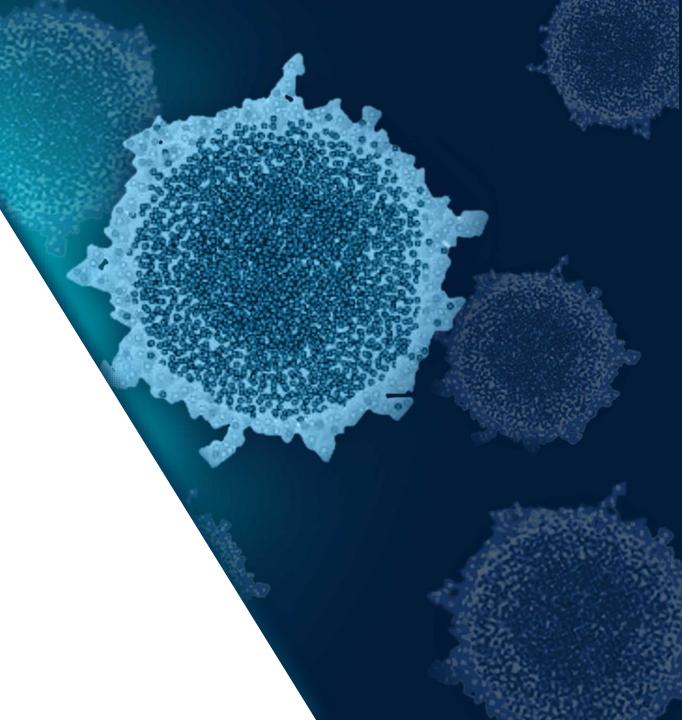


# **Corporate Presentation**

January 2023



### **Forward-Looking Statements**

Certain statements contained in this presentation regarding matters that are not historical facts, are forward-looking statements within the meaning of Section 21E of the Securities and Exchange Act of 1934, as amended, and the Private Securities Litigation Act of 1995, known as the PSLRA. These include statements regarding management's intention, plans, beliefs, expectations or forecasts for the future, and, therefore, you are cautioned not to place undue reliance on them. Forward-looking statements may include, without limitation, statements regarding: the anticipated timing of commencement, enrollment and completion of clinical trials of Aadi Bioscience, Inc. ("Aadi"); the anticipated timing for releasing data for Aadi's clinical trials; Aadi's anticipated cash runway; Aadi's potential to become a leading precision oncology company; and projected annual incidence of cancers with *TSC1* & *TSC2* alterations and in neuroendocrine tumors and endometrioid-type endometrial cancer and related market opportunities. No forward-looking statement can be guaranteed, and actual results may differ materially from those projected. Aadi uses words such as "anticipates," "believes," "plans," "expects," "projects," "intends," "may," "will," "should," "could," "estimates," "predicts," "potential," "continue," and similar expressions to identify these forward-looking statements that are intended to be covered by the safe-harbor provisions of the PSLRA.

Such forward-looking statements are based on our expectations and involve risks and uncertainties; consequently, actual results may differ materially from those expressed or implied in the statements due to a number of factors, including, but not limited to, Aadi's plans to develop and commercialize FYARRO<sup>®</sup> (*nab*-sirolimus, ABI-009); Aadi's commercialization, marketing and manufacturing capabilities and strategy; the clinical utility, potential benefits and market acceptance of FYARRO; risks related to the sufficiency Aadi's cash balance to fund operations; the timing of Aadi's clinical trials, including the timing of the availability of data from such clinical trials; uncertainties associated with the clinical development and regulatory approval of FYARRO in additional indications, including potential delays in the commencement, enrollment and completion of such clinical trials; Aadi's plans to research, develop and commercialize its current and future product candidates; Aadi's ability to identify additional products or product candidates with significant commercial potential; developments and projections relating to market size, Aadi's competitors and its industry; the impact of government laws and regulations; Aadi's ability to protect its intellectual property position; risks related to the Aadi's collaborations; and Aadi's estimates regarding future revenue, expenses, capital requirements and need for additional financing.

These risks are described in detail under the caption "Risk Factors" in Aadi's Quarterly Report on Form 10-Q for the quarter ended September 30, 2023, filed with the Securities and Exchange Commission (the "SEC") on November 8, 2023, and other documents filed from time to time with the SEC. Forward-looking statements included in this presentation are based on information available to Aadi as of the date of this presentation. Except as required by law, Aadi undertakes no obligation to revise or update any forward-looking statement, whether as a result of new information, future events or otherwise.

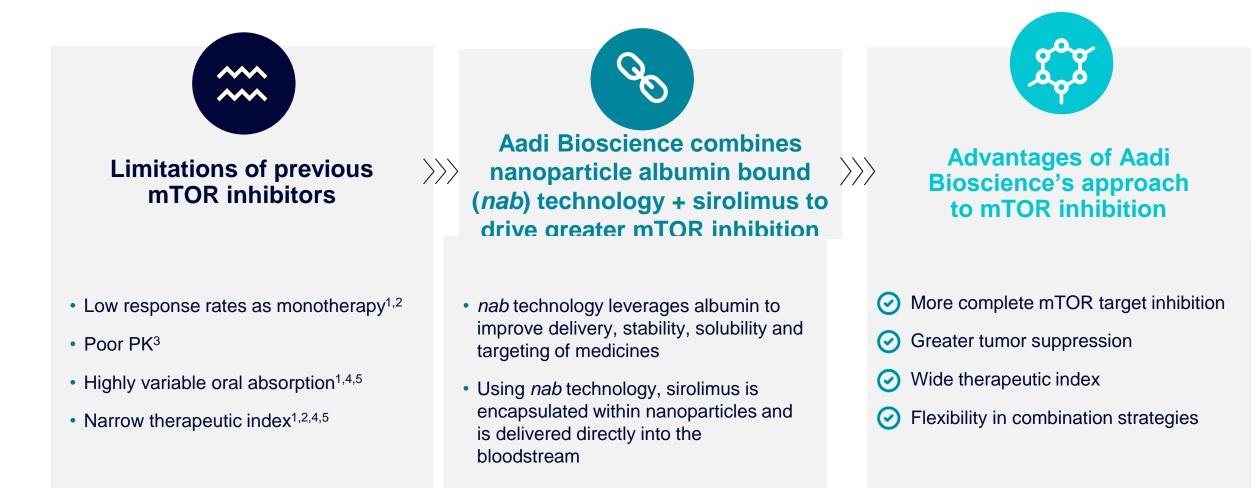


# **Our Vision**

To make bold choices in applying technology to efficiently deliver improved precision oncology therapies for people living with difficult-to-treat cancers



## Aadi Bioscience is Unlocking the Power of mTOR Inhibition





Sources: 1) AFINITOR prescribing information; 2) TORISEL prescribing information; 3) Hou et al., AACR Molecular Targets 2021 (Abstr P138); 4) ZORTRESS prescribing information; 5) RAPAMUNE prescribing information

## A Commercial-Stage Precision Oncology Company Focused on Delivering Deeper Inhibition for mTOR-Dependent Cancers



#### Successful commercial launch of FYARRO®

- Treatment for advanced malignant PEComa
- \$33M in sales achieved in first 19 months on the market\*
- Continued demand growth



### Advanced pipeline targeting multiple types of mTOR-driven tumors:

- PRECISION1 registration-directed tumor agnostic trial in solid tumors harboring TSC1 or TSC2 inactivating alterations ongoing
- Phase 2 single indication trials in endometrioid-type endometrial carcinoma and neuroendocrine tumors ongoing
- Collaboration with Mirati to evaluate combination in KRAS<sup>G12C</sup>-mutant NSCLC and other tumors ongoing



#### **Built on a strong foundation**

- Experienced management team with strong, relevant track record
- \$119.3 million in cash and cash equivalents as of September 30, 2023 with cash runway into 2025



# **Boldly Combining** *nab* **Technology** and **Therapies** to **Address Two Categories** of mTOR-dependent Tumors

### **TSC1** and **TSC2** Genetically Driven Tumors

Inactivating mutations in TSC1 and TSC2 drive mTOR pathway activation and tumor growth

### **Other mTOR-driven Tumors**

Overactivation and dysregulation of mTOR pathway is commonly found in various tumors

- *TSC1* and *TSC2* are tumor suppressor genes upstream in the mTOR pathway
- Tumors with TSC1 and TSC2 mutations occur in up to ~2% of all solid tumor cancers and across tumor types
- No approved therapies for TSC1 and TSC2 mutant patients but numerous case reports with durable responses to mTOR inhibition
- Standard CLIA-certified NGS panels already capture *TSC1* and *TSC2* mutations

- mTOR signaling pathway is overactive in many tumor types
- Known limited activity of oral mTOR inhibitors in mTOR-driven tumors like neuroendocrine tumors (NETs)<sup>1</sup>
- Combination of oral mTOR inhibitors with anti-estrogens show promise for the treatment of advanced recurrent endometrioidtype endometrial cancer (EEC)<sup>2</sup>
- Unique delivery and excellent safety profile of *nab*-sirolimus provide opportunity to treat these difficult tumors



Sources: 1) Lee, et. Al., (2018) Everolimus in the treatment of neuroendocrine tumors: efficacy, side-effects, resistance, and factors affecting its place in the treatment sequence, Expert Opinion on Pharmacotherapy, 19:8, 909-928, 2) Soliman PT et. Al., (2020) Everolimus, Letrozole, and Metformin in Women with Advanced or Recurrent Endometrioid Endometrial Cancer: A Multi-Center, Single Arm, Phase II Study. Clin Cancer Res. Feb 1;26(3):581-587

### Aadi Bioscience Advanced Oncology Development Pipeline

	Populations	Phase 1	Phase 2	Approved	Current Status		
en Tumors	Advanced malignant PEComa, AMPECT Clinical Trial	Single Agent			First FDA approved therapy for advanced malignant PEComa		
cally Driven	PRECISION1	TSC1 Arm, Single Agent			Registration directed tumor-agnostic pivotal study of <i>nab</i> -sirolimus with independent arms		
Genetically	Pan-Tumor <i>TSC1 / TSC2</i> Inactivating Alterations	TSC2 Arm, Single Agent			for <i>TSC1</i> or <i>TSC2</i> inactivating alterations; open for enrollment		
2	Advanced or recurrent endometrial cancer	<i>nab</i> -sirolimus + letrozole			Trial combining <i>nab</i> -sirolimus with letrozole for patients with endometrioid-type endometrial carcinoma; open for enrollment		
	Neuroendocrine tumors (NETs)	Single Agent			Utilizing <i>nab</i> -sirolimus as a monotherapy in neuroendocrine tumors; open for enrollment		
	Advanced solid tumors or NSCLC with <i>KRAS</i> <sup>G12C</sup> mutation	<i>nab</i> -sirolimus + adagrasib			Ongoing collaboration with MIRATI THERAPEUTICS Open for enrollment		



Evaluation of additional new single agent and combination trials ongoing



# FYARRO<sup>®</sup>: The Only Approved Treatment for Advanced Malignant PEComa



### FYARRO<sup>®</sup> First Approved Indication: Advanced Malignant PEComa





- Ultra rare sarcoma
- Estimated 100-300 new patients per year in the US<sup>1</sup>
- Biological evidence of mTOR pathway activation; cancer type with highest rate of TSC1 & TSC2 mutations
- Estimated survival of 12-16 months

- Can arise at any site but most commonly visceral (especially gastrointestinal and uterine), retroperitoneal, and abdominopelvic, with female predominance
- Mesenchymal tumor (sarcoma) consisting of perivascular epithelioid cells
  - Distinctive cells that show a focal association with blood-vessel walls
  - Usually express both melanocytic and smooth muscle markers<sup>6</sup>

Sources: 1) No formal published epidemiology information; Aadi analysis based on multiple sources including Aadi internal data and external research conducted by Tessellon Group and Corsica Life Sciences, 2) Akumalla S, et al. Oncology. 2020;98(12):905-912; 3) nab-Sirolimus AMPECT Clinical Trial mutation rates: TSC1=20%, TSC2=36%; 4) Mutation frequencies based on TCGA database "likely" and "definite" impact mutation rate and published literature rates by cancer type where available (sources available at request); 5) JS Bleeker, JF Quevedo, and AL Folpe, Sarcoma. 2012;541626; 6) Ben-Ami et al., Expert Opinion on Orphan Drugs. 2018



## **FYARRO in PEComa: Continuing Product Demand**

### **Fyarro**<sup>®</sup> sirolimus protein-bound particles

sirolimus protein-bound particles for injectable suspension (albumin-bound)

**\$6 million net sales in 3Q 2023** *40% growth Y/Y* 

**\$33 million sales to date** Achieved in first 19 months on the market\*

Sustained demand growth represents increased depth and breadth of prescribing across academic and community settings

# 

#### PREFERRED

NCCN clinical practice guidelines in oncology listed FYARRO as the only "preferred" treatment for malignant PEComa

#### ACCESSIBLE



Utilize AadiAssist, a comprehensive patient support program, to ensure access to FYARRO; national and regional payers continue to adopt coverage policies



#### ENGAGED

Experienced commercial team focused on establishing FYARRO as SOC in malignant PEComa

# > 180

Unique accounts ordering FYARRO

# + 90%

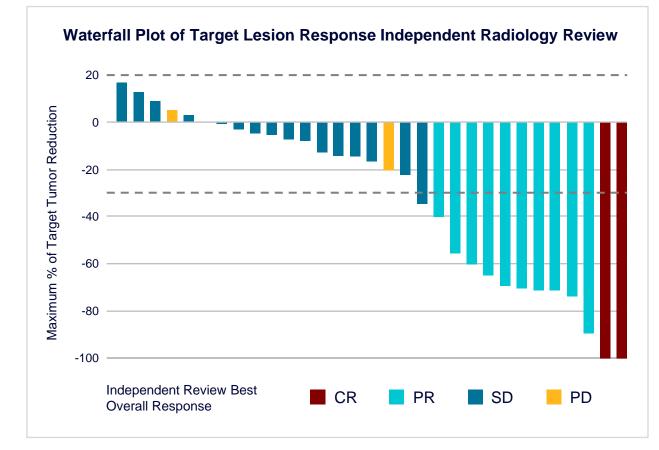
Account reorder rate

~ 50% Community adoption



## **AMPECT PEComa Registrational Trial Met Endpoints**

Highly durable responses coupled with high disease control rate and manageable toxicities showed *nab*-sirolimus effectiveness, representing an important new treatment option for patients in need



Efficacy Results in AMPECT <sup>1,2</sup>	Independent Radiology Review	
Overall Response Rate (95% CI)	<b>39%</b> (22%, 58%)	
Complete Response	7% (2/31)	
Partial Response	32% (10/31)	
Stable Disease	52%	
Progressive Disease	10%	
Disease Control Rate <sup>‡</sup>	71%	
Median Duration of Response	39.7 months	
Median Progression Free Survival	10.6 months (5.5-NR)	
Median Overall Survival <sup>†</sup>	53.1 months	

#### Safety Summary<sup>3</sup>

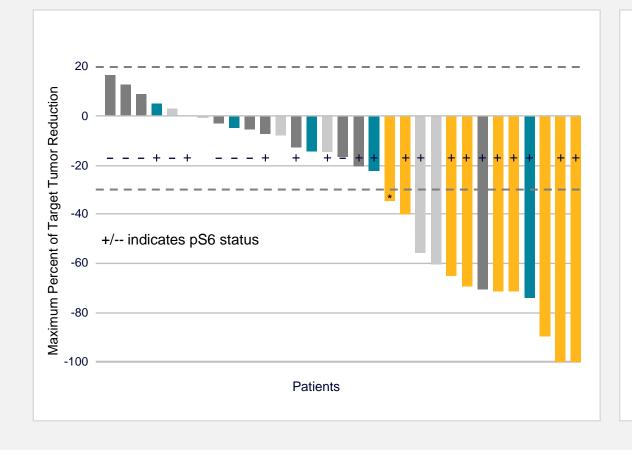
- Most treatment-related adverse events (TRAEs) grade 1 or 2 (no grade 4 or 5)
- Most common nonhematologic TRAEs: mucositis (79%), fatigue (59%), rash (56%)
- Most common hematologic TRAEs: anemia (47%) and thrombocytopenia (32%)
- Two patients discontinued due to a TRAE (grade 2 anemia and grade 1 cystitis)
- Dose reductions occurred in 13/34 (38%) of patients



Note: ‡Disease control rate defined as complete response + partial response + stable disease ≥12 weeks; Sources: 1) FYARRO® Prescribing Information; 2) AJ Wagner, CTOS 2022; 3) AJ Wagner, JCO 2021

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## Data from AMPECT in TSC1 or TSC2 Inactivating Alterations Supports Further Investigation Across Different Tumor Types



Best Overall Responses Patients with NGS* (N=25)	<u>TSC1/TSC2</u> n = 14	Non <i>TSC1/TSC2</i> n = 11
	11 = 14	
Complete or Partial Response	9/14 (64%)	1/11 (9%)
Stable Disease	4/14 (29%)	8/11 (73%)
Stable Disease ≥12 weeks	3/14 (21%)	5/11 (45%)
Progressive Disease	1/14 ( 7%)	2/11 (18%)

• 25 patients had available NGS reports

• Confirmed Responders: 9/14 (64%) pts with *TSC1/TSC2* vs 1/11 (9%) with no *TSC1/TSC2* alterations

TSC1/TSC2: 12/14 (86%) patients had Disease Control (CR or PR or SD ≥12 weeks)

**TSC2** mutation

TSC1 mutation

No TSC1 or TSC2 mutation

UNK mutational status

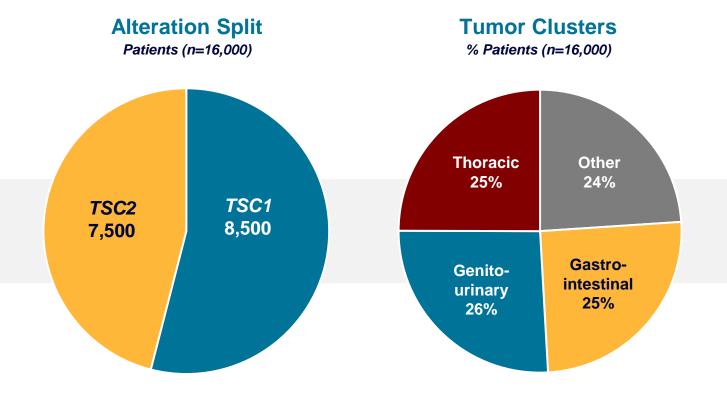


## **TSC1** and **TSC2** Inactivating Alterations Represent Significant Opportunity Across Common Cancer Types

Real-World Analysis of *TSC1* and *TSC2* Patient Population<sup>1</sup>

- Next generation sequencing (NGS) of nearly 440,000 cancer patients from the Foundation Medicine database
- 2% of patients have known or likely inactivating alterations in *TSC1* or *TSC2*
- Based on extrapolation from SEER database, ~16,000 new cancer cases each year would have actionable TSC1 or TSC2 alterations

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Approximately 16,000 patients with *TSC1* or *TSC2* inactivating alterations across varying tumor types represent a potential multi billion-dollar total addressable market in <u>each</u> alteration



Note: Methodology to determine TAM consists of applying FMI RW data (TSC1/2 mutation frequency) presented at AACR-NCI-EORTC and incident cancer volume for solid tumors in the SEER database (2023)



# PRECISION1: Registration Directed Tumor-Agnostic Trial of *nab*-sirolimus in *TSC1* or *TSC2* Inactivating Alterations



### **PRECISION1** Trial

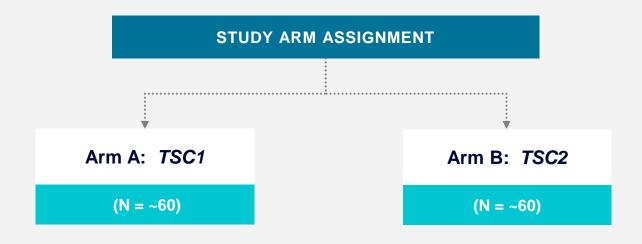
- Two independently evaluable arms, one each for *TSC1* and *TSC2*
- Primary endpoint: ORR by blinded, independent radiologic review
- Patient accrual based on local NGS results
- First patient dosed March 2022 with expected 24-month enrollment period
- 150 active clinical trial sites





## **Key Eligibility Criteria**

- Metastatic or locally advanced disease ineligible for surgery
- Naïve to mTOR inhibitor treatment
- Pathogenic TSC1 or TSC2 inactivating alterations identified through NGS
- Must have received standard therapy for the disease or in investigator opinion unlikely to benefit





# **Durable Responses Observed in Heavily Pre-Treated Patients With a Median of Three Lines of Prior Therapies**



Interim results from investigator-assessed responses in first 40 patients from TSC1 and TSC2 arms reported in December 2023

Efficacy Summary		
	<b>TSC1</b> Efficacy Evaluable <sup>1</sup> (n=19) <sup>2</sup>	TSC2 Efficacy Evaluable <sup>1</sup> (n=18)
Median prior lines of therapy	3	3.5
Partial Response (n, %) <sup>3, 4</sup>	5 (26)	2 (11)
Stable Disease (n, %) • SD • SD $\geq$ 6 mos	9 (47) 3 (16)	12 (67) 3 (17)
Progressive Disease (n, %)	5 (26)	4 (22)
Clinical Benefit Rate (n, %) (PR+SD ≥ 6 mos)	8 (42)	5 (28)
Time to response (months)	1.4	3.6

### **Safety Summary**

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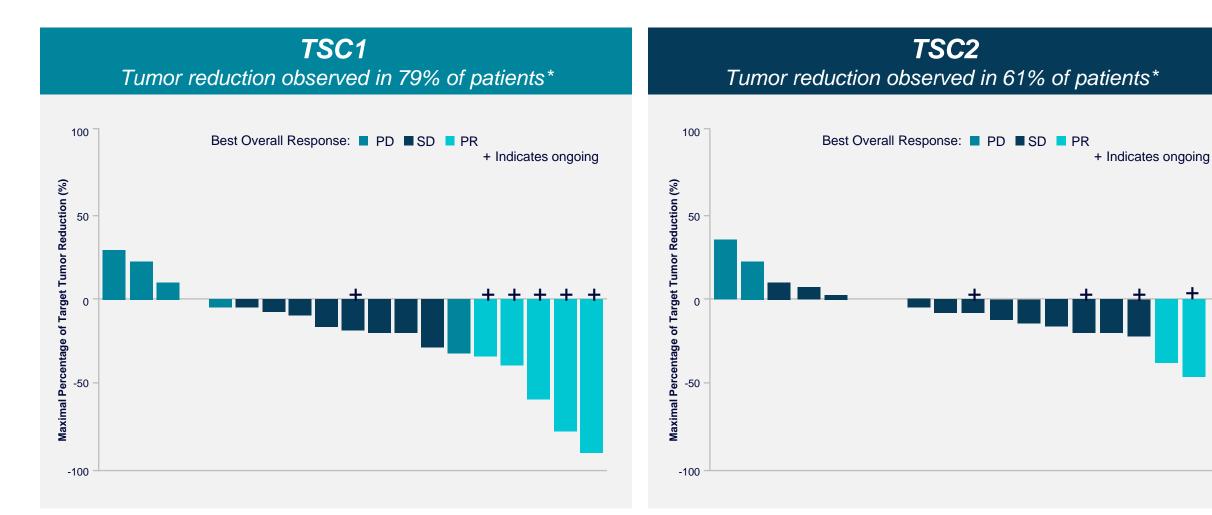
- No new safety signals
- Pattern of AEs consistent with nab-sirolimus label and mTORi class
- No grade 4 TRAEs or deaths due to study drug
- 1 patient discontinued study due to grade 2 recurrent pneumonitis

<sup>1</sup> By Investigator Assessment, <sup>2</sup> Three patients without post-baseline assessment not included, <sup>3</sup> One unconfirmed PR, patient on treatment and awaiting confirmatory scan [at the time of the data release], <sup>4</sup> One unconfirmed PR with a single PR assessment



# Majority of Patients Showed Tumor Reduction Including Deep Responses in *TSC1*-Altered Tumors







# **PRECISION1 Interim Analysis Summary**



- TSC1 arm results encouraging
  - Response rate in range of our expectations
  - Responses appear to be deep and durable in a heavily pre-treated population
  - Responses in different tumor types supportive of a tumor agnostic indication
- TSC2 arm ORR interpretation is complicated by small sample size and heavy pre-treatment
  - 50% patients received 5 or more prior therapies
- Two-third interim enrollment of 80 patients completed

Anticipated completion of PRECISION1 trial by end of 2024; results by early 2025



### Advancing Our Pipeline to Deliver New Breakthroughs in Endometrial Cancer and Neuroendocrine Tumors

### Two Phase 2 single indication trials launched in fall of 2023

Establishing a new
preferred combination
for endometrial cancer

### **Therapeutic Potential of mTOR Inhibitors in Endometrial Cancer**

- Known activity in rapalogs combined with anti-estrogens for the treatment of advanced recurrent endometrioid-type endometrial cancer (EEC)
- Unique pharmacology when combined with the standard anti-estrogen letrozole
- Recent changes in recommended first line standard of care (chemo + immunotherapy) creates potential opportunity for use in second line treatment
- Estimated addressable population\*: 10,000 EEC/year, ~7,000 2L/year

Developing *nab*sirolimus as a best-inclass mTOR inhibitor for neuroendocrine tumors

#### **Role of mTOR Inhibitors in Neuroendocrine Tumors (NETs)**

- Historically low response rate to treatment with oral rapalogs and other agents
  which nonetheless are used clinically and recommended in treatment guidelines
- In preclinical animal models, *nab*-sirolimus demonstrated improved target suppression relative to other mTORs, warranting further exploration of *nab*-sirolimus
- Estimated addressable population\*: ~3,500 patients per year



## On The Path To Becoming A Leading Precision Oncology Company

Tumor Agnostic TSC1 & TSC2 Alterations ~16,000 US Patients/yr

### Initial Interim Analysis

PRECISION1 Key Milestones Full Enrollment and Two-Thirds Interim Analysis

### PRECISION1 Trial Completion

PRECISION

#### PRECISION1 Trial Readout

#### 2023:

- \$33M in FYARRO sales since launch\*
- PRECISION1 tumor agnostic *TSC1* and TSC 2 one-third interim analysis on 40 patients
- PRECISION1 enrollment for 2/3 interim completed
- Launched additional *nab*-sirolimus trials in expanded indications

2023

#### 2024:

- PRECISION1 trial full enrollment expected in spring of 2024
- PRECISION1 2/3 interim analysis expected in 3Q 2024
- Anticipated completion of PRECISION1 trial by YE
- Ongoing commercialization of FYARRO

2024

#### 2025:

• Results of PRECISION1 trial of 120 patients expected in early 2025





\* Commercial launch on Feb 22, 2022. 19-months of sales as of close of 3Q 2023.



# Aadi Bioscience, Inc.

Pacific Palisades, CA NASDAQ: AADI www.aadibio.com



# Appendix

## Accomplished Management Team with Strong Track Record

Extensive pharma experience in building blockbuster oncology brands



**Dave Lennon** President and CEO



Scott Giacobello, CPA **Chief Financial Officer** 

Large- and small-cap biotech knowhow in effectively managing explosive growth



Loretta Itri, MD **Chief Medical Officer** 



Stephen Rodin, JD SVP & General Counsel

Strong networks enable rapid organizational scaling with top talent

Understand how to create value by building sustainable companies



**Bryan Ball** Chief Quality Officer & **SVP** Manufacturing Operations

& Corporate

Communications

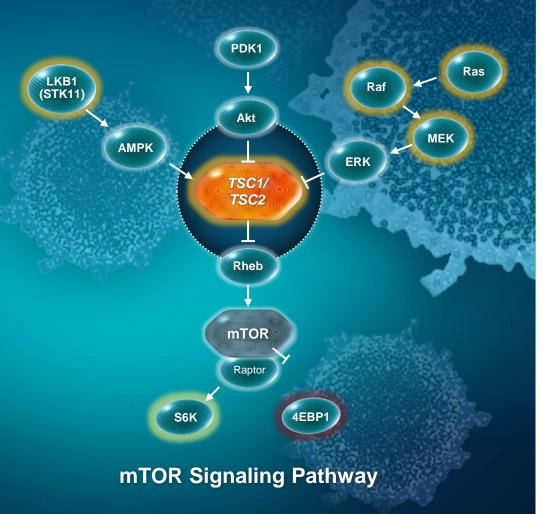


**Raymond Steitz** SVP & Chief Human **Resources Officer** 





### TSC1 and TSC2 Alterations: Key Oncogenic Drivers in the mTOR Pathway



# Inactivating mutations in *TSC1* and *TSC2* drive mTOR pathway activation and tumor growth

*TSC1* and *TSC2* are upstream regulators of mTOR activity within the PI3K/Akt/mTOR pathway

*TSC1* and *TSC2* mutations occur at a rate of approximately 1-2% each across cancers

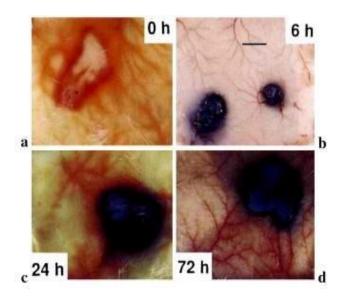
No approved therapies for *TSC1* and *TSC2* mutant patients but numerous case reports with durable responses to mTOR inhibition

Standard CLIA-certified NGS panels already capture *TSC1* and *TSC2* mutations



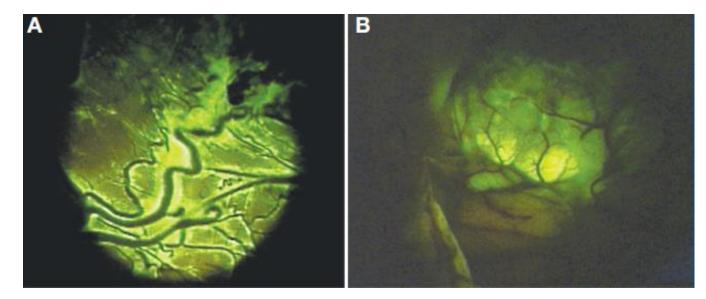
## **Role of Albumin in Tumor Targeting**

Albumin accumulation in tumors established in multiple preclinical models<sup>1</sup>



Accumulation of the evans blue albumin complex in subcutaneously growing sarcoma 180 tumors over 72 h

Labeled albumin can be used intraoperatively to guide surgical resection of tumors in humans<sup>2</sup>



- 5-Amino Fluorescein labelled albumin administered IV (0.5-1 mg/kg) in 13 patients, 0.5-4 days before surgery
- Tumor fluorescence was bright in 11 patients (84%), resulting in complete resection in 9 patients (69%)

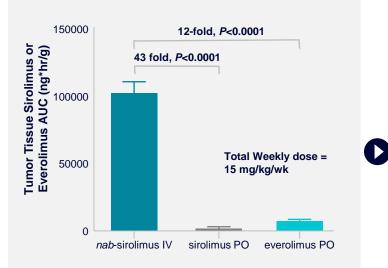
High accumulation of albumin in tumors potentially driven by tumor vessel leakiness (EPR effect); increased caveolar transport; increased albumin catabolism

bioscience

Note: EPR- Enhanced permeability and retention effect; Sources: 1) Y Shahzad et al., Curr Cancer Drug Targets. 2014;14(8):752-63; 2) P Kremer et al., Neurosurgery. 2009;64(3 Suppl):ons53-60; discussion ons60-1

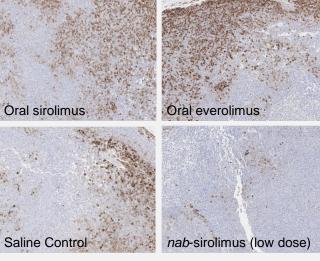
### Higher *nab*-sirolimus Intratumoral Concentrations Drive Increased Target Suppression and Tumor Growth Inhibition in a Bladder Cancer Xenograft

### Significantly Higher Intratumoral Drug Accumulation



Tumor concentration of *nab*-sirolimus, oral sirolimus, and oral everolimus measured over 7 days at equal weekly dose (15 mg/kg/wk) in mice bearing tumor xenografts

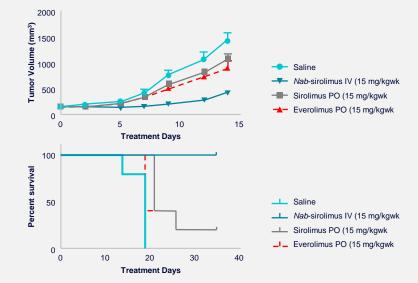
Increased mTOR Target Suppression (pS6)



Tumor IHC pS6 suppression on D7 post dose at equal doses (15 mg/kg/wk). pS6 is a downstream target of mTOR. *nab*-sirolimus vs oral sirolimus: P = 0.0001 (ANOVA)

nab-sirolimus vs oral everolimus P = 0.0034 (ANOVA)

Stronger Inhibition of Tumor Growth and Longer Survival in Animals



UMUC3 (aggressive human bladder cancer) Xenograft (n=8/group): Oral Rapamycin and Everolimus 15 mg/kg/wk (3 mg/kg, 5x/wk); IV *nab*-sirolimus 15 mg/kg/wk (7.5 mg/kg, 2x/wk) Tumor volume: *nab*-sirolimus vs oral sirolimus: P < 0.0001 (ANOVA) *nab*-sirolimus vs oral everolimus P = 0.0023 (ANOVA) Survival: *nab*-sirolimus vs oral sirolimus: P < 0.05 (Log-rank test) *nab*-sirolimus vs oral everolimus P < 0.05 (Log-rank test)

*nab*-sirolimus demonstrated enhanced anti-tumor activity vs. currently approved mTOR inhibitors in animal models at clinically relevant doses

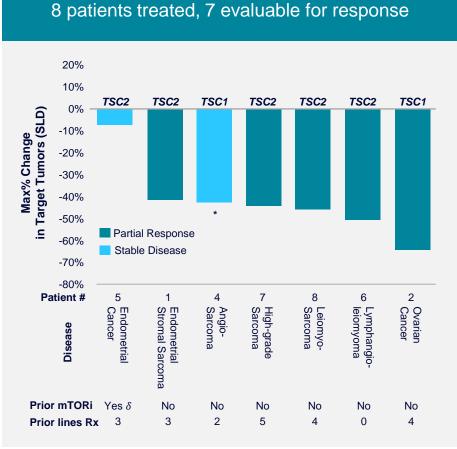


## **Expanding Beyond PEComa**

### Early Experience in Other Tumor Types with TSC1 or TSC2 Inactivating Alterations

Multi-institutional Expanded Access for an Intermediate-size Population

- =8 patients with *TSC1* or *TSC2* inactivating alterations
  - 6 mTOR-naïve
  - 2 previously treated with an mTORi
- 100 mg/m<sup>2</sup> ABI-009 (*nab*-sirolimus) given D1, D8 of a 21-day cycle
- Response Analysis: RECIST v1.1
- Tumor types: Ovarian cancer, endometrial cancer, angiosarcoma, leiomyosarcoma, lymphangio-leiomyoma, high grade sarcoma, endometrial sarcoma
- Lines of prior therapy: median 3.5 (range 0-6)



Efficacy

Safety

- Treatment-emergent AEs (≥30%) included edema, infections, mucositis, and pain (71% each), |nail changes and vomiting (57% each), and hypertension and nausea (43% each).
- Majority of events were G1/G2
- Treatment-related SAEs were reported in 2 patients and included hyperglycemia and infection (Pt#4) and acute kidney injury (Pt#7) possibly secondary to administration of contrast
- Dose reductions occurred in 3/8 patients (38%) from 100 mg/m2 to 75 mg/m<sup>2</sup>

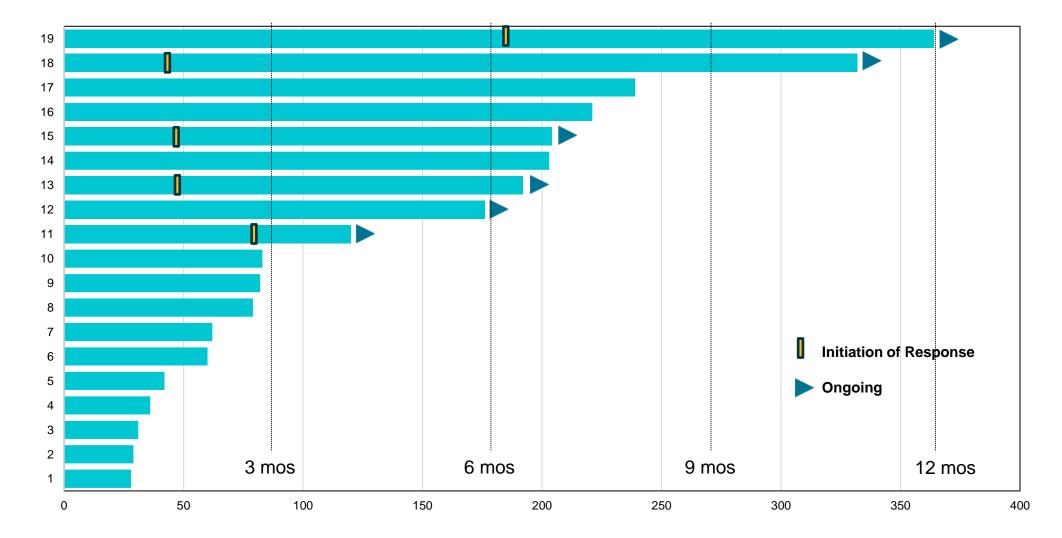


ABI-009 is an investigational new drug and has not been approved for commercial distribution in the United States. Source: MA Dickson. ASCO. 2021. Abstract # 3111

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### **TSC1** Inactivating Alterations: Patient Time on Treatment







### **TSC2** Inactivating Alterations: Patient Time on Treatment



