



## **Ironwood Lesinurad U.S. Licensing Agreement with AstraZeneca**

April 26, 2016



# Introduction

**Lisa Adler**

Senior Vice President, Corporate  
Communications

# Safe Harbor Statement

This presentation contains forward-looking statements. Investors are cautioned not to place undue reliance on these forward-looking statements, including, but not limited to, statements about the benefits anticipated from the addition of the gout franchise to Ironwood's portfolio; the timing of the closing of the lesinurad transaction; development, launch and commercialization plans for lesinurad and our product candidates; market size, growth and opportunity, including peak sales, and potential demand for lesinurad and our product candidates, as well as their potential impact on applicable markets and commercial and sales strategies; the potential indications for, and benefits of, lesinurad; the anticipated timing of regulatory developments for the fixed-dose combination of lesinurad and allopurinol; the design, timing and results of clinical and preclinical studies; the timing of filings with regulatory authorities; expected periods of patent exclusivity; our potential for rapid, sustainable, high-margin growth; and our company's financial performance and results, and guidance and expectations related thereto, including our projected cash needs and expectations regarding the need for future financings, expectations regarding the accretive nature of the transaction and the timing of such accretion, revenue growth and revenue from the transaction, commercial margin, cash flows, operating expenses, commercial expenses, LINZESS net sales, the effect of the transaction on 2016 LINZESS marketing and sales expense, and the timing of providing updated guidance on total operating expenses following closing. Each forward-looking statement is subject to risks and uncertainties that could cause actual results to differ materially from those expressed or implied in such statement. Applicable risks and uncertainties include, but are not limited to, the risk that the transaction does not close or is delayed; the risk that we are unable to successfully integrate lesinurad into our existing business or are unable to realize the anticipated benefits of the lesinurad transaction; those related to our growth strategy; decisions made by U.S. regulatory authorities, the U.S. Patent and Trademark Office and their foreign counterparts; intellectual property rights of competitors or potential competitors; efficacy, safety and tolerability of lesinurad, linaclotide and our product candidates; competition in disease states; the commercial potential of lesinurad, linaclotide, our product candidates and the other products that we promote; and the risk that we are unable to manage our operating expenses and capital expenditures due to foreseeable or unforeseeable events or occurrences. Applicable risks also include those that are listed under the heading "Risk Factors" and elsewhere in Ironwood's Annual Report on Form 10-K for the year ended December 31, 2015 and in our subsequent SEC filings. These forward-looking statements (except as otherwise noted) speak only as of the date of this presentation, and Ironwood undertakes no obligation to update these forward-looking statements.

# Lesinurad Transaction Overview

**Peter Hecht**  
Chief Executive Officer

# Lesinurad: Strategically + Financially Compelling

U.S. deal includes FDA-approved ZURAMPIC, FDC, other opportunities

## Strong Strategic Fit

- **Category:** 2M uncontrolled gout patients<sup>1</sup>; highly symptomatic<sup>2</sup>; seeking new treatment options
- **Asset:** Durable, innovative therapy with potential to help suffering patients
- Leverages strong commercial capabilities
- 5 U.S. launches expected by 2020 across the IRWD portfolio

## Financially Compelling

- \$100M cash up front plus milestones and single-digit royalty for lesinurad franchise, including ready-to-launch product
- <\$75M annual incremental commercial expenses expected initially
- Transaction expected to be cash flow accretive in 2019
- Lesinurad franchise expected >\$300M peak sales
- Lesinurad franchise commercial margins expected to be >60% by 2022
- Updating 2016 IRWD cash flow guidance to <\$70M from <\$60M cash used in operations; expect 2018 cash flow positive

# Ironwood TODAY: Successful Commercial Biotech

## Multiple blockbuster opportunities

LINZESS®: Branded  
Rx market leader

~\$455M 2015 U.S. net sales  
(~53% y/o/y growth)<sup>1</sup>

~46% 2015 commercial  
margin and expanding

**Highly successful  
primary care launch**

### IBS-C / CIC

>\$2B peak U.S. sales opportunity<sup>2</sup>

### Vascular/Fibrotic diseases

Multiple drug candidates  
with >\$1B peak sales  
opportunities<sup>2</sup>

### Refractory GERD

>\$2B  
peak sales  
opportunity<sup>2</sup>

Proven R&D and  
commercial capabilities

~\$439M 2015  
cash & investments

No further financings  
expected to fund core business

**Sustainable growth company**

1) As provided by Allergan plc.  
2) Peak U.S. sales estimates by Ironwood

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### Gout

>**\$300M**  
opportunity<sup>2</sup>

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# ZURAMPIC<sup>®</sup> (lesinurad)

**Tom McCourt**

Chief Commercial Officer

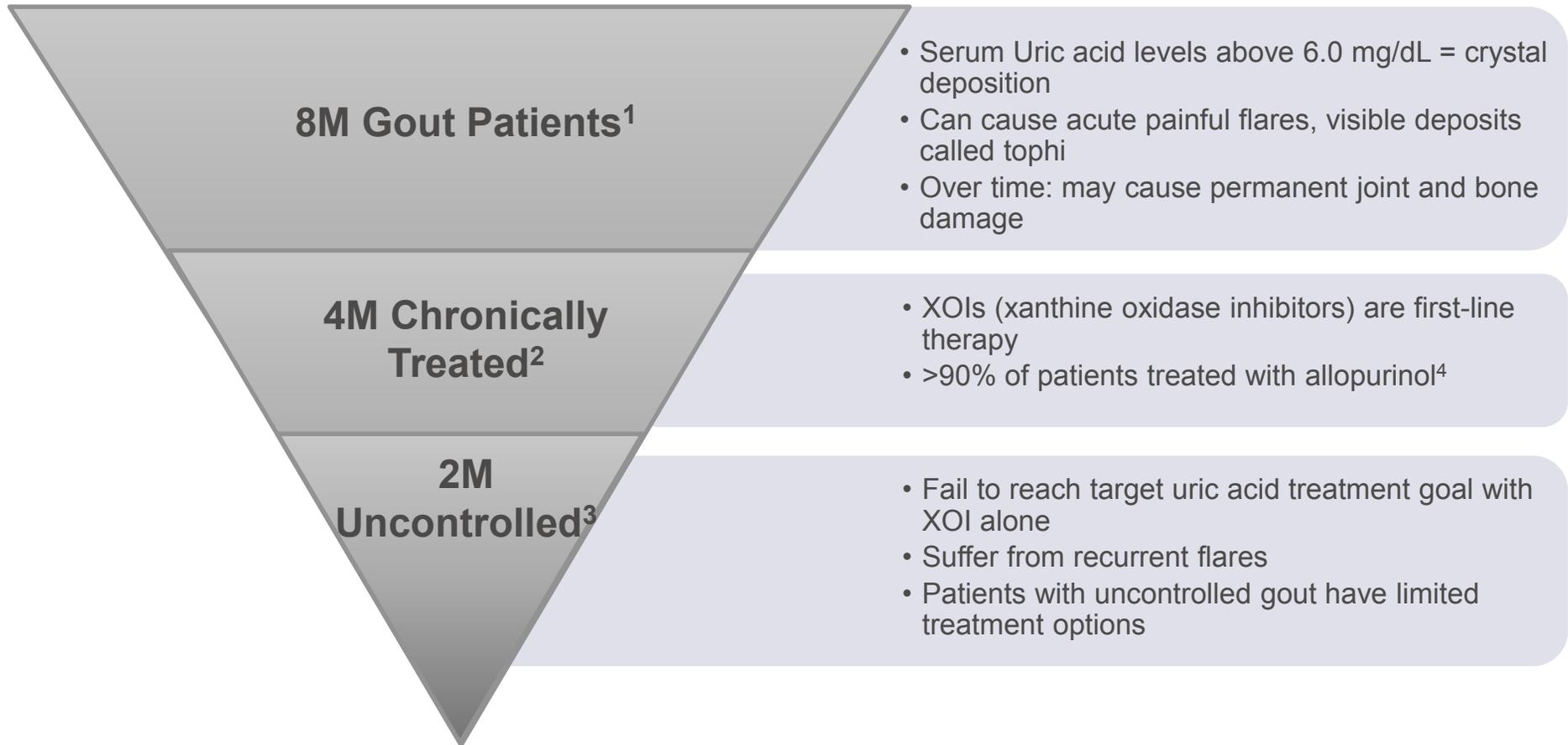


## Great Fit for Our Strong Commercial Capabilities

- **Significant Unmet Need:** 2 million<sup>1</sup> patients continue to suffer from painful flares due to uncontrolled gout despite XOI treatment
- **Innovative Asset:** 1<sup>st</sup> FDA-approved Selective Uric Acid Reabsorption Inhibitor (SURI)
  - ZURAMPIC + XOI demonstrated nearly two-fold increase in patients reaching target serum uric acid levels compared to XOI alone<sup>2</sup>
- **Highly Leverageable:** Proven sales and marketing organization
  - ZURAMPIC and LINZESS provide effective treatment options to prescriber base of 20K-30K physicians
  - Ironwood plans to strengthen sales capability by expanding deeper into primary care; focus on high-prescribers and early-adopters

# 2M Patients Suffer from Uncontrolled Gout in U.S.

Gout is a metabolic disorder caused by hyperuricemia



Sources: 1) NHANES 2007-2008 Zhu Y, et al, *Arthritis Rheum* 2011 (NHANES 07/08) 2) IMS/NHANES; IMS Rx Audit, AZ estimation, DR Gout PharmaCor 2012 3) IMS data, AZ estimation, Zhu et al, *Arthritis Rheum*, 2011, (NHANES 07/08), Becker et al, *NEJM*, 2005, Becker et al., *Arthritis Res Ther* 2010, Schumacher et al, *Arthritis Rheum* 2008 2) ZURAMPIC Prescribing Information 4) IMS/Data on file

# ZURAMPIC: First FDA-approved Selective Uric Acid Reabsorption Inhibitor (SURI)

- ZURAMPIC is indicated in combination with XOI for hyperuricemia associated with uncontrolled gout patients who have not achieve targeted serum uric levels with an XOI alone
- ZURAMPIC is not recommended for the treatment of asymptomatic hyperuricemia and should not be used as monotherapy

## Efficacy established in three Phase III trials of ZURAMPIC 200 mg + XOI versus XOI alone

- Demonstrated ability to lower serum uric acid levels; nearly doubling number of patients to target

## Boxed warning

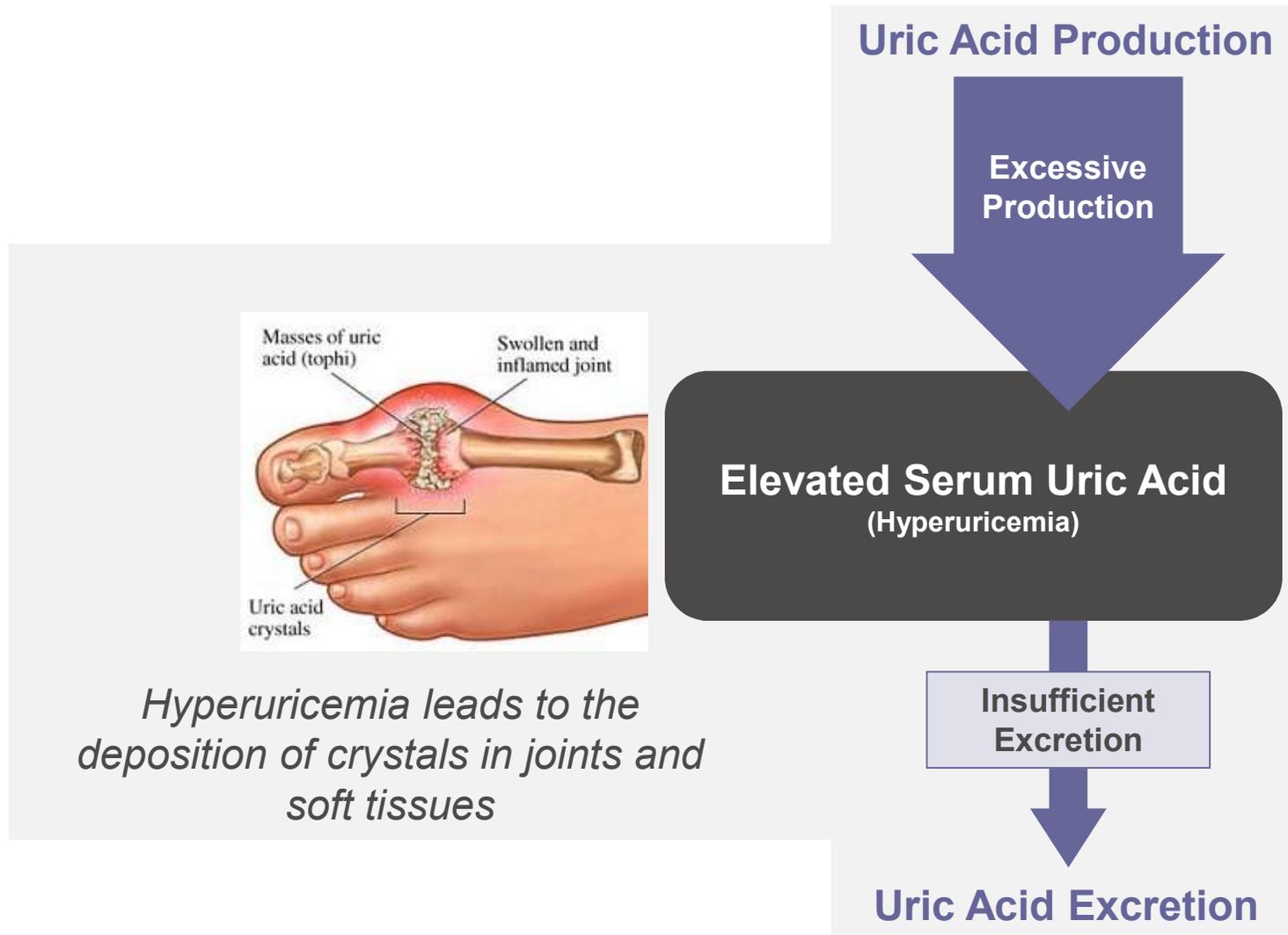
- Risk of acute renal failure has occurred with ZURAMPIC and was more common when ZURAMPIC was given alone
- ZURAMPIC should be used in combination with an XOI

## Adverse events

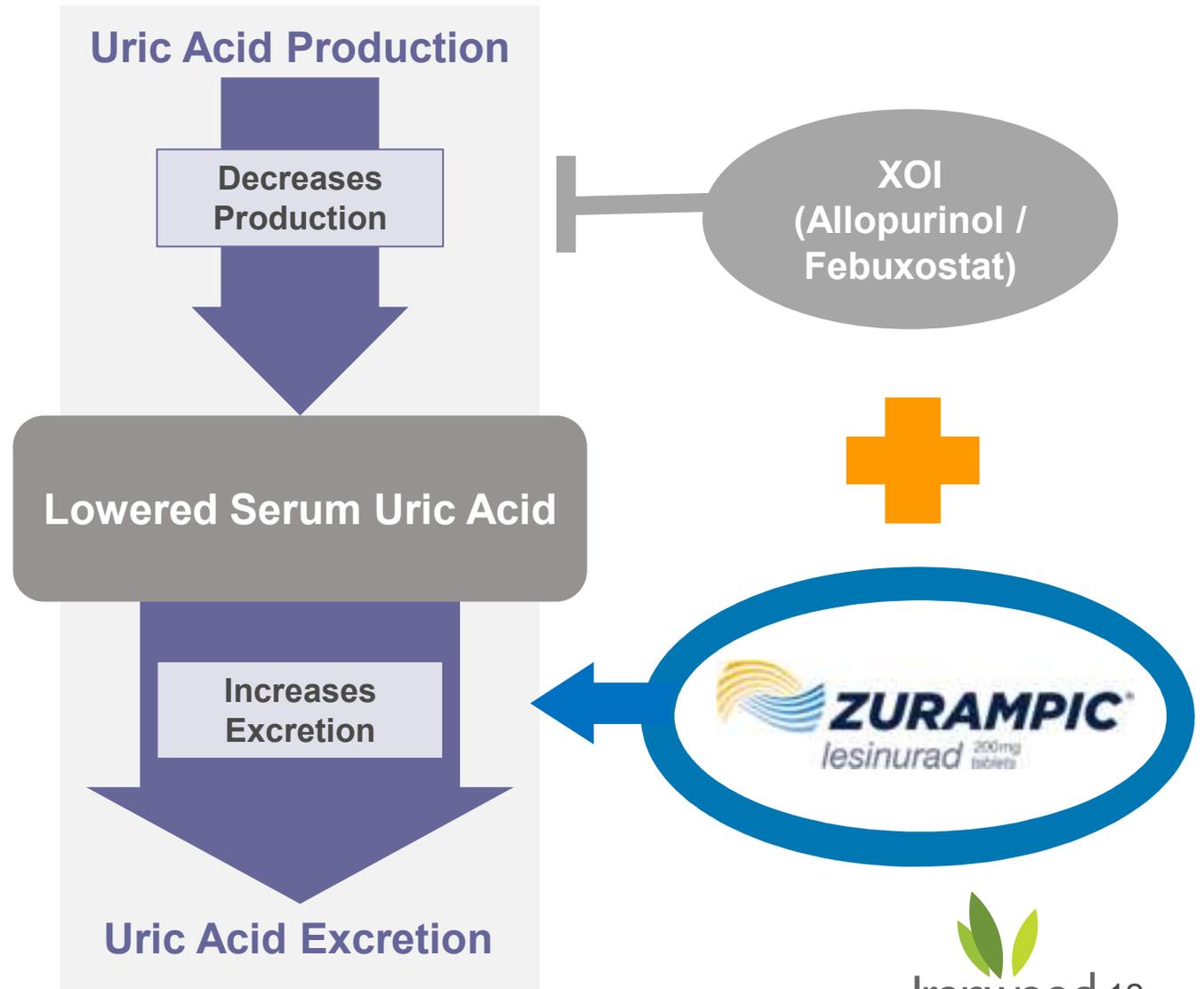
- Elevations in serum creatinine have been observed and are more common in patients with impaired kidney function
- Most common adverse events reported were headache, influenza, serum creatinine increase and gastroesophageal reflux disease



# Hyperuricemia is caused by both overproduction and inefficient excretion of uric acid



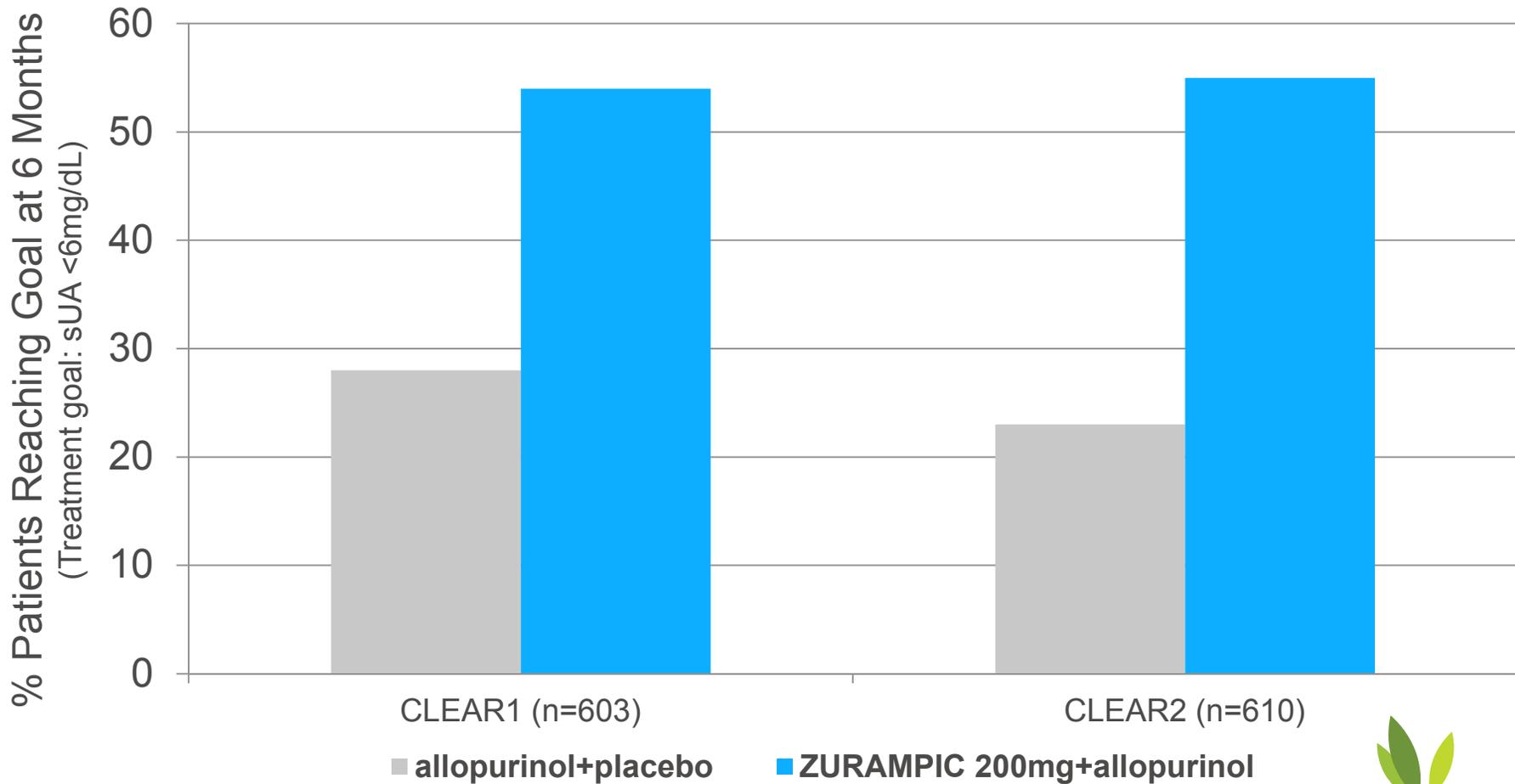
# ZURAMPIC + XO1: Complementary MOA to treat both causes of hyperuricemia in uncontrolled patients



# ZURAMPIC + Allopurinol Enabled More Patients Than Allopurinol Alone to Reach Treatment Goal

--Nearly 2X as many patients reached goal with ZURAMPIC + allopurinol--

## Proportion of Patients Achieving Target Serum Uric Acid Levels



Sources: ZURAMPIC Prescribing Information

# Key Forces to Drive ZURAMPIC Growth in Patients with Uncontrolled Gout



**MD URGENCY** to improve treatment for uncontrolled patients

- Recognize a high level of patient suffering
- Acknowledge need for additional treatment options
- Report high willingness to try ZURAMPIC



Uncontrolled patients desire more effective treatment

- Suffer recurrent painful flares
- Majority of patients not treated to sUA goal
- Actively seeking better treatment option



Payers recognize unmet need in uncontrolled gout

- Reimburse for all current treatment options
- ZURAMPIC provides strong value proposition
- IRWD objective is to provide broad access

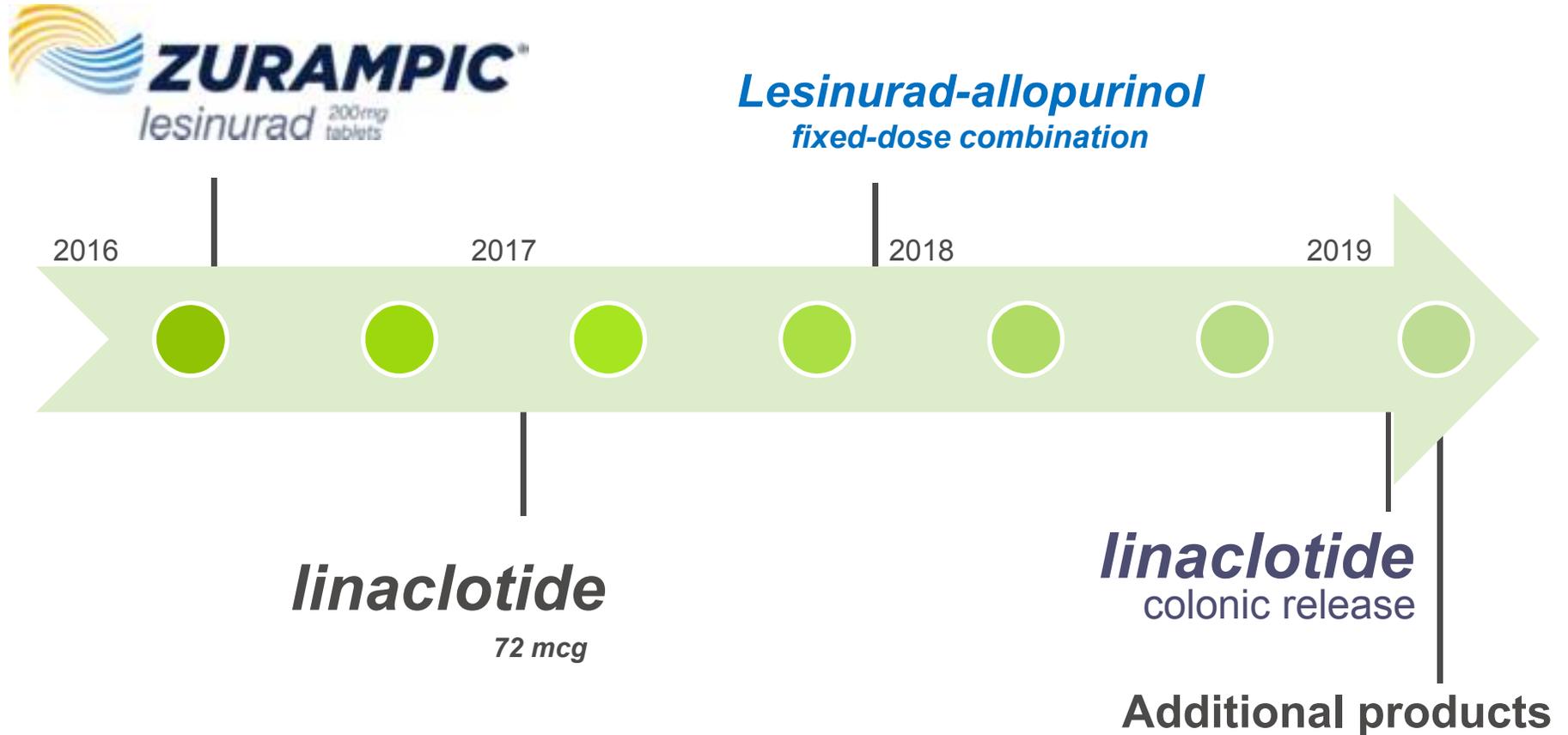
ZURAMPIC offers **COMPELLING** value to all customers

# ZURAMPIC: The Path Forward

- **ZURAMPIC launch expected mid-2H 2016: expect >\$300M peak annual net sales**
  - Targeting 2M<sup>1</sup> uncontrolled gout patients in the U.S. easily identified
  - High performing sales team will call on 20-30K high-volume, early-adopting prescribers with: LINZESS, ZURAMPIC and our existing co-promotes
    - Targets prescribers estimated to generate >65% of TRxs over first 12-18 months
  - Patients with uncontrolled disease are actively seeking better treatment
    - Opportunity to access, educate and activate patient through online promotion
- **Lesinurad-allopurinol fixed-dose combination**
  - Regulatory submission expected 2H16
- **Focused commercial investment to maximize cash flows**
  - Initial incremental commercial expenses expected to be <\$75M/yr

# Expect Five Launches by 2020

*Reflects current expectations, assuming FDA approval of investigational products*





**A COMMERCIAL BIOTECHNOLOGY COMPANY**