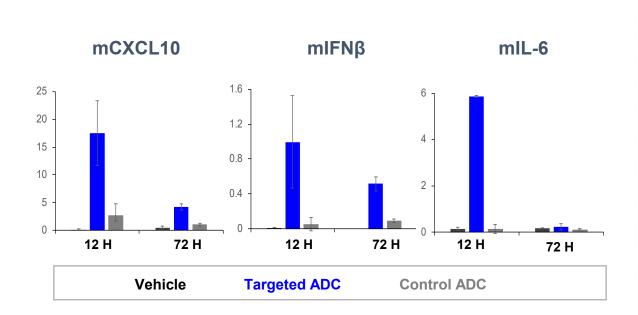




### Immunosynthen ADC Activates STING Pathway and **Induces Marked Immune Cell Infiltration in Tumors**



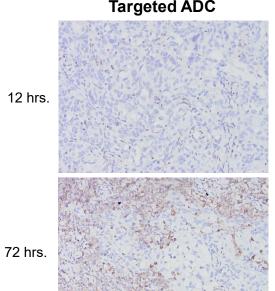
**Cytokine expression** (qPCR on FFPE samples)



Data shown for Immunosynthen ADC for Target 1 After single dose of 0.09 mg/kg by STING agonist payload

#### **CD45** Immunohistochemistry Immune cell infiltration

#### **Targeted ADC**

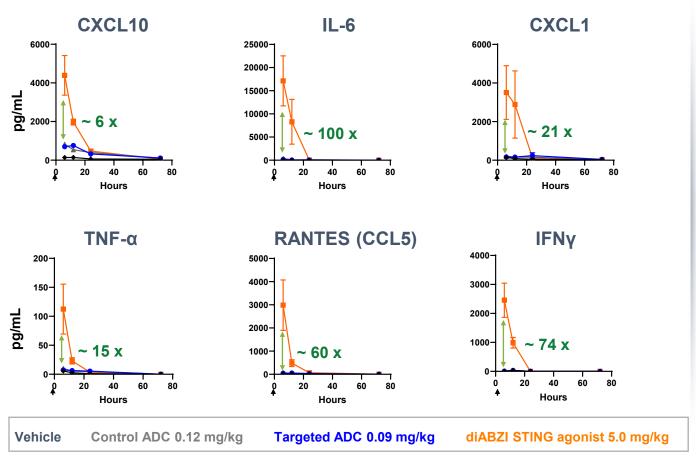


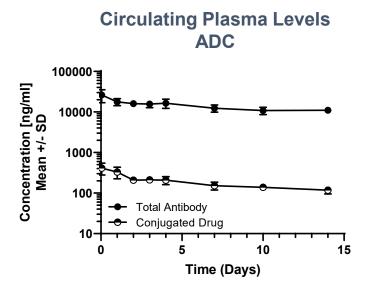
# Limited Induction of Serum Cytokines *In Vivo* by Immunosynthen ADC Despite Extended Plasma Exposure



#### **Analysis of Serum Cytokine Levels by Luminex Assay**

Data shown for Immunosynthen ADC for Target 1





Extended plasma exposure to Immunosynthen ADC does not result in extended exposure to systemic cytokines

# On Track to Select First Immunosynthen ADC Development Candidate in 2020





- ✓ Identified proprietary STING payload specifically designed for ADCs
- Demonstrated efficacy across multiple targets in a variety of models
- Confirmed tolerability in multidose exploratory NHP study
- Finalize proprietary STING ADC scaffold (linker, DAR, method and site of bioconjugation)
- Select first Immunosynthen ADC from current targets and leads



Expect to disclose data package in 2H 2020

# **2020: A Transformational Year for Mersana** with Multiple Data Readouts



### **2020 Goals & Anticipated Milestones**

XMT-1536	<ul> <li>Report dose escalation in 1H 2020</li> <li>Report interim data from OC and NSCLC expansion cohorts in 2Q 2020</li> <li>Report more mature data from expansion cohorts in 2H 2020</li> </ul>
XMT-1592	File IND and initiate clinical study in 1H 2020. Achieve rapid dose escalation
B7-H4	<ul> <li>Advance IND-enabling studies</li> <li>Disclose development candidate data package in 2H 2020</li> </ul>
Immunosynthen	<ul> <li>Select first development candidate</li> <li>Disclose development candidate data package in 2H 2020</li> </ul>
Product Engine	Continue to leverage proprietary platforms to expand pipeline
Corporate	Proactively evaluate potential for strategic collaborations that maximize value

### **Positioned to Create Value for Patients and Shareholders**



XMT-1536	<ul> <li>First-in-class NaPi2b ADC</li> <li>Completion of proof-of-concept studies in 2020</li> <li>Fast-to-market registration strategy</li> </ul>
XMT-1592	<ul> <li>Extends NaPi2b leadership</li> <li>Fast to clinical validation of preclinical differentiation</li> </ul>
Pipeline	<ul> <li>First-in-class B7-H4 and Immunosynthen ADCs</li> <li>Targeting high unmet medical needs</li> </ul>
Platforms	<ul> <li>Dolaflexin, Dolasynthen (DolaLock)</li> <li>Immunosynthen (Novel STING Agonist)</li> <li>Efficient product engines with multiple partnership opportunities</li> </ul>
Fundamentals	<ul> <li>Strong team</li> <li>Strong balance sheet</li> </ul>



