

# HC Wainwright 20<sup>th</sup> Annual Global Investment Conference

September, 2018

NASDAQ: LGND

#### **Safe Harbor Statement**

The following presentation contains forward-looking statements regarding Ligand's prospects, plans and strategies, drug development programs and collaborations. Forward-looking statements include financial projections, expectations regarding research and development programs, and other statements including words such as "will," "should," "could," "plan," etc. Actual events or results may differ from Ligand's expectations. For example, drug development program benefits may not be realized and there can be no assurance that Ligand will achieve its guidance in 2018 or thereafter or that third party research summarized herein is correct or complete.

The forward-looking statements made in the presentation are subject to several risk factors, including, statements regarding intent, belief, or current expectations of the Ligand, its internal and partnered programs, including Promacta™, Kyprolis® and EVOMELA®, Ligand's reliance on collaborative partners for milestone and royalty payments, royalty and other revenue projections based on third party research, regulatory hurdles facing Ligand's and partners' product candidates, uncertainty regarding Ligand's and partners' product development costs, the possibility that Ligand's and partners' drug candidates might not be proved to be safe and efficacious and commercial performance of Ligand's and/or its partners' products, risks related to Ligand's internal controls, its compliance with regulations, accounting principles and public disclosure, and other risks and uncertainties described in its public filings with the Securities and Exchange Commission, available at www.sec.gov. Additional risks may apply to forward-looking statements made in this presentation. Information regarding partnered products and programs comes from information publicly released by our partners. This presentation describes the typical roles and responsibilities of Ligand and our partners, and is not intended to be a complete description in all cases. Our trademarks, trade names and service marks referenced herein include Ligand and Captisol. Each other trademark, trade name or service mark appearing in this presentation belongs to its owner. The process for reconciliation between adjusted financial numbers presented on slide 37, and the corresponding GAAP figures is shown on slide 39.

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#### **Shots-on-Goal Business Model**

#### The "LIGAND MODEL"

- Realities of the pharmaceutical industry
  - Most drug research programs fail, but not all
  - Programs are not all of equal value different time to market, risk, economics
- BUT, the more quality programs you have, the higher likelihood of success
  - Diversified across a full range of industry partners
  - Diversified across a broad spectrum of therapeutic indications
- A shot-on-goal for Ligand is a fully funded partnership
  - Backed by license to Ligand's patents, know-how and/or data
  - Sharing of future economics based on partner's success



#### The "LIGAND MODEL"

#### The Balance in Our Business

#### What We Do:

- Conduct early research, discover drugs
- Provide tools that make drugs possible
- License data and patents
- Acquire new technologies and assets
- Operate with low costs and maintain lean sharecount

#### What Our Partners Do:

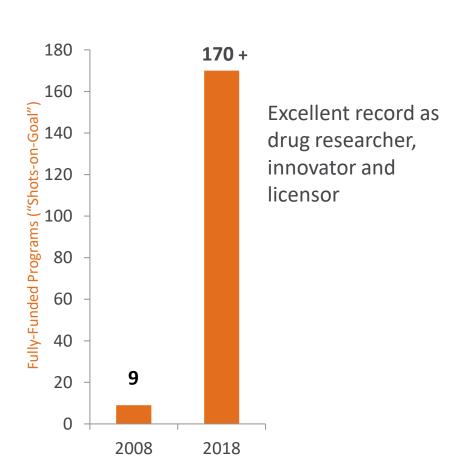
- Decide which indications to pursue
- Design studies; manage regulatory work
- Price drugs and secure reimbursement
- Market drugs
- Fund all development and commercialization



## **Ligand's Portfolio Continues to Grow**

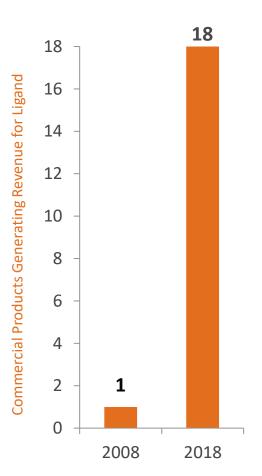
#### **Ligand's Achievement:**

#### **Portfolio Expansion**



#### Partners' Achievement:





Our partners are doing their job getting new products to the market

Latest product
approvals include
Baxdela IV,
Bryxta and
rituximab
biosimilar



## **RPT – Ligand's Foundation of Value**

Revenue

Pipeline

Technology

# **RPT – Ligand's Foundation of Value**

Revenue



High Growth
High Margin
Strong Protection

Pipeline



Large and Growing
High Quality
Many Late Stage

Technology



Best-in-Class
Leverageable
Strong IP

## **Underlying Revenue & Effective Royalty Rate**



- New approvals and increasing sales of existing partnered products are major drivers for underlying revenue growth
- Average royalty rate increasing due to mix of sales and royalty tiering
- 2018 Corporate gross margins projected to be 96%

Current 2018 outlook, underlying revenue will exceed \$2.3 billion and average royalty will exceed 5%

# **RPT – Ligand's Foundation of Value**

Revenue



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**Pipeline** 



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### **Pipeline**

# Why is Ligand's Pipeline Valuable?

- In pharmaceuticals, most programs fail; but not ALL programs
- Ligand's pipeline is:
  - Large and growing
  - Highly diversified
  - Many programs have top-tier sponsorship
- Unique economic structure of Ligand's pipeline:
  - Our deals are fully funded
  - Ligand is not generating big annual losses OR diluting shareholders to finance its pipeline
- Many of Ligand's major assets are still development-stage



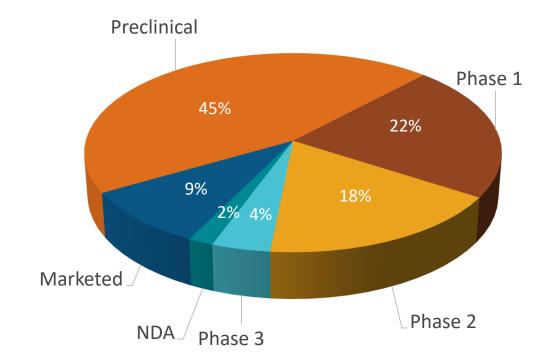
## **Ligand's Portfolio Continues to Grow**

#### **Over 170 Partnered Programs**

 Portfolio remains diversified across development stages

Over 100 different partners

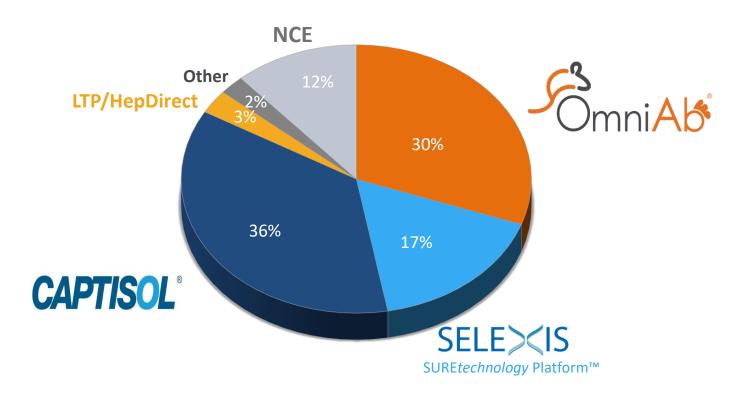
 Nearly 55% of programs in clinical development or later



11% marketed or NDA stage

# **Diversified Underlying Intellectual Property**

Partnered programs are spread across multiple IP families



- The underlying IP of Ligand's portfolio of 170+ partnered programs is spread across multiple, global families of intellectual property
- Between OmniAb and Selexis assets, antibodies/large molecules now make up
   47% of the portfolio



# **Broad Array of Partners Use Our Technologies**

Over 100 Partners Spread Throughout the Pharmaceutical Industry

#### Specialty

<u>Pharma</u>	<u>Biotech</u>		<u>Pharma</u>	
Pfizer	Amgen	Celgene	Eisai	
Novartis	Exelixis	Melinta	Spectrum	
J&J	SAGE	Gilead	Sermonix	
GSK	Viking	Aldeyra	Upsher-Smith	
Merck	MEI	Roivant	Glenmark	
Baxter	Chiva	Genmab	Ono	
BMS	Retrophin	Millennium	Sedor	
Eli Lilly	Merrimack	Meridian	CURx	
Takeda	Glenmark	C-Stone		
Otsuka	VentiRx	Marinus		
Daiichi-Sankyo	Symphogen	TG Therapeutics		
	WuXi	Five Prime		
		Seattle Genet	ttle Genetics	

#### **Generics**

Alvogen

Par

Zydus Cadila

Coherus

**OncoBiologics** 

BioCad

Beloteca

Over 40 additional companies



## **Ligand Portfolio Highlights**

- We estimate our partners will conduct over 200 studies and spend over \$2 billion in 2018 on R&D to advance our programs
- Total potential payments under existing contracts for our more than 170 partnerships exceed \$2.5 billion
- Ligand is partnered with major companies for some of the industry's most important potential medicines
- <u>Ligand-based programs are major assets for partners</u>
  - Foundational for mega-acquisitions
  - e.g., Novartis with GSK-Oncology (Promacta), Amgen with Onyx (Kyprolis), BMS with Cardioxyl

Foundational for IPO, reverse mergers or important financings

e.g., Melinta, Viking, Aldeyra, Retrophin, Sermonix, Seelos, others



# **RPT – Ligand's Foundation of Value**

Revenue



High Growth
High Margin
Strong Protection

Pipeline



Large
High Quality
Many Late Stage

**Technology** 



Best-in-Class
Leverageable
Strong IP

## **Two Major Technology Platforms**

Market Leading, Best-in-Class



Only antibody discovery platform with three species

Platform with the most partners

Strong market protection and long patent coverage for products

Fast-growing number of drug candidates moving to the clinic

# **CAPTISOL®**

Highly-pure, pharmaceutical grade ingredient with reliable supply

Broad, global patent protection

**Large Drug Master Files** 

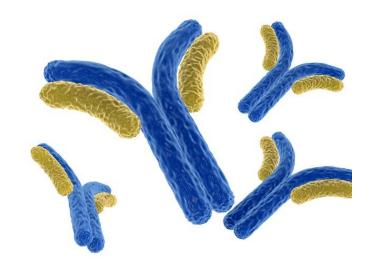
Now with most pharma partners, most approved products



## The Potential of Antibody Therapy

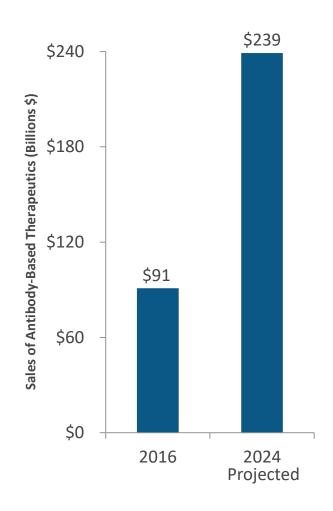
#### Harnessing the Power of the Immune System

- Antibody therapy leverages an animal's ability to make proteins that bind to specific molecules on cells in response to toxic or foreign substances
- Antibodies can kill targeted cells (ie cancer) or impact the way cells function
- Antibodies are a major, fast growing class of medicines. Ligand has valuable technology serving the space
- R&D is underway to develop antibodies for a broad array of indications, including cancer, inflammation, auto-immune, neurological, viral diseases and many others



# **Antibodies: Major Opportunity for Ligand**

- In 2016, Ligand made a major strategic investment into antibody discovery by acquiring OMT, Inc. for ~\$178 million
- In 2017, Ligand expanded its antibody discovery program by acquiring Crystal Bioscience for ~\$25 million
- Global sales of antibodies estimated to approach \$240 billion in 2024
- 520 industry-wide programs in development, more than tripled since 2008





# **OmniAb: A Best-in-Class Technology**









- Ligand's OmniAb technology is a drug discovery platform that enables drug companies to discovery antibodies
- Ligand has entered license agreements with over 30 companies
- Under the license agreements,
   Ligand is entitled to financial payments including:
  - License fees
  - Milestones
  - Royalties

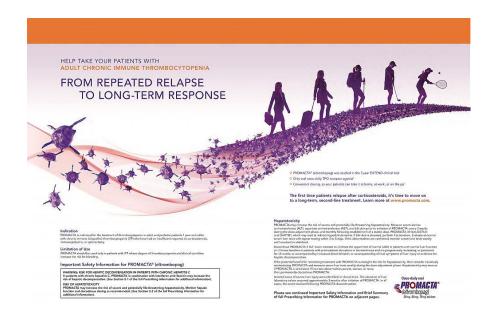


# **Promacta**®

#### **Promacta®: Blockbuster Commercial Potential**

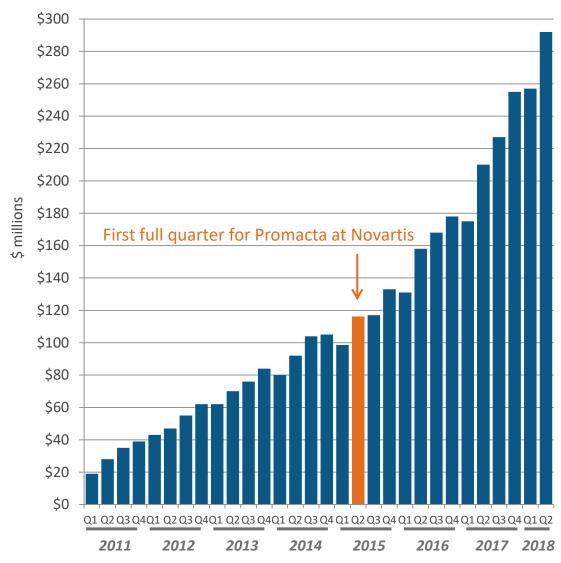
- Oral medicine that boosts platelets in patients with thrombocytopenia, or low-platelets
- Partnered with Novartis worldwide
- Sales trending to exceed \$1 billion in 2018; Consensus third-party analyst estimates project \$1.4 billion in 2021
- Approved for numerous indications involving low platelets, and multiple trials underway to support label expansion







### **Promacta: Quarterly Revenue**



- Q2'18 revenue was \$292 million, an \$82 million increase (39%) over Q2'17
- Acquisition of product by Novartis from GSK significantly increased sales and growth trends
- Sales now annualize to over \$1 billion
- Ligand earns tiered royalties between 4.7% and 9.3%



# **Kyprolis**®

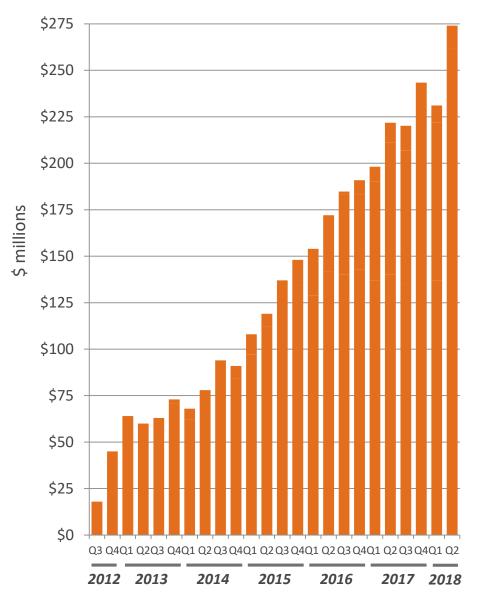
# **Kyprolis**



- Kyprolis viewed as best-in-class proteasome inhibitor for multiple myeloma (MM)
- Developed and marketed by Amgen
  - Utilizes Ligand's Captisol technology
- Approved for relapsed or refractory MM in the US, EU and Japan (Ono)<sup>1</sup>
  - As single agent, or in combination with dexamethasone or Revlimid and dexamethasone
- Kyprolis has demonstrated overall survival improvement in Phase 3 trials,
   bolstering the potential for the drug in a competitive space
- Amgen actively investing in more studies with potential to expand the label and use of the drug



# **Kyprolis: Quarterly Revenue**



- Amgen/Ono reported combined
   Q2 revenue of \$274 million
- Rest-of-world contribution becoming more substantial
- Ligand earns 1.5% to 3% royalties

# The Big 6: Major Pipeline Assets

Leading pipeline assets based on stage and/or potential value

Partner	Program (Therapy Area)	Stage	Royalty Rate	Upcoming Events
SAGE	Brexanolone (Neurology)	Pre-NDA	3.0%	December 19, 2018 PDUFA Date
<b>Re</b> trophin *	Sparsentan (FSGS- Kidney Disease	Phase 2/3	9.0%	Phase 3 Interim Data
Sermonix Pharmaceuticals	Lasofoxifene (Oncology/Women's Health)	Phase 2/3	6.0-10.0%	Phase 2 Start
Bristol-Myers Squibb	BMS986231 (Cardiovascular Disease)	Phase 2/3	2.0-3.0%	Phase 2b Data
Lilly	Prexasertib (Oncology)	Phase 2	1.5-3.0%	Phase 2 Data in various advanced cancers
ROIVANT	RVT-1502 (Diabetes)	Phase 2	Low double digit to mid teens %	Clinical Progression



# Glucagon Receptor Antagonist (GRA)

#### Roivant/Metavant Partnership - RVT-1502

- Glucagon is a hormone that stimulates the liver to produce glucose
  - GRAs are designed to lower glucose levels for treatment of diabetes



- Ligand discovered and initially developed a novel molecule that potently binds the glucagon receptor and antagonizes the actions of glucagon
  - Ligand completed successful Phase 1 and Phase 2 clinical trials
  - Global patents covering various forms of the molecule, if granted, would not be expected to expire until 2039
- Ligand entered a major deal with Roivant for GRA in March 2018
- Ligand received \$20 m at signing, and is eligible to receive \$528 m in milestone payments, and royalties ranging from low double digits to mid-teens



### **Seelos Therapeutics**

#### Focused on Novel Products to Treat CNS Diseases

- Seelos Therapeutics recently announced plan to merge with Apricus Biosciences
  - If completed, merger will result in Nasdaqlisted company with a continued focus on novel products for CNS diseases with significant unmet medical needs



Seelos Therapeutics, Inc.

- Four Ligand-partnered programs are in the Seelos pipeline
  - SLS-006: First-in-class small molecule partial dopamine agonist for Parkinson's
    Disease that has successfully completed Phase 2, plans for pivotal registration
    studies to commence in 2019
  - *SLS-008:* Once-daily oral **CRTH2** program for undisclosed pediatric orphan indication
  - *SLS-010:* Histamine 3 receptor inverse agonist for narcolepsy and related disorders
  - *SLS-012:* Captisol-enabled™ acetaminophen for post-operative pain
- Ligand eligible to receive milestones of \$144 m and royalties ranging from 4% to 10%



### **Highlighted Program**

#### VK2809



 VK2809 is a selective thyroid hormone receptor beta (TR-β) agonist



- Partnered with Viking Therapeutics in 2014
- Potential applicability in broad range of indications, including:
  - Non-alcoholic fatty liver disease (NAFLD)
  - Hypercholesterolemia
  - GSD-1a
- Enrollment currently nearing completion in Phase 2 trial in NAFLD and hypercholesterolemia
  - Viking expects to announce results in 2018
- Clinical study in GSD-1a expected to begin dosing patients in June
- Ligand eligible to receive \$75 m in development milestones (per indication),
   up to \$150 m in commercial milestones, and 3.5% to 7.5% royalty



# **Proposed Acquisition of Vernalis plc**

• On August 9, 2018, Ligand announced an offer to acquire Vernalis plc for \$43 million in cash



- Vernalis is a structure-based drug discovery biotech based in United Kingdom
- The acquisition of Vernalis would provide Ligand with the following:
  - Portfolio of more than 8 shots on goal in the respiratory, oncology and CNS sectors
  - 70-person R&D team based in Cambridge, England with an active portfolio of collaborations that have the potential to generate additional shots on goal
  - Compound library and early-stage, unpartnered programs providing BD opportunities
  - Net cash on hand after deal expenses of approximately \$36 million
  - Partners include; Corvus, Verona, Celgene, Servier, Daiichi Sankyo and others
- Transaction is subject to Vernalis shareholder approval and is expected to close in October 2018

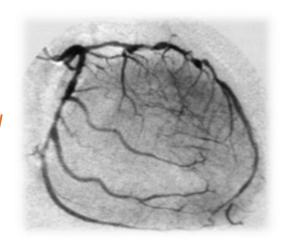


# Internal R&D

# **Captisol-enabled Iohexol**

#### Leveraging Captisol® Technology to Make Drugs Safer

- Recently established new program to develop Captisol-enabled, next generation contrast agents for diagnostic imaging
- Patented uses of Captisol to reduce acute kidney injury (AKI) during medical interventions, including:
  - **Iodinated contrast agents:** *iohexol, iopamidol, iodixanol*
  - **Anticancer agents:** *cisplatin, doxorubicin, methotrexate*
  - Aminoglycosides: gentamicin



 Captisol-enabled lohexol has successfully completed preclinical studies and Ligand plans to progress into clinical development

# **Iodinated Contrast Agents**

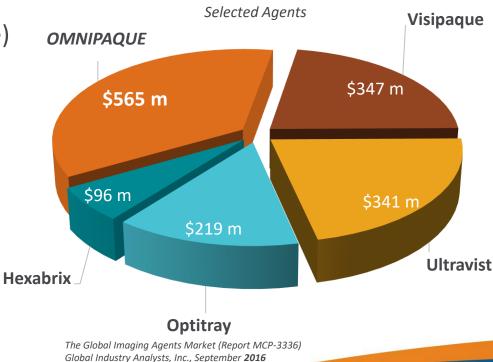
#### **Market and Needs**

- 30 million imaging procedures/yr in the US
  - lodinated contrast agents represent >60% of all X-ray imaging agents sold (~\$1.5 B market)
- Iohexol (*OMNIPAQUE™*, GE Healthcare) is the most widely-used injectable diagnostic contrast agent for X-ray imaging procedures
  - Global sales >\$500 M
  - \$250 M+ in US, ~30% market share
  - No generic competition
  - Reported incidence of contrastmedium induced nephropathy reported at 26% for lohexol<sup>1</sup>

"Acute Kidney Injury remains a concern for patients undergoing cardiac interventional procedures utilizing intravascular iodinated contrast"

McCullough, J Am College of Cardiology 2016;68:1465-73

#### **Iodinated Contrast Agents - US Sales**



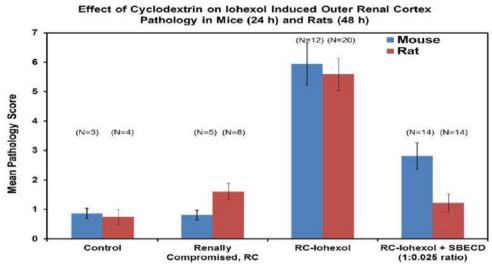
Ligand

### **Captisol-enabled Iohexol**



#### Leveraging Captisol Technology to Make Drugs Safer

- Captisol-enabled Iohexol reduces renal pathologies in animal models
  - Mean scores show >50% prevention of Iohexol-induced pathologies
  - Selected as next product for internal development, based on:
    - 1. Large number of patients affected globally
    - 2. Lack of alternatives
    - 3. Established IP portfolio
    - 4. Potential for abbreviated development and approval process



Rowe ES, et al. Journal of Neuroimaging 2016; 26(5):511-8

- Potential for high Captisol use
  - Estimated up to 10-15 metric tons annually



### **Captisol-enabled Iohexol**



#### **Commercial Landscape**

- Cardiovascular (CV) imaging is a large, growing segment of the market
- More than 50% of CV imaging procedures are performed in patients  $\geq$  65 years old, and substantial portion have risk factors for acute kidney injury
  - A continuing issue with recent and broad medical visibility<sup>1</sup>
  - No products are approved to prevent or treat acute kidney injury



- Differentiation in labeled safety, competitive strategy for potential partner(s) to gain sizeable market share
- Plan to pursue partnering opportunities following clinical work



Outcomes after Angiography with Sodium Bicarbonate and Acetylcysteine

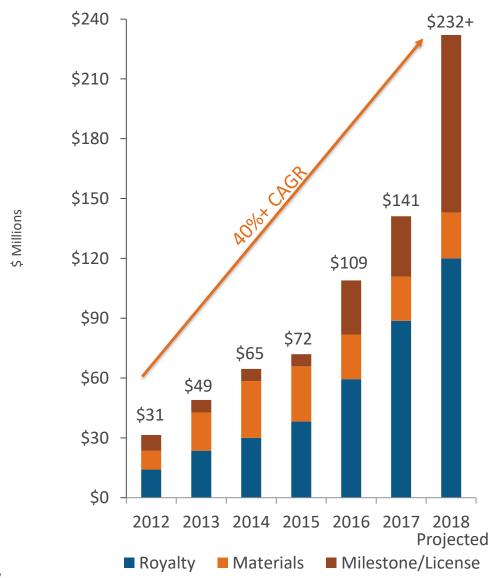
EDITORIALS

Prevention of Contrast-Associated Acute Kidney Injury

# Financial Overview and Outlook



#### **Sustained Revenue Growth**

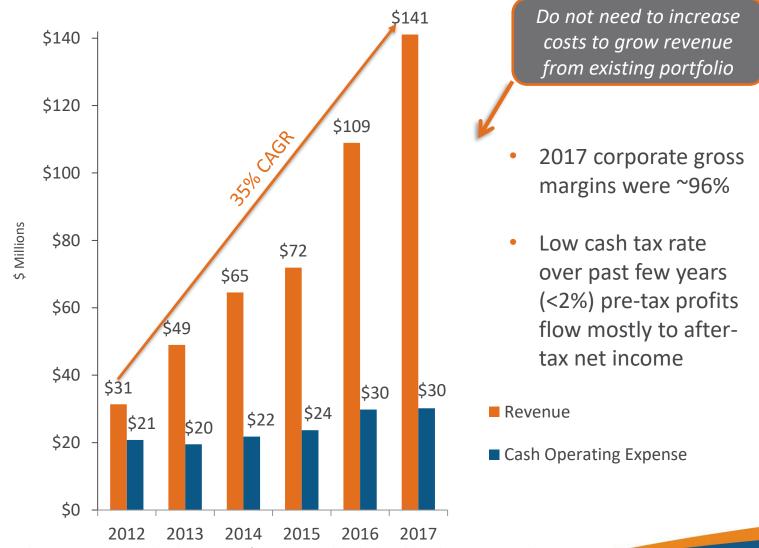


- Consistent, strong annual revenue growth driven by:
  - High royalty growth
  - Increasing contribution from milestones
  - Consistent contribution from material sales
- 2018 adjusted EPS would be approximately \$6.30 with total revenue of \$232 million
- Potential for additional \$8 million of 2018 milestones



#### **Revenue Growth Paired with Modest Costs**

#### Resulting in Increasing Operating Margins





## Reconciliation of GAAP EPS to Adjusted EPS

2017 GAAP Earnings Per Share	\$0.53	
Stock-based compensation expense	1.06	
Non-cash interest expense		
Amortization related to acquisitions		
Increase in contingent liabilities	0.11	
Loss from Viking	0.09	
Other	(0.16)	
Income tax effect of adjusted reconciling items	(0.83)	
Deferred tax asset adjustment		
Excess tax benefit from stock-based compensation	(0.20)	
Valuation allowance release	(0.18)	
2019 Senior Convertible Notes share count adjustment	0.17	
2017 Adjusted Earnings Per Share		
GAAP Shares	23.48	
Dilutive potential common shares issuable of redeemable convertible not		
Adjusted Shares		

Adjusted EPS guidance excludes stock-based compensation expense, non-cash debt-related costs, changes in contingent liabilities, transaction-related amortization, pro-rata net losses of Viking Therapeutics as well as fair value adjustments to our holdings in their common stock, convertible note receivable and warrants, mark-to-market adjustments for amounts owed to licensors, changes in contingent liabilities related to our CVRs, the excess convert shares covered by bond hedge and certain one-time non-recurring items



#### **Calendar of Potential Events**

Potential Milestones for Ligand and Partners over next ~12 months

#### Clinical Data

VK2809 Phase 2 data in Hypercholestrolemia/NASH Viking

BMS-986231 Phase 2 data in Heart Failure Bristol Myers Squibb

Baxdela Phase 3 data in community-acquired bacterial pneumonia
Melinta

Prexasertib Phase 2 in Cancer Lilly

#### **Progression**

Sparsentan Phase 3 Start in IgA Nephropathy Retrophin

VK2809 Phase 1 Start in rare glycogen storage disease
Viking

Lasofoxifene Phase 2 Start in metastatic breast cancer
Sermonix

RVT-1502 Clinical Progression in diabetes

Metavant

#### <u>Approvals</u>

Brexanalone FDA Action
Sage Therapeutics

Evomela China FDA
Action
CASI (Spectrum)

Baxdela Ex-US action Menarini/Eurofarma (Melinta)

Promacta FDA Priority review of sNDA
Novartis





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