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Syndax investment highlights

Entinostat IO

Combined with anti-PD-1:

- > Signal in melanoma
- Expanded into CRC
- Ongoing trials in NSCLC, Mel, TNBC, Ovar
- Multiple near term readouts

Entinostat HR+ Breast Cancer

Combined with exemestane:

- Breakthrough designation
- Phase 3 ongoing

SNDX-6352

CSF1-R antibody:

- Phase 1 ongoing
- Broad clinical potential

Strong management team and cash position

CRC - colorectal cancer; NSCLC - non-small cell lung cancer; Mel - melanoma; TNBC - triple negative breast cancer; Ovar - ovarian cancer



Recent progress on ENCORE 601

Enrollment accelerated

Melanoma cohort enrollment anticipated to complete in 3Q17 (vs. 4Q17) ASCO abstract presentation

Will highlight data from phase 2, stage 1 melanoma cohort

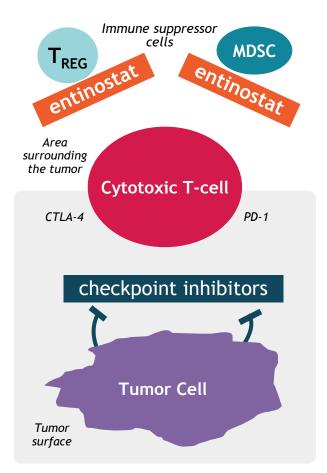
Expanding ENCORE 601

Now includes microsatellite stable colorectal cancer (MSS CRC)

Strong rationale for combining entinostat with PD-1 antagonists

Entinostat

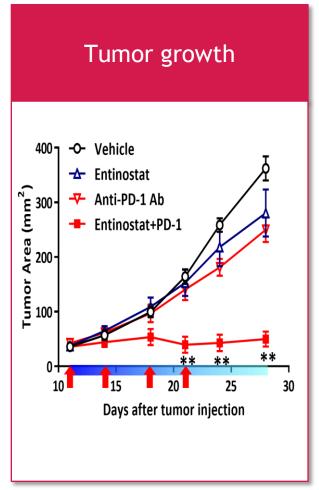
- Class I selective HDAC inhibitor
- Oral, once weekly
- Well tolerated in combinations
- Blocks MDSCs and Tregs
- Preclinical efficacy combined with anti-PD-1

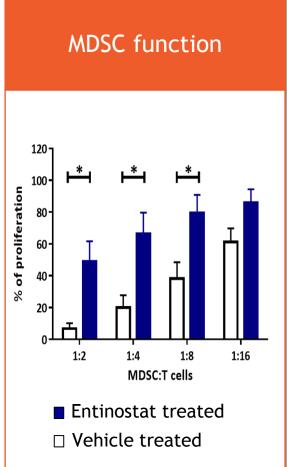


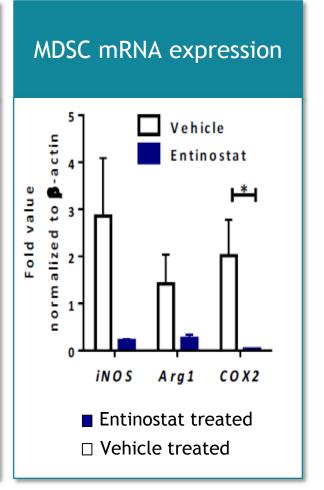
Hypothesis: Entinostat can reverse resistance to PD-1 antagonists

HDAC - histone deacetylase; MDSC - myeloid derived suppressor cell; Treg - regulatory T lymphocyte

Entinostat synergizes with anti-PD-1 through MDSC inhibition







Hashimoto et al AACR 2016

ENCORE 601/Keynote 142 expanded to include patients with colon cancer

Entinostat + KEYTRUDA®

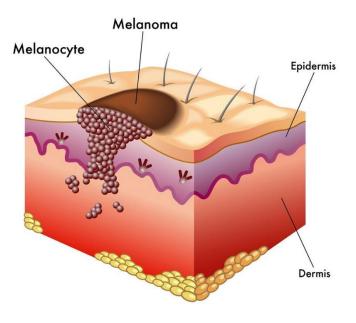
Phase 2: Simon 2-stage design

		STAGE 1	Minimal threshold to advance to Stage 2	STAGE 2 Total complete enrollment enroll	
NSCLC -	PD-(L)1 Naive	13 patients	3 responses*	Add 33 patients 46	5
	PD-(L)1 Pretreated	20 patients	2 responses*	Add 36 patients 56	5
MEL	PD-1 Pretreated	13 patients	2 responses*	Add 21 patients 34	4
CRC	PD-(L)1 Naive	13 patients	2 responses*	Add 21 patients 34	4

^{*} Response defined as confirmed PR or CR

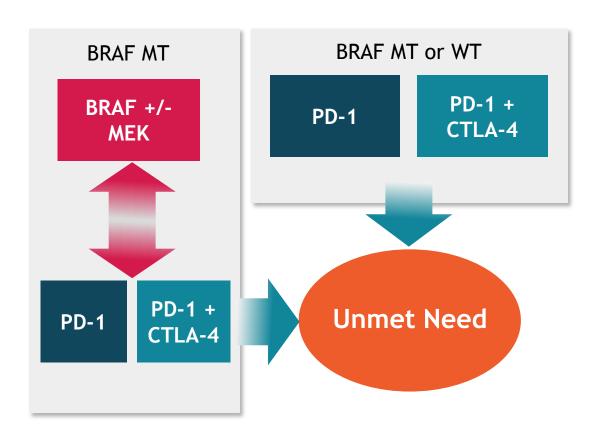
Entinostat-PD-1 antagonist promising combination for PD-1 resistant melanoma

Encore 601 melanoma cohort milestones



1Q17	Re-opened enrollment in cohort
	UPCOMING MILESTONES
2Q17	Present stage 1 data at ASCO
2Q17	FDA meeting; development path
3Q17	Complete stage 2 enrollment

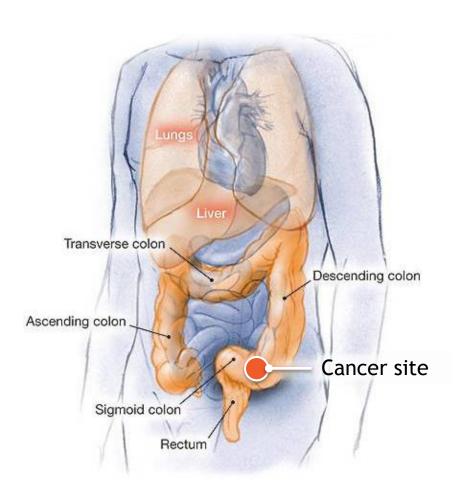
Despite important advances, unmet need exists for melanoma patients



- ENCORE 601 enrolls patients progressed on a PD-1 antagonist
- Approx. 10,000 US
 patients expected to
 require treatment
 after PD-1 antagonist

Source: Trial Trove, SEER data, Kantar 2016 Treatment Architecture report

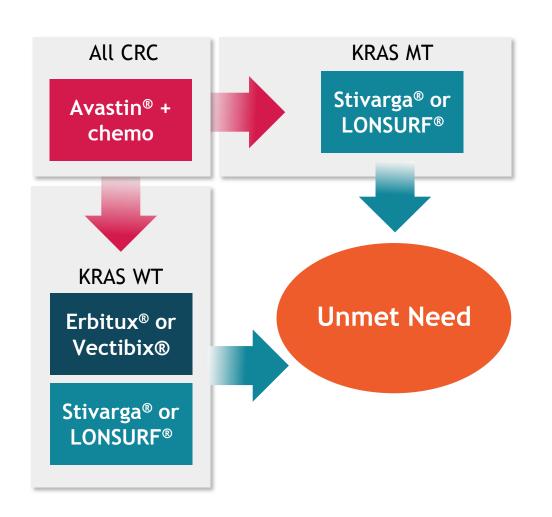
Rationale for expansion into CRC



- High unmet need population
- ENCORE 601 melanoma results suggest potential to impact immune subtypes¹
- Rapid read out

1. Dienstmann, R., Nat Rev Cancer, 2017 17, 79 - 92

Need to improve therapy for MSS CRC patients

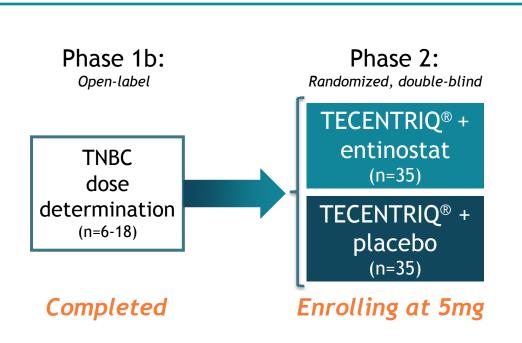


 Approx. 23,000 3L (8,000 4L) treated patients are MSS¹

 PD-(L)1 monoTx has shown minimal activity in MSS CRC²

- 1. Trial Trove, SEER data, DataMonitor, Kantar 2016 Treatment Architecture report; assumes 85% of CRC are MSS
- 2. Abstract LBA100 ASCO 2015, Abstract 479P ESMO 2016; Abstract 3502 ASCO 20161

ENCORE 602: Entinostat + TECENTRIQ® for TNBC





Primary Endpoints

Phase 1b: Phase 2 dose

Phase 2: PFS

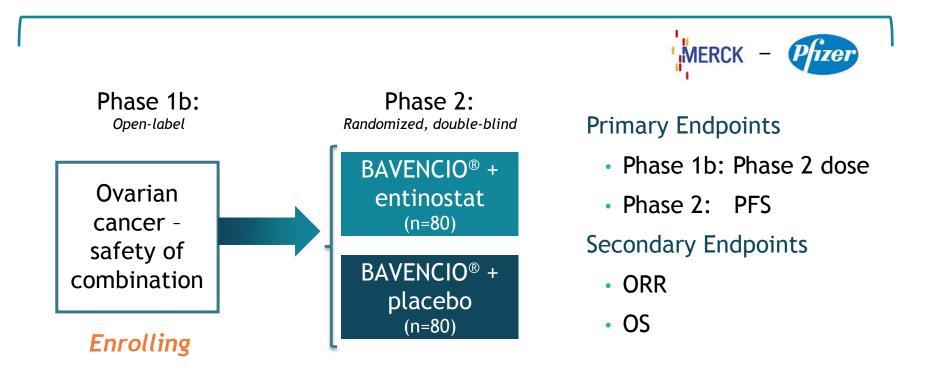
Secondary Endpoints

- Overall response rate (ORR)
- Overall survival (OS)

Primary Trial Center: UCLA Health CRO: Translational Research in Oncology Group (TRIO)

Target enrollment completion end of 2017

ENCORE 603: Entinostat + BAVENCIO® for ovarian cancer

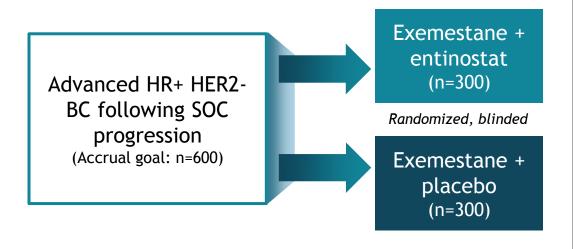


Anticipate Phase 2 to begin in 3Q17

E2112: Phase 3 registration trial in advanced HR+, HER2- breast cancer

Exemestane +/- entinostat





Treatment cycle (28 days)

- Exemestane (25mg): PO, days 1-28
- Entinostat or placebo (5mg):
 PO, days: 1, 8, 15, 22

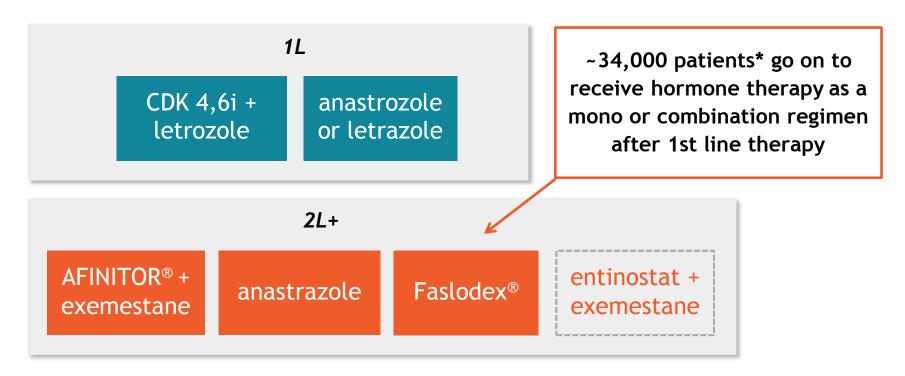
GRANTED FDA
BREAKTHROUGH THERAPY
DESIGNATION

- Two primary endpoints PFS and OS
- Potential NDA filing 2018 based upon positive PFS data
- Per ECOG-ACRIN, enrollment completion and PFS data analysis anticipated in 1H18

Entinostat: Blockbuster potential as 2nd/3rd line therapy for HR+, HER2- metastatic breast cancer

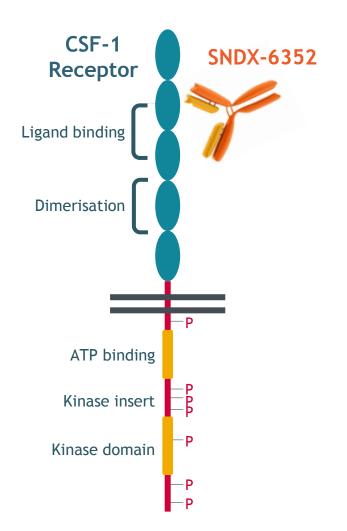
First novel MOA in HR+ BC with Phase 3 data since CDK4/6

Leading treatment options - HR+, HER2- advanced breast cancer



Source: DataMonitor 2016 Breast cancer: HR+/HER2- Disease Coverage Report

SNDX-6352: Anti-CSF-1R Ab targeting TAMs to increase tumor infiltrating lymphocytes



- High affinity, IgG4
 (K_D = 4-8 pM)
- Broad potential clinical utility
- Phase 1, single ascending dose
 (SAD) trial initiated 4Q16
 - First 3 cohorts completed dosing
- Initiate multiple ascending dose (MAD) trial (cancer patients) 3Q17

TAM - tumor associated macrophage; CSF-1R - colony stimulating factor -1 receptor

Source: Ordentlich, P. et al SITC 2016

1Q17 Financial highlights and 2017 guidance

Ticker	SNDX (NASDAQ)				
As of March 31, 2017					
Cash and short-term investments	\$92.8 million				
Common shares O/S	18.2 million				
2017 Operating Expense Guidance					
	<u>Q2</u> <u>2017</u>				
R&D	\$11-13 M \$52-57 M				
Total Operating Expen	ses \$15-17 M \$68-73 M				

Upcoming milestones

ENTINOSTAT (Class 1 specific HDAC inhibitor)		2Q17	3Q17	4Q17	1H18
ENCORE 601 - NSCLC (PD-1 preTx) decision to re-open Phase 2					
ENCORE 601 - NSCLC (PD-1 naive) decision to re-open Phase 2					
ENCORE 601 - Complete enrollment in melanoma cohort (stage 2)			•	-	
ENCORE 601 - Present stage 1 melanoma data at ASCO					
ENCORE 601 - FDA Type B meeting melanoma development path					
ENCORE 602 - Complete enrollment Phase 2 TNBC study					
E2112 - Per ECOG, complete Phase 3 enrollment; release PFS					

SNDX-6352 (anti-CSF-1R mAB)	1Q17	2Q17	3Q17	4Q17	1H18
Initiate MAD trial (cancer patients)					
Anticipate SAD trial data presentation (healthy volunteers)					

> NEW Milestone

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