

### **Corporate Presentation**

March 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. No forward-looking statement can be guaranteed, and actual results may differ materially from those projected. Aadi Bioscience, Inc. ("Aadi") undertakes no obligation to publicly update any forward-looking statement, whether as a result of new information, future events or otherwise, except to the extent required by law. We use words such as "anticipates," "believes," "plans," "expects," "intends," "may," "will," "should," "could," "estimates," "predicts," "potential," "continue," "guidance," 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 its product candidates, including 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; the timing of the availability of data from Aadi's clinical trials; Aadi's plans to research, develop and commercialize its current and future product candidates; Aadi's ability to successfully enter into collaborations, and to fulfill its obligations under any such collaboration agreements; Aadi's ability to identify additional products or product candidates with significant commercial potential; developments and projections relating to Aadi's competitors and our its industry; the impact of government laws and regulations; Aadi's ability to protect its intellectual property position; the impact of the COVID-19 outbreak on Aadi's operations, the biotechnology industry and the economy generally 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 March 31, 2022, filed with the Securities and Exchange Commission (the "SEC") on May 12, 2022, 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.



Aadi Bioscience is a Commercial-Stage Precision Oncology Company Re-engineering mTOR Inhibition



- Commercializing FYARRO<sup>®</sup> for treatment of Advanced Malignant PEComa
- Technology based on nanoparticle albumin-based (nab) platform proven with ABRAXANE<sup>®</sup>
- Focus on cancers that are highly mTOR dependent
- PRECISION 1 registrational trial in tumor-agnostic TSC1 or TSC2 inactivating alterations in solid tumors now actively enrolling
- \$72.5 million financing in Sept 2022 extends cash runway into 2025, supporting expanded development of FYARRO into potential new indications, and the continued progression of PRECISION 1 registrational trial and FYARRO commercialization



## Great Science. Great Story. Great People.

Extensive pharma experience on building blockbuster oncology brands

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

Strong networks across key functions enable rapid organizational scaling with top talent

Understand the requirements of creating value by building sustainable companies from the bottom up



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



- Ultra rare sarcoma
- Estimated 100-300 new patients per year in the US<sup>6</sup>
- Biological evidence of mTOR pathway activation; cancer type with highest rate of TSC1 & TSC2 mutations<sup>2-4</sup>
- Estimated survival of 12-16 months<sup>5</sup>

- Can arise at any site but most commonly at visceral (especially gastrointestinal and uterine), retroperitoneal, and abdominopelvic sites and with female predominance
- Mesenchymal tumor (sarcoma) consisting of perivascular epithelioid cells
  - Distinctive cells that show a focal association with blood-vessel walls<sup>1</sup>
  - Usually express both melanocytic and smooth muscle markers<sup>1</sup>

Sources: 1) Ben-Ami et al., Expert Opinion on Orphan Drugs. 2018; 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) 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 7) Primary Oncologist Market Research (N=10) conducted July and August 2019 by Corsica Life Sciences



## **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



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

## FYARRO<sup>®</sup> Inhibits Key Signaling Pathways in Cancer



#### Improvements over other Approved mTOR Inhibitors

- High drug levels in tumor result in more complete mTOR target inhibition and greater tumor suppression not achieved with other mTORi's <sup>1</sup>
- Improved PK, half-life and exposure without compromising safety – wide therapeutic index
- Flexibility in combination strategies
- Overcomes limitations of other mTORi's such as highly variable oral absorption, poor PK, narrow therapeutic index
- ✓ Unlocks full potential of mTOR inhibition



#### 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)



# Saline Control nab-sirolimus (low dose)

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



## **AMPECT PEComa Registrational Trial Basis of Accelerated Approval**

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<sup>®</sup> Prescribing Information; 2) AJ Wagner, CTOS 2022; 3) AJ Wagner, JCO 2021

## PEComa Commercial Launch (Feb 22, 2022)



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

#### 3Q 2022, \$4.2M net sales Sales to date \$10.0 M net sales

Steady product demand growth and new patient starts plus bolus of patients carried over into second quarter



#### PREFERRED

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



#### ACCESSIBLE

Launched Aadi Assist, a comprehensive patient support program, to ensure access to FYARRO; National and Regional payers continue to adopt coverage policies



#### ENGAGED

Experienced commercial team is in place with Launch execution focused on establishing FYARRO as SOC in malignant PEComa

## > 90

Accounts ordering FYARRO<sup>®</sup> with rapid site activation

## +80%

Account reorder rate

> 60% Community adoption



## FYARRO<sup>®</sup> Advanced Oncology Development Pipeline

Populations	Phase 1	Phase 2	Registrational	Approved	Current Status
Advanced Malignant PEComa, AMPECT Clinical Trial	Single Agent				First FDA approved therapy for advanced malignant PEComa
PRECISION PAn-Tumor <i>TSC1 / TSC2</i> Inactivating Alterations	<i>TSC1</i> Arm, Sing <i>TSC2</i> Arm, Sing	gle Agent gle Agent			Tumor-agnostic pivotal study with independent arms for <i>TSC1</i> or <i>TSC2</i> inactivating alterations; open for enrollment
Advanced solid tumors or NSCLC with <i>KRAS</i> <sup>G12C</sup> mutation (Phase 1/2)	Nab-sirolimus + adagrasib	)			Collaboration ongoing with MIRATI THERAPEUTICS

Evaluation of additional new single agent and combination trials ongoing

Ongoing Ong



## **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



## **TSC1** and **TSC2** Inactivating Alterations Across All Cancers Represent Significant Opportunities

Projected Annual Incidence of Cancers with *TSC1* and *TSC2* Alterations<sup>1</sup> Estimated US Patients Available for 1<sup>st</sup> Line Therapy in 2030

			EGFR26,115
	Top 5 Histold	gies by # of <i>TSC1</i> Patients	KRAS         19,911           ERBB2         17,700
TSC1 Alterations (Likely or Definite Impact)	Tumor Types	# with Definite Impact Mutations (%)	$\begin{array}{c} PIK3CA \\ BRAF \\ TSC1 + TSC2 \\ \hline \end{array} \begin{array}{c} 10,422 \\ 13,854 \\ \hline 11,654 \\ \hline \end{array}$
	Bladder	1,772 (6.33%)	FGFR3 - 8,998 TSC1 - 7.388
	NSCLC	1,297 (0.77%)	S MET
TSC1 Alterations	Endometrial	835 (2.10%)	= $TSC2$ $=$ $-4,631$
(Definite Impact)	Hepatobiliary	445 (1.27%)	a $ATM = -3,008ALK = -2,796$
	Pancreatic	344 (0.57%)	$\begin{array}{c} c \\ c$
	Top 5 Histold	gies by # of TSC2 Patients	$\begin{array}{ccc} & & & & & \\ \hline & & & & \\ \hline & & & & \\ \hline & & & &$
<b>TSC2 Alterations</b> (Likely or Definite Impact)	Tumor Types	# with Definite Impact Mutations (%)	$ROS1 \rightarrow 1,696$ $IDH1 \rightarrow 1,134$ $NTRK1 \rightarrow 566$
	NSCLC	1,945 (1.16%)	NTRK3 + 421 $KIT + 320$
	Hepatobiliary	1,157 (3.31%)	BRIP1 - 302
TSC2 Alterations	Cervix	808 (0.71%)	
(Definite Impact)	Endometrial	487 (1.22%)	BARD1 -1161 SMARCB1 -18
	Bladder	477 (1.70%)	
2,500 5,000 7,500 10,000	12,500 15,000 17,500 20	,000	Patients with drug-targetable mutations in 2030

#### Definitions:

Likely Impact Alterations (harmful missense variants): missense mutations predicted to be deleterious by SIFT or possibly or probably damaging by PolyPhen Definite Impact Alterations (truncating and deep deletions): out-of-frame frameshift insertions/deletions, nonsense mutations, splice-site mutations, and deep deletions (e.g., copy number "-2" in cBioPortal)



Incidence of TSC1 and TSC2 Alterations

vs. 26 Other Actionable Genes<sup>2</sup>

13 1) analysis of TCGA, cBioPortal, and SEER databases conducted by Tessellon Group in June 2021 2) G. Gulati, et al. AACR Annual Meeting 2022. Poster #5799

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



Best Overall Responses	TSC1/TSC2	Non TSC1/TSC2
Patients with NGS* (N=25)	n = 14	n = 11
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)

UNK mutational status



## Expanded Access Program: Efficacy of *nab*-sirolimus in Malignant PEComa Patients after progression/failure of other mTOR inhibitors



	All Patients
Best Overall Responses	n = 16
Partial Response	4/16 (25%)
Stable Disease	8/16 (50%)
Stable Disease ≥12 weeks	6/16 (38%)
Progressive Disease	4/16 (25%)

• 10/16 (63%) patients had Disease Control (CR or PR or SD ≥3 months)

• 4 nab-sirolimus responders:

- BOR on prior mTORi: 1/4 SD, 2/4 PD, 1/4 NE due to toxicity

- 2/4 had 3 prior lines of Rx

Best Overall Responses Patients with NGS* (N=13)	<u>TSC1/TSC2</u> n = 9	Non TSC1/TSC2 n = 4
Partial Response	4/9 (44%)	0
Stable Disease	3/9 (33%)	3/4 (75%)
Stable Disease ≥12 weeks	2/9 (22%)	3/4 (75%)
Progressive Disease	2/9 (22%)	1/4 (25%)

• 13 patients had available NGS reports

• Responders: 4/9 (44%) pts with TSC1/TSC2 vs 0/4 with no TSC1/TSC2 alterations

TSC1/TSC2: 6/9 (66%) patients had Disease Control (CR or PR or SD ≥3 months)



Note: NE tox- Pt came off for toxicity prior to any evaluation; NE- Not Evaluable; '+' indicates ongoing patient; \* 3 patients had Unknown mutational status; NGS reports included MSK-IMPACT, MDA Molecular Diagnostic Lab, Foundation One, Oncopanel; Source: MA Dickinson, CTOS 2021

## **Expanding Beyond PEComa**

Early Experience Other Tumor Types with TSC1 or TSC2 Inactivating Alterations

Multi-institutional Expanded Access for an Intermediate-size Population

- N=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

#### 8 patients treated, 7 evaluable for response



#### 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/m<sup>2</sup> to 75 mg/m<sup>2</sup>



## Prior Experience in Patients with *TSC1* and *TSC2* Alterations Support Rationale for Tumor-Agnostic Approach

#### **AMPECT PEComa Registrational Trial**

PEComa Patients with TSC1/TSC2 Alterations<sup>1</sup>

- All mTOR naïve
- 14 patients
- Response in 9/14 (64%)

#### FYARRO Expanded Access Program

PEComa Patients with *TSC1/TSC2* Alterations Previously Treated with mTOR inhibitors<sup>2</sup>

- Progressed on prior mTOR
- 9 patients
- Response in 4/9 (44%)

#### **FYARRO Expanded Access Program**

Non-PEComa Patients with *TSC1/TSC2* Alterations

- 6 mTOR naïve + 2 prior mTOR treated
- 8 patients
- Response in 5/8 (63%)

#### Blended data in TSC1 and TSC2 alterations: 18/31 (58%)



## PRECISION 1: *nab*-sirolimus Basket Study for *TSC1* or *TSC2* Inactivating Alterations Tumor-Agnostic Registrational Trial



- Independently evaluable arms for TSC1 and TSC2
- Primary endpoint : ORR
- Secondary endpoints : DOR, DCR
- Patient accrual based on local NGS results
- First patient dosed (March 2022)
- 24 month enrollment
- Preliminary clinical data expected 1H 2023

#### Strategies to expedite enrollment:

- Partnered with NGS providers
- Partnered with US Oncology





#### 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





## Aadi on Path to Becoming a Leading Precision Oncology Company with Approval of FYARRO and Ongoing Tumor Agnostic Registrational Trial





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