

December, 2018

# **Forward Looking Statement**

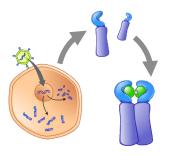
This presentation contains estimates, projections and other forward-looking statements, concerning, among other things: our research and development activities relating to our CaspaCIDe® ("iC9"), GoCAR-T® (incorporating "iMC") and related technologies; our product candidates including rivo-cel™ (previously BPX-501), BPX-601, BPX-602, BPX-603, BPX-701, and rimiducid; the effectiveness of our CaspaCIDe and GoCAR-T programs and their possible range of application and potential curative effects and safety in the treatment of diseases, including as compared to other treatment options and competitive therapies; the success of our collaborations with academic and commercial partners; the timing, progress of enrollment and success of our clinical trials; and the timing of potential European marketing authorization applications for rivo-cel and rimiducid. Our estimates, projections and other forward-looking statements are based on our management's current assumptions and expectations of future events and trends, which affect or may affect our business, strategy, operations or financial performance. Although we believe that these estimates, projections and other forward-looking statements are based upon reasonable assumptions, they are subject to numerous known and unknown risks and uncertainties and are made in light of information currently available to us. Many important factors, in addition to the factors described in this presentation, may adversely and materially affect our results as indicated in forward-looking statements. All statements of historical fact are forward-looking statements.

Estimates, projections and other forward-looking statements speak only as of the date they were made, and, except to the extent required by law, we undertake no obligation to update any forward-looking statement. These statements are also subject to a number of material risks and uncertainties that are described more fully in Bellicum's filings with the Securities and Exchange Commission, including without limitation our annual report on Form 10-K for the year ended December 31, 2017 and our quarterly report on Form 10-Q for the period ended September 30, 2018.



### **Our Platform**

Enhancing T-cell function via controllable molecular switches



### **Our Clinical Programs**

Rivo-cel\* (BPX-501)

Hematologic Malignancies & Inherited Blood Disorders **BPX-601** 

Pancreatic, Gastric, & Prostate Cancers **BPX-701** 

AML / MDS Uveal Melanoma



### Striving to deliver cures through controllable cell therapy

### **Our People**

SAN FRANCISCO
O HOUSTON

GREATER LONDON

~150 Bellicians

### **Our Capabilities**

Over 250 patients treated in clinical studies to date



Translational
Research
& Clinical
Development



GMP
Viral Vector
& Cell
Manufacturing

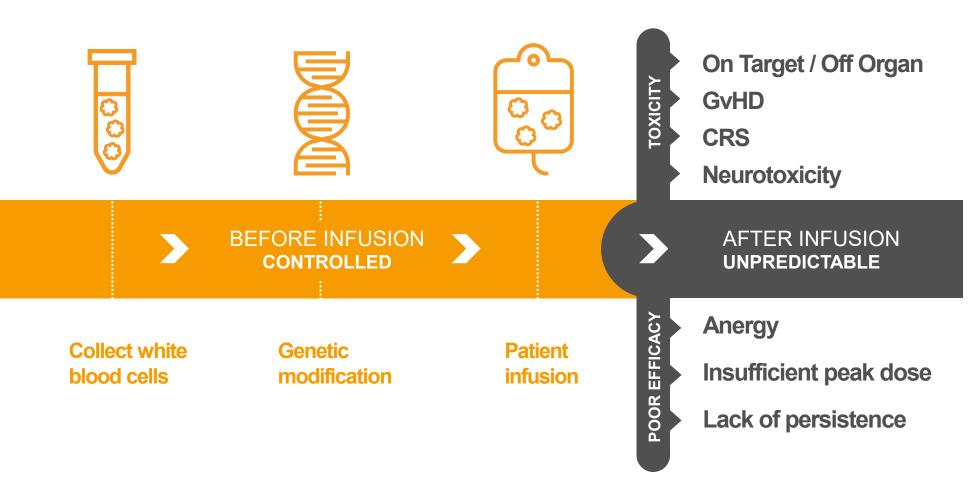


Allogeneic & Autologous Cell Therapy Supply Chain



# **Current Limitations of Cell Therapy**

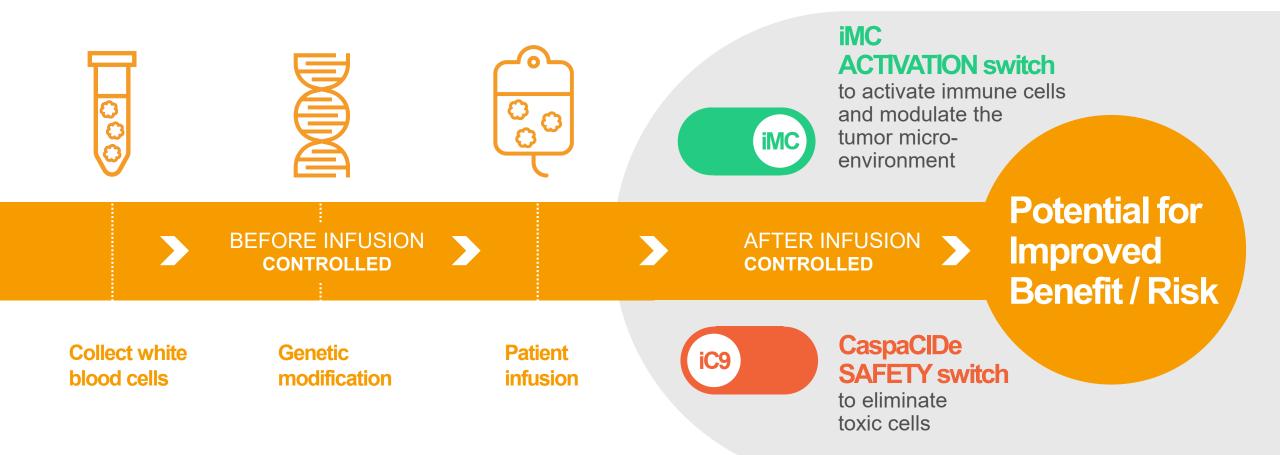
Most cell therapies can only be controlled **before** infusion





# Our Approach to Enhance Cell Therapy

Bellicum's molecular switches allow control after infusion

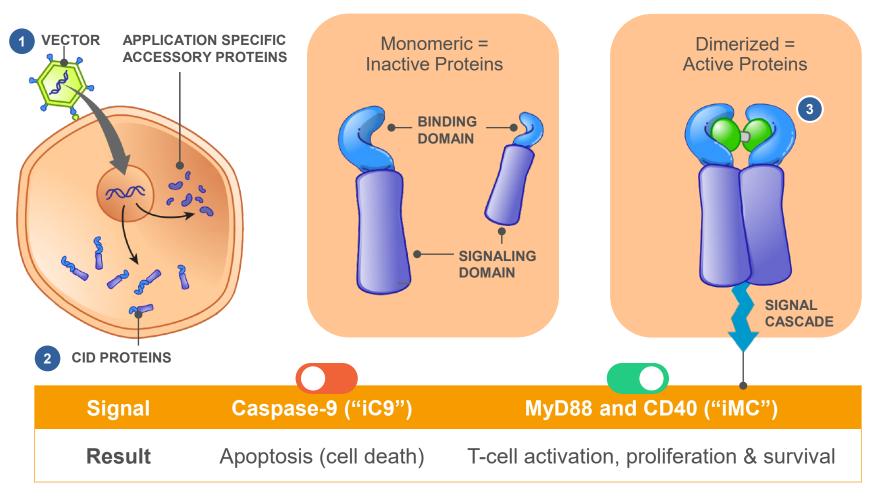




# Chemical Induction of Dimerization ("CID") Molecular Switch Platform

Rimiducid infusion activates signaling pathways to control T-cell function

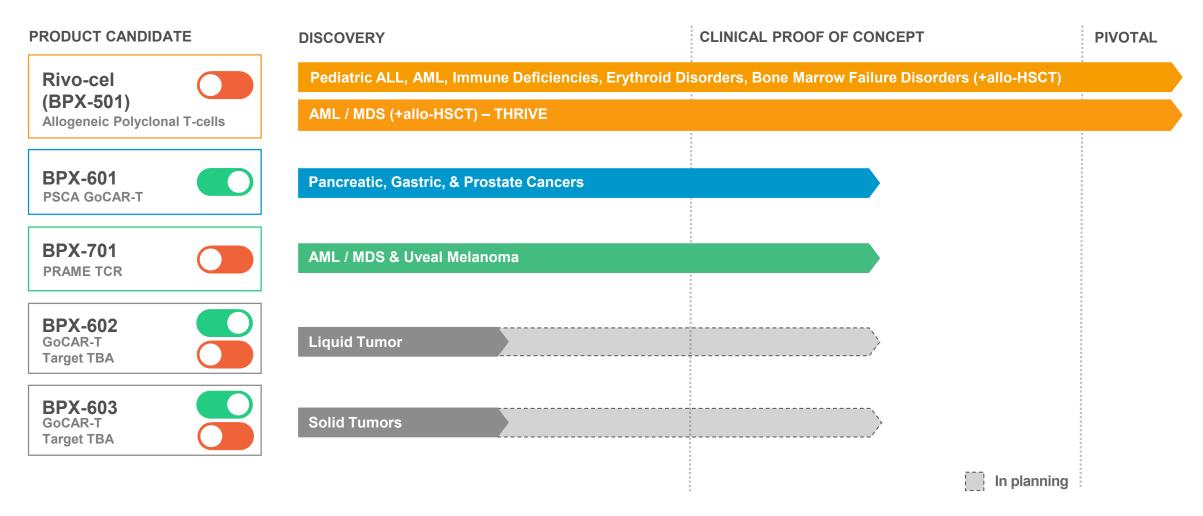
- Viral transduction transfers the DNA from a **vector** into the target cell nucleus.
- Vector-derived DNA directs expression of CID and accessory proteins.
- Rimiducid dimerizes the CID proteins, thus turning on the signal cascade.





## **Highly Differentiated Portfolio**

Control switch selected to address most critical situation-specific challenge





## Rivo-cel (rivogenlecleucel) Product Overview

Allogeneic polyclonal T-cells incorporating the CaspaCIDe safety switch (formerly BPX-501)

## Unmet Need in Leukemias, Lymphomas, and Inherited Blood Disorders

- Potentially cured by allogeneic hematopoietic stem cell transplantation (allo-HSCT)
- Allo-HSCT patients without HLA-matched related donor are at higher risk of morbidity & mortality. Leading causes:
  - Malignant relapse
  - Viral infection
  - Graft Versus Host Disease (GvHD)
- ~70% of allo-HSCT patients lack a HLA-matched related donor
  - Europe 11,700 patients / year
  - US 6,300 patients / year
- ~26,000 additional eligible patients forgo allo-HSCT annually in Europe & US in part due to risks

### **Anticipated Rivo-cel Benefits**

- Graft versus leukemia (GvL) to prevent malignant relapse and extend survival
- Reduce transplant-related mortality (TRM) due to infection
- Ability to treat GvHD via CaspaCIDe

### **Program Update**

- Enrollment complete in Phase 1/2 BP-004 pediatric basket trial – EMA filing planned for 2019
- Randomized global Phase 2/3 THRIVE AML / MDS trial in patients 12+ to initiate by year-end 2018



# **BP-004 Study: Schema and Enrolled Populations**

Phase 1/2 study of rivo-cel gene modified donor T cells following TCR  $\alpha\beta$  depleted allo-HSCT

High risk malignancies and non-malignant disorders

αβ T-cell and B-cell depleted haplo-HSCT without GvHD Prophylaxis



Rivo-cel



Rimiducid for patients who develop visceral GvHD or are refractory to SOC treatment

### **Enrolled Populations**

N = 249			
Malignant (N = 117)	Non-Malignant (N = 132)		
Diagnosis	Diagnosis		
Acute lymphocytic leukemia (ALL)	Primary Immune Deficiencies		
Acute myeloid leukemia (AML)	β Thalassemia Major		
Other	Other Erythroid Disorders		
	Bone Marrow Failure Disorders		

#### **Outcomes**

#### Rivo-cel:

- Event-free survival at 180 days (regulatory endpoint)
  - TRM/NRM, severe GvHD, life-threatening infections
- Progression-free survival
- Disease status

#### Rimiducid:

GvHD response

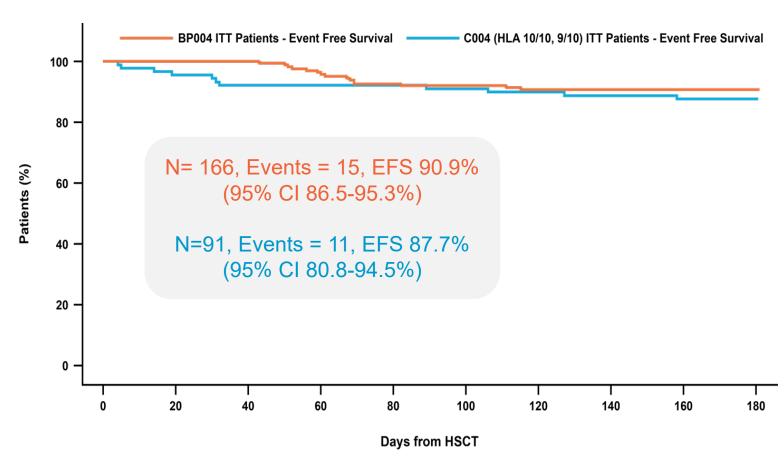


### Interim Six-Month Event-Free Survival Results

Rivo-cel interim event-free survival comparable to MUD HSCT

- C-004 is an observational trial of pediatric patients with malignant (67%) or non-malignant (33%) disease who underwent a MUD HSCT
- Non-inferiority of rivo-cel EFS at 180 days to MUD HSCT is required for EMA approval
- Full analysis with statistical comparisons of patients who received rivo-cel or a MUD HSCT planned for 2019

### **Event Free Survival at 180 days**

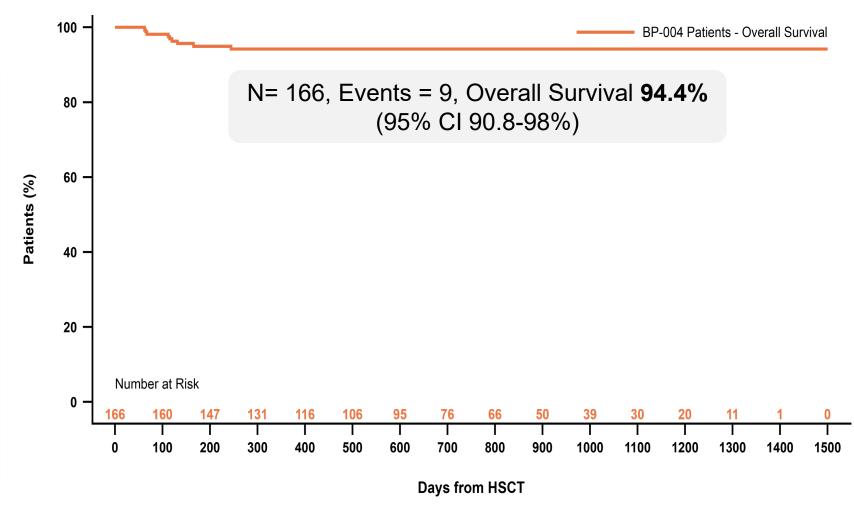


### **Interim Survival Results**

High rates of disease-free and overall survival in rivo-cel treated patients

With median 20.3 months (0.5 - 47.4 months):

- Relapse-free survival
   82.9% in malignant
   patients
- Disease-free survival
   95.2% in non-malignant patients

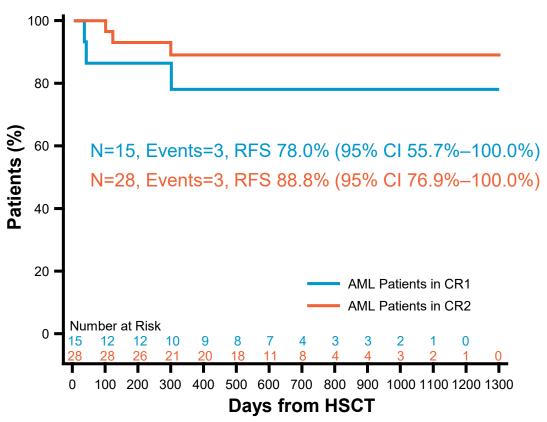




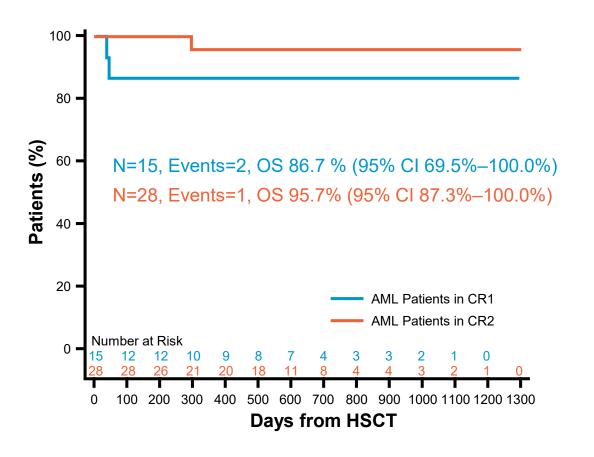
# **Interim AML Efficacy Outcomes\***

High rates of RFS and OS in patients in first or second complete remission

#### **Relapse-Free Survival by CR Status**



### **Overall Survival by CR Status**



RFS, Relapse-free survival; OS, Overall survival; CR, Complete Remission \*Efficacy evaluable population

Data cutoff date: September 17, 2018

## Interim GvHD Response to Rimiducid

High rates of response in patients refractory to standard of care treatment

# Methods & Evaluable Population

### **Efficacy Results**

### **Translational Results**

Patients who developed visceral GvHD or were refractory to SOC treatment were eligible to receive ≥1 dose (up to 3 at 48 hour intervals) of rimiducid (0.4 mg/kg)

Of 238 GvHD-evaluable patients:

- 35.7% (85/238) experienced any grade acute or chronic GvHD
- 28.2% (24/85) of patients with GvHD received rimiducid

Best overall response of 70% 7 days post-rimiducid

- 9 CR and 7 PR
- Median time to response of 1 day (1 - 4 days)

Four patients in PR or not evaluable at day 7 achieved CR within 30 days post-rimiducid

Reduction in rivo-cel serum levels observed in all patients receiving rimiducid<sup>1</sup>

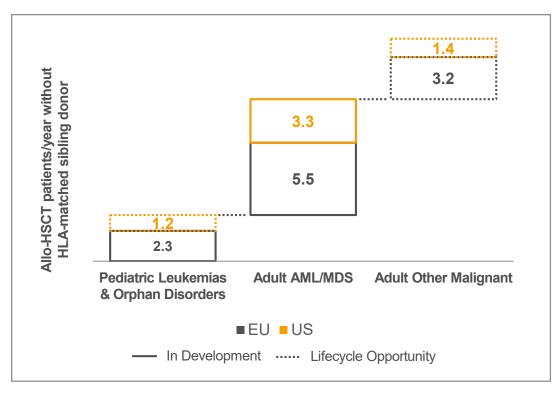
Rimiducid eliminates the most highly activated rivo-cel T cells which express the highest level of iC9<sup>2</sup>, leaving remaining cells to re-expand

 79% (11/14) malignant patients receiving rimiducid remain relapse free



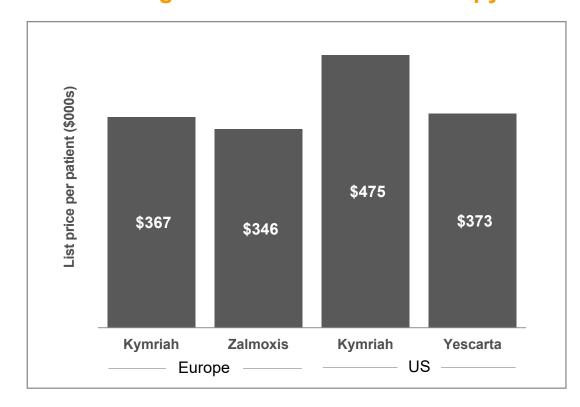
# **Rivo-cel: Compelling Commercial Opportunity**

#### Large addressable patient population (000's)



Additional ~26k eligible patients per year without HLA-matched donor who forgo transplant represent market growth opportunity

### Pricing reflects value in cell therapy





## Rivo-cel: Compelling Value Proposition

Potential Rivo-cel Benefits

# Potential to address the leading causes of morbidity & mortality in curative allo-HSCT

- Malignant relapse
- Viral infection
- GvHD



# May reduce healthcare costs for the most complex allo-HSCTs: those without HLA-matched sibling

- May shorten hospital length-ofstay and lower readmission
- May reduce infectious and GvHD complications during and post-discharge
- Eliminates MUD graft procurement costs



# Potential to reduce disease burden and associated costs

- May reduce rate of malignant relapse
- Potential to enable curative allo-HSCT in patients without HLAmatched donor





### **BPX-601 Product Overview**

GoCAR-T targeting Prostate Stem Cell Antigen (PSCA)

**Unmet Need** 

### **Strategic Rationale**

### **Program Update**

# High unmet need in solid tumors expressing PSCA

	Incidence (US)	Annual Deaths (US)	% Expressing PSCA
Pancreatic	55k	44k	~60%
Prostate	165k	29k	75-90%
Gastric	26k	11k	76-89%

# Attractive first-in-class solid tumor CAR-T opportunity

- Clinically validates the GoCAR-T platform, designed to:
  - Drive T-cell activation, proliferation, and persistence
  - Modulate the tumor microenvironment to enhance immune activity

# Phase 1 trial enrollment ongoing

- Trial amended Q3 2018
  - Standardized Cy/Flu conditioning
  - Added gastric & prostate cancers
- Initial data presentation planned for December, 2018



# **GoCAR-T: Designed to Enhance Efficacy**

Broad immunological effects of inducible MyD88/CD40 (iMC) activation switch

Proliferation/ CAR **TCR** Resistance to **Targeting Targeting** Self-renewal **Inhibitory Factors GoCAR-T HOST IMMUNITY** Costimulation Tumor **Antigen Immune Cell Presentation Visibility** Recruitment



### **BPX-601 Phase 1 Trial Dose Escalation**

BP-012 trial in relapsed/refractory pancreatic, gastric, and prostate cancers

	Cohort 0 (Lead-in)	Cohort 3	Cohort 4	Cohort 5a	Cohort 5b
Patient Population	3L+ Pancreatic			2L Pancreatic 2L Gastric HR-Refractory Prostate	
BPX-601 Dose x10 <sup>6</sup> cells/kg @ Day 0	1.25	1.25	2.5	5	.0
Rimiducid Dose mg/kg @ Day 7	None	0.4	0.4	0.4	
Conditioning	Cytoxan 1g/m² @ Day -3			Cytoxan 1g/m² @ Day -3	Cytoxan 0.5g/m² Fludarabine 30mg/m² @ Days -5, -4, -3
Status	Enrolled			Active	

#### **Trial Highlights and Updates**

- Standard 3+3 dose escalation / de-escalation design to establish MTD or RP2D
- Q3 amendment updated conditioning regimen and adds gastric and prostate cancer patients
- Schedule for <u>repeat dosing</u> of <u>rimiducid</u> to be evaluated after cohort 5
- Abstract accepted for oral presentation at ESMO Immuno-Oncology meeting in December



### **BPX-701 Product Overview**

TCR targeting Preferentially Expressed Antigen in Melanoma (PRAME) incorporating CaspaCIDe

**Unmet Need** 

### **Strategic Rationale**

**Program Update** 

# Several hematologic and solid tumors express PRAME

 Predominantly expressed in AML, uveal melanoma, sarcomas and neuroblastomas

# Attractive first-in-class opportunity targeting a cancer/testis antigen

 Supports further proof-of-concept of CaspaCIDe in T-cell therapy

# Phase 1 trial enrollment ongoing

- Adding sites beginning Q4 2018 to accelerate enrollment
- Initial data presentation planned for 2019



# **Anticipated Program Milestones**

	2018	2019
Rivo-cel	Initiation of randomized Phase 2/3 study in AML / MDS (patients 12+)	Final analyses of BP-004 and C-004 trials  MAA submissions for rivo-cel and rimiducid for pediatric patients
BPX-601	Abstract accepted for oral presentation of initial Phase 1 results at ESMO Immuno-Oncology Congress	Presentation of updated Phase 1 results, including pancreatic, gastric & prostate cancers
BPX-701		Presentation of initial Phase 1 results
PIPELINE		IND submissions for two new dual-switch GoCAR-T programs



# **Financial Highlights**

# Cash and Investments

\$118.4 million of cash, restricted cash and investments as of September 30, 2018



# **Cash Guidance**

Expect that current cash resources will be sufficient to meet operating requirements through 2019



# **Shares Outstanding**

**43.4 million shares** of common stock at **September 30, 2018** 





# Bellicum

Striving to deliver cures for cancer and rare diseases through controllable cell therapy

### Industry-leading CID switch platform for controlling cell therapy

- iMC activation switch to enhance efficacy, particularly in solid tumors
- CaspaCIDe safety switch to manage toxicity

# Growing portfolio of differentiated oncology & hematology programs

- Rivo-cel Best-in-class allogeneic T-cell product in hematologic malignancies and inherited blood disorders
- BPX-601/701 First-in-class GoCAR-T and TCR products
- BPX-602/603 First controllable
   "dual-switch" GoCAR-Ts to enter clinic in 2019

# Fully integrated cell therapy capabilities support future growth

- Robust R&D, clinical, and manufacturing capabilities
- Ongoing collaborations with other leaders in the field



## **Bellicum Leadership Team**



**Rick Fair** President & CEO Rick has a 20-year track record as a strategist and commercial leader in the biopharmaceutical industry. Rick joined Bellicum in 2017 from Genentech/Roche.



**David Spencer Chief Technology Officer** Dave is the inventor of CID technology, and codeveloped the first clinical applications of the technology, DeCIDe® and CaspaCIDe®.



**Thierry Darcis** General Manager, Europe Thierry has over 20 years of experience in European and global marketing, product development, and operations. He joined Bellicum in 2018.









































William Grossman **Chief Medical Officer** Bill has an extensive background in the development of cancer immunotherapies and ioined Bellicum in 2018 from Genentech/Roche.



**Aaron Foster** Vice President Translational Research & **New Product Development** Aaron leads the CAR and TCR gene-modified T-cell programs that are developing systems for controlling T-cell behavior in vivo using molecular switch technology.



**Shane Ward** General Counsel & **Corporate Secretary** Shane is a seasoned public company executive with over 20 years of biotechnology and pharmaceutical industry experience. Shane joined Bellicum in 2018.



**Gregory Naeve** Chief Business Officer Greg joined in 2017 from Pfizer, where he was ImmunoOncology & Cell Therapy Lead in their External Research and Development Unit. He was previously a Principal at The Column Group.



**Alan Smith Exec. Vice President Tech Operations** Alan has over 30 years of experience in R&D, Manufacturing and Quality roles in cellular therapeutics.



Vice President Finance & Controller Rosie is a CPA and has over 30 years of experience in finance and accounting at Arthur Andersen and in a variety of companies and industries. She ioined Bellicum in 2014





# **APPENDIX**

### Uptake of Haplo HSCT has accelerated in the last decade

