

CHIMERIX

DISCOVERING, DEVELOPING, AND COMMERCIALIZING NOVEL MEDICINES THAT IMPROVE OUTCOMES FOR IMMUNOCOMPROMISED PATIENTS

Year-End Earnings and Update Call March 2, 2017

Forward-Looking Statements

These slides and the accompanying oral presentation contain forward-looking statements and information within the meaning of the Private Securities Litigation Reform Act of 1995 that are subject to risks, uncertainties and other factors, including the possibility that there may not be a viable continued development path for brincidofovir, that any clinical trials we may conduct will not demonstrate adequate efficacy and safety of brincidofovir, that the FDA and other regulatory authorities may not approve brincidofovir or brincidofovir-based regimens, and that marketing approvals, if granted, may have significant limitations on their use. As a result, brincidofovir may never be successfully commercialized. In addition, Chimerix may be unable to file for regulatory approval for brincidofovir with other regulatory authorities. These risks, uncertainties and other factors could cause actual results to differ materially from those referred to in the forward-looking statements. The reader is cautioned not to rely on these forwardlooking statements. These and other risks are described in detail in Chimerix's Annual Report on Form 10-K for the year ended December 31, 2016 and other documents subsequently filed with or furnished to the U.S. Securities and Exchange Commission. All forward-looking statements are based on information currently available to Chimerix, and Chimerix assumes no obligation to update any such forward-looking statements.



Value Drivers for Chimerix

Data Generation and Publications

Oral BCV EU Launch 2020 AdV treatment IV BCV
Launch US & EU 2021
Multi-viral prevention

IV BCV BKV CMV JCV 2023-2030

Smallpox NDA
US Procurement, Ex-US procurement
2019

CMRX Proprietary Library & Discovery Program

CMX521 for Norovirus IND in 2017

CMX521 Clinical Data 2018 -2020

CMX521 for Norovirus NDA/MAA 2022



Chimerix Pipeline

	Program	Discovery	Pre-clinical	Phase 1	Phase 2	Phase 3	Anticipated Approval
Short-course Oral BCV	AdV Pediatric Treatment	Study 999 in EU (+	-/- US) to start i	n 2H 2017			2020
	Smallpox	Data from second	animal model ir	2017			2019
IV BCV	Multiviral Prevention	Data from MAD in	2017	Ph 2/3 in	Peds HCT		2021
	BKV Treatment	Initiate Phase 2b la	ate 2017	Ph 2/3 in	Kidney Tx		2023
	CMV Treatment	Initiate Phase 2b la	ate 2017				
CMX521	Norovirus	IND 2H 2017		FTIH 4Q2017	> POC Challe	enge Study	2022
CMX157	HBV Treatment	*Licensed to Contr	aVir				
Business Development		Ongoing Diligence		/			



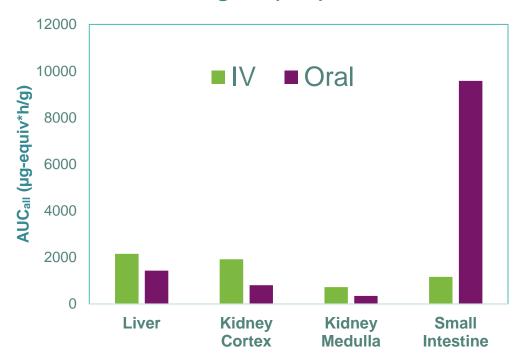
2016 Brincidofovir Learnings

- Gastrointestinal toxicity of oral brinci was related to GI over exposure and duration
- 2. High brinci plasma exposures delivered via the intravenous route in animal studies did not result in the gut injury observed with oral brinci dosing
- 3. IV brinci provides higher drug levels in plasma and difficult-to-reach compartments in animals allows exploration of treatment indications not possible with oral dosing
- 4. Oral brinci was associated with rapid AdV clearance in AdVise
- Rapid clearance of AdV was associated with a decrease in overall and AdVassociated mortality in AdVise



Key Learning #1: Oral BCV GI Tox is Related to GI Exposure and Duration

Total Drug Exposure (AUC_{all}) in Clearance Organs (rats)



- Oral BCV in rats resulted in significantly higher exposures in the intestine vs other organs, providing explanation for GI toxicity
- In rats, IV BCV delivered comparable drug exposure to key organs, including the liver, kidney, and small intestine
- IV BCV without GI tox could support longer duration of therapy

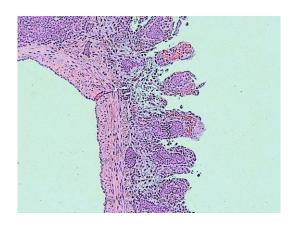
IV BCV expected to prevent "over-exposure" of gut



Learning #2: IV BCV Delivers High Exposures without GI Tox

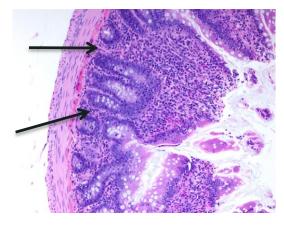
28-day rat studies of IV BCV:

- No in-life clinical findings at highest dose: IV BCV 15 mg/kg
- No animals had diarrhea
- All animals gained expected weight during study
- No liver enzymes elevations
- Minimal GI findings in intestines at final pathology



Rat intestine after oral BCV

 Significant loss of epithelium in intestinal villi



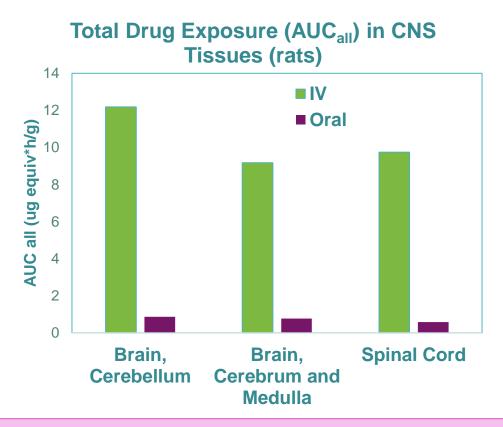
Rat intestine after IV BCV

Minimal single-cell effects noted for IV BCV

IV BCV for 28 days avoided gut injury observed with oral BCV



Learning #3: IV BCV Raises the Exposure Ceiling, Delivers More Drug to Plasma and to Difficult-to-Reach Compartments



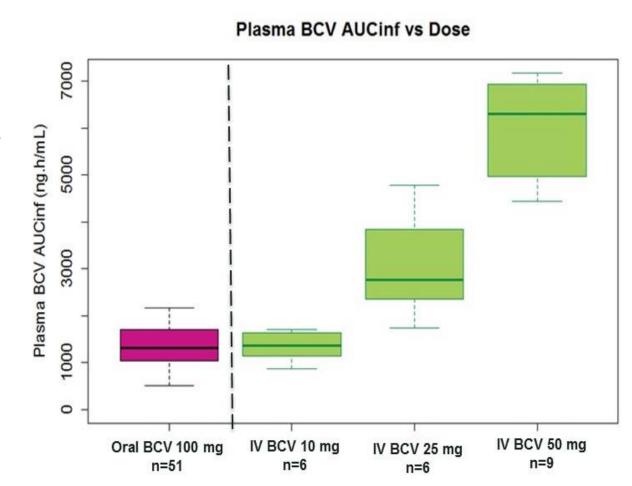
- IV BCV provides higher exposures which may be needed for some treatment indications (BK nephropathy, CMV disease)
- Higher CNS exposures with IV BCV could support testing for viral infections in the brain, e.g.:
 - Herpes encephalitis
 - HHV-6 encephalitis
 - JC virus/PML in transplant recipients or patients with Multiple Sclerosis

IV BCV enables exploration of treatment indications not possible with oral BCV



Ongoing IV BCV Single Ascending Dose Study: Drug Levels

- IV BCV/placebo 10 mg, 25 mg, & 50 mg cohorts have completed dosing in healthy subjects
- IV BCV 10 mg provides similar exposure as oral BCV 100 mg
- Drug exposure was linear as dose was increased
- IV BCV 50 mg provided plasma drug exposures higher than achieved with oral BCV, and at or above the range of exposures targeted for treatment indications such as CMV and BK nephropathy





IV BCV Single Ascending Dose: Preliminary Safety & Tolerability

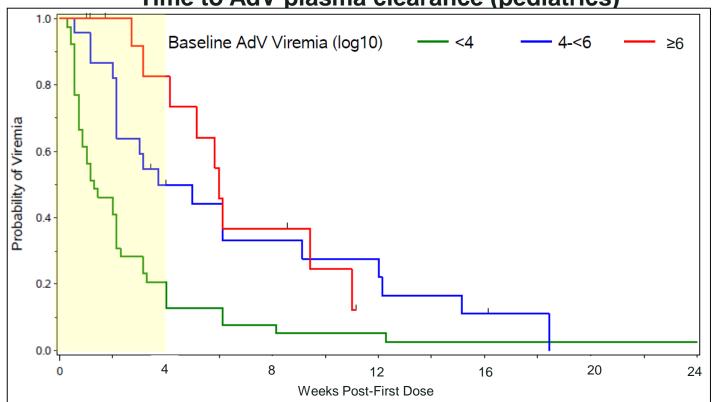
- Cohorts 1-3 IV BCV/placebo doses have been generally safe and well tolerated:
 - Grade 1-2 lab abnormalities in Cohorts 1, 2 & 3, none clinically significant
 - NO myelotoxicity or nephrotoxicity
 - Study-drug related AEs were mild, and limited to Cohort 3
 - 3 subjects with IV site bruising/discomfort
 - 2 subjects with headache (resolved spontaneously)
 - 1 subject with loose stools (resolved spontaneously)

Observed GI tolerability of single dose IV BCV thru Cohort 3 in humans is improved compared to single dose oral BCV



Learning #4: Early Intervention Resulted in Undetectable AdV Viral Loads in the First 4 Weeks of Dosing in AdVise

Time to AdV plasma clearance (pediatrics)



- Peds pts with lower AdV viral loads at baseline (<10,000 or 4 log₁₀ c/mL) cleared within a median 8 days of BCV
- Peds pts with AdV plasma <5 log₁₀
 (<100,000 c/mL): 72% cleared within
 4 weeks
- Study 999 will focus on sites with regular screening for AdV who are likely to detect and treat AdV viremia while viral loads are ~5 log₁₀ or lower

Short-course oral BCV should result in clearance of AdV in majority of patients



Learning #5: Virologic Response Is Associated with a Decrease in Overall and AdV-Associated Mortality in AdVise

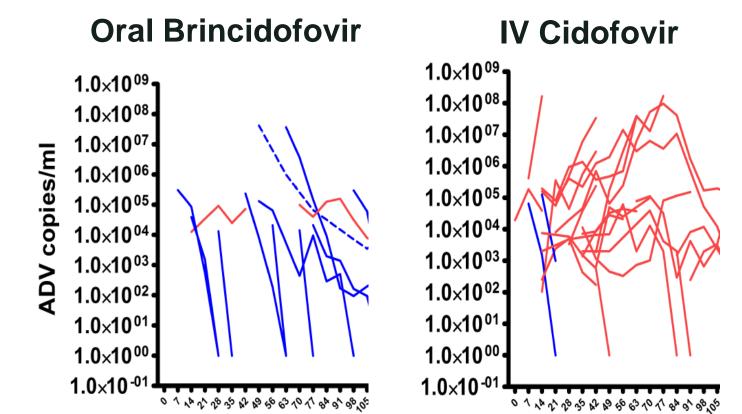
Patients with Disseminated AdV Disease		Mortality	AdV-Associated Mortality	
Pediatric	Responder*	7/28 (25%) p=0.031	1/28 (4%)	
	Non-responder	7/13 (54%)	2/13 (15%)	
Adult	Responder*	5/10 (50%)	0/10 (0%)	
	Non-responder	p=0.0004 13/14 (93%)	10/14 (71%)	

^{*} Responders are subjects with baseline AdV viremia still on study at week 6 who had undetectable plasma AdV at week 6; non-responders are subjects who did not achieve the specified cut-off

Undetectable AdV was associated with nearly double the overall survival



Greater Viral Load Reduction for Brinci vs Cidofovir in First 100 Days Post-Transplant from UK Consortium



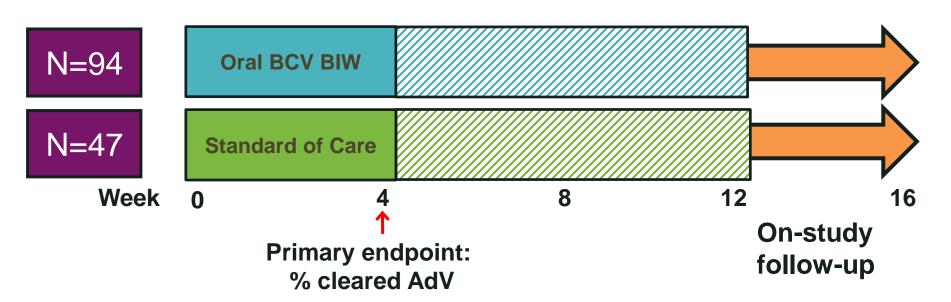
- AdV 2 log₁₀ reduction in 2 weeks (solid blue lines) observed with BCV in 13/18 (72%) vs. 2/23 (9%) with IV cidofovir
- Complete response observed in 13 BCV patients (80%) vs. 8 patients with cidofovir (35%)
- Differences greatest in first 100 days (before immune reconstitution)

Day after Transplant



Proposed Study 999: Short-course Oral BCV for Treatment of AdV in Pediatric HCT in First 100 Days Post-Transplant

- Small, open label, comparative study of BCV vs. standard of care
 - Inclusion: pediatric T-cell depleted or cord blood HCT recipients with confirmed > 1000 c/mL AdV DNA in plasma, <100d from HCT
- Duration: Treat until AdV cleared from plasma (minimum 4 weeks, maximum 12 weeks)
- Primary endpoint: % undetectable plasma AdV at Week 4
 - N~140 (2:1, 90% power) for 70% vs. 40% response rate
 - Superiority of BCV in clearance of AdV from plasma could enable conditional or full EU Approval





Next Steps for IV and Oral BCV

Oral BCV

- Short-course dosing for treatment of AdV and smallpox continue in development
- Small comparative study in pediatric HCT recipients at high risk of AdV disease anticipated to begin in 2H 2017, potential for conditional or full approval in the EU based on positive data

IV BCV

- Single Ascending Dose Study in healthy subjects continues, with one additional cohort nearing completion
- Multiple-ascending dose study in healthy subjects to initiate in 1H 2017
- Phase 2 dose-ranging studies in treatment of CMV and BKV infections anticipated
- Opportunity to explore a broad range of additional indications, esp CNS infections
- Ability to dose for longer durations with lower risk of GI tox allows pursuit of multi-viral prevention

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Our ability to provide BCV in oral and IV formulations enables development across multiple indications and populations, with the potential to address conditions without any approved therapies

Demonstrating IV BCV's Multi-Viral Prevention

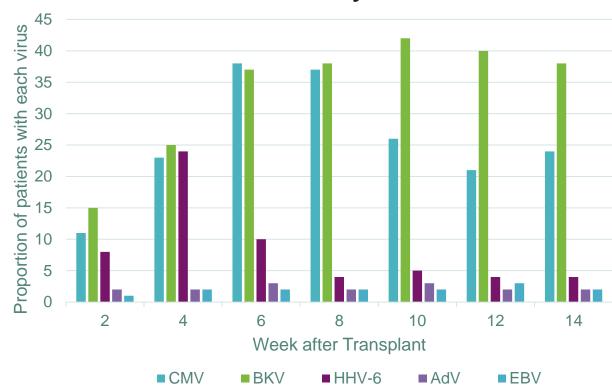
- Pediatric patients are at high risk for multiple DNA virus infections, with adenovirusrelated mortality a particular concern
- The lower risk of GI toxicity with IV BCV may allow longer duration of dosing throughout high-risk period
- Proposal: placebo-controlled trial of IV BCV in pediatric allogeneic HCT recipients
 - Primary endpoint prevention of adenovirus; allows placebo control
 - Secondary endpoint of CMV, as high risk pediatric HCT recipients tend to reactivate AdV earlier than CMV
 - Other secondary endpoints to include other DNA virus prevention and health outcomes
- Benefits:
 - Superiority design allows smaller study than head to head trial for CMV



Adult and Pediatric HCT Recipients Face Risks Beyond CMV

- Multiple DNA viruses commonly reactivate as early as the first week after transplant
- Of the HCT recipients who reactivated CMV, >75% had at least one other DNA virus identified and were at increased risk of mortality
- 1 in 3 HCT recipients had ≥3 DNA viral infections detected

DNA virus reactivation by Week after HCT



More DNA viruses reactivating = higher risk of death



Unmet Need Drives Demand for Brincidofovir

More than 300 patients received oral BCV for life-threatening AdV in 2016

	Pediatrics		Adı	TOTAL	
	Asymptomatic	Symptomatic ^a	Asymptomatic	Symptomatic ^a	All
EINDs/NPP	-	344	-	210	566 ^b
Study 202	24	3	10	2	39
Study 304	51	79	23	48	201
Study 350	10	19	5	23	68 ^c
Study 351	3	62	1	26	92 ^d
Total	88	465	39	321	966

alncludes local and disseminated AdV disease



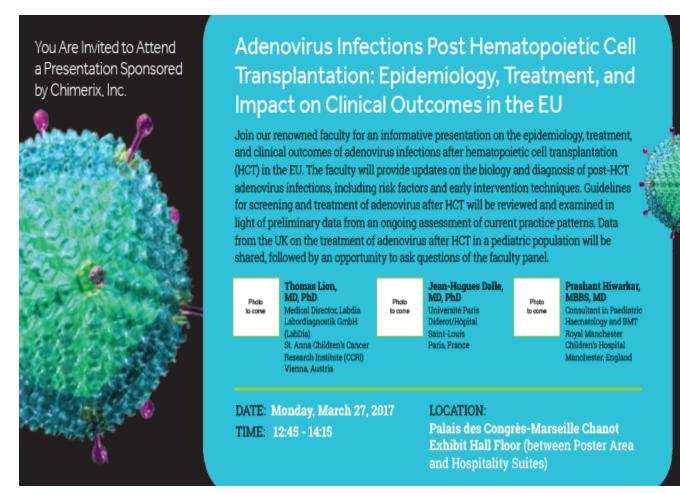
blncludes 12 pts whose age is not known; assumes that EIND/NPP pts are symptomatic; Includes 9 patients who received open-label BCV

[°]Grimley et al. EBMT 2013; total numbers includes all subjects with AdV identified as a primary or secondary DNA viral infection that could not be classified (N=68)

dOngoing expanded access protocol, n=92 as of 22 Feb 2017

Educational Efforts Continue to Build

- Multiple activities scheduled for the European BMT Meeting in March
 - KOL Advisory Board
 - Chimerix supported educational presentation on March 27, 2017
- Several oral presentations and abstracts of interest including:
 - Data from the Chimerix long-term registry study of patients from our trials
 - Data from the UK, France, Spain and the Netherlands summarizing their experiences using brincidofovir to treat AdV infections





Year-end 2016 Financial Position Remains Strong

12/31/16 Cash and investments: \$ 278M

2016 Net cash burn: \$65M

Net cash burn for 2016 reflected a 35% decrease over 2015

We currently expect expenses to trend upward modestly in 2017



Chimerix: 2017 Potential Catalysts



- IV BCV: Present final IV BCV clinical data from single ascending dose study, initiate multiple dose study
- CMX521 for Norovirus: IND-enabling studies
- IV BCV: multiple ascending dose clinical data
- Oral BCV: Initiate Study 999, small comparative AdV trial in pediatric HCT recipients for potential approval
- Initiate Dose-Ranging Studies in patients with CMV and BKV
- Data from second animal efficacy model for smallpox
- CMX521: Submit IND and initiate first clinical studies

\$278 million in capital is sufficient to fund operations through anticipated catalysts in 2017

