



# TAKEDA ONCOLOGY

WE ASPIRE TO CURE CANCER

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Head, Oncology Therapeutic Area

## ORIENTATION TO OUR ONCOLOGY R&D OVERVIEW

### Focused Oncology R&D Strategy

- Building on foundational expertise in hematologic malignancies and a growing portfolio in lung cancer

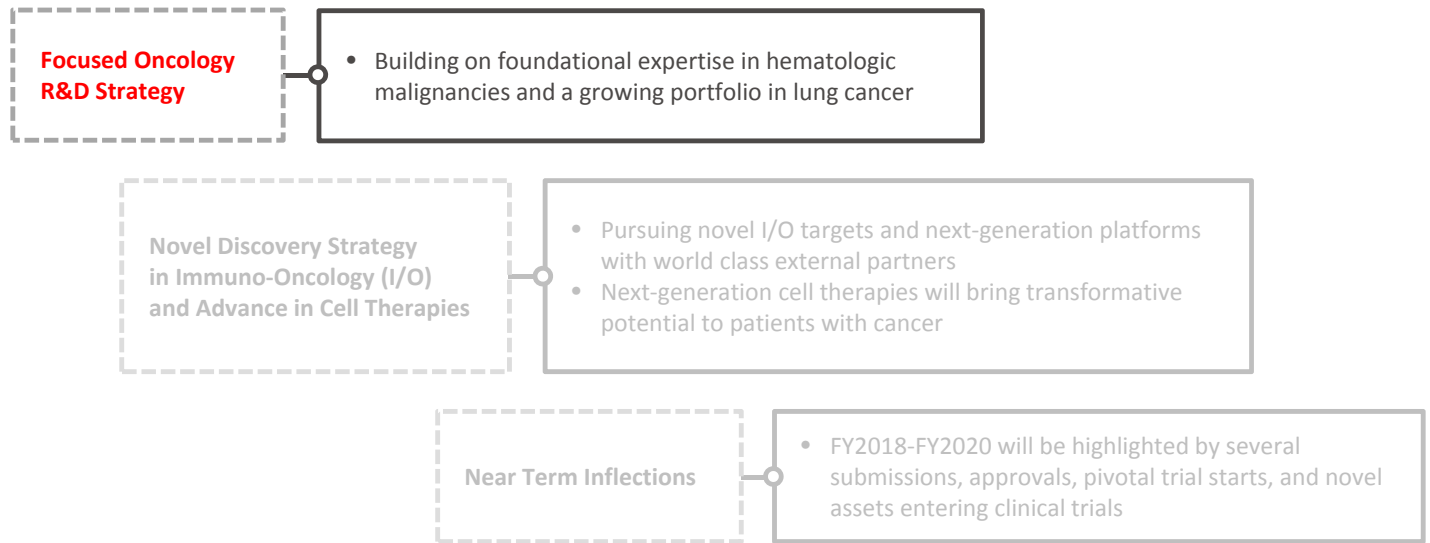
### Novel Discovery Strategy in Immuno-Oncology (I/O) and Advance in Cell Therapies

- Pursuing novel I/O targets and next-generation platforms with world class external partners
- Next-generation cell therapies will bring transformative potential to patients with cancer

### Near Term Inflections

- FY2018-FY2020 will be highlighted by several submissions, approvals, pivotal trial starts, and novel assets entering clinical trials

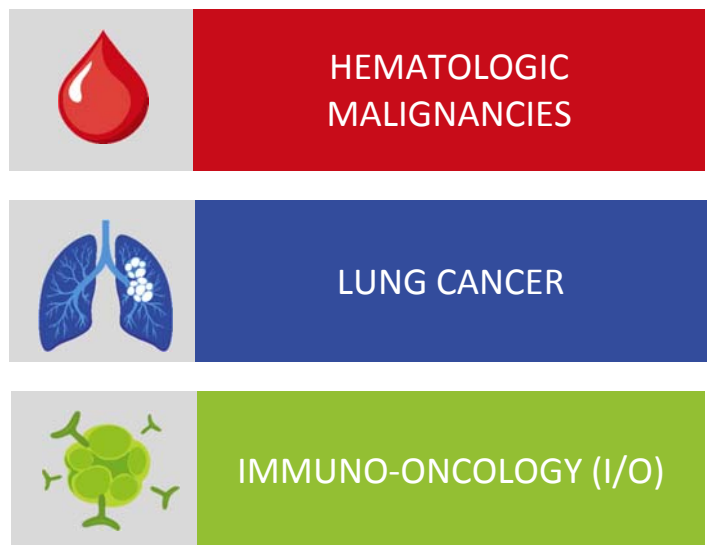
## ORIENTATION TO OUR ONCOLOGY R&D OVERVIEW



## WE ASPIRE TO CURE CANCER

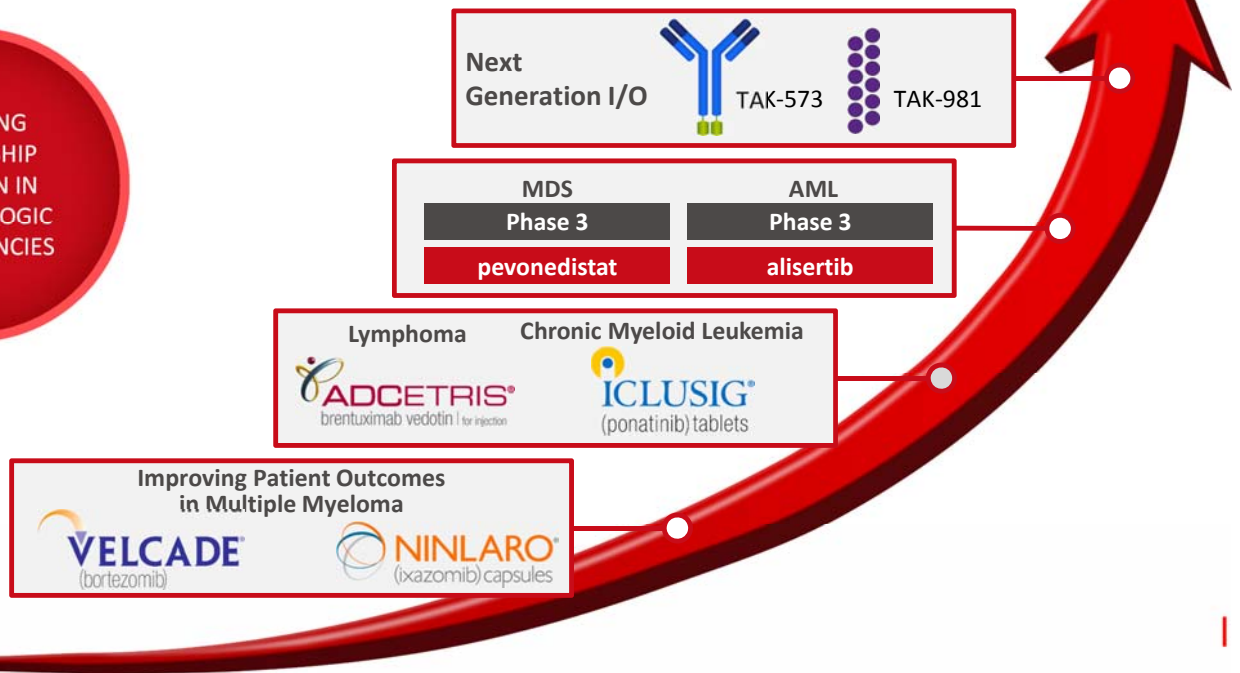
### OUR MISSION

We endeavor to deliver novel medicines to patients with cancer worldwide through our commitment to science, breakthrough innovation, and passion for improving the lives of patients.



# BUILDING ON THE TAKEDA ONCOLOGY FOUNDATION IN HEMATOLOGIC MALIGNANCIES

GROWING LEADERSHIP POSITION IN HEMATOLOGIC MALIGNANCIES



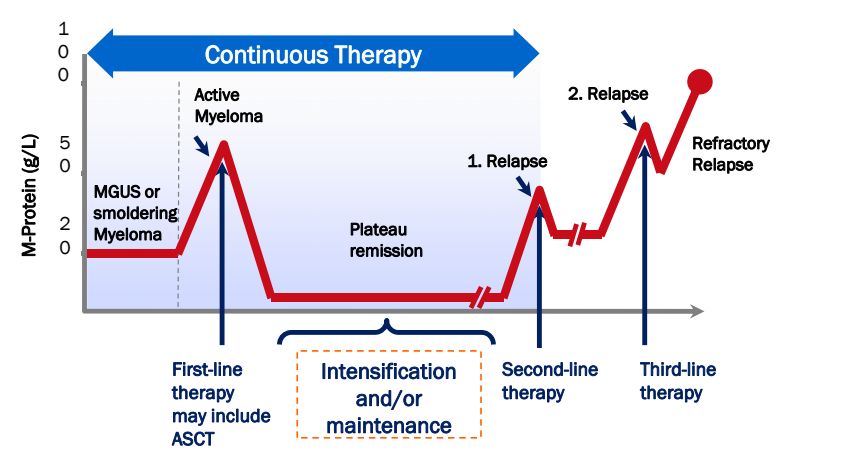
## RECENT PROGRESS AND NEXT STEPS



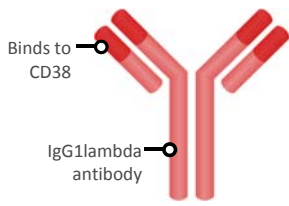
**Current Status**  
 Approved in 59 countries for Relapsed/Refractory Multiple Myeloma  
 First Phase 3 maintenance readout (post-transplant)

**Looking Forward**  
 2019 Data Inflections:  
 MM2 (newly diagnosed)  
 MM4 (non-transplant maintenance)  
 AL1 (amyloidosis)  
 Evolution of real world evidence

**Ideal Maintenance Therapies in Multiple Myeloma:**  
 ✓ Easy to administer    ✓ Minimal toxicity    ✓ Maintain response

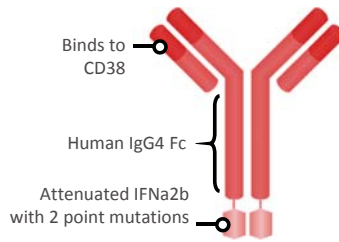


# ADVANCE CD38 BIOLOGY FOR REFRACTORY MULTIPLE MYELOMA



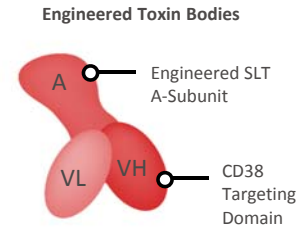
**TAK-079**

- A fully human, anti-CD38 cytolytic IgG1lambda antibody
- Potent and selective reduction of plasmablasts and NK cells
- Potential for convenient subcutaneous delivery
- Currently in Phase 1 for refractory multiple myeloma



**TAK-573**

- Novel immuno-cytokine approach
- Potential to overcome toxicity of unmodified interferon  $\alpha$  and realize the true benefit in oncology
- Compelling pre-clinical data; Phase 1 enrolling for patients with refractory multiple myeloma



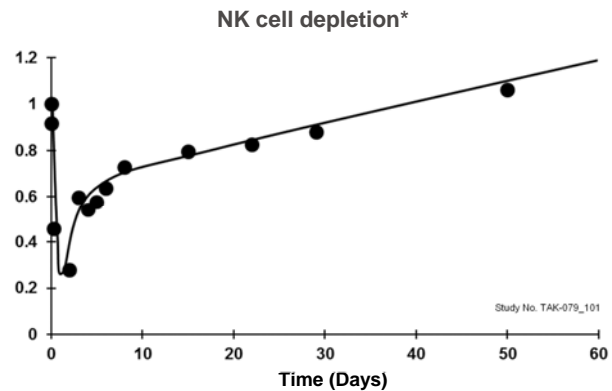
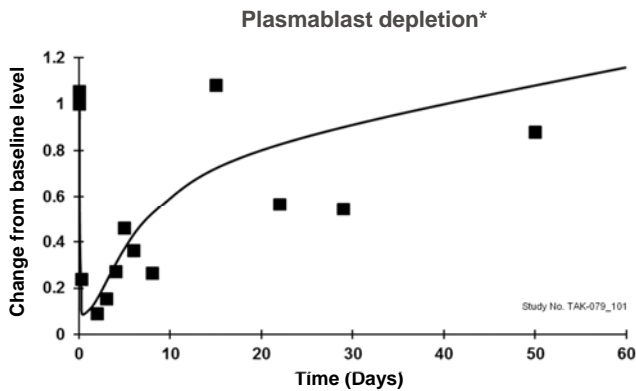
**TAK-169**

- 2<sup>nd</sup> generation Molecular Templates platform
- pM activity against CD38+ cells plus activity in daratumumab-resistant cells
- IND planned in 2019

## TAK-079: IMPROVING UPON FIRST GENERATION ANTI-CD38 mAb FOR REFRACTORY MULTIPLE MYELOMA PATIENTS



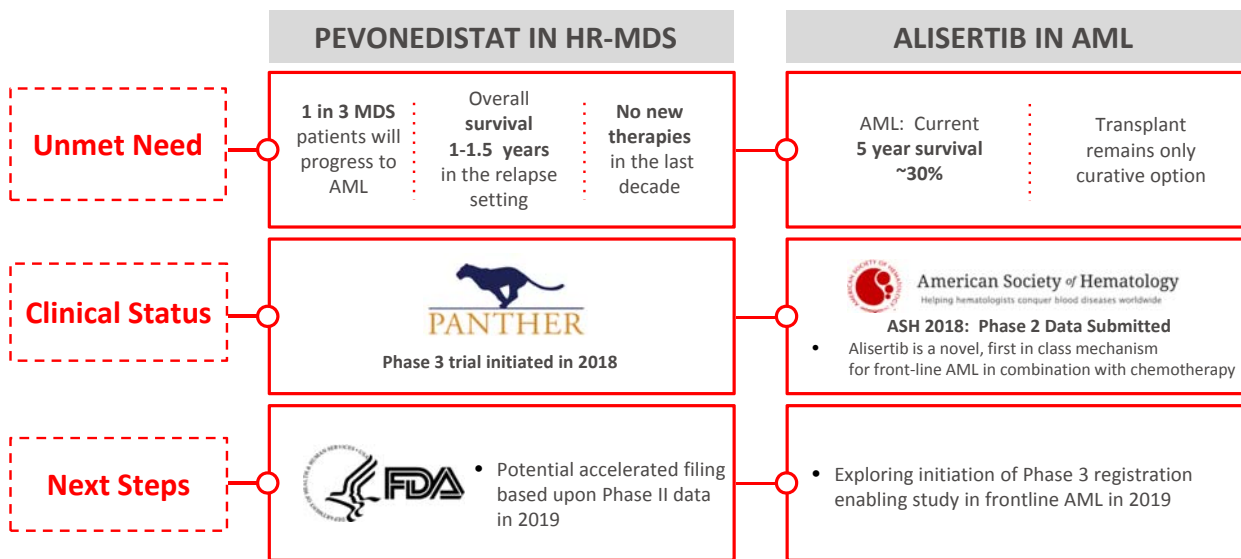
**A potent anti-CD38 mAb administered as a low volume subcutaneous (SC) injection**



\* After a single SC injection of 0.6 mg/kg into healthy volunteers (n=6)

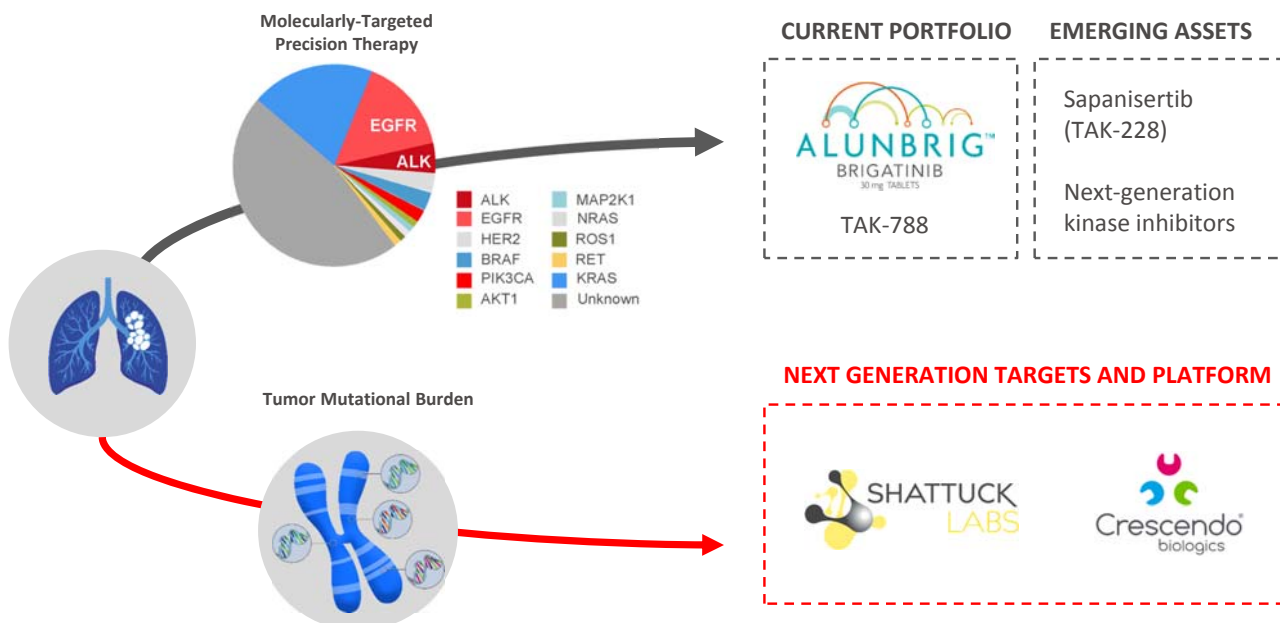
**Novel pharmacokinetic properties enhance potency and enable convenient administration**

# BRINGING NOVEL THERAPIES TO MDS AND AML

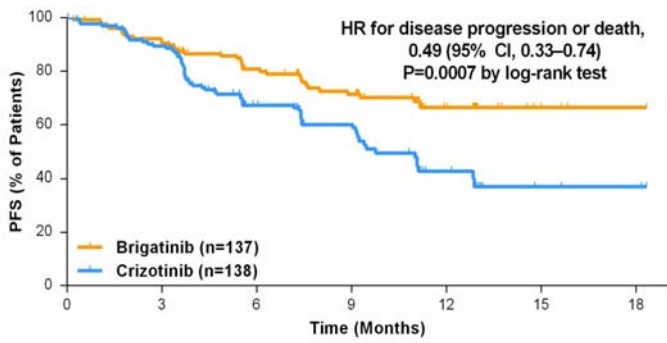


American Cancer Society – Survival Statistics for Myelodysplastic Syndromes, Tamamyan et al. *Critical Reviews in Oncology/Hematology* 2017, Yeung et al. *Biology of Blood and Marrow Transplantation* 2015, Courville et al. *BMC Clinical Pathology* 2017.

# DUAL STRATEGY IN LUNG CANCER: TARGETING DRIVER MUTATIONS AND NEXT-GENERATION I/O



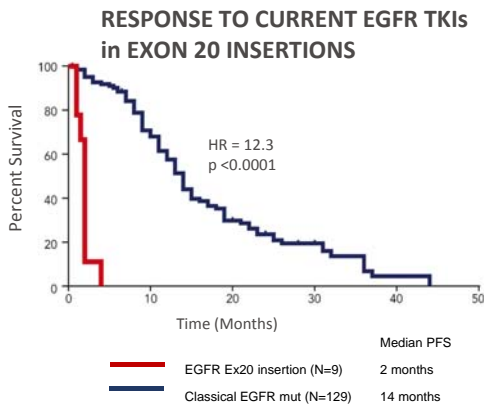
# ALUNBRIG ALTA 1L— POTENTIAL BEST-IN-CLASS PROFILE IN ALK+ NSCLC



Camidge R., WCLC 2018

- Clear superiority to crizotinib and early separation in PFS curve
- Primary endpoint (PFS) hazard ratio is 0.49
- Risk/benefit profile consistent with the expectations of a best-in-class therapy

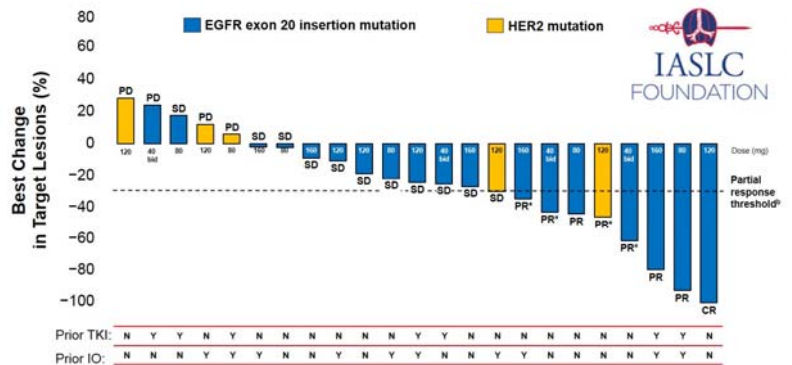
# TAK-788: ADDRESSING UNMET NEED IN EGFR EXON20 MUTATIONS



Overall survival <6 months for exon 20 insertions

Current therapies ineffective for these mutations

## ANTITUMOR ACTIVITY IN ALL PATIENTS TREATED WITH TAK-788 AT A TOTAL DAILY DOSE OF ≥80-160 mg<sup>a</sup>

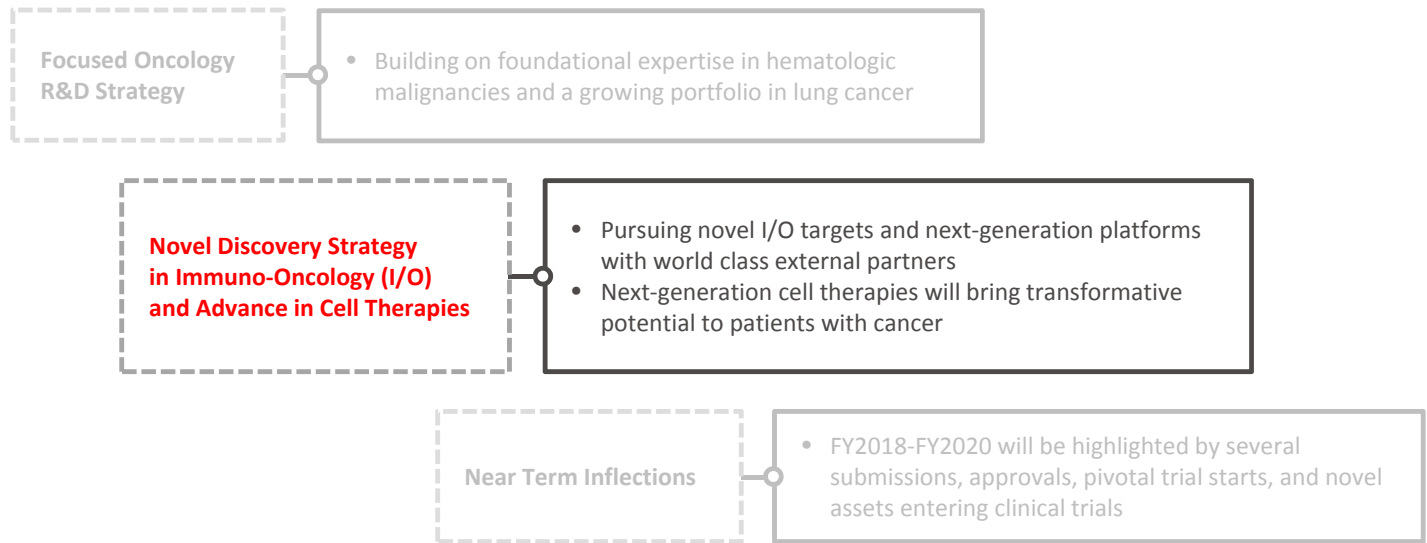


CR, complete response; PD, progressive disease; PR, partial response; SD, stable disease  
<sup>a</sup> Includes 40 mg bid, 80 mg qd, 60 mg bid, 120 mg qd, and 160 mg qd dose groups  
<sup>b</sup> Per RECIST v1.1  
<sup>c</sup> Response awaiting confirmation

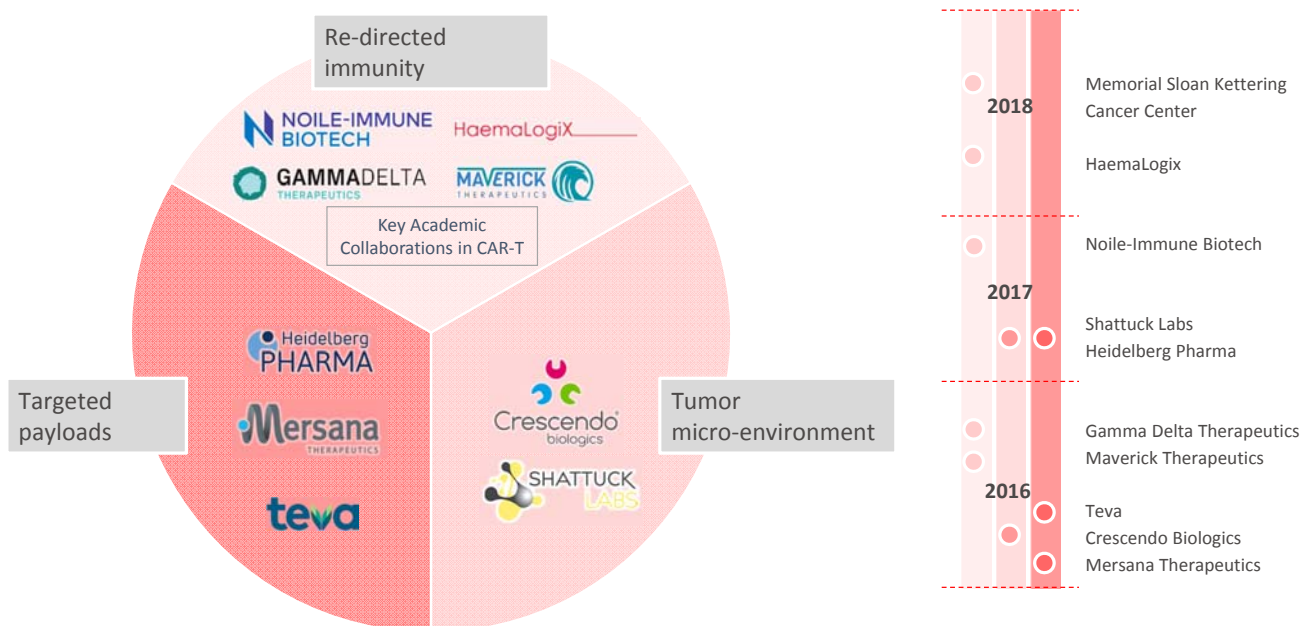
Neal et al., WCLC 2018

Expected to begin registration-enabling Phase 2 trial in FY2018

# ORIENTATION TO OUR ONCOLOGY R&D OVERVIEW



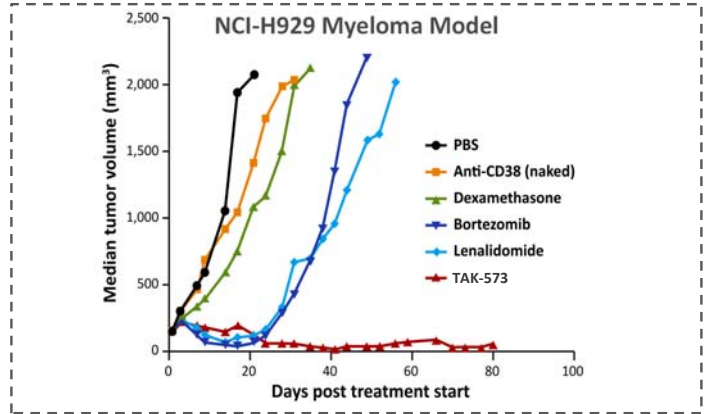
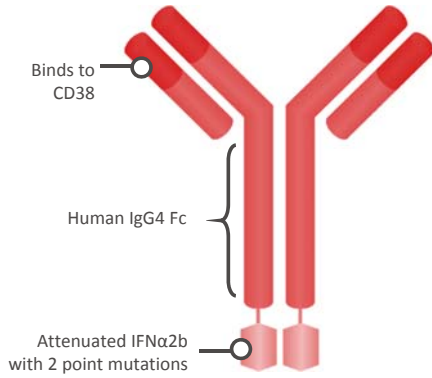
# WORLD CLASS PARTNERS FUELING THE I/O PIPELINE



# TAK-573: BRINGING A NOVEL IMMUNO-CYTOKINE APPROACH TO MULTIPLE MYELOMA



Targeted delivery of attenuated interferon  $\alpha$  to CD38 - a known target in multiple myeloma

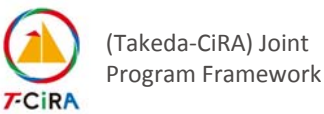


Highly compelling pre-clinical data with TAK-573 in a core area of our clinical development expertise in multiple myeloma Ph 1 currently enrolling for patients with refractory multiple myeloma

# TAKEDA ONCOLOGY AIMS TO BECOME A LEADER IN CELL THERAPIES



TRANSFORMATIVE POTENTIAL UTILIZING NEXT GENERATION CELL THERAPY PLATFORMS



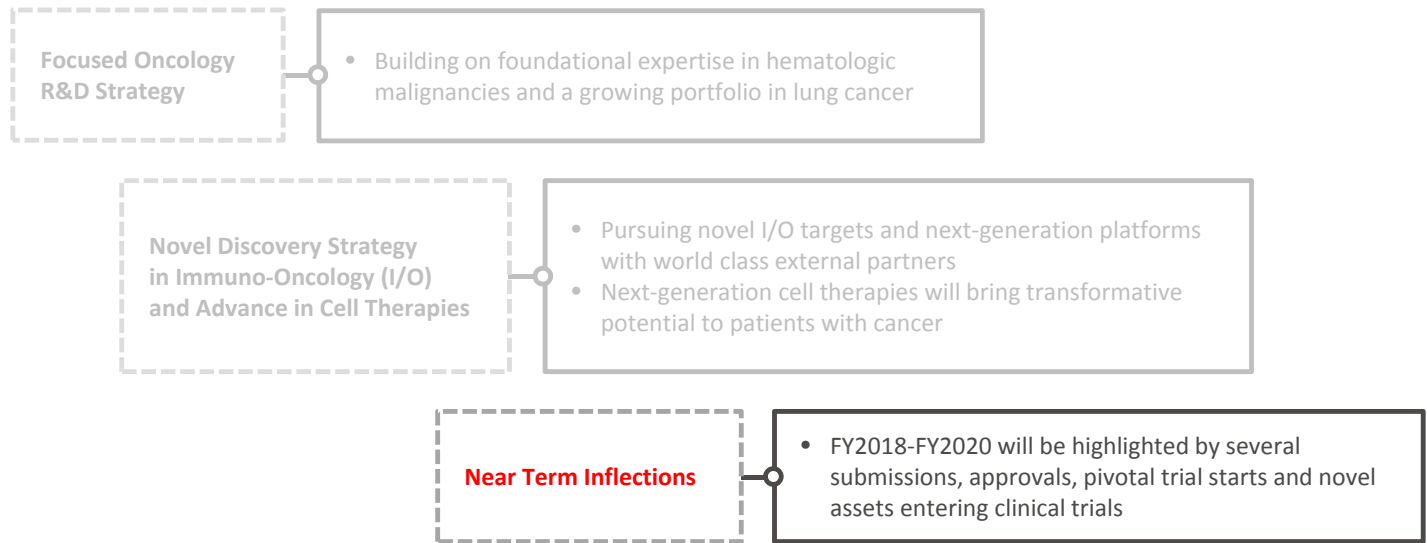
Key Academic Collaborations in CAR-T

Cell therapy engine for Takeda R&D

FY2019: Differentiated CAR-Ts in Phase I  
FY2020+: Other Hematologic Malignancy and Solid Tumor CAR-Ts



# ORIENTATION TO OUR ONCOLOGY R&D OVERVIEW



# AN INNOVATIVE PIPELINE ENHANCED WITH EXTERNAL PARTNERSHIPS

	Discovery/preclinical*	Phase 1	Phase 2	Phase 3	Approved**
<b>Hematologic Malignancies</b>	TAK-169 CD38 SLTA	TAK-079 RR MM, SLE CD38 mAb	TAK-659 Lymphoma SYK, FLT-3 Small Molecule  Alisertib AML AURORA A Small Molecule	Pevonedistat HR-MDS/AML NEDD 8 Small Molecule	NINLARO Amyloidosis, ND MM, R/R MM dara combo, R/R MM Ninlaro/dex,, Maint. MM post-SCT PROTEASOME Small Molecule  ADCETRIS FL HL, FL PTCL, CTCL (JP) R/R HL (CN), sALCL (CN) CD30 mAb ADC  ICLUSIG 2nd-Line Chronic Phase CML, Ph+ ALL BCR-ABL Small Molecule
<b>Lung Cancer</b>		TAK-788 NSCLC Exon 20 EGFR/HER2 Small Molecule	Sapanisertib Endometrial Cancer Lung Cancer mTORC1/2 Small Molecule		ALUNBRIG 2L post-crizotinib ALK+NSCLC (EU, JP, CN), FL ALK+ NSCLC ALK Small Molecule
<b>Immuno-Oncology</b>	TAK-252 PD-1/OX40L  TAK-676 STING	TAK-573 RR MM CD38 Attenukine mAb Fusion Protein  TAK-981 SUMOYLATION Small Molecule			
<b>Solid Tumors</b>		TAK-522 Solid Tumors HER2 mAb ADC  TAK-164 Solid Tumors GCC mAb ADC	TAK-931 Solid Tumors CDC7 Small Molecule	relugolix Prostate Cancer (JP) GnRH antagonist Small Molecule	niraparib*** Ovarian Cancer. PARP 1/2 Small Molecule  cabozantinib*** 1L/2L RCC, 2L HCC Multi-RTK Small Molecule

Pipeline as of September 23, 2018 \* Assets shown in discovery/preclinical and Phases 1-3 explicitly refer to new molecular entities

\*\* Some with active development seeking new or supplemental indications, or approvals in new territories

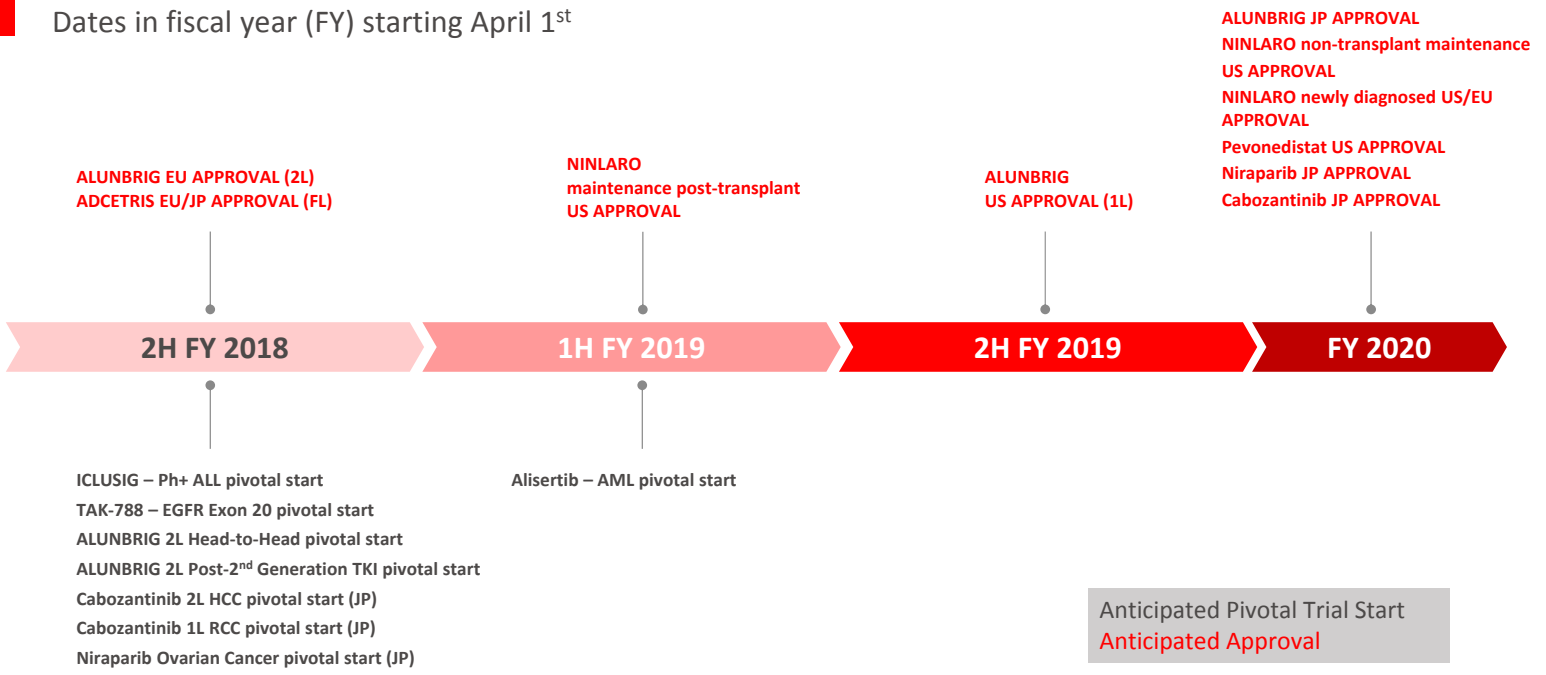
\*\*\* In pivotal trial for Japan approval

Note: Takeda holds the right to develop and commercialize Adcetris in ex-US/Canada. For Niraparib and Cabozantinib, Takeda holds the right to develop and commercialize in Japan and selected Emerging Markets

External collaboration

# EXPECTED KEY ONCOLOGY PORTFOLIO INFLECTION AND MILESTONES

Dates in fiscal year (FY) starting April 1<sup>st</sup>



Projected timelines as of September 23, 2018, subject to change

## CONCLUSION

**1**

Focused on delivering the next approvals for NINLARO, ALUNBRIG, and pevonedistat

**2**

Expanding transformative treatment options in our focus areas of hematologic malignancies and lung cancer with alisertib, TAK-788 and novel CD38 targeted mechanisms

**3**

Harnessing the power of external innovation with a diverse set of world-class partnerships, accelerating novel therapies into the clinic