ZENITH

Advanced Epigenetic Technology Biotech Showcase Conference San Francisco, CA

January 9, 2017

50 ml

Todays Agenda for Zenith Capital Corp.



- 1. Corporate Profile & Structure
- 2. Epigenetic Mechanism
- **3. Prostate Cancer Rationale**
- 4. Phase 1 Details & Early Results
- 5. Enzalutamide Combination Trial Phase 1b
- 6. Next Steps
- 7. Intellectual Property

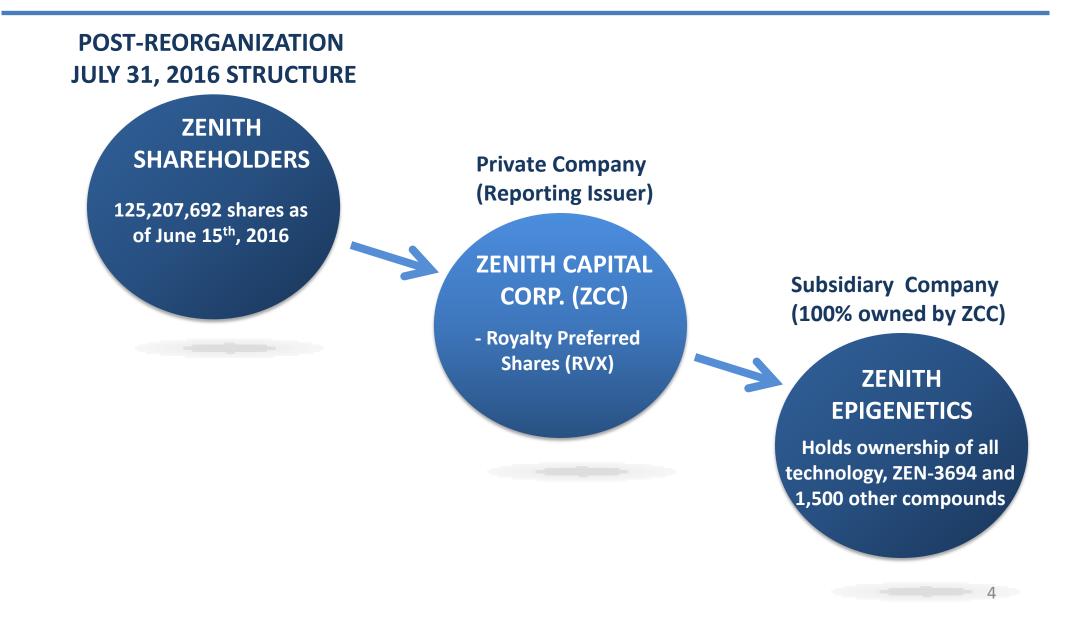


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Suite 300, 4820 Richard Road S.W. Calgary, AB, Canada T3E 6L1
Tel: (403) 254-9252, Fax:(403) 256-8495, http://www.zenithepigenetics.com



Founded	Corporate spin out from Resverlogix in June 2013
Status	Unlisted Possible US market IPO when conditions permit
Cash Raised 2014-2016	Approx. US\$44MM @ \$1.00 USD per share
Enterprise Value est.	\$250 MM
Shares Outstanding	125.2 MM 134.0 MM fully diluted
Cash Burn	\$2 MM per quarter - Current

Post July 31, 2016 Corporate Structure



Historical Timeline & Strategic Progression

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Company formation

Spun out of Resverlogix to focus on oncology/auto-immune

FDA approved IND for **ZEN-3365**

Top investigators & institutions recruited for Phase 1 Jul 2014

Challenges

Prior to the Phase 1 launch, overlapping IP published by another group, ZEN-3365 discontinued

Oct 2014

Change of plans

ZEN-3694 selected as DC, superior properties, mutiple back ups, IP published May 2015

2015 2014 []] \square 2015 2016 **First patient dosed Biology Expansion**⁺ **Financing Focused clinical strategy**

IND accepted

MSKCC/UCSF selected as

Raised \$25M Mar 2016

Jun 2016

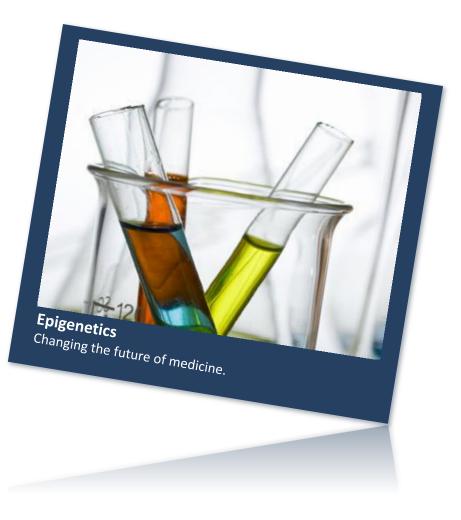
Immuno-Oncology TNBC, NSCLC, ER+ Breast, next indications

Ongoing

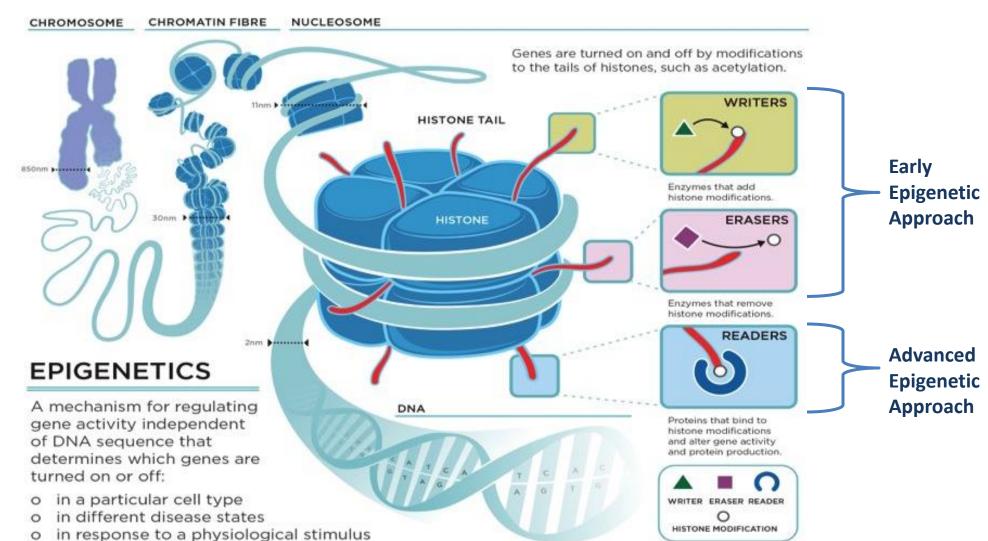
Epigenetics Mechanism



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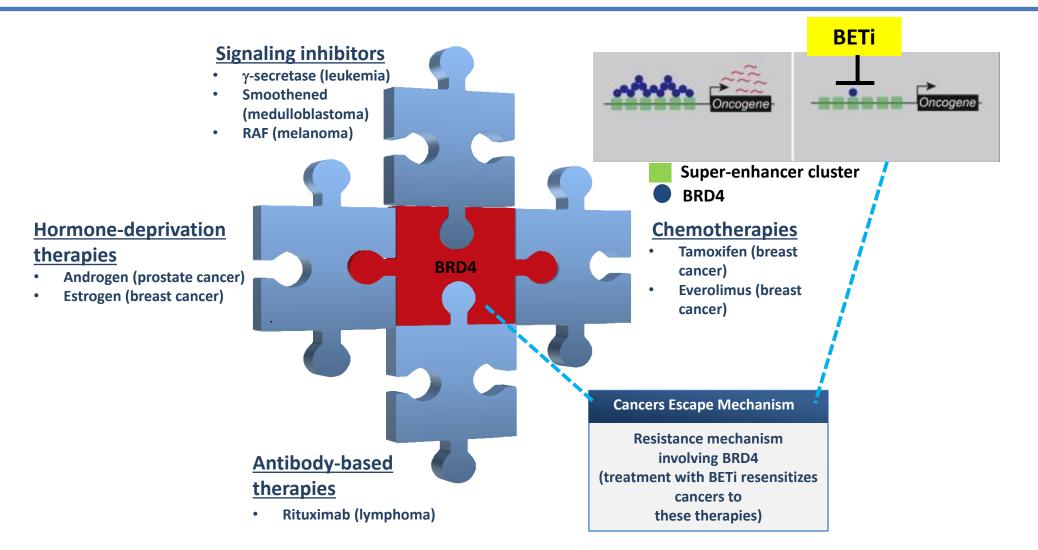


Epigenetics, the Mechanism Behind Our Approach



Zenith's BRD4 Targets Resistance Mechanisms



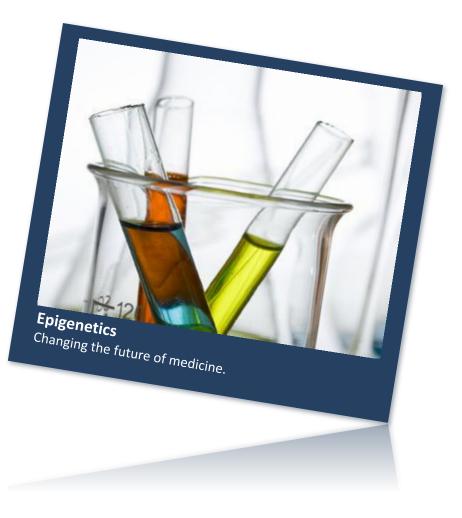


Resistance to several standard of care treatments does not impede sensitivity to BETi

Prostate Cancer Rationale



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Current Market and Unmet Need

- ~135,000 annual mCRPC patients in the US/EU alone majority receive enzalutamide or abiraterone as first-line treatment
- Over \$4B in sales in 2015 for first-line enzalutamide and abiraterone
- Patients become resistant to these therapies, no effective second-line therapy yet
- Continuing high mortality rate in resistant mCRPC (50% 1 year survival, 28% in 5 years)

Opportunity for ZEN-3694

- Second-line single-agent treatment
 - key opinion leaders agree that there is no effective second-line treatment
 - ~60,000 second-line treatment eligible patients in US/EU alone
- Expand into first-line treatment in combination with enzalutamide or abiraterone

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Prostate Cancer Epidemiology & Market



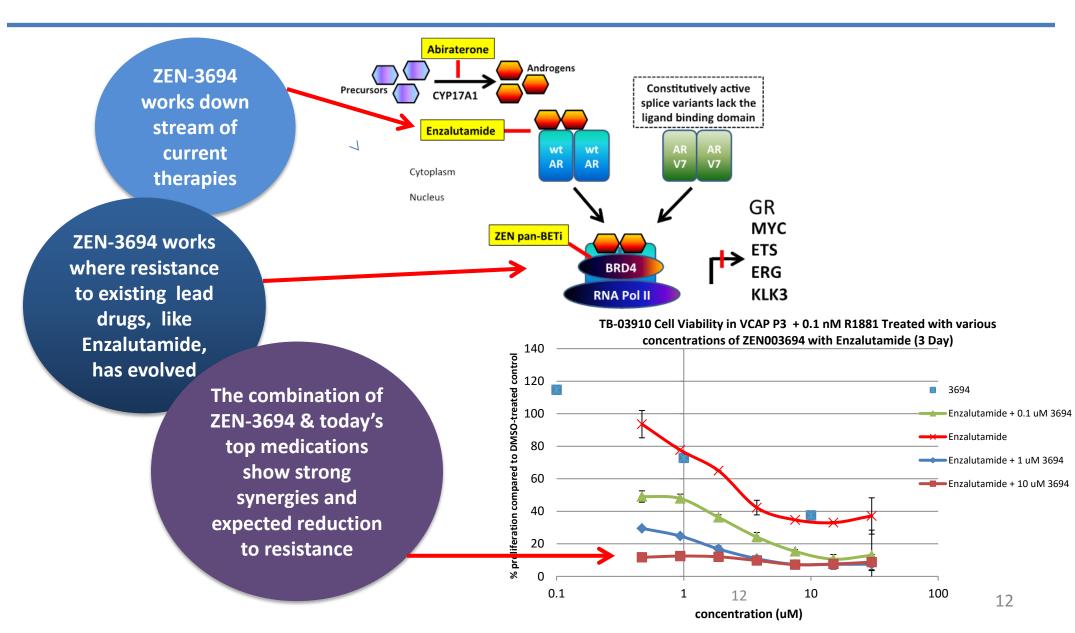
	US/EU Prevalence (2015)	5 Year Survival Rate	Japan Prevalence	Korea Prevalence	China Prevalence
mCRPC	~ 134,000	28%	~26,000	~3300	~16,000



The global prostate cancer WW market is expected to reach \$11B by 2019, driven by Zytiga and Xtandi.

ZEN-3694 Potential in Patients Developing mCRPC Resistance to Enzalutamide





Potential Resistance Pathways in CRPC in response to Enzalutamide and/or Abiraterone

ENZALUTAN	/IIDE/ABIRATERONE	THERAPY	CRPC		
Resistance mechanism	Alterations of AR (Mutations, amplification, splice-variant)	GR up-regulation	<u>NF-kB bone</u> <u>metastasis</u>	Loss of AR	Other mechanisms MYC/MYCN, BCL-2, FOXA1, ERG, ETS, CHD1, SRC, AKR1C3
BETi dependent mechanism	Inhibition of AR signaling	Down-regulation of GR	Inhibition of NF-kB activity	Decrease in NE markers (activity in AR- cells)	BETi downregulates MYC, ERG, BCL-2, BRD4 interacts with ERG

ZEN-3694 shows good efficacy in different CRPC models that are resistant to AR antagonists

Phase 1 Details & Early Results



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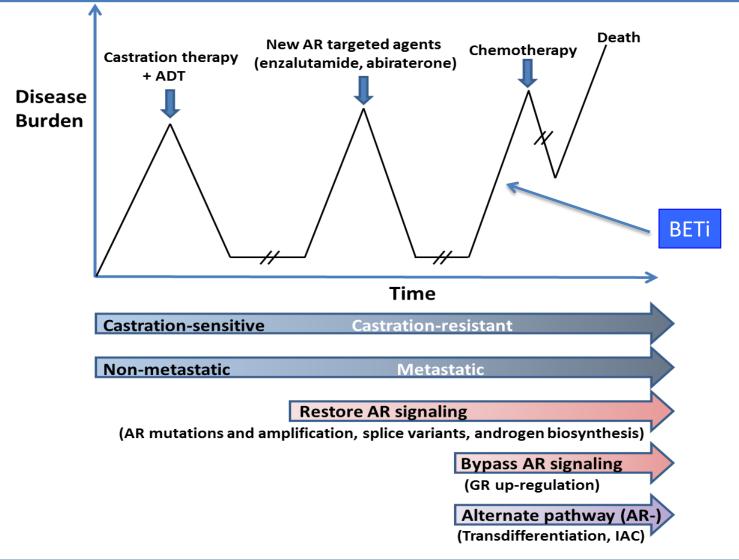


Name	Institution	Comments
Eric Small, MD Chief, Dept. of Medicine	University of California, San Francisco (UCSF)	Developed abiraterone - #2 CRPC drug, owned by J&J.
Rahul Aggarwal, MD Developmental Therapeutics Specialist, Genitourinary Oncologist		
Howard Scher, MD Chief, Genitourinary Oncology Wassim Abida, MD, PhD Medical Oncologist	Memorial Sloane Kettering Cancer Center (MSKCC)	Developed enzalutamide - #1 CRPC drug, now owned by Pfizer. Developing ARN-509 for J&J
Joshi Alumkal, MD Associate Professor	Oregon Health Sciences University (OHSU)	Expert in epigenetics in prostate cancer research
Allan Pantuck, MD Professor, Dept. of Urology	University of California Los Angeles (UCLA)	Involved in enzalutamide and provenge development
Elizabeth Heath, MD Professor, Dept. Hematology/Oncology	Karmanos (Wayne State)	Genitourinary oncology specialist
Tom Fleming, MD Oncologist	Virginia Oncology Associates	Community site for high enrollment

Castration-resistant Prostate Cancer (CRPC)

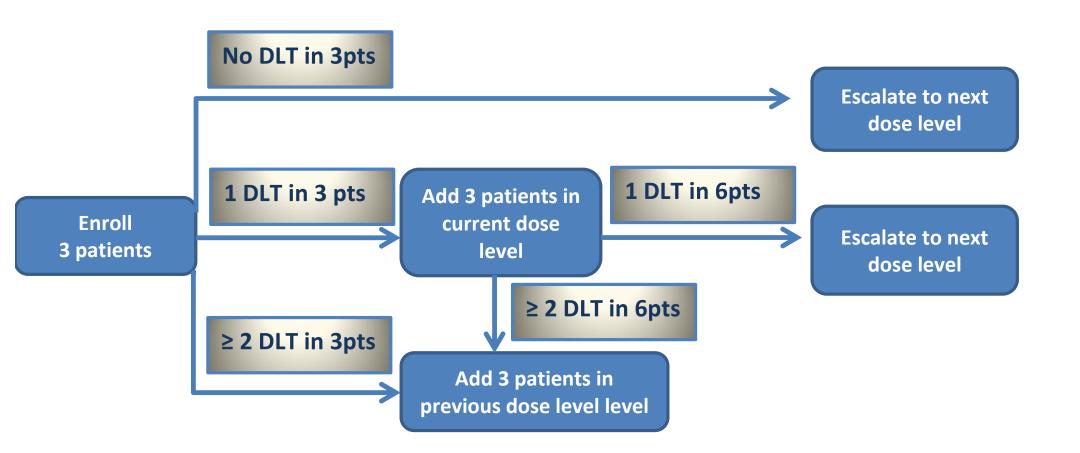


Treatment Algorithm



Medical need for targeting patients resistant to AR targeted agents; Need for targeting downstream AR signaling and alternate resistance pathways

3 + 3 Dose Escalation Design



Note: MTD (Maximum tolerated dose) is the highest dose with <1 DLT in 6 patients. DLT is Dose Limiting Toxicity



Primary

• Safety, tolerability, maximum tolerated dose (MTD), and recommended Phase 2 dose (RP2D) of ZEN-3694

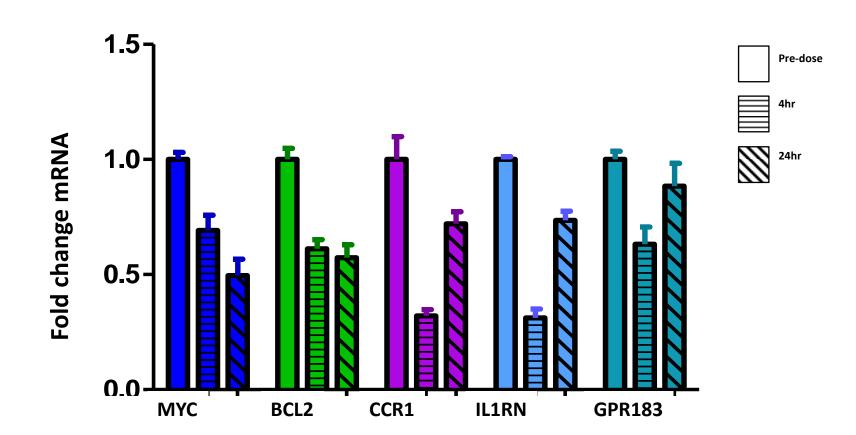
Secondary

- Pharmacokinetics (PK)
- Preliminary clinical activity
 - PCWG2 Criteria: PSA response rate, Radiographic response rate, PFS
 - Circulating Tumor Cell (CTC) response rate



ZEN-3694 single-agent study ongoing

- Maximum Tolerated Dose (MTD) confirmation
- Good PK, exposures reach IC₅₀ cell proliferation values
- Target modulation shown
- On-target safety profile
- Longest patient on drug now beyond 4 months
- Intermittent dosing schedule cohorts may be initiated in Q1/17 for recommended Phase 2 dose (in discussion)

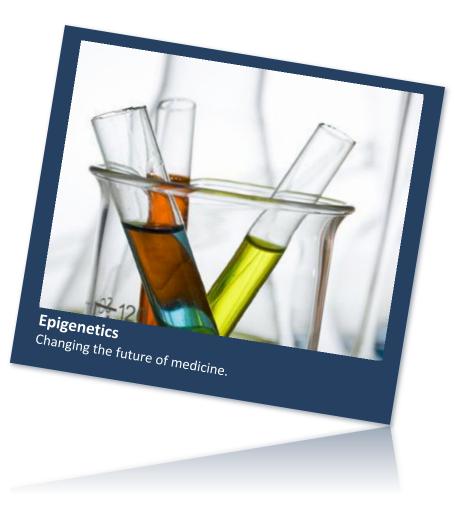


Robust target modulation for 24h

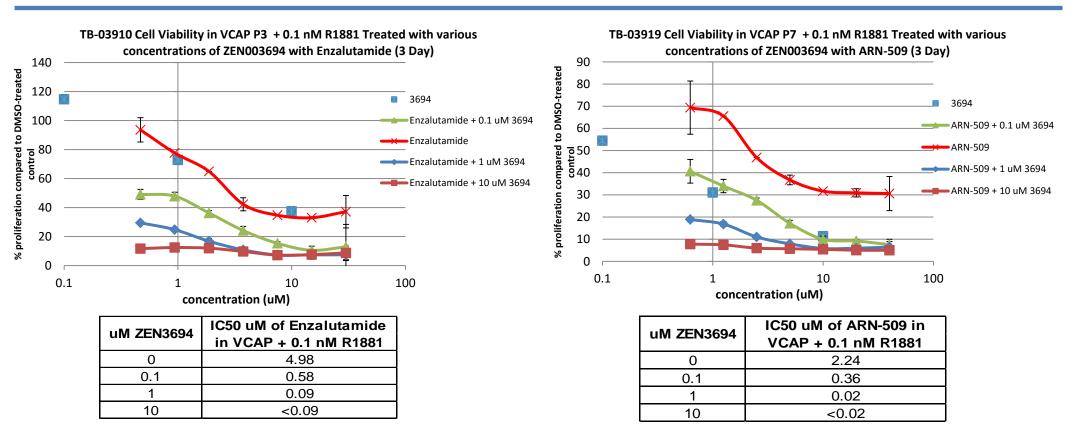
Enzalutamide Combination Trial – Phase 1b



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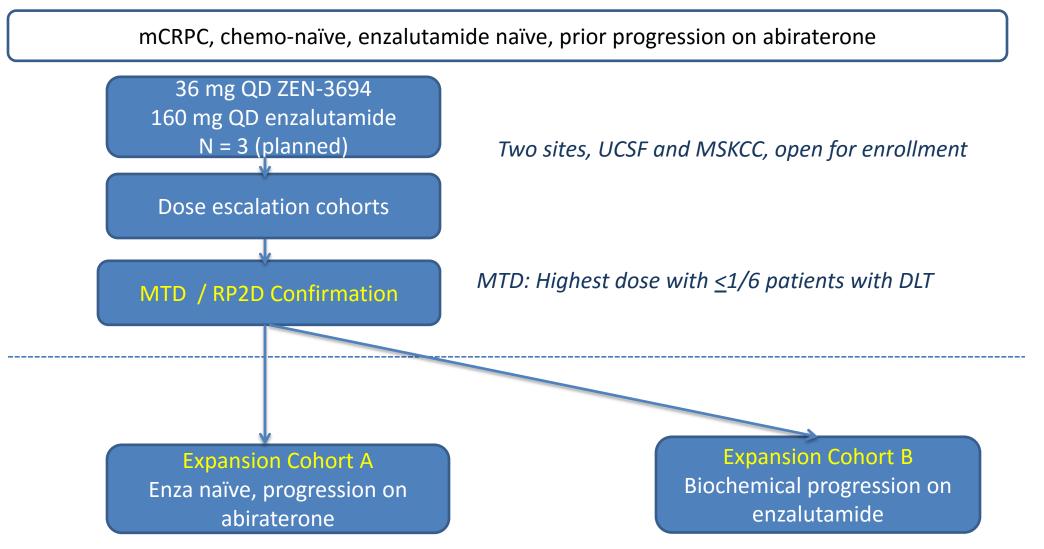
ZEN-3694 Synergizes With Enzalutamide & ARN-509



VCAP curve shift: Enzalutamide and ARN-509 sensitive, ZEN003694 highly synergistic

ZEN-3694 Phase 1b Study Design

Phase 1b, open label, combination, 3x3 dose escalation/confirmation



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Primary

• Safety, tolerability, MTD, and RP2D of ZEN-3694 in combination with enzalutamide

Secondary

- Pharmacokinetics (PK) af ZEN-3694 and enzalutamide when given in combination
- Preliminary clinical activity
 - PCWG2 Criteria: PSA response rate, Radiographic response rate, Median PFS
 - Circulating Tumor Cell (CTC) response rate

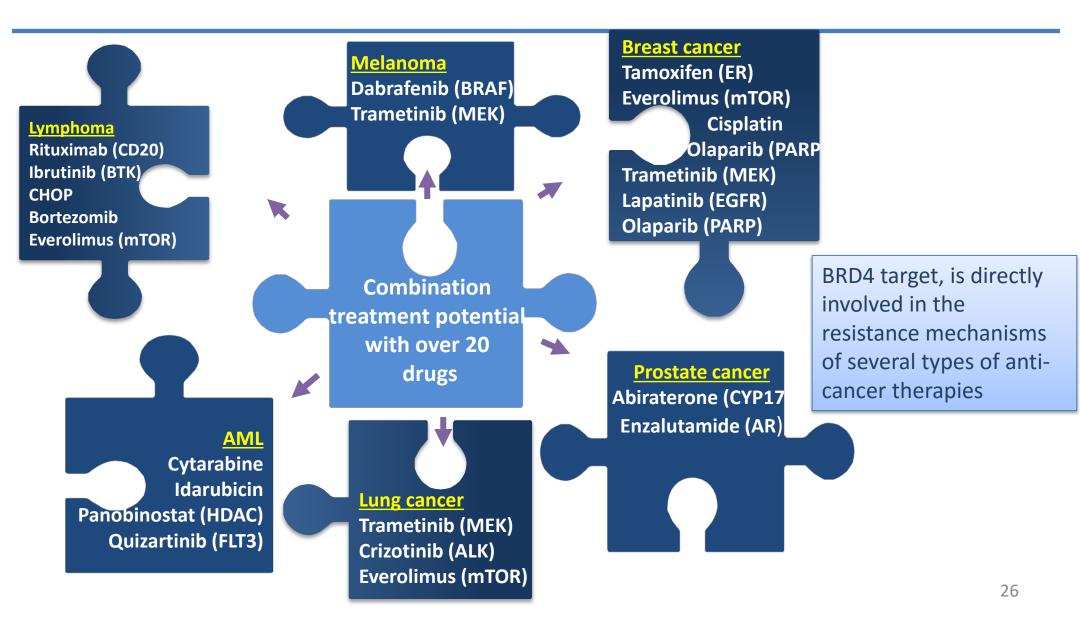
Next Steps



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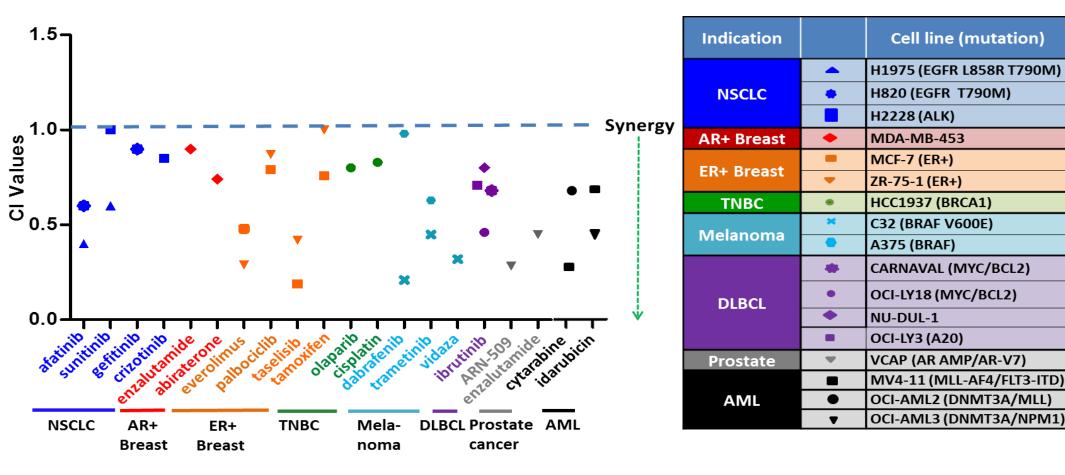


BET Inhibitors Potential as Combination Agents

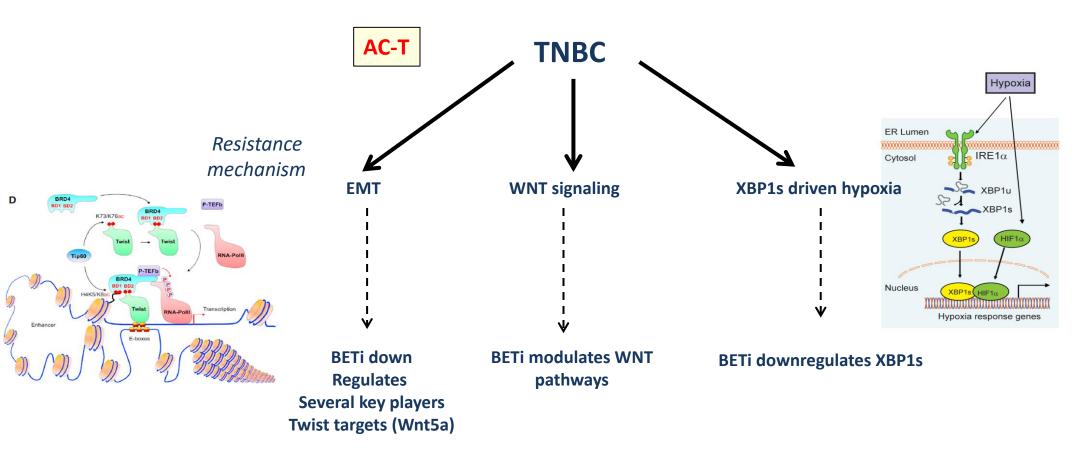


ZEN-3694 Synergizes With Several Standard of Care Cancer Drugs



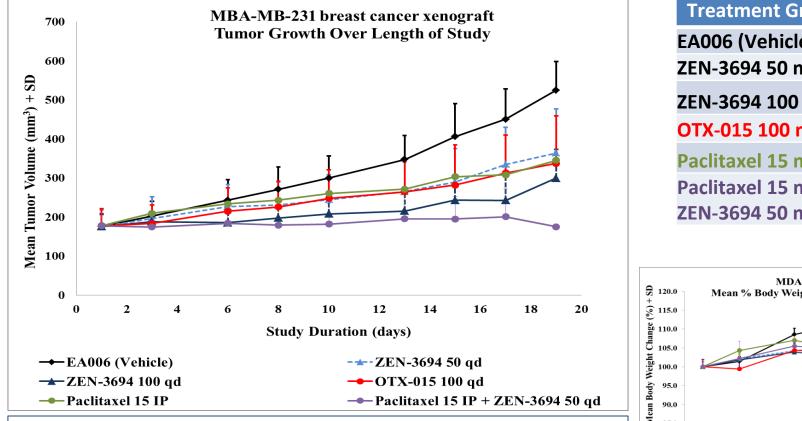


Potential Resistance Pathways in TNBC in Response to Chemotherapy



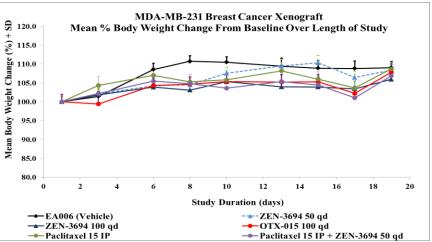
Rationale for ZEN-3694 to show activity in different TNBC models that are resistant to chemotherapy

ZEN-3694 is Synergistic With Paclitaxel in Triple-negative Breast Cancer Models



- Combination regimen is well tolerated
- ZEN-3694 is more potent than OTX at equivalent dose
- ZEN-3694 is synergistic in combo with Paclitaxel (5/12 regressed tumors)

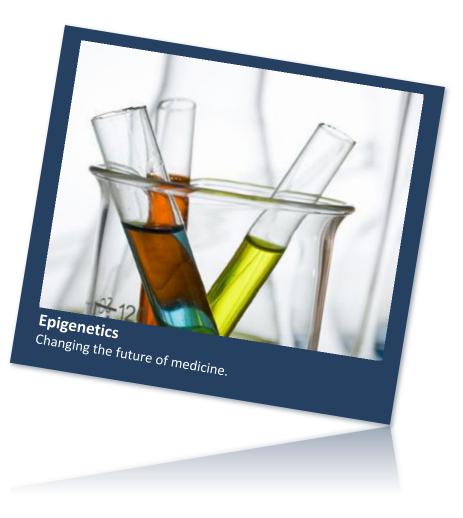
Treatment Groups	TGI
EA006 (Vehicle)	0%
ZEN-3694 50 mg/kg qd	46%
ZEN-3694 100 mg/kg qd	64%
OTX-015 100 mg/kg qd	54%
Paclitaxel 15 mg/kg IP	52%
Paclitaxel 15 mg/kg IP + ZEN-3694 50 mg/kg qd	101%



Intellectual Property



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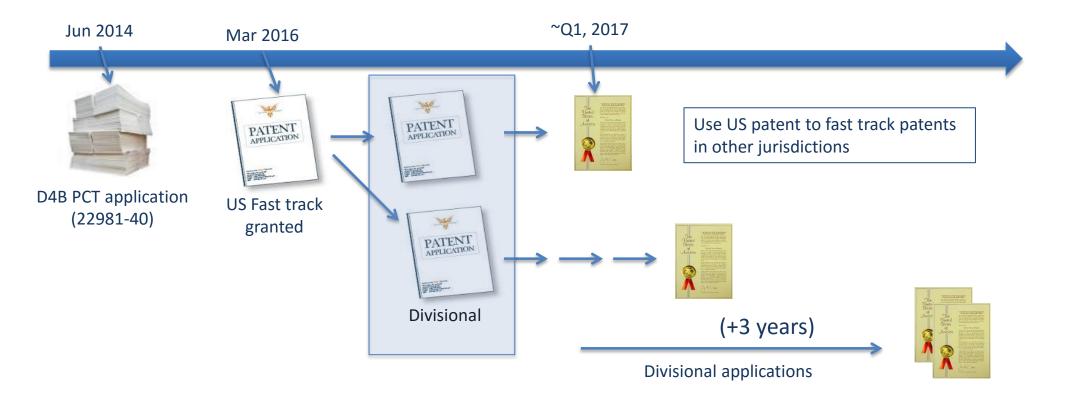


Zenith Epigenetics Ltd. owns numerous patent families, including three issued US patents and 60 pending applications. The portfolio includes a number of US applications and world-wide equivalents.

Composition Patents							
Zenith Reference Number	Provisional Patent Application	Patent Application	Publication	National Stage	Examination	Issuance	
22981-36							
22981-37						2x	
22981-38							
22981-39							
22981-40							
22981-41							
22981-45							
22981-46				National phase mid 2017			
22981-47				National phase mid 2017			
22981-49				National phase mid 2017			
22981-50				National phase mid 2017			
22981-51				National pha	ase mid 2017		
22981-57		Convert mid Dec.2016				21	

ZEN-3694 Patent Application





How can <u>you</u> determine the true potential of a new clinical drug candidate in oncology?



There are hundreds of biotech companies with potential drug candidates

Drug candidates require 3rd party Principle Investigators (PI's) to act as independent clinical investigators

The best oncology units and principle investigators in the United States are highly sought after

Zenith's cutting edge technology has attracted the top two U.S. PI's in prostate cancer research as well as the Prostate Cancer Clinical Trials Consortium (PCCTC)

Zenith has confirmed PI's -Dr. Eric Small - Univ. of California, San Francisco -Dr. Howard Scher at Memorial-Sloan Kettering, NY Both Dr.'s Small & Scher were involved in the development of the top 2 current prostate cancer drugs in use abiraterone & enzalutamide respectively

Four of the last five FDA approved prostate drugs have come from the PCCTC which is highly selective and only champions the most promising drugs

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