



ONCONOVA
T H E R A P E U T I C S

EXECUTIVE SUMMARY

October 2018 | Nasdaq: ONTX

FORWARD LOOKING STATEMENTS

This presentation contains forward-looking statements about Onconova Therapeutics, Inc. based on management's current expectations which are subject to known and unknown uncertainties and risks. Onconova has attempted to identify forward-looking statements by terminology including "believes," "estimates," "anticipates," "expects," "plans," "intends," "may," "could," "might," "should," "approximately" or other words that convey uncertainty of future events or outcomes. This presentation assumes the Company raises capital for disclosed product development plans. Our actual results could differ materially from those discussed due to a number of factors, including, but not limited to, our ability to raise additional financing on favorable terms, the success of our clinical trials and our ability to obtain regulatory approvals and other risk factors outlined in our annual and quarterly reports filed with the Securities and Exchange Commission. We are providing this information as of the date of this presentation and do not undertake any obligation to update any forward-looking statements, whether written or oral, that may be made from time to time, as a result of new information, future events or otherwise except as required by law.

ONCONOVA THERAPEUTICS, INC.

- Founded in 1998; IPO in 2013 (Nasdaq: ONTX)
- Phase 3 stage clinical candidate: rigosertib
 - Focused on Myelodysplastic Syndromes (MDS)
- Rigosertib partnered in Japan and Latin America
 - Additional partnerships to come
- Broad pipeline of drug candidates
 - Larger opportunities in solid tumor indications



PORTFOLIO: RIGOSERTIB AND OTHER OPPORTUNITIES

Lead

- Phase 3 INSPIRE trial progressing to completion after promising interim analysis and enhanced powering
- Trial completion projected in H2-2019

Oral

- Oral rigosertib provides two large-market opportunities
- Combination trial for front-line high risk MDS ready to advance to Phase 3 protocol in Q4-2018

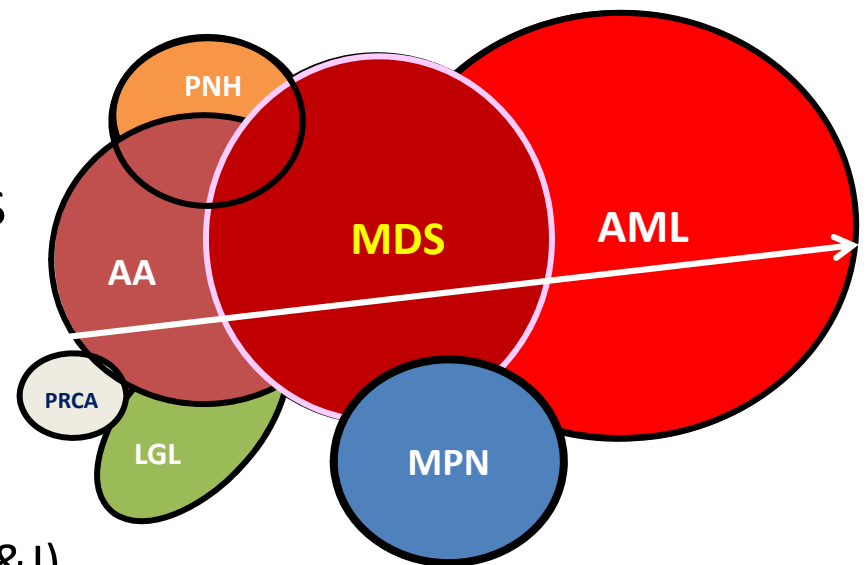
More

- NCI funded RASopathies trial for rare pediatric indications
- New CDK inhibitor presents opportunities for the future



MDS IS RELATED TO OTHER BONE MARROW DISEASES

- MDS: malignant bone marrow disorder characterized by:
 - Acquired cytogenetic and genomic abnormalities, but typically only in the marrow.
- US prevalence is 59,000
 - ~13,000 have higher risk (HR) MDS
 - ~10,000 second-line patients
- Available Treatments limited to hypomethylating agents
 - Vidaza (Celgene); Dacogen (Eisai/J&J)
 - Approved >decade ago; now off-patent
 - No approved therapy following HMA failure
 - New therapy could have \$billions opportunity



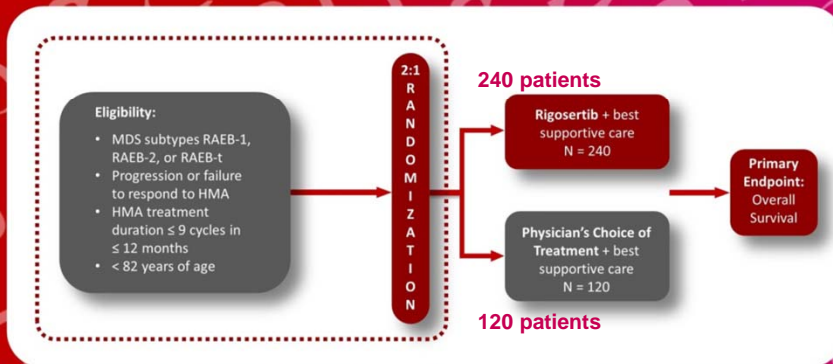
INSPIRE PHASE 3 TRIAL RESULTS EXPECTED IN 2019

The Pivotal MDS **INSPIRE** Trial is Now Recruiting Patients

International Study of Phase III Intravenous Rigosertib

STUDY DESCRIPTION

A Phase 3, international, randomized, controlled study of Rigosertib + best supportive care versus physician's choice of treatment + best supportive care in patients with myelodysplastic syndrome (MDS) after failure of a hypomethylating agent (HMA).



PRIMARY ENDPOINTS

Overall survival in the intention-to-treat population and in patients with very high risk per the Revised International Prognostic Scoring System (Greenberg et al, *Blood* 2012).

INTERNATIONAL TRIAL

More than 170 trial sites

INSPIRE start
December 2015



Interim Analysis
January 2018

Trial size increased after "promising" signal



Top-line Data
H2-2019 (projected)



INITIAL RESPONSE DATA FOR ONGOING COMBINATION TRIAL

An additional 45 patients are enrolled in the expanded Phase 2 trial at an increased dose of oral rigosertib (1120 mg) to determine optimal efficacy and safety

Response Criteria	Response per IWG 2006	
	No prior HMA (N=20)	HMA resistant (N=13)
Complete Remission*	7 (35%)	1 (8%)
Marrow CR + Hematologic Improvement (HI)	6 (30%)	4 (31%)
Marrow CR alone	3 (15%)	3 (23%)
Stable Disease	3 (15%)	5 (38%)
Overall IWG Response	17 (85%)	8 (62%)

**All responders had CR and no PR was noted in this study*



COMBINATION THERAPY: NEXT STEPS AND TIMELINES

Step	Start	Complete	Remarks
Phase 2 expansion <i>Fully enrolled</i>	Q1-2017	Q2-2018	<ul style="list-style-type: none"> Incidence of hematuria reduced (to date) in the trial Dose and schedule of 1120 mg daily dose explored*
Phase 3 protocol	Q1-2018	Q4-2018	<ul style="list-style-type: none"> Synopsis created SPA and BTM submissions contemplated after complete efficacy assessment
Phase 3 trial	2019	2021	<ul style="list-style-type: none"> Rapid enrollment expected All patients to receive active therapy Response endpoint can be achieved in <6-9 months after patient is enrolled

*Dose justification based on oral rigosertib optimal transfusion independence rate data in Lower-Risk MDS (ASH 2017)



EXPANDING AND EXTENDING RIGOSERTIB PATENT COVERAGE

- Strong existing patent estate
 - Existing coverage of composition of matter (e.g. U.S. 7,598,232), formulations, combinations and methods in US and many countries worldwide
- Supplemented by Orphan Designation for MDS in US, Europe and Japan
- **New issued US patent 10,098,862 extends IP runway to 2037**

- **US Patent 10,098,862**
 - Pending in PCT and non-PCT countries worldwide
 - Covers injectable and oral products

(12) United States Patent Maniar	(10) Patent No.: US 10,098,862 B1
	(45) Date of Patent: Oct. 16, 2018
(54) FORMULATIONS WITH ENHANCED STABILITY AND BIOAVAILABILITY FOR ADMINISTRATION OF (E)-2,6-DIALKOXYSTYRYL 4-SUBSTITUTED BENZYL SULFONES	(56) References Cited
	U.S. PATENT DOCUMENTS
	7,598,232 B2 10/2009 Reddy et al. 514,710
	8,063,109 B2* 11/2011 Bell A61K 9/0019 514,710
(71) Applicant: ONCONOVA THERAPEUTICS, INC., Newtown, PA (US)	8,476,320 B2* 7/2013 Bell A61K 9/0019 514,710
(72) Inventor: Manoj Maniar, Fremont, CA (US)	2010/0305059 A1 12/2010 Reddy et al.
(73) Assignee: ONCONOVA THERAPEUTICS, INC., Newtown, PA (US)	OTHER PUBLICATIONS
(*) Notice: Subject to any disclaimer, the term of this patent is extended or adjusted under 35 U.S.C. 154(b) by 0 days.	Advani et al., Indian Journal of Cancer (2014), 51(1), pp. 40-44.*
(21) Appl. No.: 15/688,320	Garcia-Manero, G. et al. "Comprehensive Analysis of Safety: Rigosertib in 557 Patients with Myelodysplastic Syndromes (MDS) and Acute Myeloid Leukemia (AML)," Blood 128:2011-(2016).
(22) Filed: Aug. 28, 2017	Navada, S. et al. "Combination of Oral Rigosertib and Injectable Azacitidine in Patients with Myelodysplastic Syndromes (MDS): Results from a Phase II Study," Blood 128:3167-(2016).
	Dash, A.K., et al. "Preformulation Development of a Parenteral Formulation for ON 01210.Na, a Radioprotectant," Presentation Abstract AAPS Annual Meeting and Exposition, Nov. 5-10, 2005.
	Strickley, R. G., "Solubilizing Excipients in Oral and Injectable Formulations," Pharmaceutical Research vol. 21(2) pp. 201-230 (2004).



ON 123300: NEXT GENERATION CDK4/6 INHIBITOR

Also targets ARK5 (NUAK1)

Differentiation for a Competitive Field

- Recently launched Ibrance[®], Kisquali[®] and Verzenio[®] have been hailed as potential breakthroughs in cancer therapy
 - First FDA approval for CDK 4/6 inhibitor is for breast cancer
- ON 123300 differentiated features
 - Also targets ARK5 controlling cellular metabolism and survival
 - Potential to act as single agent
 - May be active in resistant cells

Partnership with HanX Biopharmaceuticals

- License for Greater China
 - Onconova retains ROW rights
- HanX to fund IND-enabling studies
- Upfront, milestones, royalties
- HanX a specialty Oncology company
 - Phase 1 stage PD-1 checkpoint antibody
 - Checkpoint blockade and CDK inhibition believed to be synergistic
- Pre-IND consultation with the FDA
 - Guidance for manufacturing
 - Development plan for an IND application
- Next Milestone is IND
 - US IND anticipated in H1-2019



ONCONOVA BUSINESS DEVELOPMENT OPPORTUNITIES

Patent protected, differentiated small molecule compounds

Compound	Target	Stage	Next Step	Competition	Patents	Licensing Territories Available
<i>Clinical Stage</i>						
Rigosertib	<ul style="list-style-type: none"> RAS pathway MDS initial indication 	Phase 3	Top-line data in 2019	Only HMAs approved for MDS	Worldwide issued and pending	Europe Asia, except Japan and Korea North America
Briciclib	eIF4E (Cyclin D)	Phase I*	Phase II Dose	4EGI-1	Issued US	Worldwide
Recilisib	GSK-3, Akt	Phase I	Primate efficacy	CBLB502	Issued WW	Ex-US rights
<i>Advanced pre-IND stage</i>						
ON 123300	CDK4/6; ARK5	IND in 2019	Toxicology underway	Palbociclib	Issued US, EP	Ex-China rights
<i>Pre-clinical stage</i>						
ON 150030	FLT3 + Src	Pre-clinical	Animal studies	Quizartinib	Issued US, EP	Worldwide
ON 1231320	PLK2	Formulation	Pre-IND	Volasertib	Issued	Worldwide
ON 108600	CK2	Formulation	Pre-IND	CX-4945	Issued	Worldwide
ON 146040	PI3K α/δ	Pre-clinical	Toxicology	IPI-145	In process	Worldwide

**Trial on hold, pending new manufacturing batch*



FINANCIAL DETAILS & SUMMARY

Onconova founded in 1998; public since 2013

Ticker	Nasdaq ONTX	Debt	\$0
Stock Information	<ul style="list-style-type: none"> 5.7 million common shares outstanding 1:15 reverse split effective 9/26 allows for continued Nasdaq listing Public float ~95% YTD average daily volume: 102,000 		Cash and cash equivalents of \$29.5 million as of 06-30-2018
Ownership	683 Capital, EcoR1 Capital, Armistice Capital, Tyndall, Sabby; insiders including Board and management	Burn-rate	~\$5.5 million per quarter over the last 8 quarters
Analyst Coverage*	H.C. Wainwright, Laidlaw, Maxim, Dawson James, Van Leeuwenhoeck Research (VLR)	Partnerships	<ul style="list-style-type: none"> Rigosertib is partnered with SymBio Pharmaceuticals in Japan/Korea and Pint Pharma in Latin America CDK 4/6 & ARK-5 compound ON 123300 partnered with HanX for Greater China Onconova retains rights to the rest of the world

**Reports available upon request*

