# UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

### FORM 8-K

CURRENT REPORT
Pursuant to Section 13 or 15(d) of The Securities Exchange Act of 1934

Date of Report (Date of Earliest Event Reported): January 9, 2017

## bluebird bio, Inc.

(Exact name of registrant as specified in its charter)

	DELAWARE	001-35966	13-3680878
	(State or other jurisdiction of incorporation)	(Commission File Number)	(I.R.S. Employer Identification No.)
	150 Second Street Cambridge, MA		02141
	(Address of principal executive offi	ices)	(Zip Code)
	Registra	nt's telephone number, including area code (339) 49	9-9300
		Not Applicable Former name or former address, if changed since last report)	
Check	11 1	ng is intended to simultaneously satisfy the filing ob	ligation of the registrant under any of the following
	Written communications pursuant to Rule 42	5 under the Securities Act (17 CFR 230.425)	
	Soliciting material pursuant to Rule 14a-12 u	under the Exchange Act (17 CFR 240.14a-12)	
	Pre-commencement communications pursuan	nt to Rule 14d-2(b) under the Exchange Act (17 CFR	240.14d-2(b))
	Dra common coment communications number	et to Dula 12a 4(a) undantha Evahanga Act (17 CED	240.12 - 4(a))

#### Item 2.02 Results of Operations and Financial Condition.

The Company intends to share with investors the amount of cash, cash equivalents and marketable securities it had on hand as of December 31, 2016. Although the Company has not finalized its financial results for the twelve months ended December 31, 2016, the Company currently anticipates that its cash, cash equivalents and marketable securities were approximately \$885 million as of December 31, 2016. This information is unaudited and does not present all information necessary for an understanding of the Company's financial condition as of December 31, 2016 and its results of operations for the twelve months ended December 31, 2016. The Company expects to announce its full results for the twelve months ended December 31, 2016 on or before March 1, 2017.

#### Item 7.01 Regulation FD Disclosure.

The Company will be conducting meetings with investors attending the 35th Annual J.P. Morgan Healthcare Conference in San Francisco beginning on January 9, 2017. As part of these meetings, the Company will deliver the slide presentation furnished to this report as Exhibit 99.1 and which is incorporated herein by reference.

See Item 2.02 above, which is incorporated by reference herein.

The information in this report furnished pursuant to Items 2.02 and 7.01 shall not be deemed "filed" for the purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), or otherwise subject to the liabilities of that section. It may only be incorporated by reference in another filing under the Exchange Act or the Securities Act of 1933, as amended, if such subsequent filing specifically references the information furnished pursuant to Items 2.02 and 7.01 of this report.

#### Item 9.01 Financial Statements and Exhibits.

(d) Exhibits

Exhibit No. Description

99.1 Investor presentation furnished by bluebird bio, Inc. on January 9, 2017.

#### **SIGNATURES**

Pursuant to the requirements of the Securities Exchange	Act of 1934, the registrant has duly caused this report to be signed on its beha	alf by the
undersigned hereunto duly authorized.		

Date: January 9, 2017 bluebird bio, Inc.

By:/s/ Jason F. Cole Jason F. Cole Chief Legal Officer

#### EXHIBIT INDEX

Exhibit No. 99.1

Description
Investor presentation furnished by bluebird bio, Inc. on January 9, 2017.



Corporate Overview

January 2017

Nasdaq : BLUE

## Forward Looking Statements

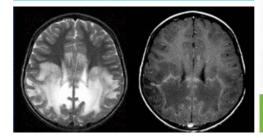
These slides and the accompanying oral presentation contain forward-looking statements and information. The use of words such as "may," "might," "will," "should," "expect," "plan," "anticipate," "believe," "estimate," "project," "intend," "future," "potential," or "continue," and other similar expressions are intended to identify forward-looking statements. For example, all statements we make regarding the initiation, timing, progress and results of our preclinical and clinical studies and our research and development programs, our ability to advance product candidates into, and successfully complete, clinical studies, and the timing or likelihood of regulatory filings and approvals are forward looking. All forward-looking statements are based on estimates and assumptions by our management that, although we believe to be reasonable, are inherently uncertain. All forward-looking statements are subject to risks and uncertainties that may cause actual results to differ materially from those that we expected. These statements are also subject to a number of material risks and uncertainties that are described in our most recent quarterly report on Form 10-Q, as well as our subsequent filings with the Securities and Exchange Commission. Any forward-looking statement speaks only as of the date on which it was made. We undertake no obligation to publicly update or revise any forward-looking statement, whether as a result of new information, future events or otherwise, except as required by law.

**OUR VISION:** 

# Make Hope a Reality



**OUR PATIENTS** 





**BLUE MOJO** 



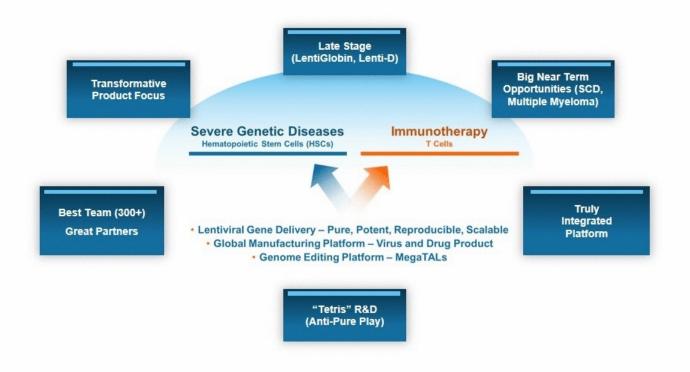
TRUE BLUE



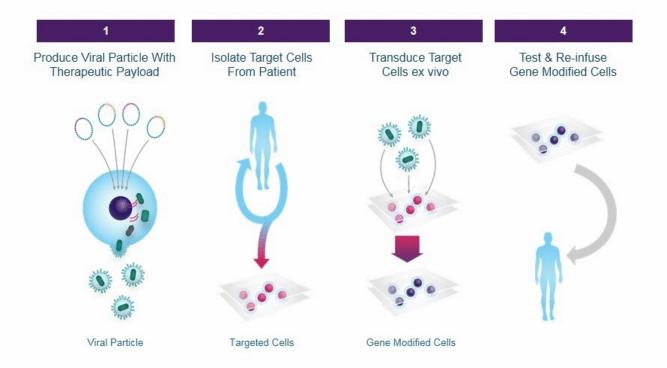
**OUR PEOPLE** 



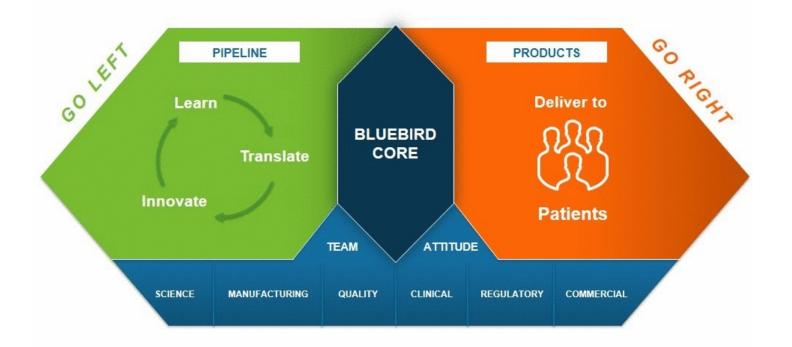
## Our Strategic Intent



## How Our Gene Therapy Approach Works



## Focused on Building Left & Right Around the Core



## bluebird Pipeline Overview



'The current clinical trials for LentiGlobin are Phase 1/2 studies that may provide the basis for early conditional approval in some jurisdictions















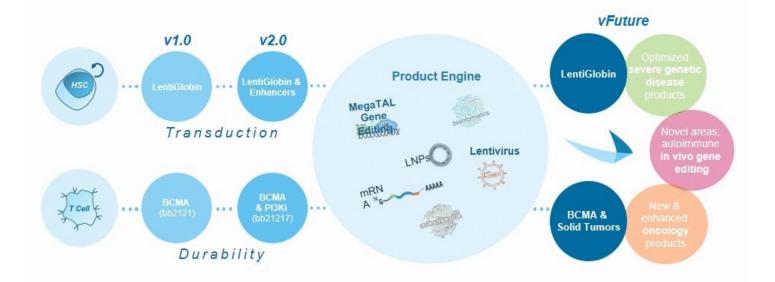




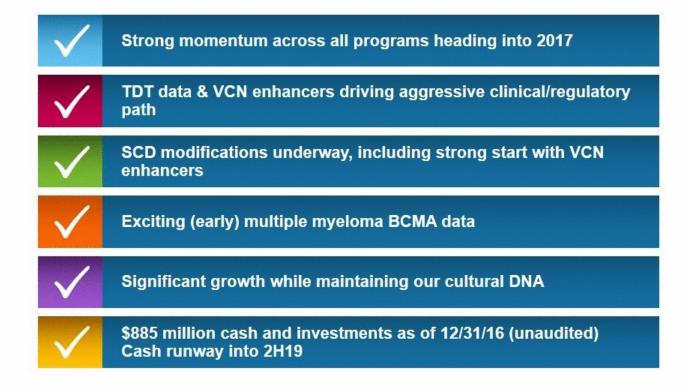




## Good Is Never Good Enough For Patients: BLUE Toolbox Strategy



### Where We Ended 2016



# How Do We Get There? Data, Execution and Development in 2017



## Context for 2017



## **World-class Gene Therapy Platform** and Integrated Global Capabilities



on the Market



**Program Summaries** 

Nasdaq : BLUE

### bb2121 Current Status

- Preliminary results suggest bb2121 demonstrated objective responses in heavily pretreated patients with multiple myeloma
  - Patients in the second cohort achieved stringent complete responses and/or elimination of minimal residual disease
  - 100% ORR (6/6) with doses above 5x107 CAR+ T cells
- bb2121 has been well tolerated, with mild-to-moderate cytokine release syndrome reported as of November data cut-off
  - No dose-limiting toxicities yet identified and dose escalation continues
- Dose escalation and expansion will continue to identify recommended phase 2 dose

bb2121 anti-BCMA CAR T therapy may offer a promising new treatment paradigm for patients suffering from multiple myeloma

Data as of Nov 18, 2016

### LentiGlobin Current Status: TDT\*

- LentiGlobin treatment shows promising results in TDT
  - Patients with non-β<sup>0</sup>/β<sup>0</sup> genotypes and ≥12 months follow-up remain free of RBC transfusions
  - Clinically meaningful reductions in transfusion volume and frequency in patients with  $\beta^0/\beta^0$  genotypes
- Toxicity profile remains consistent with single-agent busulfan conditioning, with no evidence of clonal dominance
- LentiGlobin VCN correlated with HbAT87Q level
- LentiGlobin manufacturing process using transduction enhancement for ongoing and planned clinical studies
  - Goal to increase drug product VCN and total hemoglobin production in all patients, regardless of genotype
- Pivotal HGB-207 study launched

\*As of September 2016 data cut-off

### LentiGlobin Current Status: Severe SCD

- Results in HGB-205 subject 1204 demonstrate promise of LentiGlobin autologous gene therapy for severe SCD
  - ~50% anti-sickling hemoglobin with sustained absence of severe sickle cell disease-related symptoms
- Initial findings from HGB-206 confirm feasibility of autologous HSC gene therapy in severe SCD
  - Successful bone marrow harvests and centralized drug product manufacturing
  - Safety profile consistent with procedural requirements
  - No gene therapy-related AEs
  - HbA<sup>T87Q</sup> production in all treated patients
- Changes implemented in protocol and manufacturing with goal of achieving higher levels of anti-sickling hemoglobin to optimize clinical benefit

Data as of Sept 9, 2016 [HGB-205] and Nov 9, 2016 [HGB-206]

### Lenti-D Current Status

- Initial interim Starbeam results suggest early treatment with Lenti-D gene therapy may halt neuro-inflammation and demyelination in most CALD patients, with promising safety
  - All subjects were free of Major Functional Disabilities (MFD) as of March 31, 2016 data cut-off
  - Stabilization of NFS achieved in 94% (16/17) and Loes score achieved in 82% (14/17)
  - Resolution of gadolinium enhancement by month 6 in 94% (16/17)
  - Re-appearance of diffuse gadolinium enhancement in 5 subjects, resolved in those (n=2) who have later follow-up
- No deaths, graft failure, or GvHD reported as of March 31, 2016
- AE profile consistent with myeloablative conditioning with bulsulfan and cyclophosphamide
- Lenti-D gene therapy may offer an alternative to allogeneic bone marrow transplant, particularly for patients with no matched sibling donor
  - Additional follow-up is needed to fully assess efficacy, durability of effect and long-term safety
  - Eight additional patients to be enrolled: same enrollment criteria
    - · Gain experience manufacturing and delivering Lenti-D in Europe
    - · Bolster data package for US and EU regulatory filings

Data presented at AAN 2016. Data as of March 31, 2016



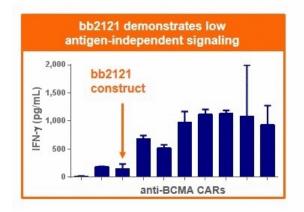
Multiple Myeloma

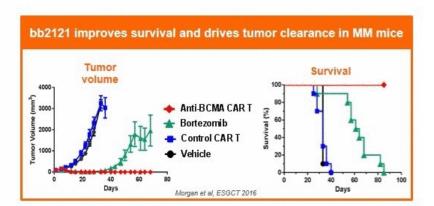
Nasdaq : BLUE

# bb2121: Anti-BCMA Chimeric Antigen Receptor T Cell Product Candidate



- Autologous T cells transduced with a lentiviral vector encoding a novel anti-BCMA CAR
- 4-1BB co-signaling motif selected to promote proliferation and persistence
- Construct demonstrated potent preclinical in vivo activity with low tonic signaling





## CRB-401 Phase 1 Study in Relapsed / Refractory Multiple Myeloma

#### CRB-401 Open-label Phase 1 Clinical Study of bb2121

- Objectives: Determine preliminary safety and efficacy and recommended phase 2 dose
- N = 50 patients, standard 3+3 dose escalation + expansion cohort
- Eligibility
  - Relapsed / refractory MM with ≥ 3 prior lines of therapy (including PI and IMiD), or double refractory
  - Measurable disease
  - ≥ 50% BCMA expression
  - Adequate bone marrow, renal and hepatic function

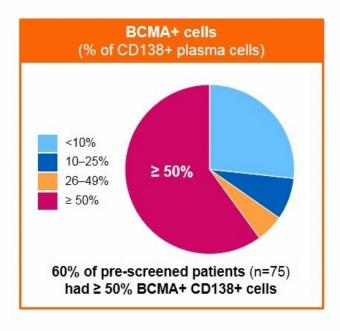
#### 3 + 3 Dose Escalation of CAR + T Cells

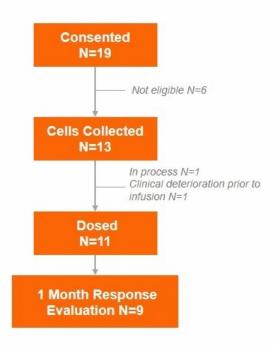


Up to 5 dose cohorts planned, fixed dose of CAR + T Cells

9 U.S. Clinical Sites, 1 Centralized Manufacturing Site

## Study Status as of November 18, 2016





Data as of Nov 18, 2016

## Demographics and Disease History in Treated Patients

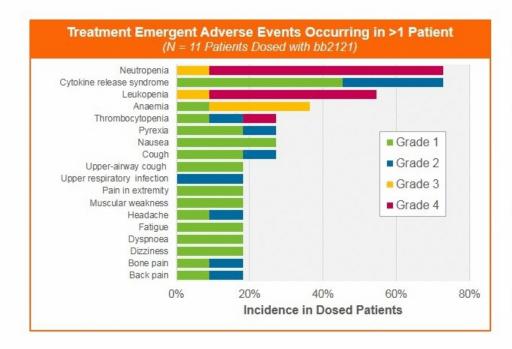
Parameter	Statistic	N=11 Dosed Patient
Age years	Median (range)	58 (41-74)
Male gender	N (%)	7 (64%)
Time since diagnosis years	Median (range)	5 (1-9)
ECOG1 = 0	N (%)	6 (55%)
ISS <sup>2</sup> Stage I II III	N (%)	5 (45%) 4 (36%) 2 (18%)
High-risk cytogenetics (del17p, t(4;14), t(14;16), 1q, del 13)	N (%)	5 (45%)

Parameter	Statistic	N=11 Dosed Patients
Prior lines of therapy	Median (range)	6 (5-13)
Prior autologous SCT	N (%)	11 (100%)
Prior therapies	N (%)	
IMiD lenalidomide pomalidomide		<b>11 (100%)</b> 11 (100%) 9 (82%)
proteasome inhibitor bortezomib carfilzomib		<b>11 (100%)</b> 11 (100%) 9 (82%)
daratumumab / CD38 antibody		7 (64%)

Data as of Nov 18 2016

1. Eastern Cooperative Oncology Group Performance Score. 2. International Staging System

# Adverse Events Generally Mild, No ≥ Grade 3 CRS\* or Neurotoxicity

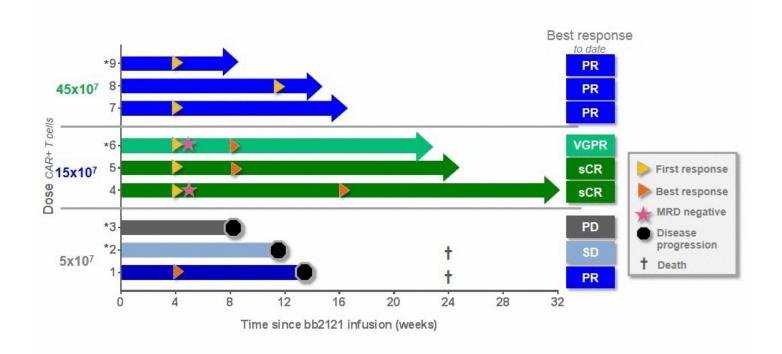


- No dose-limiting toxicities as of data cut-off
- Cytopenias related to fludarabine/ cyclophosphamide lymphodepletion, as expected
- No ≥ Grade 3 cytokine release syndrome or neurotoxicity as of data cut-off

\*CRS uniformly graded according to Lee et al., Blood 2014;124:188-195

Data as of Nov 18, 2016

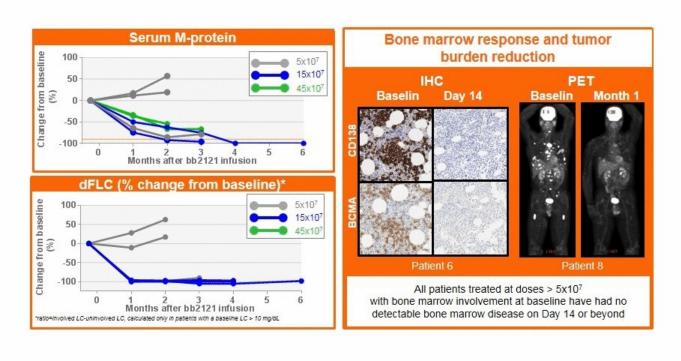
## Best Response and Time Since bb2121 Infusion



Data as of Nov 18, 2016

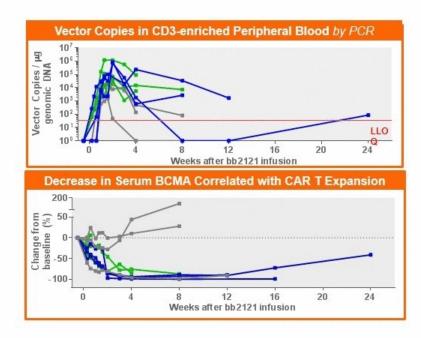
\* Patient with ≥50% bone marrow involvement

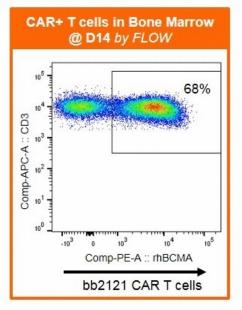
## Responses to bb2121 Infusion



Data as of Nov 18, 2016

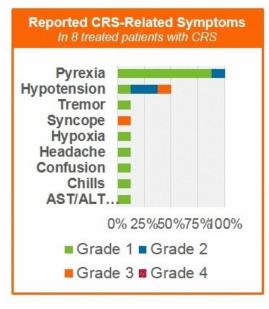
## CAR T Cell Expansion at Every Dose



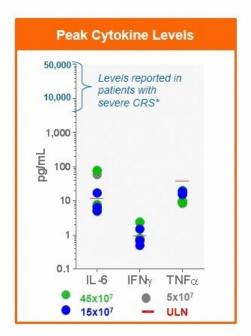


Data as of Nov 18, 2016

## Cytokine Release Syndrome Summary



- 8/11 (73%) with cytokine release syndrome (CRS)
  - CRS severity Grades 1 & 2
  - Including patients in all dose groups and those with ≥50% bone marrow involvement
- CRS-related symptoms mostly Grade 1
- No patients received tocilizumab or steroids



Data as of Nov 18, 2016

<sup>\*</sup> In anti-BCMA and anti-CD19 CAR T studies. Ali et al., Blood 2016 128: 1688. Maude et al., NEJM 2014



Transfusion Dependent β-Thalassemia

Nasdaq : BLUE

## TDT Studies: Status

#### HGB-204 multicenter study of LentiGlobin in TDT

**CURRENT STATUS** 

All 18 patients treated, with ≥ 6 months follow-up

2 patients have completed 2-year analysis

#### HGB-205 single center study of LentiGlobin in TDT and severe SCD

**CURRENT STATUS** 

4 TDT patients treated, with 11 – 33 months follow-up

Data as of Sept 16, 2016

## Patient and Drug Product Characteristics HGB-204

N=18 treated patients

	Genotype	
	$\beta^0/\beta^0$ (n=8)	Non-β <sup>0</sup> /β <sup>0</sup> (n=10)
Genotype	8	10
$\beta^{E}/\beta^{0}$	-	6
Other $(\beta^+/\beta^0, \beta^+/\beta^+, \beta^x/\beta^0)$		4
Age at start of regular transfusions Age at consent Median (range) years	0 (0 – 7) 23 (12 - 35)	6 (0 – 26) 19.5 (16 – 34)
Median (range) pre-study pRBC transfusion vol annualized median (range) mL/kg/year	184.9 (128.7 - 261.3)	146.3 (117.0 – 234.5)
Splenectomy	3	3
Drug Product Parameters	Media	n (range)
Drug product VCN <sup>1</sup>	0.7 (range 0.3 - 1.5)	0.8 (range 0.3 - 1.1)
Drug product cell dose CD34+ cells x10 <sup>6</sup> /kg	11.0 (range 6.1-18.1)	7.1 (range 5.2-13.0)

Data as of Sept 16, 2016 1. VCN: vector copy number (vector copies per diploid genome)

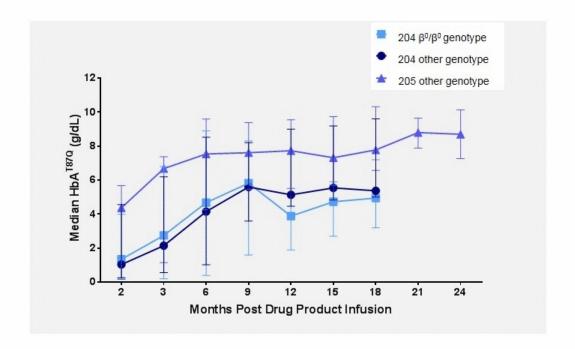
## Patient and Drug Product Characteristics HGB-205

	1201	1202	1203	1206
Age at Enrollment (years)	18	16	19	17
Genotype	β□/βΕ	β0/β <b>E</b>	homozygous IVS1 nt 110 G>A	β⁰/βΕ
Pre-Treatment pRBC Transfusions (mL/kg/year)¹	139	188	176	197
VCN in Drug Product <sup>2</sup>	1.5	2.1	0.8	1.1
CD34+ Cell Dose (x10e/kg)	8.9	13.6	8.8	12.0
Busulfan AUC (average, uM/min)	4,967	5,212	4,670	4,930
Follow-up (months)	33.5	30.3	14.6	11.6

<sup>1</sup>mean pRBC requirement per year, over the past 2 years prior to consent; <sup>2</sup>VCN = number of vector copies per diploid genome

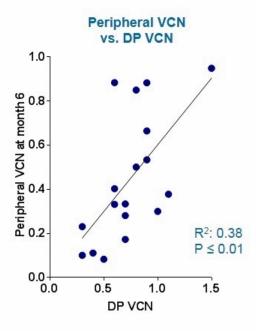
Data as of Sept 16, 2016

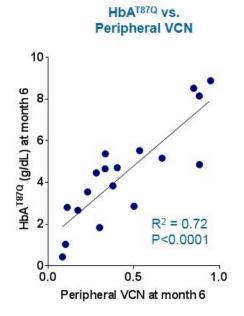
# HbA<sup>T87Q</sup> production increases to month 9, then stabilizes



Data as of Sept 9, 2016 [HGB-205], Sept 16, 2016 [HGB-204]

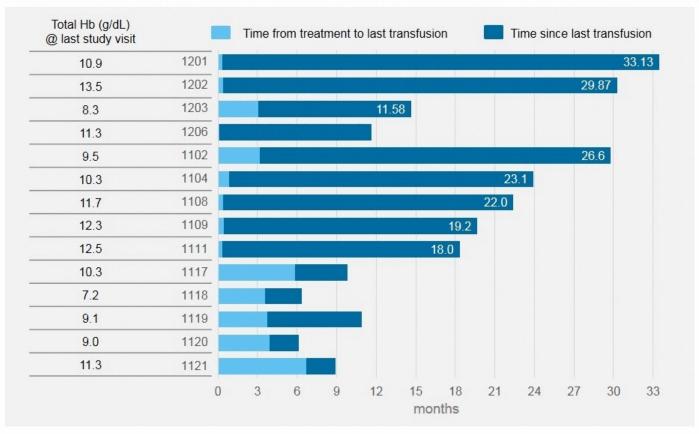
# Peripheral VCN correlates with DP VCN and HbA<sup>T87Q</sup> level at Month 6





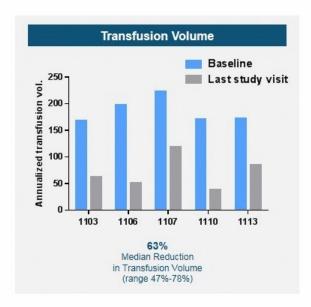
Data as of Sept 16, 2016

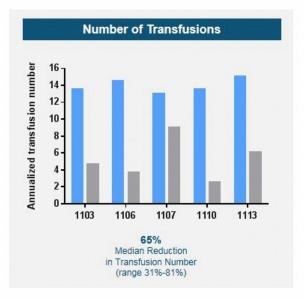
# Patients with non-β<sup>0</sup>/β<sup>0</sup> genotypes and ≥12 months follow-up have stopped RBC transfusions



Data as of Sept 9, 2016 [HGB-205], Sept 16, 2016 [HGB-204]

# Transfusion reduction seen in patients with β<sup>0</sup>/β<sup>0</sup> genotypes with ≥12 months follow-up



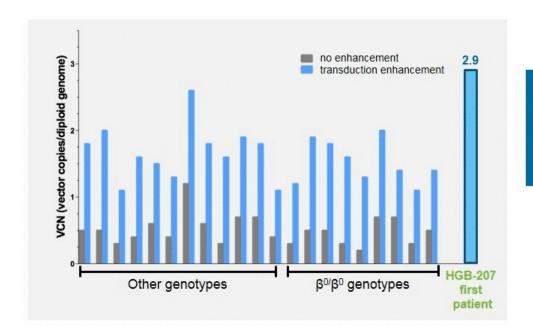


Post-treatment: annualized on-study volume and number of transfusions based on observed values starting at month 6 through data cut-off

Median follow-up for patients with  $\beta^0/\beta^0$  genotypes (N=8) 17.3 months (range 6.7-25.4)

Data as of Sept 16, 2016

# Research-scale Results Demonstrate Increase in Drug Product VCN Across Genotypes



Percent of cells transduced: 77%

Data as of Nov 30, 2016 Exploratory in vitro analysis conducted at research scale

## Next steps: Pivotal clinical studies of LentiGlobin therapy in TDT

## N\*RTHSTAR-2

HGB-207 Non-β<sup>0</sup>/β<sup>0</sup> genotypes

Phase 3, multi-center, global study

- N=15 adults and adolescents, and N=8 pediatric patients
- · Open and enrolling

## N\*RTHSTAR-3

HGB-212 β<sup>0</sup>/β<sup>0</sup> genotypes

Phase 3, multi-center, global study

- N=15 adults, adolescents and pediatric patients
- Initiation planned for 2017

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## U.S. Registration Strategy

General agreement with U.S. regulators on the registration path for LentiGlobin BB305 for the treatment of transfusion-dependent β-thalassemia



Pursue approval in adults and adolescents based on data from ongoing pivotal HGB-207 trial

Pediatric
population to be
included as a
cohort of HGB207, rather than
separate study

Submission for approval in  $\beta^{0}/\beta^0$  patients to be based on planned HGB-212 study



BREAKTHROUGH THERAPY DESIGNATION

## **EU Registration Strategy**

General agreement with EU regulators on the registration path for LentiGlobin BB305 for the treatment of transfusion-dependent β-thalassemia





### ADAPTIVE PATHWAYS



### PRIME

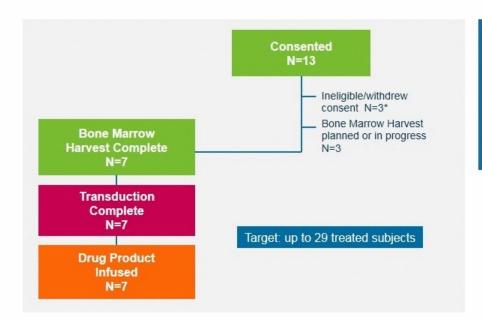
Plan to include available data from HGB-207 and HGB-212 studies at the time of filing in the marketing authorization application



Sickle Cell Disease

Nasdaq : BLUE

## Current Status of HGB-206 Study



### **Enrollment criteria**

- 18+ years of age
- History of symptomatic SCD
- Adequate organ function/performance status
- · No previous HSCT or gene therapy

Data as of Nov 9, 2016
\* 2 screen failures (bilirubin levels/fertility concerns), 1 withdrew consent

## **Patient and Treatment Characteristics**

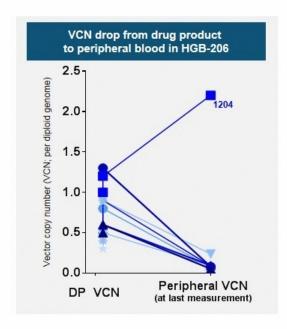
# All treated patients (n=8) have a **history of severe SCD** in 2 years prior to enrollment, despite hydroxyurea therapy

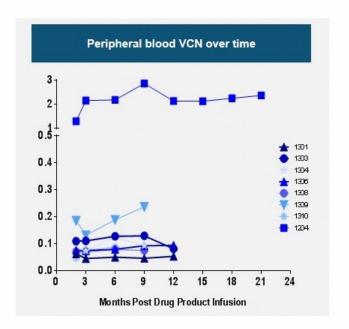
SCD History (n=8)					
Recurrent VOCs	Stroke	Acute Chest Syndrome	Regular pRBC Transfusions		
6	2	6	2		

Treatment Characteristics						
Parameter	<b>HGB-206</b> (n=7) Median (range)	<b>HGB-205</b> (n=1)				
Age at Enrollment (years)	26 (18 – 42)	13				
Bone Marrow Harvests	2 (1 – 4)	2				
Target daily busulfan AUC (µM/min)	5000 (4400 – 5400)	4841 (actual)				
LentiGlobin DP cell dose (CD34+x10 <sup>6</sup> cells/kg)	2.1 (1.6 – 5.1)	5.6				
LentiGlobin DP vector copy number (VCN)	0.6 (0.3 – 1.3)	1.0, 1.2				

Data as of Sept 9, 2016 [HGB-205] and Nov 9, 2016 [HGB-206]

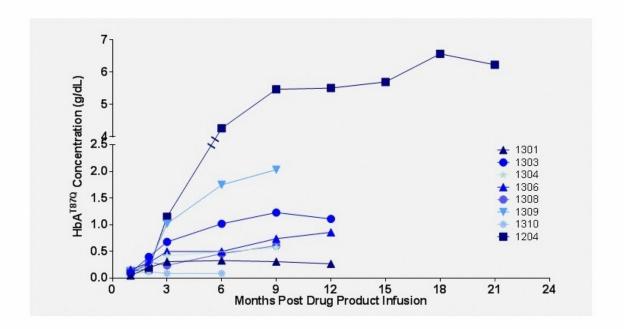
## Vector Copy Number (VCN) in Drug Product and Peripheral Blood





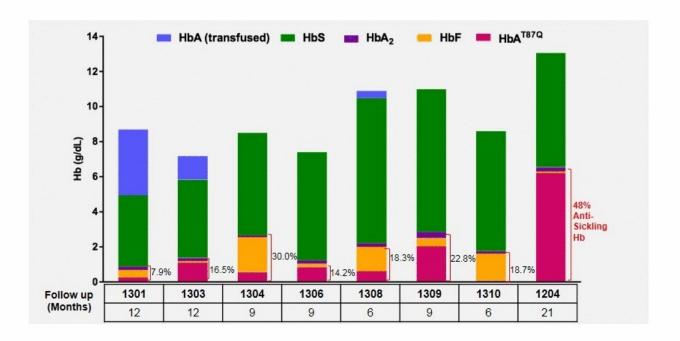
Data as of Sept 9, 2016 [HGB-205] and Nov 9, 2016 [HGB-206]

## All Treated Patients Produce Measurable HbAT87Q



Data as of Sept 9, 2016 [HGB-205] and Nov 9, 2016 [HGB-206]

## 8% to 48% Anti-Sickling Hemoglobin at Last Follow Up



Data as of Sept 9, 2016 [HGB-205] and Nov 9, 2016 [HGB-206]

### Clinical Outcomes 21 Months After Treatment in HGB-205 Patient 1204

Pre-Treatment 21 Months After Treatment Transfusions Weaned off transfusions Chronic transfusions Last transfusion on Day + 88 (> 18 months ago) Multiple hospitalizations before No hospitalizations or starting transfusion regimen acute SCD-related events Hemolysis Baseline while on transfusions · Reticulocytes 132 x 109/L Reticulocytes 238.3 x 10<sup>9</sup>/L LDH 287 U/L • Bilirubin 11 µmol/L

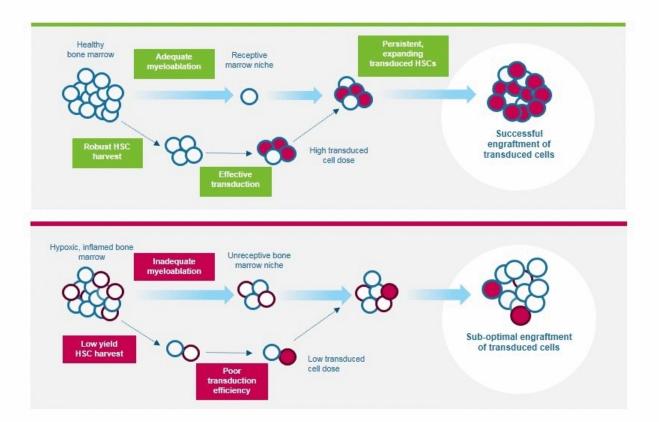
Data as of Nov 9, 2016 [HGB-206]

## Interim Summary – Where are we today?

- Results in HGB-205 subject 1204 demonstrate promise of LentiGlobin autologous gene therapy for severe SCD
  - ~50% anti-sickling hemoglobin with sustained absence of severe sickle cell disease-related symptoms
- Initial findings from HGB-206 confirm feasibility of autologous HSC gene therapy in severe SCD
  - Successful bone marrow harvests and centralized drug product manufacturing
  - Safety profile consistent with procedural requirements
  - No gene therapy-related AEs
  - HbAT87Q production in all treated patients
- Challenges remain to achieve target level of anti-sickling hemoglobin in all patients
- Higher levels of anti-sickling hemoglobin are needed to optimize clinical benefit

Data as of Sept 9, 2016 [HGB-205] and Nov 9, 2016 [HGB-206]

# Stem Cell Transduction and Engraftment – The Challenge of Sickle Cell Disease



# Protocol and Process Changes to Potentially Improve Outcomes in SCD Patients

Hypoxic, inflamed marrow

Pre-harvest transfusions to reduce marrow inflammation, hypoxia

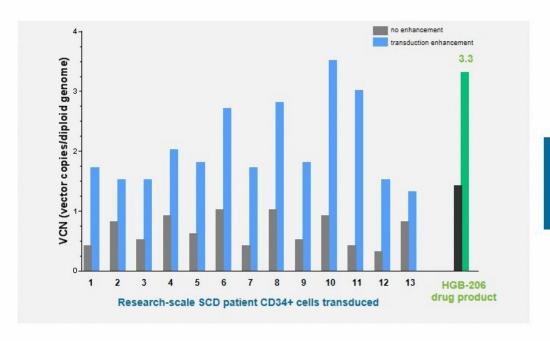
Additional changes to manufacturing process to increase cell dose

VCN Enhancers (Process 2)

Inadequate myeloablation

Increased exposure to myeloablative agent

## LentiGlobin Manufacturing Process with Transduction Enhancers Increases DP VCN in SCD CD34+ Cells



Percent of cells transduced: 83%

Data as of Nov 30, 2016



Closing

Nasdaq : BLUE

## The 2017 Plan is Clear and Catalyst Rich

### **β-THALASSEMIA**

Ongoing Data from HGB 204 & 205

Data from HGB-207 & Initiate HGB-212

VCN Enhancers Impact & Filing Strategy

### SICKLE CELL DISEASE

HGB-205

HGB-206

Protocol & VCN Enhancers Impact

### CALD

Starbeam Study Expansion

Allo Transplant Study

Confirm Regulatory Strategy

## MULTIPLE

Ongoing Data from CRB-401

Next Gen bb21217 IND

Study Expansion & Clin/Reg Path Forward

#### **PIPELINE**

Platform Improvements

> Innovative Science

Deliver New Programs to the Clinic

\$885 million cash and investments as of 12/31/16 (unaudited)

Cash runway into 2H19

## 2022 Vision – The Gene Therapy Product Platform



## **∞** | Patient Impact

2+ Products on the Market

2+ Programs Nearing Commercialization

4+ Additional Programs in the Clinic

## **Bringing & Valuing Hope**

Go TRUE BLUE





## Closing

Nick Leschly, chief bluebird

Nasdaq : BLUE