

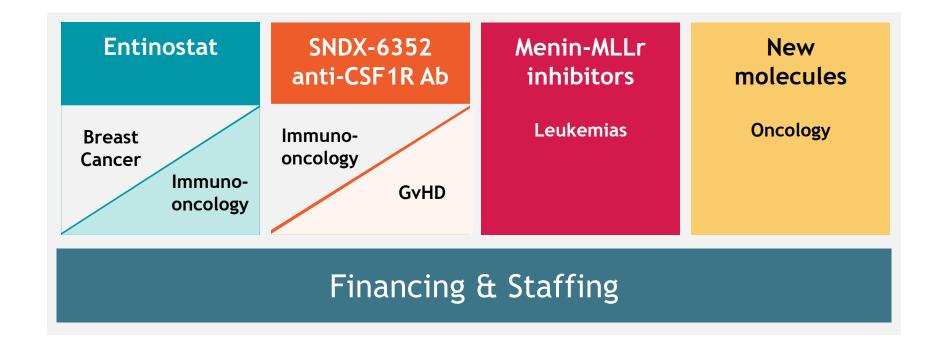


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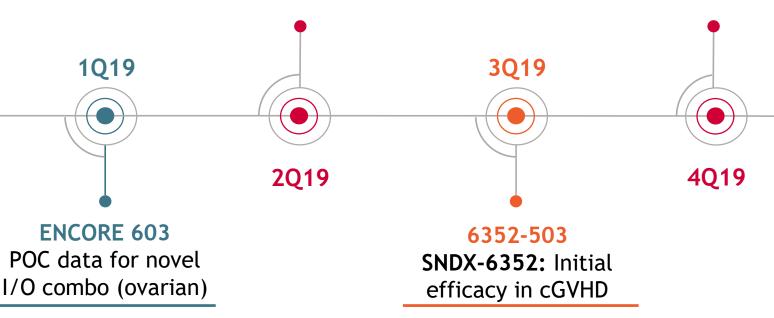
## **Company strategy**



## Potential value drivers throughout 2019

- E2112: Potential for positive overall survival in \$\$B market (HR+ mBC)
- ENCORE 602: POC data in novel I/O combo (TNBC)

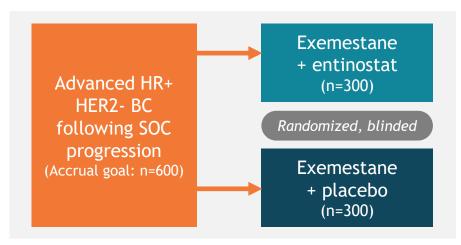
- E2112: Potential for positive overall survival in \$\$B market (HR+ mBC)
- SNDX-5613: Initial clinical efficacy in AML



Triple negative breast cancer (TNBC); acute myeloid leukemia (AML); chronic graft versus host disease (cGVHD)

#### Phase 3 E2112: Focused on overall survival

#### E2112: Exemestane +/- entinostat



Primary endpoint: OS



#### **E2112 Trial Milestones**



4Q18: Accrual completes, PFS and interim OS analyses shared

**2Q19:** Next interim OS analysis

**4Q19:** Additional interim OS analysis

**2020:** Final OS analysis (if needed)







A positive OS result allows filing for full regulatory approval

## Blockbuster potential as 2<sup>nd</sup>/3<sup>rd</sup> line agent

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

1<sup>st</sup> line hormone Tx

Anastrozole or letrazole +/CDK4,6 inhibitor

2<sup>nd</sup>/3<sup>rd</sup>/4<sup>th</sup> line hormone Tx

Anastrazole, Faslodex +/- CDK4,6 inhibitor or Afinitorexemestane

34,000 pts

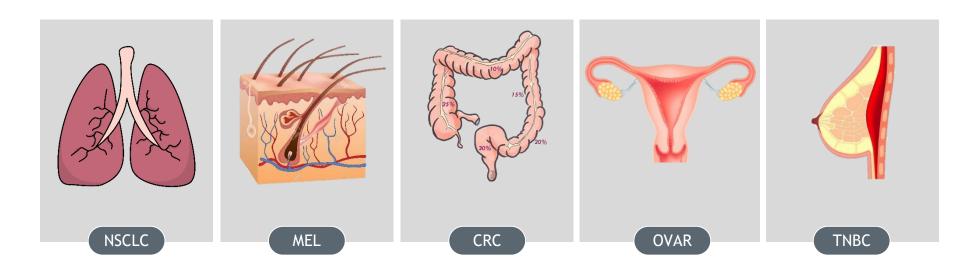
Entinostat-exemestane target population

Chemo-Tx

Capecitabine, gemcitabine, eribulin

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

# ENCORE Clinical Trial Program: Evaluating entinostat's potential to enhance anti-PD-(L)1 efficacy



PD-(L)1

Immune cells

Tumor mutational burden (TMB)

Nanostring

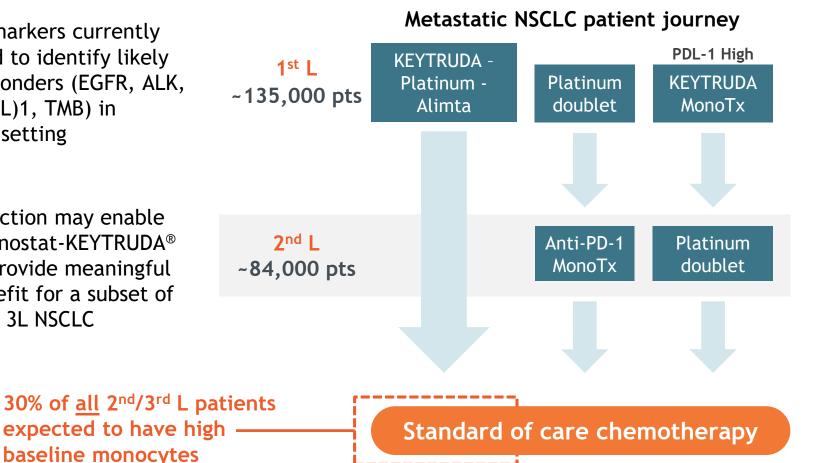
Focused on early signs of efficacy and biomarkers that predict clinical benefit

## Alternative treatment options required to benefit significant NSCLC patient population

Biomarkers currently used to identify likely responders (EGFR, ALK, PD-(L)1, TMB) in this setting

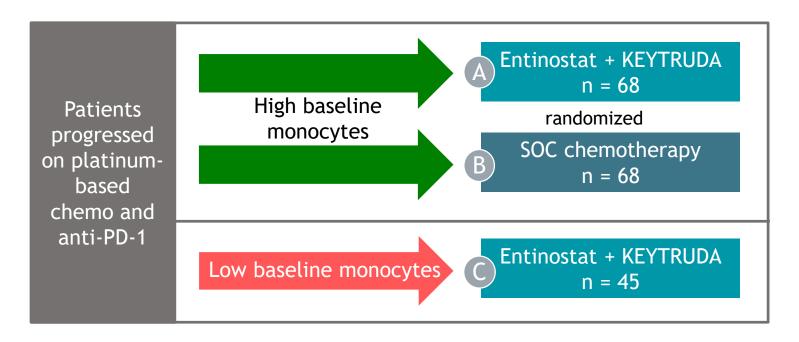
 Selection may enable entinostat-KEYTRUDA® to provide meaningful benefit for a subset of 2L / 3L NSCLC

(~25,000 pts)



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

## Proposed NSCLC trial to validate monocyte-based selection and confirm benefit of ENT-KEYTRUDA



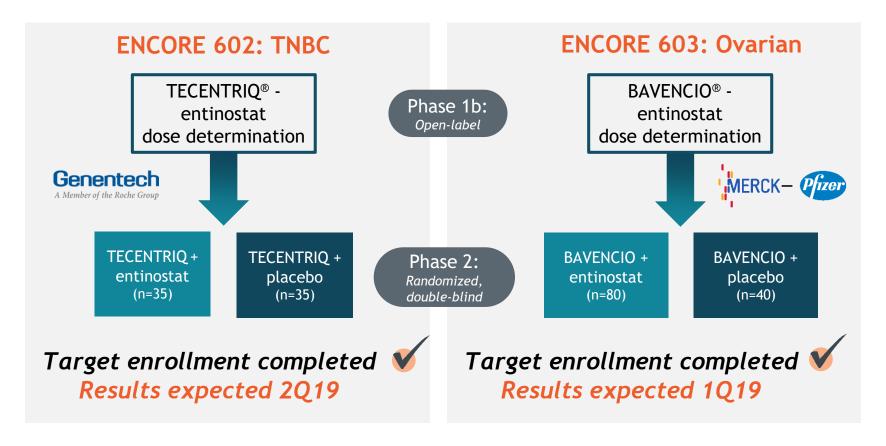
#### Primary Endpoint: PFS

- High baseline monocytes compared to low baseline monocytes
- Entinostat + KEYTRUDA compared to SOC chemotherapy

Secondary endpoints: ORR, DOR, OS

Progression free survival (PFS), objective response rate (ORR), duration of response (DOR), overall survival (OS)

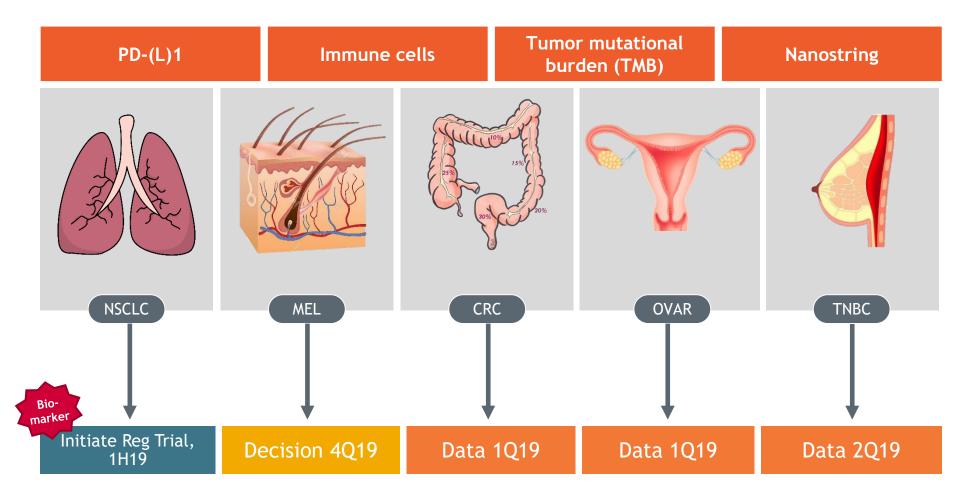
# ENCORE 602, 603: Near-term topline data readouts



#### Phase 2 Endpoints:

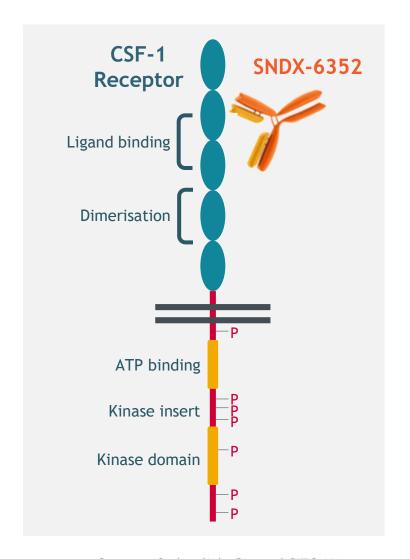
- Primary endpoint PFS
- Secondary endpoint Overall response rate (ORR)
- Secondary endpoint Overall survival (OS)

# ENCORE Clinical Trial Program: Evaluating entinostat's potential to enhance anti-PD-(L)1 efficacy



## Update on SNDX-6352: pursuing novel indication

- High affinity, IgG4
   (K<sub>D</sub> = 4-8 pM)
- Multiple ascending dose (MAD, solid tumors) ongoing
- Combination study with IMFINZI (durvalumab, AZ) commenced
  - RP2D expected in 2Q19
- Chronic graft versus host disease (cGVHD) study initiated
  - RP2D expected in 2H19



Source: Ordentlich, P. et al SITC 2016

CSF-1R - colony stimulating factor -1 receptor; RP2D - recommended Phase 2 dose

### SNDX-5613 (Menin-MLLr inhibitor) IND expected 2Q19

#### MLL-r driver of subset of AML, and ALL

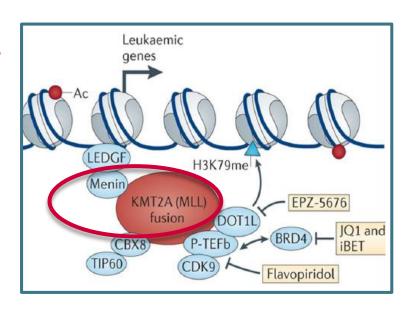
WW incidence: 4,000/yr

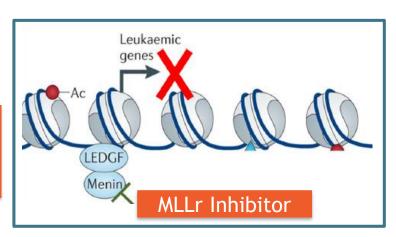
#### NPM1<sup>mut</sup> driver of a subset of AML

WW incidence (30% of AML): ~25,000/yr

MLLr and NPM1 leukemias sensitive to Menin-MLLr inhibition

Development program to be featured in two presentations at ASH Annual Meeting





### 3Q18 financial highlights and 4Q18, 2019 guidance

Ticker	SNDX (N	SNDX (NASDAQ)		
As of Septemb	er 30, 2018			
Cash and short-term investments	\$89.6 million			
Shares Outstanding*	26.1 million			
4Q 2018 and 2019 Operating Expense Guidance				
	Q4 2018	2019		
Research and Development	\$13 - 15 M	\$54-58 M		
Total Operating Expenses^	\$17 - 19 M	\$68-73 M		

 $<sup>^*</sup>$  Includes 24.1 million common shares and pre-funded warrants to purchase 2.0 million common shares  $^{\hat{}}$  Includes \$1.5 and \$6 million non-cash stock compensation expense for Q4 2018 and for 2019, respectively

## **Upcoming milestones**

ENTINOSTAT (Class 1 specific HDAC inhibitor)	4Q18	1Q19	2Q19	2H19
E2112 - Interim OS analysis				
ENCORE 601 - Registration trial decision for melanoma				
ENCORE 601 - Go / No go decision, Stage 1 of MSS CRC cohort				
ENCORE 603 - Report topline ovarian results				
ENCORE 602 - Report topline TNBC results				
SNDX-6352 (anti-CSF-1R mAB)	4Q18	1Q19	2Q19	2H19
SNDX-6352 (anti-CSF-1R mAB)  Identify recommended Phase 2 dose and schedule	4Q18	1Q19	2Q19	2H19
	4Q18	1Q19	2Q19	2H19
Identify recommended Phase 2 dose and schedule Preliminary efficacy in chronic GVHD			•	•
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