

**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 11, 2016**

**AMICUS THERAPEUTICS, INC.**  
(Exact Name of Registrant as Specified in Its Charter)

**Delaware**  
(State or Other Jurisdiction of  
Incorporation)

**001-33497**  
(Commission File Number)

**71-0869350**  
(IRS Employer Identification No.)

**1 Cedar Brook Drive, Cranbury, NJ**  
(Address of Principal Executive Offices)

**08512**  
(Zip Code)

Registrant's telephone number, including area code: **(609) 662-2000**

(Former Name or Former Address, if Changed Since Last Report.)

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:

- Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)
- Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)
- Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))
- Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

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

On January 11, 2016, Amicus Therapeutics, Inc. (the "**Company**") issued a press release (the "**Press Release**") regarding its financial condition for the year ended December 31, 2015. A copy of the Press Release is furnished as Exhibit 99.1 to this Current Report on Form 8-K and is incorporated herein by reference.

**Item 7.01. Regulation FD Disclosure**

*Press Release*

The Press Release also includes information regarding the Company's 2015 accomplishments and its strategic outlook and financial guidance for the year ending December 31, 2016.

A copy of the Press Release is furnished as Exhibit 99.1 to this Current Report on Form 8-K and is incorporated herein by reference.

*Corporate Presentations*

The Company has updated its corporate presentation as of January 11, 2016. The slides from this presentation are attached hereto as Exhibit 99.2. The attached materials will be posted on the Company's website at [www.amicusrx.com](http://www.amicusrx.com). The Company does not undertake to update this presentation.

The Company presented a corporate presentation at the 3<sup>rd</sup> Annual Dermatology Summit: SD-101 for Epidermolysis Bullosa on Sunday, January 10, 2016. The slides from this presentation are attached hereto as Exhibit 99.3. The attached materials will be posted on the Company's website at [www.amicusrx.com](http://www.amicusrx.com). The Company does not undertake to update this presentation.

**Item 9.01. Financial Statements and Exhibits.**

(d) Exhibits:

<u>Exhibit No.</u>	<u>Description</u>
99.1	Press Release dated January 11, 2016
99.2	Corporate Presentation
99.3	Corporate Presentation

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### SIGNATURES

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

Amicus Therapeutics, Inc.

Date: January 11, 2016

By: /s/ William D. Baird III  
William D. Baird III  
Chief Financial Officer

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### EXHIBIT INDEX

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99.1	Press Release dated January 11, 2016
99.2	Corporate Presentation
99.3	Corporate Presentation

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## Amicus Therapeutics Provides Full-Year 2016 Strategic Outlook and Financial Guidance

### *Migalastat for Fabry Disease Moving Toward CHMP Opinion in European Union*

### *Pompe Disease Clinical Study Initiated to Investigate Novel Enzyme Replacement Therapy*

### *Epidermolysis Bullosa (EB) Phase 3 Study Advances*

**CRANBURY, NJ, January 11, 2016** — Amicus Therapeutics (Nasdaq: FOLD), a biopharmaceutical company at the forefront of therapies for rare and orphan diseases, today provided its full-year 2016 strategic outlook and financial guidance.

John F. Crowley, Chairman and Chief Executive Officer of Amicus Therapeutics, Inc., stated, “During 2015 we established a global infrastructure with a leading commercial team, acquired a Phase 3 rare disease program in EB, submitted our first marketing application in Fabry in the EU, and advanced our first biologic medicine into the clinic in Pompe. These achievements have transformed Amicus and strongly position us to build one of the world’s leading biotechnology companies focused on rare and devastating diseases. In 2016 we will continue to focus on advancing our three lead programs in Fabry, Pompe and EB. The vision for Amicus continues to be to advance our pipeline and to acquire capabilities, programs and technologies that deliver on our mission for patients and that create substantial shareholder value.”

#### Key 2015 Accomplishments

- **MAA Submission** - submitted marketing authorization application (MAA) to European Medicines Agency (EMA) for approval of migalastat in Fabry patients with amenable genetic mutations
- **Global Commercial Capabilities** - established global commercial infrastructure with key leadership in place for EU migalastat launch
- **Late-Stage Product Acquisition** - strengthened pipeline with the acquisition of SD-101, a Phase 3 program for Epidermolysis Bullosa (EB)
- **Biologics Manufacturing** - completed successful manufacturing scale up to supply clinical study of ATB200 — a novel ERT for Pompe Disease
- **Pompe Clinical Study Initiation** - initiated clinical study of ATB200 and oral chaperone (AT2221) in Pompe patients

Mr. Crowley will discuss Amicus’ corporate objectives and key milestones in a presentation at the 34th Annual J.P. Morgan Healthcare Conference on Tuesday, January 12, 2016 at 9:00 a.m. PT (12:00 p.m. ET). A live webcast of the presentation can be accessed through the Investors section of the Amicus Therapeutics corporate web site at <http://ir.amicustherapeutics.com/events.cfm>, and will be archived for 90 days.

#### Program Highlights

##### **Migalastat for Fabry Disease**

Migalastat is an oral personalized medicine intended to treat Fabry disease in patients who have amenable genetic mutations. Amicus has built a commercial organization in the EU that is prepared to launch migalastat upon approval. The EMA’s review of the MAA for migalastat is currently in progress. Amicus continues to expect an opinion from the Committee for Medicinal Products for Human Use (CHMP) in early 2016. As previously reported, the timing of a new drug

application (NDA) submission in the U.S. will be based on the determination of the optimal regulatory pathway. The Company expects to provide an update on the U.S. strategy for migalastat in the first quarter of 2016.

##### Anticipated 2016 Fabry Program Milestones

- CHMP Opinion in Europe (early 2016)
- Oral presentations and posters at the 12th Annual *WORLDsymposium™* 2016 (February 29 - March 4, 2016 in San Diego, CA). Oral presentations will include:
  - Podocyte globotriaosylceramide (GL-3) content in male adult patients with Fabry disease reduces following 6-12 months of treatment with migalastat
  - Comparison of integrated white blood cell alpha-galactosidase A activity exposure between every-other-day orally administered migalastat and biweekly infusions of agalsidase beta or agalsidase alfa
- U.S. regulatory update (1Q16)
- Phase 3 data publications

##### **Novel ERT for Pompe Disease (ATB200 + Chaperone)**

Amicus has initiated the clinical study (ATB200-02) in Pompe patients with a novel treatment paradigm that consists of ATB200, a uniquely engineered recombinant human acid alpha-glucosidase (rhGAA) enzyme with an optimized carbohydrate structure to enhance uptake, administered with a pharmacological chaperone (AT2221) to improve activity and stability.

Amicus completed good manufacturing practice (GMP) production runs of ATB200 during 2015, and has sufficient supply for this clinical study in Pompe patients. Amicus finalized the design of the ongoing study following in-person meetings with regulatory authorities in both the U.S. and EU.

##### Anticipated Pompe Program Milestones

- Dosing of first patient in clinical study (early 2016)
- Oral presentations and posters at *WORLDSymposium™*. Oral presentation will include:
  - Co-administration of the pharmacological chaperone AT2221 with a proprietary recombinant human acid alfa-glucosidase leads to greater plasma exposure and substrate reduction compared to alglucosidase alfa
- Data from clinical study ATB200-02

## **SD-101 for Epidermolysis Bullosa (EB)**

SD-101 is a novel, late-stage, proprietary topical treatment and potential first-to-market therapy for EB. This investigational product was granted FDA breakthrough therapy designation in 2013 based on results from a Phase 2a study for the treatment of lesions in patients suffering with EB. SD-101 is the first-ever treatment in EB clinical studies to show improvements in wound closure across all major EB subtypes.

SD-101 is currently being investigated in a Phase 3 study (SD-005) to support global regulatory submissions. Amicus continues to open additional clinical sites for this Phase 3 study and expects enrollment to be complete by mid-year with data in the second half of 2016. Amicus Chief Business Officer Dipal Doshi has assumed the additional responsibilities as General Manager of the Scioderm Division of Amicus.

### **Anticipated EB Program Milestones**

- Completion of enrollment in Phase 3 study (mid-2016)
- Top-line Phase 3 data (2H16)

## **2016 Financial Guidance**

Cash, cash equivalents, and marketable securities totaled \$214.0 million at December 31, 2015 compared to \$169.1 million at December 31, 2014. The Company's balance sheet was strengthened during 2015 with a \$258 million public offering. Amicus expects full-year 2016 net cash spend between \$135 million and \$155 million. The current cash position is projected to fund operations into mid-2017.

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## **About Amicus Therapeutics**

Amicus Therapeutics (Nasdaq: FOLD) is a biotechnology company at the forefront of therapies for rare and orphan diseases. The Company has a robust pipeline of advanced therapies for a broad range of human genetic diseases. Amicus' lead programs in development include the small molecule pharmacological chaperone migalastat as a monotherapy for Fabry disease, SD-101 for Epidermolysis Bullosa (EB), as well as novel enzyme replacement therapy (ERT) products for Fabry disease, Pompe disease, and other lysosomal storage disorders.

## **Forward-Looking Statements**

This press release contains "forward-looking statements" within the meaning of the Private Securities Litigation Reform Act of 1995 relating to preclinical and clinical development of Amicus' candidate drug products, the timing and reporting of results from preclinical studies and clinical trials evaluating Amicus' candidate drug products and the projected cash position for the Company. Words such as, but not limited to, "look forward to," "believe," "expect," "anticipate," "estimate," "intend," "potential," "plan," "targets," "likely," "may," "will," "would," "should" and "could," and similar expressions or words identify forward-looking statements. Such forward-looking statements are based upon current expectations that involve risks, changes in circumstances, assumptions and uncertainties. The inclusion of forward-looking statements should not be regarded as a representation by Amicus that any of its plans will be achieved. Any or all of the forward-looking statements in this press release may turn out to be wrong. They can be affected by inaccurate assumptions Amicus might make or by known or unknown risks and uncertainties. For example, with respect to statements regarding the goals, progress, timing and outcomes of discussions with regulatory authorities and the potential goals, progress, timing and results of preclinical studies and clinical trials, actual results may differ materially from those set forth in this release due to the risks and uncertainties inherent in the business of Amicus, including, without limitation: the potential that results of clinical or pre-clinical studies indicate that the product candidates are unsafe or ineffective; the potential that it may be difficult to enroll patients in our clinical trials; the potential that regulatory authorities may not grant or may delay approval for our product candidates; the potential that preclinical and clinical studies could be delayed because we identify serious side effects or other safety issues; the potential that we will need additional funding to complete all of our studies and, our dependence on third parties in the conduct of our clinical studies. Further, the results of earlier preclinical studies and/or clinical trials may not be predictive of future results. With respect to statements regarding projections of the Company's cash position, actual results may differ based on market factors and the Company's ability to execute its operational and budget plans. In addition, all forward looking statements are subject to other risks detailed in our Annual Report on Form 10-K for the year ended December 31, 2014 and Form 10-Q for the quarter ended June 30, 2015. You are cautioned not to place undue reliance on these forward-looking statements, which speak only as of the date hereof. All forward-looking statements are qualified in their entirety by this cautionary statement, and Amicus undertakes no obligation to revise or update this news release to reflect events or circumstances after the date hereof. This caution is made under the safe harbor provisions of Section 21E of the Private Securities Litigation Reform Act of 1995.

## **CONTACTS:**

### **Investors/Media:**

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(973) 271-6085





34<sup>th</sup> Annual J.P.  
Morgan  
Healthcare  
Conference

John F. Crowley, Chairman and Chief Executive Officer  
January 12, 2016

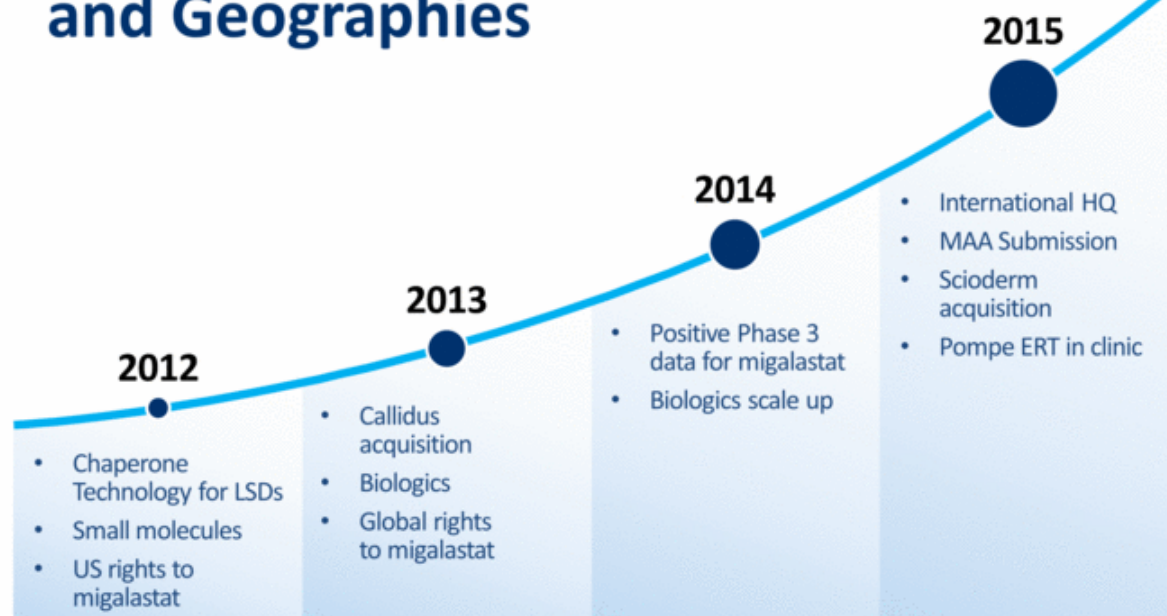
## Safe Harbor

*This presentation will contain, "forward-looking statements" within the meaning of the Private Securities Litigation Reform Act of 1995 relating to preclinical and clinical development of Amicus' candidate drug products, the timing and reporting of results from preclinical studies and clinical trials evaluating Amicus' candidate drug products, financing plans, and the projected cash position for the Company. Words such as, but not limited to, "look forward to," "believe," "expect," "anticipate," "estimate," "intend," "potential," "plan," "targets," "likely," "may," "will," "would," "should" and "could," and similar expressions or words identify forward-looking statements. Such forward-looking statements are based upon current expectations that involve risks, changes in circumstances, assumptions and uncertainties. The inclusion of forward-looking statements should not be regarded as a representation by Amicus that any of its plans will be achieved. Any or all of the forward-looking statements in this press release may turn out to be wrong. They can be affected by inaccurate assumptions Amicus might make or by known or unknown risks and uncertainties. For example, with respect to statements regarding the goals, progress, timing and outcomes of discussions with regulatory authorities, and in particular the timing of an NDA submission for migalastat monotherapy, and the potential goals, progress, timing and results of preclinical studies and clinical trials, actual results may differ materially from those set forth in this release due to the risks and uncertainties inherent in the business of Amicus, including, without limitation: the potential that results of clinical or pre-clinical studies indicate that the product candidates are unsafe or ineffective; the potential that it may be difficult to enroll patients in our clinical trials; the potential that regulatory authorities may not grant or may delay approval for our product candidates; the potential that preclinical and clinical studies could be delayed because we identify serious side effects or other safety issues; the potential that we will need additional funding to complete all of our studies and, our dependence on third parties in the conduct of our clinical studies. Further, the results of earlier preclinical studies and/or clinical trials may not be predictive of future results. With respect to statements regarding projections of the Company's cash position, actual results may differ based on market factors and the Company's ability to execute its operational and budget plans. In addition, all forward looking statements are subject to other risks detailed in our Annual Report on Form 10-K for the year ended December 31, 2014 and Form 10-Q for the quarter ended June 30, 2015. You are cautioned not to place undue reliance on these forward-looking statements, which speak only as of the date hereof. All forward-looking statements are qualified in their entirety by this cautionary statement, and Amicus undertakes no obligation to revise or update this news release to reflect events or circumstances after the date hereof. This caution is made under the safe harbor provisions of Section 21E of the Private Securities Litigation Reform Act of 1995.*

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## Amicus 2016 – Looking Back

# Amicus Has Greatly Expanded Product Pipeline, Technologies and Geographies





# Amicus 2016 – Continuing the Momentum

## Significant Milestones in 2016



# Amicus Vision

Amicus Therapeutics is a global biotechnology company at the forefront of developing advanced therapies to treat a range of devastating rare and orphan diseases

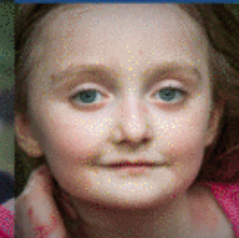
Rare &  
Devastating  
Diseases



Potential  
First-in-Class  
/ Best-in-  
Class



Meaningful  
benefits for  
patients



## Key Drivers of Value

3 Novel Product Candidates Each with \$500M to \$1B+ Market Potential

### Fabry

- Migalastat Personalized Medicine (Small Molecule)
- MAA Submitted
- CHMP Opinion Anticipated Early 2016
- Prepared for EU Launch\*

### Epidermolysis Bullosa (EB)

- Phase 3 Novel Topical Cream (SD-101)
- U.S. Breakthrough Therapy Designation
- Rolling NDA
- Phase 3 Data Expected in 2H16

### Pompe

- Novel ERT + Chaperone Treatment Paradigm
- Biologics Manufacturing
- Clinical Study Initiated with Data Anticipated in 2016

R&D Engine and Continued Business Development Activity

\*Pending Approval



# Migalastat Personalized Medicine for Fabry Disease

# Fabry Disease Overview – For Meetings

## Fabry Disease is a Fatal Genetic Disorder that Affects Multiple Organ Systems

### Leading Causes of Death

#### TRANSIENT ISCHEMIC ATTACK (TIA) & STROKE<sup>1</sup>

#### HEART DISEASE<sup>2</sup>

- Irregular heartbeat (fast or slow)
- Heart attack or heart failure
- Enlarged heart

#### KIDNEY DISEASE<sup>3</sup>

- Protein in the urine
- Decreased kidney function
- Kidney failure

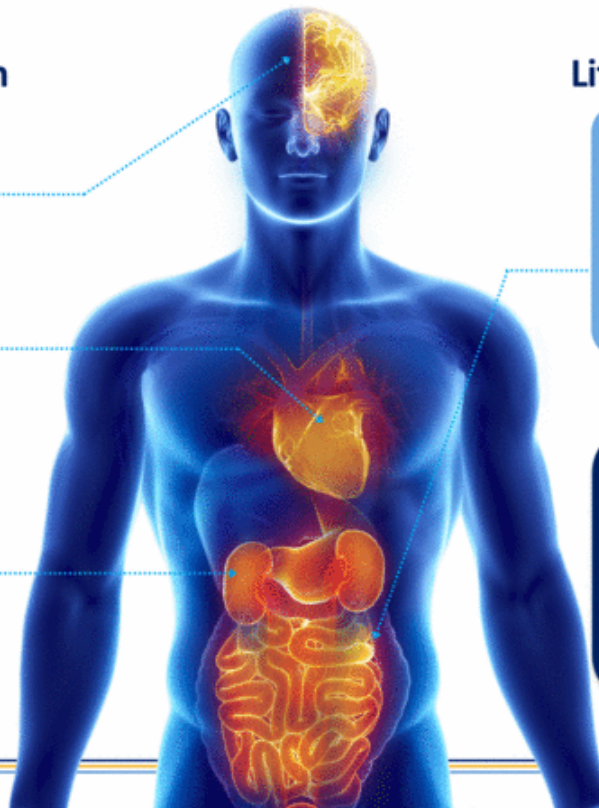
### Life-Limiting Symptoms

#### GASTROINTESTINAL<sup>3</sup>

- Nausea, vomiting, cramping, and diarrhea
- Pain/bloating after eating, feeling full
- Constipation
- Difficulty managing weight

### Key Facts

- Deficiency of  $\alpha$ -Gal A enzyme leading to GL-3 accumulation
- >800 known mutations
- 5-10K diagnosed WW (51% female/49% male<sup>4</sup>)
- Newborn screening studies suggest prevalence of ~1:1000 to ~1:4000



1. Desnick R, et al. *Ann Intern Med.* 2003 2. Yousef Z, et al. *Eur Heart J.* 2013 3. Germain D. *Orphanet J Rare Dis.* 2010 4. Fabry Registry 2011

# Fabry Disease Overview – For Podium

Fabry Disease is a Fatal Genetic Disorder that Affects Multiple Organ Systems

## Leading Causes of Death

TRANSIENT ISCHEMIC  
ATTACK (TIA) & STROKE<sup>1</sup>

HEART DISEASE<sup>2</sup>

KIDNEY DISEASE<sup>3</sup>

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# Summary of Clinical Data

## Favorable Efficacy and Safety Data in Two Largest Phase 3 Studies Ever Completed in Fabry Disease



### Reduction in Disease Substrate

IC GL-3 (Study 011<sup>4</sup>)  
Plasma Lyso Gb-3 (Study 011<sup>3,4</sup> and 012<sup>2</sup>)

### Stability of Kidney Function

Estimated Glomerular Filtration Rate (eGFR) and measured GFR  
(Study 011<sup>1</sup> and Study 012<sup>1,2</sup>)

### Reduction in Cardiac Mass

Left Ventricular Mass Index (LVMI) (Study 011<sup>3</sup> and 012)

### Improvement in GI Symptoms

Gastrointestinal Symptoms Rating Scale (GSRS) (Study 011<sup>4</sup>)

### Low Rate of Fabry-Associated Clinical Events

Renal, Cardiac and Cerebro-Vascular Events (Study 012<sup>2</sup>)

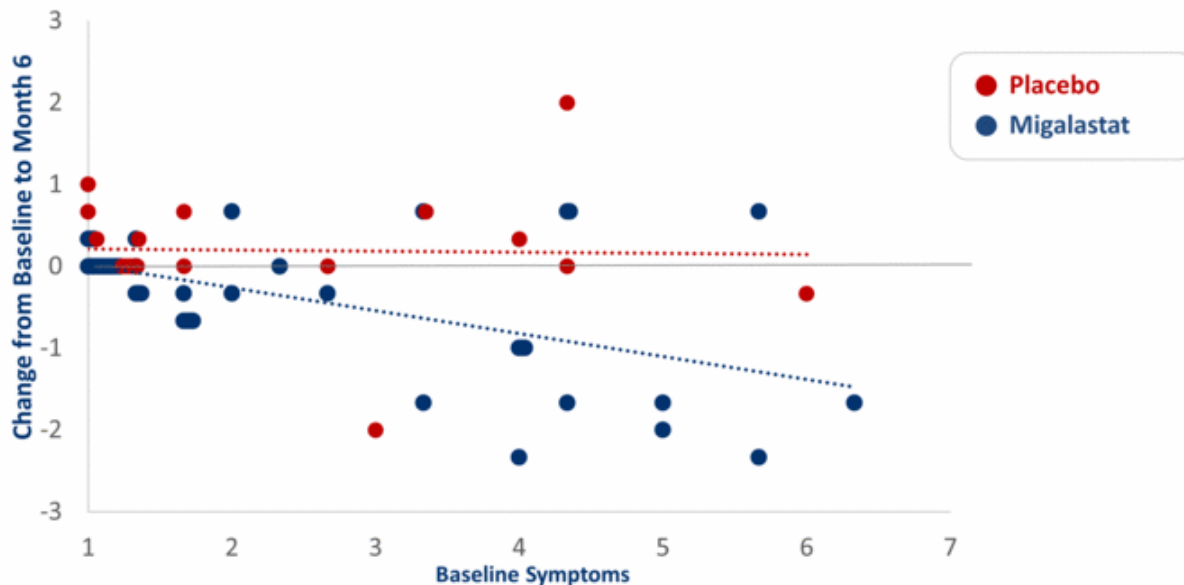
1: Stabilization from baseline over 18 months with favorable comparison to natural history in literature  
2: Comparable to ERT over 18 months  
3: Improvement from baseline over 18+ months  
4: Improvement versus placebo over 6 months in amenable patients

# Additional Phase 3 Data on Diarrhea Symptoms

**NEW**

**Migalastat has Generated Promising Data on Diarrhea Improvement, One of the Most Life-Limiting Symptoms of Fabry Disease**

Statistically Significant Change - Migalastat vs. Placebo from Baseline to Month 6 (nominal p=0.026)



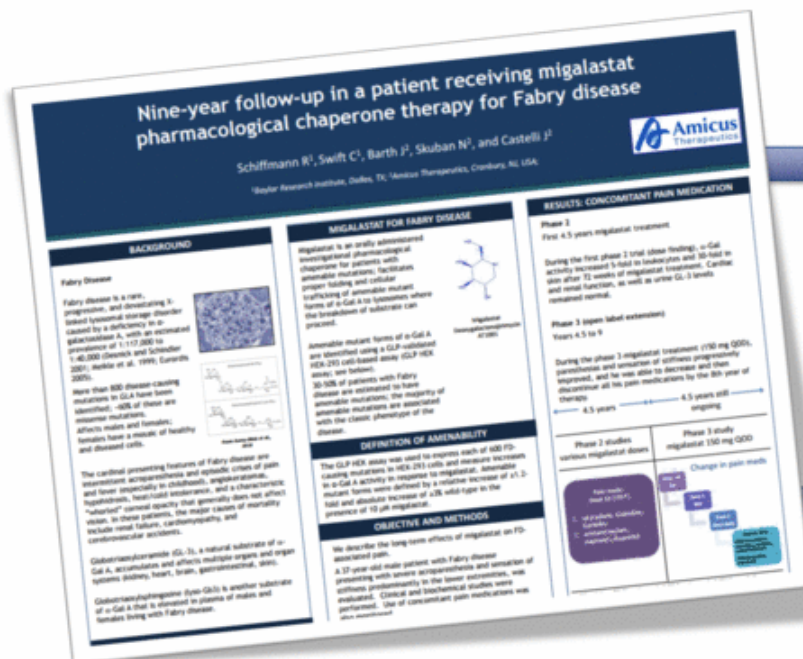
Note: Month 6 used as baseline for patients in placebo arm switching to migalastat; MID from Chan 2006 in kidney transplant. Minimal important difference (MID) for the GRSRS diarrhea domain is 0.4 (Chan 2006, renal transplant patients)





# Fabry Patient Perspective

## Case Report from Long-Term Treatment with Migalastat Shows Improvement in Pain and Return to Everyday Activities<sup>1</sup>



### Patient Journey to Diagnosis

- Chronic pain
- Weakness and fatigue
- Pain medication

### Phase 2 Study + OLE for 4.5 Years

- Enzyme activity increased
- Cardiac and renal functions remained normal

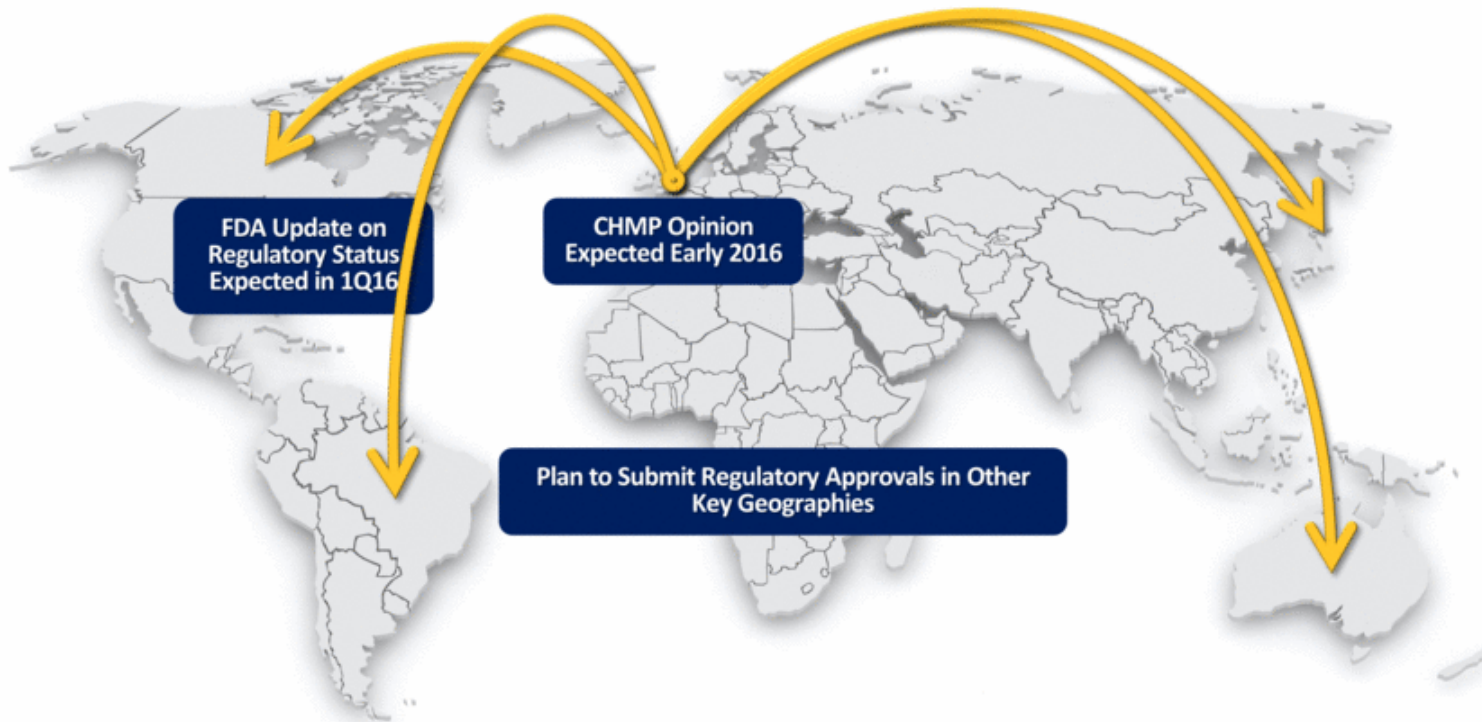
### Phase 3 OLE for 4.5 Years (Still Ongoing)

- Paresthesias and sensation of stiffness improved
- Feels well, goes to gym and works
- Discontinued pain medication

1. Schiffmann, et al. [Nine-year follow-up in a patient receiving migalastat pharmacological chaperone therapy for Fabry disease](#). SSIEM 2015.

# Global Regulatory Strategy

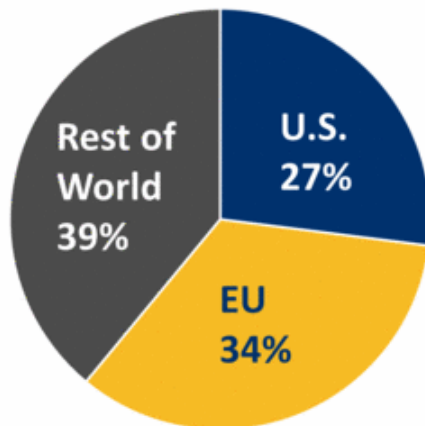
**EU Approval Will Lay the Foundation to Address ~70% of Global Fabry Market**



# Fabry Market Today

Migalastat has Potential to Offer a Number of Important “Firsts” for Fabry Patients

**\$1.1B in FY14 ERT Sales<sup>1</sup>**



- 40-50% of Diagnosed Patients Not on ERT Today
- Market Continues to Grow > 10% / Year
- ERT Infused Once Every 2 Weeks

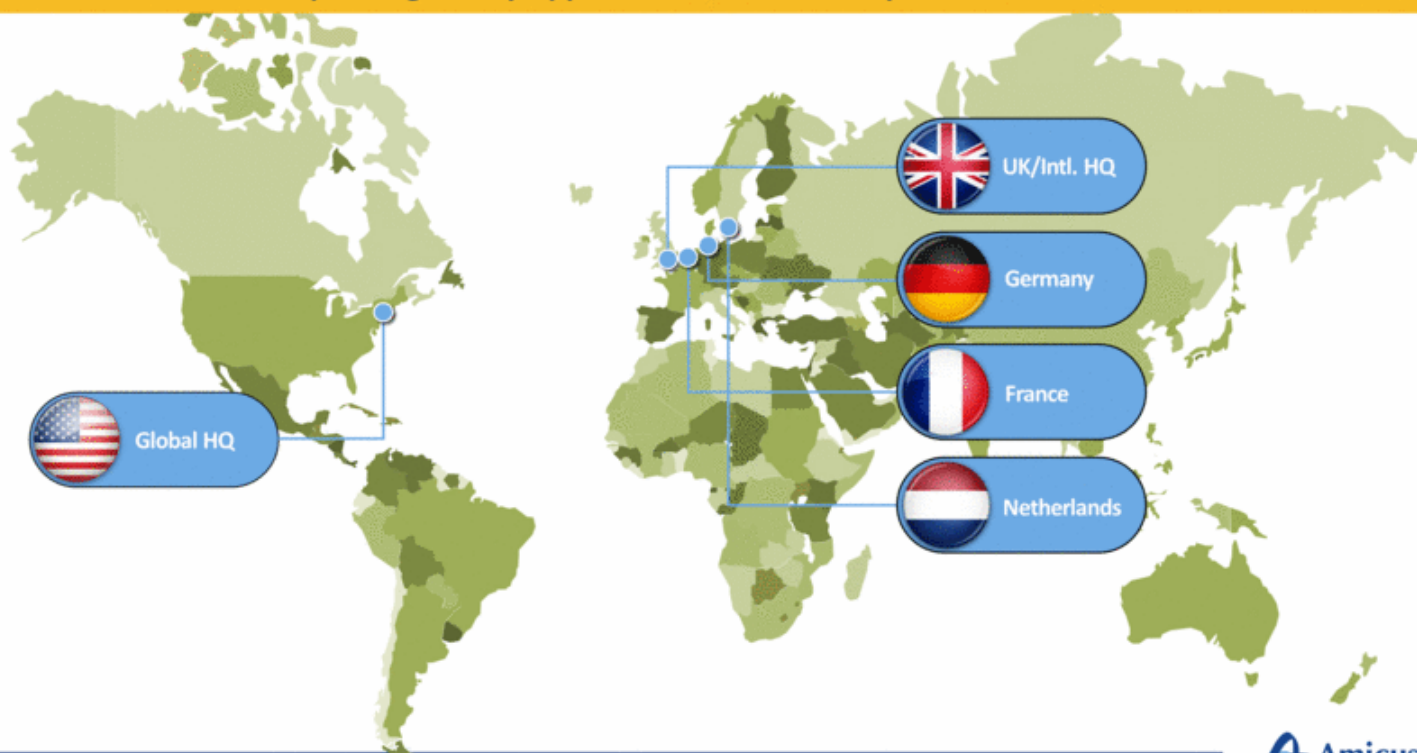
- **First** new product in > 10 years
- **First** oral therapy
- **First** targeted therapy for amenable patients (30%-50% of population)



1. Company filings and Amicus estimates

# Global Infrastructure and International Team

**World-Class Global Commercial Team to Support Migalastat Launch  
Upon Regulatory Approvals with Further Expansion in 2016**



# Global Infrastructure and International Team

**World-Class Global Commercial Team to Support Migalastat Launch  
Upon Regulatory Approvals with Further Expansion in 2016**



## Commercial Launch Preparation Activities



Medical education and patient advocacy ongoing on behalf of Fabry patients



Experienced commercial leadership team with established international operations



Patient and physician mapping



Global value dossier complete and local submissions in development



International distribution system

### Amicus is Preparing for Potential Launches in 2016





# SD-101 for Epidermolysis Bullosa (EB)

Poised to deliver pivotal data for a  
devastating rare disease in 2016

# EB Disease Overview

## Rare, Devastating, Connective Tissue Disorder with No Approved Treatments



- Multiple genes cause disease which results in fragility of skin and can affect internal organs
- Diagnosed from infancy to adulthood
- Severe blistering, open wounds and scarring in response to minor friction to the skin
- Disfiguring, excruciatingly painful, and can be fatal
- Given the lack of approved treatment options, any reduction in disease symptoms would be considered meaningful
- 30,000 – 40,000 diagnosed patients in major global regions



# Three Major EB Subtypes Represent ~99% of EB Population

Multiple Subtypes...Single Devastating and Fatal Genetic Disorder

**Simplex**



~75% of EB Population

**Dystrophic**



~20% of EB Population

**Junctional**



~5% of EB Population

INCREASING SEVERITY

No Approved Therapies Today

SD-101 in Development for All 3 Major Subtypes

30,000-40,000+ Diagnosed in Major Markets

# U.S. Breakthrough Therapy Designation

## Positive Early Results from Phase 2a Study Led to Breakthrough Therapy Designation

- Open-label, 8 patient proof-of-concept study<sup>1</sup>
- Ages 6 months – 9 years
- All baseline target wounds  $\geq 10 \text{ cm}^2$
- SD-101 3% applied once daily for 3 months

### Key Findings

**87.5%**

of patients experienced complete closure of target wounds within 1 month

**57%**

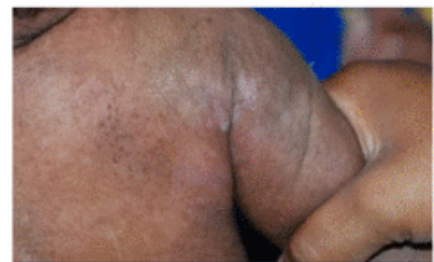
reduction in affected body surface area by month 3

Daily administration generally safe and well-tolerated

### 1-Year-Old Girl with EB Simplex



Baseline



Following 2 months of treatment

1. Simplex (n=3), Junctional (n=3), Recessive Dystrophic (n=2)

# Phase 2b Design (Study 003)

## 3-Month Double-Blind Treatment Period<sup>1</sup>

SD-101 6% (n=15)

SD-101 3% (n=16)

Placebo (n=17)

### Primary Efficacy Endpoint: Target Wound Healing at Month 1

- Baseline wound: Chronic ( $\geq 21$  days), size 5-50 cm<sup>2</sup>

### Secondary Efficacy Endpoints Include:

- Time to target wound closure
- Change in Body Surface Area (BSA) of lesional skin

## Optional Extension (SD-004)

Open-Label Zorblisa (6%)

42/44 Patients entered extension study

\$400K FDA Grant for Extension Study

## 48 EB patients (age $\geq 6$ months)<sup>1</sup> - 1:1:1 Randomization - Daily Topical Application

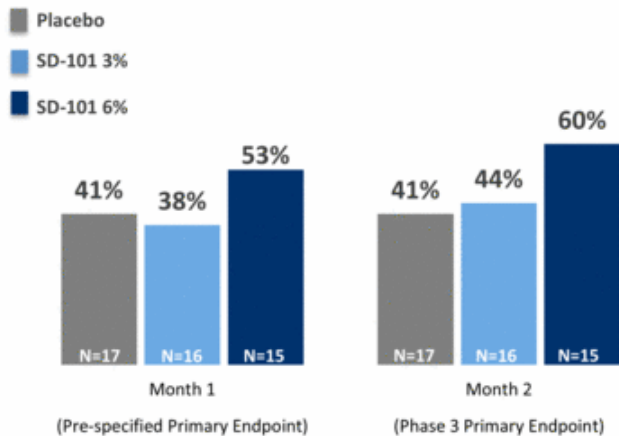
1. Assessments: 0, 14, 30, 60, 90 Days. 2. Initial Disease Severity: Mean target lesion size (cm<sup>2</sup>) 14.0 (range 5-39); mean lesional BSA: 19.4% (range 0.4-48%); mean wound age (days): 182 (range 21-1,639). EB Subtypes enrolled: Simplex (n=11), Recessive Dystrophic (n=29), and Junctional (n=8)

# Phase 2b Results

## SD-101 6% Demonstrated Higher Proportion of Complete Target Wound Closure

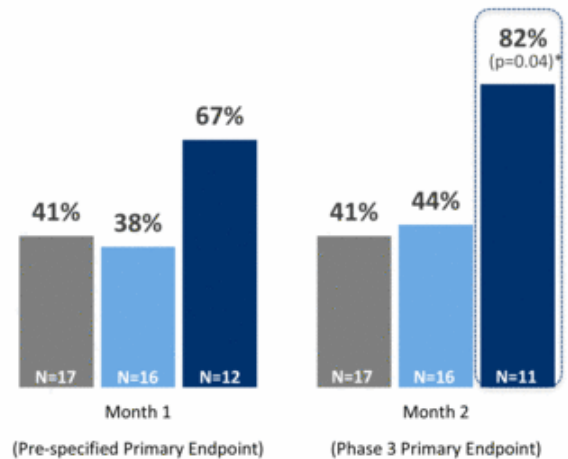
### ITT Population (n=48)

Proportion of Complete Target Wound Closure (%)



### Evaluable Population<sup>1</sup> (n=45)

Proportion of Complete Target Wound Closure (%)

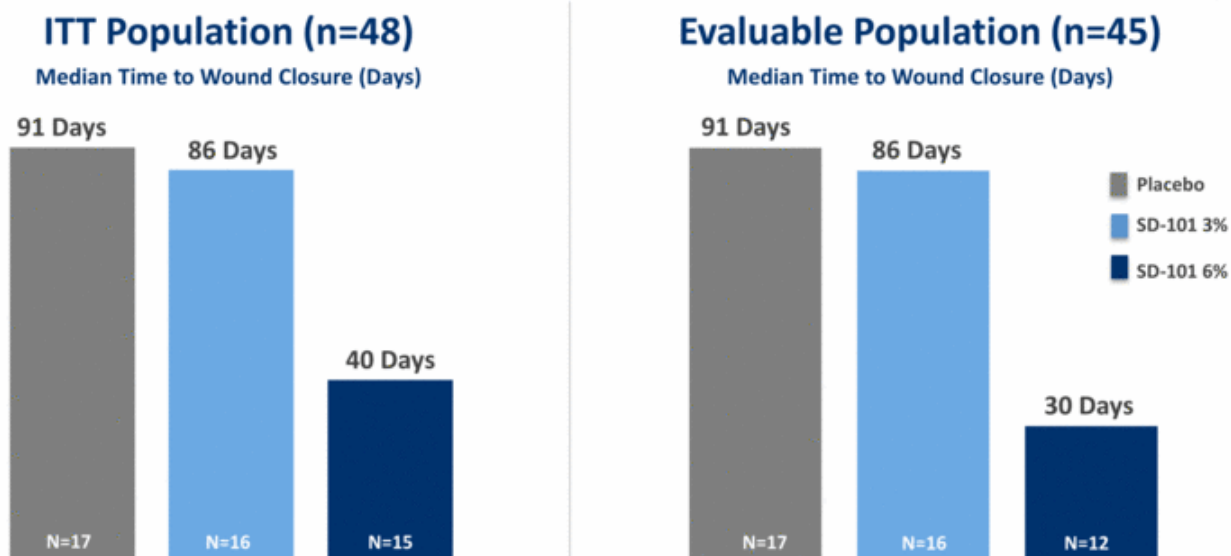


\*SD-101 6% vs placebo, unadjusted p=0.04

1. Excluded from Evaluable population: 1 patient (due to lost to follow-up), 2 patients (did not have single identified and qualified target lesion). 1 additional patient lost to follow up after Month 1 visit and is excluded from target wound assessment at later time points

## Phase 2b Results – Secondary Endpoint

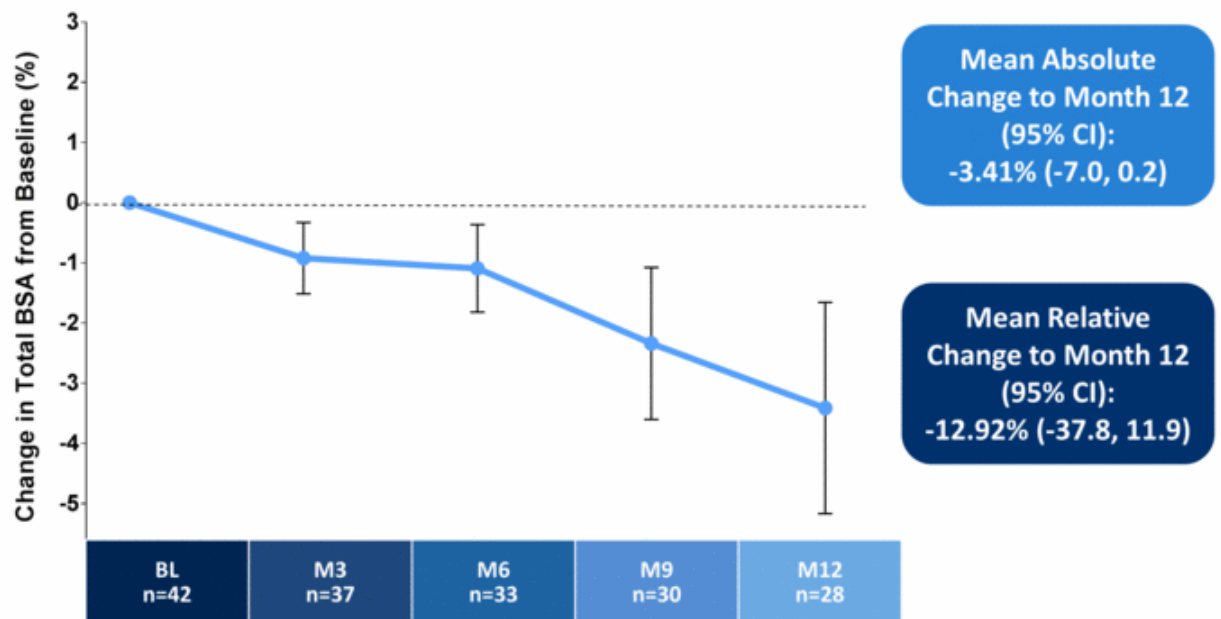
**SD-101 6% Showed Fastest Time to Wound Closure;  
SD-101 Generally Safe and Well-Tolerated**



**Adverse Events Similar Across Treatment Arms of Placebo, SD-101 3%, and SD-101 6%**

# Phase 2b Extension (Study 004) Results

## Results on Total Body Surface Area (BSA) Affected by Wounds and Lesions



Note: Mean and SEM on change from baseline are plotted. Study 004 Total BSA baseline values are: N=42 Baseline population: 11.3. N=28 population used for Month 12 comparison: 10.9

## Phase 3 Design (SD-005)

Phase 3 Initiated in 2Q15 and ~50% Enrolled  
Top-Line Data Expected 2H16

### 3-Month Double-Blind Treatment Period<sup>1</sup>

SD-101 6%

~150 EB patients (age ≥ 1 month)

Placebo

### Optional Extension (SD-006)

Open-Label Zorblisa (6%)

44/44 Patients Have  
Continued in Open-Label  
Extension  
(Jan. 8, 2015)

#### Primary Endpoint: Target Wound Healing at Month 2

- US and EU regulatory authorities agreed on primary endpoint
- Baseline wound: Chronic (≥ 21 days), size ≥10 cm<sup>2</sup>

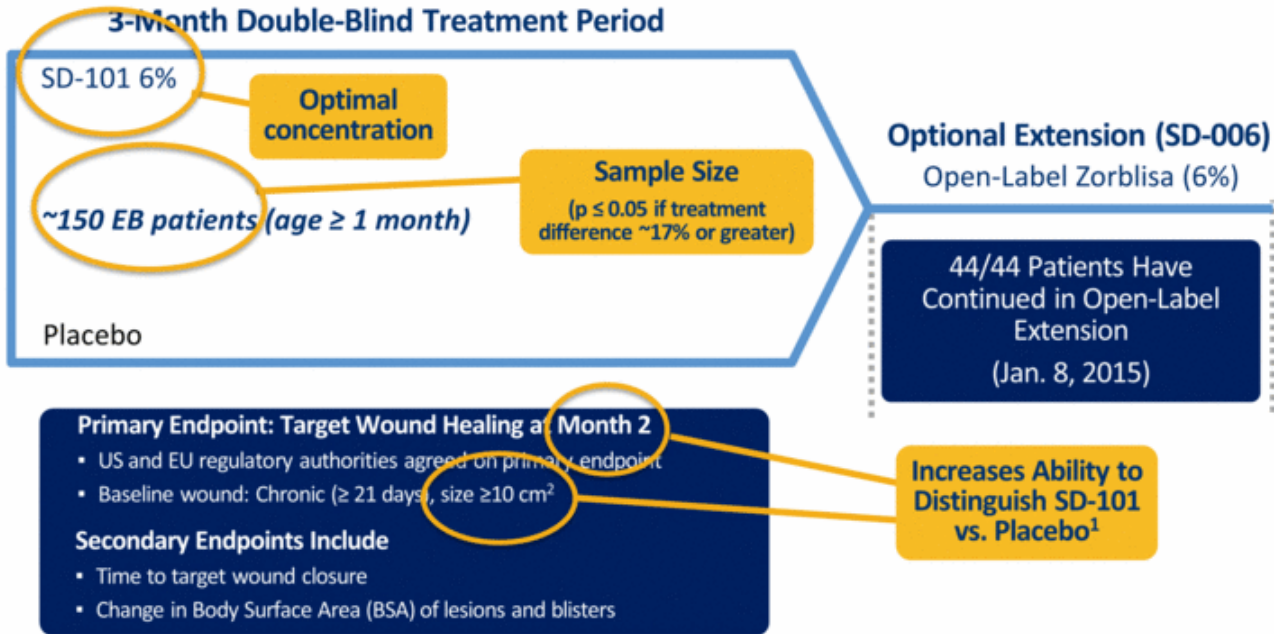
#### Secondary Endpoints Include

- Time to target wound closure
- Change in Body Surface Area (BSA) of lesions and blisters

1. Assessments: 0, 14, 30, 60, 90 Days. 1:1 randomization, daily topical application

# Phase 3 Design (SD-005)

## Study Design Incorporates Key Learnings from Phase 2b Study

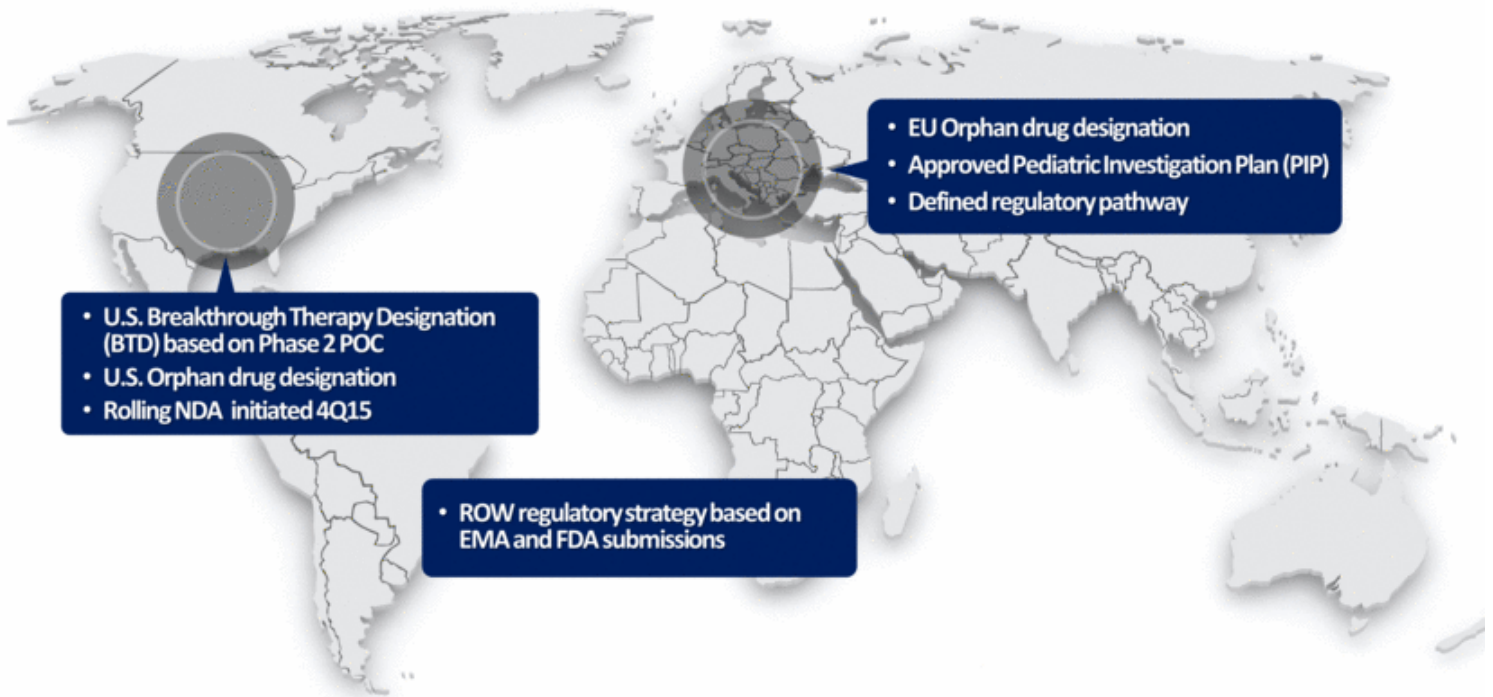


1. Complete target wound closure in patients with target wounds  $\geq 10$  cm<sup>2</sup> at Month 2 in Phase 2b: SD-101 6% - 50% (n= 4) vs. Placebo - 12.5% (n=8)



# Global Regulatory Strategy

## Positive FDA and EMA Feedback on Phase 3 Study Design

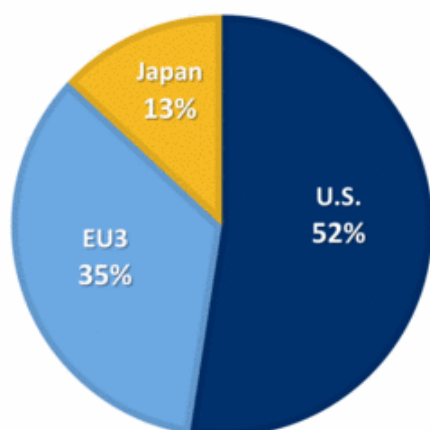


# \$1B+ Commercial Potential

## KOL Feedback Supports Profound Unmet Medical Need and Broad Usage in All EB Subtypes

### Diagnosed EB Patients by Geography

(US, EU3, Japan)



#### Significant Unmet Clinical Need

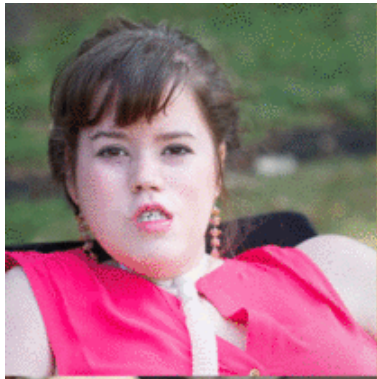
- No approved treatments, opportunity for first-in-class
- Promising proof-of-concept in all EB subtypes

#### Strong Support Among Surveyed Stakeholders

- Physicians indicate usage in 100% patients
- Payers indicate support for broad reimbursement if approved

#### Large Commercial Opportunity

- 30,000 – 40,000 diagnosed patients in major markets
- KOLs expect diagnosis rates to increase



# ATB200 Novel ERT for Pompe Disease

A Proprietary, Clinical-Stage Biologics Program

# Pompe Disease Overview

## Severe, Fatal Genetic Disorder with Significant Unmet Medical Need



- Deficiency of GAA leading to glycogen accumulation
- Age of onset ranges from infancy to adulthood
- Symptoms include muscle weakness, respiratory failure and cardiomyopathy
- Respiratory and cardiac failure are leading causes of morbidity and mortality
- 5,000 – 10,000 patients diagnosed WW<sup>1</sup>
- ~\$700M+ Global Pompe ERT sales in FY14<sup>2</sup>

1. National Institute of Neurological Disorders and Stroke (NIH). 2. Sanofi Press Release & 10-K

# Pompe Patient Perspectives

**Very Significant Unmet Need Despite Availability of Currently Marketed Therapy**



# Pompe ERT - 3 Challenges

## Amicus Technology Platforms with Potential to Address Challenges with Existing Pompe ERT

### Activity/ Stability

Rapid denaturation of ERT in pH of blood<sup>1</sup>

Protein Aggregation



CHAPERONE-ADVANCED  
REPLACEMENT THERAPY

### Tolerability / Immunogenicity

Infusion-associated reactions in >50% of late-onset patients<sup>3</sup>

Antibody titers shown to affect treatment outcomes<sup>4,5</sup>

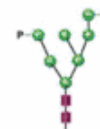


CHAPERONE-ADVANCED  
REPLACEMENT THERAPY

### Uptake/ Targeting

Low M6P receptor uptake into skeletal muscle<sup>2</sup>

Vast majority of rhGAA not delivered to lysosomes<sup>2</sup>



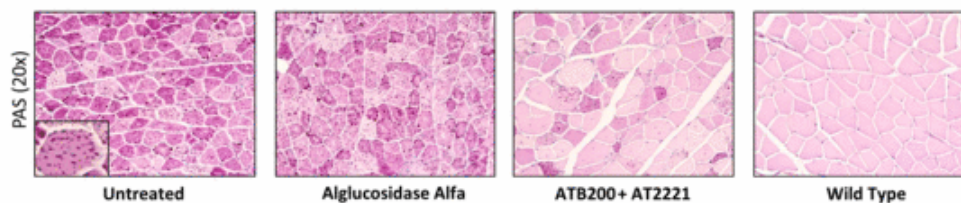
Uniquely Engineered  
rhGAA Optimized M6P  
& Carbohydrates

<sup>1</sup>Khanna et al., PLoS ONE, 2012; <sup>2</sup>Zhu et al., Amer. Soc. Gene Therapy, 2009 June; <sup>3</sup>Banati et al., Muscle Nerve, 2011 Dec.; <sup>4</sup>Banugaria et al., Gen. Med., 2011 Aug.; <sup>5</sup>Sde Vries et al., Mol Genet Metab., 2010 Dec.

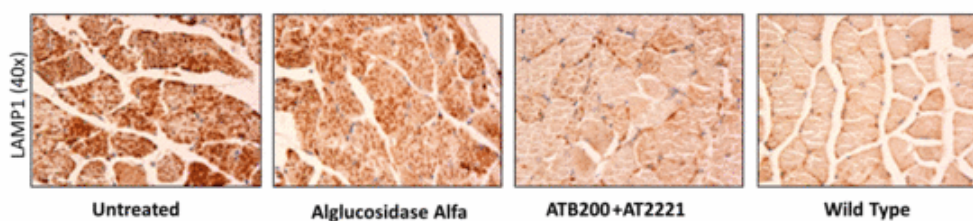
# Preclinical Proof-of-Concept

## ATB200 + Chaperone Results in Improved Substrate Clearance in Preclinical Models<sup>1</sup>

### PAS-glycogen staining in Quadriceps



### LAMP1 Immunohistochemical staining in Soleus

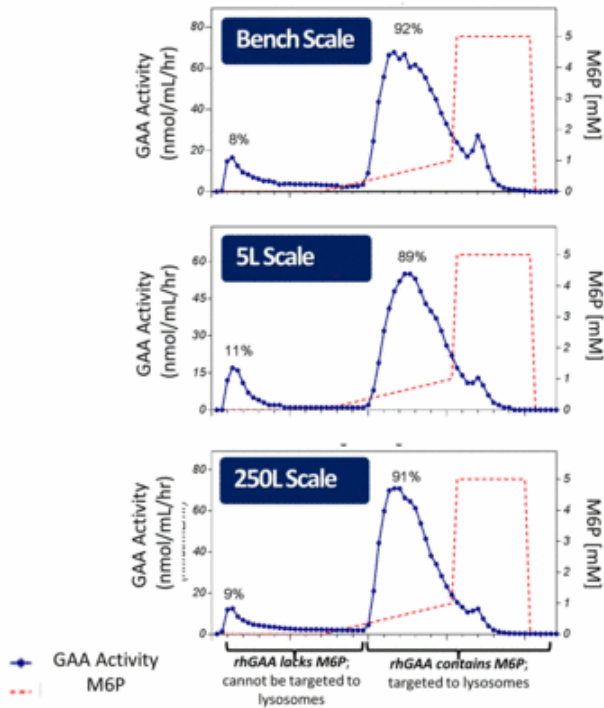


1. Following 2 doses of 20mg/kg alglucosidase alfa or ATB200 + AT2221 in Gaa KO mice, skeletal muscles evaluated for glycogen clearance and proliferated lysosomes. Treatment with alglucosidase alfa modestly reduced glycogen or proliferated lysosomes while ATB200, co-administered with AT2221 significantly decreased the muscle pathology associated with Pompe disease

# Biologics Manufacturing Capabilities

## Optimized Glycosylation and Key Quality Attributes Maintained Through Scale Up

### CI-MPR Receptor Chromatography



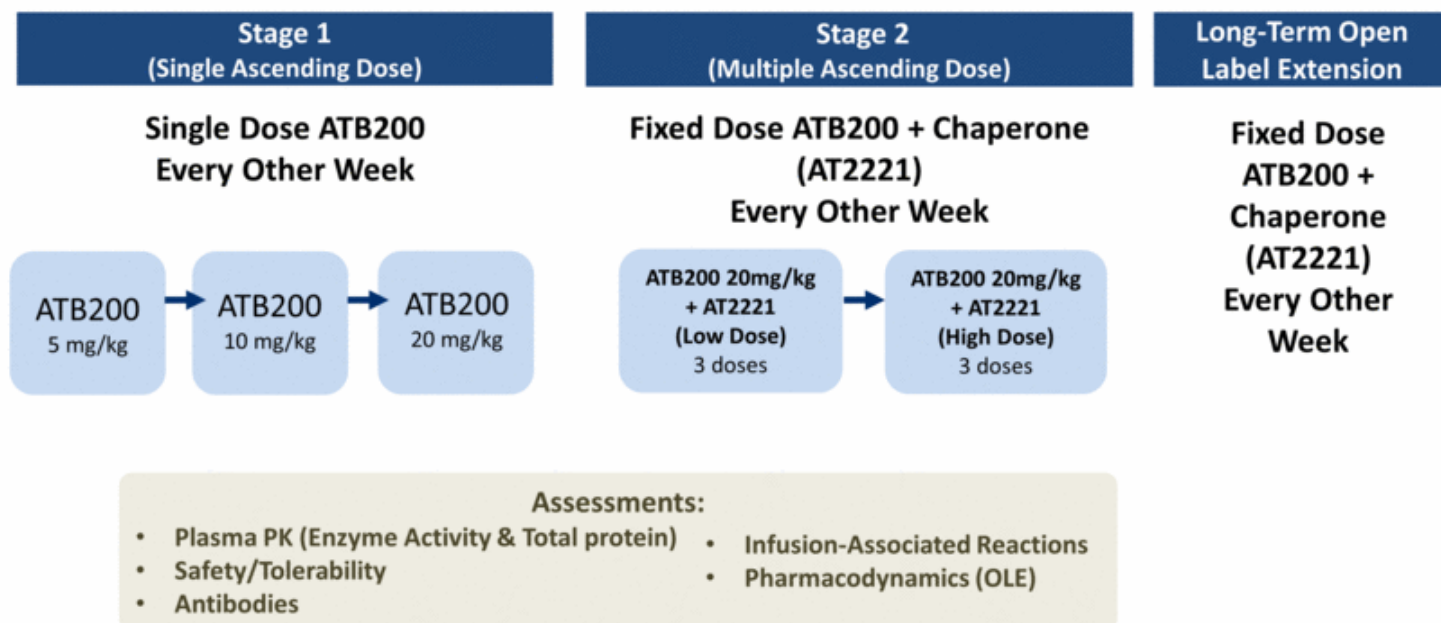
### Lyophilized Vial of ATB200

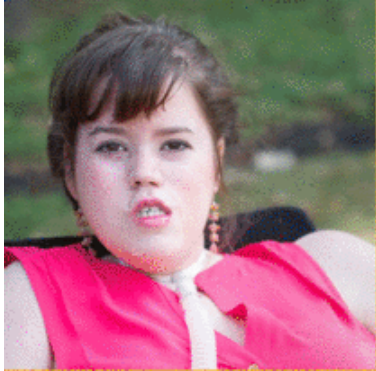




# Clinical Study in Pompe Patients

## Study Design Supported by US and EU Regulators





# Financial Summary

Strong Balance Sheet to Invest in Rare Disease Pipeline

# Strong Balance Sheet

Cash Position Provides Runway Under Current Operating Plan into 1H17

Financial Position	December 31, 2015
Current Cash:	\$214M
Current Debt	\$50M
FY16 Net Cash Spend Guidance:	\$135M-\$155M
Cash Runway	Mid-2017
<b>Capitalization</b>	
Shares Outstanding	125,027,034

## Key Drivers of Value

### 3 Novel Product Candidates Each with \$500M to \$1B+ Market Potential

#### Fabry

- Migalastat Personalized Medicine (Small Molecule)
- MAA Submitted
- CHMP Opinion Anticipated Early 2016
- Prepared for EU Launch\*

#### Epidermolysis Bullosa (EB)

- Phase 3 Novel Topical Cream (SD-101)
- U.S. Breakthrough Therapy Designation
- Rolling NDA
- Phase 3 Data Expected in 2H16

#### Pompe

- Novel ERT + Chaperone Treatment Paradigm
- Biologics Manufacturing
- Clinical Study Initiated with Data Anticipated in 2016

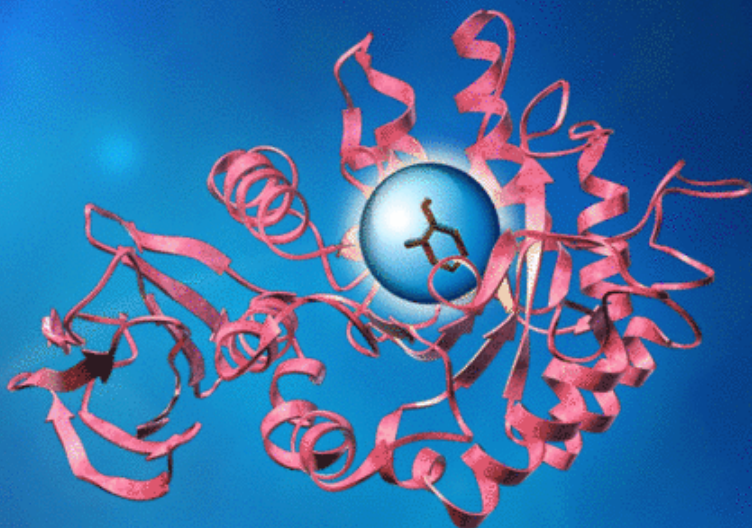
### R&D Engine and Continued Business Development Activity

\*Pending Approval

Thank You

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***3<sup>rd</sup> Annual Dermatology Summit  
SD-101 for Epidermolysis Bullosa (EB)***

*Jay Barth, MD, Chief Medical Officer  
at the forefront of therapies  
for rare and orphan diseases*

# Safe Harbor

*This presentation will contain, "forward-looking statements" within the meaning of the Private Securities Litigation Reform Act of 1995 relating to preclinical and clinical development of Amicus' candidate drug products, the timing and reporting of results from preclinical studies and clinical trials evaluating Amicus' candidate drug products, financing plans, and the projected cash position for the Company. Words such as, but not limited to, "look forward to," "believe," "expect," "anticipate," "estimate," "intend," "potential," "plan," "targets," "likely," "may," "will," "would," "should" and "could," and similar expressions or words identify forward-looking statements. Such forward-looking statements are based upon current expectations that involve risks, changes in circumstances, assumptions and uncertainties. The inclusion of forward-looking statements should not be regarded as a representation by Amicus that any of its plans will be achieved. Any or all of the forward-looking statements in this press release may turn out to be wrong. They can be affected by inaccurate assumptions Amicus might make or by known or unknown risks and uncertainties. For example, with respect to statements regarding the goals, progress, timing and outcomes of discussions with regulatory authorities, and in particular the timing of an NDA submission for migalastat monotherapy, and the potential goals, progress, timing and results of preclinical studies and clinical trials, actual results may differ materially from those set forth in this release due to the risks and uncertainties inherent in the business of Amicus, including, without limitation: the potential that results of clinical or pre-clinical studies indicate that the product candidates are unsafe or ineffective; the potential that it may be difficult to enroll patients in our clinical trials; the potential that regulatory authorities may not grant or may delay approval for our product candidates; the potential that preclinical and clinical studies could be delayed because we identify serious side effects or other safety issues; the potential that we will need additional funding to complete all of our studies and, our dependence on third parties in the conduct of our clinical studies. Further, the results of earlier preclinical studies and/or clinical trials may not be predictive of future results. With respect to statements regarding projections of the Company's cash position, actual results may differ based on market factors and the Company's ability to execute its operational and budget plans. In addition, all forward looking statements are subject to other risks detailed in our Annual Report on Form 10-K for the year ended December 31, 2014 and Form 10-Q for the quarter ended June 30, 2015. You are cautioned not to place undue reliance on these forward-looking statements, which speak only as of the date hereof. All forward-looking statements are qualified in their entirety by this cautionary statement, and Amicus undertakes no obligation to revise or update this news release to reflect events or circumstances after the date hereof. This caution is made under the safe harbor provisions of Section 21E of the Private Securities Litigation Reform Act of 1995.*

# Epidermolysis Bullosa (EB)

Rare, Devastating, Connective Tissue Disorder with No Approved Treatments



- Multiple genes cause disease which results in fragility of skin and can also affect internal organs
- Diagnosed from infancy to adulthood
- Severe blistering, open wounds and scarring in response to minor friction to the skin
- Disfiguring, excruciatingly painful, and can be fatal
- Given lack of treatment, any reduction in disease symptoms would be considered meaningful
- 30,000 – 40,000 **diagnosed** patients in major global regions

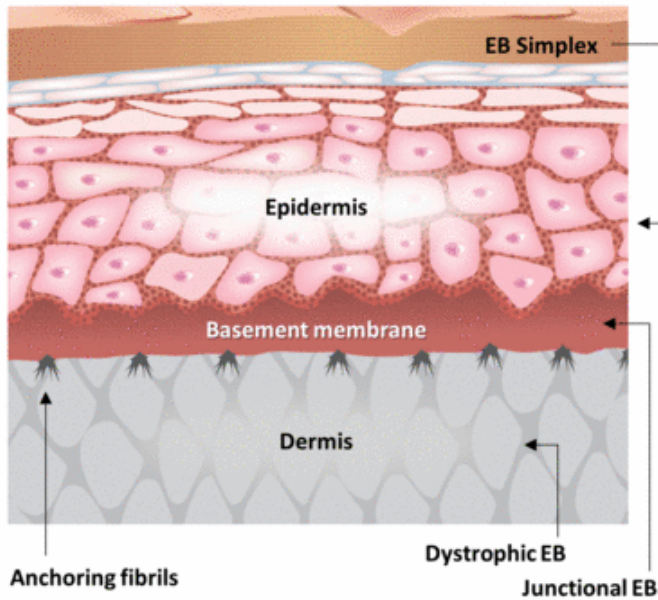


# Three Major EB Subtypes

Three Major EB Subtypes Differ By Physical Manifestations, Genetic Makeup, and Prognosis

## Skin structure

Sites of primary blister formation



## EB subtypes

Represent ~99% of EB Population

Subtypes	Symptoms	Frequency	Mortality risk
Junctional	<ul style="list-style-type: none"> <li>External blistering</li> <li>Internal blistering (oral tract, internal organs)</li> <li>Severe complications can become fatal early in life</li> </ul>	~5%	
Dystrophic	<ul style="list-style-type: none"> <li>External blistering</li> <li>Narrowing of esophagus</li> <li>Higher risk of aggressive skin cancer</li> <li>Associated with mortality</li> </ul>	~20%	
Simplex	<ul style="list-style-type: none"> <li>Localized and generalized external blistering</li> </ul>	~75%	

SD-101 being developed for all major EB subtypes

# SD-101 Overview

*Patented High Concentration Allantoin with Breakthrough Therapy Designation*

**Novel, Proprietary Topical Cream Promotes Healing of Wounds in EB and is Differentiated by Applicability for All Major EB Subtypes**

Active Ingredient & ROA

Proprietary topical cream containing 6% allantoin, applied to entire body once daily

Proposed Indication

All major EB subtypes (Simplex, Dystrophic, Junctional)

Development Phase

Phase 3 registration study (SD-005) ongoing

Proposed MOA\*

Aids inflammatory response, bactericidal effects, loosens protein bridges, promotes collagen

Formulation

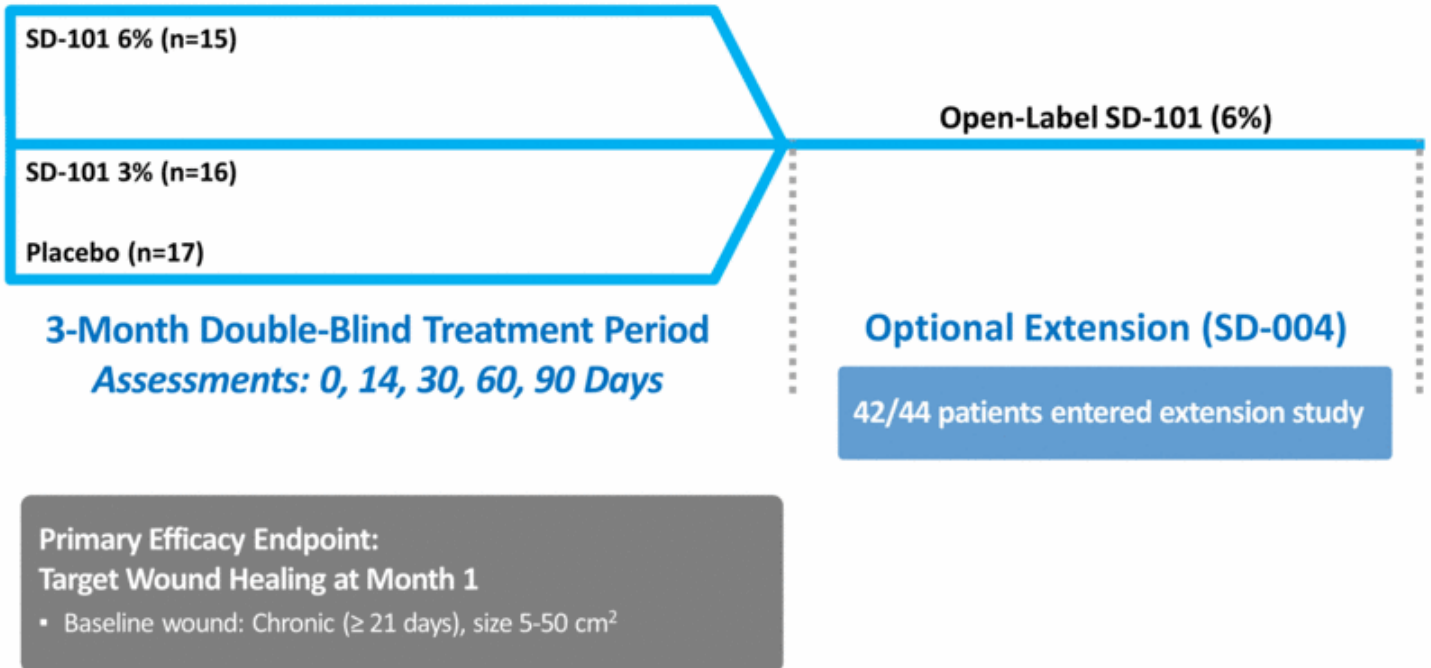
Patented formulation to deliver high concentration in highly stable, soluble form



\*Margraf and Covey 1977; Meixell and Mecca 1966; Settle 1969; Flesch 1958; Fisher 1981; Cajkovic et al., 1992; Medda 1976

# Phase 2b (Study 003) Design

48 EB patients (age  $\geq$  6 months)\* - 1:1:1 Randomization - Daily Topical Application



*\*Initial Disease Severity: Mean target lesion size (cm<sup>2</sup>) 14.0 (range 5-39); mean lesional BSA: 19.4% (range 0.4-48%); mean wound age (days): 182 (range 21-1,639)*  
*EB Subtypes enrolled: Simplex (n=11), Recessive Dystrophic (n=29), and Junctional (n=8)*

# Phase 2b (Study 003) Safety Summary

Adverse Events Similar Across Treatment Arms of Placebo, SD-101 3%, and SD-101 6%

- Treatment-emergent adverse events (TEAE) generally similar across treatment groups
- No deaths and no severe TEAEs
- No serious adverse events reported in SD-101 6% group

## Treatment Emergent Adverse Events $\geq 10\%$ Frequency

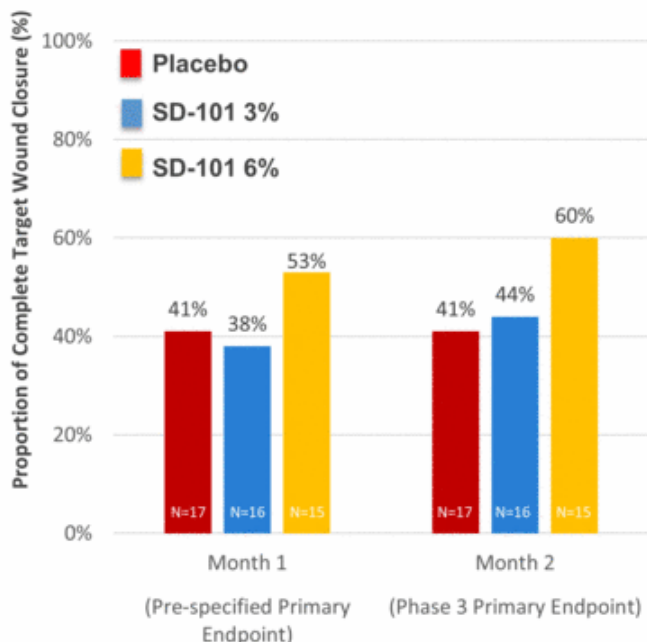
	Placebo	SD-101 3%	SD-101 6%
N subjects	17	16	15
N subjects with TEAEs (%)	12 (70.6)	13 (81.3)	9 (60.0)
Nasopharyngitis	12%	25%	7%
Pyrexia	12%	19%	33%
Application Site Pain	6%	19%	13%
Pain	-	-	13%
Skin and Subcutaneous Tissue Disorders	35%	19%	20%
Pruritus	6%	13%	13%
Rash	12%	-	7%
Rash Erythematous	12%	-	-
Cough	6%	-	13%
Oropharyngeal Pain	12%	-	-
Rhinorrhea	-	-	13%
Vomiting	6%	6%	13%
Headache	12%	-	7%

# Phase 2b (Study 003) Primary Endpoint Results

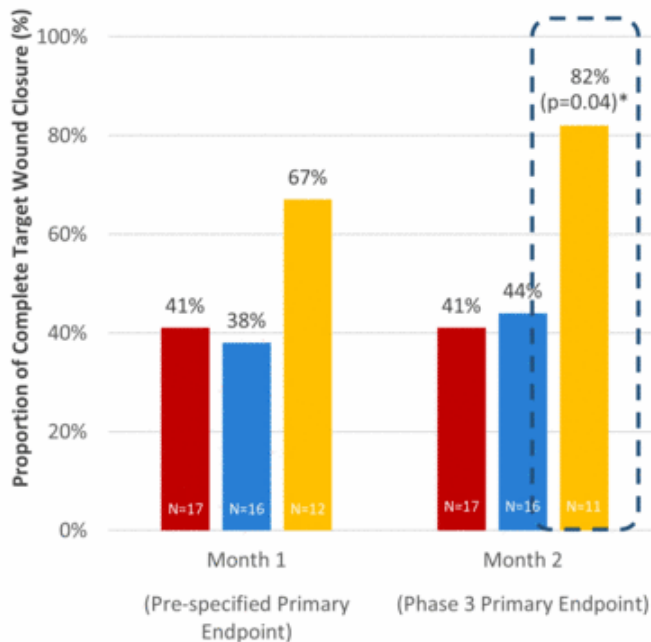
## % Patients with Complete Closure of Target Wounds

**SD-101 6% Demonstrated Higher Proportion of Complete Target Wound Closure at Pre-Specified Endpoint and Subsequently During the Study**

**ITT Population (n=48)**



**Evaluable Population (n=45)**

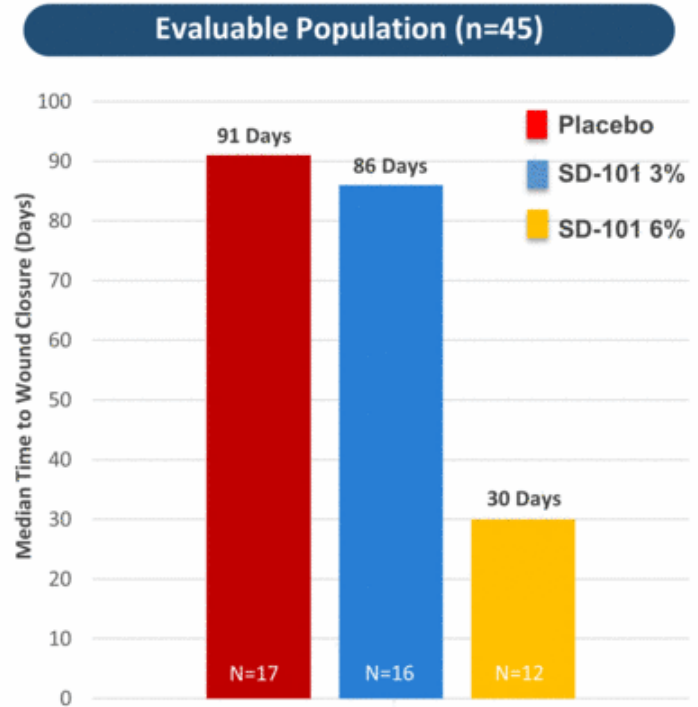
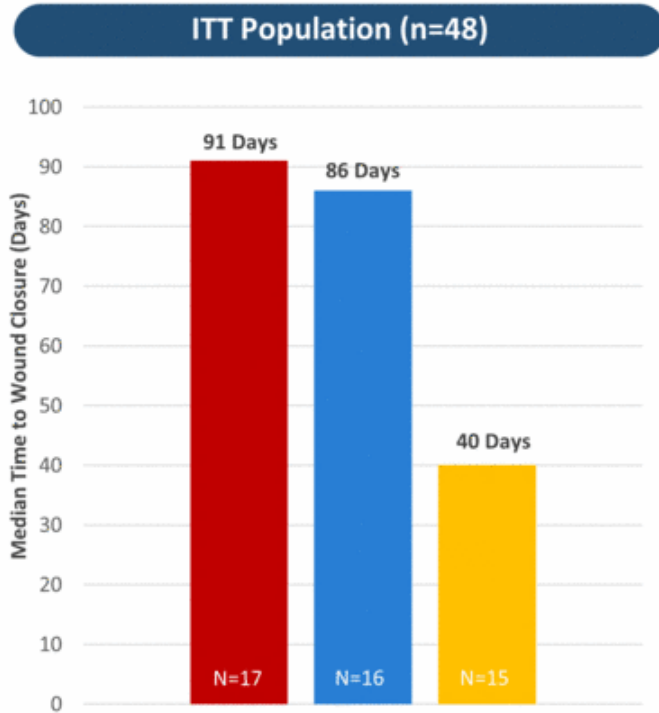


\*SD-101 6% vs placebo, unadjusted p=0.04

Excluded from Evaluable population: 1 patient (due to lost to follow-up), 2 patients (did not have single identified and qualified target lesion), 1 additional patient lost to follow-up after Month 1 visit and is excluded from target wound assessment at later time points

# Phase 2b (Study 003) Secondary Endpoint Median Time to Wound Closure

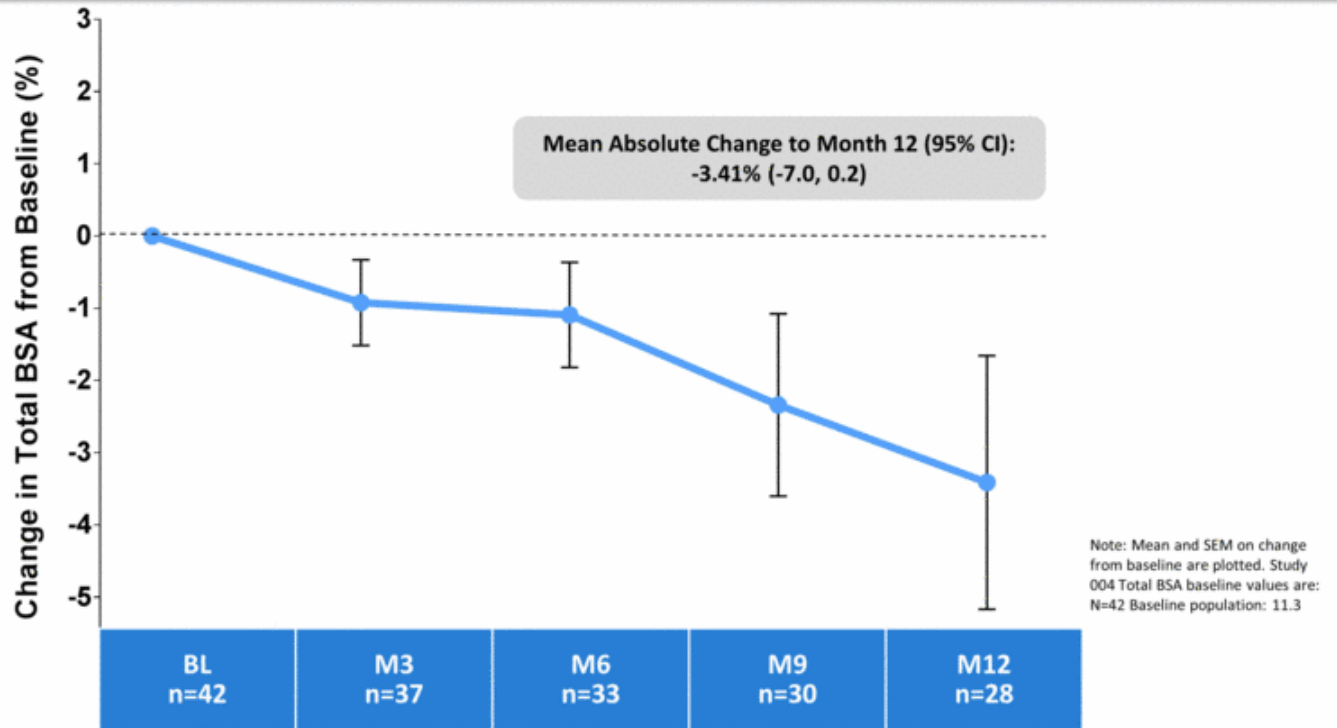
**SD-101 6% Showed Fastest Time to Wound Closure in Both ITT and Evaluable Populations**



Excluded from Evaluable population: 1 patient (due to lost to follow-up), 2 patients (did not have single identified and qualified target lesion)

# Phase 2b Open-Label Extension (Study 004) Total Body Surface Area of EB Lesions

Reductions in Total BSA of EB Lesions Observed  
Through Month 12 in Patients Receiving SD-101 6%



# Phase 2b (Study 003): Results Summary and Key Learnings

## Phase 2b Learnings Informed Dose Selection, Patient Population, and Primary Endpoint for Phase 3 Trial

- SD-101 6% concentration selected for Phase 3 study based on Phase 2b dose response
- Placebo response minimized by analyzing subgroup of patients with wounds  $\geq 10 \text{ cm}^2$ 
  - Complete target wound closure SD-101 6% - 50% (n= 4) vs. Placebo - 12.5% (n=8) at Month 2
- Phase 2b results used to calculate appropriate sample size in Phase 3 study
  - $p \leq 0.05$  if treatment difference  $\sim 17\%$  or greater
- Wound closure at Month 2 (versus Month 1) is optimal time to measure primary endpoint
  - Increases ability to distinguish SD-101 vs placebo
  - Endpoint accepted by FDA and EU regulators
- Defined approval pathway with Phase 3 study design based on EMA and FDA feedback



# Pivotal Phase 3 (Study 005) Underway

*Study Design Supported by Both FDA and EMA*

Phase 3 Initiated in 2Q15 and Currently Enrolling Patients  
Top-line data expected 2H 2016

SD-101 6%

~150 EB patients (age ≥ 1 month)  
1:1 Randomization - Daily Topical Application

Placebo

**3-Month Double-Blind Treatment Period**  
**Assessments: 0, 14, 30, 60, 90 Days**

Optional Extension (SD-006)

Open-Label SD-101 (6%)

38/38 Patients Who Completed Study 005 Continued in Open-Label Extension (Dec. 2015)

#### Primary Efficacy Endpoint: Target Wound Healing at Month 2

- US and EU regulatory authorities agreed to target wound healing as primary endpoint
- Baseline wound: Chronic (≥ 21 days), size ≥10 cm<sup>2</sup>

#### Secondary Endpoints

- Time to target wound closure; Change in Body Surface Area (BSA) of lesions and blisters; itching; pain

# SD-101 Regulatory Pathway

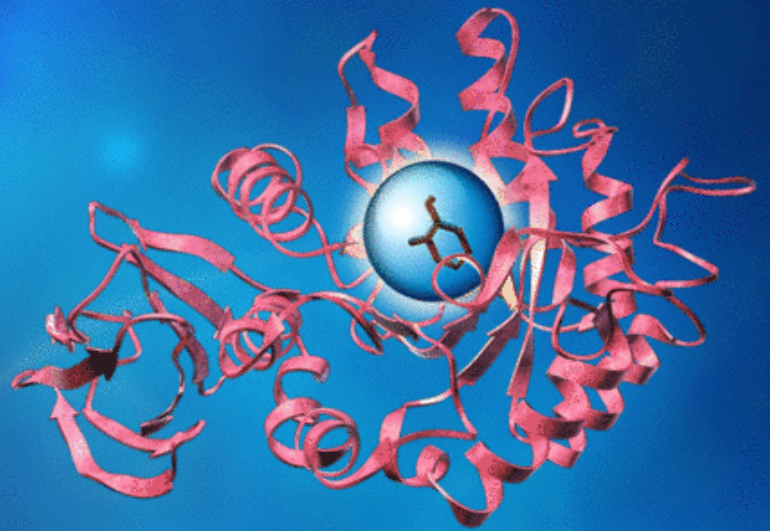
## *Rolling NDA Initiated 4Q15*

FDA and EMA Aligned on Phase 3 Study Design and Feedback to Date Provides Defined Registration Pathway for SD-101 in Major Subtypes of EB

- Breakthrough Therapy Designation (BTD) based on Phase 2 POC
- Orphan drug designation
- Rolling NDA initiated 4Q15

- Orphan drug designation
- Approved Pediatric Investigation Plan (PIP)
- Defined registration pathway

- ROW regulatory path based on EMA and FDA submissions



***3<sup>rd</sup> Annual Dermatology Summit  
SD-101 for Epidermolysis Bullosa (EB)***

*Jay Barth, MD, Chief Medical Officer  
at the forefront of therapies  
for rare and orphan diseases*