

# **Corporate Overview**

February 27, 2019

© 2019 Jazz Pharmaceuticals all rights reserved.



# **Forward-Looking Statements**

"Safe Harbor" Statement Under the Private Securities Litigation Reform Act of 1995

This slide deck and the accompanying oral presentation contain forward-looking statements, including, but not limited to, statements related to Jazz Pharmaceuticals' future financial and operating results, including 2019 financial guidance and goals; the company's corporate development efforts; the company's growth strategy and expectations for growth; future product sales and volume; planned sales and marketing and related efforts; planned, ongoing and future clinical trials and other product development activities, including clinical trials, data read-outs and pre-clinical activities; regulatory events, such as the potential approvals of the NDA and the MAA for solriamfetol and additional planned regulatory submissions; future product launches; the timing of such events and activities; and other statements that are not historical facts. These forward-looking statements are based on the company's current plans, objectives, estimates, expectations and intentions and inherently involve significant risks and uncertainties. Actual results and the timing of events could differ materially from those anticipated in such forward-looking statements as a result of these risks and uncertainties, which include, without limitation, risks and uncertainties associated with: maintaining or increasing sales of and revenue from Xyrem<sup>®</sup> (sodium oxybate) oral solution; effectively commercializing the company's other products and product candidates; the time-consuming and uncertain regulatory approval process, including the risk that the company's regulatory submissions, including the NDA and the MAA for solriamfetol, may not be approved by applicable regulatory authorities in a timely manner or at all; pharmaceutical product development and the uncertainty of clinical success, including risks related to failure or delays in initiating or completing clinical trials; protecting and enhancing the company's intellectual property rights; delays or problems in the supply or manufacture of the company's products and product candidates; complying with applicable U.S. and non-U.S. regulatory requirements; government investigations and other actions, including the risk that the company may not ultimately reach a final settlement with the U.S. Department of Justice to resolve an investigation relating to the company's support of 501(c)(3) organizations that provide financial assistance to Medicare patients; obtaining and maintaining appropriate pricing and reimbursement for the company's products; identifying and acquiring, in-licensing or developing additional products or product candidates, financing these transactions and successfully integrating acquired businesses; the ability to achieve expected future financial performance and results and the uncertainty of future tax and other provisions and estimates; and other risks and uncertainties affecting the company, including those described from time to time under the caption "Risk Factors" and elsewhere in Jazz Pharmaceuticals plc's Securities and Exchange Commission filings and reports (Commission File No. 001-33500), including the company's Annual Report on Form 10-K for the year ended December 31, 2018 and future filings and reports by the company. Other risks and uncertainties of which the company is not currently aware may also affect the company's forward-looking statements and may cause actual results and the timing of events to differ materially from those anticipated. The forward-looking statements made in this slide deck and the accompanying oral presentation are made only as of the date hereof or as of the dates indicated in the forward-looking statements, even if they are subsequently made available by the company on its website or otherwise. The company undertakes no obligation to update or supplement any forward-looking statements to reflect actual results, new information, future events, changes in its expectations or other circumstances that exist after the date as of which the forward-looking statements were made.



# A Strong Foundation To Support Global Growth



# Transforming Investment Opportunity

STRONG FINANCIAL EXECUTION	DIVERSE AND GROWING PIPELINE	DISCIPLINED CAPITAL ALLOCATION TO OPTIMIZE SHAREHOLDER RETURNS
<ul> <li>2018 revenues of \$1.9B</li> <li>Doubling of revenues over past 5 years</li> <li>Non-GAAP adjusted net income CAGR of 17% (2013-2018)<sup>1,4</sup></li> <li>\$3.3B of cash generation (2013-2018)<sup>2</sup></li> </ul>	<ul> <li>Multiple late- and mid-stage assets</li> <li>Growing early-stage opportunities</li> <li>Potential to drive significant growth</li> </ul>	<ul> <li>\$4.4B deployed in corporate development transactions since 2012</li> <li>\$1.1B invested in share repurchases (2013-2018)</li> <li>\$708M in R&amp;D investments<sup>3,4</sup> (2013-2018)</li> </ul>

<sup>1</sup> In 2Q16, Jazz modified the calculation of its non-GAAP income tax provision and effected this modification in the non-GAAP results from 2014 onwards. <sup>2</sup> Net cash provided by operating activities. <sup>3</sup> Non-GAAP adjusted R&D expense for period from January 1, 2013 through December 31, 2018. <sup>4</sup> Reconciliations of GAAP to non-GAAP can be found in the Appendix at the end of this presentation.



### **Corporate Development Alliances**

Acquisitions	Collaboration and Licensing Partnerships	Venture Investments
ORPHAN MEDICAL AZUR PHARMA	MDAnderson Cancer Center	<b>Corrivo</b> BioVentures LLC
EUSA Pharma (2012)	OHARA XL-protein	onera
Sentium Alizé Pharma	Superior Biopharmaceuticals Porton Biopharma Ltd	
Celator <sup>®</sup> Pharmaceuticals	CONCERT Pharmaceuticals Inc.	

) Jazz Pharmaceuticals

### **R&D** Transformation

	PRE-CLINICAL	PHASE 1	PHASE 2	PHASE 3
2013	JZP-386	JZP-416	IV Erwinaze	Leukotac



### **R&D** Transformation

	PRE-CLINICAL	PHASE 1	PHASE 2	PHASE 3		
2013	JZP-386	JZP-416	IV Erwinaze	Leukotac		
2019	Oxybate / once nightly	Vyxeos + gemtuzumab <sup>3</sup> /R/R AML	Solriamfetol / EDS PD	JZP-258 / narcolepsy		
	CombiPlex / solid tumors	Vyxeos + venetoclax <sup>2</sup> / Low intensity unfit AML	Defitelio / paGvHD	JZP-258 / IH		
	CombiPlex / hem/onc	IMGN779 <sup>1</sup> / CD33+ AML	Defitelio / TA-TMA <sup>2</sup>	Defitelio / pVOD		
	Asparaginase / ALL	IMGN632 <sup>1</sup> / CD123+ hem/onc	Defitelio / CAR-T associated neurotoxicity <sup>2</sup>	Vyxeos / AML18 <sup>4</sup>		
	Recombinant pegaspargase <sup>1</sup> / hem/onc		Vyxeos + venetoclax <sup>3</sup> / R/R AML	Vyxeos / AML19 <sup>4</sup>		
	Defitelio / exploratory activities		Vyxeos / MDS <sup>2,4</sup>			
	Exosome NRAS <sup>5</sup> / hem/onc		Vyxeos / R/R AML <sup>4</sup>			
	Exosome STAT3 <sup>5</sup> / hem/onc					
	Exosome candidates <sup>5</sup> / hem/onc	<sup>1</sup> Opt-in opportunity, <sup>2</sup> Planned study, <sup>3</sup> Jazz & MD Anderson collaboration study, <sup>4</sup> Cooperative Group study, <sup>5</sup> Jazz & Codiak collaboration				

# **R&D** Programs



### Defitelio Clinical Development Strategy

Pursue prevention and treatment of serious diseases associated with endothelial cell damage

PREVENTION OF VOD	PREVENTION OF aGvHD	TREATMENT OF TA-TMA	PREVENTION OF CAR-T ASSOCIATED NEUROTOXICITY
<ul> <li>Phase 3, randomized, open- label, multi-center study in high-risk patients vs best supportive care</li> <li>Interim analysis in 2019 to determine final enrollment goal of 400 or up to 600 patients</li> </ul>	<ul> <li>Phase 2 proof of concept</li> <li>FPI 1Q18</li> <li>N = 150</li> </ul>	<ul> <li>Phase 2 single arm, open label study in high-risk patients</li> <li>Expect to activate study sites in 2019</li> <li>N ~40</li> </ul>	<ul> <li>Expect to initiate exploratory Phase 2 in 2019</li> <li>N ~35</li> </ul>



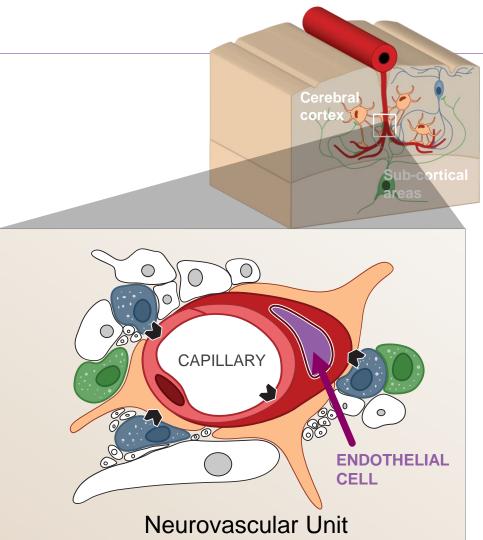
# Rationale for Prevention of CAR-T Associated Neurotoxicity with Defitelio

### Pathophysiology of CAR-T Associated Neurotoxicity

- Endothelial cell damage leading to BBB disruption seems to play a role in CAR-T associated neurotoxicity, as clinical evidence reports:
  - Diffusion of cytokines into the CSF
  - Trafficking of T-cells into the CNS
  - Serum endothelial biomarker elevations (vWF and ANG2)

### **Rationale for Prevention with Defitelio**

- Endothelial cell damage can start with lymphodepletion prior to CAR-T infusion
- The BBB consists of <u>endothelial</u> cells, pericytes and astrocytes
- Defibrotide may prevent the initiation and/or progression of damage to these "gatekeepers" thereby minimizing the cytokine surge





# Vyxeos Clinical Development Strategy

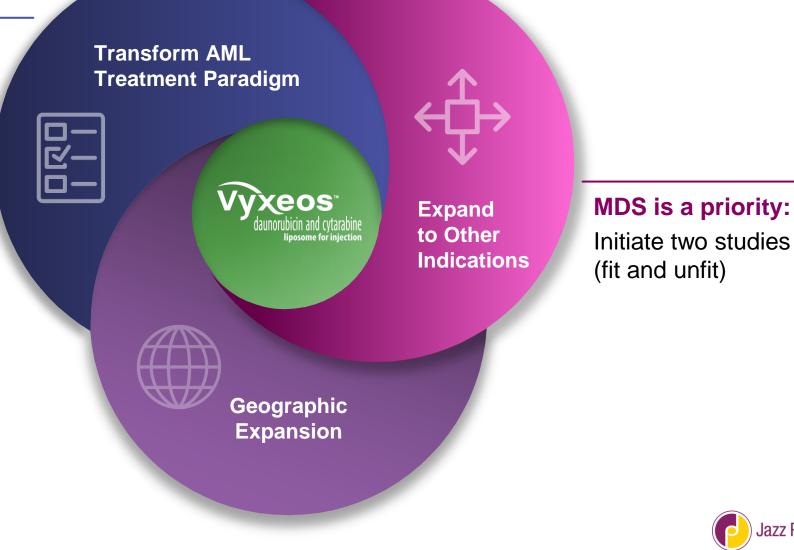
Become the chemo backbone for AML

Target new patient segments across AML landscape: Prolong survival and increase potential of curative intent

### Pursue broad combinations approach: Synchronizing Jazz studies, collaborations and ISTs to generate clinical data

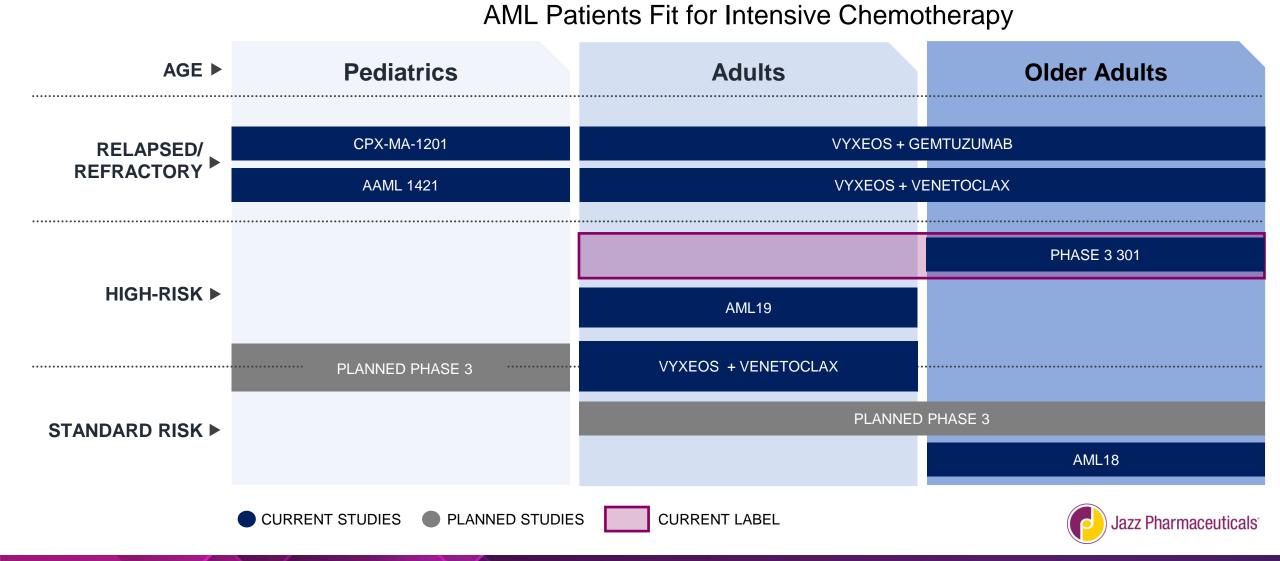
**Extend clinical benefits:** 

Lower intensity and higher intensity dose/schedule, MRD, genetics/mutations



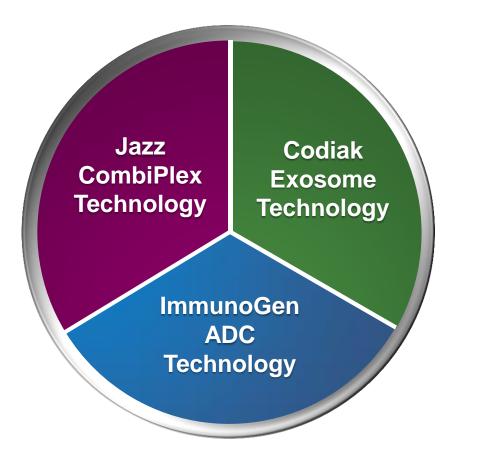


### Vyxeos Clinical Development Program in Fit AML Current and Planned Studies



## Jazz Precision Oncology Strategy

Leverage Multiple Technologies to Potentially Improve Therapeutic Index of Cancer Agents



Precision oncology therapeutics may result in better outcomes for targeted subgroups of patients than nontargeted approaches<sup>1</sup>

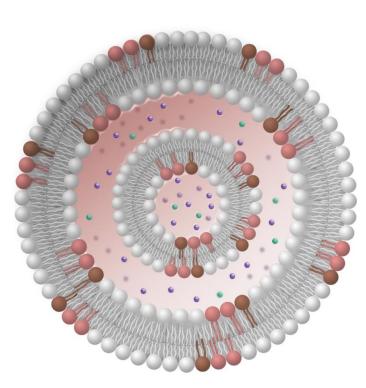
Jazz Pharmaceuticals

<sup>1</sup> JAMA Oncol. 2018;4(2):210-216. doi:10.1001/jamaoncol.2017.4427

### **CombiPlex Strategy**

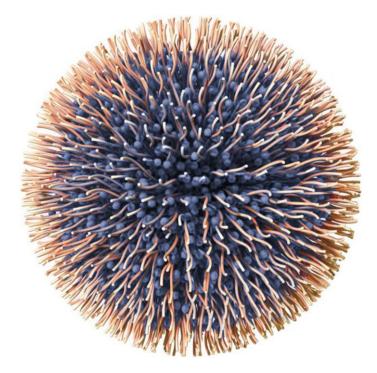
# Transform Patient Outcomes with the Innovative CombiPlex Delivery Technology

### LIPOSOME DELIVERY PLATFORM Delivery of water soluble drugs



### NANOPARTICLE DELIVERY PLATFORM

Delivery of hydrophobic drugs in polymer core



### Solid Tumors Candidate I

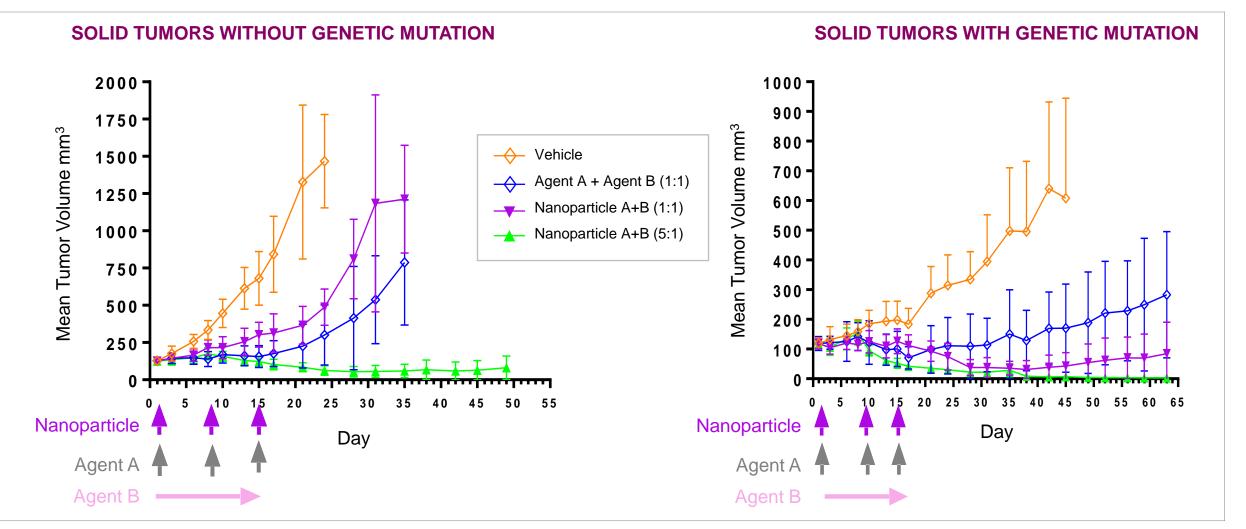
- Novel combination
- IND-enabling activities
   ongoing

### Hematology/Oncology Opportunities

 Exploratory pre-IND enabling activities ongoing



# CombiPlex Nanoparticles Generate Robust Anti-Tumor Activity in Murine Xenografts





### ImmunoGen Collaboration



### **IMGN779**

- Anti-CD33 ADC
- First ADC to employ an IGN, a new type of cancer-killing agent (DGN462)
- Potential indications: AML
- Received orphan drug designation by FDA for the treatment of AML in May 2018
- Phase 1 data presented at ASH 2018



### IMGN632

- Anti-CD123 ADC
- Employs a novel IGN cancer-killing agent (DGN549)
- Potential indications: hematological malignancies, including AML and BPDCN
- Received orphan drug designation by FDA for the treatment of AML in October 2018
- Phase 1 data presented at ASH 2018

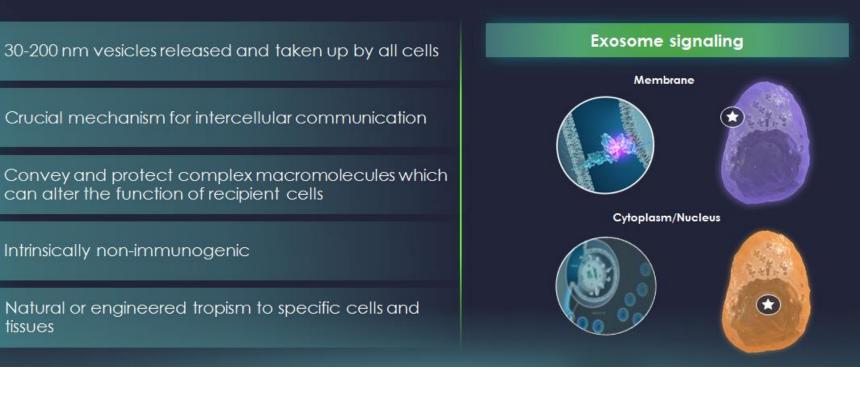
- Collaboration and option agreement
- Worldwide rights to develop and commercialize three ADC programs
  - IMGN779
  - IMGN632
  - Undisclosed program
- May exercise opt-in prior to a pivotal study or prior to a BLA
- ImmunoGen responsible for development up to Jazz opt-in



### Codiak Collaboration

tissues

Research collaboration and license agreement to develop exosome-based oncology or hematology/oncology therapeutics



- Exosome platform can be targeted broadly throughout the body
- Platform to control the signals • on the surface of, or within, the exosome with the goal of delivering precise signals to promote a therapeutic effect
- Jazz has rights to five targets
  - NRAS
  - STAT3
  - 2 Undisclosed
  - One to be selected at later date
- Codiak to fund early stage research activities for all five targets
- Jazz to fund IND-enabling and • Phase 1/2 studies for three targets
- Codiak to fund IND-enabling and Phase 1/2 studies for two targets

Jazz Pharmaceuticals

# Growing R&D Pipeline

PRE-CLINICAL	PHASE 1	PHASE 2	PHASE 3	REGULATORY
Oxybate Once Nightly Formulation	Vyxeos + gemtuzumab <sup>3</sup> R/R AML or HMA Failure MDS	Solriamfetol EDS PD	JZP-258 Cataplexy & EDS in narcolepsy	Solriamfetol U.S. EDS in OSA and Narcolepsy
CombiPlex Solid tumors candidate I	Vyxeos + venetoclax <sup>4</sup> Low Intensity Dosing for unfit AML	Defitelio Prevention of aGvHD	JZP-258 Idiopathic hypersomnia	Solriamfetol EU EDS in OSA and Narcolepsy
CombiPlex Hem/Onc exploratory activities	IMGN779 <sup>1</sup> CD33+ AML	Defitelio Treatment of TA-TMA <sup>4</sup>	Defitelio Prevention of VOD	
Asparaginase ALL/other hematological malignancies	IMGN632 <sup>1</sup> CD123+ Hematological malignancies <sup>5</sup>	Defitelio Prevention of CAR-T associated neurotoxicity <sup>4</sup>	Vyxeos AML or HR-MDS (AML19) <sup>6</sup>	
Recomb. Pegaspargase <sup>1</sup> Hematological malignancies		Vyxeos + venetoclax <sup>3</sup> <i>de novo</i> or R/R AML	Vyxeos AML or HR-MDS (AML18) <sup>6</sup>	
Defitelio Exploratory activities		Vyxeos MDS <sup>4,6</sup>		
Exosome NRAS candidate <sup>2</sup> Hematological malignancies		Vyxeos R/R AML <sup>6</sup>		
Exosome STAT3 candidate <sup>2</sup> Hematological malignancies			-	
Exosome-based candidates <sup>2</sup> Solid tumors/Hematological malignancies				SLEEP HEMATOLOGY/ONCOLOGY



<sup>1</sup> Opt-in opportunity, <sup>2</sup> Jazz & Codiak collaboration, <sup>3</sup> Jazz & MD Anderson Cancer Center collaboration study, <sup>4</sup> Planned , <sup>5</sup> Including AML and BPDCN, <sup>6</sup> Cooperative group study

# Sleep R&D Clinical Development and Regulatory Strategy

### Continued investment to deliver therapeutic options for unmet medical needs in sleep medicine

JZP-258 EDS & Cataplexy for Narcolepsy	JZP-258 Idiopathic Hypersomnia	XYREM EDS & Cataplexy for Pediatric Narcolepsy	SOLRIAMFETOL EDS for Narcolepsy/OSA	SOLRIAMFETOL EDS For PD
<ul> <li>Expect top-line data in spring 2019</li> </ul>	<ul> <li>Initiated Phase 3 study in idiopathic hypersomnia in 4Q18</li> </ul>	<ul> <li>FDA Approval 4Q18</li> <li>Expect to launch 1Q19</li> </ul>	<ul> <li>MAA submission November 2018</li> <li>FDA PDUFA goal date: March 20, 2019</li> <li>Expect to launch mid-2019</li> </ul>	<ul> <li>Completed enrollment 3Q18 in Phase 2 proof of concept study</li> <li>Phase 2 data at AAN (May 2019)</li> </ul>

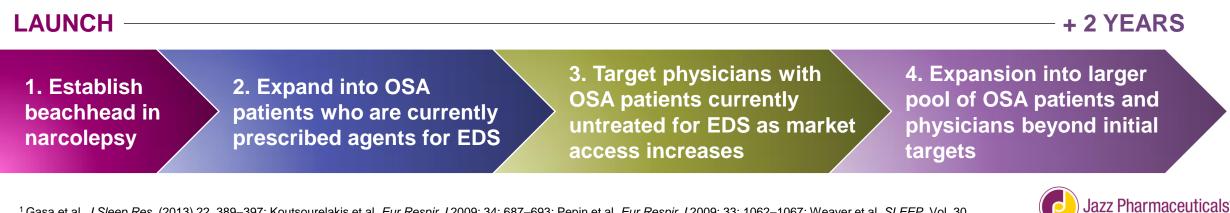


# Solriamfetol Planned Launch Strategy

Building a market over time

### **Key Considerations**

- OSA large market opportunity, but need to build market as <10% are drug treated with wake-promoting agents
- 7% of all physicians who diagnose and manage OSA patients cover 60% of the OSA population and over 40% of drug treated OSA patients
- The prevalence of EDS is high in OSA, even in CPAP-compliant patients, ranging from 12%-65%<sup>1</sup>
- Generic wake-promoting agents available
- Consolidation in the payer industry has concentrated the majority of commercial lives with a few payers/PBMs
- Open formularies have declined significantly; ~15% of commercial lives in open formularies during initial launch phase<sup>2</sup>
- Retail pharmacy non specialty (electronic adjudication)
- Co-pay coupon program/patient assistance program



<sup>1</sup> Gasa et al, J Sleep Res. (2013) 22, 389–397; Koutsourelakis et al, Eur Respir J 2009; 34: 687–693; Pepin et al, Eur Respir J 2009; 33: 1062–1067; Weaver et al, SLEEP, Vol. 30, No. 6, 2007, <sup>2</sup> McKinsey September 2017



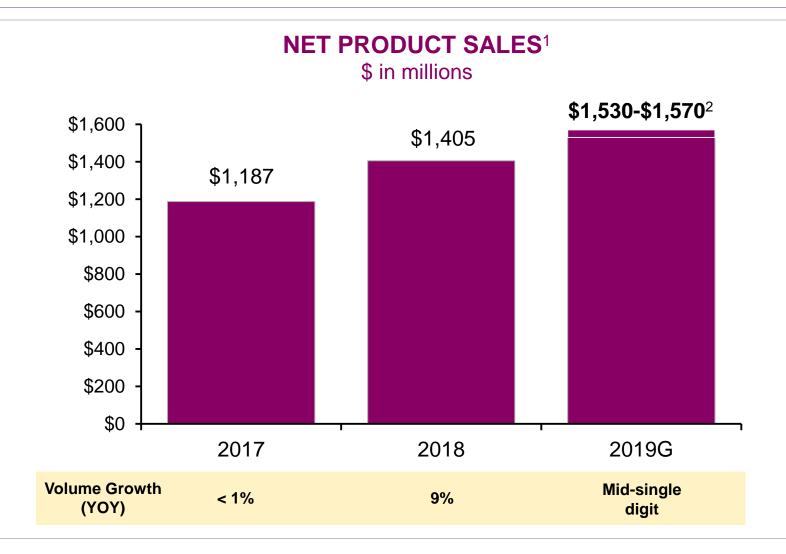
# **Commercial Portfolio**



Jazz Pharmaceuticals

John AML patient



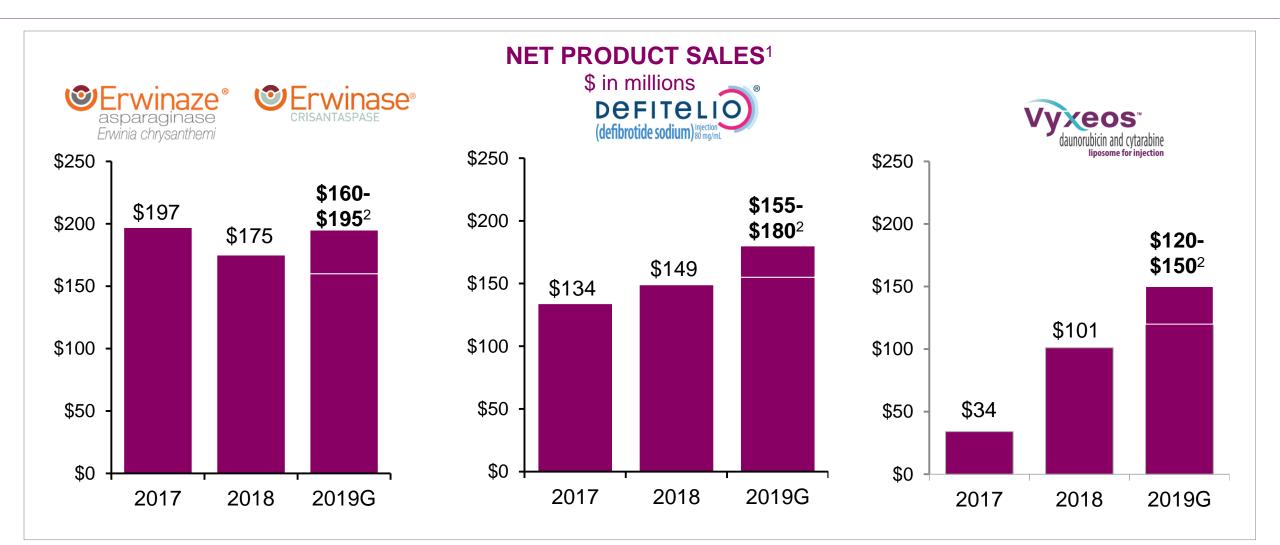


### <sup>1</sup> 2017 and 2018 audited. <sup>2</sup> G=Guidance. Guidance provided by Jazz Pharmaceuticals plc on and as of February 26, 2019. Jazz Pharmaceuticals plc is not confirming or updating that guidance and actual results may differ.

- FDA-approved for the treatment of cataplexy and EDS in narcolepsy patients ≥ 7 years of age
- Disease awareness efforts to improve narcolepsy awareness and diagnosis rates
- Received pediatric exclusivity in October 2018 - expect to launch pediatric indication 1Q19



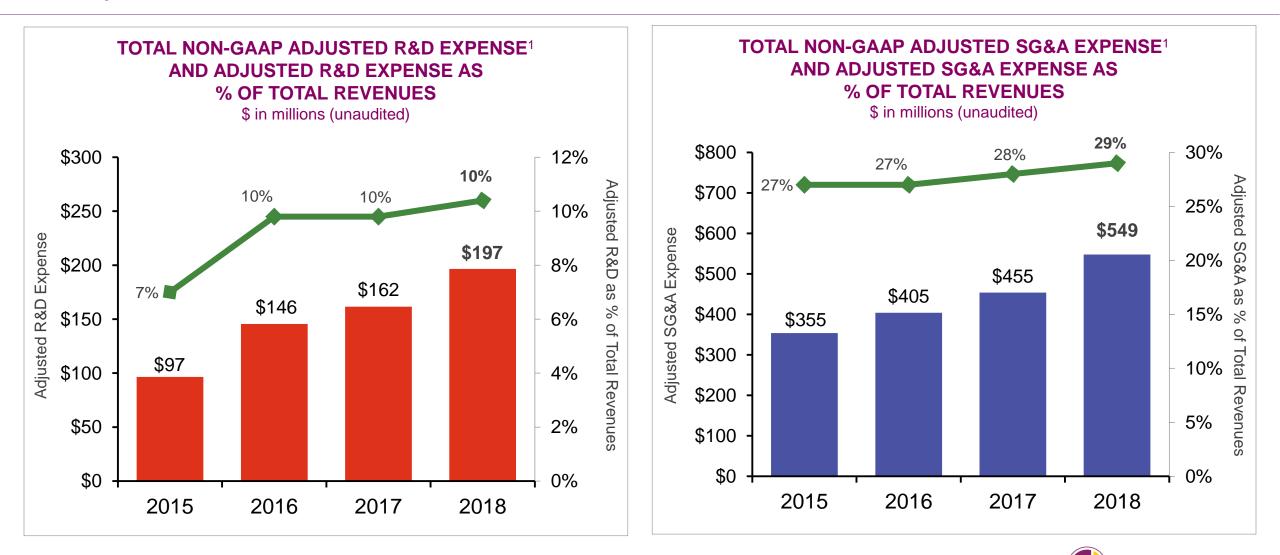
## Hematology/Oncology Revenues



<sup>1</sup> 2017 and 2018 audited. <sup>2</sup> G=Guidance. Guidance provided by Jazz Pharmaceuticals plc on and as of February 26, 2019. Jazz Pharmaceuticals plc is not confirming or updating that guidance and actual results may differ.



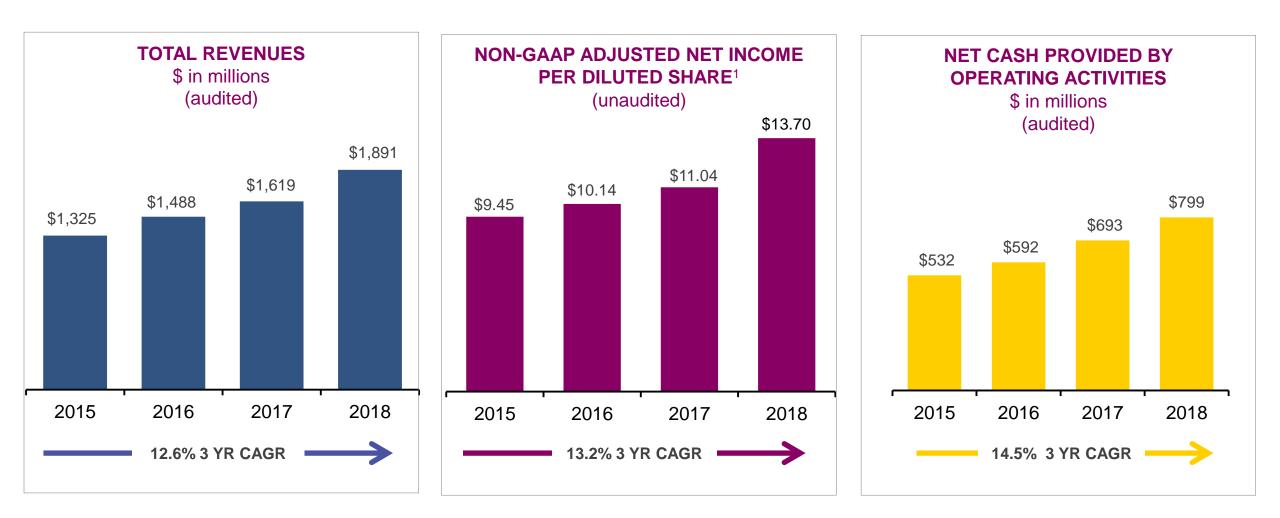
### Disciplined Resource Allocation to Fuel R&D Pipeline and Support Multiple Product Launches



<sup>1</sup> Reconciliations of GAAP to non-GAAP can be found in the Appendix at the end of this presentation.

Jazz Pharmaceuticals

## **Strong Financial Execution**





<sup>1</sup> Reconciliations of GAAP to non-GAAP can be found in the Appendix at the end of this presentation.

### 2019 Goals

#### Xyrem

- Launch pediatric narcolepsy indication
- Deliver mid-single digit volume growth for 2019

#### JZP-258

• Deliver top-line Phase 3 narcolepsy data spring 2019

### **Solriamfetol**

- Gain U.S. approval by PDUFA goal date of March 20, 2019
- Obtain DEA scheduling decision in 2Q19
- Launch in U.S. for EDS in narcolepsy and OSA
- Deliver top-line data for EDS for PD in early 2019
- Gain EU approval for EDS in narcolepsy and OSA in 4Q19

### CombiPlex

 Continue IND-enabling activities for one solid tumor combination and progress exploratory activities for other candidates

#### Vyxeos

- Obtain data read-outs
  - COG R/R pediatric AML study data
  - Interim combination data from MD Anderson collaboration
- Commence Jazz-sponsored combination studies
  - AML combination study with multiple targeted agents
  - Lower intensity dosing study for unfit AML
- Initiate studies in MDS (fit and unfit)
- Continue collaborations with key cooperative groups for Phase 3 studies

#### Defitelio

- Conduct interim analysis for prevention of VOD Phase 3 study
- Complete enrollment in prevention of aGvHD Phase 2 study
- Initiate Phase 2 study in TA-TMA
- Initiate exploratory Phase 2 study in CAR-T associated neurotoxicity

#### **Corporate Development**

• Expand our commercial and/or development portfolio through multiple acquisitions or partnerships



# Appendix

Eve VOD patient



### **Non-GAAP Financial Measures**

To supplement Jazz Pharmaceuticals' financial results and guidance presented in accordance with U.S. GAAP, the company uses certain non-GAAP (also referred to as adjusted or non-GAAP adjusted) financial measures in this presentation and the accompanying tables. In particular, the company presents non-GAAP adjusted net income (and the related per share measure) and its line item components, as well as certain non-GAAP adjusted financial measures derived therefrom. Non-GAAP adjusted net income (and the related per share measure) and its line item components exclude from reported GAAP net income (and the related per share measure) and its line item components exclude from reported GAAP net income (and the related per share measure) and its line item components exclude from reported GAAP net income (and the related per share measure) and its line item components exclude from reported GAAP net income (and the related per share measure) and its line item components exclude from reported GAAP net income (and the related per share measure) and its line item components exclude from reported GAAP net income (and the related per share measure) and its line item components exclude from reported GAAP net income (and the related per share measure) and its line item components exclude from reported GAAP net income (and the related per share measure) and its line item components exclude from reported GAAP adjusted net income (and the related per share measure), adjust for the income tax effect of non-GAAP adjustments, and the U.S. Tax Cut and Jobs Act impact. In this regard, the components of non-GAAP adjusted net income are income statement line items prepared on the same basis as, and therefore components of, the overall non-GAAP adjusted net income measure.

The company believes that each of these non-GAAP financial measures provides useful supplementary information to, and facilitates additional analysis by, investors and analysts. In particular, the company believes that each of these non-GAAP financial measures, when considered together with the company's financial information prepared in accordance with GAAP, can enhance investors' and analysts' ability to meaningfully compare the company's results from period to period and to its forward-looking guidance, and to identify operating trends in the company's business. In addition, these non-GAAP financial measures are regularly used by investors and analysts to model and track the company's financial performance. Jazz Pharmaceuticals' management also regularly uses these non-GAAP financial measures internally to understand, manage and evaluate the company's business and to make operating decisions, and compensation of executives is based in part on certain of these non-GAAP financial measures are important internal measurements for Jazz Pharmaceuticals' management, the company also believes that these non-GAAP financial measures are useful to investors and analysts since these measures allow for greater transparency with respect to key financial metrics the company uses in assessing its own operating performance and making operating decisions.

These non-GAAP financial measures are not meant to be considered in isolation or as a substitute for comparable GAAP measures; should be read in conjunction with the company's consolidated financial statements prepared in accordance with GAAP; have no standardized meaning prescribed by GAAP; and are not prepared under any comprehensive set of accounting rules or principles. In addition, from time to time in the future there may be other items that the company may exclude for purposes of its non-GAAP financial measures; and the company has ceased, and may in the future cease, to exclude items that it has historically excluded for purposes of its non-GAAP financial measures. Likewise, the company may determine to modify the nature of its adjustments to arrive at its non-GAAP financial measures. Because of the non-standardized definitions of non-GAAP financial measures, the non-GAAP financial measures as used by Jazz Pharmaceuticals in this presentation and the accompanying tables have limits in their usefulness to investors and may be calculated differently from, and therefore may not be directly comparable to, similarly titled measures used by other companies.



### Reconciliation of GAAP to Non-GAAP Adjusted Net Income

In millions, except per share amounts (unaudited)	2013	2014	2015	2016	2017	2018	2019 Financial Guidance <sup>3</sup>
GAAP net income <sup>1</sup>	\$ 216.3	\$ 58.4	\$ 329.5	\$ 396.8	\$ 487.8	\$ 447.1	\$395 - \$495
Intangible asset amortization	79.0	126.6	98.2	102.0	152.1	201.5	250 - 270
Share-based compensation expense	44.6	69.6	91.6	98.8	106.9	102.4	110 - 125
Estimated loss contingency						57.0	
Impairment charges and disposal costs		39.4	31.5			44.0	
Upfront and milestone payments	5.0	202.6	25.0	23.8	101.5	11.0	56 - 90
Transaction and integration related costs	6.2	28.8	18.2	13.6			
Acquisition accounting inventory fair value step-up adjustments	3.8	10.5					
Changes in fair value of contingent consideration	15.2						
Expenses related to certain legal proceedings and restructuring	1.5	1.9	1.6	6.1	6.0		
Non-cash interest expense	4.6	13.7	22.7	22.1	30.0	44.0	40 - 50
Loss on extinguishment and modification of debt	3.7		16.8	0.6			
Income tax effect of above adjustments <sup>2</sup>	5.3	(53.8)	(39.6)	(36.7)	(58.8)	(60.9)	(75) - (95)
U.S. Tax Cuts and Jobs Act impact					(148.8)	(7.5)	
Amount attributable to noncontrolling interests		(1.5)					
Non-GAAP adjusted net income <sup>2</sup>	\$ 385.2	\$ 496.3	\$ 595.5	\$ 627.2	\$ 676.7	\$ 838.6	\$835 - \$875
GAAP net income per diluted share <sup>1</sup>	\$ 3.51	\$ 0.93	\$ 5.23	\$ 6.41	\$ 7.96	\$ 7.30	\$6.80 - \$8.50
Non-GAAP adjusted net income per diluted share <sup>2</sup>	\$ 6.26	\$ 7.93	\$ 9.45	\$ 10.14	\$ 11.04	\$ 13.70	\$14.30 - \$15.00
Weighted-average ordinary shares used in diluted per share calculation	61.6	62.6	63.0	61.9	61.3	61.2	58

#### Note: Amounts may not total due to rounding.

<sup>1</sup>2013, 2014, 2015, 2016, 2017 and 2018 audited. <sup>2</sup> In 2Q16 Jazz modified the calculation of its non-GAAP income tax provision and effected this modification in the non-GAAP results from 2014 onwards. <sup>3</sup> Guidance provided by Jazz Pharmaceuticals plc on and as of February 26, 2019. Jazz Pharmaceuticals plc is not confirming or updating that guidance and actual results may differ.



### Reconciliation of GAAP R&D to Non-GAAP Adjusted R&D Expense

In millions (unaudited)	2013	2014	2015	2016	2017	2018
GAAP R&D expense <sup>1</sup>	\$41.6	\$85.2	\$135.3	\$162.3	\$198.4	\$226.6
Share-based compensation expense	(6.7)	(12.2)	(13.4)	(15.3)	(17.9)	(19.0)
Transaction and integration related costs	(0.6)	(1.2)		(0.5)		
Upfront and milestone payments			(25.0)		(18.5)	(11.0)
Restructuring charges			(0.2)			
Non-GAAP adjusted R&D expense	\$34.3	\$71.8	\$96.7	\$146.5	\$162.1	\$196.6



In millions (unaudited)	2015	2016	2017	2018
GAAP SG&A expense <sup>1</sup>	\$449.1	\$502.9	\$544.2	\$683.5
Share-based compensation expense	(74.7)	(79.0)	(83.2)	(76.8)
Estimated loss contingency				(57.0)
Expenses related to certain legal proceedings and restructuring	(0.9)	(6.0)	(6.0)	(6.0)
Transaction and integration related costs	(18.2)	(13.1)		
Disposal costs				(1.1)
Non-GAAP adjusted SG&A expense	\$355.4	\$404.8	\$454.9	\$548.7



### **Glossary of Abbreviations**

AAML = COG AML Study Identifier AAN = American Academy of Neurology ADC = Antibody Drug Conjugate aGvHD = Acute Graft vs Host Disease ALL = Acute Lymphoblastic Leukemia AML = Acute Myeloid Leukemia ANG2 = Angiopoietin-2 ASH = American Society of Hematology BBB = Blood Brain Barrier **BLA = Biologics License Application** BPDCN = Blastic Plasmacytoid Dendritic Cell Neoplasm CAGR = Compounded Annual Growth Rate CAR-T = Chimeric Antigen Receptor T-cell Therapy CNS = Central Nervous System COG = Children's Oncology Group CPAP = Continuous Positive Airway Pressure CPX-MA-1201 = University of Cincinnati Vyxeos Study Identifier CSF = Cerebrospinal Fluid DEA = U.S. Drug Enforcement Administration EDS = Excessive Daytime Sleepiness EU = European Union FDA = U.S. Food and Drug Administration FPI = First Patient In GAAP = Generally Accepted Accounting Principles GHB = Gamma Hydroxybutyrate HMA = Hypomethylating Agent HR-MDS = High-Risk MDS

IGN = New class of cancer-killing agents (DNA-alkylating indolinobenzodiazepines) developed by ImmunoGen IH = Idiopathic Hypersomnia IMGN = ImmunoGen IND = Investigational New Drug Application IST = Investigator Sponsored Trial LR-MDS = Low-Risk MDS MAA = Marketing Authorization Application MDS = Myelodysplastic Syndrome MOA = Mechanism of Action MRD = Minimal Residual Disease OSA = Obstructive Sleep Apnea paGvHD = Prevention of acute Graft vs Host Disease PBM = Pharmacy Benefit Manager PD = Parkinson's Disease PDUFA = Prescription Drug User Fee Act pVOD = Prevention of VOD R&D = Research & Development REMS = Risk Evaluation Mitigation Strategies R/R = Relapsed/Refractory SG&A = Selling, General & Administrative TA-TMA = Transplant Associated Thrombotic Microangiopathy VOD = Hepatic Veno-occlusive Disease vWF = Von Willebrand Factor YOY = Year Over Year YR = Year



# Xyrem<sup>®</sup> (sodium oxybate) Boxed Warning

#### WARNING: CENTRAL NERVOUS SYSTEM DEPRESSION and ABUSE AND MISUSE.

#### **Central Nervous System Depression**

Xyrem (sodium oxybate) is a CNS depressant. In clinical trials at recommended doses, obtundation and clinically significant respiratory depression occurred in adult patients treated with Xyrem. Many patients who received Xyrem during clinical trials in narcolepsy were receiving central nervous system stimulants.

#### Abuse and Misuse

Xyrem® (sodium oxybate) is the sodium salt of gamma-hydroxybutyrate (GHB). Abuse or misuse of illicit GHB, either alone or in combination with other CNS depressants, is associated with CNS adverse reactions, including seizure, respiratory depression, decreases in the level of consciousness, coma, and death.

Because of the risks of CNS depression and abuse and misuse, Xyrem is available only through a restricted program under a Risk Evaluation and Mitigation Strategy (REMS) called the Xyrem REMS Program. Further information is available at www.XYREMREMS.com or 1-866-XYREM88® (1-866-997- 3688).



### Vyxeos<sup>®</sup> (daunorubicin and cytarabine) liposome for injection Boxed Warning

### WARNING: DO NOT INTERCHANGE WITH OTHER DAUNORUBICIN- AND/OR CYTARABINE-CONTAINING PRODUCTS

VYXEOS has different dosage recommendations than daunorubicin hydrochloride injection, cytarabine injection, daunorubicin citrate liposome injection, and cytarabine liposome injection. Verify drug name and dose prior to preparation and administration to avoid dosing errors (5.1).

