

Company Presentation

September 2020



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This presentation contains forward-looking statements within the meaning of the US Private Securities Litigation Reform Act of 1995. The forward-looking statements involve substantial risks and uncertainties. All statements, other than statements of historical facts, contained in this presentation, including statements regarding our strategy, future operations, future financial position, future revenue, projected costs, prospects, plans and objectives of management and expected market growth are forward-looking statements. The words "anticipate," "believe," "continue," "could," "estimate," "expect," "intend," "may," "might," "plan," "potential," "predict," "project," "should," "target," "would" and similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words. Such statements include, but are not limited to, statements about regulatory approvals, clinical trial timing and plans, the achievement of clinical and commercial milestones, future financial and operating results. business strategies, market opportunities, financing, and other statements that are not historical facts. Our product candidates and related technologies are novel approaches to cancer treatment that present significant challenges. Actual results may differ materially from those indicated by such forward-looking statements as a result of various factors, including but not limited to: risks associated with the Company's development work, including any delays or changes to the timing, cost and success of our product development activities and clinical trials including if we encounter difficulties enrolling patients in our clinical trials; the risks of delays in FDA and/or EU approval of our drug candidates or failure to receive approval; the risks related to commercializing any approved new pharmaceutical product including the rate and degree of market acceptance of our product candidates; development of our sales and marketing capabilities and risks associated with failure to obtain sufficient reimbursement for our products; our inability to enter into collaboration or alliances with partners; risks associated with protection of our intellectual property rights; and other risks and uncertainties affecting the Company including those described in the "Risk Factors" section included in documents the Company files from time to time with the Securities and Exchange Commission. Any forward-looking statements contained in this presentation speak only as of the date hereof, and the Company undertakes no obligation to update any forward-looking statement, whether as a result of new information, future events or otherwise.

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# **MISSION**



# Investment Highlights

FDA acceptance of BLA¹ for Danyelza™ (naxitamab) for NB. PDUFA date in Nov 2020. Submission for omburtamab completed Aug 2020. Both products have BTD²

Potential to expand into other indications and other lines of therapy, multiple studies ongoing. Expansion includes <sup>177</sup>Lu-omburtamab-DTPA, opening Phase 1/2 in Q4 2020

First BsAb product candidate, nivatrotamab, using the Y-BiClone technology in Phase 1/2. Second BsAb product candidate CD33xCD3

Novel SADA technology platform, a concept also referred to as Liquid Radiation™

GD2-GD3 Vaccine - ongoing Phase 2 Study in high-risk NB patients in remission

Financial strength – secured financing through the end of 2022





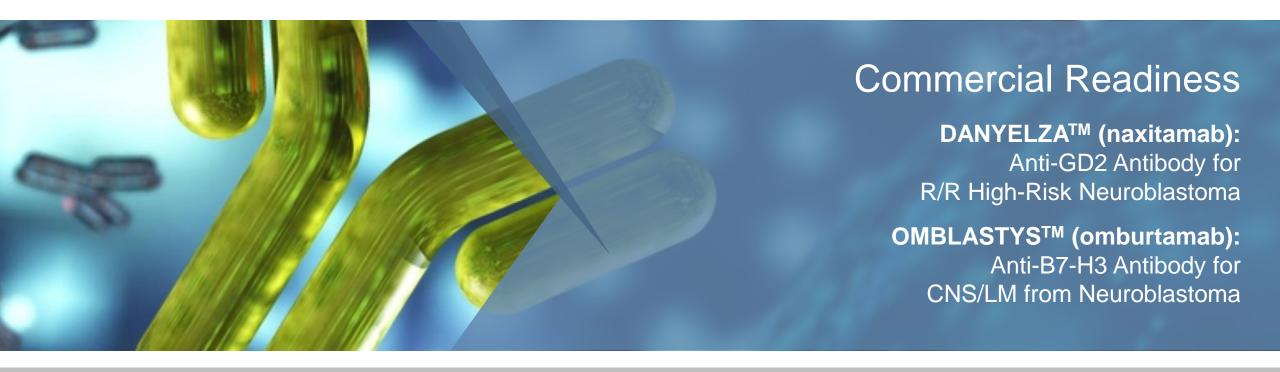
# Strong Pipeline

| Programs R P D D              | rreciinicai                         | Phase 1 | Phase 2/Pivotal Study | Next Anticipated<br>Milestones                    |
|-------------------------------|-------------------------------------|---------|-----------------------|---|
| Lead Development Candidates 🗸 | Naxitamab (GD2)                     |         |                       | PDUFA date in Nov 2020. Priority review           |
| <b>✓</b>                      | <sup>131</sup> I-omburtamab (B7-H3) |         |                       | Rolling BLA submission completed in Aug 2020      |
| Vaccine                       | GD2-GD3 Vaccine                     |         |                       | Multicenter Phase 2 study to open Q1 2021         |
| Y-BiClone                     | Nivatrotamab                        |         |                       | Small Cell Lung Cancer - IND filing in Q4<br>2020 |
| Bispecific Platform           | CD33xCD3                            |         |                       | AML Pediatric Cancer IND being prepared           |
| Early Stage RIT               | 177Lu-omburtamab-<br>DTPA           |         |                       | Medulloblastoma study to open in Q4 2020          |
| SADA Technology               | GD2-SADA                            |         |                       | GD2 Positive solid tumors, IND 2021               |
|                               | GPA33-SADA                          |         |                       | Colon Therapeutic/Diagnostic, IND 2022            |
|                               | HER2-SADA                           |         |                       | Breast Cancer, IND 2022                           |
|                               | B7-H3-SADA                          |         | •                     | Prostate Cancer, IND 2022                         |

<sup>&</sup>lt;sup>1</sup>Indicates eligibility for a Priority Review Voucher (PRV) on approval







# Development Programs Approaching Registration and Commercialization

| Compound            | Indication  | Total Incidence per Year<br>(US) | Addressable Patient<br>Population per Year (US) |
|---------------------|---|----------------------------------|---|
|                     | Neuroblastoma – 2 <sup>nd</sup> Line                                    | 300                              | 300   |
| GD2<br>naxitamab    | Neuroblastoma – Front Line  | 800                              | 450   |
|                     | Osteosarcoma – 2 <sup>nd</sup> Line                                     | 450                              | 200   |
|                     | Neuroblastoma Metastatic to the Central Nervous System (CNS/LM from NB) | 80                               | 80  |
| B7-H3<br>omburtamab | Diffuse Intrinsic Pontine Glioma (DIPG)                                 | 300                              | 300   |
|                     | Desmoplastic Small Round Cell Tumors (DSRCT)                            | 100                              | 100   |



# In preparation for launch, commercial activities focused on three key areas:

Build best in class, right-sized commercial organization

- Small universe of pediatric cancer centers treat majority of neuroblastoma
- Lean and efficient commercial organization to align with targeted launch



Launch planning and execution focused on driving:

- Rapid uptake at launch
- Optimal pricing and reimbursement coverage
- Supportive stakeholder experience along neuroblastoma treatment journey



Building awareness of Y-mAbs

- Outreach & engagement with KOLs and top pediatric cancer centers
- Engagement with key neuroblastoma advocacy groups
- Medical congress presence to raise profile of Y-mAbs



# Naxitamab - Primary and Secondary Refractory Patients

Investigator evaluated responses

## **Study 12-230 (SIOP October 2019)**

- 23 evaluable patients with primary refractory high-risk
   NB: 78% ORR
- 50% two-year progression free survival (PFS) was observed
- Study population of 35 secondary refractory patients with relapsed NB resistant to salvage therapy: 37% ORR
- 36% two-year PFS was observed

### Study 201 (investigator assessed dataset for BLA)

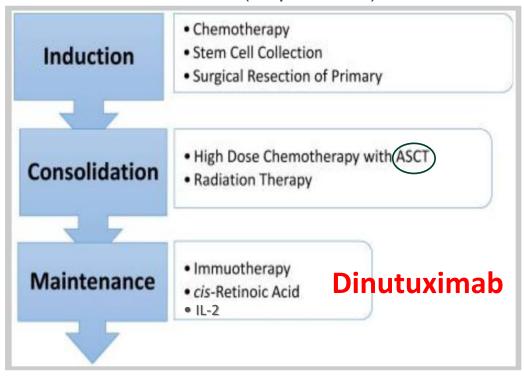
- 24 patients included in BLA filing: 79% ORR and 71% CR
- In 13 of 14 patients with bone marrow disease, bone marrow was cleared after treatment



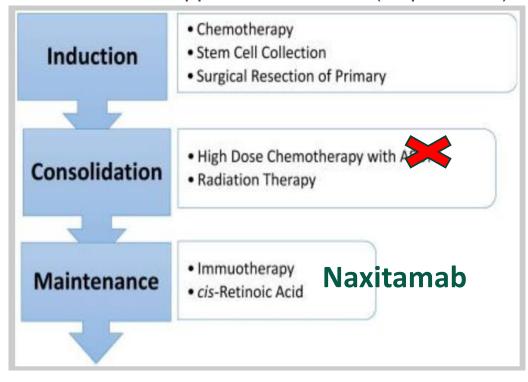
# High-Risk Neuroblastoma Treatment Recommendation

COG and MSK/Y-mAbs

COG – 8-20 h infusion (x4 per week)

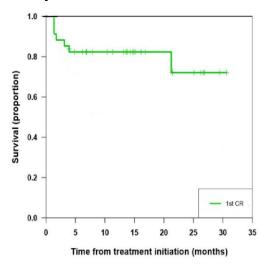


MSK/Y-mAbs – app 30 min infusion (x3 per week)



# Naxitamab: Frontline NB data without standard ASCT

## 2-year Event Free Survival:

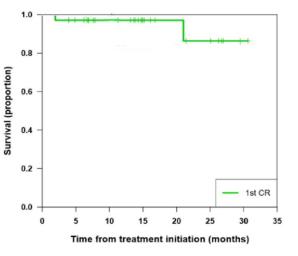


72.1% (95% CI = (53.1%, 97.7%))

VS.

dinutuximab 63%

## 2-year Overall Survival:



86.3% (95% CI = (68.0%,100.0%))

VS.

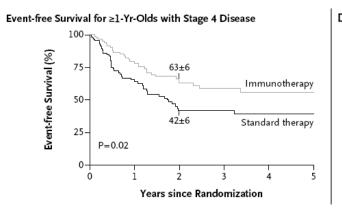
dinutuximab 84%

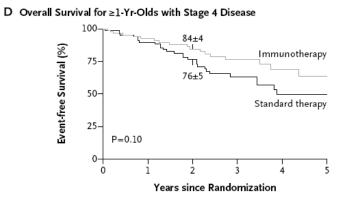
Data from Dr. J. Mora, Y-mAbs R&D Day Dec 11, 2019

The NEW ENGLAND JOURNAL of MEDICINE

#### ORIGINAL ARTICLE

Anti-GD2 Antibody with GM-CSF, Interleukin-2, and Isotretinoin for Neuroblastoma





# Naxitamab: Key Takeaways

Addresses Significant Unmet Needs in R/R High-Risk NB • Potential to Expand to Broader Populations

Studies 12-230 and 201 formed primary basis of BLA, which was completed in March 2020. PDUFA date in November 2020.

US commercialization in high-risk NB being planned for 2020 – Frontline studies ongoing

Granted ODD, BTD, and RPDD1

ct

Multiple potential advantages over other GD2 targeting antibody-based therapies: Modest toxicity, shorter infusion time, ability to be administered in outpatient setting

<sup>1</sup>Indicates eligibility for a Priority Review Voucher (PRV) on approval



# Omburtamab: Regulatory Path to BLA Approval

## Regulatory

Studies 03-133 and 101 formed basis of BLA submission:

OS data accepted by FDA for accelerated approval PK and dosimetry comparison required

Data from Study 101 multicenter supports BLA submission

Qualifies for accelerated approval

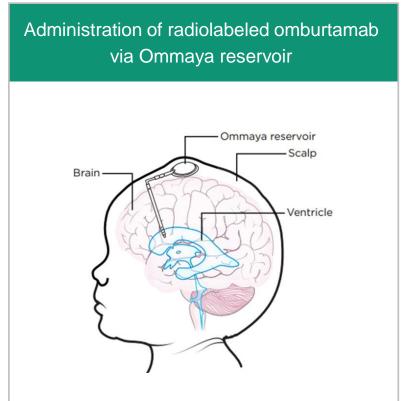
BLA submission completed in August 2020

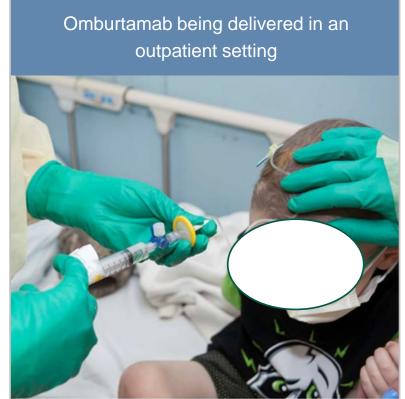
ODD, BTD, and RPDD

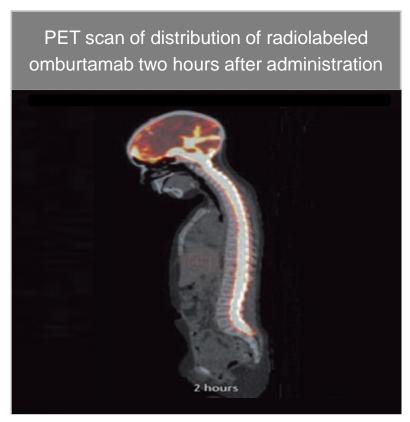


# Omburtamab: Delivered in an Outpatient Setting – 2 Doses per Patient

CNS/LM from NB patients





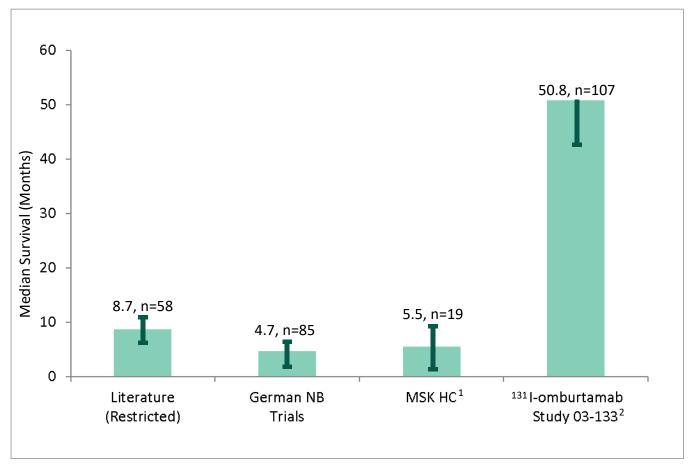


After induction treatment including all or some of the three treatments (chemotherapy, surgery, and radiation) patients will receive radiolabeled omburtamab



## Omburtamab: Clinical Overview

Study 03-133: <sup>131</sup>I-omburtamab Improves Survival in CNS/LM from NB Patients



These results demonstrate the opportunity for <sup>131</sup>I-omburtamab to address the lack of an established, effective therapy for patients with CNS/LM from NB

<sup>&</sup>lt;sup>2</sup> <sup>131</sup>I-omburtamab = Patients with CNS/LM treated under Study 03-133



<sup>&</sup>lt;sup>1</sup> MSK HC = neuroblastoma patients with CNS/LM treated at MSK prior to 2003

# Omburtamab: Key Takeaways

Addresses Significant Unmet Needs and has the Potential to Expand its Application to Broader Populations

- No approved products for patients with R/R NB who have CNS/LM from NB
- Goal of treatment is generally palliative

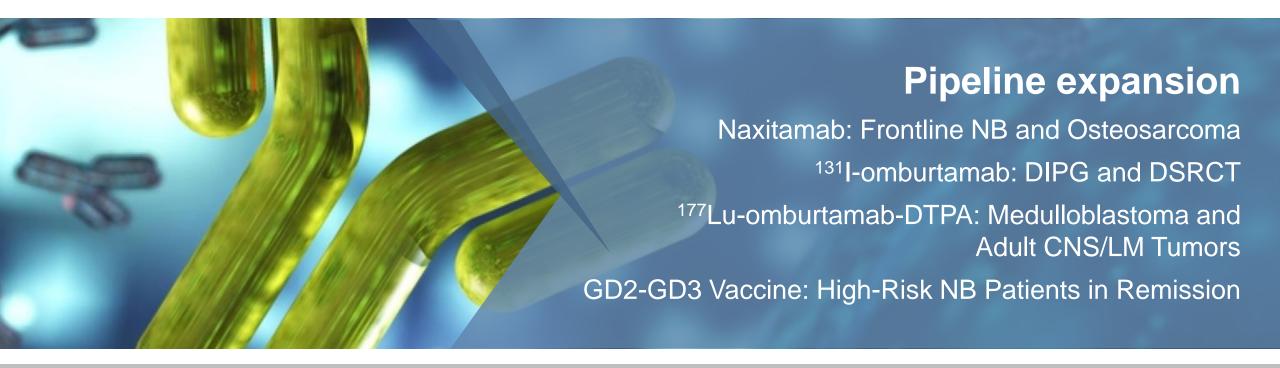


Historical median OS of ~six months and no expected five-year survival

- Granted ODD, BTD, and RPDD; May qualify for a sBLA for DIPG and DSRCT assuming positive pivotal data
- Studies 03-133 and 101 form primary basis for rolling BLA submission for CNS/LM from NB – Completed in August 2020
- Large potential market opportunity for the treatment of LM from tumors expressing B7-H3

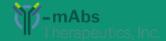






# Naxitamab: Targets GD2 with Expanding Clinical Program

| Naxitamab (GD2)   | Phase 1   | Phase 2/Pivotal Study | Highlights  |
|---|---|-----------------------|---|
| Accelerated Pathway   | Phase 2: Primary R/R High-Risk NB (Pediatric) – Study 201       |                       | BLA submission completed in March 2020. PDUFA date in November 2020. Priority review. |
| 7.000.01a.tod 1 atriway                                     | Phase 2: Primary R/R NB (Pediatric) – Study 12-230              |                       | Single-center study – part of BLA pivotal data package                                |
| Phase 2: Frontline High-Risk NB (Pediatric) – Study 16-1643 |   | Study 16-1643         | Ongoing Phase 2 study   |
| Expanding to Frontline                                      | Phase 2: Frontline naxitamab – Study 202                        |                       | Frontline Phase 2 study to initiate in 2020   |
|   | Phase 2: Chemoimmunotherapy for R/R High-Risk NB – Study 17-251 |                       | Heavily pre-treated, high-risk NB patients  |
| Label Expansion   | Phase 2: Combo naxitamab plus chemo – Stu                       | dy 203                | Combo Phase 2 study to initiate in 2020   |
|   | Phase 2: Relapsed Second-line Osteosarcoma                      | a – Study 15-096      | If successful, may form part of support for future sBLA in Osteosarcoma               |



# Omburtamab: Broad Clinical Platform

| Omburtamab<br>(B7-H3) | Phase 1                                 | Phase 2/Pivotal Study | Highlights  |
|-----------------------|---|-----------------------|---|
| Accelerated           | Phase 2: CNS/LM from NB (Pediatric) – S | tudy 101              | Multi-center PK study; Rolling BLA submission completed in Aug 2020 |
| Pathway               | Phase 1: CNS/LM – Study 03-133          |                       | MSK single-center efficacy data                                     |
|                       | Phase 2: DIPG multi-center - Study 102  |                       | Multi-center study to initiate in 2020                              |
| Label Expansion       | Phase 1: DIPG – Study 11-011            |                       | Study update presented at ASCO 2019                                 |
|                       | Phase 2: DSRCT – Study 19-182           |                       | Study update from Phase 1 presented at CTOS Nov 2019                |
|                       |   |                       |   |



# <sup>177</sup>Lu-omburtamab-DTPA - Pediatric and Adult Strategy

| <sup>177</sup> Lu-omburtamab-DTPA | Phase 1                              | Phase 2/Pivotal Study | Highlights                     |
|-----------------------------------|--------------------------------------|-----------------------|--------------------------------|
| Pediatric                         | Phase 1: Medulloblastoma - Study 301 |                       | Phase 1 to initiate in Q4 2020 |
| Adult                             | Phase 1: B7-H3 Positive - Study 302  |                       | Phase 1 to initiate in Q4 2020 |

#### Adult

- First indication: Basket study of B7-H3 positive CNS/LM tumors
- Prior experience from compartmental treatment of adult patients with <sup>131</sup>I-omburtamab

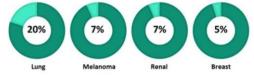
#### **Pediatric**

- First indication: **Medulloblastoma.** IND submitted Dec 2019.
- Prior experience from compartmental treatment with <sup>131</sup>I-omburtamab – 27 pediatric patients treated

Clinical Testing (Adult)

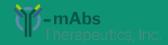
- Experience using <sup>131</sup>I-omburtamab in 68 patients with tumors such as sarcoma, melanoma and medulloblastoma
- cGMP production established





Incidence of Brain Metastases - Top 4 Tumors

| SURVIVAL              |            |  |  |  |
|-----------------------|------------|--|--|--|
| Primary Tumor 5 Years | 32.9%      |  |  |  |
| Metastatic            | 3-6 months |  |  |  |



## GD2-GD3 Vaccine: A Naxitamab Add-On in Phase 2 at MSK

| GD2-GD3 Vaccine | Phase 1                                    | Phase 2/Pivotal Study | Highlights                               |
|-----------------|--|-----------------------|--|
| MSK             | Phase 2: NB - Second and later remission - | - Study 05-075        | Phase 2 ongoing at MSK                   |
| Multicenter     | Phase 2: NB - Second remission - Study 60  | 1                     | <br>Phase 2 multicenter to start Q1 2021 |



More than 230 patients on study drug – ODD granted – RPDD granted in 2019



84 high-risk NB patients received the GD2-GD3 Vaccine, all of whom were in second or later remission



PFS of approximately 51% and OS of approximately 90% at two years



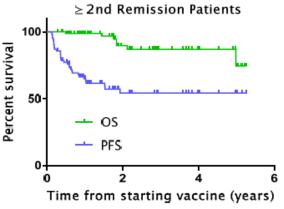
Study now also enrolling patients in first remission



The GD2-GD3 Vaccine appears to be well tolerated, with no reported grade 3 or grade 4 toxicities

# Initial Focus on 2<sup>nd</sup> and Later Remission Group:

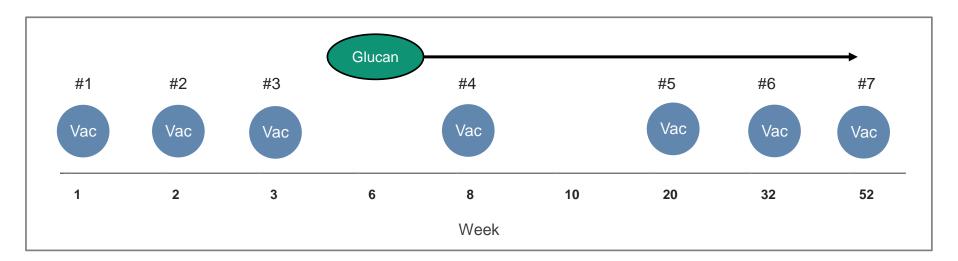
Y-mAbs multi-center Study 601 in NB patient 2<sup>nd</sup> CR

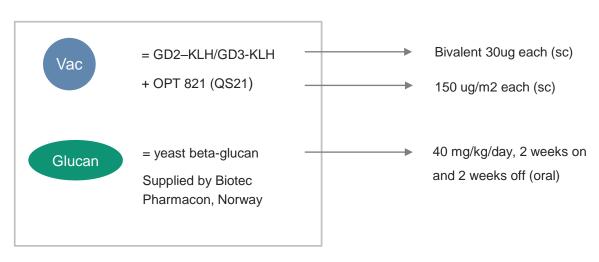


# Phase 2 Vaccine Study at MSK

Clinicaltrials.gov NCT00911560

7 cycles



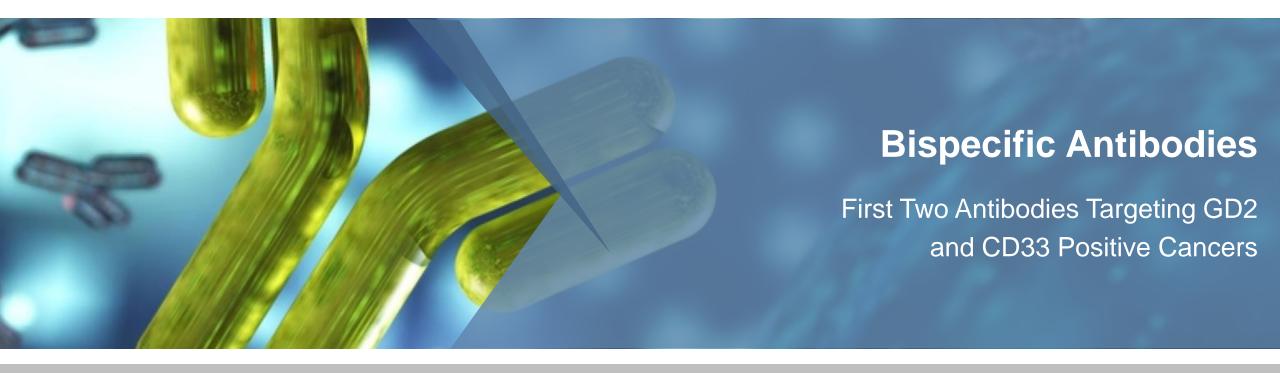


| Seroconversion = antibody response |       |       |  |  |
|------------------------------------|-------|-------|--|--|
| % patients with positive           |       |       |  |  |
| Anti-GD2 titer Anti-GD3 titer      |       |       |  |  |
| Pre-vaccine                        | 13.3% | 29.4% |  |  |
| During vaccine/follow-up           | 82.7% | 70.4% |  |  |

I. Cheung et al., Phase II Trial of GD2-KLH/GD3-KLH Vaccine for Stage 4 Neuroblastoma in 2<sup>nd</sup> or later Remission ANR, San Francisco, May 2018

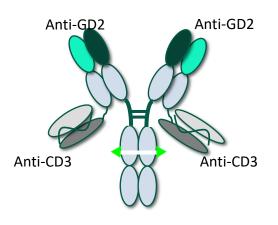






# Nivatrotamab – Planning for three Phase 2 studies

| GD2-BsAb | Phase 1                                      | Phase 2/Pivotal Study | Highlights   |
|----------|--|-----------------------|--|
|          | Phase 1: Basket trial - Study 18-034         |                       | Ongoing since Q1 2019, Cohort 6 recruiting           |
|          | Phase 2: SCLC - Study 402                    |                       | Study planned for Q4 2020                            |
|          | Phase 2: Third line NB – Based on Study 18-0 | 034                   | Expansion of MSK study into Phase 2 study in Q1 2021 |
|          | Phase 2: Refractory Osteosarcoma – Based o   | n Study 18-034        | Expansion of MSK study into Phase 2 study in Q1 2021 |



- Expanding into three Phase 2 studies
- Expanding into adult indications
- Multicenter studies underway





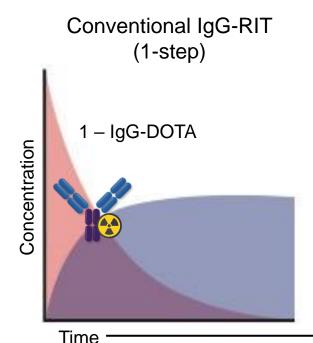
# Conventional RadioImmunoTherapy (RIT)

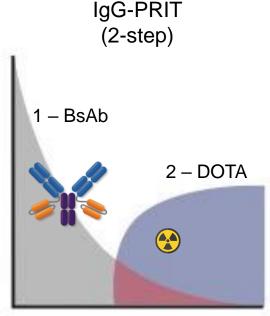
Limited by high levels of unwanted radiation exposure to non-target tissues, like the blood

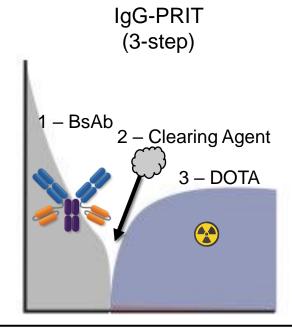
## Blood exposure (payload)

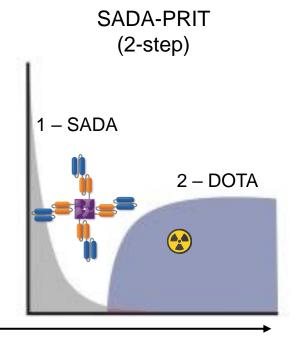
Tumor uptake (payload)

Antibody clearance









Low Tumor:Blood Ratio High toxicity

Mid to Hi Tumor:Blood Ratio Some toxicity

High Tumor:Blood Ratio Low toxicity Needs clearing agent

High Tumor:Blood Ratio
Low toxicity
No clearing agent

# Proof of principle for GD2-SADA

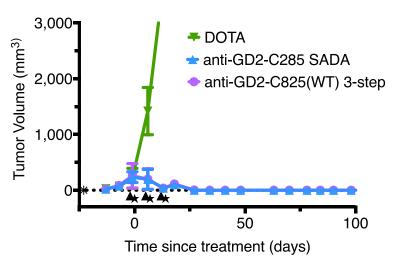
2-step GD2-SADA can successfully treat GD2(+) small cell lung cancer - ablating large, established tumors thereby potentially opening application beyond neuroblastoma for the GD2 construct

### **Major observations:**

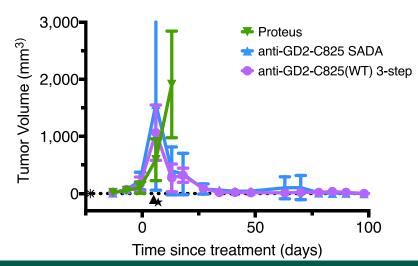
### SCLC is highly sensitive to pre-targeted radioimmunotherapy

- a) First time seeing complete responses in highly aggressive LX22 PDX tumor model
- b) 1 dose of 1uCi/mouse of <sup>225</sup>Ac was sufficient to completely ablate very large established subcutaneous tumors (5/5 CR)
- c) 3 doses of 1mCi/mouse of <sup>177</sup>Lu provided very comparable efficacy as well (5/5 CR)
- d) 2-step GD2-SADA showed equivalent efficacy to conventional 3-step approach, without needing any clearing agent

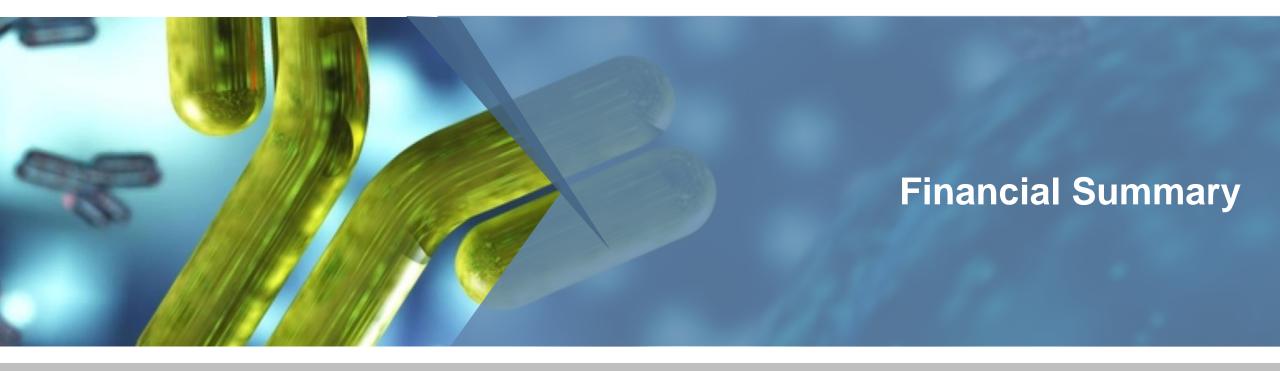
### 177Lu-DOTA Treatment of LX22 (SCLC PDX)



225Ac-Proteus Treatment of LX22 (SCLC PDX)

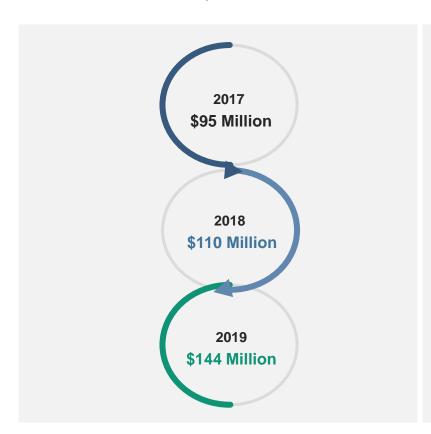






# Strong Financial Position with Blue Chip Investors

Y-mAbs Has Completed a Series of Successful Financing Rounds, with \$374 Million Raised to Date





IPO: September 2018 \$110 Million

Follow on: November 2019 \$144 Million



3 RPDDs
Received for leading compounds

\$158.1 Million

of cash and cash equivalents as of June 30, 2020



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