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

FORM 8-K

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

Date of Report (Date of earliest event reported): December 15, 2020

PIERIS PHARMACEUTICALS, INC.
(Exact name of registrant as specified in its charter)

Nevada	**001-37471**	**30-0784346**
(State or other jurisdiction of Incorporation)	**(Commission File Number)**	**(IRS Employer Identification No.)**

225 State Street, 9th Floor	**02109**
Boston, MA	
(Address of principal executive offices)	**(Zip Code)**

Registrant's telephone number, including area code: 857-246-8998
N/A
(Former name or former address, if changed since last report.)

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:

Title of each class	Trading Symbol(s)	Name of each exchange on which registered
Common Stock, $0.001 par value per share	**PIRS**	**The Nasdaq Capital Market**

Indicate by check mark whether the registrant is an emerging growth company as defined in Rule 405 of the Securities Act of 1933 (17 CFR §230.405) or Rule 12b-2 of the Securities Exchange Act of 1934 (17 CFR §240.12b-2).

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. ☐

Item 7.01 Regulation FD Disclosure.

Attached hereto as Exhibit 99.1 and incorporated by reference herein is the December 2020 Investor Presentation of Pieris Pharmaceuticals, Inc.

The information set forth under this "Item 7.01. Regulation FD Disclosure," including Exhibit 99.1 attached hereto, shall not be deemed "filed" for any purpose, and shall not be deemed incorporated by reference into any filing under the Securities Act of 1933, as amended, or the Securities Exchange Act of 1934, as amended, regardless of any general incorporation language in any such filing. except as shall be expressly set forth by specific reference in such filing.

Item 9.01 Financial Statements and Exhibits

(d) *Exhibits.*

99.1 Investor Presentation, Dated December 2020.

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.

PIERIS PHARMACEUTICALS, INC.

Dated: December 15, 2020

/s/ Tom Bures
Tom Bures
Vice President, Finance

INVESTOR PRESENTATION

DECEMBER 2020

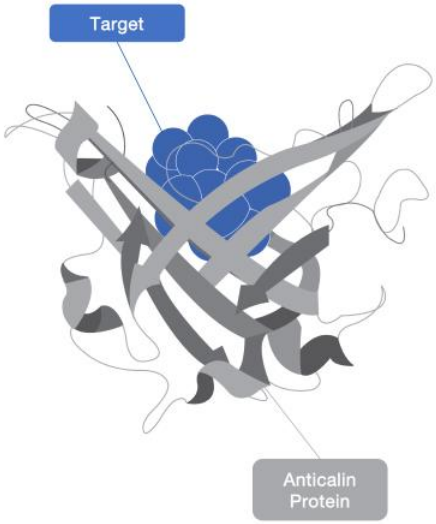
Forward Looking Statements

This presentation contains forward-looking statements as that term is defined in Section 27A of the Securities Act of 1933 and Section 21E Securities Exchange Act of 1934. Statements in this presentation that are not purely historical are forward-looking statements. Such forward- statements include, among other things, the timing for, and outcome of, the additional in-use and compatibility study for PRS-343 as requested FDA; whether data from patients enrolled to date will be sufficient to inform the recommended phase 2 dose for the Company's planned p concept study of PRS-343 in gastric cancer; the expected timing and potential outcomes of the reporting by the Company of key clinical data programs, references to novel technologies and methods and our business and product development plans, including the advancement proprietary and codevelopment programs into and through the clinic and the expected timing for reporting data, making IND filings or achievir milestones related to our programs, including PRS-060/AZD1402, PRS-343, PRS-344, and PRS-352 and the expected timing of the initiation of t stage of PRS-343's development in gastric cancer. Actual results could differ from those projected in any forward-looking statements due to nu factors. Such factors include, among others, our ability to raise the additional funding we will need to continue to pursue our business and development plans; the inherent uncertainties associated with developing new products or technologies and operating as a developmer company; our ability to develop, complete clinical trials for, obtain approvals for and commercialize any of our product candidates, including ou to recruit and enroll patients in our studies; our ability to address the requests of the FDA, including with respect to the additional in-u compatibility study for PRS-343, and the resolution of the partial clinical hold relating to that drug candidate; competition in the industry in w operate; delays or disruptions due to COVID-19; and market conditions. These forward-looking statements are made as of the date of this prese and we assume no obligation to update the forward-looking statements, or to update the reasons why actual results could differ from those proj the forward-looking statements, except as required by law. Investors should consult all of the information set forth herein and should also refe risk factor disclosure set forth in the reports and other documents we file with the SEC available at www.sec.gov, including without limita Company's Annual Report on Form 10-K for the fiscal year ended December 31, 2019 and the Company's Quarterly Reports on Form 10-Q.

The Anticalin® Protein Platform

A Novel Therapeutic Class with Favorable Drug-Like Properties

- **Human** – Derived from lipocalins (human extracellular binding proteins)
- **Small** – Monomeric, monovalent, small size (~18 kDa vs 150kDa mAbs)
- **Stable** – Inhalable delivery
- **Simple** – Bi/multispecific constructs
- **Proprietary** – Broad IP position on platform and derived products



Powerful Drug Discovery Pla

- Highly diverse libraries
- Automated screening
- Protein engineering know-how

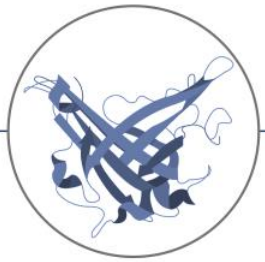
Translational Science Expert Deploy Platform in Meaningf

- Immunology expertise underpi and respiratory focus
- A leader in 4-1BB-related effor
- Patient phenotyping efforts for improved stratification and nov intervention points in, e.g., asth

Company Snapshot

Pipeline Highlights

- **PRS-060:** Inhaled IL4-Rα antagonist for moderate-to-severe asthma (partnered with AstraZeneca)
- **PRS-343:** 4-1BB/HER2 bispecific for solid tumors
- **Next-generation respiratory:** Includes 4 discovery-stage inhaled therapeutics programs (2 proprietary, 2 partnered with AstraZeneca)
- **4-1BB-based bispecifics:** Multiple proprietary and partnered 4-1BB-based programs for IO



Anchor Partnerships

- Validation through three anchor partnerships
- $120+M in upfront payments and milestones since January 2017
- Each partnership includes options for co-development & US-focused commercialization rights
- Value-driving opt-in for PRS-060 after phase 2a completion

Catalysts

- **Respiratory:**
  - ✓ PRS-060 phase 2a trial initiation
  - ❑ Data and rationale for advancement i IND-enabling studies for wholly-owne inhaled program
- **IO:**
  - ✓ PRS-343 complete monotherapy and combination with atezolizumab phase escalation data at ESMO
  - ❑ PRS-343 initiation of 2nd line HER2+ gastric cancer PoC study, additive to
  - ❑ PRS-344 IND

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Improving Lives

Partnerships

AstraZeneca

- PRS-060 + 4 additional novel inhaled Anticalin protein programs
- Retained co-development and co-commercialization (US) options on PRS-060 and up to 2 additional programs
- $57.5M upfront & 2017 milestone
- ~$2.1B in milestone potential, plus double-digit royalties
- AZ funds all PRS-060 development costs through post-phase 2a co-development opt-in decision
- Access to complementary formulation and device know-how for inhaled delivery

SERVIER

- Immuno-oncology partnership based on antibody-Anticalin bispecific protein fusions
- PRS-344: PD-L1/4-1BB antibody-Anticalin bispecific
  - ✓ Pieris opted in for full U.S. rights
- PRS-352: n.d. antibody-Anticalin bispecific
  - Pieris planning handover to Servier in 2020
  - Pieris to receive royalties
- ~$31M upfront payment with significant milestone potential
  - ✓ Two preclinical milestones achieved for PRS-344

SeattleGenetics

- 3-program partnership based on tum localized costimulatory bispecific fus proteins
- Pieris retains opt-in rights for 50/50 g profit split and U.S. commercializatio rights on one of the programs
- $30M upfront payment, ~$1.2B miles potential
  - ✓ Achieved $5M milestone paym for first program, a bispecific t targeted costimulatory agonist
- Up to double-digit royalties on non-c developed products

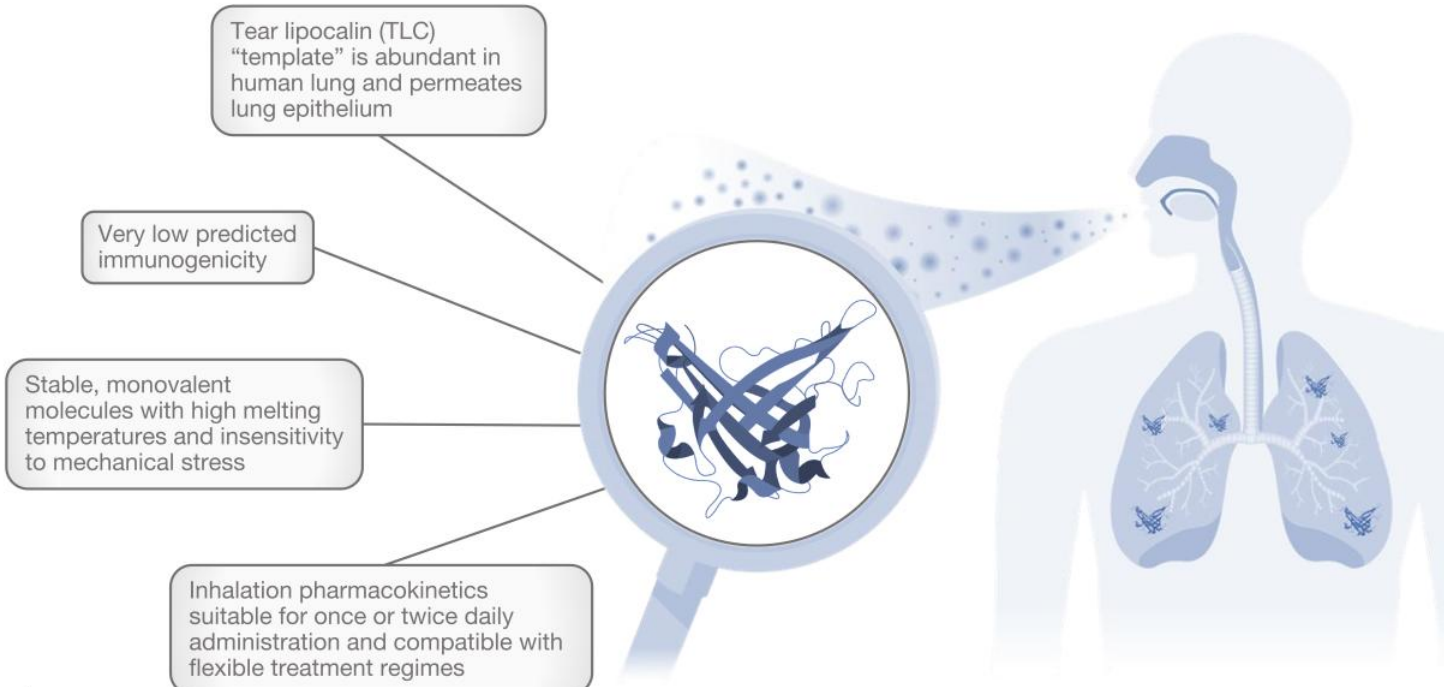
Strong Partners • Significant Cash Flow • Retained Commercial Rights

Pipeline

RESPIRATORY							
CANDIDATE	TARGETS	PARTNER	COMMERCIAL RIGHTS	DISCOVERY	PRECLINICAL	PHASE I	PH
PRS-060/AZD1402	IL4-Rα	AstraZeneca	Pieris Worldwide Profit-Share Option				
AstraZeneca Programs*	n.d.	AstraZeneca	Pieris Worldwide Profit-Share Option*				
Proprietary Programs	n.d.	n/a	Pieris Worldwide				
*4 additional respiratory programs (3 active, 1 forthcoming) in collaboration with AstraZeneca, 2 of which carry co-development and co-commercialization options for Pieri							

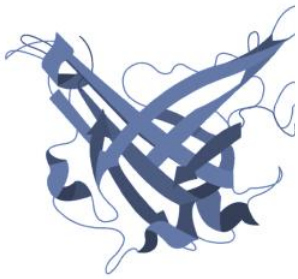
IMMUNO-ONCOLOGY							
CANDIDATE	TARGETS	PARTNER	COMMERCIAL RIGHTS	DISCOVERY	PRECLINICAL	PHASE I	PH
PRS-343	HER2/4-1BB	n/a	Pieris Worldwide				
	+ Anti-PD-L1	n/a					
PRS-344	PD-L1/4-1BB	SERVIER	Pieris U.S. Rights				
PRS-352	n.d.	SERVIER	SERVIER				
Proprietary IO Programs	n.d.	n/a	Pieris Worldwide				
Seattle Genetics Program‡	co-stim agonist	SeattleGenetics	Pieris U.S. Option‡				
‡3 bispecific programs (1 active, 2 forthcoming) in collaboration with Seattle Genetics, with Pieris retaining US rights for 1 program							

Anticalin Technology Advantages: Differentiated Respiratory Platform



PRS-060: IL-4Rα Antagonist

Candidate	PRS-060
Function/MoA	Inhibiting IL4-Rα (disrupts IL-4 & IL-13 signaling)
Indications	Moderate-to-severe asthma
Development	Phase 2a in moderate asthmatics
Commercial Rights	Co-development and U.S. co-commercialization rights, including gross margin share



PRS-060

PRS-060 Phase I Multiple Ascending Dose Trial

Strategic Objectives	Ascertain PK/PD with a reliable biomarker to confirm local target engagemer and inform Phase II dosage regimen
Trial Design Highlights	Dosing patients with mild asthma with elevated FeNO levels (≥35 ppb), to rec inhaled PRS-060 or pbo b.i.d.* over a 10-day period

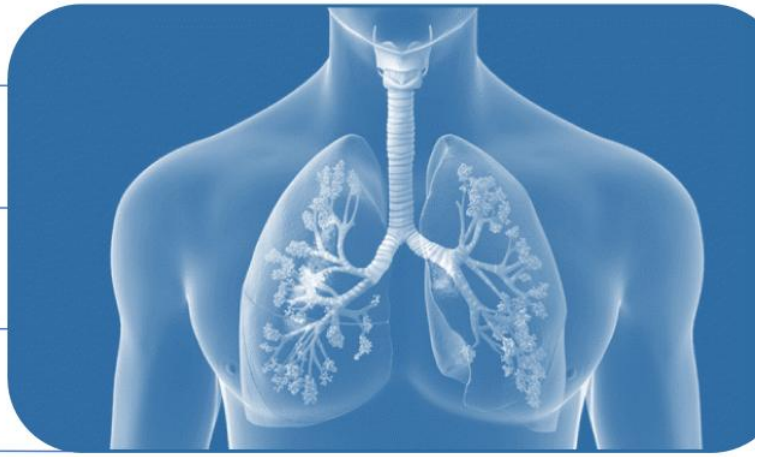
*q.d. or

Initiated in July 2018

Evaluating safety, tolerability, PK, PD and will also evaluate FeNO reduction vs. placebo

Measuring safety, tolerability and FeNO changes days 1-10,17, and 40

Pieris is sponsoring the trial, AstraZeneca is reimbursing Pieris for all associated costs



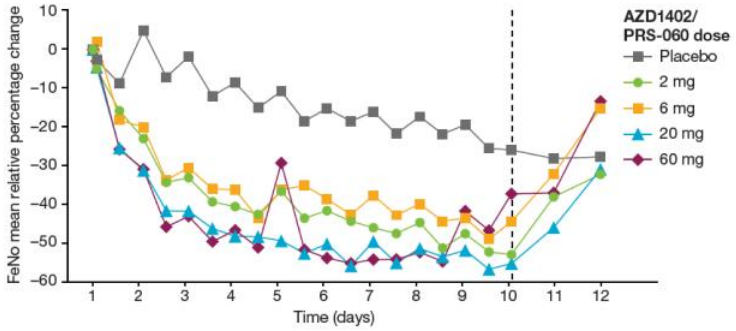
Phase 1b Interim Results: Favorable Safety Profile

- All doses of AZD1402/PRS-060 tested in the study were well tolerated
- No treatment-related serious AEs were observed

System organ class AE Preferred Terms[b]	Placebo (N = 12) n (%) m	AZD1402/PRS-060[c] (N = 30) n (%) m	Overall (N = 42) n (%) m
Gastrointestinal disorders	4 (33.3) 4	13 (43.4) 14	17 (40.5) 18
Dry mouth	1 (8.3) 1	2 (6.7) 2	3 (7.1) 3
Nausea	1 (8.3) 1	3 (10.0) 3	4 (9.5) 4
Infections and infestations	1 (8.3) 1	7 (23.3) 8	8 (19.0) 9
Upper respiratory tract infection	1 (8.3) 1	3 (10.0) 4	4 (9.5) 5
Nervous system disorders	5 (41.7) 9	13 (43.4) 18	18 (42.9) 27
Headache	3 (25.0) 6	5 (16.7) 7	8 (19.0) 13
Presyncope	0	4 (13.3) 6	4 (9.5) 6
Respiratory, thoracic and mediastinal disorders	6 (50.0) 6	14 (46.7) 15	20 (47.6) 21
Cough	1 (8.3) 1	4 (13.3) 4	5 (11.9) 5
Rhinorrhoea	2 (16.7) 2	1 (3.3) 1	3 (7.1) 3
Wheezing	2 (16.7) 2	4 (13.3) 5	6 (14.3) 7

Phase 1b Interim Results: Robust FeNO Reduction

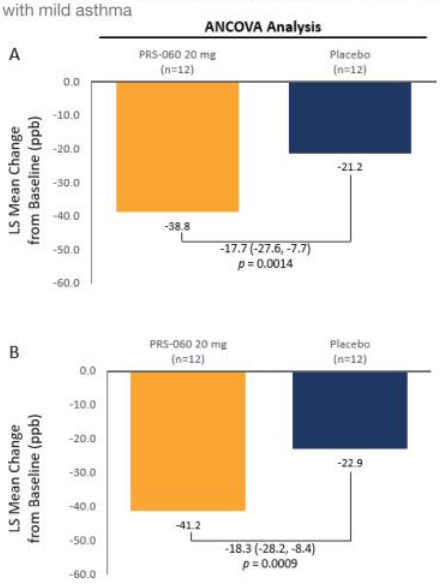
PRS-060 Relative FeNO Reduction (Emax Analysis)



PRS-060, mg (delivered)	n	Reduction vs. placebo, % (95% CI)	p-value
2	6	24.0 (1.8–41)	0.04
6	6	24.3 (2.7–41)	0.03
20	12	36.4 (22–48)	<0.0001
60	6	30.5 (10–46)	0.005
Placebo	12		

PRS-060 Relative FeNO Reduction (ANCOVA A

Mean change from baseline in FeNO levels at 0.5h (A) and 2h (B) post-dose on participants with mild asthma



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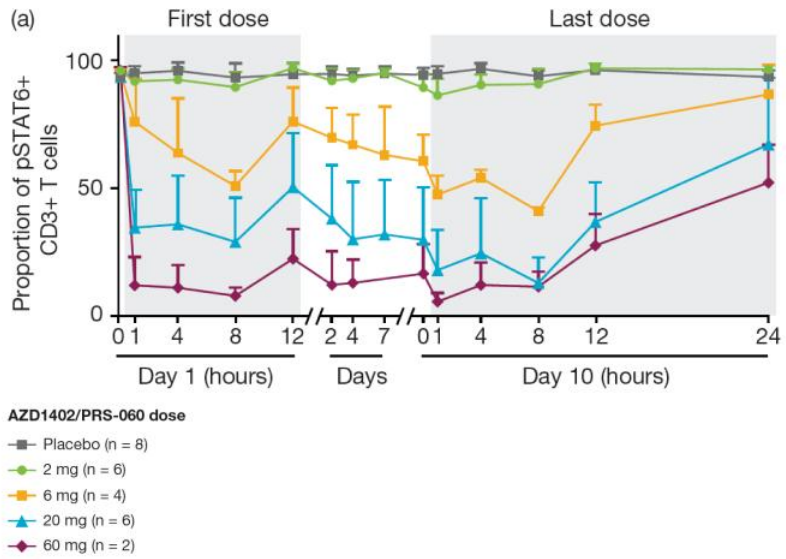
Improving Lives

Phase 1b Interim Results: Pharmacological Versatility

pSTAT6 levels over time following inhalation of PRS-060



No systemic target engagement minimal systemic exposure observed at the 2mg dose, sugges that local target engagement by drug is sufficient to reduce air inflammation

Pharmacological versatility, given dose FeNO reduction with no obse systemic activity (pSTAT6) versus high-dose FeNO reduction systemic activity

PRS-060 Phase 2a Trial

Part 1	**Patient Population:** Moderate controlled asthmatics **Primary Endpoint:** Safety and tolerability **Number of Patients:** ~45
Part 2	**Patient Population:** Moderate uncontrolled asthmatics with blood eosinophil coun ≥ 150 cells/µL and FeNO ≥ 25 ppb at screening **Primary Endpoint:** Improvement of FEV1 over four weeks relative to placebo **Number of Patients:** ~360

- First patient dosed expected 1Q2021
- Dry powder formulation, administered b.i.d. over four weeks
- Up to three dose levels plus placebo
- Study is sponsored and funded by AstraZeneca

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Improving Lives

4-1BB Agonism Offers Promise of Material Clinical Benefit

Unique Attributes of 4-1BB Agonism on Tumor-specific T Cells

- ✓ Increases T-cell proliferation to bolster immune repertoire
- ✓ Enhances cytotoxicity for tumor killing
- ✓ Improves metabolic fitness for increased survival of T cells
- ✓ Drives T-cell memory phenotype for increased durability

Pieris' 4-1BB bispecifics recognize that 4-1BB agonists have proven clinica
potency, yet must be kept out of the liver to ensure suitable therapeutic inde

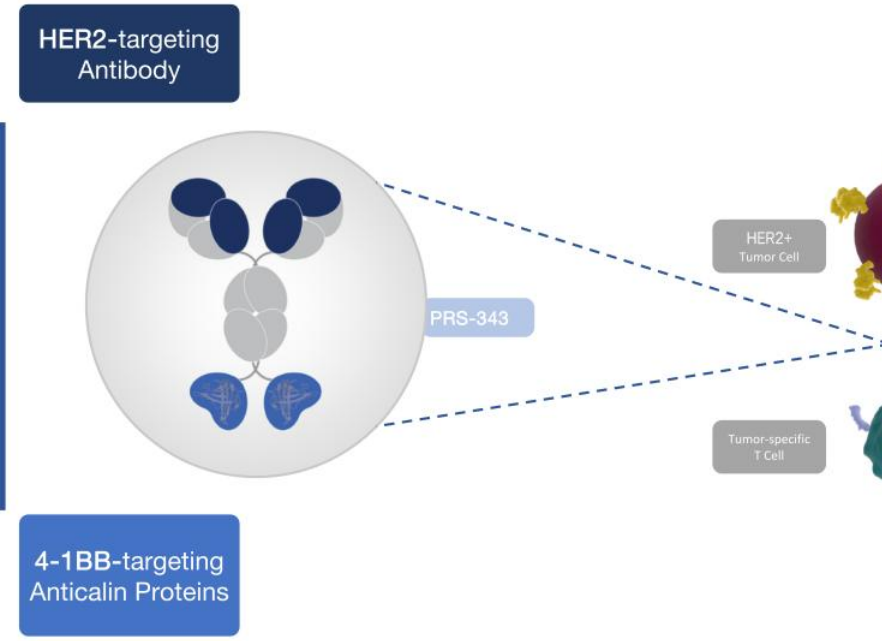
PRS-343: Proprietary Lead IO Asset

Candidate	PRS-343
Function/MoA	Tumor-targeted 4-1BB agonism and HER2 antagonism
Indications	HER2+ solid tumors
Development	Initiating phase 2 in combination with ramucirumab and paclitaxel in second line gastric
Commercial Rights	Fully proprietary

PRS-343 Localizes 4-1BB Agonism Within HER2+ Tumors

HER2-targeting moiety of the drug localizes to the tumor microenvironment and facilitates 4-1BB cross-linking

4-1BB cross-linking ameliorates T-cell exhaustion and is critical for T-cell expansion



Monotherapy and Combination Studies: Enabled Meaningful Clinical Characterization of PRS-343

Snapshot

- Patients with HER2+ solid tumors
- Monotherapy and combination with atezolizumab
- Data updates presented at ESMO 2020

Primary Objectives

- Characterize safety profile
- Identify MTD or RP2D

Secondary Objectives

- Characterize PK profile
- Investigate dosing schedule
- Assess potential immunogenicity and PD effects
- Investigate efficacy

ACTIVE SCHEDULES

Schedule 1: Q3W dosing on day 1; 21-day cycle

Schedule 2 (b): Q2W dosing on days 1, 15; 28-day cycle

Schedule 3 (c): Q1W dosing on days 1, 8, 15; 21-day cycle

In combination with atezolizumab: Q3W dosing on day 1; 21-day cycle

Mono Dose Cohort*	Combo Dose Cohort**	Do (mg
1		0.0
2		0.0
3		0.0
4		0.0
5	1	0.(
6	2	0.1
7	3	0.
8	4	1
9	5	2.
10	6	5
11	7	8
11 (b)		8
11 (c)		8
12 (b)		12
13 (b)		18
Obinutuzumab + 11(b)		8

9-13b: active dose cohorts in mono study

*Additional dose cohorts enrolling in monotherapy study

**1200mg flat dose of atezolizumab

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Improving Lives

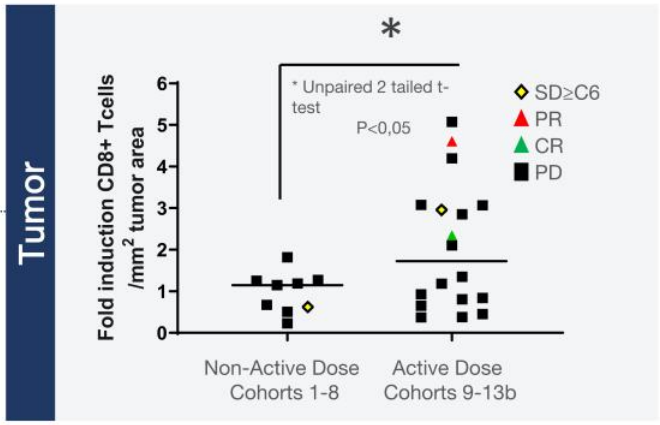
Summary of Responses of PRS-343 in Monotherapy

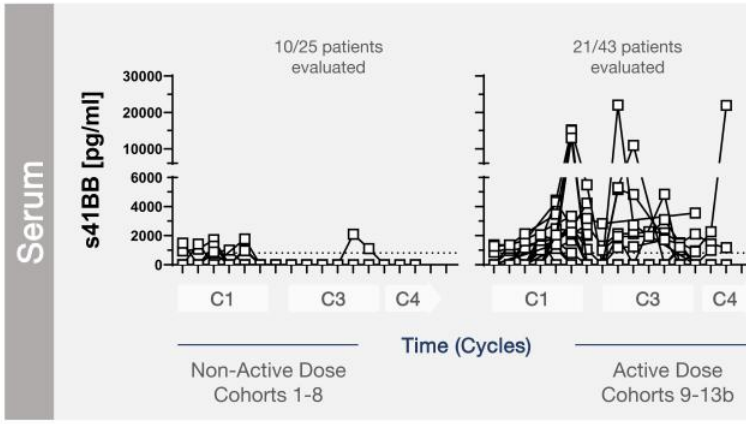
Based on clinical data, serum concentration of > 20 µg/ml defines active dose range (beginning at Cohort

Cohort	13b	12b	11c	Obi	11b	11	10	9	T
Best Response	18 mg/kg, Q2W	12 mg/kg, Q2W	8 mg/kg, QW	8 mg/Kg, Q2W	8 mg/kg, Q2W	8 mg/kg, Q3W	5 mg/kg, Q3W	2.5 mg/kg, Q3W	
Evaluable Patients	3	2	4	2	7	4	6	5	
CR	1	-	-		-	-	-	-	
PR	-	-	-		3	-	-	-	
SD	-	-	1	1	3	3	3	2	
ORR	33%	0%	0%	0%	43%	0%	0%	0%	1
DCR	33%	0%	25%	50%	86%	75%	50%	40%	5

Data cut-off: 27-Jul-20

Increase in CD8+ T Cells and Circulating Soluble 4-1BB Support 4-1BB Engagement by PRS-343





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Phase 2 PoC Study in 2nd Line Gastric Cancer

Phase 2 Initiation

PRS-343 in combination with ramucirumab and paclitaxel for 2nd-line HER2+ gastric cancer

Clinical trial collaboration with Eli Lilly; Lilly to supply ramucirumab

Single-arm, up to 60 patients

Primary endpoints: ORR and DCR

HER2+ subgroup from RAINBOW trial as comparator enriched w/ RWD

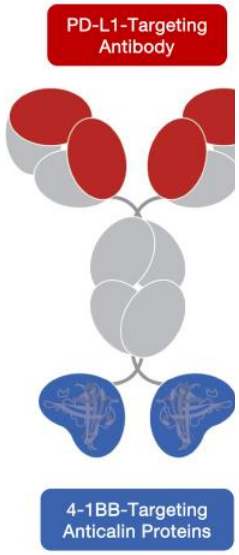
GC 2L PIVOTAL TRIAL

PRS-343 PoC Trial Considers Several Value-driving Elements

Factor	Impact
Biology: **Synergistic MoA in IO-amenable Patients**	• Vasculature normalization from ramucirumab for improved environment for T-cell infiltration • Tumor debulking and antigen release from paclitaxel for improved disease control and enhanced immune priming
Regulatory: **Additive to Standard of Care**	• Straightforward path from PoC to pivotal • Reduced patient enrollment hurdles compared to monotherapy study
Commercial: **Meaningful Beachhead Indication**	• Second line gastric cancer market size in US, EU5, and JP is up to $1.7B • Upside in several other tumors

PRS-344: Meaningfully Building on Localized MoA of PRS-343

Candidate	PRS-344
Function/MoA	Localized 4-1BB agonism with PD-L1 antagonism
Indications	N.D.
Development	2021 IND expected (in co-dev with Servier)
Commercial Rights	Opt-in for co-development with full U.S. commercial rights; royalty on ex-U.S. sales



Financial Overview (As of 9/30/20)

$82.6 M	$0.0	54.3 M
Cash & Cash Equivalents	Debt	CSO

$125+ M non-dilutive capital from anchor partnerships

Appendix

Baseline Characteristics : Monotherapy and Combination with Atezolizumab

All Subjects (n = 74, 41)

Characteristic	Monotherapy; n (%)	In Combination with Atezolizumab; n (%)
Age, Median (range)	63 (24–92)	59 (26-87)
Gender		
F	44 (59%)	23 (56%)
M	30 (41%)	18 (44%)
ECOG PS*		
0	19 (26%)	12 (29%)
1	55 (74%)	18 (44%)
Prior Therapy Lines		
1	9 (12%)	5 (12%)
2	10 (14%)	7 (17%)
3	15 (21%)	6 (15%)
4	11 (15%)	6 (15%)
5+	28 (38%)	17 (41%)
Median no. of anti-HER2 Treatments		
Breast	7	3-4
Gastric	3	1

Primary Cancer Type	Monotherapy; n (%)	In Combinatio Atezolizumab;
Gastroesophageal	27 (36%)	7 (17%)
Breast	16 (22%)	12 (29%)
Colorectal	10 (14%)	5 (12%)
Gynecological	9 (12%)	4 (10%)
Biliary Tract	7 (9%)	6 (15%)
Non-Small Cell Lung	-	4 (10%)
Bladder	2 (3%)	1 (2%)
Pancreatic	1 (1%)	1 (2%)
Other – Cancer of Unknown Origin	1 (1%)	1 (2%)
Other – Salivary Duct	1 (1%)	-

*Combination trial enrolled ECOG 2 patients as well (not shown on this chart)

Data cut-off

Monotherapy

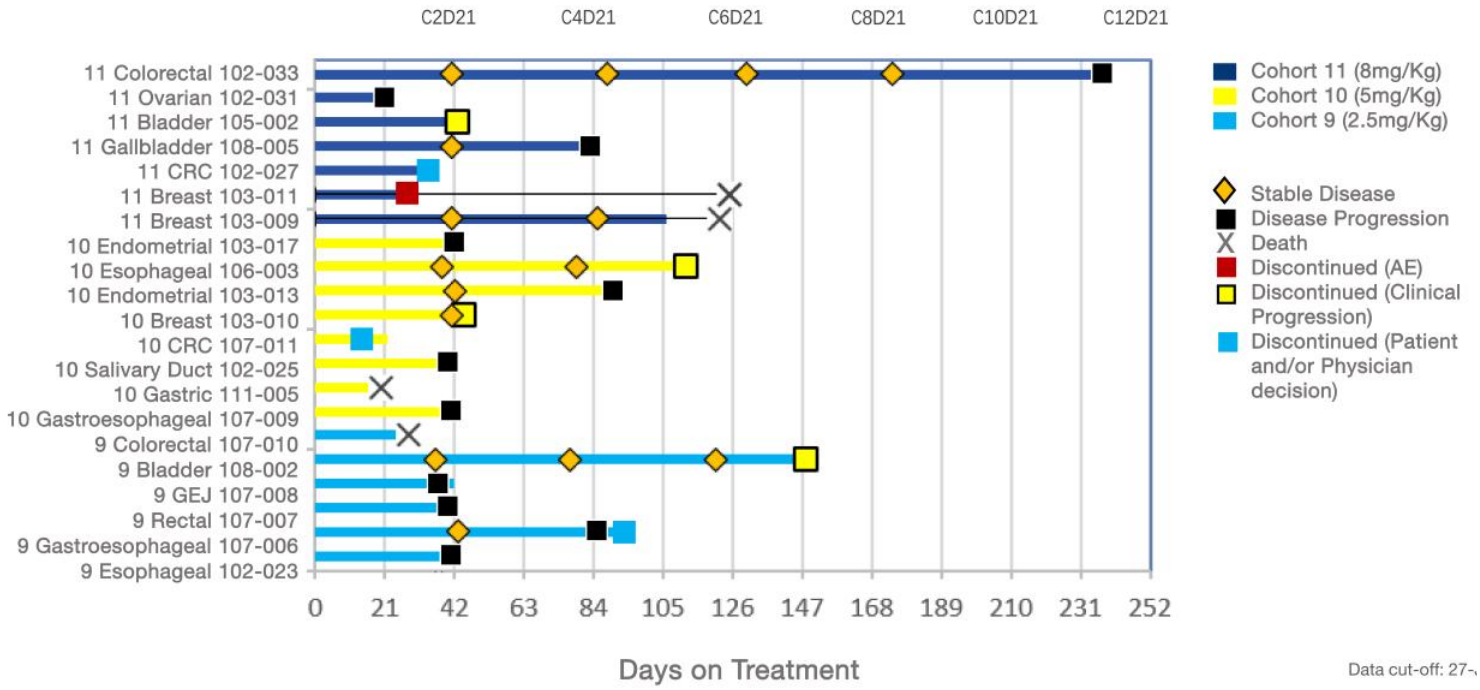
Treatment-Related Adverse Events (Monotherapy Trial)
All Subjects

Occurred in > 1 Patient	Monotherapy	
	n = 145 (%)	% Grade 3
Infusion Related Reaction	27 (19%)	3 (2%)
Fatigue	11 (8%)	1 (1%)
Nausea	11 (8%)	
Vomiting	8 (6%)	
Chills	8 (6%)	
Anemia	2 (1%)	1 (1%)
Arthalgia	2 (1%)	
Asthenia	2 (1%)	
Cough	2 (1%)	
Decreased appetite	2 (1%)	
Diarrhea	6 (4%)	
Dizziness	2 (1%)	
Dyspnoea	3 (2%)	
Flushing	5 (3%)	2 (1%)
Non-cardiac chest pain	4 (3%)	
Paraesthesia	3 (2%)	1 (1%)
Pruritis	3 (3%)	
Rash	2 (1%)	

One TRAE above Grade 3: Grade 4 Infusion Related Reaction in cohort 10 (5mg/kg PRS-343, Q3W).

Data cut-off

Average Time on Treatment with PRS-343
Cohorts 9-11a



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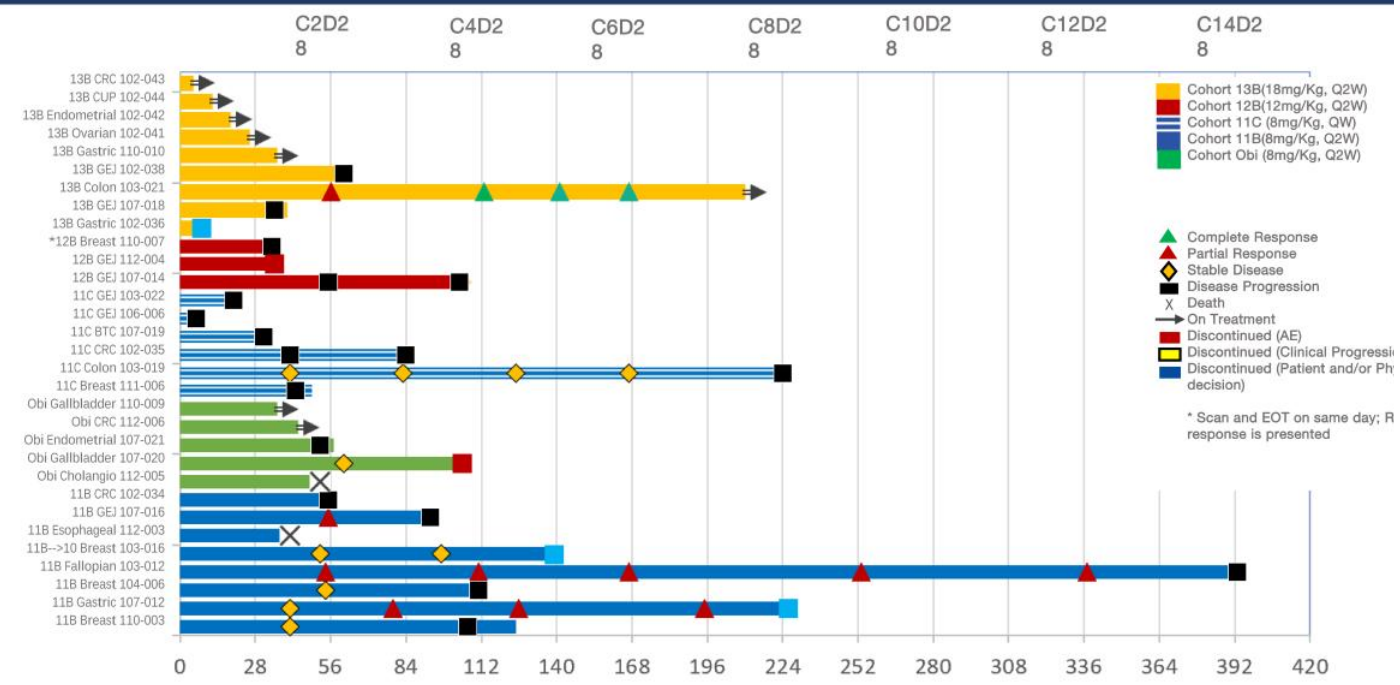
Improving Lives

Average Time on Treatment with PRS-343
Cohorts 11b-13b



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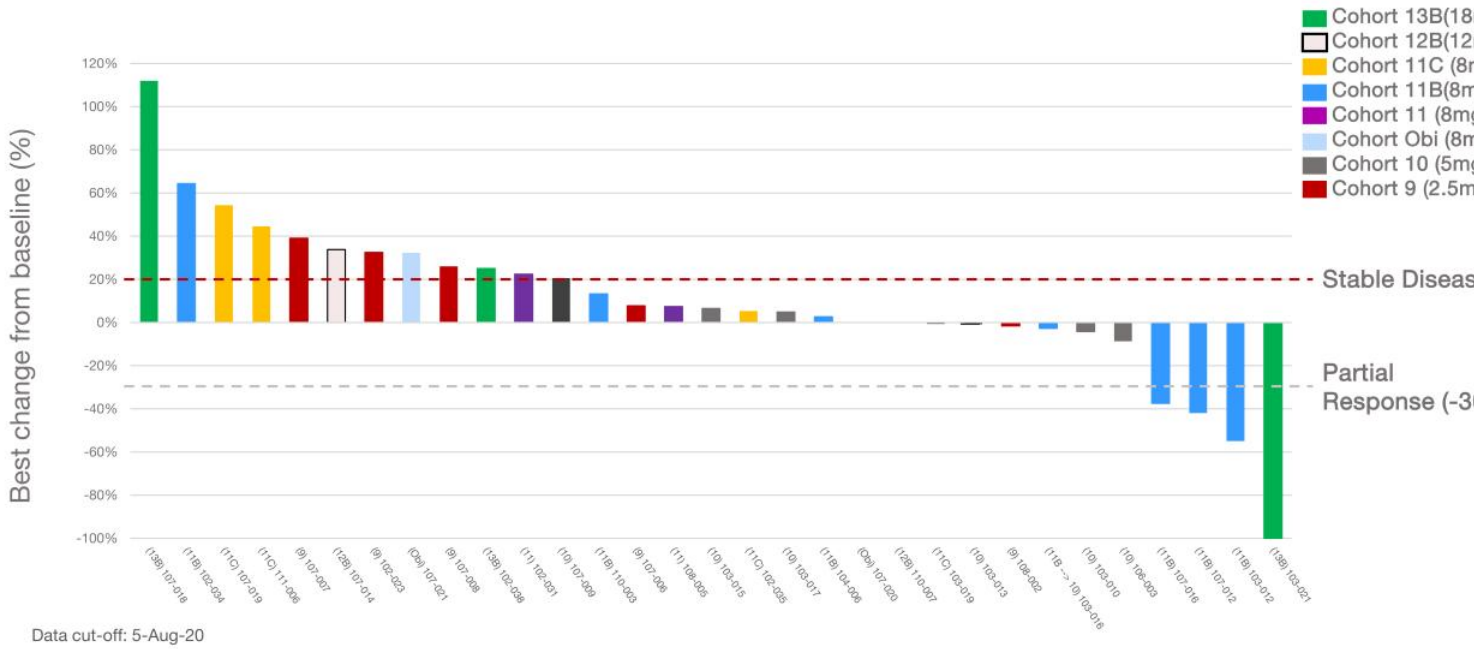
Improving Lives

Best Response in Target Lesions (Monotherapy Trial)
Cohorts 9-13b



Data cut-off: 5-Aug-20

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Improving Lives

Case Study: Gastric Cancer Patient with Confirmed Partial Resp

Patient Profile, Treatment History and Treatment Outcome

Patient Profile

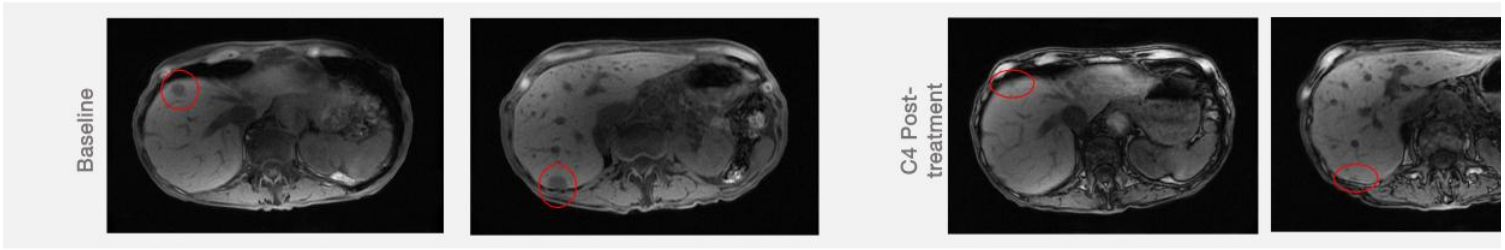
- Cohort 11b | 8 mg/kg every two weeks
- 80-year old woman; initial diagnosis in June 2017
- Stage IV gastric adenocarcinoma
- Metastases to liver, lymph node and adrenal glands
- HER2 IHC 3+; PD-L1 positive (CPS=3)
- NGS: ERBB2 amplification, TP53 mutation, alteration of CDK12 and SF3B1

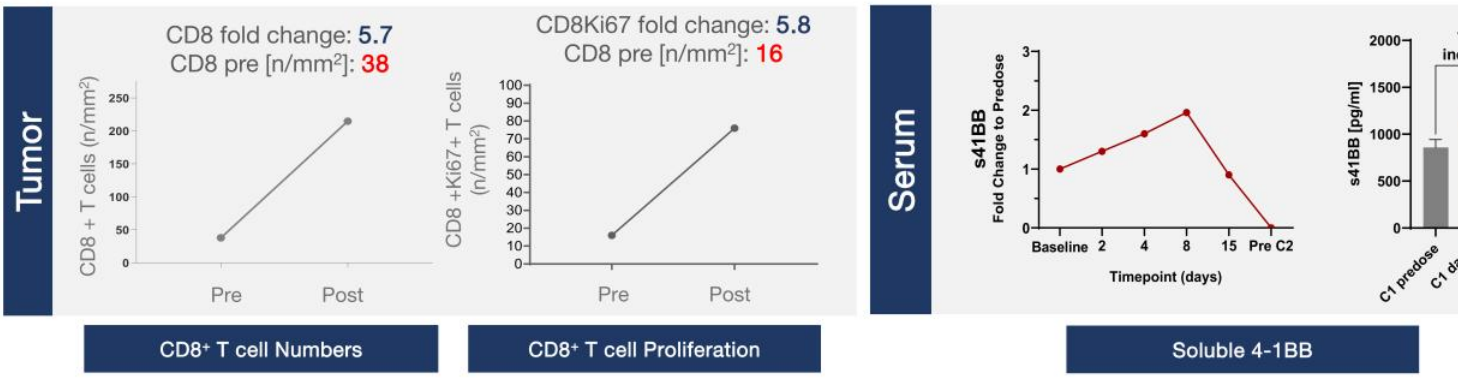
Oncology Treatment History	Duration	Best
Trastuzumab, Pembrolizumab + Capecitabine/oxaliplatin	July 2017 – June 2018	Stabl
Nivolumab with IDO1 inhibitor (investigational drug)	Aug 2018 – Jan 2019	Stabl

Lesions	Lesion Site	Lesion Size (mm)				
		Baseline	C2 Post-treatment	C3 Post-treatment	C4 Post-treatment	C6 Post-trea
Target 1	Liver	14	12	10	9	8
Target 2	Liver	20	16	10	8	9
Target 3	Pancreas	19	16	14	14	14
% Change from Baseline			-17%	-36%	-42%	-42%
Non-target 1	Lung	Present	Present	Present	Present	Presen
Non-target 2	Stomach	Present	Present	Present	Present	Absent
Non-target 3	Stomach	Present	Present	Present	Present	Absent

Data cut-off:

CD8+ T Cell Numbers in the Tumor and Circulating s4-1BB Incr Post-Treatment in Responding Gastric Cancer Patient





CD8⁺ T cell Numbers | CD8⁺ T cell Proliferation | Soluble 4-1BB

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Improving Lives

Case Study #2: Rectal Cancer Patient with Confirmed Complete Response

Patient Profile, Treatment History and RECIST

Patient Profile

- Cohort 13b | 18 mg/kg Q2W
- 59-year-old male; initial diagnosis March 2017
- Stage 4 rectal adenocarcinoma cancer; metastasized to heart and lung
- FoundationOne Her2 amplification; in-house testing IHC 3+
- MSS, TMB low (2 mt/Mb)

Oncology Treatment History	Duration
Capecitabine + XRT	Apr-May 2017
Neoadjuvant Folfox	May-Sep 2017
Resection	Dec 2017
Folfiri/Avastin	Mar-Jul 2018
5FU/Avastin maintenance	Aug 2018-May 201
Irinotecan/Avastin	May-Nov 2019
SBRT	Nov 2019

Lesions	Lesion Site	Lesion Size (mm)			
		Baseline	C2 Post-treatment	C4 Post-treatment	C6 Post-tre
Target 1	Lung	22	13	0	0
% Change from Baseline			-41%	-100%	-100
Non-target 1	-	Present	Present	Absent	Abse

Data cut-off: 27-Jul-20

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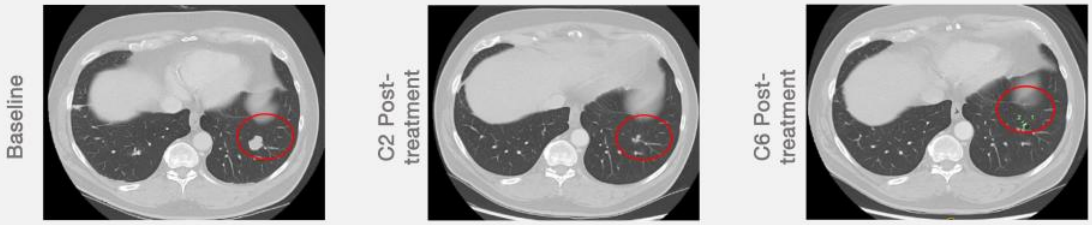
Improving Lives

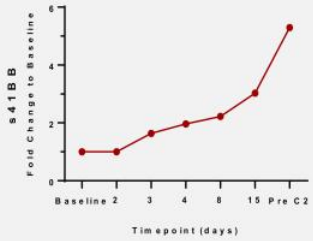
CD8+ T Cell Numbers in the Tumor and Circulating s4-1BB Increase Post-Treatment in CR Rectal Cancer Patient

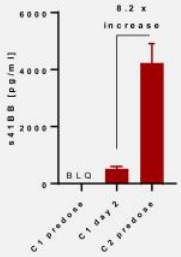


Tumor

CD8 fold change: 2.3 CD8 pre [n/mm²]: 238

Serum





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Improving Lives

Combination Therapy with Atezolizumab

Treatment-Related Adverse Events (Combination Trial)
All Subjects

Occurred in > 1 Patient	Combination with Atezolizumab	
	n = 148 (%)	% Grade 3
Infusion Related Reaction	38 (26%)	3 (2%)
Fatigue	12 (8%)	
Nausea	8 (5%)	
Vomiting	38 (26%)	
Abdominal pain	2 (1%)	
Anemia	4 (3%)	2 (1%)
Anorexia	2 (1%)	
Arthalgia	2 (1%)	
Diarrhea	5 (3%)	1 (1%)
Dry mouth	3 (2%)	
Fever	3 (2%)	
Lightheadness	2 (1%)	
Lymphocyte count decreased	3 (2%)	1 (1%)
Neutrophil count decreased	3 (2%)	1 (1%)
Peripheral sensory neuropathy	2 (1%)	
Pruritis	4 (3%)	

Two TRAEs above Grade 3: Grade 4 AST increase, Grade 3 transaminitis, and eventually Grade 5 hepatic failure in cohort 7 (8mg/kg + 1200mg atezolizumab); Grade 4 hemolytic anemia (unrelated to PRS-343, related to atezolizumab) in cohort 7.

Data cut-o

Summary of Responses of PRS-343 in Combination with Atezolizumab

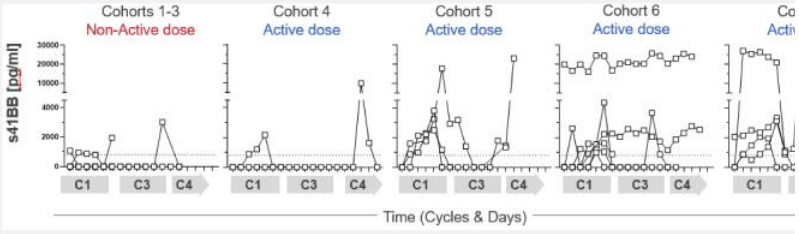
Cohort	7	6	5	4	Total
Best Response	8mg/kg, Q3W	5mg/kg, Q3W	2.5mg/kg, Q3W	1mg/kg, Q3W	
Evaluable Patients	8	8	8	3	27
PR	1	2	-	1	4
SD	4	1	1	0	6
ORR	13%	25%	0%	33%	15%
DCR	63%	38%	13%	33%	37%

Data cut-off: 27-Jul-20

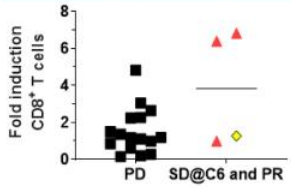
Soluble 4-1BB Increases in Active Dose Cohorts & Clinical Benefit is Associated with Tumoral Immune Cell Activation

4-1BB target engagement

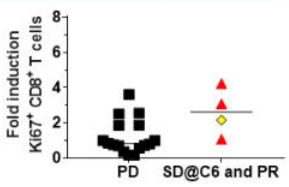
Soluble 4-1BB in serum



CD8⁺ T cell Numbers



CD8⁺ T cell Proliferation



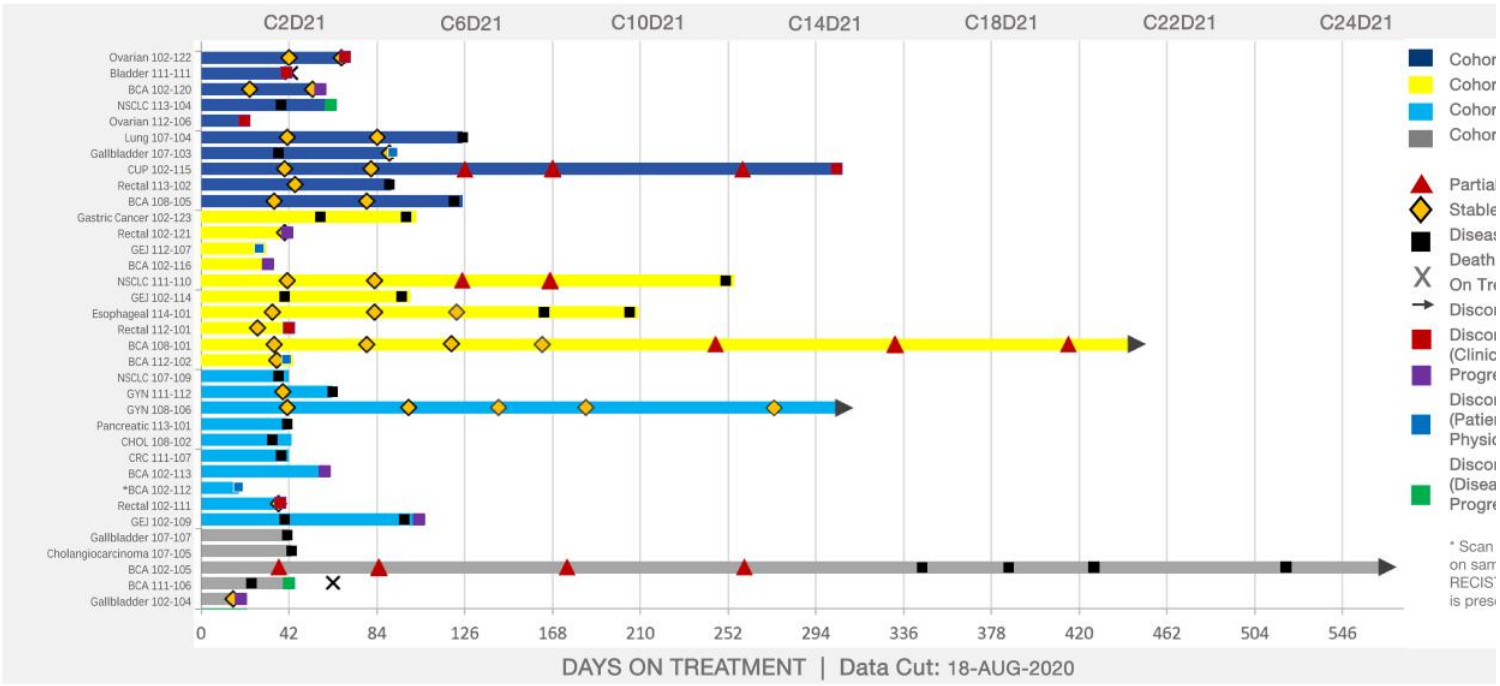
Tumor-localized activity

IHC on tumor tissue

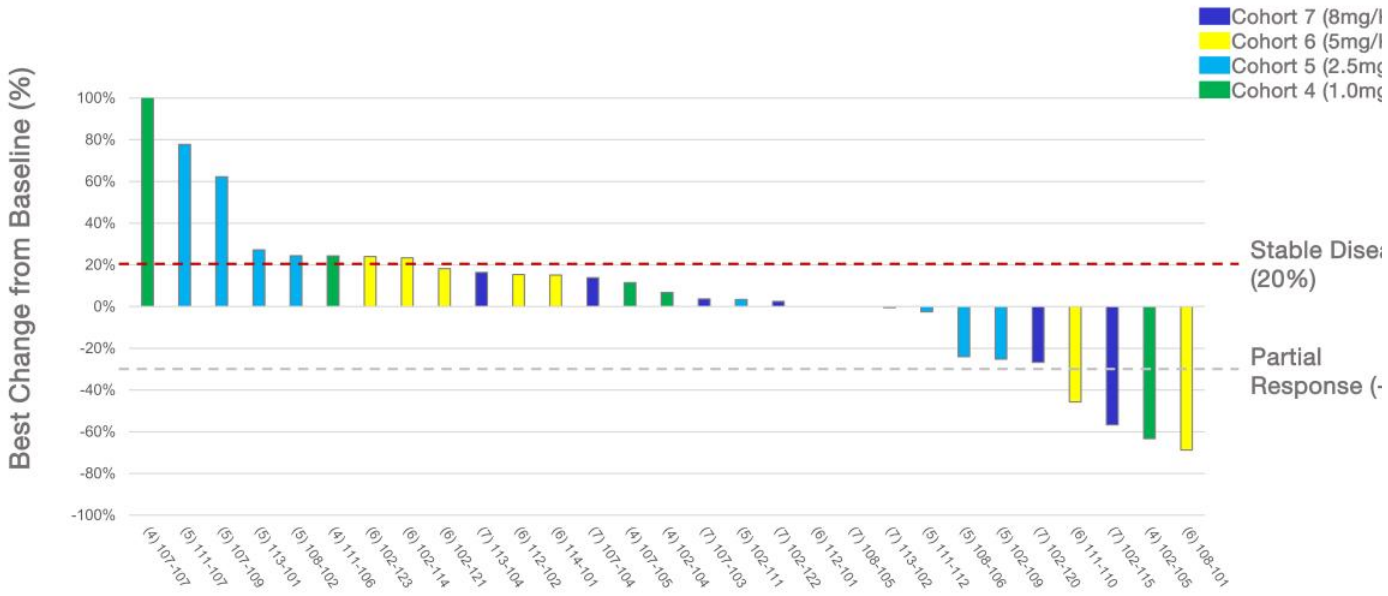
Patients with prolonged clinical benefit show a trend of increased CD8⁺ T cell numbers, proliferation and elevated cytolytic function in tumor biopsies

Substantial increase of s4-1BB is observed in active dose cohorts (4-7), suggesting PRS-343-mediated target engager

PRS-343 + Atezolizumab Duration of Exposure



Best Response in Target Lesions (Combination Study)
Cohorts 4-7



Data cut-off: 27-Jul-20

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Improving Lives

Case Study: Breast Cancer Patient with Stable Disease (Update

Patient Profile, Treatment History and RECIST

Patient Profile:

- Cohort 6 | 5 mg/kg Q3W + 1200mg atezolizumab
- 52-year-old male; Initial diagnosis July 2011
- Stage 2 Invasive Ductal Breast Cancer
- FISH HER2/CEP17 ratio 2.4, HER2 copy number 4.8
  In-house testing IHC2+, FISH+
- PD-L1 low in pre-treatment and high in post treatment biopsy

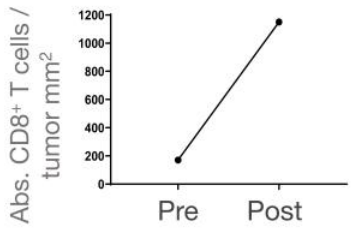
Oncology Treatment History	Duration
Trastuzumab/Docetaxel/ Tamoxifen/Carboplatin	Sep 2011-Jul 2013
Trastuzumab/Pertuzumab/Vinorelbine	Aug 2013-Jan 201
T-DM1/Fulvestrant	Nov 2017-Mar 201
Capecitabine/Lapatinib	Mar 2018
Palbociclib/Arimidex	Apr-May 2019

Lesions	Lesion Site	Lesion Size (mm)						
		Baseline	C2 Post-treatment	C4 Post-treatment	C6 Post-treatment	C8 Post-treatment	C12 Post-treatment	C16 treat
Target 1	right pulmonary ligament lymph node	16	18	15	13	13	6	
% Change from Baseline			+12.5%	-6%	-19%	-19%	-63%	-6
Non-target 1-4	-	Present	Present	Present	Present	Present	Present	Pre

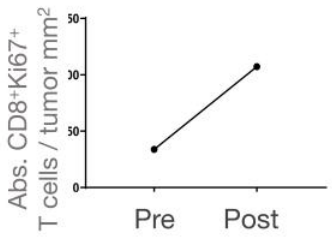
Data cut-off: 27-Jul-20

Tumoral and Circulating s4-1BB Increase Post-Treatment in PR Breast Cancer Patient

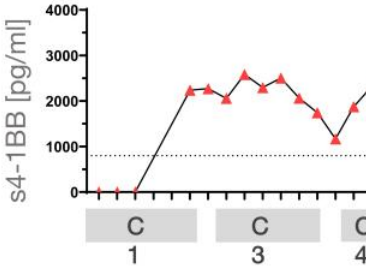
CD8+ T cell Numbers



CD8+ T cell Proliferation



Soluble 4-1BB



CD8+ T cell numbers, proliferation, cytolytic molecules and s4-1BB increase post-treatment, demonstrating 4-1BB arm activity of PRS-343

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