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**UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION**  
Washington, D.C. 20549

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**FORM 8-K**

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**CURRENT REPORT**  
**Pursuant to Section 13 or 15(d)**  
**of the Securities Exchange Act of 1934**

**Date of Report (Date of earliest event reported): March 24, 2026**

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**ABSCI CORPORATION**  
(Exact name of registrant as specified in its charter)

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**Delaware**  
(State or other jurisdiction  
of incorporation)

**001-40646**  
(Commission  
File Number)

**85-3383487**  
(I.R.S. Employer  
Identification No.)

**18105 SE Mill Plain Blvd**  
**Vancouver, WA 98683**  
(Address of principal executive offices, including zip code)

**(360) 949-1041**  
(Registrant's telephone number, including area code)

**Not Applicable**  
(Former Name or Former Address, if Changed Since Last Report)

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Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:

- Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)
- Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)
- Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))
- Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

Securities registered pursuant to Section 12(b) of the Act:

<u>Title of each class</u>	<u>Trading Symbol(s)</u>	<u>Name of each exchange on which registered</u>
Common Stock, \$0.0001 par value per share	ABSI	The Nasdaq Global Select Market

Indicate by check mark whether the registrant is an emerging growth company as defined in Rule 405 of the Securities Act of 1933 (§ 230.405 of this chapter) or Rule 12b-2 of the Securities Exchange Act of 1934 (§ 240.12b-2 of this chapter).

Emerging growth company

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act.

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**Item 2.02. Results of Operations and Financial Condition.**

On March 24, 2026, Absci Corporation (the “Company”) announced its financial results for the fourth quarter ended December 31, 2025. A copy of the press release is being furnished as Exhibit 99.1 to this Current Report on Form 8-K.

The information contained in Item 2.02 of this Current Report on Form 8-K, together with Exhibit 99.1 hereto, is being furnished and shall not be deemed to be “filed” for the purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the “Exchange Act”), or otherwise subject to the liabilities of that section and shall not be incorporated by reference in any filing under the Securities Act of 1933, as amended, or the Exchange Act, except as shall be expressly set forth by specific reference in such filing.

**Item 8.01. Other Events.**

From time to time, the Company presents and/or distributes slides and presentations to the investment community to provide updates and summaries of its business. On March 24, 2026, the Company released a presentation which includes certain internal pipeline program updates, which is available on the “News & Events” section of the Company’s website. A copy of this presentation is filed as Exhibit 99.2 to this Current Report on Form 8-K and is incorporated by reference herein.

**Item 9.01. Financial Statements and Exhibits.**

(d) Exhibits

[99.1 Press Release issued by the Company on March 24, 2026, furnished herewith.](#)

[99.2 Absci Corporate Presentation March 2026](#)

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**SIGNATURE**

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

Absci Corporation

Date: March 24, 2026

By: /s/ Shelby Walker  
Shelby Walker  
Chief Legal Officer



### **Absci Reports Business Updates and Fourth Quarter and Full Year 2025 Financial and Operating Results**

*Successfully dosed first three cohorts in SAD portion of ongoing ABS-201™ HEADLINE trial; well-tolerated with favorable emerging safety data*

*Unveiled human ex vivo data demonstrating that ABS-201 stimulates hair growth and regenerates follicle stem cell niche*

*Appointed seasoned biopharmaceutical executive Ransi Somaratne, M.D., FACC, MBA as Chief Medical Officer*

*Cash, cash equivalents, and marketable securities sufficient to fund operations into the first half of 2028*

**VANCOUVER, Wash. and NEW YORK, March 24, 2026** – Absci Corporation (Nasdaq: ABSI), a clinical-stage biopharmaceutical company advancing breakthrough therapeutics designed with generative AI, today reported financial and operating results for the quarter and full year ended December 31, 2025.

“Over the past year, we advanced ABS-201 from preclinical to three dosed cohorts in our HEADLINE trial with favorable emerging safety data, extending our track record of moving from AI design to clinic in approximately two years at a fraction of industry cost,” said Sean McClain, Founder and CEO. “Dr. Ransi Somaratne has joined as our first Chief Medical Officer to lead clinical execution and strategy. We are focused on delivering interim proof-of-concept data in the second half of 2026 and initiating our endometriosis Phase 2 by year-end. In AGA and endometriosis, there is a significant unmet need, as no approved disease-modifying therapeutic options exist for these patients. This is the kind of whitespace where we believe our AI-native drug creation strategy can generate the most value.”

#### **Recent Highlights**

- Successfully dosed first three cohorts in single ascending dose (SAD) portion of ongoing Phase 1/2a HEADLINE trial. ABS-201 has been well tolerated to date, with favorable emerging safety data.
- Unveiled human *ex vivo* data demonstrating that ABS-201 effectively stimulates hair growth by regenerating the stem cell niche as well as promoting additional key growth modulators. In these studies, ABS-201 treatment significantly inhibited the PRLR signaling pathway (STAT5 phosphorylation), which correlated with prolongation of anagen and restoration of growth

signaling, preservation and expansion of the stem cell niche, and potential for follicle reconversion.

- Released manuscript on Origin-1: a generative AI platform that designs full-length monoclonal antibodies (mAbs) against "zero-prior" epitopes. "Zero-prior" means that to our knowledge there are no published reports describing a protein that binds to the target at the selected epitope. In contrast to traditional screening methods, Origin-1 generated potential lead candidates by screening fewer than one hundred designs per target. This platform is potentially the first demonstration of *de novo* design of full-length mAbs against "zero-prior" epitopes with atomically accurate complex structures and functional activity.
- Appointed seasoned biopharmaceutical executive Ransi Somaratne, M.D., FACC, MBA as Chief Medical Officer to spearhead the clinical strategy and execution for Absci's expanding pipeline of AI-designed therapeutics through clinical development. Dr. Somaratne joins Absci from Vertex Pharmaceuticals, where he served as Senior Vice President of Clinical Development and Translational Medicine, and previously held various roles at BioMarin Pharmaceutical and Amgen.

#### Internal Pipeline Updates and 2026 Outlook

- **ABS-201 (anti-PRLR antibody) for androgenetic alopecia:** ABS-201 is an anti-PRLR antibody, currently undergoing Phase 1/2a studies, in development for androgenetic alopecia (AGA), commonly known as male and female pattern hair loss. Absci believes that ABS-201, if successfully developed and approved, could provide a significant new category of AGA treatment that offers potentially durable hair growth with a convenient administration profile. Today, Absci announced that it has successfully dosed the first three cohorts in the SAD portion of its ongoing Phase 1/2a HEADLINE trial. ABS-201 has been well tolerated to date, with favorable emerging safety data. Absci anticipates reporting preliminary safety, tolerability, and pharmacokinetic (PK) data in the first half of 2026, with interim proof-of-concept data in the second half of 2026 and full proof-of-concept data in early 2027.
- **ABS-201 (anti-PRLR antibody) for endometriosis:** Absci is also developing ABS-201 for endometriosis, a large, underserved market with significant unmet medical need and poor standard of care. Endometriosis is prevalent in up to 10% of women worldwide, including an estimated 9 million women in the U.S., and there is currently no FDA-approved disease-modifying therapy. ABS-201 for endometriosis represents a novel mechanism (non-sex steroid hormone), with potential to be disease-modifying, act on both pain and lesion growth, and offer an improved safety profile. Absci anticipates initiation of a Phase 2 clinical trial for endometriosis in the fourth quarter of 2026, with potential proof-of-concept data in the second half of 2027.

- **ABS-101 (anti-TL1A antibody):** Absci continues to explore potential partnership and outlicensing opportunities for ABS-101, as well as first-in-class indication expansion opportunities for this target.
- **ABS-301 (potential first-in-class antibody for undisclosed immuno-oncology target):** ABS-301 is a fully human antibody designed to bind to a novel target discovered through Absci's Reverse Immunology platform. Absci has presented data for this program showing that expression of ABS-301's target suggests broad potential in squamous cell carcinomas and beyond.
- **ABS-501 (novel AI-designed anti-HER2 antibody):** For this program, Absci has identified antibody leads using its zero-shot *de novo* AI technology with the following characteristics: novel epitope interactions, increased or equivalent affinity to *trastuzumab* in preclinical settings, efficacious against a *trastuzumab*-resistant xenograft tumor, and good developability.
- **Drug Creation Partnerships:** Absci continues to make further progress on its existing drug creation partnerships and anticipates signing one or more partnerships, including with a Large Pharma company, in 2026.

Based on the company's current plans, Absci believes its existing cash, cash equivalents, and marketable securities will be sufficient to fund its operations into the first half of 2028.

#### Fourth Quarter 2025 Financial Results

Revenue was \$0.7 million for the three months ended December 31, 2025 compared to \$0.7 million for the three months ended December 31, 2024.

Research and development expenses were \$25.3 million for the three months ended December 31, 2025 compared to \$18.4 million for the three months ended December 31, 2024. This increase was primarily driven by advancement of Absci's internal programs, including direct costs associated with external preclinical and clinical development of ABS-101 and ABS-201.

Selling, general, and administrative expenses were \$8.6 million for the three months ended December 31, 2025 compared to \$8.8 million for the three months ended December 31, 2024.

Operating expenses for the three months ended December 31, 2025 were offset by a \$5.1 million gain recorded on settlement of the Company's contingent consideration during the fourth quarter 2025, which resulted in the receipt of \$8.7 million in unrestricted cash.

Net loss was \$29.6 million for the three months ended December 31, 2025, as compared to \$29.0 million for the three months ended December 31, 2024.

**Full Year 2025 Financial Results**

Revenue was \$2.8 million for the twelve months ended December 31, 2025 compared to \$4.5 million for the twelve months ended December 31, 2024.

Research and development expenses were \$81.4 million for the twelve months ended December 31, 2025 compared to \$63.9 million for the twelve months ended December 31, 2024. This increase was primarily driven by advancement of Absci's internal programs, including direct costs associated with external preclinical and clinical development.

Selling, general, and administrative expenses were \$35.1 million for the twelve months ended December 31, 2025 compared to \$36.2 million for the twelve months ended December 31, 2024. This decrease was primarily due to a reduction in personnel-related costs.

Operating expenses for the twelve months ended December 31, 2025 were offset by a \$5.1 million gain recorded on settlement of the Company's contingent consideration during the fourth quarter 2025.

Net loss was \$115.2 million for the twelve months ended December 31, 2025, as compared to \$103.1 million for the twelve months ended December 31, 2024.

Cash, cash equivalents, and marketable securities as of December 31, 2025 were \$144.3 million, compared to \$152.5 million as of September 30, 2025.

Based on the company's current plans, Absci believes its existing cash, cash equivalents, and marketable securities will be sufficient to fund its operations into the first half of 2028.

**Webcast Information**

Absci will host a conference call to discuss its fourth quarter and full year 2025 business updates and financial and operating results on Tuesday, March 24, 2026 at 4:30 p.m. Eastern Time / 1:30 p.m. Pacific Time. A webcast of the conference call can be accessed at [investors.absci.com](https://investors.absci.com). The webcast will be archived and available for replay for at least 90 days after the event.

**About Absci**

Absci is advancing the future of drug discovery with generative design to create better biologics for patients, faster. Our Integrated Drug Creation™ platform combines cutting-edge AI models with a synthetic biology data engine, enabling the rapid design of innovative therapeutics that address challenging therapeutic targets. Absci's approach leverages a continuous feedback loop between advanced AI algorithms and wet lab validation. Each cycle refines our data and strengthens our models, facilitating rapid innovation and enhancing the precision of our therapeutic designs. Alongside collaborations with top pharmaceutical, biotech, tech, and academic leaders, Absci is advancing its own pipeline of AI designed therapeutics including ABS-201™, a groundbreaking innovation in hair regrowth with the potential to redefine treatment possibilities for androgenetic alopecia, commonly known as male

and female pattern hair-loss. ABS-201 is also being investigated as a potential “best-in-class” therapeutic for endometriosis, a condition with significant unmet medical need and market potential. Absci is headquartered in Vancouver, WA, with AI Research Labs in New York City and Serbia, and an Innovation Center in Switzerland. Learn more at [www.absci.com](http://www.absci.com) or follow us on LinkedIn (@absci), X (@Abscibio) and YouTube.

#### **Forward-Looking Statements**

Statements contained in this press release regarding matters that are not historical facts are “forward-looking statements” within the meaning of the Private Securities Litigation Reform Act of 1995. Because such statements are subject to risks and uncertainties, actual results may differ materially from those expressed or implied by such forward-looking statements. Such statements include, but are not limited to, statements regarding any or all of the following: (i) Absci’s preclinical studies, clinical trials, as well as partnered and internally developed programs, including, without limitation, manufacturing capabilities, status of such studies and trials and expectations regarding data, safety and efficacy generally; (ii) data included in the above-described oral presentation, as well as the ability to use data from ongoing and planned clinical trials for the design and initiation of further clinical trials; (iii) projections regarding potential market opportunity, potential regulatory approval, the final approved label, and evolving competitive landscapes, as well as certain research findings based on participant responses to a hypothetical product profile and not any clinical results for ABS-201; (iv) Absci’s strategy, goals, anticipated financial performance and the sufficiency of its cash resources; (v) regulatory submissions and authorizations, including timelines for and expectations regarding any anticipated regulatory agency decisions; (vi) the expected benefits of its collaborations with partners; and (vii) the therapeutic value, development, and commercial potential of antibody therapies, as well as other technologies. Risks that contribute to the uncertain nature of the forward-looking statements include, without limitation, the risks and uncertainties discussed under the heading “Risk Factors” in Absci Corporation’s most recent annual report on Form 10-K and in any other subsequent filings made by Absci Corporation with the U.S. Securities and Exchange Commission. Existing and prospective investors are cautioned not to place undue reliance on these forward-looking statements, which speak only as of the date they are made. We disclaim any obligation or undertaking to update or revise any forward-looking statements contained in this press release, other than to the extent required by law.

#### **Investor Contact:**

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Corporate Vice President  
Head of Investor Relations  
[investors@absci.com](mailto:investors@absci.com)

**Media Contact:**

[press@absci.com](mailto:press@absci.com)

**Absci Corporation**  
**Consolidated Statements of Operations**

(In thousands, except for share and per share data)	(Unaudited)			
	For the Three Months Ended December 31,		For the Years Ended December 31,	
	2025	2024	2025	2024
Revenues				
Partner program revenue	\$ 650	\$ 665	\$ 2,800	\$ 4,534
Operating expenses				
Research and development	25,347	18,377	81,418	63,859
Selling, general and administrative	8,617	8,828	35,058	36,174
Depreciation and amortization	2,828	3,234	11,742	13,389
Gain on settlement of contingent consideration	(5,101)	—	(5,101)	—
Total operating expenses	31,691	30,439	123,117	113,422
Operating loss	(31,041)	(29,774)	(120,317)	(108,888)
Other income (expense)				
Interest expense	(29)	(109)	(209)	(565)
Other income, net	1,346	921	5,412	6,417
Total other income, net	1,317	812	5,203	5,852
Loss before income taxes	(29,724)	(28,962)	(115,114)	(103,036)
Income tax expense	162	(21)	(69)	(70)
Net loss	\$ (29,562)	\$ (28,983)	\$ (115,183)	\$ (103,106)
Net loss per share:				
Basic and diluted	\$ (0.20)	\$ (0.25)	\$ (0.84)	\$ (0.94)
Weighted-average common shares outstanding:				
Basic and diluted	150,610,966	114,929,962	136,776,885	110,239,870

**Absci Corporation**  
**Consolidated Balance Sheets**

(In thousands, except for share and per share data)	December 31, 2025	December 31, 2024
<b>ASSETS</b>		
Current assets:		
Cash and cash equivalents	\$ 20,025	\$ 41,213
Restricted cash	—	15,947
Marketable securities	124,267	71,212
Prepaid expenses and other current assets	5,281	5,459
Total current assets	149,573	133,831
Operating lease right-of-use assets	2,914	3,968
Property and equipment, net	20,860	29,167
Intangibles, net	41,514	44,883
Restricted cash, long-term	1,053	1,054
Other long-term assets	383	705
<b>TOTAL ASSETS</b>	<b>\$ 216,297</b>	<b>\$ 213,608</b>
<b>LIABILITIES AND STOCKHOLDERS' EQUITY</b>		
Current liabilities:		
Accounts payable and accrued expenses	\$ 19,348	\$ 10,449
Accrued expenses	—	12,750
Contingent consideration	—	2,733
Long-term debt	873	1,608
Operating lease obligations	1,805	739
Deferred revenue	739	1,116
Total current liabilities	22,765	28,656
Long-term debt, net of current portion	—	1,257
Operating lease obligations, net of current portion	2,624	4,429
Deferred revenue, long-term	436	—
Other long-term liabilities	1,023	133
<b>TOTAL LIABILITIES</b>	<b>26,848</b>	<b>34,475</b>
<b>STOCKHOLDERS' EQUITY</b>		
Preferred stock	—	—
Common stock	15	12
Additional paid-in capital	813,627	688,726
Accumulated deficit	(624,784)	(509,601)
Accumulated other comprehensive income (loss)	591	(4)
<b>TOTAL STOCKHOLDERS' EQUITY</b>	<b>189,449</b>	<b>179,133</b>
<b>TOTAL LIABILITIES AND STOCKHOLDERS' EQUITY</b>	<b>\$ 216,297</b>	<b>\$ 213,608</b>

GENERATIVE AI

# DRUG CREATION

Corporate Presentation  
March 2026

absci.

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## Disclaimers




### 1 FORWARD-LOOKING STATEMENTS

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### 2 MARKET AND STATISTICAL INFORMATION

This presentation also contains estimates and other statistical data made by independent parties and by us relating to market size and growth and other industry data. These data involve a number of assumptions and limitations, and you are cautioned not to give undue weight to such estimates. We have not independently verified the data generated by independent parties and cannot guarantee their accuracy or completeness.

### 3 TRADEMARK USAGE

This presentation/document/webpage contains references to our trademarks and service marks and to those belonging to third parties. Absci®, , SoluPro®, Bionic SoluPro® and SoluPure® are Absci registered trademarks with the U.S. Patent and Trademark Office. We also use various other trademarks, service marks and trade names in our business, including the Absci AI logo mark () , the Unlimit with us mark () , Denovium, Integrated Drug Creation, HiPrBind, and IgDesign. All other trademarks, service marks or trade names referred to in this presentation/document/webpage are the property of their respective owners. Solely for convenience, the trademarks and trade names in this presentation/document/webpage may be referred to with or without the trademark symbols, but references which omit the symbols should not be construed as any indicator that their respective owners will not assert, to the fullest extent under applicable law, their rights thereto.

# From Code to Clinic

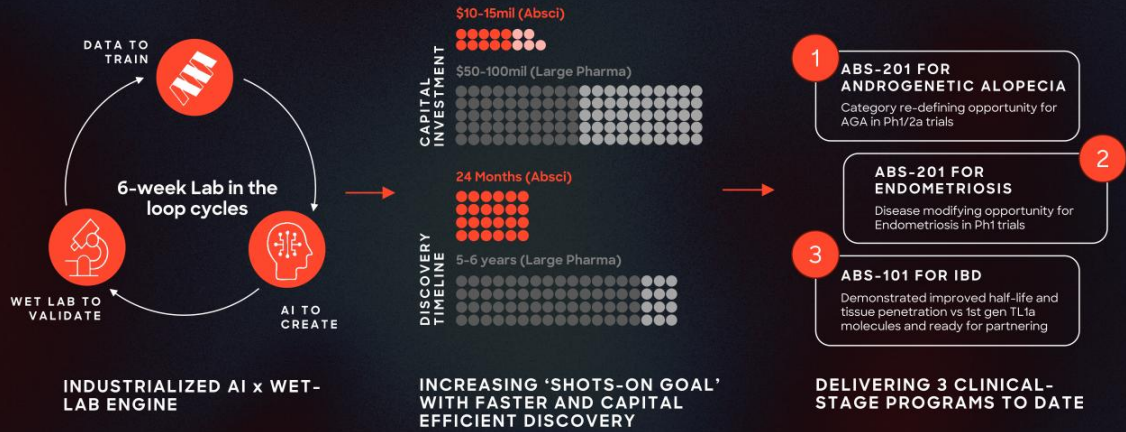
## 1. LEADING AI PLATFORM FOR CREATING ANTIBODY-BASED THERAPEUTICS

- Interdisciplinary Team with 10+ approved drugs and AI expertise
- Integrated Lab-in-the-Loop leveraging 77k ft<sup>2</sup> automated wet-lab
- Leading AI platform for *de novo* design and AI optimization of antibody-based therapeutics

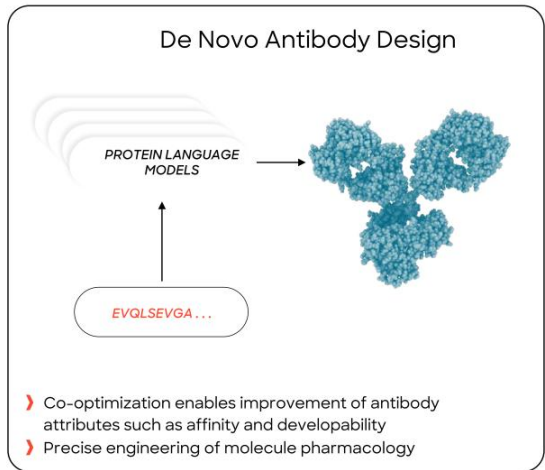
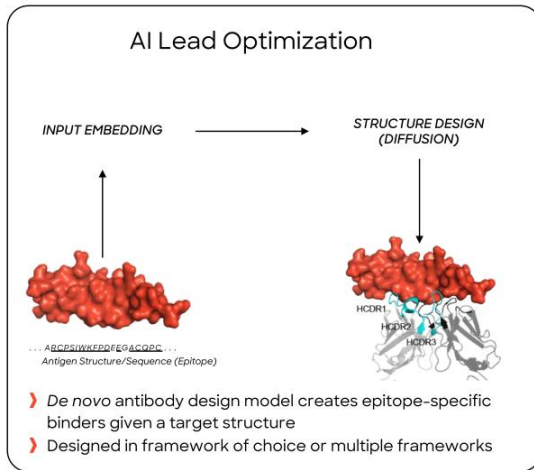
## 2. DIFFERENTIATED PIPELINE OF AI-DESIGNED THERAPEUTICS

- ABS-201 (anti-prolactin receptor)
  - Androgenetic Alopecia (AGA): Accelerated Ph1/2a trial on track to initiate December 2025, with interim POC readout 2H 2026
  - Endometriosis (Endo): Indication expansion into endometriosis with anticipated Ph2 initiation in 4Q2026 with PoC readout as early as 2H 2027
- **Preclinical pipeline** focused on metabolism and I&I

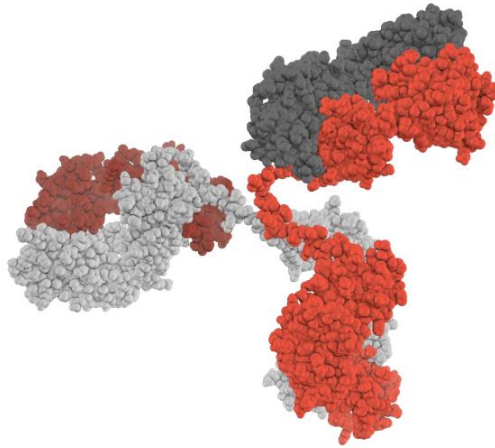
# Industrializing Drug Discovery



# Leadership in AI de novo design of antibody-based therapeutics



## We use AI to create novel & differentiated therapeutics



✓ EPI-TOPE-SPECIFIC DESIGN +  
EPI-TOPE INTERFACE OPTIMIZATION

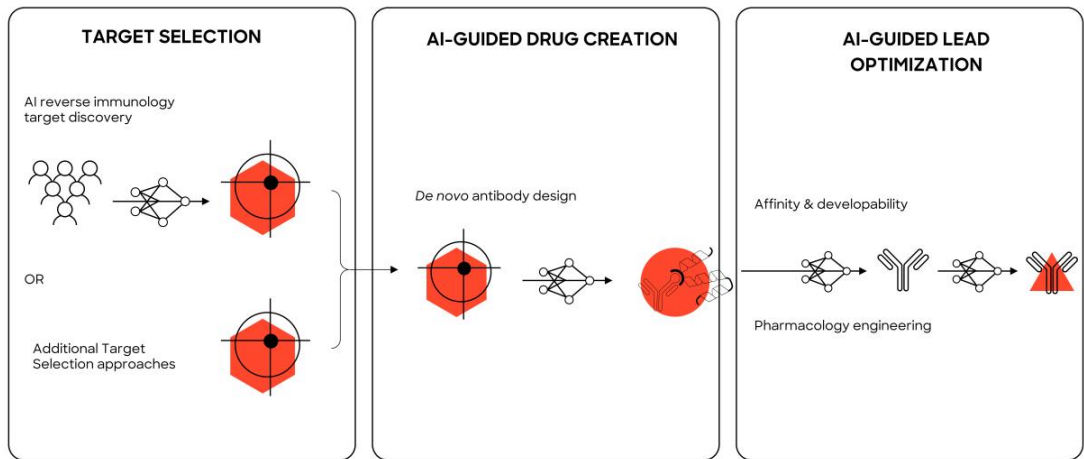
✓ ENHANCED POTENCY AND MOA

✓ ABILITY TO ADDRESS DIFFICULT  
TARGET CLASSES, E.G. GPCRS

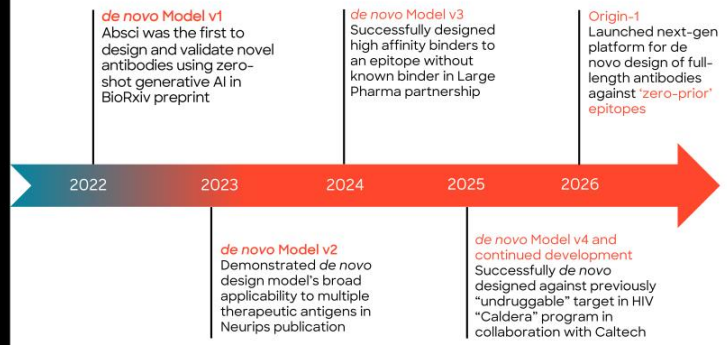
✓ ENABLING FEATURES: MULTI-VALENCY,  
pH-DEPENDENT BINDING

✓ POTENTIAL TO CREATE MEANINGFUL  
IP: 100S TO 10,000S OF FUNCTIONALLY  
VALIDATED SEQUENCES ENABLED BY  
PROPRIETARY WET-LAB VALIDATION

# Leveraging AI throughout the end-to-end drug discovery process



Since publishing the first work in **AI de novo antibody design**, Absci has continued to rapidly progress and lead the field

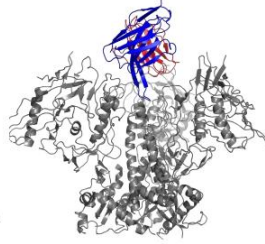


## Platform case studies

### De Novo Antibody Design

De novo antibody design program in collaboration with Caltech funded by the Gates Foundation

Caltech BILL & MELINDA GATES FOUNDATION

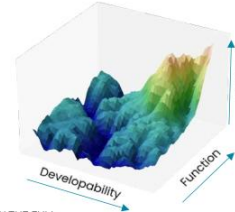


VIEW THE FULL CASE STUDY

- › **Goal:** Create universally neutralizing HIV antibody by binding conserved epitope within "Caldera" region of HIV gp120
- › Absci's *de novo* design platform can successfully address difficult to drug target epitopes

### AI Lead Optimization

AI lead optimization for PH sensitivity which may reduce toxicity and/or improve efficacy of therapeutic mAbs



Model searches a massive space of  $\sim 10^{19}$ , identifying functional and developable antibodies in one step.



VIEW THE FULL CASE STUDY

- › **Goal:** Co-optimize antibodies for pH sensitive binding to increase efficacy and reduce toxicity
- › Absci's lead optimization platform enables molecules with differentiated pharmacology

“Multilingual”  
team with  
expertise in AI  
and drug  
creation

EXPERTISE AND  
BACKGROUND FROM:



Leadership Team:



Sean McClain Founder, CEO & Board Director  
Zach Jonasson, PhD Chief Financial Officer & Chief Business Officer  
Andreas Busch, PhD Chief Innovation Officer  
Ransi Somaratne, MD Chief Medical Officer  
Shelby Walker, JD Chief Legal Officer  
Christine Lemke SVP, Portfolio & Growth Strategy  
Amir Shanehsazzadeh SVP, Chief AI Officer

Board Of Directors:





Frans Van Houten Chairman of the Board Former CEO, Royal Philips  
Sean McClain Founder, CEO & Board Director  
Sir Mene Pangalos, PhD Former EVP R&D AstraZeneca  
Mary Szela CEO & President TriSalus Life Sciences  
Joseph Sirosch, PhD Former CVP AI Microsoft  
Dan Rabinovitsj VP Hardware Engineering, Meta  
Karen McGinnis, CPA Former Chief Accounting Officer, Illumina

Scientific Advisory Board:

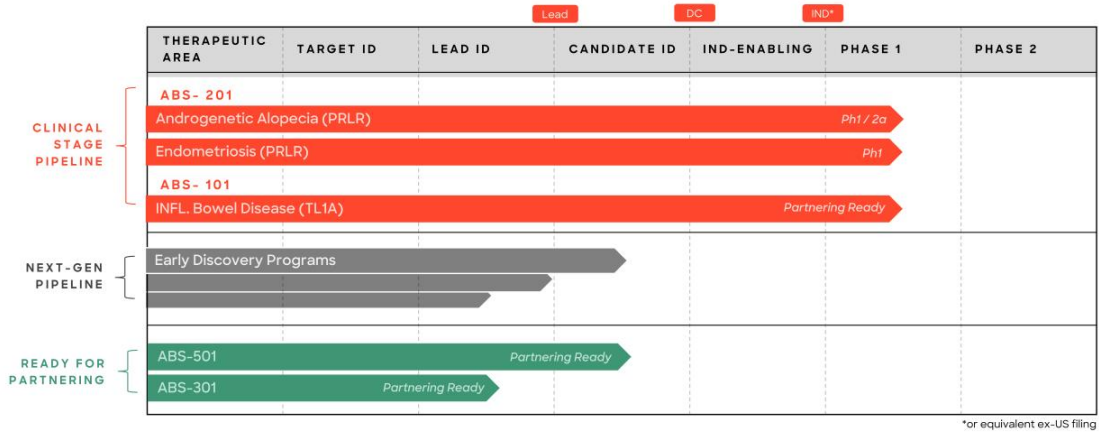


Sir Mene Pangalos, PhD Co-Chair SAB Former EVP R&D AstraZeneca  
Andreas Busch, PhD Co-Chair SAB Chief Innovation Officer  
Ian McInnes, PhD Vice Principal and Head of College University of Glasgow  
Luis Diaz, MD Head, Division of Solid Tumor Oncology Memorial Sloan Kettering Cancer Center  
John Wherry, PhD Director, Institute for Immunology & Immune Health, University of Pennsylvania  
Victor Greiff, PhD Associate Professor University of Oslo  
Hubert Truebel, MD, PhD, MBA Chief Medical Officer Acuris

## Absci at a Glance

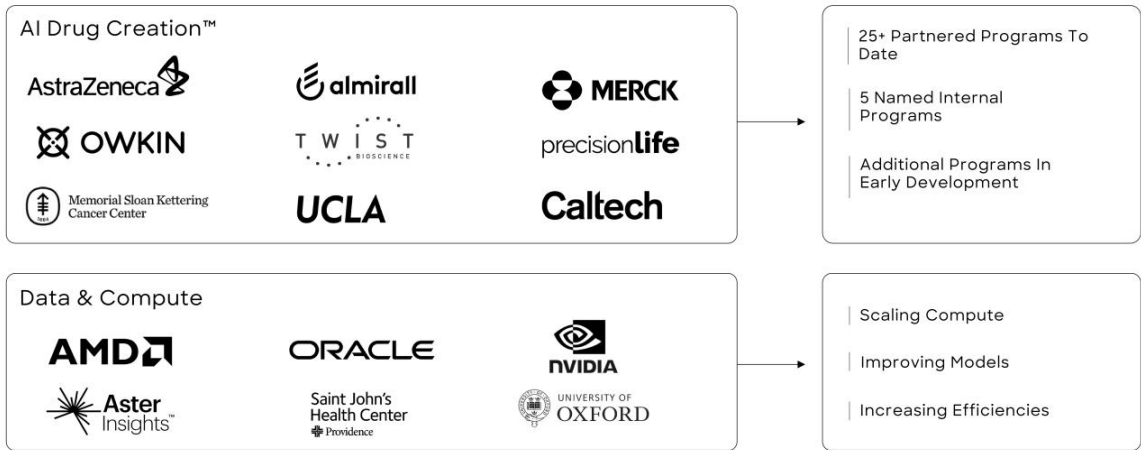
<p><b>140</b> <b>EMPLOYEES</b></p> <p>"Multi-lingual" AI + Drug Discovery expertise AI team drawing on experience from tech leaders:</p>  <p>Biologics drug discovery expertise from:</p> 	<p><b>77,000</b> <b>SQUARE FEET</b></p> <ul style="list-style-type: none"> <li>State-of-the-art wet and dry lab in Vancouver WA</li> <li>Absci AI Research (AAIR) lab in NYC</li> <li>Innovation Centre in Zug Switzerland</li> </ul>	<p><b>10+</b> <b>PARTNERS, INCLUDING</b></p> 
	<p><b>3</b> <b>CLINICAL STAGE PROGRAMS</b></p> <p>In Androgenetic Alopecia, Endometriosis, and Inflammatory Bowel Disease</p>	<p><b>&gt;\$600M</b> <b>CAPITAL RAISED TO DATE</b></p>

# Advancing and expanding our pipeline of novel & differentiated assets designed using AI



\*for equivalent ex-US filing

## Track record of industry-leading partnerships



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# Leading AI x Bio platform driving 2 Phase 2 readouts in the next 24 months

## ABS-201 in AGA

- Ph1/2a Study initiated Dec 2025
- Safety, Tolerability, and PK readout expected 1H 2026
- Interim PoC Readout - anticipated 2H 2026

## ABS-201 in ENDO

- Ph1/2a Study initiated Dec 2025
- Phase 2 initiation expected in 2H2026

## Absci's progress in Drug Creation

### Acceleration and Expansion of Lead Program

#### ABS-201 AGA

Accelerated development of ABS-201 in androgenetic alopecia

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- › Interim POC readout anticipated 2H 2026

#### ABS-201 Endo

Indication expansion for ABS-201 in Endometriosis

Ph2 study anticipated to initiate 4Q2026

### Differentiated AI designed pipeline

#### ABS-101

Phase 1 interim results reported with extended half-life vs 1<sup>st</sup> gen TL1A competitors

Advancing partnership & out-licensing discussions, including in potential 'first-in-class' indications

#### ABS-301 & 501

ABS-301: Potential first-in-class asset discovered through Reverse Immunology Platform

ABS-501: Candidate ID phase for novel HER2 program designed using de novo AI

## ABS-201 has the potential to unlock a wholly new category of therapy in hair “re-growth”

1. Significant clinical and commercial unmet need in androgenetic alopecia
2. Strong scientific rationale, with validated target, de-risked Mode of Action, and pharmacology
3. Straightforward development path with objective endpoints



## Underserved patient population looking for therapeutic innovation

~80 million Americans live with androgenetic alopecia (AGA)



**MALE AGA**

~50M men in the U.S.

Only 2 FDA approved drug therapies

**FEMALE AGA**

~30M women in the U.S.

Only 1 FDA approved drug therapy for women



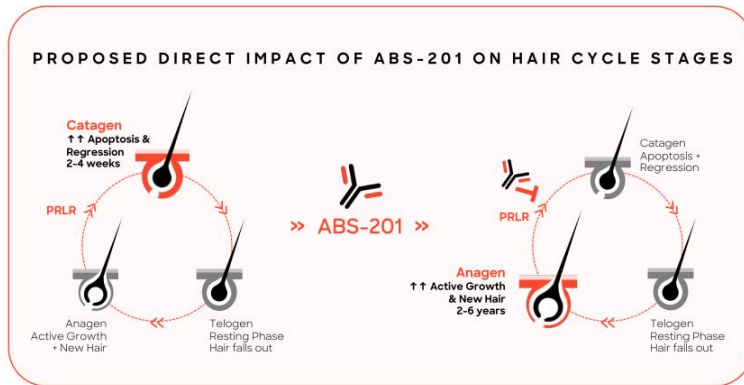
Growing patient population with limited therapeutic options and concerns of adverse side-effects

Last FDA approved therapy for androgenetic alopecia was in the 1990s

Patients and clinicians need better treatment options for "hair re-growth"

- › Hair re-growth, not just slowing of hair loss
- › Safe and minimal side effects
- › Durable effect
- › Convenient administration frequency
- › FDA approved

## PRLR inhibition for androgenetic alopecia is an innovative alternative to current treatment options

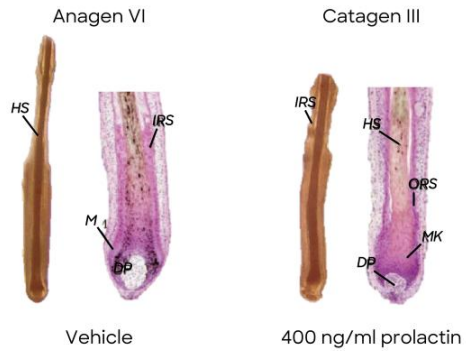


### ABS-201 HAS THE POTENTIAL TO:

- Shift the balance in hair cycle stage towards anagen phase<sup>1,2</sup> with:
  - Active and new hair growth
  - Prevention of telogen effluvium
- Promote a long-lasting effect after treatment cessation
- Block cessation of pigmentation, which may lead to the restoration of hair pigmentation<sup>2</sup>

## Prolactin impacts on organ-cultured human hair follicles

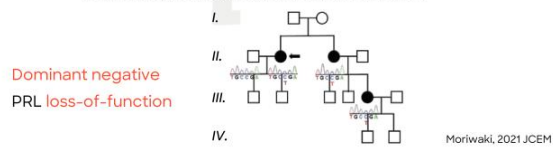
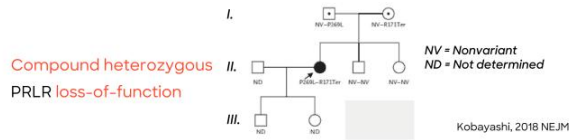
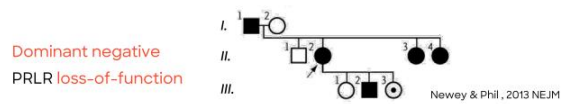
Prolactin drives hair follicle regression in human ex vivo culture



Prolactin prematurely induces a catagen-like stage in organ-cultured human hair follicles<sup>1</sup> characterized by:

- › Condensed shape of the dermal papilla (DP)
- › Diminishment of the hair matrix volume
- › Apparent cessation of pigmentation
- › Inhibition of hair shaft elongation

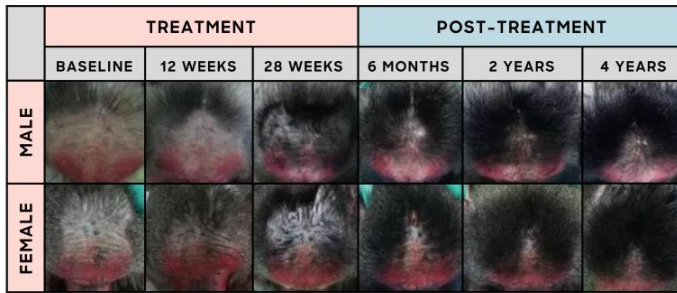
## PRLR inhibition anticipated to be safe & well tolerated as supported by human genetics



Reduced/Loss of PRL or PRLR Signaling

- › Postpartum agalactia
- › Otherwise in good health:
  - No apparent impact on fertility
  - No report on erectile dysfunction in male
  - Normal breast development and menses in females
  - Normal serum electrolytes and hormone levels (except elevated PRL in PRLR mutation carrier)
  - No reported abnormalities of other hypothalamic-pituitary axes

### Top head view of Stumptailed Macaque's showing phenotypic change over time

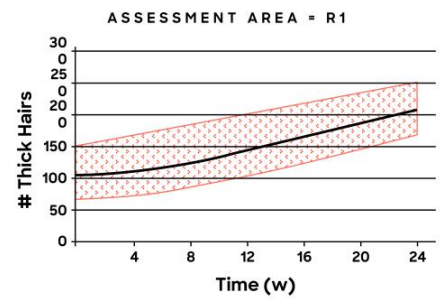


40mg/kg s.c. Q2W for 28 weeks

Study commissioned by Absci CIO Andreas Busch while at Bayer. Disclosure from competitor

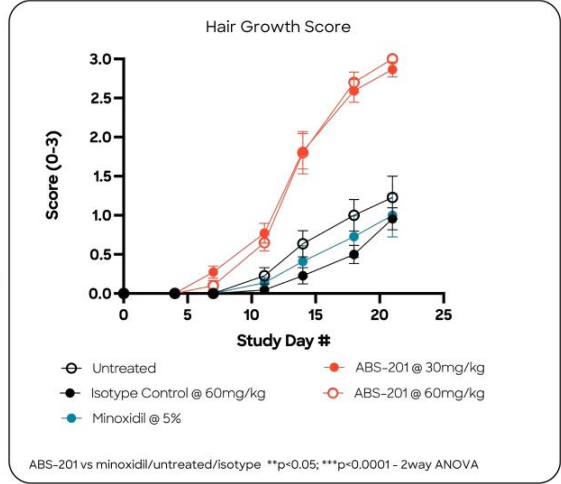
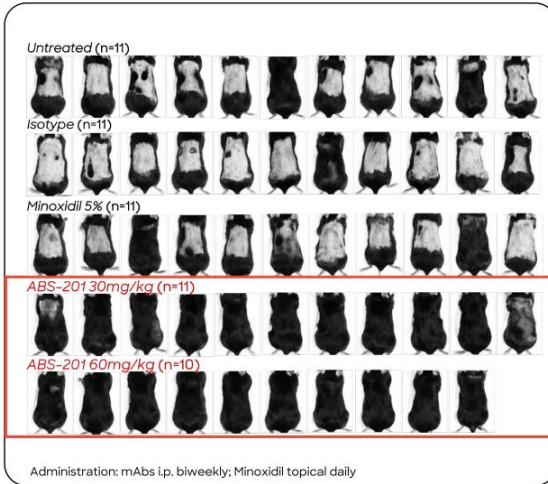
- Hair density & thickness improved with short treatment duration in primate model of androgenetic alopecia
- Hair growth remains and improves several years post cessation

### Terminal hair count "Thick Hairs" in prior bald areas

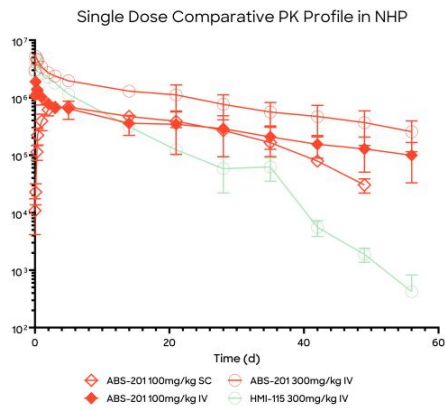


- Hair regrowth observed for both male and female animals (>100 hairs/cm2 increase in bald area at week 28 of treatment\*)

# ABS-201 shows superior efficacy vs 5% topical minoxidil in 21d hair regrowth model



## 56 Day NHP PK data confirms extended half-life profile and high SC bioavailability



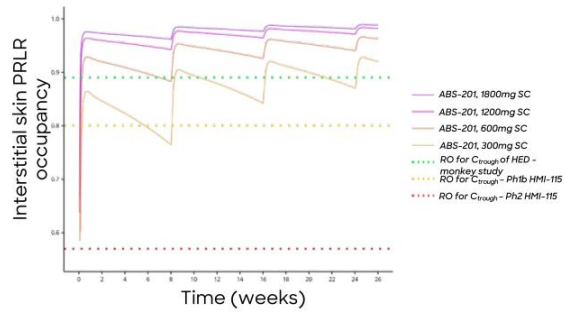
### NHP-PK 56 Day Results

- >3x extended half-life in NHPs compared to HMI-115
- High subcutaneous bioavailability in NHPs at >90%
- In silico prediction of Q8W-Q12W dosing intervals anticipated in humans
- Manufacturability & developability profile believed to enable future high concentration formulation targeting >150mg/mL

Based on PK/PD modeling, ABS-201 is anticipated to likely require only 2-3 doses over a 6-month treatment period, compared to HMI-115, which would likely require 6-12+\* doses in the same period, assuming the AGA indication is pursued.

\*assumption on HMI-115: 60mg/mL formulation and Q2W or Q4W dosing interval

## Modelling shows superiority of ABS-201 vs HMI-115 on PK & Receptor Occupancy

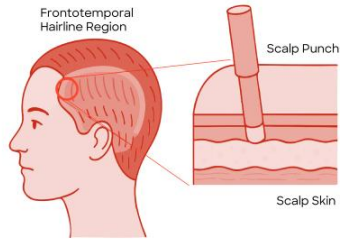


### Preliminary in Silico Modeling

- >3x extended half-life in NHPs predicted to translate in humans to Q8W-Q12W dosing intervals
- PK profile predicted to translate into higher interstitial skin concentrations resulting in higher receptor occupancy

Modelling assumptions include published NHP and Ph1b PK data on HMI-115 (formerly BAY 1158061), as well as in house generated in vitro and in vivo data. Parameters incl. 0.2 skin exposure coefficient,  $2.6 \times 10^{-2}$  nM interstitial PRLR concentration

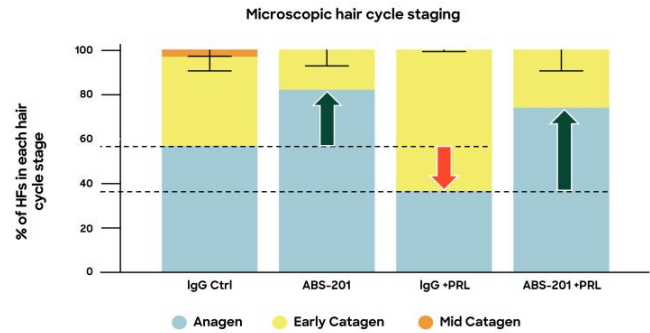
## ABS-201 in human ex vivo culture study supports MOA in human scalp follicles

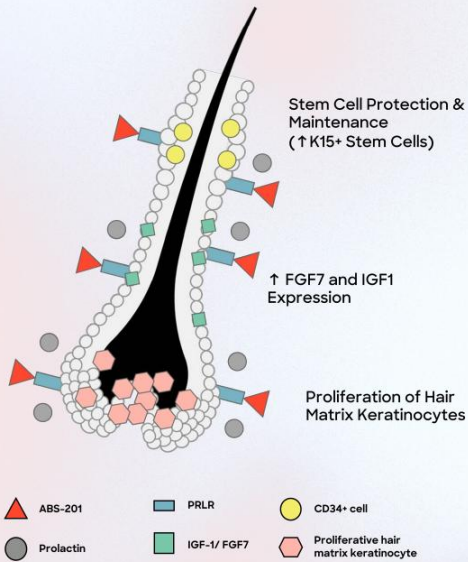


### MODEL SYSTEM:

- Frontotemporal male scalp skin is the most androgenetic alopecia affected skin region
- Organ culture is the most relevant human preclinical hair research tool ex vivo

### ABS-201 significantly prolongs anagen/inhibits catagen and stimulates hair matrix proliferation





## Additional ABS-201 ex vivo study found:

- Prolonging anagen phase and blocking catagen, thereby inhibiting telogen effluvium
- Protecting and promoting hair follicle stem cells and restoring CD34+ progenitor cells
- Stimulating key hair growth factors (IGF1, FGF7)
- Decreasing catagen driver TGFβ-2
- Increasing hair shaft and hair shaft keratin production

## Phase 1/2a trial designed to provide readouts on safety, tolerability, and PoC in AGA

### HEADLINE

#### Design Elements:

- Double-Blind, Placebo-controlled, FIH
- Multi-site study in Australia
- Dose range ensures predicted >90% RO

#### Population:

- Up to 227 male & female healthy participants
- SAD; n= 32 healthy volunteers
- MAD; n= 147 AGA subjects (Norwood Scale IIIv-V)
- Optional AGA cohorts in SAD/MAD; n= 48
- 3:1 randomization

#### Endpoints:

- **Primary:** Safety & Tolerability
- **Secondary:**
  - PK/PD
  - Efficacy readouts include target area hair count, width, and darkness (pigmentation)



#### Single Ascending Dose

Cohort 1  
150mg IV  
n=8

Cohort 2  
450mg IV  
n=8

Cohort 3  
900mg IV  
n=8

Cohort 4  
1800mg IV  
n=8

- First cohort fully enrolled and dosed - Enrollment of remaining cohorts ongoing
- **Dec 2025:** Initiated
- **1H 2026:** PK and interim safety expected



#### Multiple Ascending Dose (26 weeks)

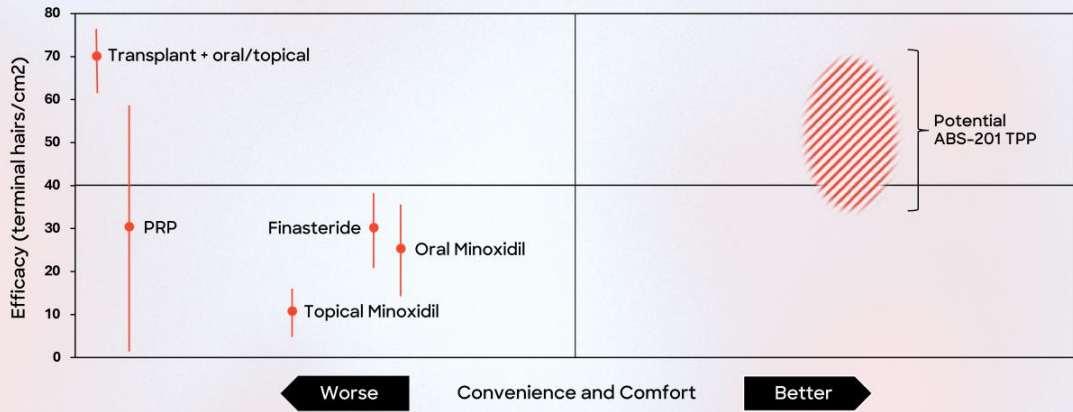
Cohort 1  
300mg SC  
n=49

Cohort 2  
600mg SC  
n=49

Cohort 3  
1200mg SC  
n=49

- MAD design enabling **PoC for AGA**
- **2Q 2026:** Expected initiation
- **2H 2026:** Expected 13-week interim PoC readout
- **Early 2027:** Expected 26-week topline PoC readout

## ABS-201 TPP aims to offer a new treatment category in AGA based on efficacy and convenience



\* Based on 2-3 injections during first 6 months for >2 years of hair growth  
 Efficacy at 24w for Vertex terminal hair count in male subjects: Oral Minoxidil (5mg/day); Panchaprateep 2020 (10.1007/s13555-020-00448-x) and Penha 2024 (doi:10.1001/jamadermatol.2024.0284); PRP; Dervishi 2019 (10.1111/jocd.13113);  
 Finasteride and Topical Minoxidil: Gupta 2022 (doi:10.1001/jamadermatol.2021.5743). Transplants based on KOL interviews.

## Consumer Research Commissioned by Absci Validates Market Potential of ABS-201

### Significant Unmet Need:

Driven by psycho-social impacts (loss of confidence, self-esteem) from AGA

### Strong Commercial Demand:

Nearly all men and roughly 90% of women are inclined to ask their doctors about ABS-201

### High Value Proposition:

Significant share of respondents willing to pay a premium for the ABS-201 TPP

### Disruptive Potential:

Over 1/3 of respondents would select ABS-201 before their current treatment, suggesting ABS-201 can effectively compete as first-line therapy

## 610 Participants:

306 Men | 304 Women

\*ALL PARTICIPANTS EXPERIENCING HAIR LOSS

UP TO  
**97%**  
MEN

UP TO  
**88%**  
WOMEN

EXTREMELY OR VERY  
LIKELY TO ASK HCP  
ABOUT ABS-201

**37%**  
MEN

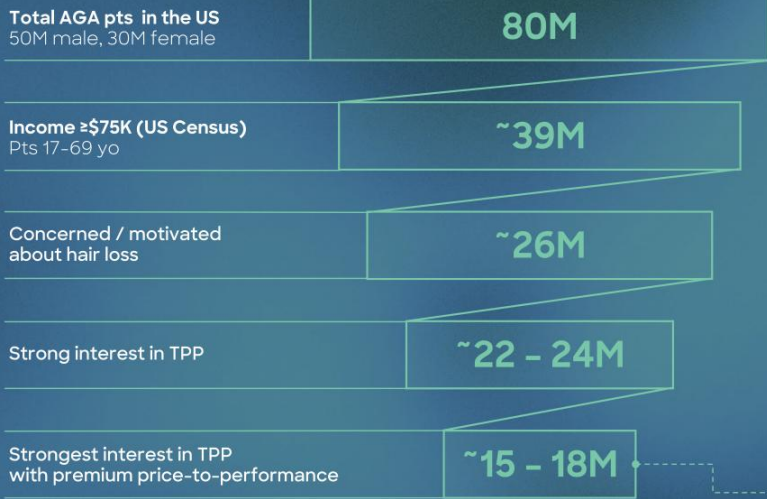
**36%**  
WOMEN

WOULD TRY ABS-201  
FIRST (FIRST LINE)

**80%**  
MEN

**81%**  
WOMEN

REPORT NEGATIVE  
PSYCHOLOGICAL  
IMPACT



Patient Funnel

>\$25B

ESTIMATED U.S. TAM

>\$40B

POTENTIAL GLOBAL TAM

5-9M Pts Treated/Year  
Assuming 2-3 Year Durability

## Absci's progress in Drug Creation

### Acceleration and Expansion of Lead Program

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Ph2 study anticipated to initiate 4Q2026

### Differentiated AI designed pipeline

#### ABS-101

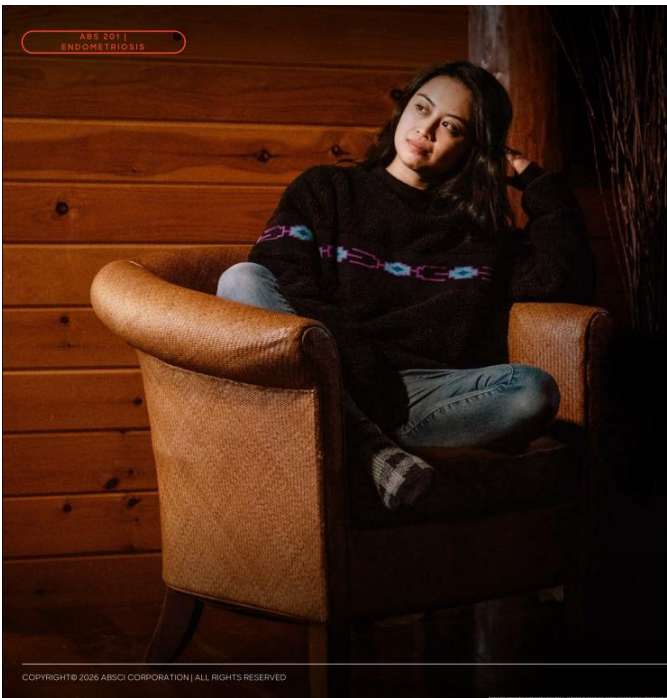
Phase 1 interim results reported with extended half-life vs 1<sup>st</sup> gen TL1A competitors

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#### ABS-301 & 501

ABS-301: Potential first-in-class asset discovered through Reverse Immunology Platform

ABS-501: Candidate ID phase for novel HER2 program designed using de novo AI

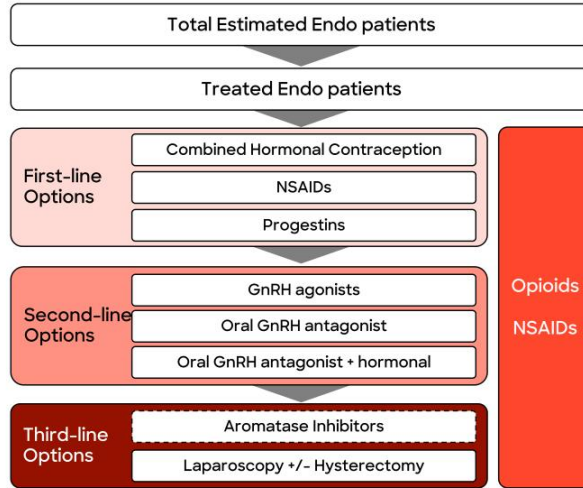


## Development of **ABS-201** in **Endometriosis**

1. Addresses Long-standing Unmet Medical Need and Poor standard of care
2. Strong Biological And Clinical Rationale: Including POC for PRLR mechanism in humans
3. Large, untapped market offers significant upside potential

# Hormonal therapies and surgical intervention make up the treatment paradigm for Endo

## Current Treatment Paradigm (US)

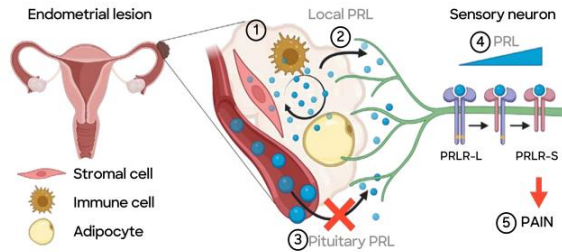


- Endo is a chronic condition with currently no medical or surgical cure
- Roughly, 75% of patients are estimated to use opioids and/or NSAIDs throughout their disease course
- Up to 33% of patients do not respond to hormonal treatment alone
  - Additionally, patients will pause treatment when seeking pregnancy
- GnRH therapies are typically prescribed by Gynecologist and often require formal Dx (surgical confirmation)
  - Due in part to higher cost, and AE profile which limit long-term use
- Notably, aromatase inhibitors are not FDA approved for Endo, but are used off-label
- Up to 40% of Endo patients undergo a laparoscopy and 12% receive hysterectomies
- Even after hysterectomy ~15% of patients still report pain symptoms



# PRLR antagonism is a novel and differentiated MoA in Endometriosis

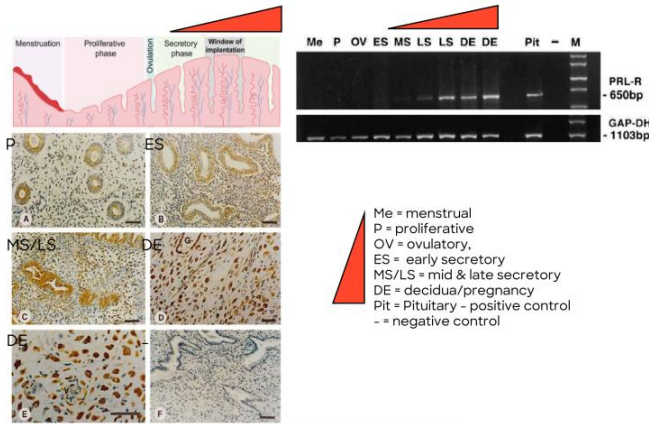
PRL and PRLR play a dual role in endometrial lesion development and pain response



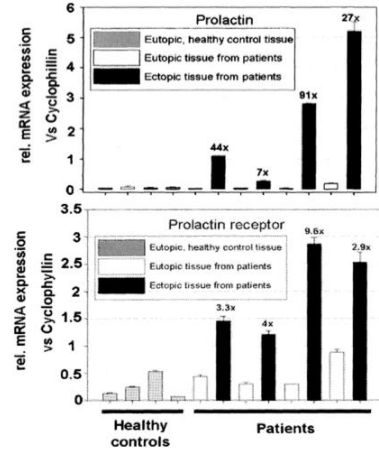
- Endometriotic lesions produce prolactin under estrogen/progesterone control.
- Excess prolactin promotes lesion growth and sensitizes pain-sensing nerves, contributing to chronic pelvic pain.
- Prolactin signaling is independent of sex-hormone pathways, offering a differentiated, non-hormonal treatment modality vs current therapies.

# PRL and PRLR increase during secretory phase in healthy tissue, and is overexpressed in endometrium of patients with endometriosis

Localization And Temporal Expression Of Prolactin Receptor In Human Endometrium



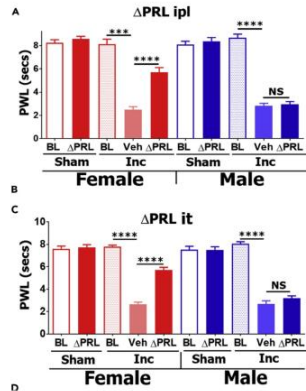
PRL & PRLR is Elevated Ectopic Endometriotic Lesions



Jones et al. Journal of Clinical Endocrinology and Metabolism, 1998; Otto et al. WO 2011/069795 A4

# PRLR antagonism suppresses postoperative pain in female mice and inhibits endometriosis interna formation

Prolactin Regulates Pain Responses via a Female-Selective Nociceptor-Specific Mechanism

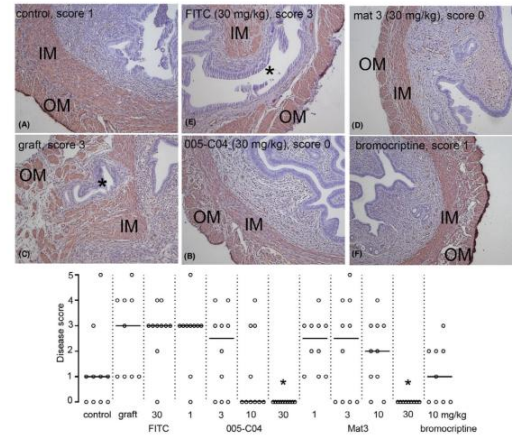


Depicted: pain incision (Inc) model for heat sensitivity & ΔPRL administration

Patil et al. iScience 2019; Otto et al. Pharmacol Res Perspect. 2022

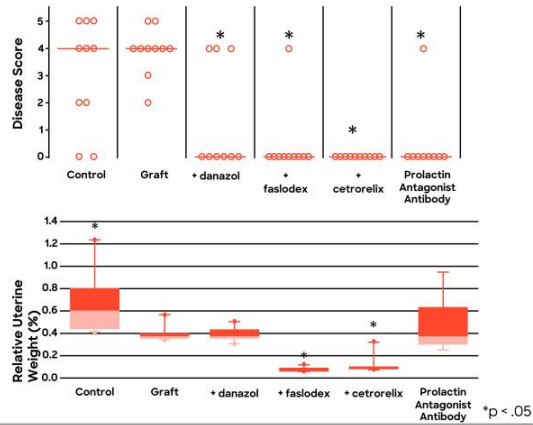
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The Effects of Prolactin Receptor Blockade in a Murine Endometriosis Interna Model

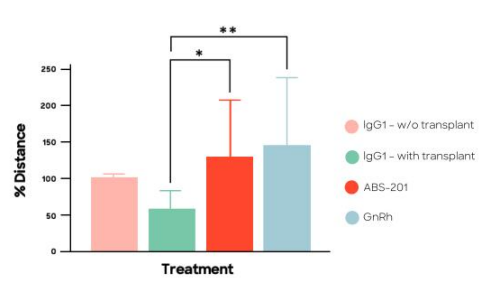


# PRLR antagonism reduces lesion formation and pain in endometriosis mouse model

Prolactin inhibition decreases endometrial lesion formation in female mouse interna



% Distance travelled at Week 7 (compared to baseline)



○ ABS-201 and GnRh modulator increase distance traveled relative to placebo over time as surrogate for pain reduction

\*\* p<0.01; \*\*\* p<0.001

# ABS-201: a potentially differentiated profile targeting a large underserved market opportunity

NOVEL TREATMENT OPTION FOR ~9M PATIENTS  
IN THE U.S. ALONE WITH ENDOMETRIOSIS

- **Novel Mechanism:** Non-sex-steroid (peptide) hormone
- **Potential for Improved Safety Profile:** Potential improved AE profile & longer use than GnRH
- **Dual Action:** Potential on both pain and lesion growth
- **Best-in-class Potential:** Superior developability and expected half-life
- **Disease Modifying:** Potential to treat cause
- **Clinically validated:** through HMI-115 Ph2 study

Potential to generate

>**\$4.5B**  
at peak sales

## Absci's progress in Drug Creation

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ABS-301: Potential first-in-class asset discovered through Reverse Immunology Platform

ABS-501: Candidate ID phase for novel HER2 program designed using de novo AI

ABS-101 | TL1A



## AI Designed TL1A antibody in Ph1 Clinical development; ready for partnering

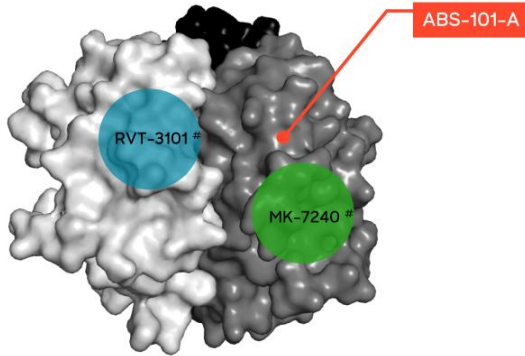
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### ABS-101 summary

- › Interim results from the first cohorts of Phase 1 trial demonstrated extended half-life as compared to 1st-generation anti-TL1A competitor programs
- › No apparent impact of ADA on PK and the overall safety profile was favorable with no serious adverse events reported to date
- › Trial on track to complete treatment period in Q1 2026 and we will no longer pursue additional internal clinical development of this asset following completion of the Phase 1 trial
- › Advancing partnership discussions, some of which leverage unique properties of ABS-101 and are focused on first-in-class indications outside of IBD

# Successful application of AI platform to generate high affinity variants

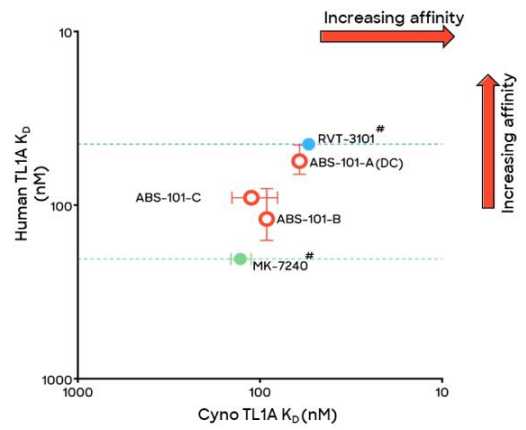
Epitope bins on TL1A\*



› Absci AI-designed and optimized leads span multiple unique epitopes on a single TL1A subunit

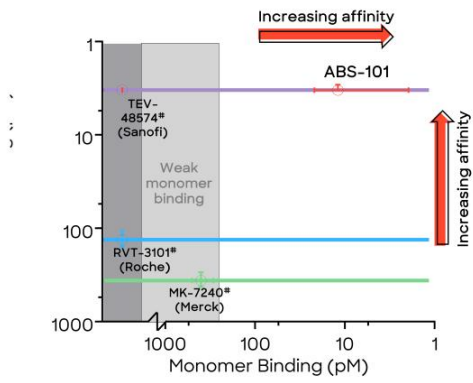
\*Estimated performance of a putative clinical competition molecule generated for in-house comparison

High Affinity mAbs with Preserved Cross-Reactivity

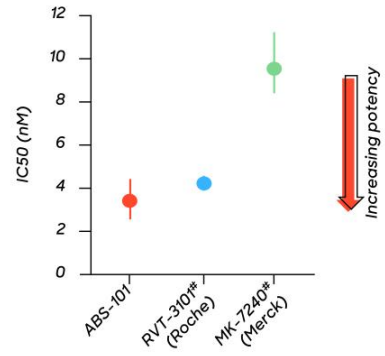


# AI-designed candidate with high affinity and potential for superior potency

Affinity by Biolayer Interferometry (BLI)

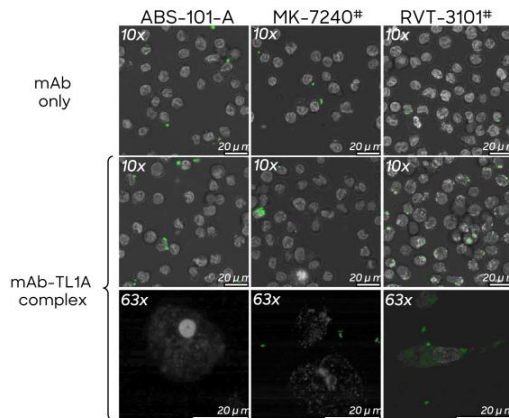


Apoptosis Inhibition Assay in TF-1 Cells



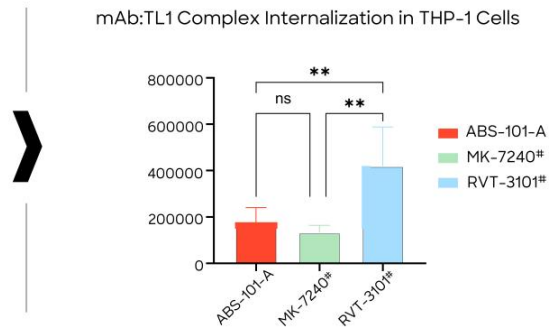
\*Estimated performance of a putative clinical competition molecule generated for in house comparison

## Internalization of mAb:TL1A complexes potentially contribute to immune activation and formation of ADA



doi: 10.1053/j.gastro.2019.08.009

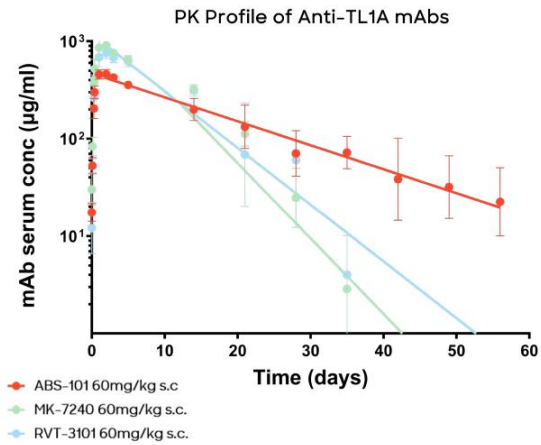
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ABS-101 and MK-7240# show reduced TL1A complex internalization versus RVT-3101#

\*Estimated performance of a putative clinical competition molecule generated for in house comparison

## NHP Pharmacokinetics & CMC Data



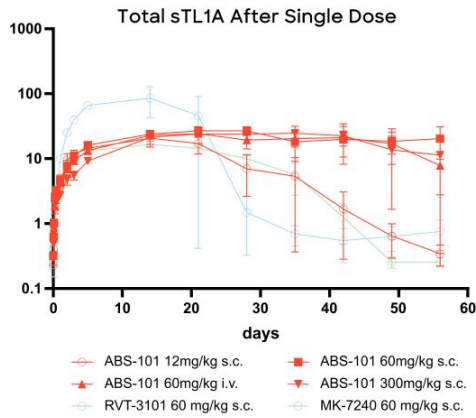
### NHP-PK & Preliminary 13-Week NHP GLP-TOX

- › 2-3x extended half-life in NHPs over 1<sup>st</sup> generation clinical competitors
- › ABS-101 shows enhanced biodistribution in NHPs, compared to antibodies in clinical development based on in silico modelling
- › High subcutaneous bioavailability in NHPs at ~80%
- › Preliminary 13-week GLP-tox shows no treatment-related adverse findings during in-life phase and necropsy

### CMC—High Concentration Formulation

- › Optimal developability profile allowed successful development of high-concentration formulation at 200mg/mL suitable for subcutaneous injection

## ABS-101 Non-Human Primate (NHP) data



ABS-201 shows dose dependent and sustained target engagement

- › Data confirm engagement of soluble TL1A (sTL1A) in non-human primates.
- › Target engagement is dose-dependent with a ceiling effect.
- › ABS-101's extended half-life translates into sustained target engagement compared to first generation TL1A antibodies at comparable dose and route of administration.

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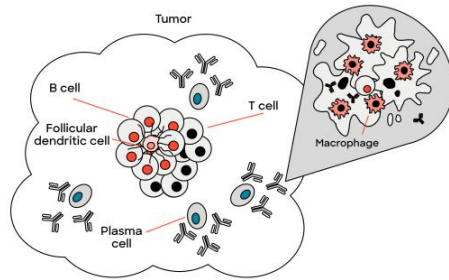
Advancing partnership & out-licensing discussions, including in potential 'first-in-class' indications

#### ABS-301 & 501

ABS-301: Potential first-in-class asset discovered through Reverse Immunology Platform

ABS-501: Candidate ID phase for novel HER2 program designed using de novo AI

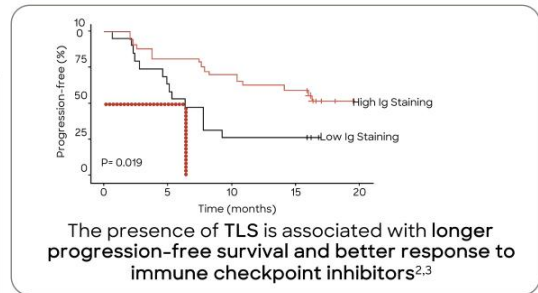
## Tertiary Lymphoid Structures (TLS): The focus of Absci's Reverse Immunology approach



Tertiary lymphoid structures (TLS) are centers of immune activity, such as B-cell proliferation and antibody production, that develop in chronically inflamed tissues<sup>1</sup>.

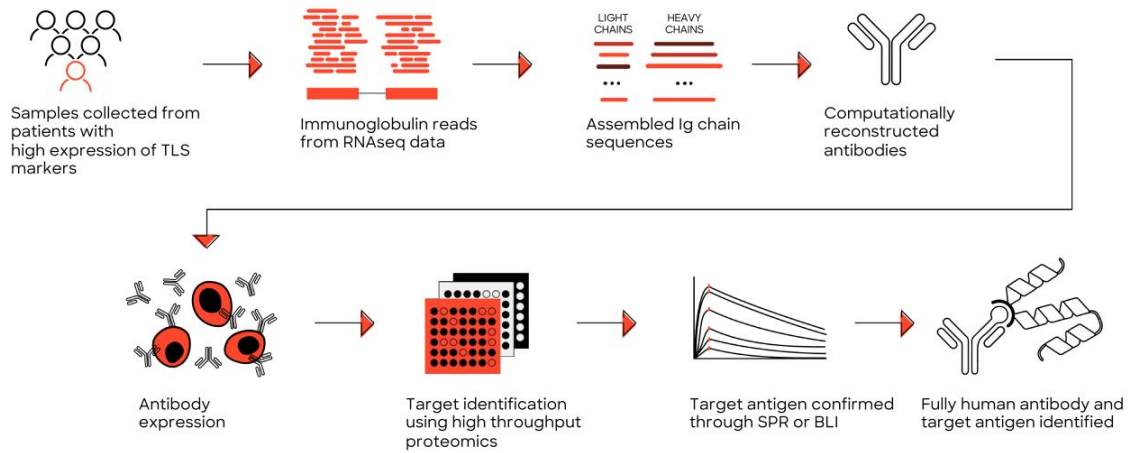
Antibodies from TLS are specialized for local antigens and play a significant role in the progression of chronic diseases and cancer, setting them apart from the general population of antibodies in the peripheral blood<sup>2</sup>.

<sup>1</sup> doi: 10.3389/fimmu.2018.01952; <sup>2</sup> doi: 10.1016/j.immuni.2022.02.001; <sup>3</sup> doi: 10.1038/s41586-019-1922-8



- › Rapidly growing evidence illustrates correlation between **TLS-derived antibodies** in the tumor microenvironment and **positive clinical outcomes**<sup>2</sup>
- › **TLS-derived antibodies** have been shown to be associated with apoptosis of cancer cells in patients<sup>2</sup>

## Reverse Immunology platform identifies the antigens targeted by endogenous antibodies produced in tumor lymphoid structures (TLS)



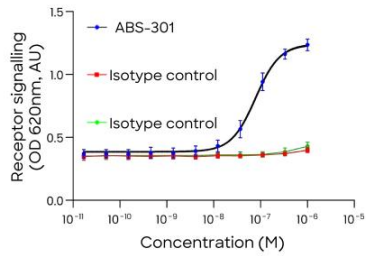
doi: 10.1101/2021.02.06.430058

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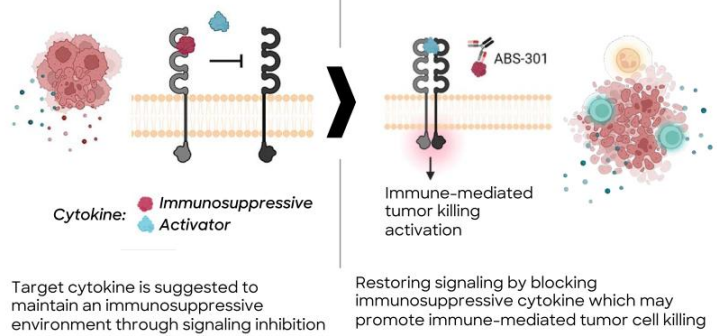
49

# A patient-derived antibody discovered by reverse immunology blocks an immunosuppressive cytokine

ABS-301 Rescues Pro-Inflammatory Signaling Through Inhibition of Immunosuppressive Cytokine



Target Biology and Proposed ABS-301 Mechanism of Action



doi: 10.1101/2021.02.06.430058

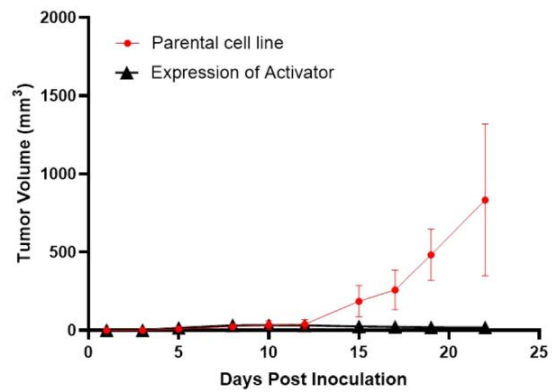
## Key Findings

- › Activation of the ABS-301-targeted pro-inflammatory pathway triggers a robust anti-tumor immune response.

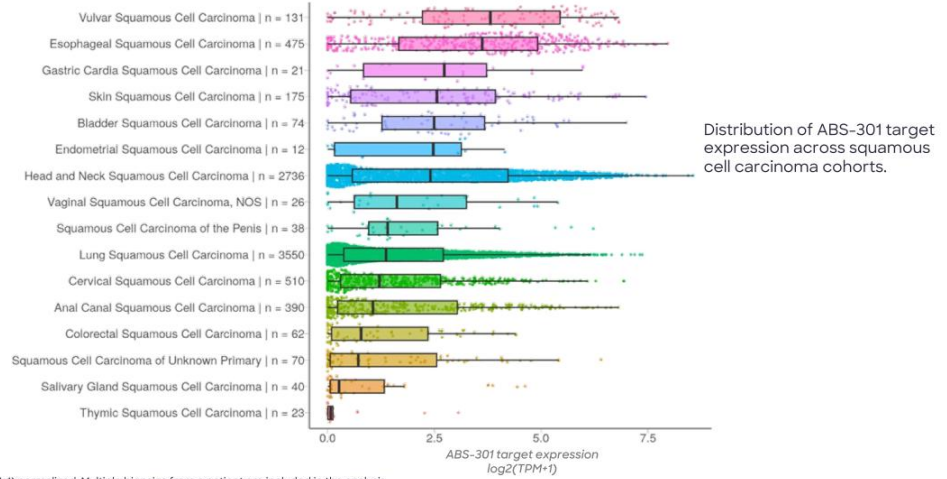
## Study Overview

- › Mouse melanoma cells were genetically modified to activate the ABS-301-targeted pro-inflammatory pathway via Activator expression.
- › Tumor progression was assessed in immunocompetent mice injected with either engineered cells or unmodified parental cells.

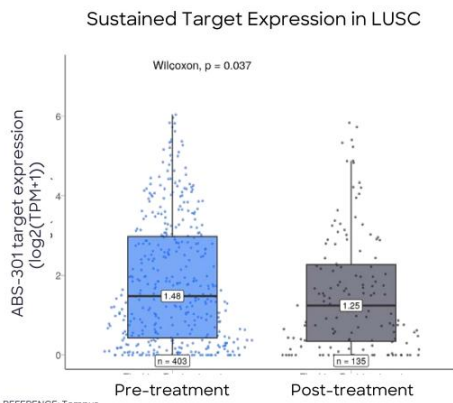
Tumor Growth Mouse Model



# Expression of ABS-301's target suggests broad potential in squamous cell carcinomas

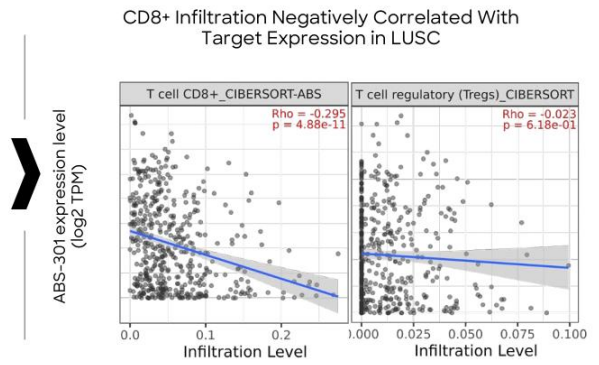


# Expression in Lung Squamous Cell Carcinoma (LUSC): no change with treatment and strong negative correlation with CD8+ T cell infiltration



REFERENCE: Tempus

In LUSC, univariate analysis of ABS-301 expression indicate only a minor change in expression between pre- and post-treatment suggesting opportunity for combination therapy.



ABS-301 target expression shows a strong negative correlation with CD8+ T cell infiltration with a minimal effect on Treg infiltration supporting immunosuppressive activity of target in vivo.

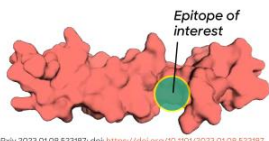
## Ongoing preclinical studies exploring broad application in immuno-oncology

Indication	US Prevalence	Estimated 5-year survival rate*	Projected US Sales in 2030
NSCLC	Calculated: ~202K in 2023	28%	\$27B
	SCC 30% of NSCLC cases Calculated: ~61K	24%	Calculated Sales: \$8.1B
Head and Neck SCC	~54K in 2022	68.5%	Calculated Sales: \$2.3B
Esophageal Cancer	~21K in 2022	20%	\$1.5B
	SCC ~20% of cases Calculated: ~4.2K		Calculated Sales: \$0.3B
Cervical Cancer	~14K in 2023		\$0.6B
	SCC 90% of cases Calculated: ~13K	67%	Calculated Sales: \$0.6B
Skin Cancer, non-melanoma	Incidence = ~3,300K	95-100%	\$1.0B
	SSC Incidence = ~700K	95%	Calculated Sales: \$0.2B

\*dependent on stage of diagnosis  
References provided in appendix

## Deploying de novo AI model on HER2 led to discovery of antibodies displaying molecular interactions distinct from trastuzumab

Zero shot de novo AI discovery on HER2



bioRxiv 2023.01.08.523187; doi: <https://doi.org/10.1101/2023.01.08.523187>

- › Hits with edit distance of up to 12 amino acids in HCDR3 region (13 aa, search space of 2013) were screened
- › Selected 50 hits with <10 nM affinity were expressed as mAbs for binding affinity determination
- › Top 11 antibodies were characterized in vitro and 3 leads evaluated in vivo

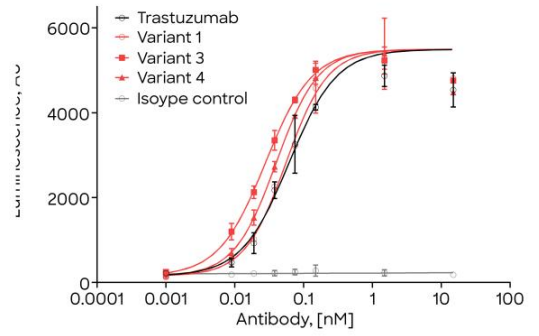
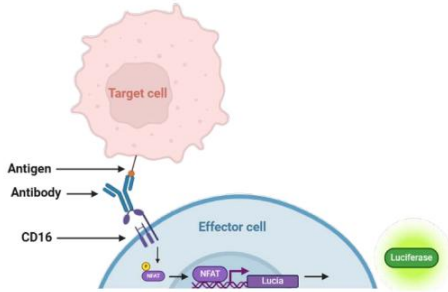
AI-designed antibodies: same epitope, different HER2 contact preferences

Variant #	Edit distance	K <sub>d</sub> (nM)	Epitope mapping view	Loop 581-590
Trastuzumab	0	1.07		
1	7	4.16		
3	7	9.75		
4	2	6.66		

■ Not critical  
■ Partial  
■ Critical

# AI-designed antibodies demonstrate measurable enhancement of ADCC activity compared to trastuzumab

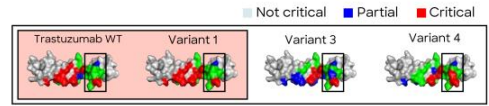
**ADCC Assay Principle**  
Luciferase signal driven by NFAT transcription factor positively correlates to ADCC activation against JIMT-1



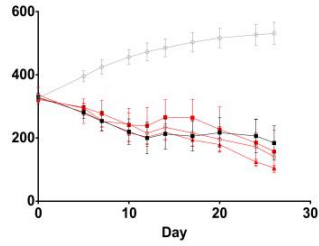
	Trastuzumab	Variant 1	Variant 3	Variant 4
EC50 (nM)	0.062	0.056	0.028	0.040
R squared	0.93	0.97	0.97	0.95
P value	N/A	Not significant	<0.0001	0.0015

\*Estimated performance of a putative clinical competition molecule generated for in-house comparison  
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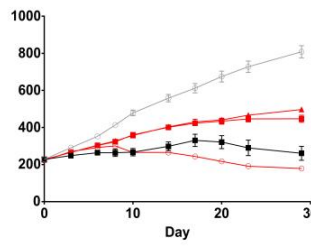
# AI-designed antibodies suppress growth of trastuzumab-sensitive & resistant HER2+ breast tumors



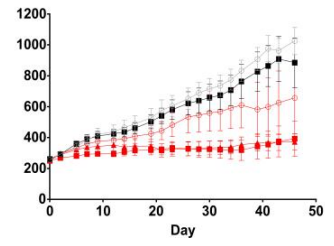
Mouse xenograft model using EFM192A (HER2+ BC; Tz sensitive)



Mouse xenograft model using MDA-MB-361 (HER2+ BC; Tz sensitive)



Mouse xenograft model using JIMT-1 (HER2-amp BC; Tz resistant)



Trastuzumab-sensitive EFM192A and MDA-MB-361 tumors respond to both trastuzumab (Tz) & AI-designed antibodies

JIMT-1 tumors are trastuzumab resistant but sensitive to variants 3 and 4

Xenograft studies conducted by Dr. Dennis Slamon's team at UCLA  
2-way ANOVA \*\*P<0.001 and \*\*\*P<0.0001 vs Isotype control

## AI-designed antibodies create opportunities to address unmet medical need

Currently exploring breast cancer as opportunity: alternative to or post Enhertu®

- › Despite Enhertu's good efficacy, leading oncologists are only moderately satisfied due to toxicity (e.g. interstitial lung disease)
- › Less toxic therapy and effective treatment post-Enhertu are key unmet needs.

*"Post-Enhertu is really where the action is right now in the field. I think the first company that comes up with something that has significant benefit in Enhertu progressive disease is going to win."*

- KOL

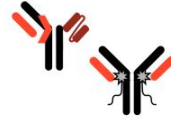
### Multiple paths possible for therapeutic development

Modality switch or combination opportunities under consideration to address unmet medical needs



- › Later-line treatment regimens for HER2-positive cancer:

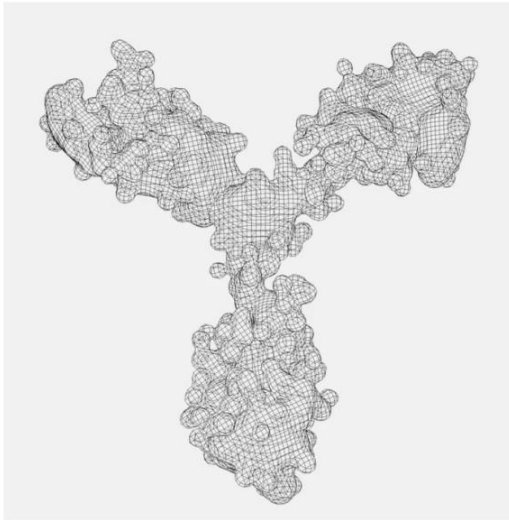
- Monotherapy
- Combination therapy with targeted small molecules



- › Enhancing efficacy and expanding indications (e.g. Enhertu resistance):

- Antibody-drug conjugates (ADCs)
- Multi-specific antibodies

## Leading AI models to create novel & differentiated therapeutics



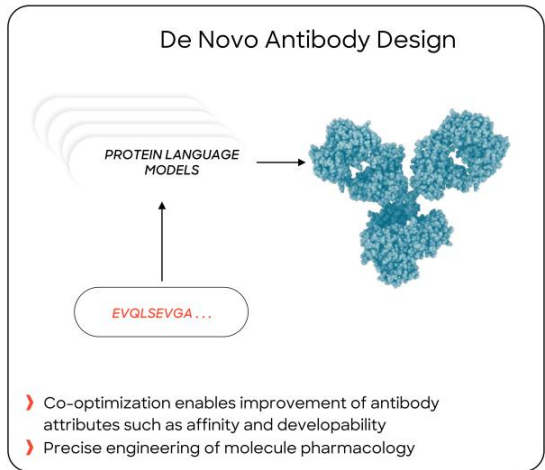
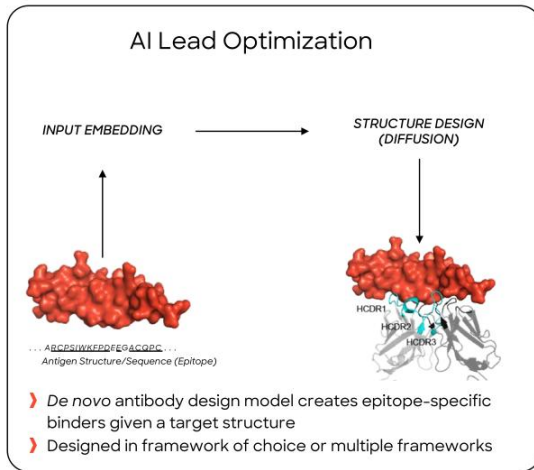
### > ADDRESS COMPLEX AND PREVIOUSLY "HARD TO DRUG" TARGETS

- | Bind Specific extracellular domains
- | Target Specific conformations
- | Address difficult target classes e.g. GPCRs

### > INTRODUCE PRECISE CONTROL OVER ANTIBODY DESIGN

- | "Smart" biologics
- | Enhanced Potency & MOA
- | Engineer selectivity, minimizing off target toxicity
- | Agonism vs. Antagonism

# Leadership in AI de novo design of antibody-based therapeutics

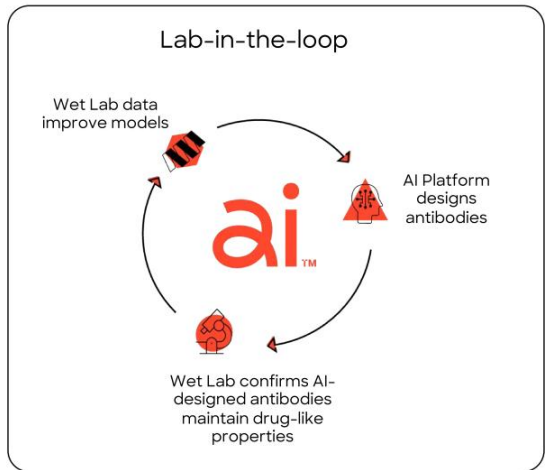



# Our AI platforms are enabled by our 6-week 'lab-in-the loop' active learning cycles

### AI Platforms

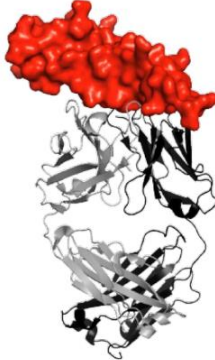
**DE NOVO ANTIBODY DESIGN**  
de novo design of epitope-specific antibodies against targets without requiring a known binder

**LEAD OPTIMIZATION**  
AI guided lead optimization enables tunable pharmacology



# AbSciDesign comprises two categories of AI models for *de novo* antibody design

**AbSciGen**  
*antibody-<=>antigen complex structure and sequence design*



**Design 1**  
HCDR1: GFNIKDTY  
HCDR2: IYPTNGYT  
HCDR3: SRWGGDGFYAMDY

LCDR1: QDVNTA  
LCDR2: SAS  
LCDR3: QQHYTTPPT

•

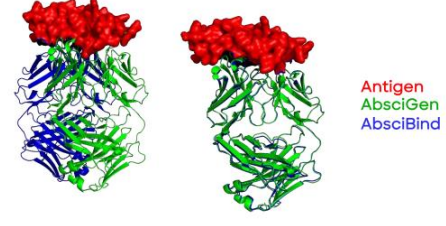
•

•

**Design N**  
HCDR1: GFNIKDTW  
HCDR2: IYPSNGYT  
HCDR3: ARWGGDGFYAMDY

LCDR1: QDVNTA  
LCDR2: SAS  
LCDR3: QQHYTTPPT

**AbSciBind**  
*Antibody design scoring and filtering*



Antigen  
AbSciGen  
AbSciBind

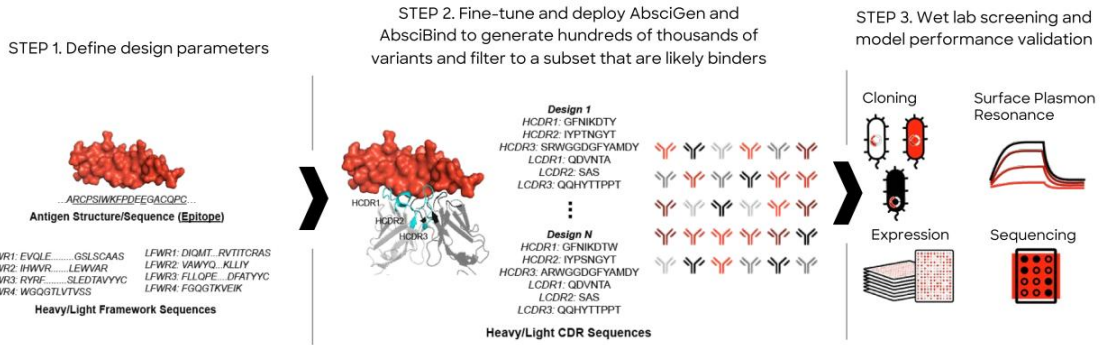
**AbSciBind**  
Low Rank

RMSD = 5.3 Å  
Confidence = 0.64

**AbSciBind**  
High Rank

RMSD = 2.3 Å  
Confidence = 0.95

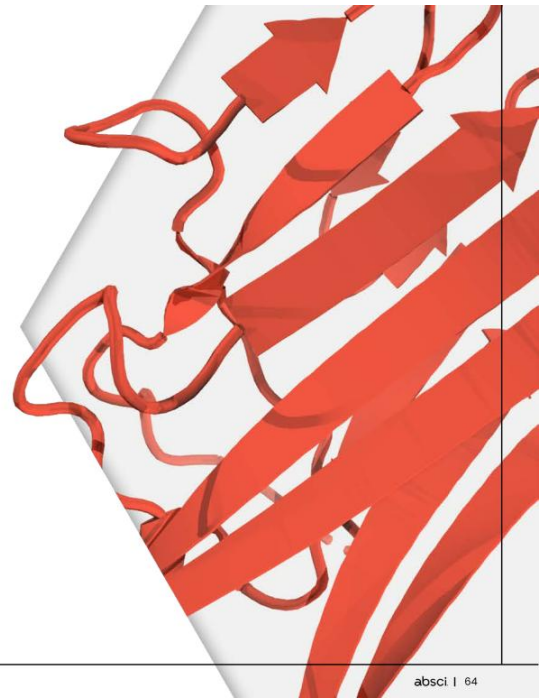
# The AbsciDesign AI platform delivers de novo antibodies via an end-to-end design-validation workflow



**CASE STUDY**

*de novo* design of an antibody that binds the Caldera region of HIV-1 trimer

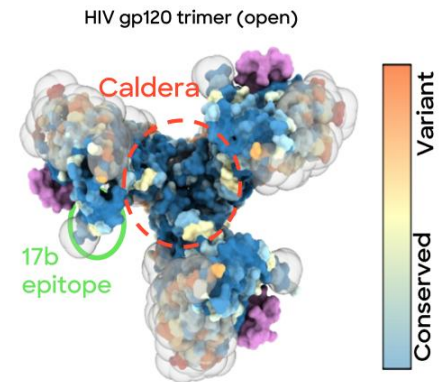
**Caltech** **absci.** BILL & MELINDA  
GATES foundation



**DE NOVO DESIGN**

**de novo design antibody that binds to the highly conserved caldera region of HIV gp120**

- › No natural or synthetic antibody for HIV exists today because immune system cannot derive an antibody that is universally neutralizing against HIV
- › Design challenge: create universally neutralizing HIV antibody by binding unique and conserved epitope within “caldera” of open conformation of gp120 to prevent HIV from entering host cells
- › Numerous attempts to target this epitope have failed—previous efforts have identified antibodies, but none bind the “caldera” and none are universally neutralizing.



**HIV Env Trimer Challenge :**

- Highly glycosylated
- Extremely high sequence diversity among isolates
- High mutation rate at common neutralizing epitopes

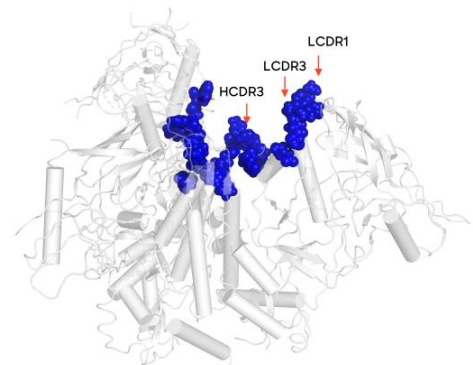
**Model inputs:**

1. Antigen structure
2. Framework of 17b
3. Epitope selected conserved across HIV strains (Clades A, B, and C)

**Design of CDRs:**

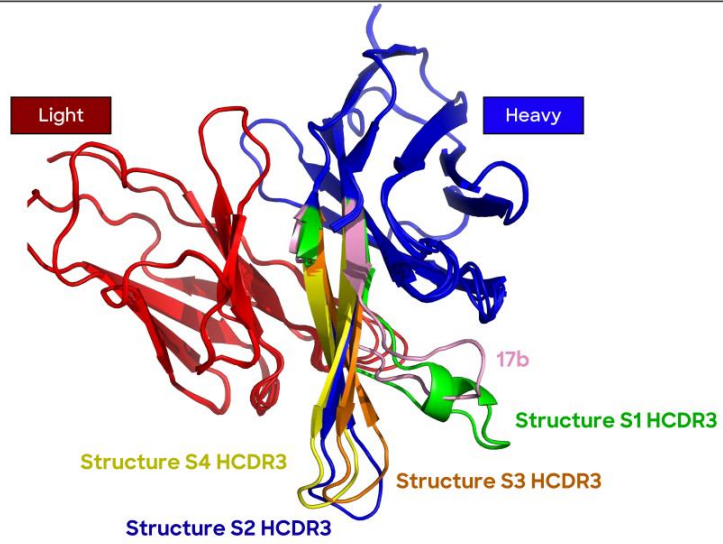
- Condition the model to design long HCDR3s to reach into open caldera region (>20 residues)
- Designed HCDR2 and LCDR3 to bind to HIV surface

HIV Env trimer (open)

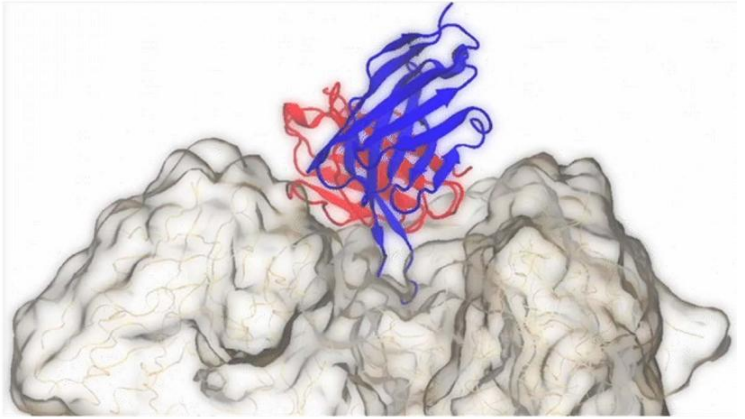


**DE NOVO DESIGN**

**4 best structures selected from 10,000+ structures generated by *de novo* model**

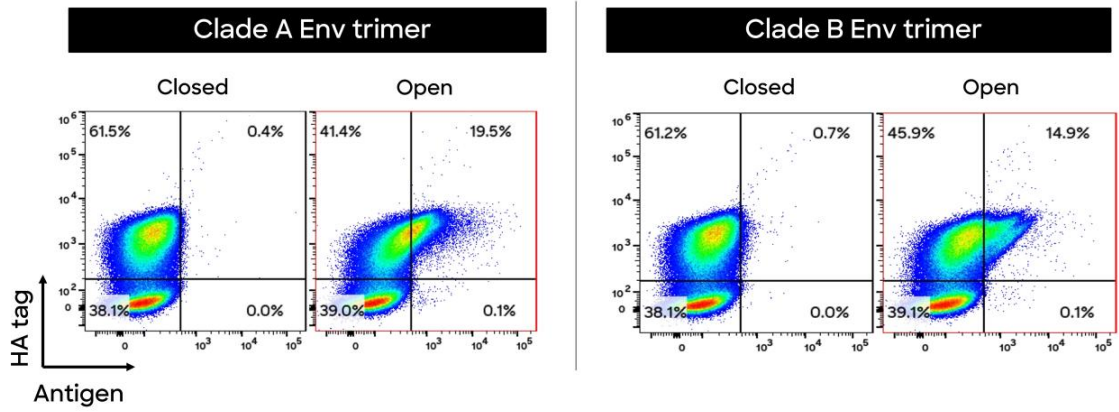


Applied molecular  
dynamics simulation  
to *de novo* designed  
antibodies



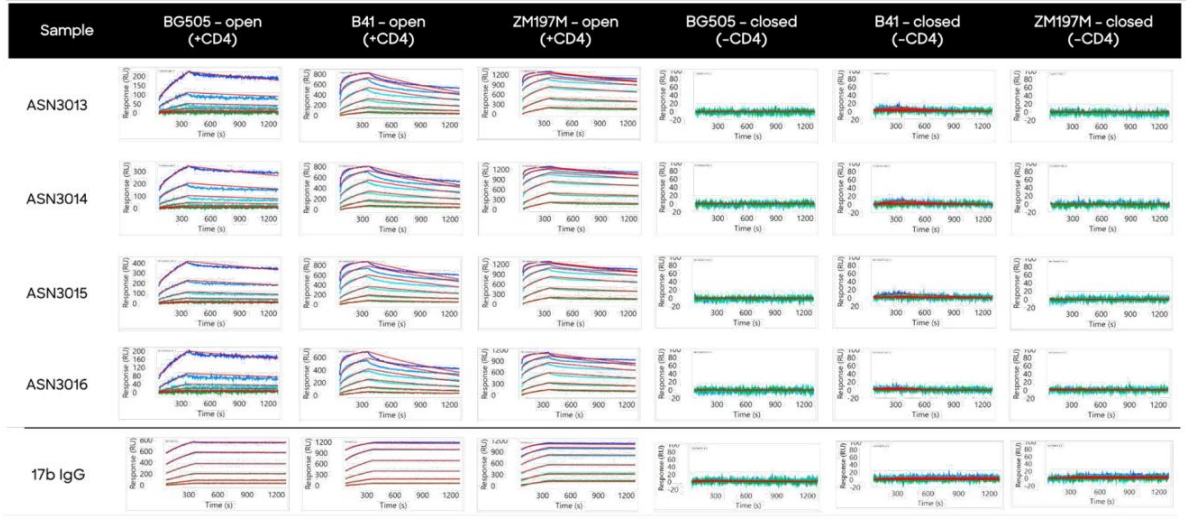
**DE NOVO DESIGN**

Enriched de novo library binds open, not closed, Env trimer conformation in YSD



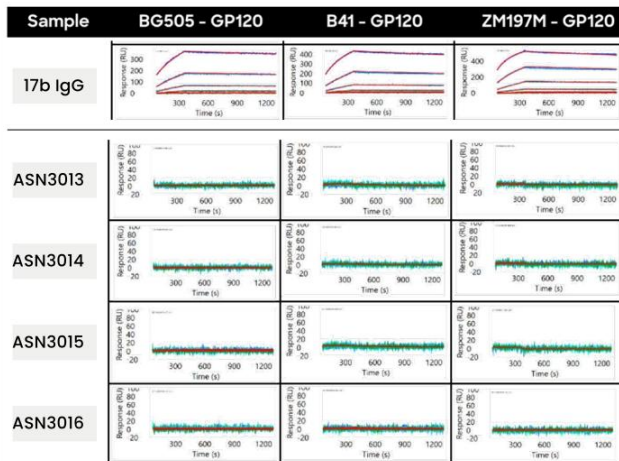
**DE NOVO DESIGN**

**SPR data demonstrate binding characteristics consistent with binding of caldera**



**DE NOVO DESIGN**

**HIV-Caldera: SPR demonstrates no binding of *de novo* designs to GP120 monomer**



**Hypothesis:**

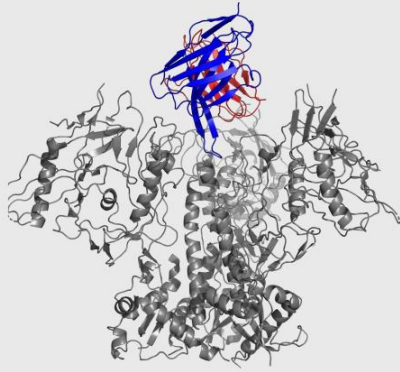
If the designed mAbs are binding to the caldera region we should not observe binding to monomeric GP120 since the caldera is only present in the Env trimer

**Key results:**

- ✓ 17b showed high affinity binding to monomeric GP120 as expected
- ✓ Absci mAbs showed no binding to monomeric GP120, suggesting these binders are targeting an epitope that is only present in the Env trimer

## HIV DE NOVO DESIGN

### HIV-Caldera: demonstrating AI de novo design for challenging target



#### SUMMARY

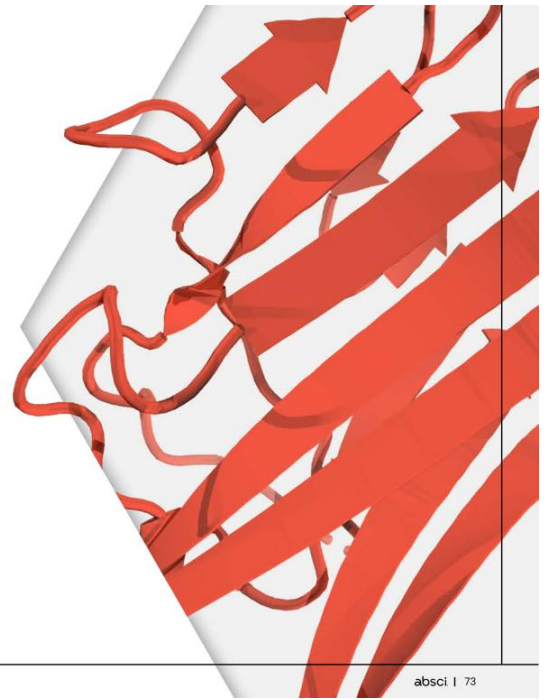
- › *de novo* design model created a novel and diverse antibody which binds multiple clades of HIV indicating successful targeting of the caldera epitope
- › Screening cascade enabled selection of differentially binding variants

#### NEXT STEPS

- › Binders from this study will be selected for affinity maturation
- › Structure of *de novo* binder and epitope specificity will be experimentally solved to confirm fidelity with designed structure and targeted epitope

## CASE STUDY

AI Optimization for pH sensitivity



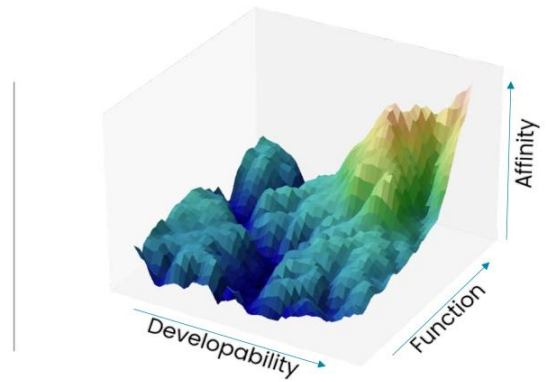
## AI lead optimization platform for 'smart biologics'

### The Challenge

The diversity of antibodies is vast, making it impossible for traditional methods to explore effectively.

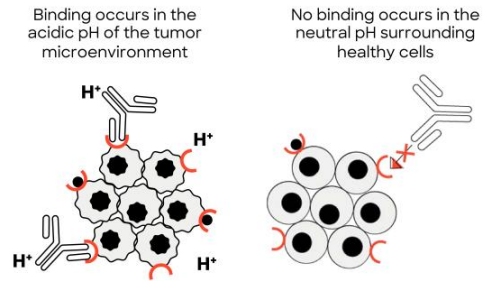
### Absci Solution

Our AI can search a space of  $\sim 10^{19}$ , a million times larger than traditional methods, identifying functional, developable antibodies in one step.

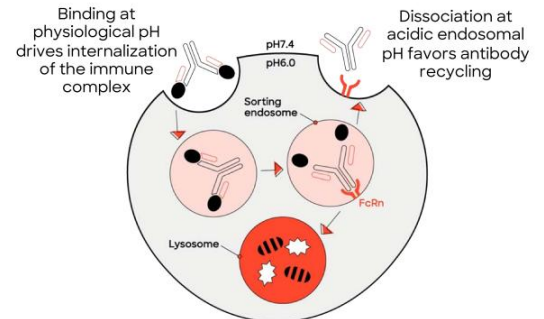


## pH sensitivity may reduce toxicity and/or improve efficacy of therapeutic mAbs

Tumor specificity improves efficacy and reduces "on-target off-tumor" toxicities

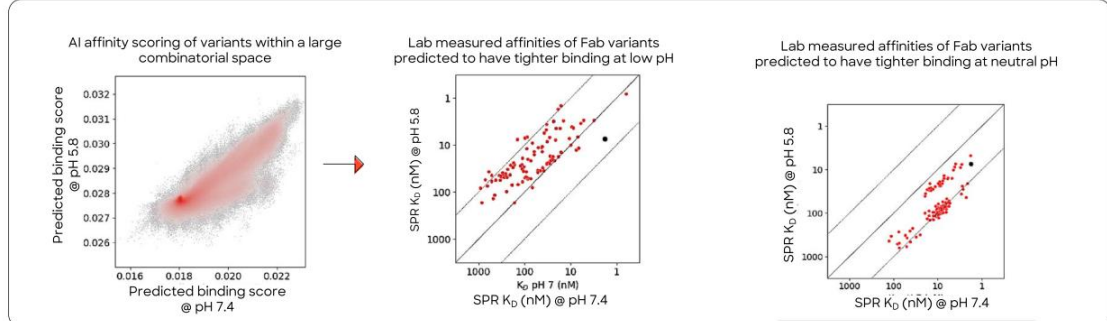


Disassociation in the endosome drives antibody recycling and efficient clearance of soluble targets



## Models identify pH sensitive Fab variants from the same lead for either indication

1. Library for model training sampled 60 positions on heavy chain framework and CDRs with up to 7 substitutions biased for ionizable residues (H, K, R, D, E)
2. Library screened for antigen binding at pH 7.4 and pH 5.8
3. Model trained and used to generate antibodies with tuned pH dependency

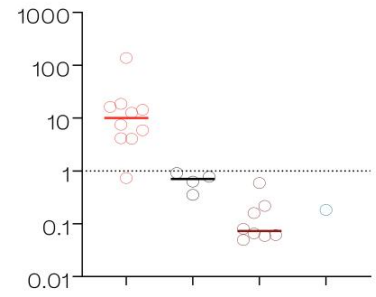


## Hits reformatted as mAbs show desired binding profiles

### SUMMARY

- › AI optimized leads achieves variants with pH sensitive binding up to 100x differential
- › pH-sensitive leads had no liabilities for stability, aggregation and polyreactivity<sup>1</sup>
- › Model proposed mutations use all 6 ionizing residues in heavy chain CDRs and framework region
- › Sequences were proposed from a >10<sup>13</sup> combinatorial space

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Modeling Strategy

## Summarized platform case studies

### De Novo Design

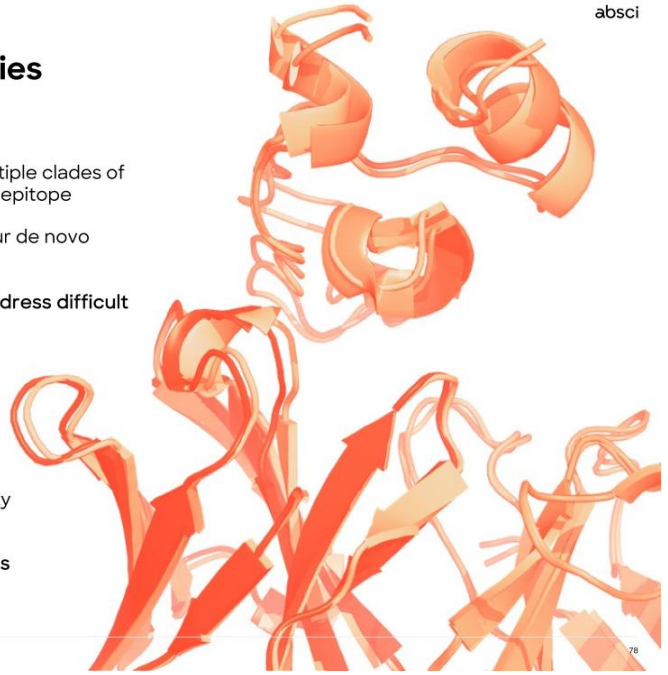
- › de novo design model created molecule binds multiple clades of HIV suggesting successful targeting of the caldera epitope
- ›
- › Represents second disclosed target success for our de novo platform in the 2nd half of this year

**Absci's de novo design platform can successfully address difficult to drug target epitopes**

### AI Optimization

- › Models identify unseen variants with 10x-20x pI sensitivity in both directions, and up to 100x differential compared to parental molecule after only one round
- › Designed leads had no liabilities indicating the ability to successfully search a fitness landscape

**Absci's lead optimization platform enables molecules with differentiated pharmacology**



# Generative AI Re(Generative) Biology

absci.

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