

Determined to realize a future in which people with cancer live longer and better than ever before



4Q18 EARNINGS PRESENTATION | MARCH 2019

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2019: Portfolio prioritization to drive value



Entinostat - exemestane

Oral, Class I HDAC in HR+ mBC

- Potential positive OS data 2019
- Efficacy post-CDK4,6 Tx
- Potential NDA filing in 2019/20
- Blockbuster potential

Potential first combo to demonstrate survival benefit

SNX-5613

Oral, Menin inhibitor

- Blocks MLL-fusion protein activity
- IND filing 2Q, clinical data '19/'20
- Benefit expected in high need AML, ped ALL populations
- Blockbuster potential

Targeted therapy provides fast to market opportunity

Entinostat - KEYTRUDA

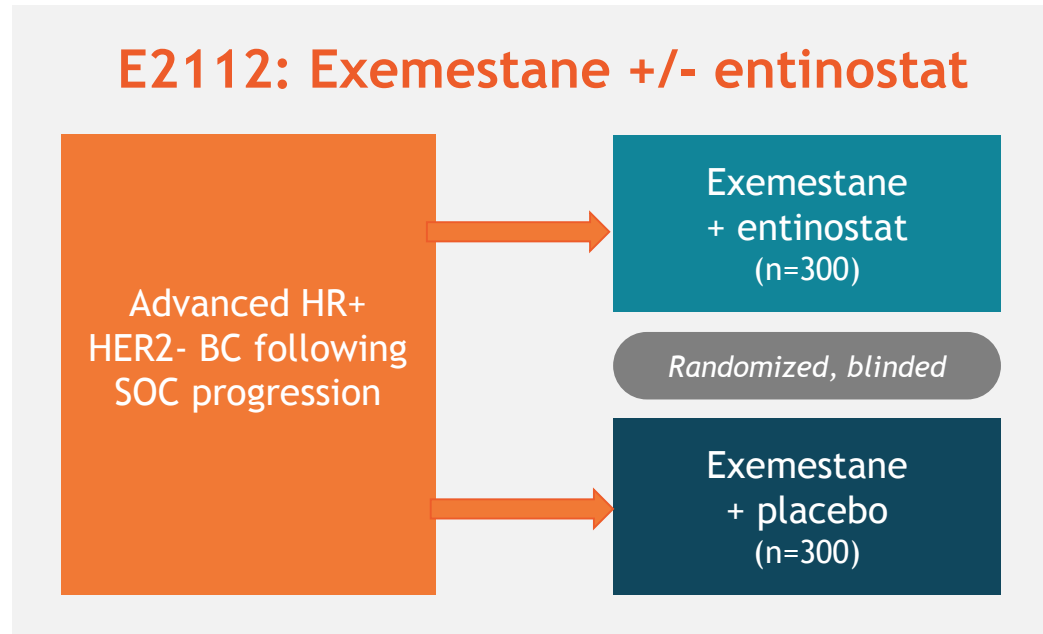
Oral, Class I HDAC, solid tumors

- Strong data in NSCLC and MEL
- 2 oral presentations at AACR
- Blockbuster potential

Registration trial deferred until positive OS in E2112

NSCLC - non-small cell lung cancer; MEL - melanoma; AML - acute myeloid leukemia; ALL - acute lymphoblastic leukemia

Phase 3 E2112: Focused on overall survival



Primary endpoint: OS



E2112 Trial Milestones

- ✓ **4Q18:** Accrual completed (n=608), PFS and interim OS analyses shared
- **2Q19:** Next interim OS analysis
- **4Q19:** Additional interim OS analysis
- **2Q20:** Final OS analysis (if needed)

Expect to file NDA ~6 months after positive OS data

A positive OS result allows filing for full regulatory approval

Blockbuster potential as 2nd/3rd line agent

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

1st line hormone Tx

Anastrozole or
letrozole +/-
CDK4,6 inhibitor

2nd/3rd/4th line hormone Tx

Anastrozole, Fulvestrant +/-
CDK4,6 inhibitor or
Afinitor-exemestane

34,000 pts

Entinostat-exemestane
target population

Chemo-Tx

Capecitabine, gemcitabine,
eribulin

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

SNDX-5613 targets novel fusion protein: Fusion proteins proven to be good candidates for targeted therapies

Advantages

- Strong target validation
- Precise patient selection
- Big effect in small studies
- Molecular markers of disease status
- Rapid regulatory path

Therapies targeting fusion proteins

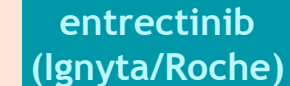
BCR-ABL



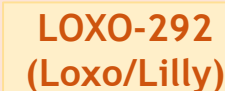
EML4-ALK



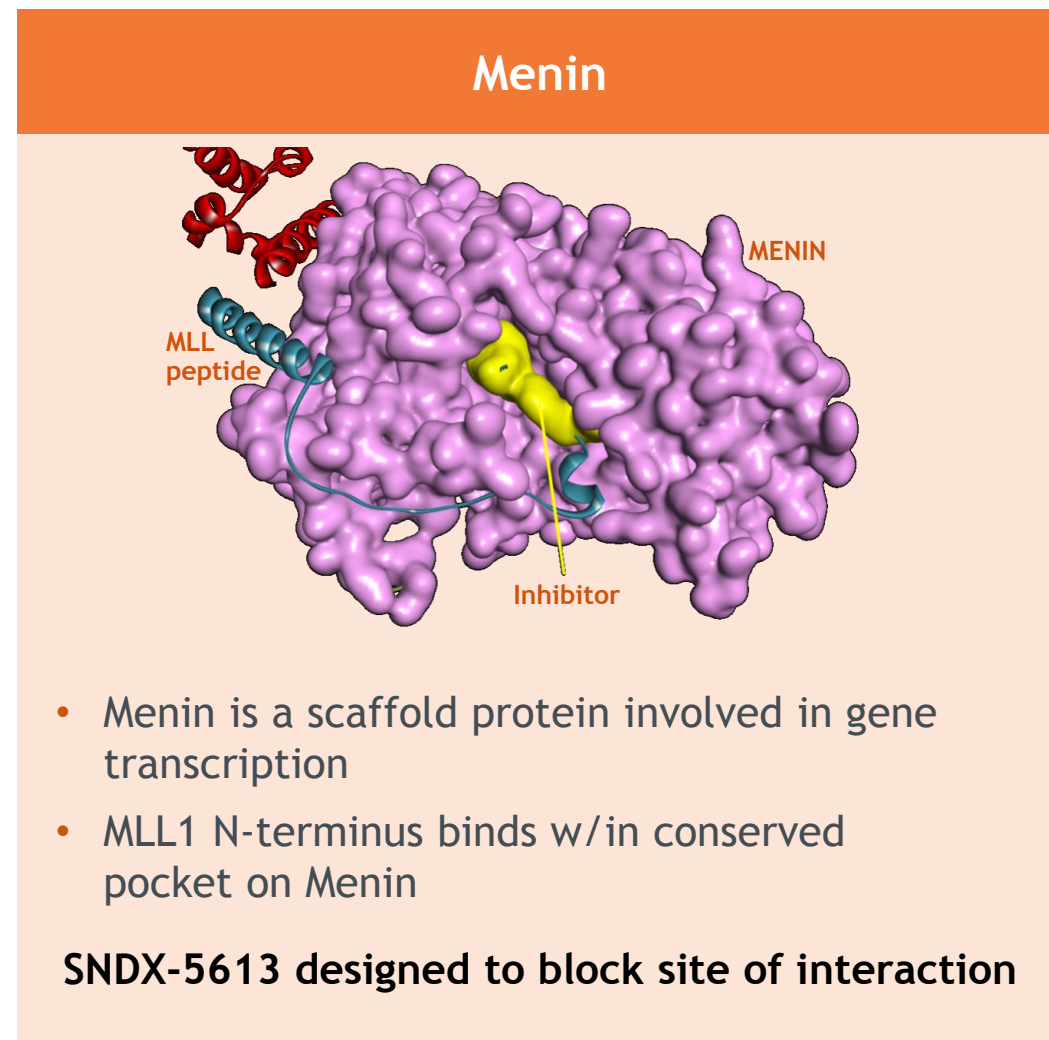
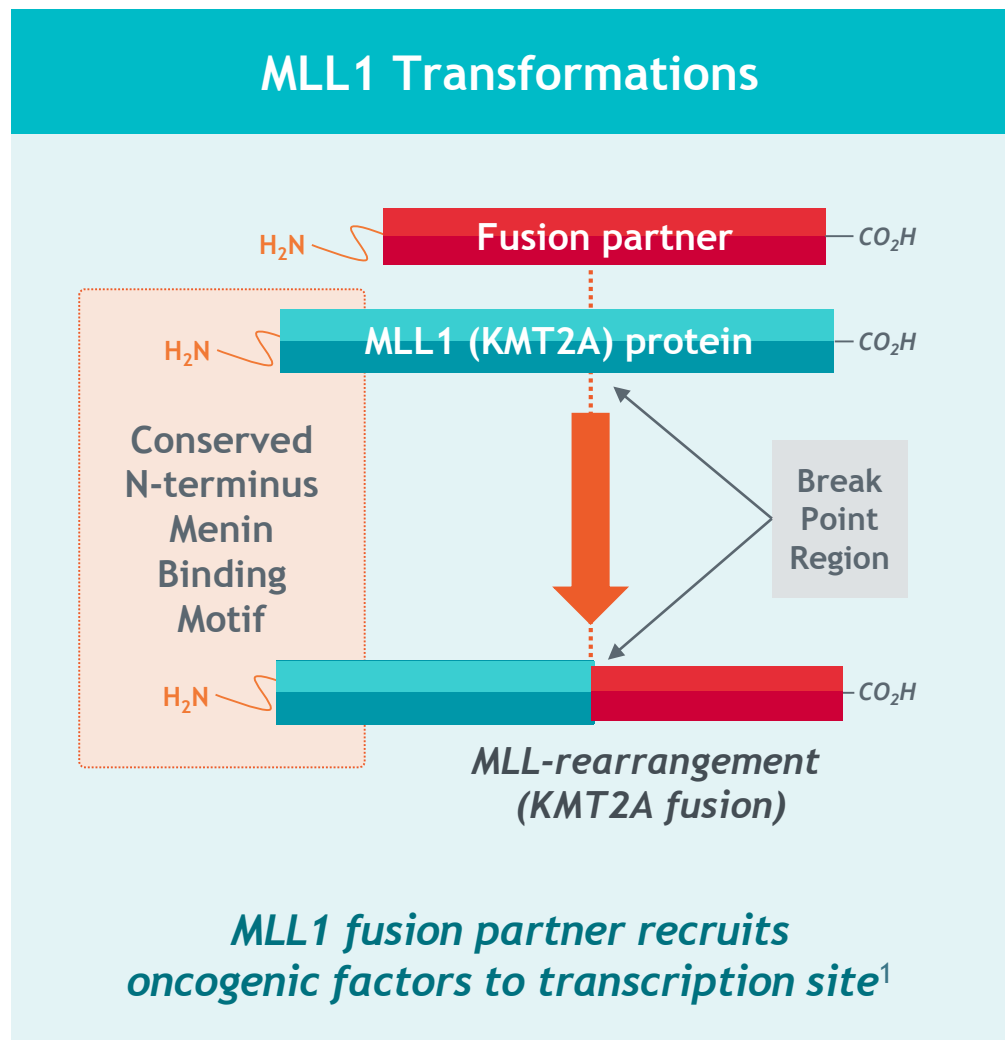
NTRK Fusions



RET Fusions

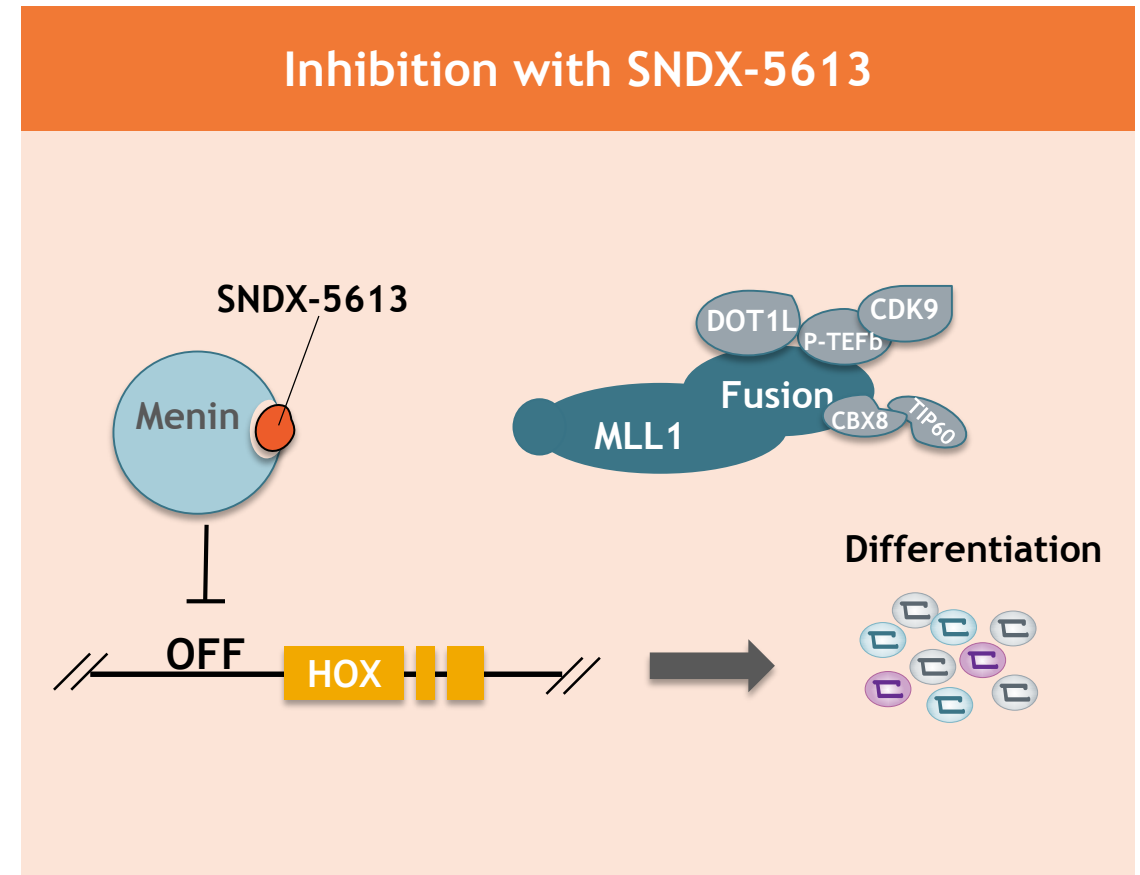
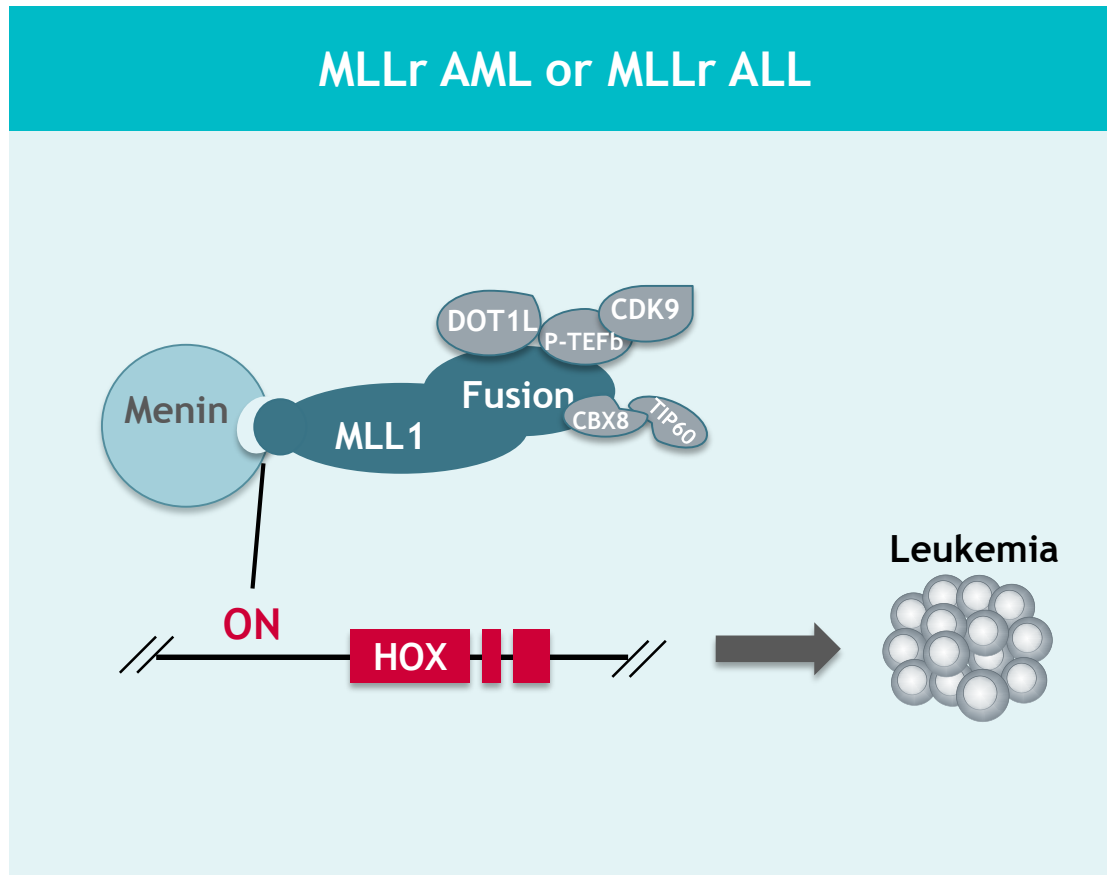


In MLLr, leukemic transformation is highly dependent on the Menin-MLL interaction



Source: Yokoyama A, Cell. 2005 Oct 21;123(2):207-18; Caslini C, Cancer Res. 2007 Aug 1;67(15):7275-83.

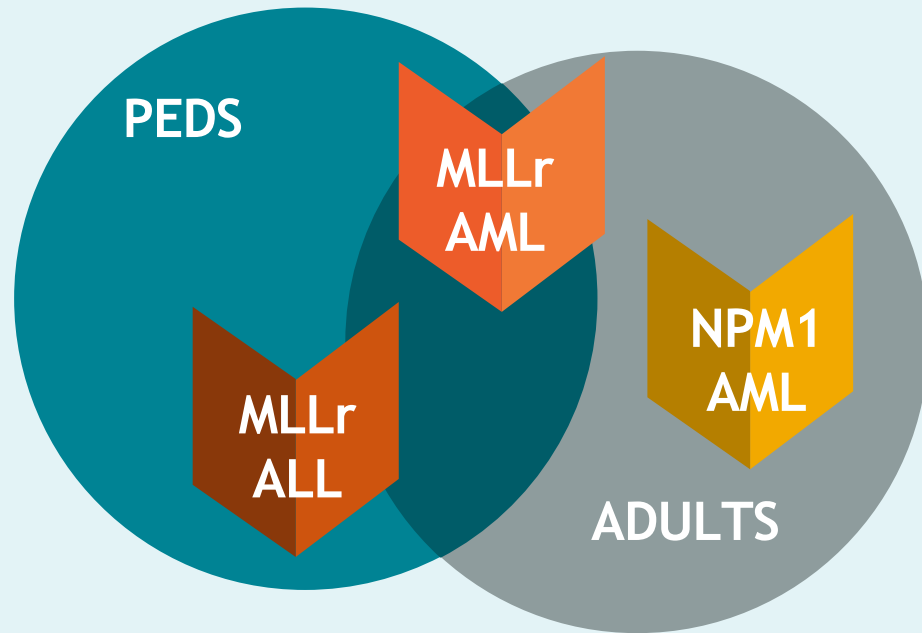
Binding of Menin to MLL1 leads to upregulation of HOX gene transcription and leukemia in MLLr AML and MLLr ALL



Adopted from: Uckelmann HJ, et al. Presented at ASH Annual Meeting, 2018.

SNDX-5613: potential best-in-class, targeted, oral agent with single agent activity and fast to market potential

Phase 1/2 trial population:
MLLr adult, MLLr peds, NPM1 mut AML



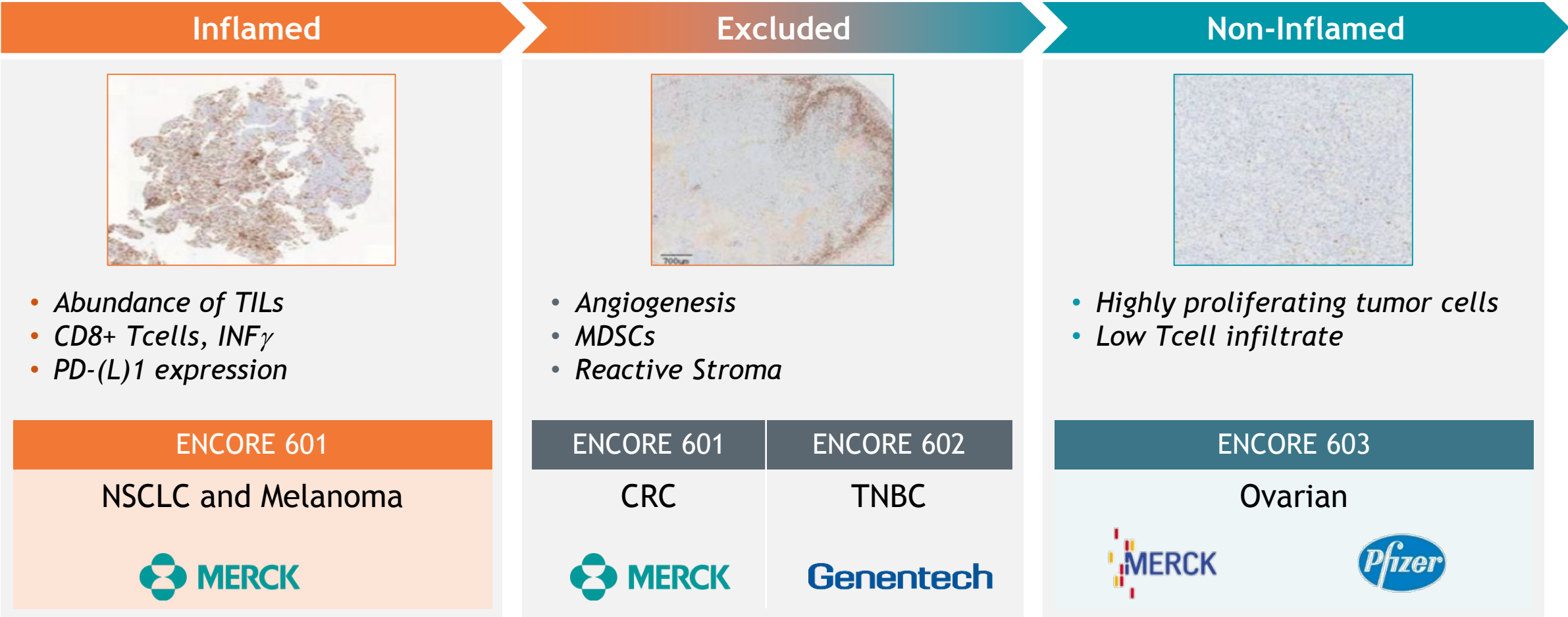
Defined fast to market pathway

- IND filing est. 2Q19; Phase 1 to follow
 - Early efficacy possible as early as year-end 2019
- MLLr and NPM1 identified today with standard screening protocols
- No approved therapies targeting MLLr or NPM1 acute leukemias
 - \$\$B commercial opportunity

ENCORE program testing combos across immune phenotypes

Responds to PD-(L)1

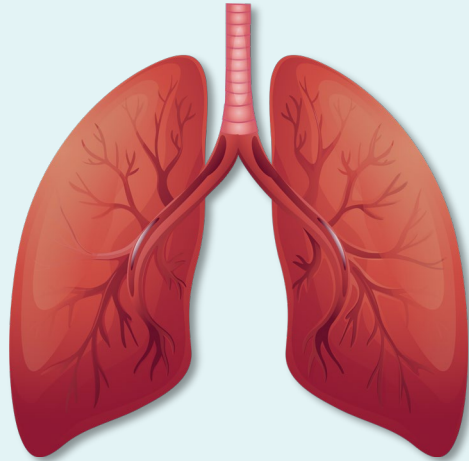
Convert to “inflamed” with combinations



Source: Hedge, et al. *Clin Cancer Res*; 22.8 (2016): 1865-1874.

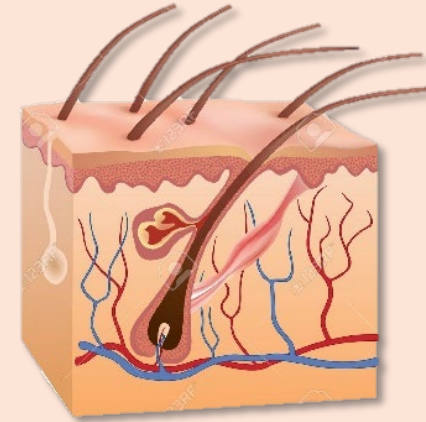
ENCORE clinical proof of concept program complete; positive signal observed in NSCLC and melanoma

NSCLC



*Strong signal in
select population*

Melanoma



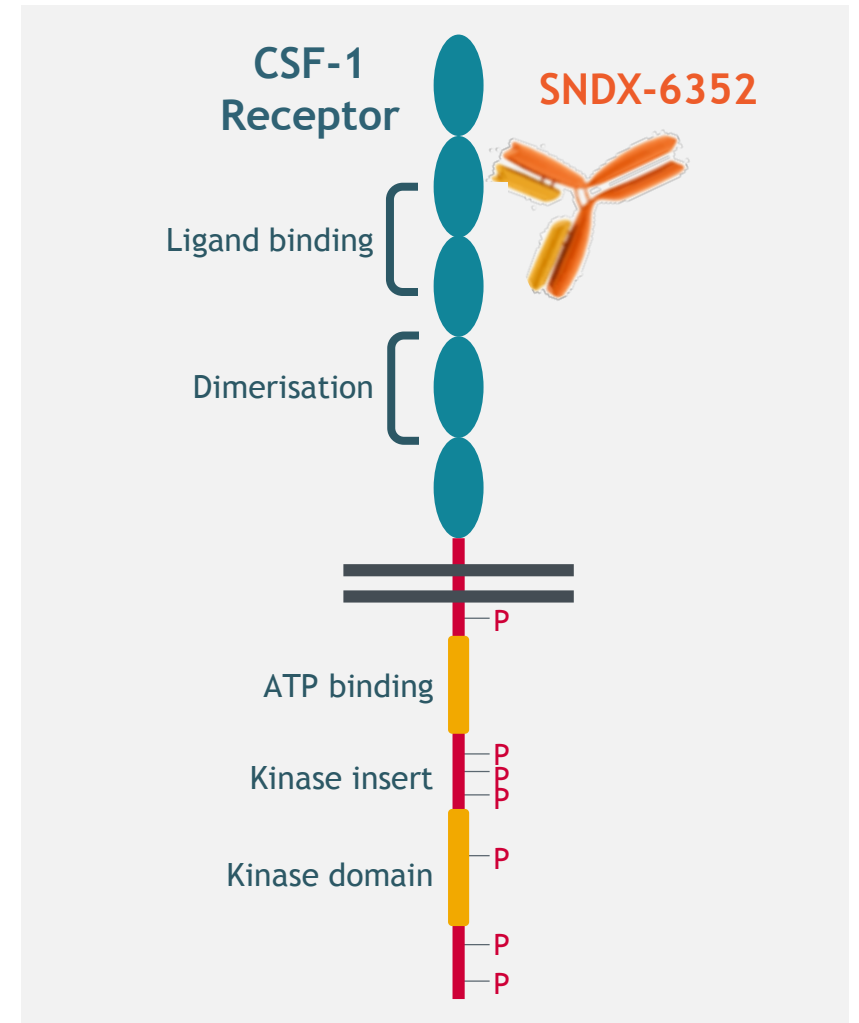
*Strong signal in
PD-1/CTLA-4 preTx pop*

Oral presentations on NSCLC biomarker and melanoma results at AACR

Update on SNDX-6352: pursuing novel indication

High affinity, IgG4 ($K_D = 4-8$ 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



CSF-1R - colony stimulating factor -1 receptor; RP2D recommended Phase 2 dose.
Source: Ordentlich, P. et al SITC 2016.

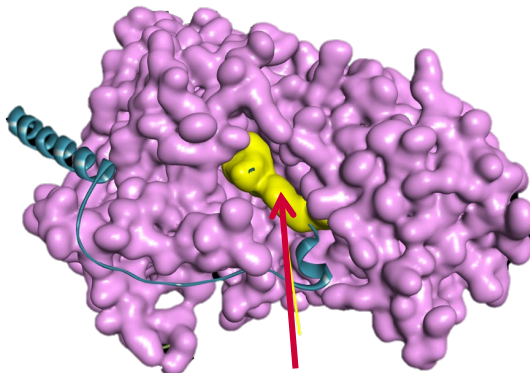
Proven ability to build the pipeline

3Q16: UCB



SNDX-6352

4Q17: Allergan/Vitae



Menin-MLLr
inhibitors

- Established relationships enhance identification and access to quality assets
- Clinical development leadership enables competitive advantage
- Business development continues to be a core strength of our business

2018 financial highlights and 1Q, full-year 2019 guidance

Ticker		SNDX (NASDAQ)	
As of December 31, 2018			
Cash and short-term investments		\$80.9 million	
Shares Outstanding*		26.8 million	
2019 1Q and full year Operating Expense Guidance			
		1Q 2019	2019
Research and Development		\$11 - 13 M	\$46 - 50 M
Total Operating Expenses^		\$15 - 17 M	\$60 - 64 M

* Includes 24.8 million common shares and pre-funded warrants to purchase 2.0 million common shares

^ Includes \$1.5 and \$6 million non-cash stock compensation expense for 1Q 2019 and for 2019, respectively

Upcoming milestones

ENTINOSTAT (Class 1 specific HDAC inhibitor)	1Q19	2Q19	3Q19	4Q19	1H20
E2112 - Interim OS analysis		●		●	●
ENCORE 601 - Present final results for melanoma cohort	●				
ENCORE 601 - Present biomarker analysis for NSCLC cohort	●				

SNDX-5613 (Menin inhibitor)	1Q19	2Q19	3Q19	4Q19	1H20
Investigational New Drug (IND) application		●			
Potential for early efficacy in relapsed refractory AML				▬	

SNDX-6352 (anti-CSF-1R mAB)	1Q19	2Q19	3Q19	4Q19	1H20
Identify recommended Phase 2 dose and schedule		●			
Preliminary efficacy in chronic GVHD			●		

Thank you. Questions?

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