

# Investor Presentation

Striving to deliver cures through controllable cell therapy

April 2019

# 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-603, BPX-802, 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 other than 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, 2018.

# Investment Summary

## Rivo-cel

Allogeneic polyclonal T-cells for hematologic malignancies and inherited blood disorders (+HSCT)

### European pediatric opportunity clinically de-risked

- 249 patients enrolled in Phase 1 / 2 study
- Late interim results presented at ASH in Dec. 2018 trend toward meeting primary endpoint
- Expect topline data in 1H 2019; MAA filings in 2H 2019
- European HQ and leadership team in place for commercialization prep

### Global trial underway to broaden label

- Enrolling Phase 2/3 THRIVE study in AML and MDS in patients 12+ years old

## GoCAR-T Pipeline

Controllable CAR-T cells designed to optimize efficacy and safety

### BPX-601 GoCAR-T promising early clinical data

- Phase 1 / 2 study enrolling in pancreatic, gastric and prostate cancers
- Initial safety data on 12 pancreatic patients presented at ESMO-IO in Dec. 2018 indicate attractive safety profile and early clinical activity
- Trial amendments to lymphodepletion regimen and activation molecule administration to enhance potential clinical response

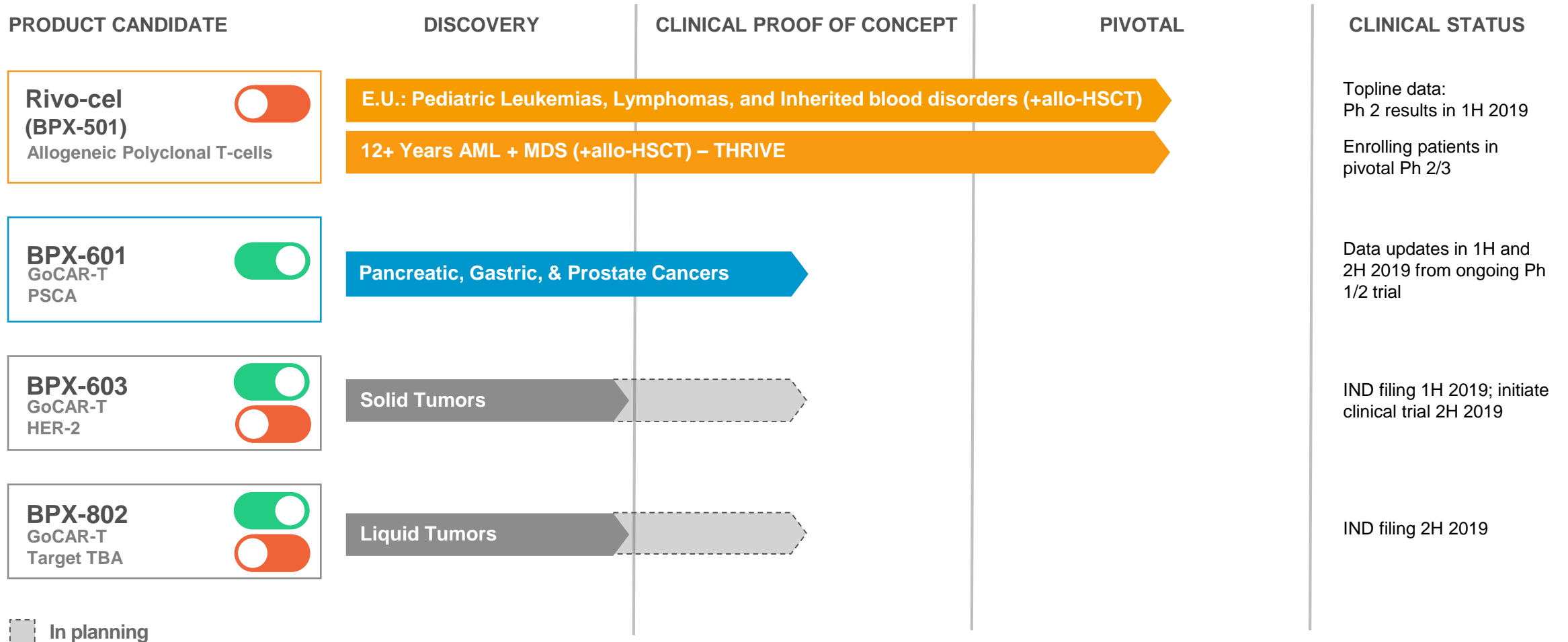
### Two dual-switch GoCAR-T candidates to IND in 2019

- BPX-603 targeting HER2 antigen in solid tumors
- BPX-802 targeting liquid tumors, target antigen TBA

**Cash of \$98.0MM as of December 31, 2018; Cash Runway Through 2019**

# Development Pipeline: Rivo-cel and GoCAR-T

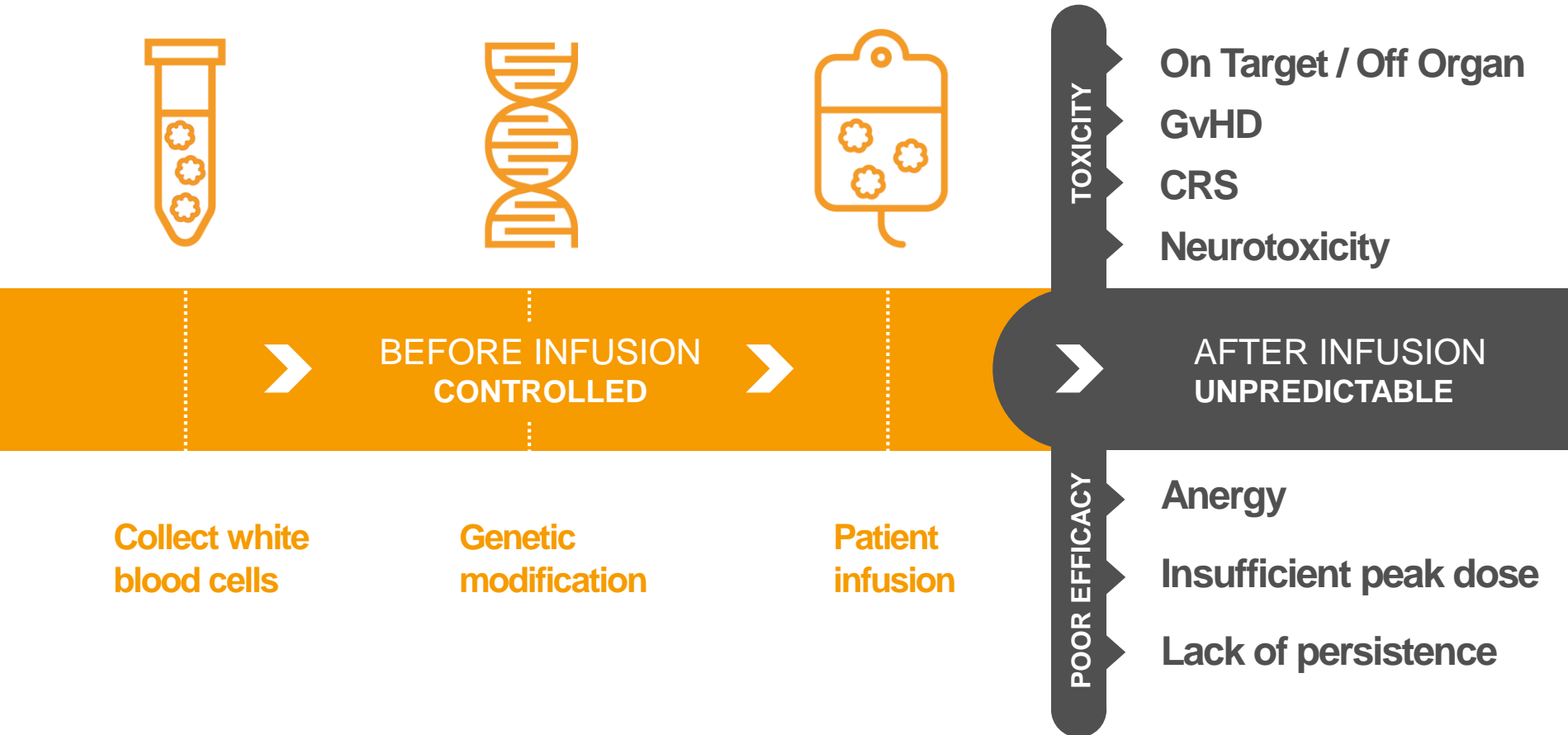
Controllable cell therapies that may represent major advances in liquid and solid tumors



# Technology Overview

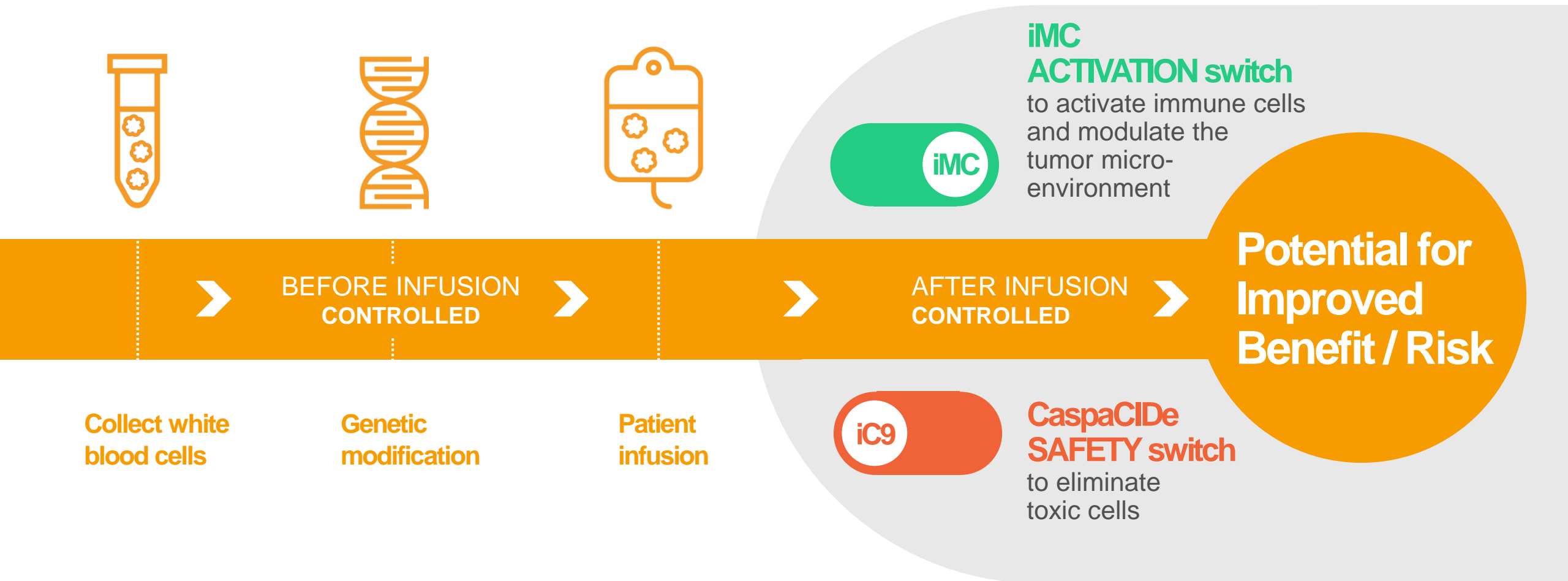
# Most Cell Therapies Only Controlled Before Infusion

Limited ability to expand a narrow therapeutic window



# Bellicum Platform Enables Control After Infusion

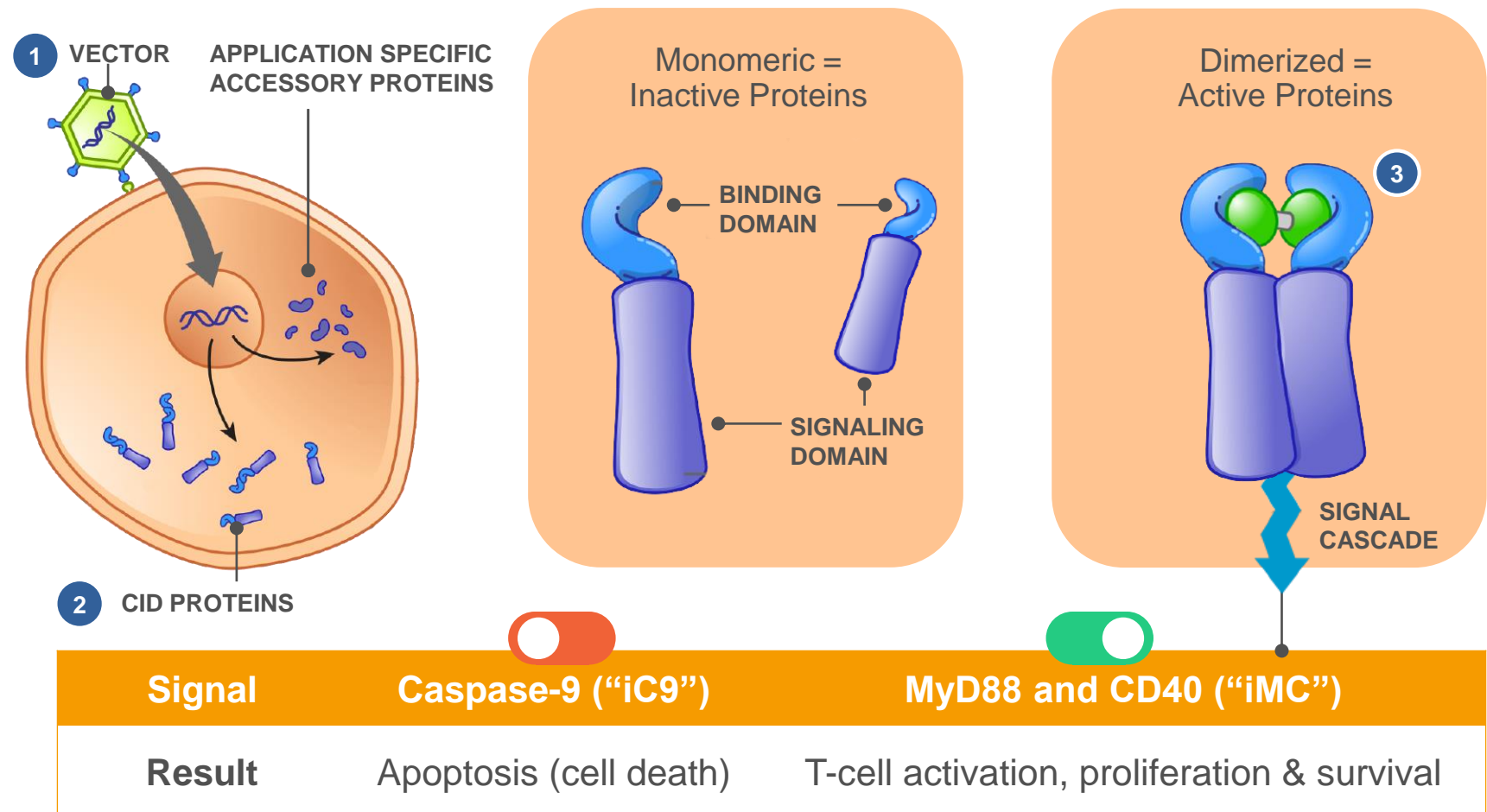
Provides physicians ability to expand the therapeutic window in each patient



# Chemical Induction of Dimerization (“CID”) Molecular Switch Platform

Rimiducid infusion activates signaling pathways to control T-cell function

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





# GoCAR-T Pipeline

# GoCAR-T: Differentiated Approach to Cell Therapy

## Current Challenges in Cell Therapy

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### ⊗ Limited efficacy in solid tumors

- Inadequate cell proliferation and persistence to sustain efficacy
- Inability to overcome immune suppressive factors in tumor microenvironment (TME)

### ⊗ Potential safety issues with more potent approaches

## GoCAR-T Benefits

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### ✓ Potential for enhanced efficacy in solid tumors via iMC signaling

- MyD88 and CD40 are superior co-stimulatory molecules with potential for greater cell expansion and persistence
- Modulates the tumor microenvironment, overriding common inhibitory pathways (PD-1, PGE2, TGF-β)
- Enhances host immune activity by inducing pro-inflammatory cytokines and chemokines

### ✓ Potential for enhanced safety

- iMC provides control over timing and frequency of co-activation
- CaspaCIDE rapidly eliminates a majority of CAR-T cells to manage acute toxicities

# BPX-601 GoCAR-T Targeting PSCA

## Product Summary

- Attractive first-in-class solid tumor CAR-T opportunity
- First-in-human experience with iMC
- Initial Phase 1 results presented in Dec 2018 demonstrate:
  - Safety
  - iMC-driven T cell activation
  - Biologic activity
- Phase 1 enrollment ongoing

## Unmet Need

High unmet need in solid tumors expressing Prostate Stem Cell Antigen (PSCA)

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

Incidence and annual deaths: Noone AM, Howlader N, Krapcho M, Miller D, Brest A, Yu M, Ruhl J, Tatalovich Z, Mariotto A, Lewis DR, Chen HS, Feuer EJ, Cronin KA (eds). SEER Cancer Statistics Review, 1975-2015, National Cancer Institute. Bethesda, MD, [https://seer.cancer.gov/csr/1975\\_2015/](https://seer.cancer.gov/csr/1975_2015/), based on November 2017 SEER data submission, posted to the SEER web site, April 2018.  
PSCA expression: Argani et al, Cancer Res 2001; Reiter et al., PNAS 1998; Abate-Daga et al, HGT 2014; Data on file

# BPX-601: Phase 1 Trial Progression

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

|   | Lead-in                               | Dose Escalation | Full Conditioning   | Efficacy Optimized Regimen |
|---|---------------------------------------|-----------------|---|----------------------------|
| Patient Population                                      | 2L to 6L Pancreatic                   |                 | 2L Pancreatic<br>2L Gastric<br>HR-Refractory<br>Prostate                            |                            |
| BPX-601 Dose<br><i>x10<sup>6</sup> cells/kg @ Day 0</i> | 1.25                                  | 1.25, 2.5, 5.0  | 5.0   |                            |
| Conditioning  | Cytoxan 1g/m <sup>2</sup><br>@ Day -3 |                 | Cytoxan 0.5g/m <sup>2</sup><br>Fludarabine 30mg/m <sup>2</sup><br>@ Days -5, -4, -3 |                            |
| Rimiducid Dose<br>@ Day 7                               | None                                  | Single Dose     | Single Dose   | Scheduled Repeat Dosing    |
| Status  | Enrolled & Presented                  |                 | Active  | Pending                    |

## Dose escalation designed conservatively to evaluate safety

- Partial conditioning with Cytoxan monotherapy
- Single dose of rimiducid to activate iMC

## Currently evaluating impact of full conditioning

- Standard cy/flu regimen
- Data presentation: mid 2019

## Next step: efficacy-optimized regimen

- Standard cy/flu regimen plus repeat rimiducid dosing
- Data presentation: late 2019

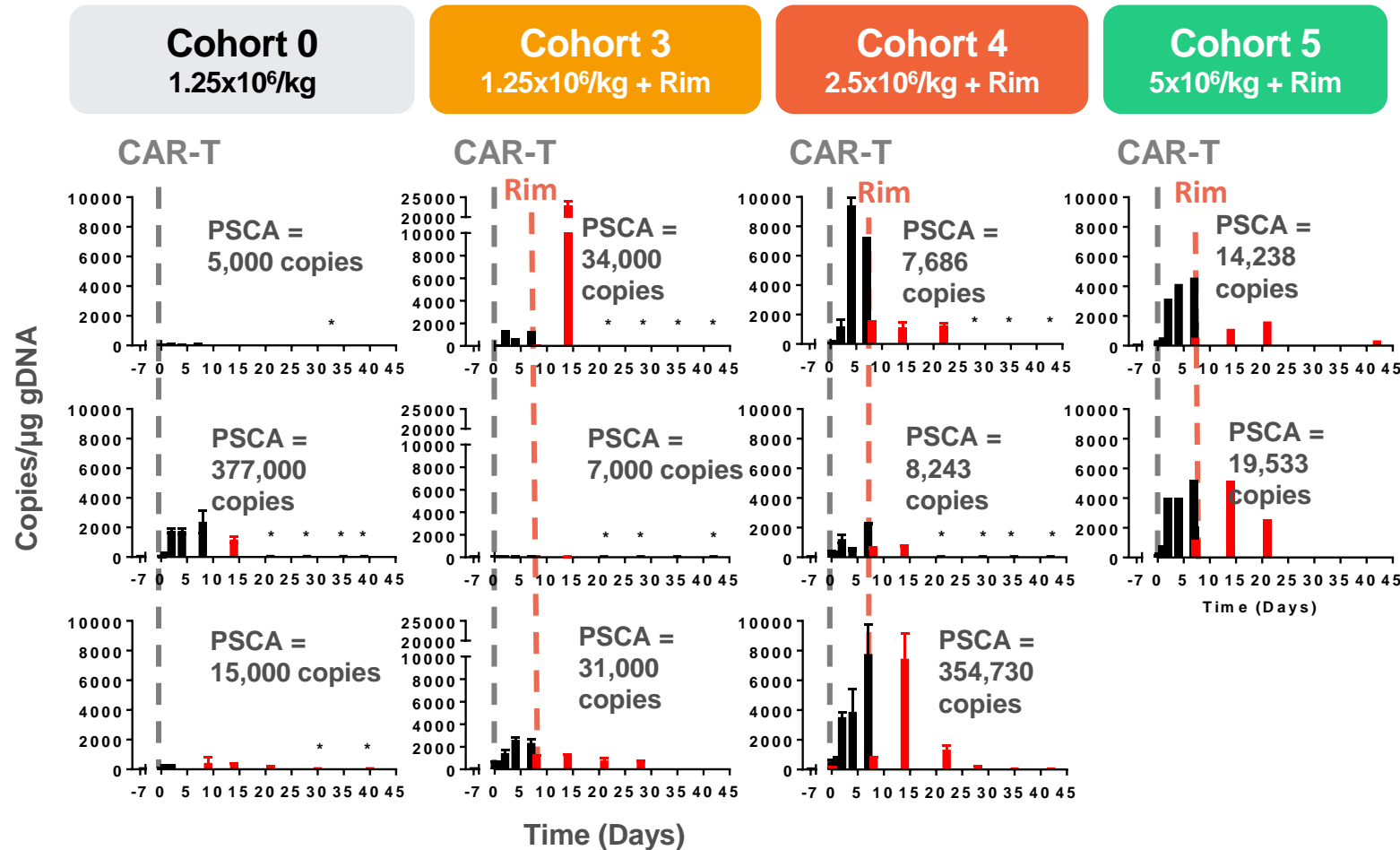
# BPX-601: No Dose Limiting Toxicities Observed

Data presented at ESMO Immuno-Oncology Congress 2018 – clinical cut-off October 29, 2018

| Most common AEs reported by > 1 patient | Total (N=12) |
|---|--------------|
| Any Event, n (%)                        | 12 (100)     |
| Fatigue                                 | 4 (33)       |
| Abdominal pain upper                    | 3 (25)       |
| Hypotension                             | 3 (25)       |
| Abdominal pain                          | 2 (17)       |
| Back pain                               | 2 (17)       |
| Diarrhea                                | 2 (17)       |
| Flatulence                              | 2 (17)       |
| Nausea                                  | 2 (17)       |
| Pyrexia                                 | 2 (17)       |

- **No dose limiting toxicities were observed**
- **Pyrexia was the only treatment-related AE reported by >1 patient (n=2)**
  - Grade 1–2 on Day 0 following BPX-601 infusion
  - Both events resolved within 24–36 hours with supportive care

# BPX-601: iMC-Driven T Cell Expansion & Persistence



- Limited evidence of LD with CTX-only regimen (79% ± 25% of cells remained)
- Rapid cell expansion by Day 4, but no persistence without Rim
- With single-dose Rim:
  - Cell expansion of 3 to 20-fold within 7 days in 4 patients
  - Cell persistence of >3 weeks in 3 patients

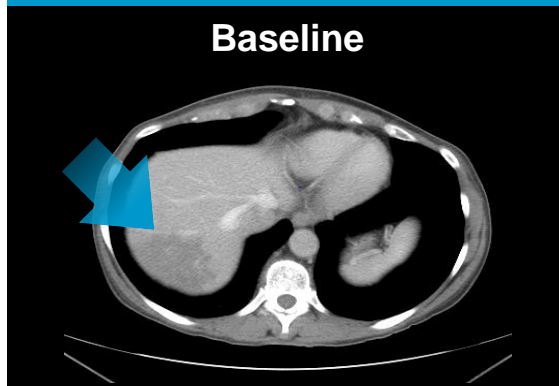
# BPX-601: Evidence of Anti-Tumor Activity

| Cohort | Best Response (RECIST) |    |    |    |
|--------|------------------------|----|----|----|
|        | CR                     | PR | SD | PD |
| 0      | 0                      | 0  | 1  | 2  |
| 3      | 0                      | 0  | 2  | 1  |
| 4      | 0                      | 0  | 2  | 1  |

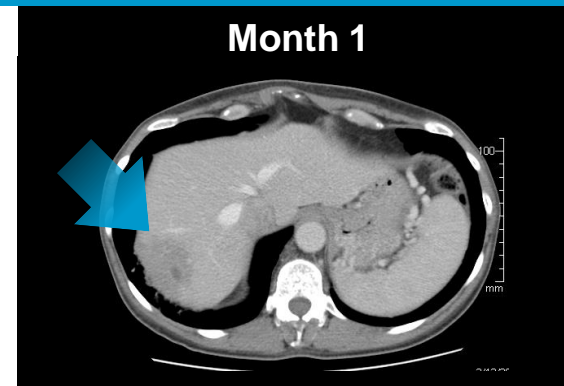
Two patients with SD had tumor shrinkage >20%

Disease control without new therapies: 16 and >18 weeks in 1 and 3 patients, respectively

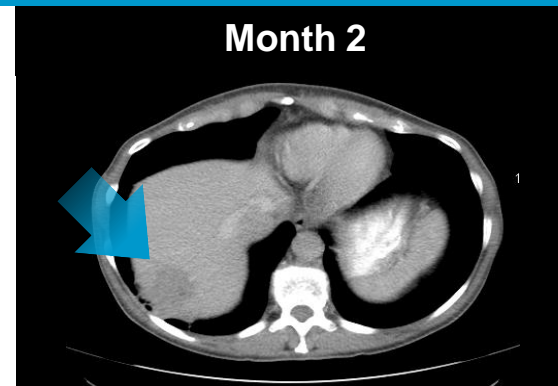
## Patient 3A | 2 prior therapies; PSCA = 34,000 copies



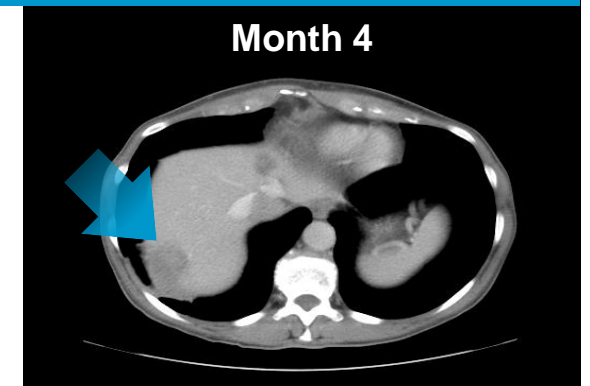
- Lesion longest diameter: 70 mm
- **CA19-9: 294**



- Lesion longest diameter: 57 mm
- **CA19-9: 152.6**
- Overall response: SD (-15%)



- Lesion longest diameter: 49 mm
- **CA19-9: 207.2**
- Possible new lesion
- Overall response: SD (-25%)



- Lesion longest diameter: 40 mm
- **CA19-9: 641.4**
- New lesion confirmed
- Overall response: PD

# BPX-603 Dual Switch GoCAR-T Targeting HER2

## Product Summary

- HER2 is a validated tumor antigen and is expressed on numerous solid tumors with high unmet need
- Historical HER2 CAR-T studies have shown modest overall activity and off-tumor / on-target toxicity
- BPX-603 may address these limitations
  - iMC may increase cell proliferation & persistence, modulate the TME, and enhance host immunity
  - CaspaCIDE may mitigate treatment emergent toxicities

## Unmet Need

| Indication              | Incidence <sup>1</sup> | HER2 <sup>+</sup>   | 5-year OS (Stage IV) <sup>1</sup> |
|-------------------------|------------------------|---------------------|-----------------------------------|
| Gastric                 | 28,000                 | 10-30% <sup>3</sup> | <20%                              |
| Colorectal              | 145,000                | 10% <sup>4</sup>    | <15%                              |
| Ovarian                 | 22,000                 | 20-30% <sup>5</sup> | <30%                              |
| Uterine/<br>Endometrial | 61,000                 | 50-80% <sup>6</sup> | 14-69%                            |
| Glioblastoma            | 12,000                 | 20-30% <sup>2</sup> | <20%                              |

<sup>1</sup>National Cancer Database, American Cancer Society, <https://www.cancer.org>, accessed 21 December 2018; <sup>2</sup>Lui et al., Cancer Res 2004; <sup>3</sup>Gravolos et al., Annals Oncol 2008; <sup>4</sup>Tu et al., Exp Ther Med 2018; <sup>5</sup>Berchuck et al., Cancer Res 1990, Bartlett et al., Brit J Cancer 1996; <sup>6</sup>Grushko et al., Gynecologic Oncol 2008



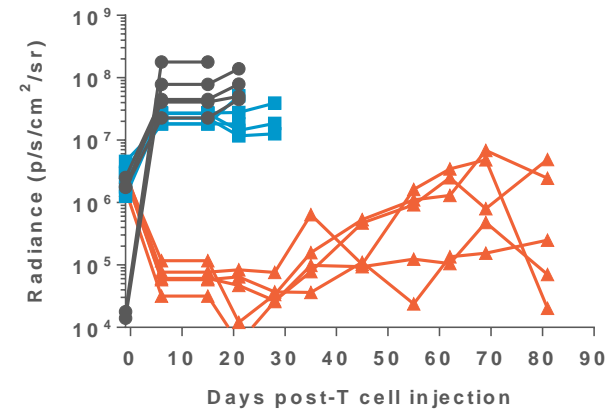
# Historical HER2 Studies: Modest Clinical Outcomes

| Study Properties       | Morgan, 2010                | Ahmed, 2015                       | Hegde, 2017     | Feng, 2017      | Ahmed, 2017                       |
|------------------------|-----------------------------|-----------------------------------|-----------------|-----------------|-----------------------------------|
| <b>Construct</b>       | 4D5-28-BB-z                 | FRP5-28-z                         | FRP5-28-z       | Her2-BB-z       | FRP5-28-z                         |
| <b>Indication(s)</b>   | Metastatic colon            | Sarcomas                          | Sarcomas        | CCA and PCa     | GBM                               |
| <b>Patient number</b>  | 1                           | 19                                | 6               | 11              | 17                                |
| <b>HER2 expression</b> | ≥2+ (IHC)                   | ≥1+ (IHC)                         | ≥1+ (IHC)       | >50% positive   | ≥1+ (IHC)                         |
| <b>CAR-T dose</b>      | 10 <sup>10</sup>            | 10 <sup>4</sup> - 10 <sup>8</sup> | 10 <sup>8</sup> | 10 <sup>6</sup> | 10 <sup>6</sup> - 10 <sup>8</sup> |
| <b>CAR-T expansion</b> | NE                          | Negligible                        | >10,000 copies  | >1,000 copies   | Negligible                        |
| <b>Toxicity</b>        | Lung reactivity             | No DLTs                           | Mild AEs        | Mild AEs        | Mild AEs                          |
| <b>Outcome</b>         | Grade 5 toxicity            | 1 PR, 3 SD, 13 PD                 | 2 CR, 2 SD      | 1 PR, 5 SD      | 1 PR, 7 PD                        |
| <b>Total Responses</b> | 2 CR, 3 PR, 5/54 (9.3% ORR) |                                   |                 |                 |                                   |

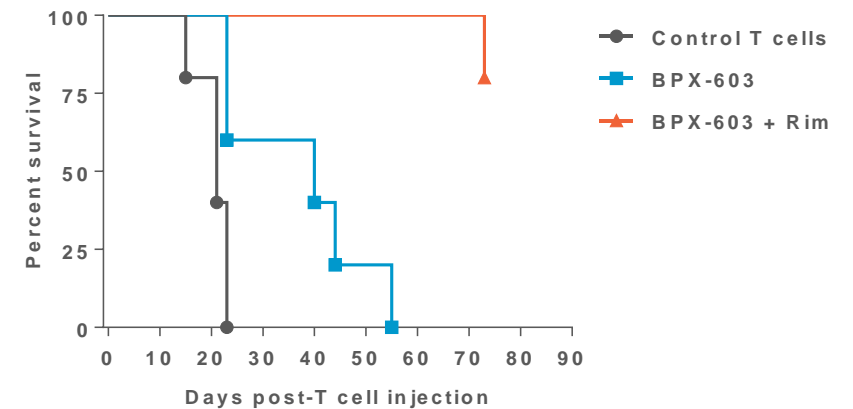
# BPX-603 Pre-Clinical Studies Demonstrate Potential Clinical Benefits

HER2+ A549  
Lung Carcinoma  
( $1 \times 10^4$  T cells)

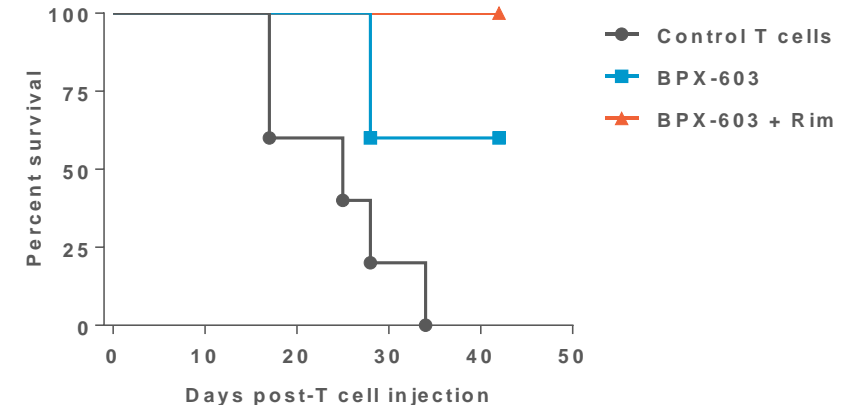
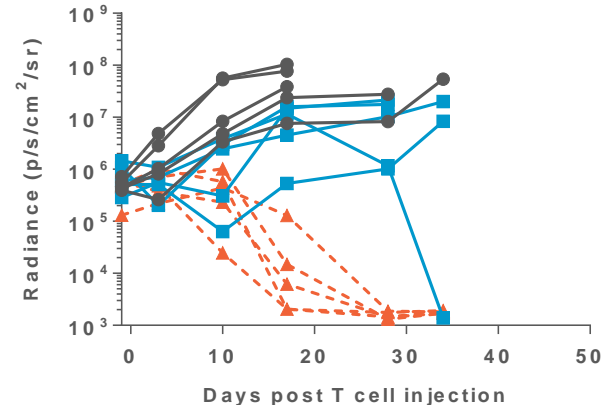
## Tumor growth



## Survival



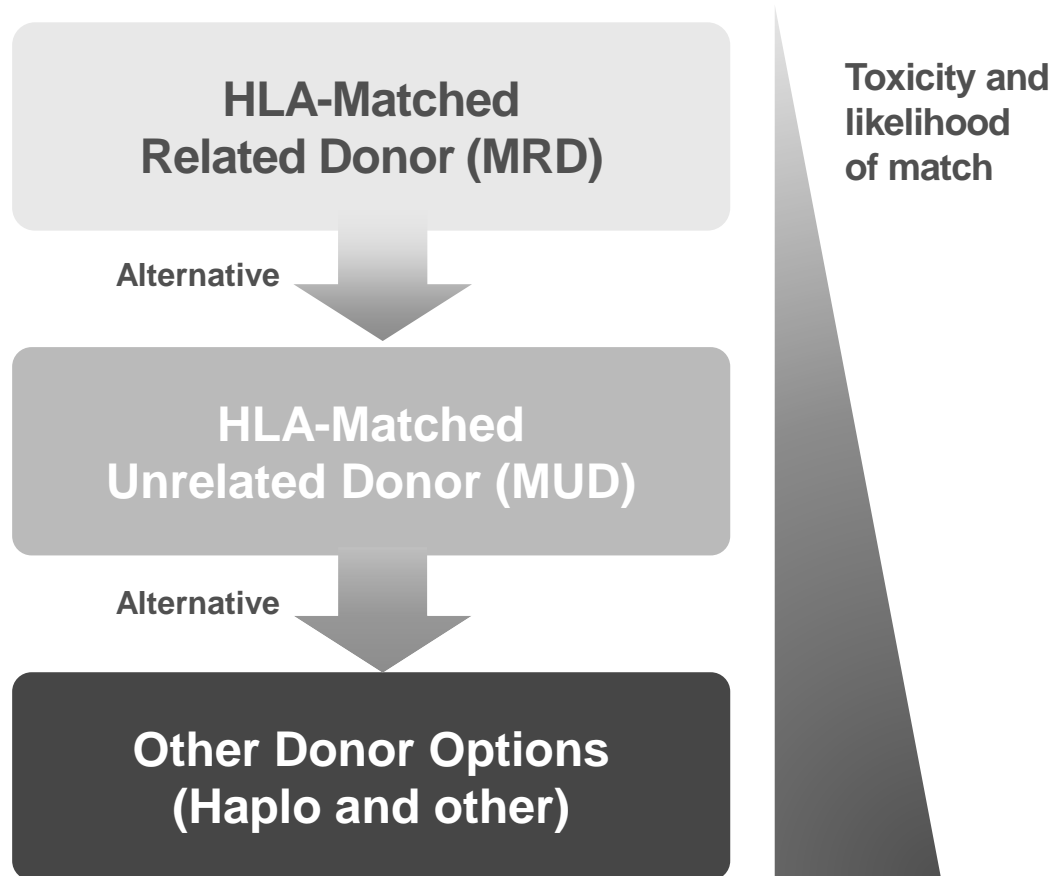
HER2+ OE19  
Esophageal Carcinoma  
( $5 \times 10^6$  T cells)



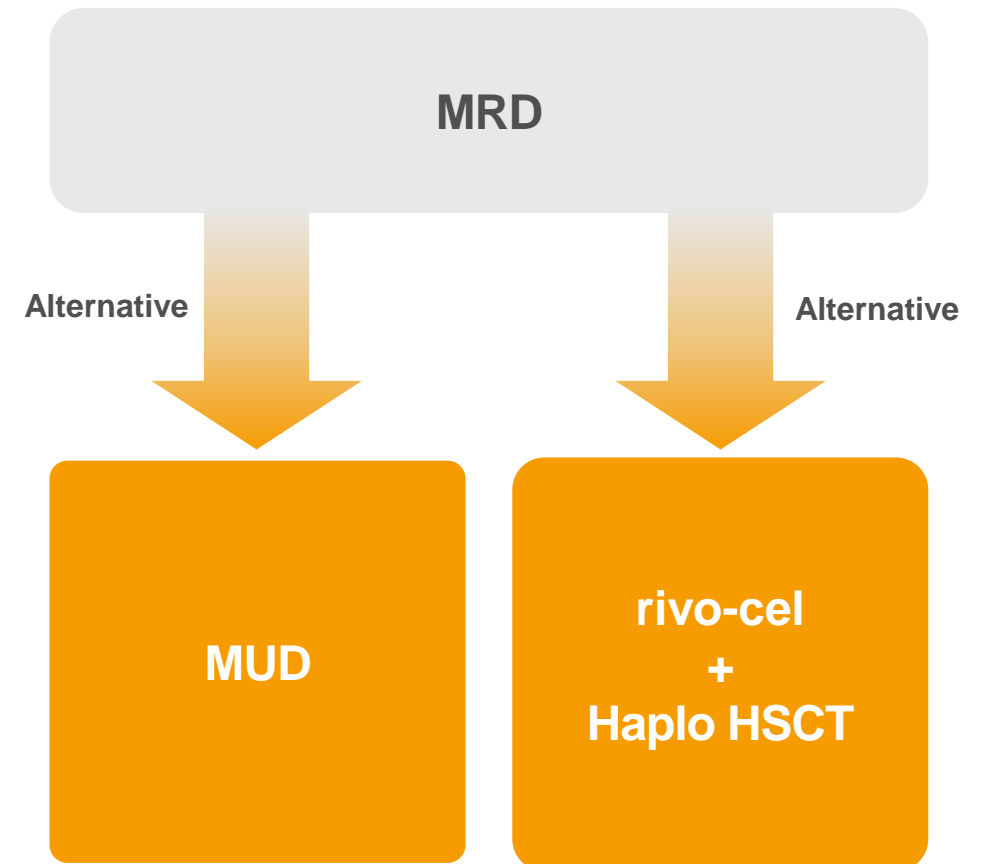
# RIVO-CEL

# Rivo-cel: Opportunity To Transform Treatment Paradigm

## Current HSCT Treatment paradigm



## Potential Future HSCT Treatment Paradigm



# BP-004 Study: Basis for European Pediatric Approval

Phase 1/2 study of rivo-cel in pediatric patients following TCR  $\alpha\beta$  depleted allo-HSCT



## 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, and life-threatening infections
- Progression-free survival
- Disease status

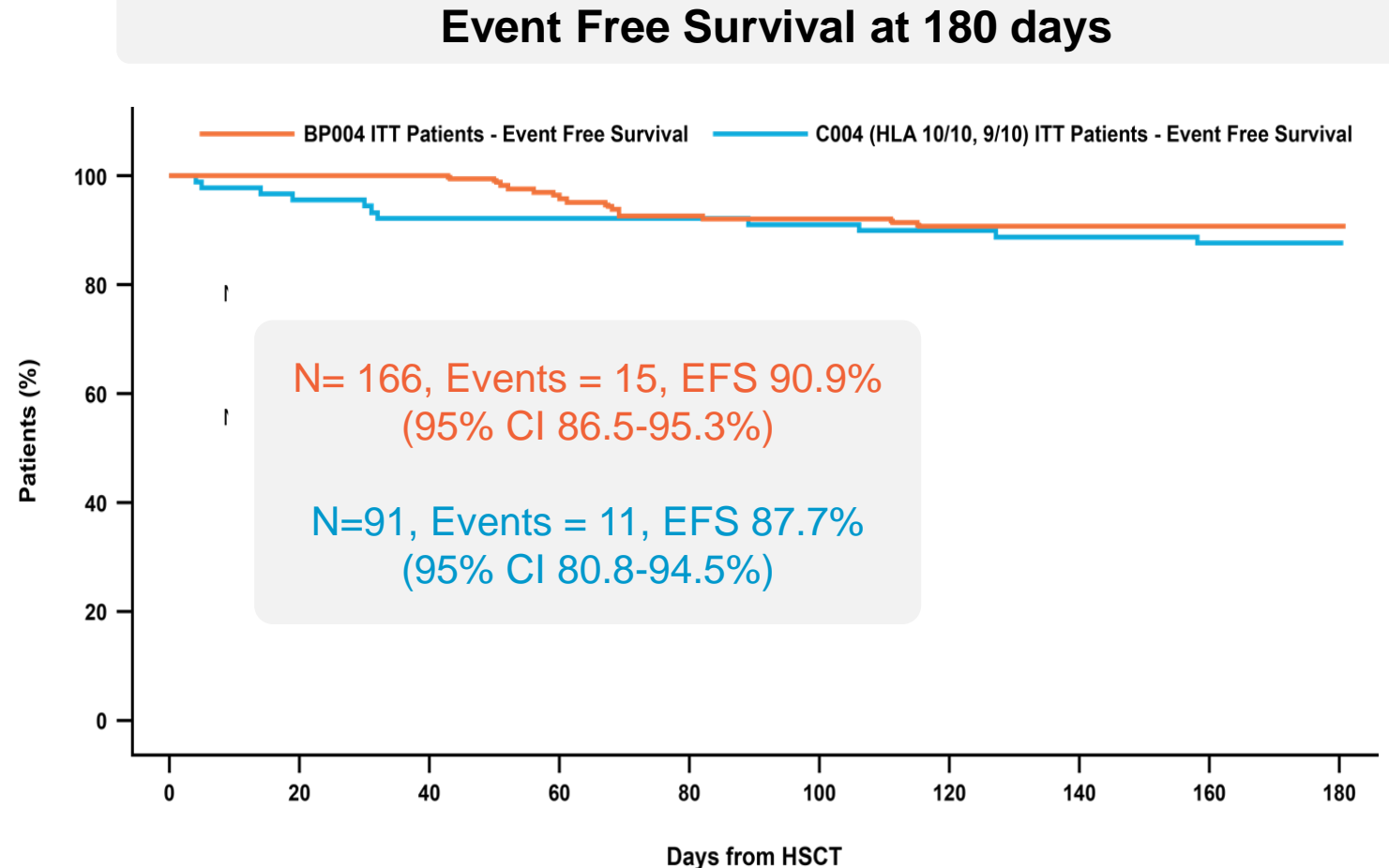
### Rimiducid:

- GvHD response

# Rivo-cel Interim Results Trend Towards Meeting Primary Endpoint

Interim six-month 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

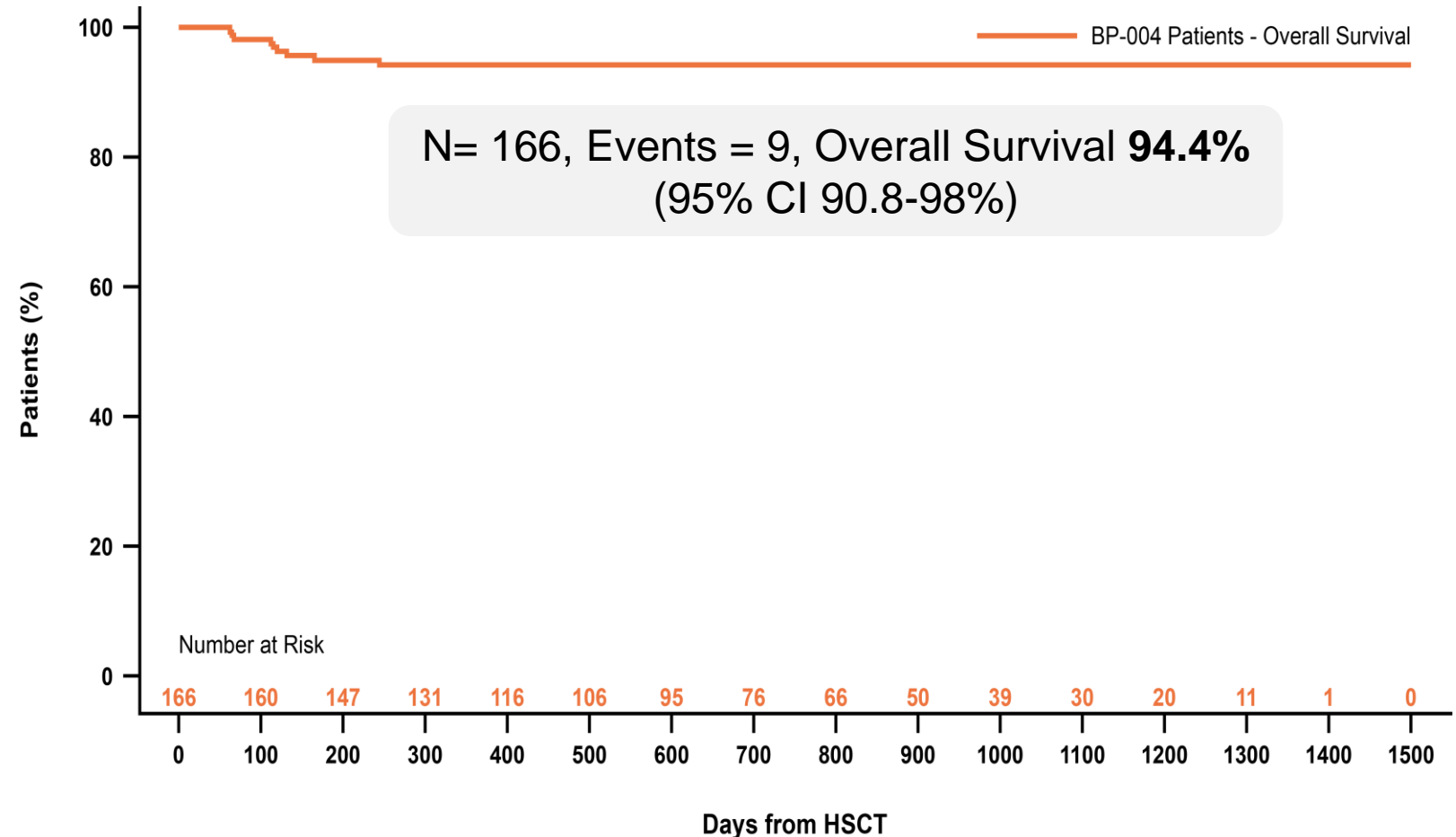


# Rivo-cel: High Rates of Disease-Free and Overall Survival

## Interim survival results

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



# Rivo-cel: High Rates of GvHD Response to Rimiducid

Interim results of response in patients refractory to standard of care treatment

## Methods & Evaluable Population

Patients who developed visceral GvHD or were refractory to SOC treatment were eligible to receive  $\geq 1$  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

## Efficacy Results

### **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

## Translational Results

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

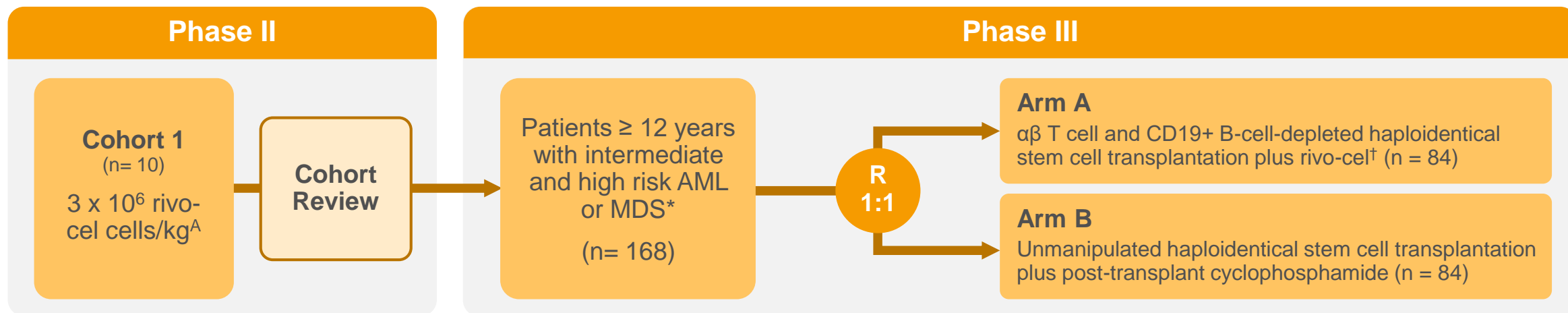
Rimiducid eliminates the most highly activated rivo-cel T cells which express the highest level of iC9<sup>2</sup>, leaving remaining cells to re-expand

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



# THRIVE: Registrational Trial in Adults & Adolescents

Phase 2/3 study of rivo-cel in intermediate and high risk AML & MDS in patients 12+ years old



## Primary Outcome

Overall Survival

## Secondary Outcomes

- Graft-versus host disease and relapse-free survival (GRFS) at time from randomization until Grade 3-4 acute GvHD, chronic GvHD requiring systemic immunosuppression, disease relapse or death, whichever comes first
- Relapse free survival (RFS)
- Non-relapse mortality (NRM)
- Time to resolution of GvHD after administration of rimiducid


















<sup>A</sup> If dose level 1 exceeds the MTD, alternative dose levels (dose level -1: 1 x 10<sup>6</sup> BPX-501 cells/kg) will be explored

<sup>†</sup> No GvHD prophylaxis will be given. Rimiducid will be administered to inactivate rivoteglecleucel (rivo-cel) in the event of GVHD not responsive to standard of care treatment

Updated 8 Nov 2018. Clinicaltrials.gov identifier: NCT03699475

# Rivo-cel Addresses Key Shortcomings

Rivo-cel addresses shortcomings of stem cell transplants to treat hematological malignancies and inherited blood disorders

|   |                 | Rivo-cel Target Market  |   |   |   |
|---|-----------------|---|---|---|---|
| % of Current Market                       |                 | Matched Related Donor (MRD)<br>25-30%   | Matched Unrelated Donor (MUD)<br>~50%   | Haplo and Other<br>~20-25%  | Rivo-cel +HSCT  |
| Leading Causes of Mortality and Morbidity | Disease Relapse |  |  |  |   |
|   | Infection       |  |  |  |   |
|   | GvHD            |  |  |  |   |
| Likelihood to Find Donor                  |                 | Low   | Low-Medium  | High  |  High  |
| Time to Identify Donor                    |                 | Short   | Long  | Short   |  Short   |

# Rivo-cel: Significant Market Opportunity

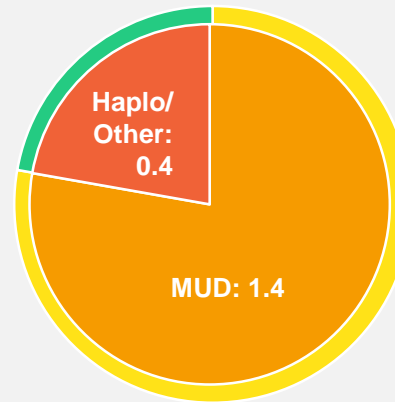
## Potential List Price



## Patient Population

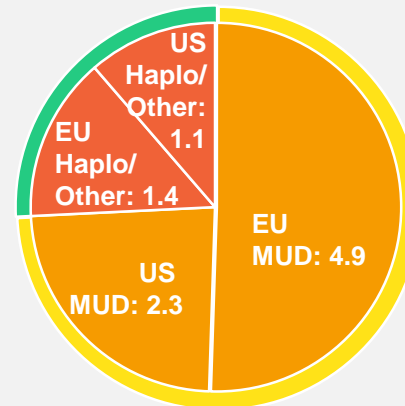
**Initial market:**  
**EU Pediatric\***  
(in thousands)

**TOTAL: 1.8**



**Next market:**  
**Adult & Adolescent AML/MDS\***  
(in thousands)

**TOTAL: 9.7**



## Market Opportunity

**European Pediatric**

**\$0.5-0.7 billion**

**Adult & Adolescent AML/MDS**

**\$3-4 billion**

## Additional Opportunities

- **Geographic expansion**
  - U.S. Pediatric
  - Asia
- **Patient population growth**
- **Expansion of HSCT eligibility**
- **Development in other malignancies**

# Rivo-cel: Significant Opportunity for EU Pediatric Launch

## Demand

- Encouraging early KOL feedback
- Compelling clinical value proposition
- Broad and rapid haplo donor availability

## Pricing & Reimbursement

- Early payer market research encouraging at target pricing
- Outstanding clinical profile and strong health economics

## Manufacturing & Supply Chain

- Robust manufacturing process developed
- Commercially experienced cell therapy CMO
- High touch supply chain & customer service solution being co-developed

## Team

- Efficient: ~75 transplant centers represent ~80% of opportunity
- Experienced: outstanding team in place with relevant track record

# Execution of Key Objectives

# Substantial Progress Achieved in 2018

Delivered on commitments and strengthened the organization

| 2018 To-Do List |   |   |
|-----------------|---|---|
| BPX-601         | Complete enrollment in cell dose escalation portion of BP-012 Phase 1/2 study | ✓ |
|                 | Present initial clinical data at medical meeting                              | ✓ |
| Rivo-cel        | Complete enrollment & present IA on BP-004 and C/CP-004 comparator studies    | ✓ |
|                 | Initiate Phase 2/3 study in adult & adolescent AML & MDS                      | ✓ |
|                 | Confirm pediatric approval pathway in US                                      | ✗ |
|                 | Initiate commercial launch preparation in Europe                              | ✓ |
| BPX-701         | Present initial clinical data at medical meeting                              | ✗ |
| PIPELINE        | Complete dual-switch constructs for two new GoCAR-T candidates                | ✓ |
| ORG             | Complete build-out of Houston cell & viral vector manufacturing facility      | ✓ |
|                 | Establish site in San Francisco Bay Area and European HQ                      | ✓ |
|                 | Strengthen the leadership team  | ✓ |

# Anticipated 2019 and 2020 Key Program Milestones

|                | 1H'19   | 2H'19   | 2020  |
|----------------|---|---|---|
| BPX-601        | <p>Presentations of updated Phase 1 results (Cy/flu regimen)</p> <p>Amend BP-012 to allow for scheduled dosing of rimiducid to reactivate iMC</p> | <p>Presentation of updated Phase 1 results (repeat rimiducid dosing)</p>                | <p>Updated Phase 1 and Phase 2 results</p>              |
| CAR-T PIPELINE | <p>IND submission for BPX-603</p>   | <p>First patient treated in BPX-603 Phase 1 trial</p> <p>IND submission for BPX-802</p> | <p>BPX-603 Phase 1 data</p> <p>BPX-802 Phase 1 data</p> |
| Rivo-cel       | <p>Final analyses of BP-004 and C/CP-004 trials</p>   | <p>MAA submissions for rivo-cel and rimiducid for pediatric patients</p>                | <p>MAA Approval</p> <p>THRIVE Phase 2 interim data</p>  |

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- Expect topline data in 1H 2019; MAA filings in 2H 2019
- European HQ and leadership team in place for commercialization prep

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