

# **Perseus Proteomics Inc.** **(Security code:4882)**

**FY2021 First Half Business Results**  
**November 15, 2021**

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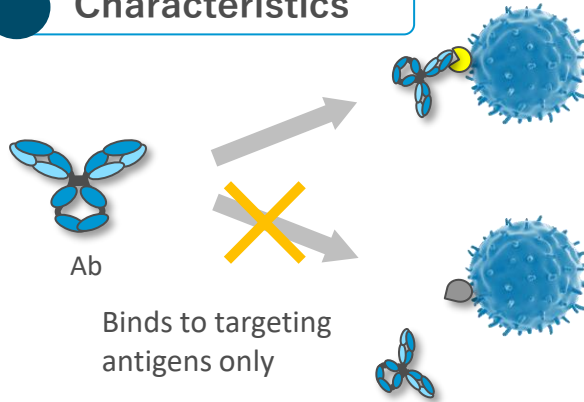
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# What are Ab drugs?

Abs are substances that remove foreign objects in human body

Ab drugs are Abs obtained against targets expressed on cancers or pathogens

## Characteristics



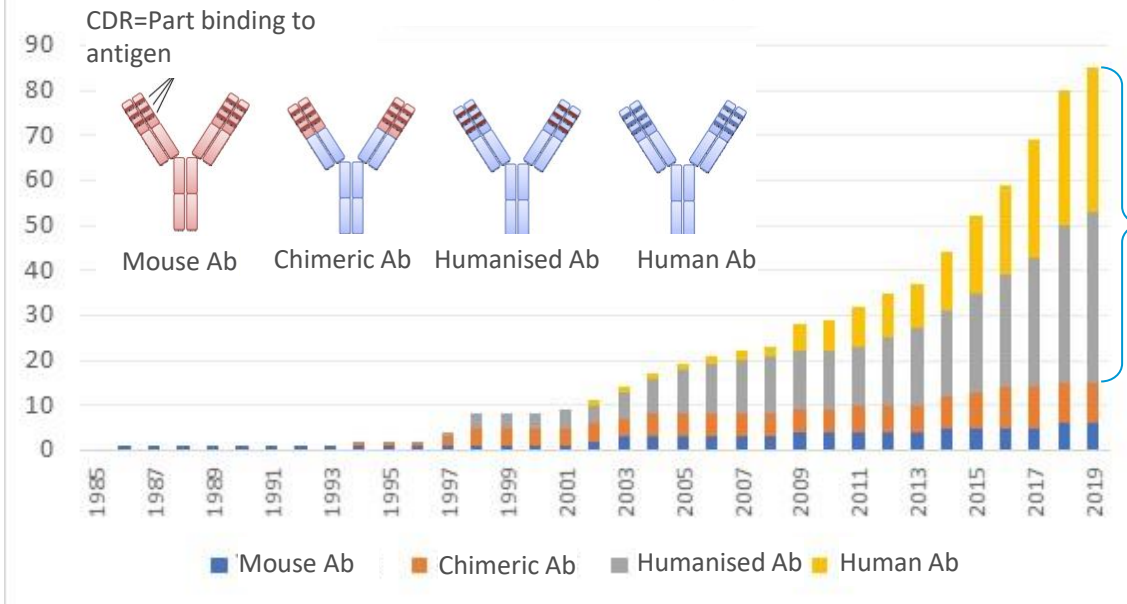
## Expected effects

- Blocks signal transmission and inhibits multiplication functions, etc.
- Activates immune cells including T cells to induce cytotoxicity
- Activates physiological functions
- Transmits drugs to cells where targets are expressed

## Indication of Ab drugs

- Cancer
- Rheumatism
- Inflammatory bowel disorder
- Psoriasis
- Infectious diseases
- Bronchial asthma
- Atopic dermatitis
- Seasonal allergic rhinitis
- Complement deficiencies
- Hypercholesteremia
- Macular degeneration
- Rare diseases
- Urticaria

Number of Ab drugs approval



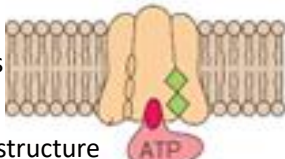
Humanised or human Abs are in mainstream

# Ab creation technology now required

## Difficulty=High antigen

The most important targets  
Still untouched

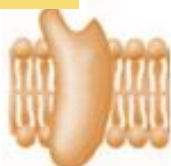
Quaternary structure



## Difficulty=Medium antigen

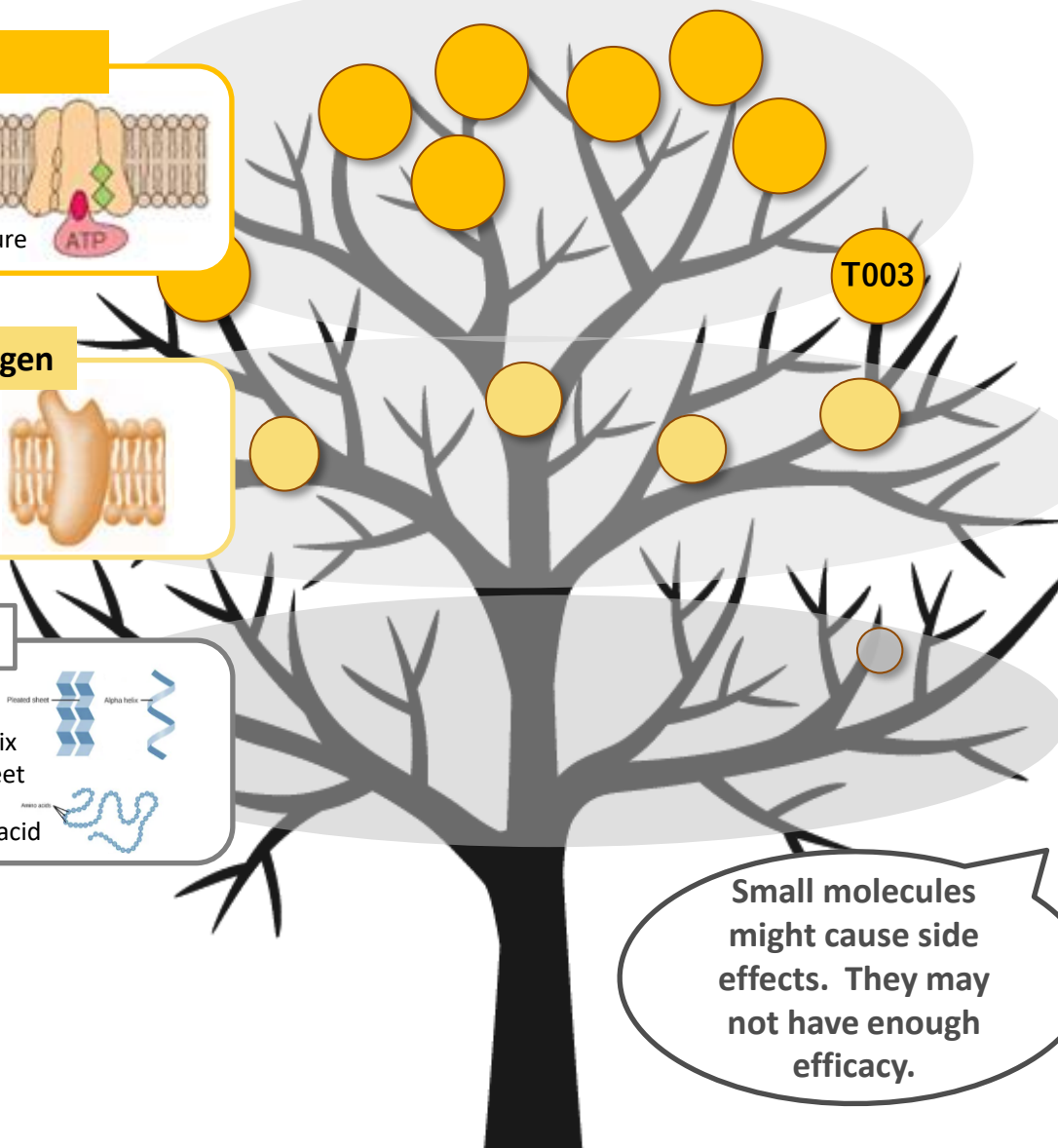
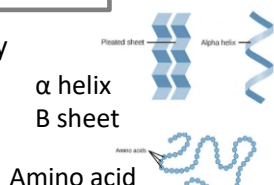
Receptor-type targets  
Needs functional Ab

Tertiary structure



## Difficulty=Easy antigen

Antigen as targets already  
developed



Important targets  
are not easily  
reachable.

Any technology to  
help us to get the  
fruit on the treetop  
easily?

Small molecules  
might cause side  
effects. They may  
not have enough  
efficacy.

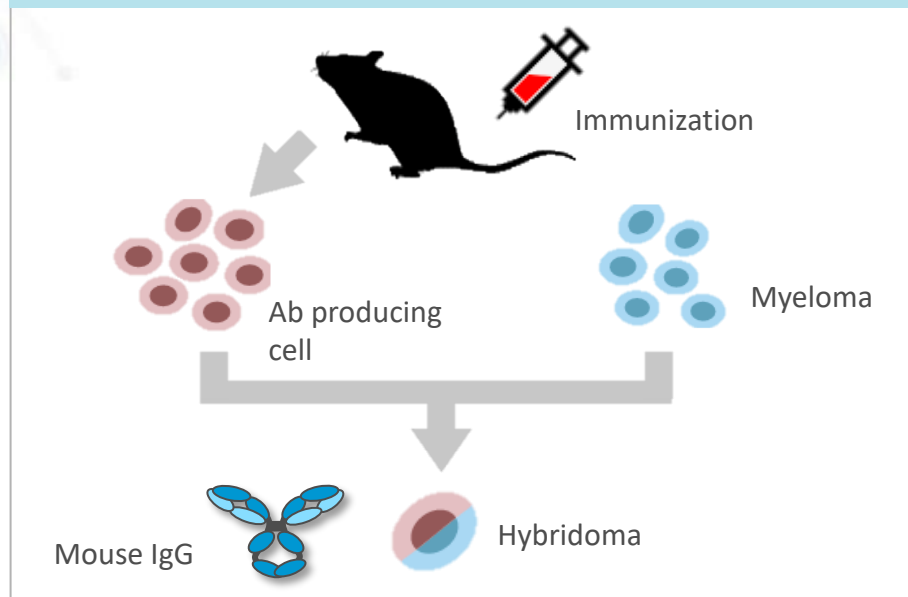
Antigen  
preparation is the  
core task!



**Technology required for obtaining Abs  
efficiently against medium to high level antigens**

# Our technology to obtain Abs

## 1) Hybridoma method



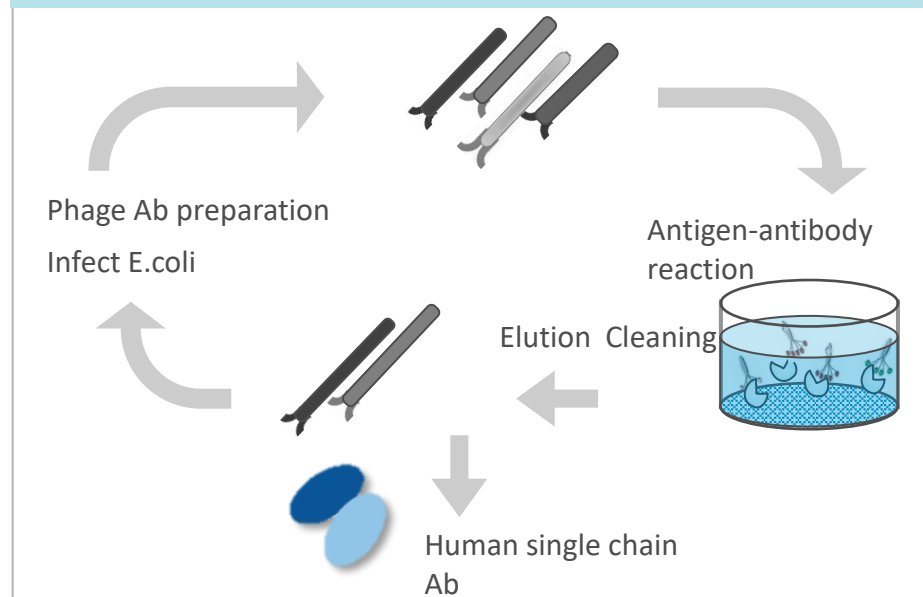
### Merit

- Easy method, established technique
- Increased biological affinity
- Low cost

### Problem

- Abs with species crossing are hard to obtain
  - Needs humanisation due to immunnogenicity
  - Abs against complex antigens are hard to obtain
  - Easy-to-obtain Abs already developed
- ⇒ Focusing on new targets and modified Abs including ADC\*1 and RIT\*2

## 2) Phage display method



- Possible to obtain human Ab
- No animals used
- No need to consider biological toxicity
- Rich in screening conditions

- Needs skills in creation of libraries
  - More expensive than animal immunization
  - Low affinity of antigen-antibody
- ⇒ Conquered this problem by maximising library diversity

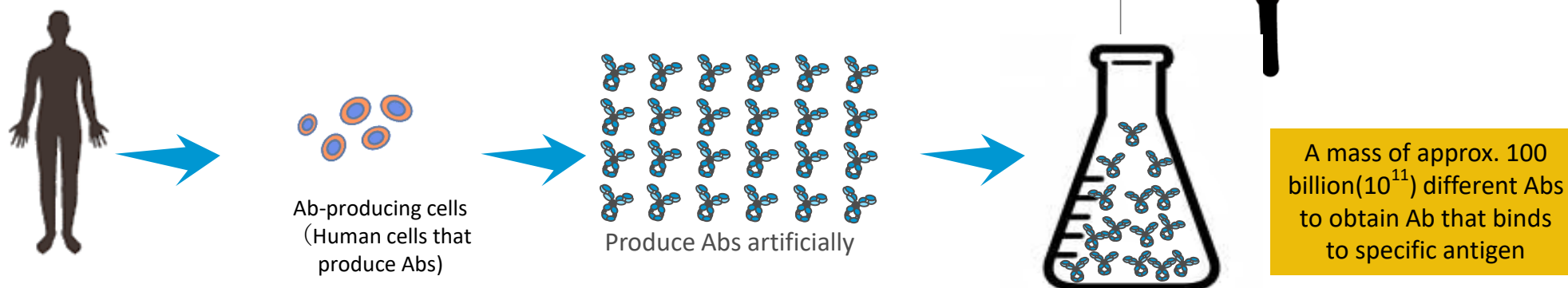
\*1 ADC: Antibody drug conjugate. It delivers drug combined with Ab by utilizing Ab function.

\*2 RIT: Radioimmunotherapy. Radioisotope combined with Ab irradiates cancer cells by utilizing Ab function.

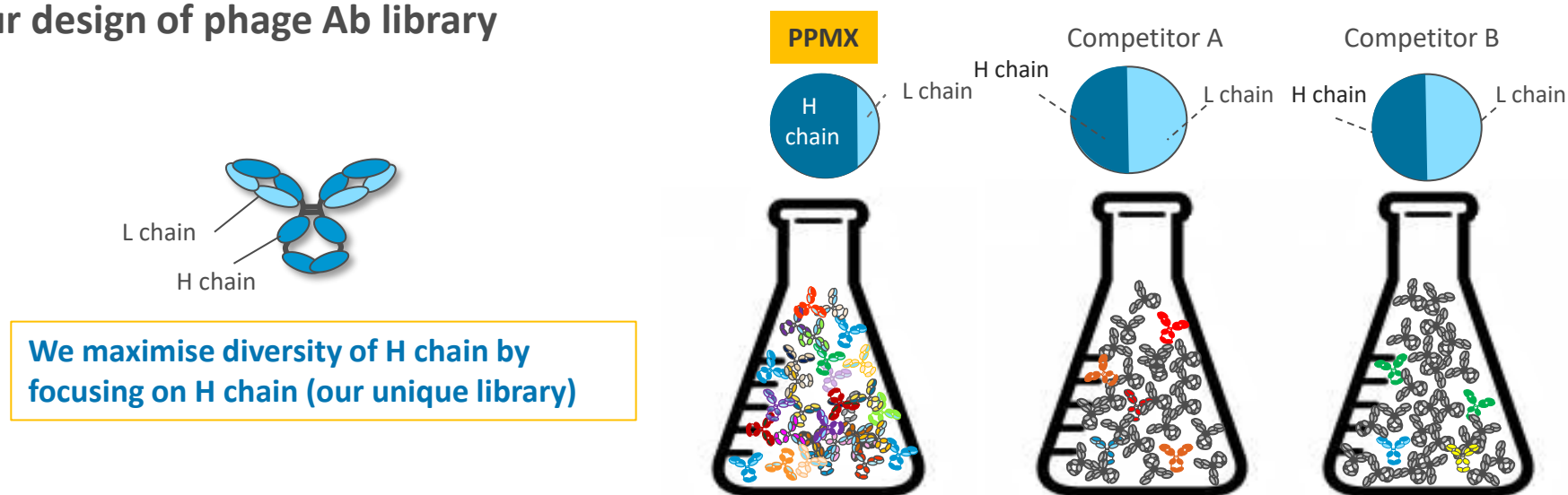


# Our strength: Phage Ab library

## 1 What is phage Ab library?



## 2 Our design of phage Ab library



While numbers of Abs are the same 10 billion, diversities are different

**Phage display method utilizing maximised diversity of Ab library**

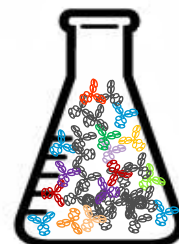
# Our strength: Ab screening using cell (PPMX exclusive method)

## Problem 1

During preparation of antigen, steric structure is lost.



**Ab screening using living cells**

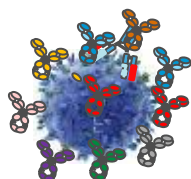


## Solution



- Reflects complex steric structure through using living cells
- Directly obtains Abs against antigens on cell membrane

## Problem 2



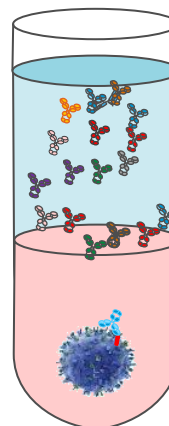
Numerous unrelated Abs also bind to cells.



**ICOS\* method:  
Ab screening utilizing organic solvent**

Water layer

Organic layer



## Solution

- Obtains Abs that bind to antigen only
- Patent registered

**Efficiently separates Abs difficult to obtain by targeting cells**

### Our technology on Ab drug development

Our unique technology platform sophisticated to aim at drug discovery for highly difficult targets

**Hybridoma  
method**



Mature and reliable  
technology

**Phage display  
method**



Maximizing human Ab  
diversity

**ICOS method**



Reflecting complex steric  
structure on cell  
membrane

Showing our maximum value  
in developing anti-cancer drugs

**PPMX's sophisticated Ab obtaining platform**



01 Topics

02 Pipeline details

03 Financial results of first half FY2021

About us



# 01 Topics

**1**

**PPMX-T003, Ab under development in house:  
Started phase I clinical trial among polycythemia vera patients**

**2**

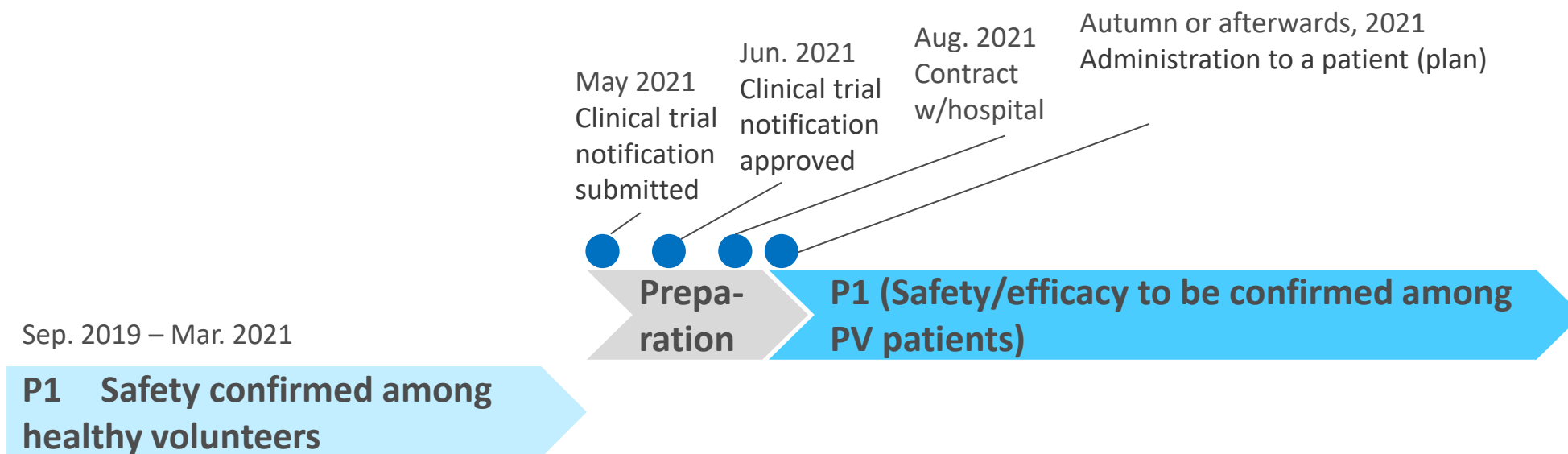
**PPMX-T001:  
Phase I clinical trial of ERY974 in combination with atezolizumab and  
bevacizumab by Chugai Pharmaceutical**

1

## PPMX-T003, Ab under development in-house:

### Started phase I clinical trial among polycythemia vera patients

Code No.	PPMX-T003
Applicable disease	Polycythemia vera (PV), various blood cancers
Stage	Phase I clinical trial (JP: preparation for first patient in)
Out-licensed to	-



#### Clinical trial information

[jRCT](#)

jRCT2051210083: <https://jrct.niph.go.jp/en-latest-detail/jRCT2051210083>

[clinicaltrials.gov](https://clinicaltrials.gov)

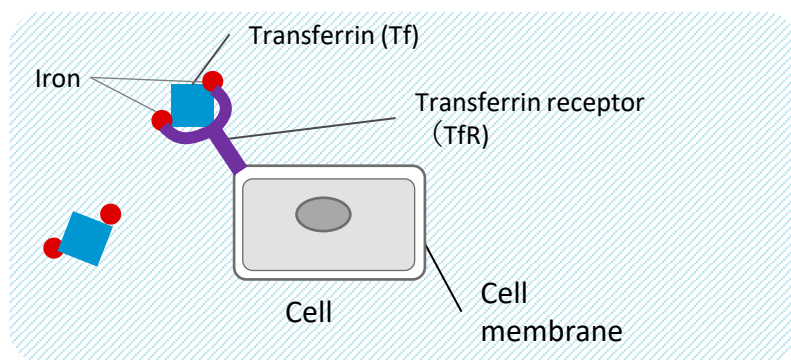
NCT05074550 : <https://clinicaltrials.gov/ct2/show/NCT05074550>

## First-in-class anti-cancer drug candidate targeting transferrin receptor

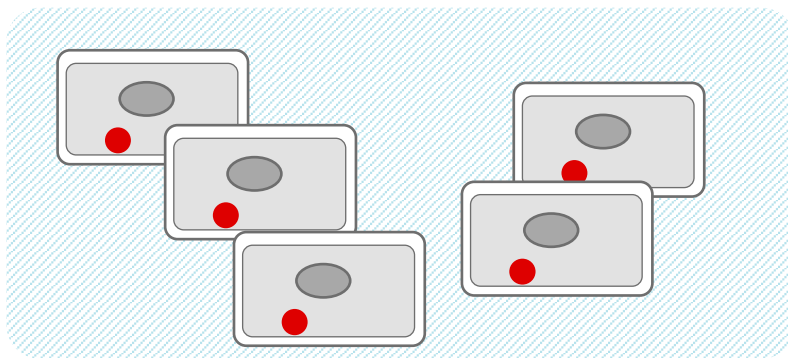
### Transferrin receptor (TfR):

- Strong target molecule for anti-cancer drug
- Expressed on cell membrane. Binds to transferrin (Tf) carrying iron for cellular iron uptake

#### 1 TfR binds to Tf



#### 2 Cell proliferation



#### [ Cells where TfR is highly expressed ]

- Erythroblast (normal cell, RBC producing cell)
- Cancer cell (especially acute cancer which is actively proliferating)

#### Well-known concept

**Blocking iron**

**⇒ Death or proliferation inhibition of cells**

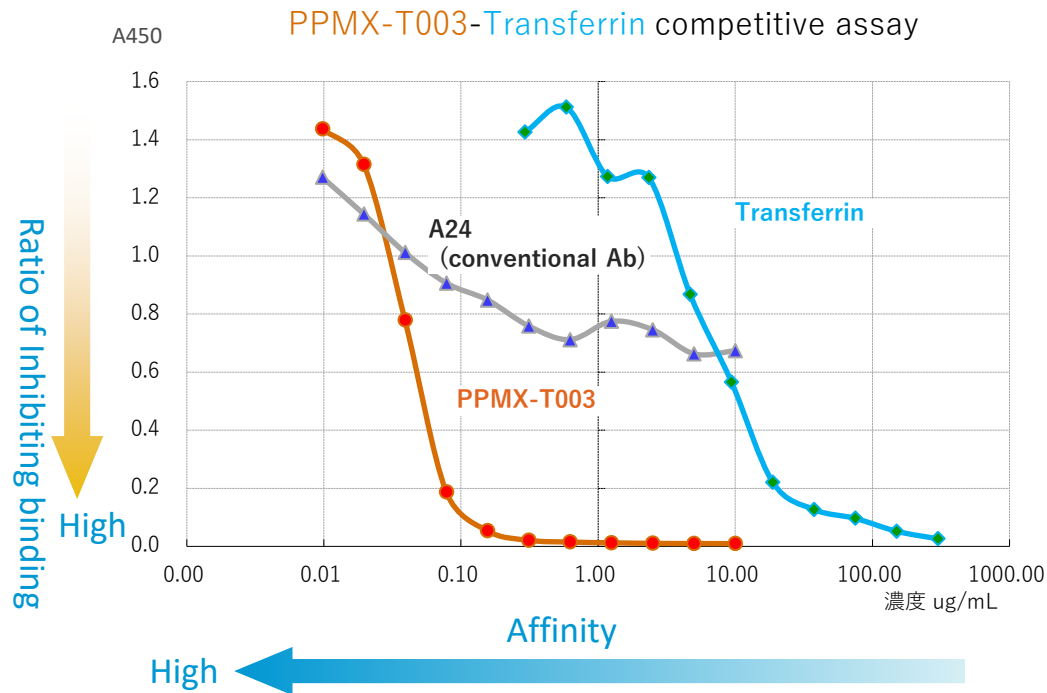
**Inhibiting cellular iron uptake leads to death/proliferation inhibition of cancer cells**

# PPMX-T003

## Highly functional Ab obtained by our phage display technology

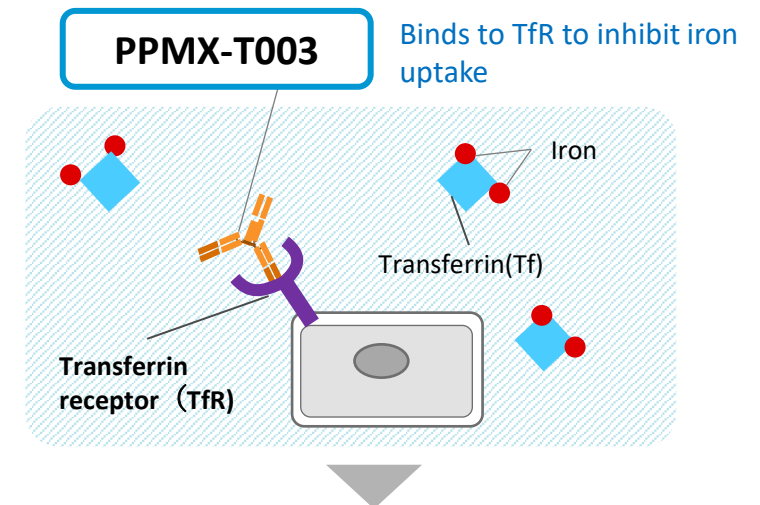
Shows unprecedented result in inhibiting ratio of binding Tf to TfR

Inhibits iron uptake into **erythroblast** and **cancer cells** and leads to cell death/proliferation inhibition

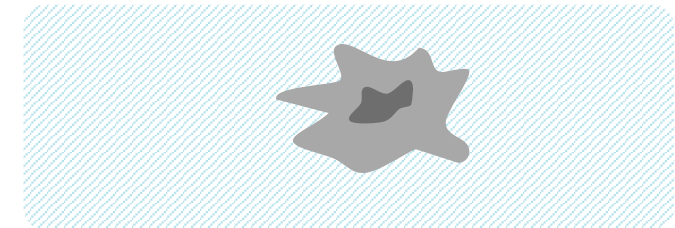


Inhibition of iron uptake has been difficult, however, PPMX-T003 is expected to bring it to reality as the first therapeutic drug for cancer and PV.

### 1 PPMX-T003 binds to TfR more tightly than Tf



### 2 Iron uptake inhibited. Death or proliferation inhibition of cells



**Anti-Transferrin receptor Ab with incomparable function of inhibiting binding**



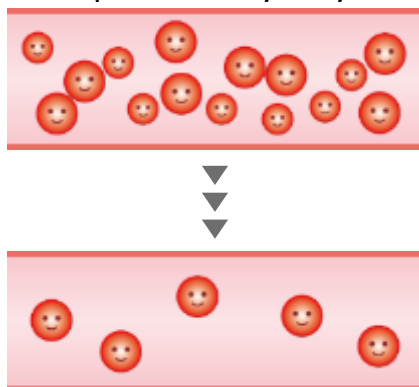
## Indication: Polycythemia vera (PV)

- RBC increases to an abnormal level.
- Thrombosis is easily formed due to thick and slow blood flow. Various organs are affected by thrombosis.
- 2 out of 100,000 people develop this disease. Number of patients in Japan: 30,000 (estimated by PPMX. Average life expectancy: 16 yrs)

### Current therapeutics

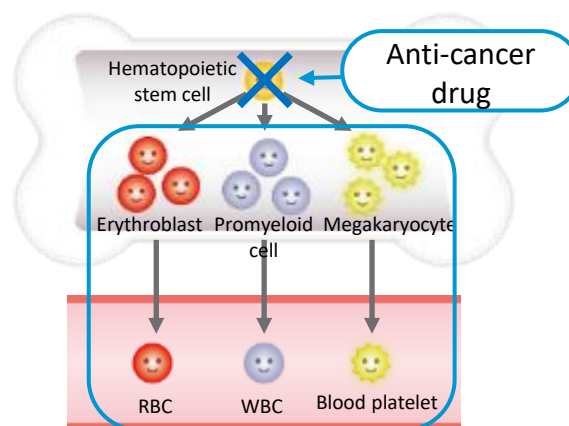
#### Therapeutic phlebotomy

Half of patients are treated by therapeutic phlebotomy only.



- Anemia
- Lassitude
- Depression
- Restless hands and legs
- Other diseases by iron deficiency

#### Anti-cancer drug, etc.

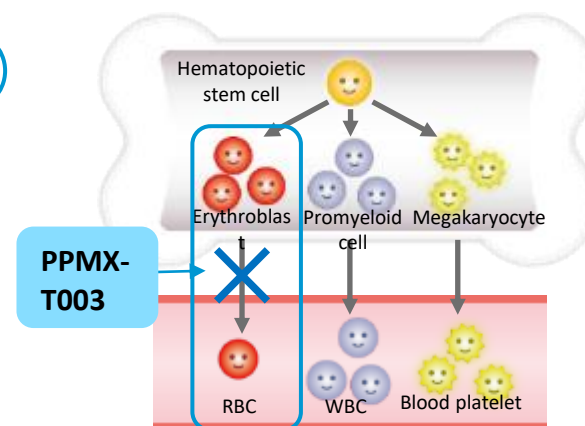


- Entire hematopoietic stem cell affected
- Secondary cancer risk
- Many side effects

### New candidate

PPMX

#### PPMX – T003



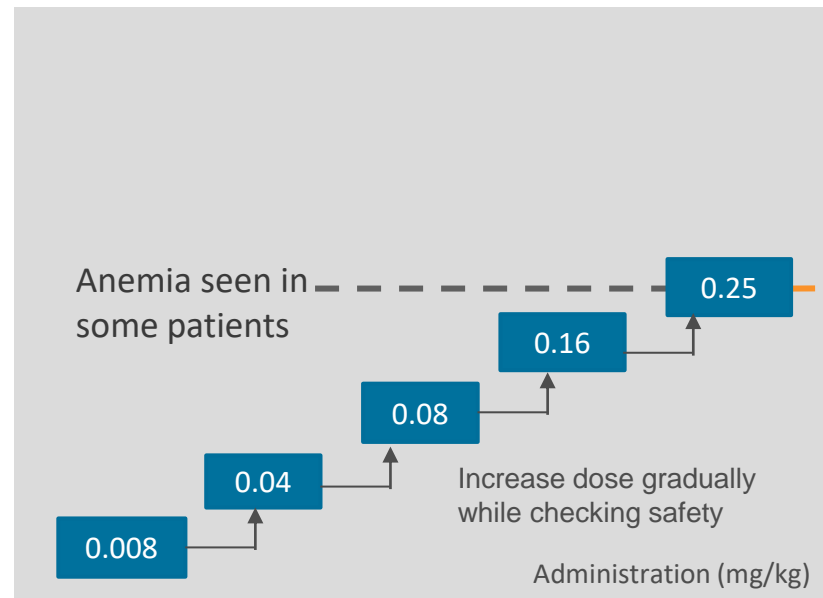
- Acts only on erythroblast
- Few side effects
- Safe to use

**PPMX-T003: effects on inhibiting abnormal proliferation of RBC expected**

# PPMX-T003: Development status

## P1 details

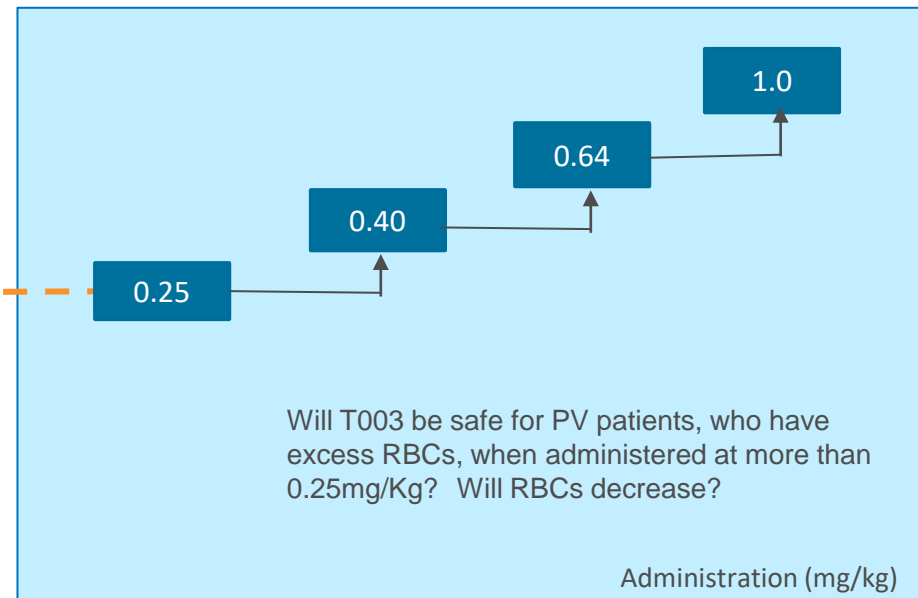
### P1 among healthy volunteers (finished)



2019.11 (administration)- 2021.3 (CSR)

Subject: Healthy volunteers  
Total number of patients: 40  
Cohort : 5 (double-blind, placebo)

### P1 among PV patients



2021.autumn (to be administered)- 2022.12 (plan)

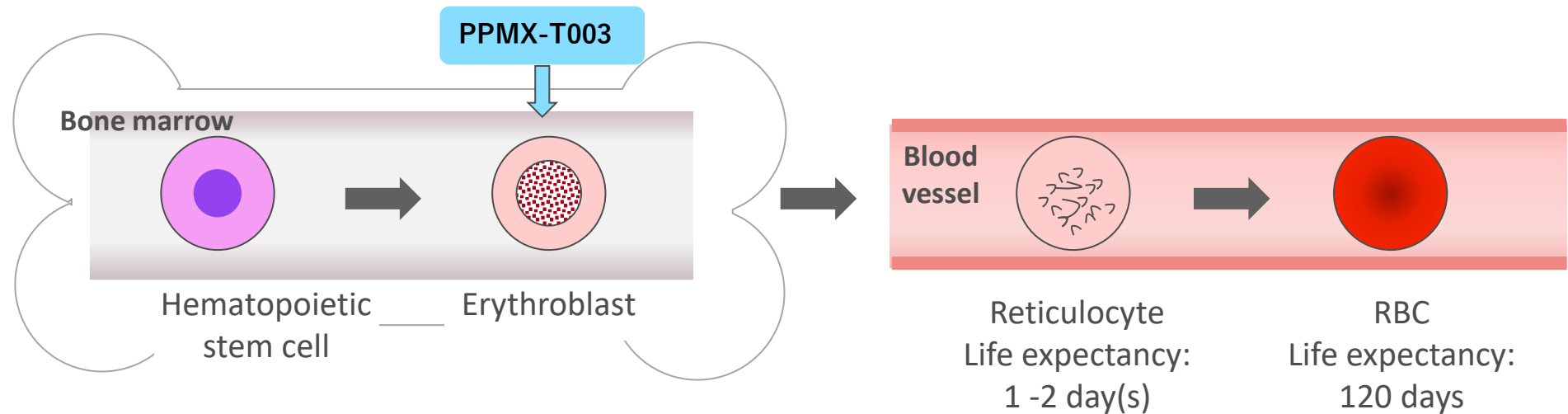
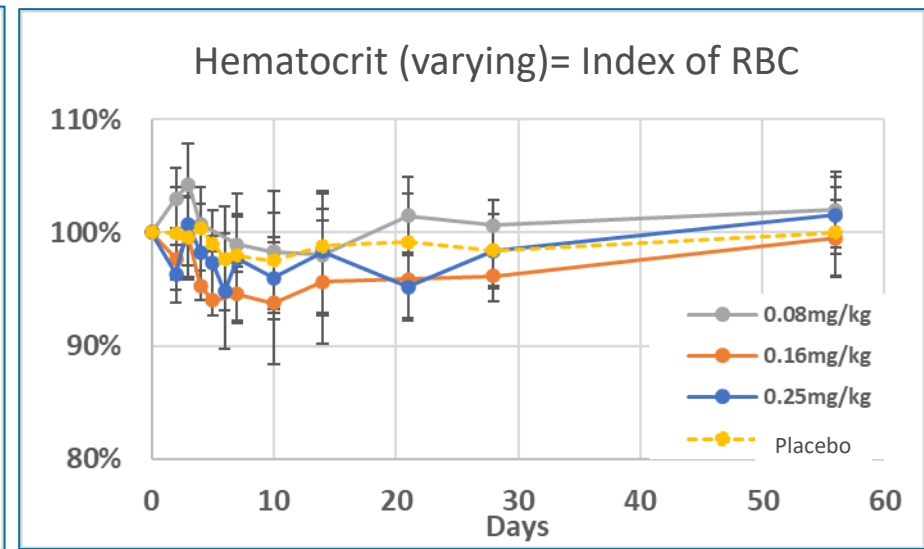
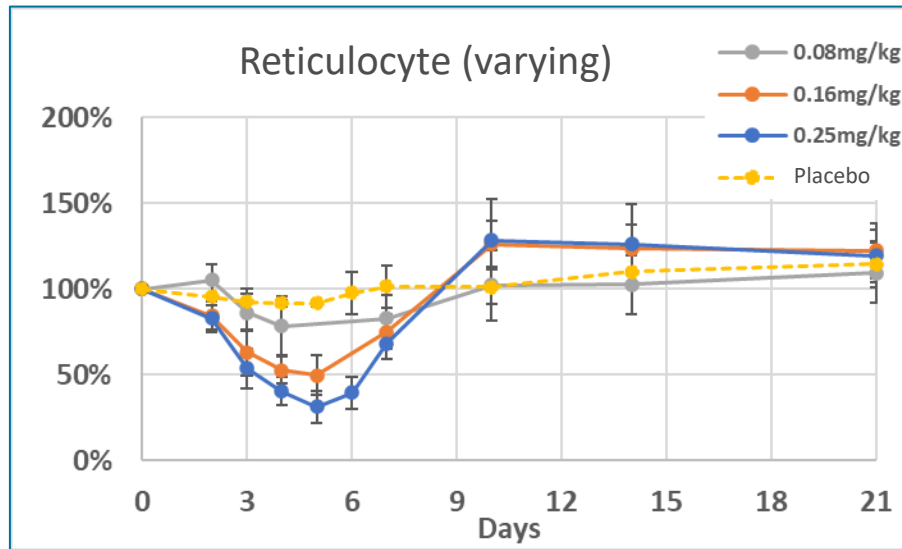
Subject: Polycythemia vera (PV) patients  
Total number of patients: 6  
Cohort : Ascending dose regimen (open)

**Administration to PV patients to start in or after autumn 2021**

# PPMX-T003: Results of P1 among healthy volunteers

Decrease in reticulocyte (immature RBC) and hematocrit (percentage of RBC)  $\Rightarrow$  Efficacy confirmed

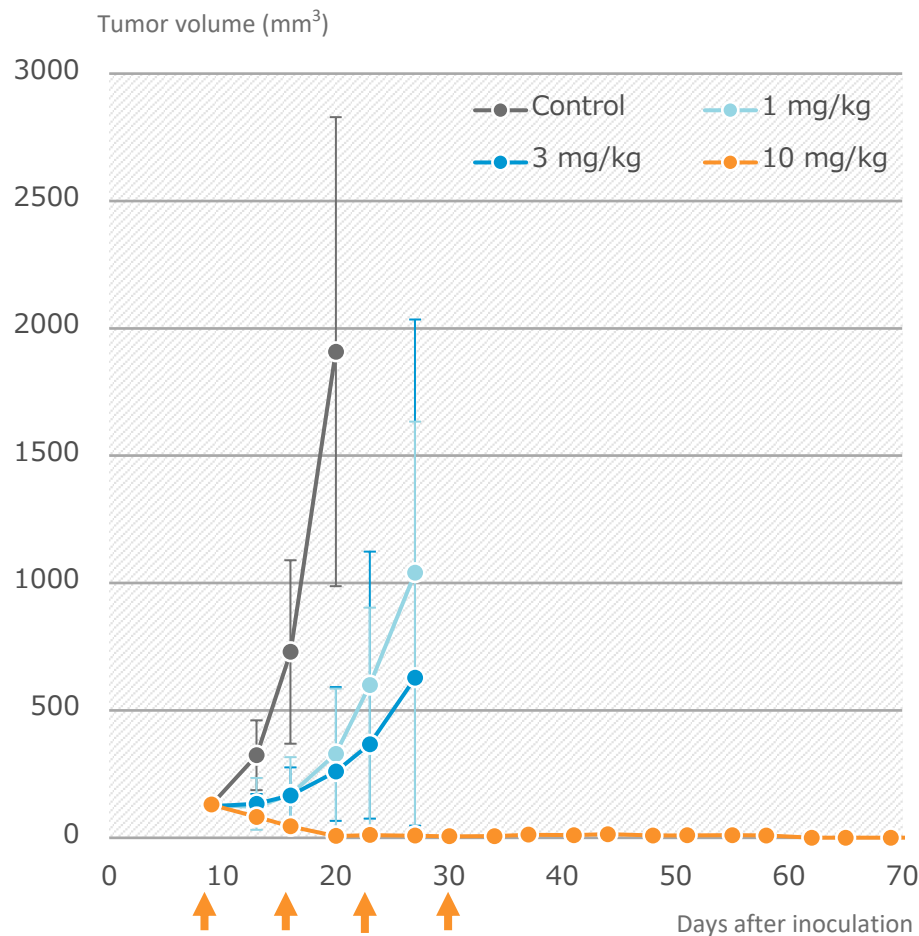
Human  
(HV)  
n=6



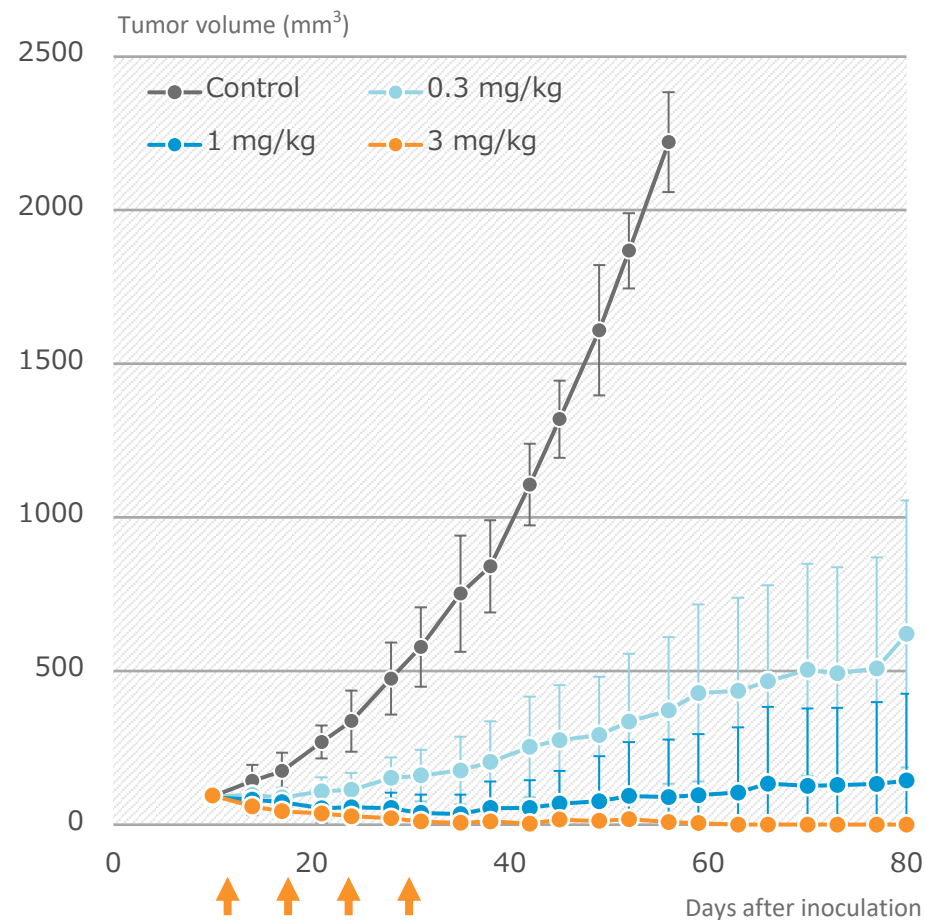
**Confirmed decrease in RBC on human body PPMX-T003 dose-dependently**

# PPMX-T003: Confirmed efficacy against blood cancers in mice

## ● AML



## ● Malignant Lymphoma



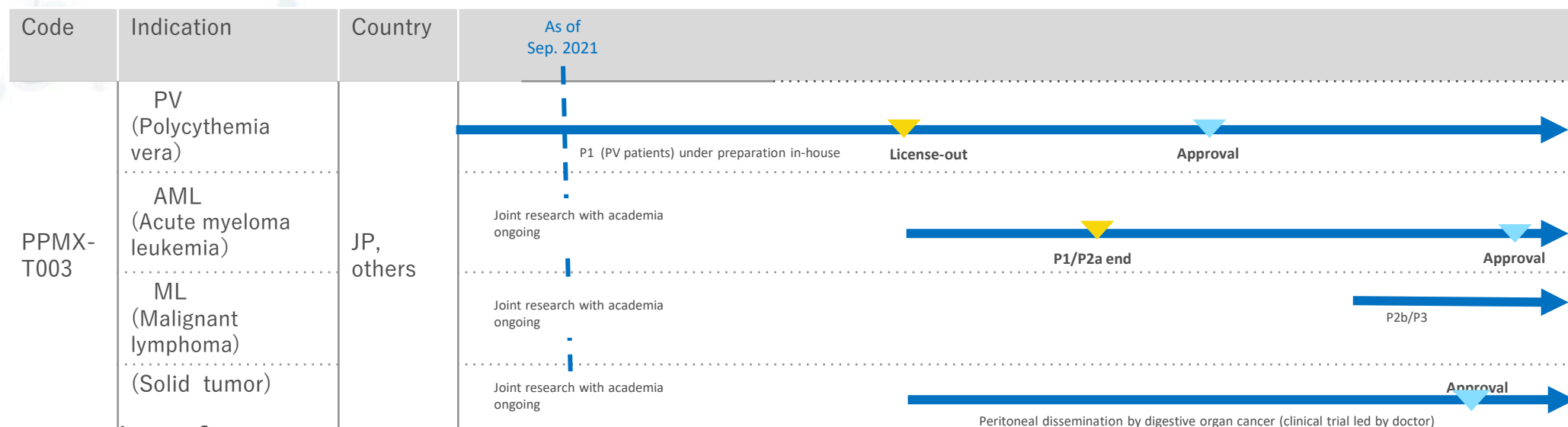
**Excellent efficacy against AML and various blood cancers is confirmed**

# PPMX-T003: Development plan

創薬

抗体研究  
支援研究

抗体・  
試薬販売



## Number of patients

Indication		Number of patients ww (rounded)	Note
PV (Polycythemia vera)	Chronic blood disease	280,000	Calculated with onset risk rate at 2 in 100,000*, life expectancy at 14 years*, population at 1 billion (developed countries)
AML (Acute myeloma leukemia)	Blood cancer	200,000	WHO data (assumes 40% of leukemia)
Malignant lymphoma	Blood cancer	590,000	WHO data (number of non-Hodgkin lymphoma patients)
Multiple myeloma	Blood cancer	190,000	WHO data
Peritoneal dissemination of cancer	Solid tumor	N/A	Over 10,000 and several thousand new patients annually in Japan

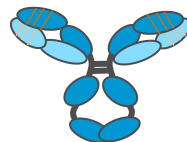
\* This chart is based on our assumption and does not guarantee the progress as shown here.

\* All the development after out-licensing is determined by the development strategies of licensing partners.

## Phase I clinical trial of ERY974 in combination with atezolizumab<sup>\*1</sup> and bevacizumab<sup>\*2</sup> by Chugai Pharmaceutical

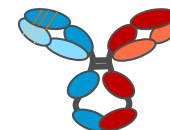
Code No.	PPMX-T001
Applicable disease	Liver cancer, solid tumor
Stage	<ul style="list-style-type: none"> <li>GC33 in combination with immune checkpoint inhibitor (ICI): P1 ongoing (JP, TW)</li> <li>ERY974 monotherapy: P1 finished (US, EU), P1 ongoing (JP)</li> <li><b>New</b> ERY974 in combination with ICI and angiogenic inhibitor: P1 started (JP, TW)</li> </ul>
Out-licensed	Chugai Pharmaceutical

Chugai Pharmaceutical development code: GC33, ERY974



GC33

● GPC3 Ab  
Binds to cancer cell



● CD3 Ab  
Binds to T cell

ERY974 (bispecific Ab)

2 arms respectively bind to different antigens.

\*1: immune checkpoint inhibitor      \*2: angiogenic inhibitor

Patents on PPMX-T001 will expire in June 2022.



## 02 Pipeline details

# Pipeline progress

Code	Indication	Region	Drug discovery/ Research	Preclinical	P1	P2	P3	Out-licensed
PPMX-T002	Solid tumor	USA Japan	RIT					FUJIFILM
PPMX-T003	Blood cancer	Japan						—
PPMX-T004	Solid tumor		ADC					FUJIFILM
PPMX-T001	Liver cancer	Japan USA Europe	GC33 Monotherapy					Chugai Pharmaceutical
		Japan Taiwan			GC33 w/ICI			
	Solid tumor	USA Europe Japan	ERY974 monotherapy					
	Liver cancer	Japan Taiwan			ERY974 w/ICI, angiogenic inhibitor			

Clinical trial of GC33 monotherapy is not ongoing.

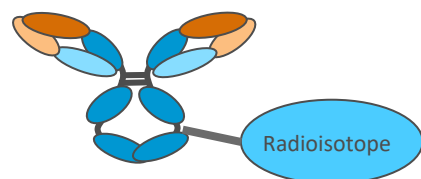
Patents on PPMX-T001 will expire in June 2022.

## PPMX-T002 :

### Ab labeled with radioisotope

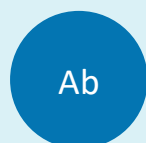
Code	PPMX-T002
Applicable disease	Biliary tract cancer, ovarian cancer, head and neck cancer, etc.
Stage	Enhanced P1 ongoing (USA) , P1 ongoing (JP)
Out-licensed	Fujifilm

Fujifilm development code: FF-21101



Armed Ab labelled with radioisotope.

Binds to CDH3, which is often expressed on various cancers, and kills cancer cells with beta ray (RIT)



Accumulates on  
cancer only



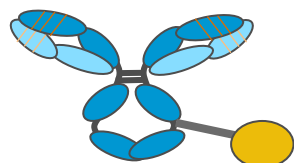
Known effect  
against cancer



**Patient-friendly anti-cancer drug**

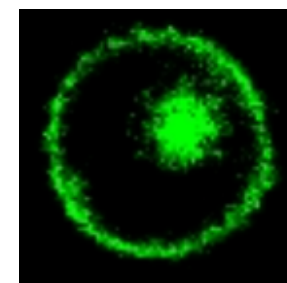
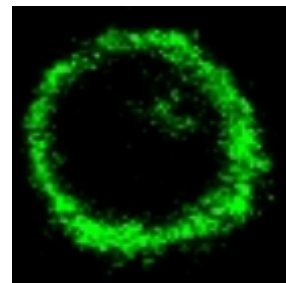
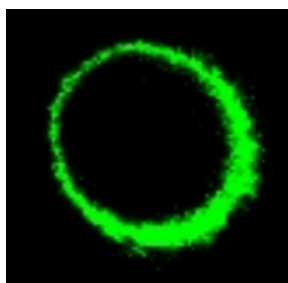
# PPMX-T004 : Ab-drug conjugate (ADC)

Code	PPMX-T004
Applicable disease	Solid tumor expressing CDH3
Stage	Drug discovery
Out-licensed	Fujifilm



Small molecule anti-cancer drug

Armed Ab labeled with small molecule anti-cancer drug.  
Designed to make cancer cells take up ADC so that the drug released inside may kill cancer cell.



Uptake of PPMX-T004 and drug into a cancer cell.  
Fluorescence-labelled PPMX-T004 and cancer stem cells derived from human are made coexist and are observed with a confocal microscope.

## 03 Financial results of first half FY2021

Develops seeds of academia through our technologies and provides for patients after licensing out

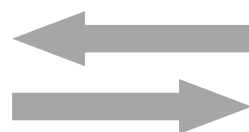
**Drug  
discovery**  
(Ab drugs)

**Ab creation,  
Contract  
research,  
Sequencing  
analysis**

Promotes research of academia, etc. through our technologies. Contributes to enhance network and sales.

**Academia**

Co-research, research support, sales of Abs/reagents



Research seeds, network

**Strong network**



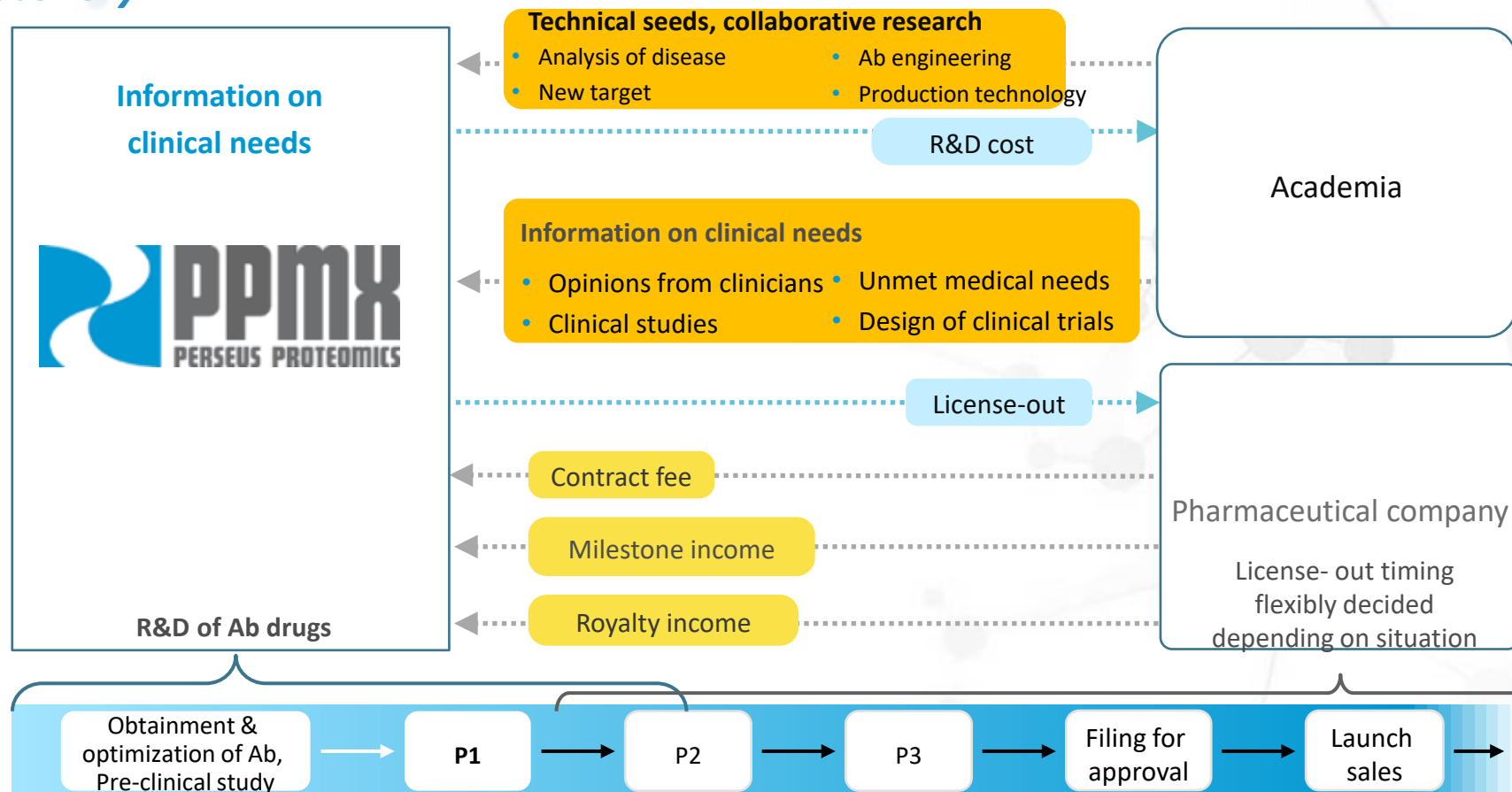
**Sales of  
Abs/reagents**

Provides nuclear receptor Abs, etc. for global researchers in life science, drugs and basic research on Abs field. A stable state of the selling line forecasted.

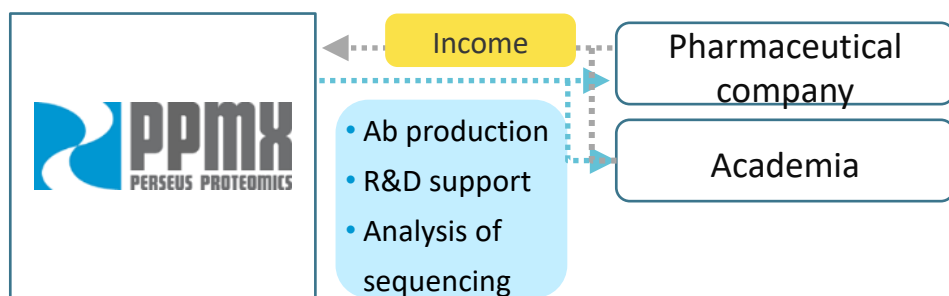


# Sales/Profit creating structure

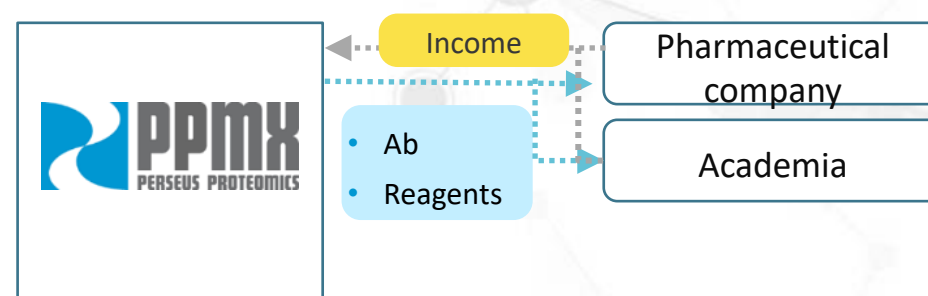
## 1. Drug discovery



## 2. Support of Ab research



## 3. Sales of Abs/reagents



# FY2021 first half business results and forecast

## ● Profit & loss

		(million yen)	
	FY2020 result	FY2021 1 <sup>st</sup> half	FY2021 forecast
<b>Sales</b>	<b>67</b>	<b>29</b>	<b>70</b>
<b>Gross profit</b>	64	29	65
R&D cost	313	154	411
Other	162	120	219
<b>SG &amp; A</b>	475	275	630
<b>Operating income</b>	<b>-411</b>	<b>-245</b>	<b>-564</b>
<b>Ordinary income</b>	<b>-410</b>	<b>-263</b>	<b>-583</b>
Extraordinary income	1	9	40
<b>Net income</b>	<b>-413</b>	<b>-274</b>	<b>-625</b>

Ab/reagent sales

PPMX-T003  
Other drug  
discovery

- Sales/Profit: Support research on Ab sales were weak, but Ab/reagent sales were almost as planned
- FY2021 forecast: No change in both business results and P1

# FY2021 first half financial status

## ● Balance sheet

Assets		
	2021/3/31	2021/9/30
Cash & deposits	1,069	3,506
Accounts receivable - trade	8	4
Other	31	29
<b>Total current assets</b>	<b>1,108</b>	<b>3,540</b>
<b>Non-current assets</b>	<b>9</b>	<b>9</b>
<b>Total assets</b>	<b>1,118</b>	<b>3,550</b>

(million yen)

Liabilities		
	2021/3/31	2021/9/30
<b>Current liabilities</b>	34	70
<b>Total liabilities</b>	<b>34</b>	<b>70</b>
Share capital	604	1,939
Capital surplus	889	2,225
Retained earnings	-413	-687
<b>Total shareholders' equity</b>	<b>1,080</b>	<b>3,477</b>
<b>Total net assets</b>	<b>1,083</b>	<b>3,479</b>
<b>Total liabilities and net assets</b>	<b>1,118</b>	<b>3,550</b>

- Cash & deposits, share capital, capital surplus: increased due to IPO
- Capital ratio: 97.9%

# About us

# Company name and corporate philosophy

## Company name



Perseus and Andromeda by François Lemoyne

Perseus

Hero in Greek myths

X

Proteomics

Study of structure and functions of protein

“Perseus Proteomics” represents the mission of the Company that we save patients (Andromeda) by fighting with refractory diseases including cancer (monster) through our antibody technology (Perseus’ weapon).

## Corporate philosophy

**We are committed to contributing to global medical care with our cutting-edge antibody technologies.**

# Company outline

## Business

- Develop Ab drugs
- Support research on Ab
- Sales of Abs/reagents

## Company name

Perseus Proteomics Inc.

## Established

February 2001

## Office

HQ : 4-7-6 Komaba, Meguro-ku, Tokyo, Japan  
Nagoya : 2-22-8 Chikusa-ku, Nagoya-shi, Aichi, Japan

## Capital

1,939 million yen

## Employee

22 (R&D: 16, Administration: 6) as of 30 Sep. 2021



## Directors



### **Takuya Yokokawa (President & CEO)**

Director, FUJIFILM Medical Drug Laboratories (drug discovery, research); Deputy Division Manager, FUJIFILM Medical Drug Div.(Clinical development, business development)



### **Shinichi Suzukawa**

Director-general, KDDI Global ICT; President, Telehouse Europe; President, overseas subsidiaries of KDDI



### **Tadashi Matsuura (Manager, R&D Dept)**

Faculty of Medicine, Shinshu University; Faculty of Medicine, Dartmouth University; National Institute of Bioscience and Human-Technology, Agency of Industrial Science and Technology, MITI



### **Kinichiro Kominami (CEO, Tech & Fin Strategy Co., Ltd)※**

Institute of Cancer Research, Nomura Securities, Mizuho Securities)



### **Toshikazu Ban (Executive Director, FUJIFILM) ※**

Takeda Pharmaceutical (Preclinical development), 15-year-experience of business development in USA



### **Nobuo Hanai (Outside director, Shimadzu Corporation)※**

Kyowa Kirin (President & CEO, Chairman)  
Development and license-out of Potelligent technology)

## Directors, Audit & Supervisory Committee Member



### **Kazuo Miwa (Standing Statutory Auditor)**

Head of East Asia Region and President of KDDI China, KDDI Global Business Division



### **Takao Hamakubo ※ (Professor, Nippon Medical School)**

Faculty of Medicine, Kyoto University; Ex-professor, Research Center for Advanced Science and Technology, University of Tokyo; Doctor; our founder



### **Tadashi Horiuchi (Professor, Clinical Research Promotion Center, Keio-gijuku University Hospital) ※**

Chief Manager, Daiichi Sankyo Drug Discovery Development Research Center, Standing Statutory Auditor, Asubio Pharma)



### **Takashi Ohno ※**

Representative, Ohno Certified Public Accountant's Office

## Technical Advisors

### **Fuyuki Ishikawa**

Professor, Graduate School of Biostudies, Kyoto University;  
Member, Science Council of Japan

### **Kohei Tsumoto**

Professor, Graduate School of Engineering, University of Tokyo;  
Director, Center for Drug Design Research, National Institutes of Biomedical Innovation, Health and Nutrition

# History

Ab drugs development  
1975  
Hybridoma method invented

Ab drug as anti-cancer drug  
1998  
Herceptin, 1<sup>st</sup> monoclonal Ab approved

Humanised Ab drug  
1990  
Phage display method proposed

2000  
Mylotarg, 1<sup>st</sup> Ab drug conjugate approved

2002  
Humira, 1<sup>st</sup> drug by phage display method approved; Zevalin, 1<sup>st</sup> RIT approved

2009  
Removab, 1<sup>st</sup> bispecific approved

Many of Abs against target proteins expressed on cancer cell surface already developed (Abs against new targets or hard to obtain are left.)

2000.4  
“Drug discovery against targets found by analysis of human genome project” starts at LSBM at U of T starts

2001.2  
Established

2005.9  
Sales of Ab against 48 nuclear receptors starts

Humanised Ab  
2006.9  
PPMX-T001  
Licensed out to Chugai Pharmaceutical

2008.9  
PTX3  
Sales of ELISA kit starts

2009.1  
FUJIFILM becomes parent company; stock holding ratio : 77%

Chimeric Ab  
2011.1  
PPMX-T002  
licensed out to FUJIFILM

Human Ab by phage display method  
2014.12  
PPMX-T003  
selected as JST drug discovery project (940 M yen)

Humanised Ab  
2015.9  
PPMX-T004  
licensed out to FUJIFILM

2018.3  
FUJIFILM becomes other related company; stock holding ratio : 49%

2019.1  
Nagoya Laboratory opens

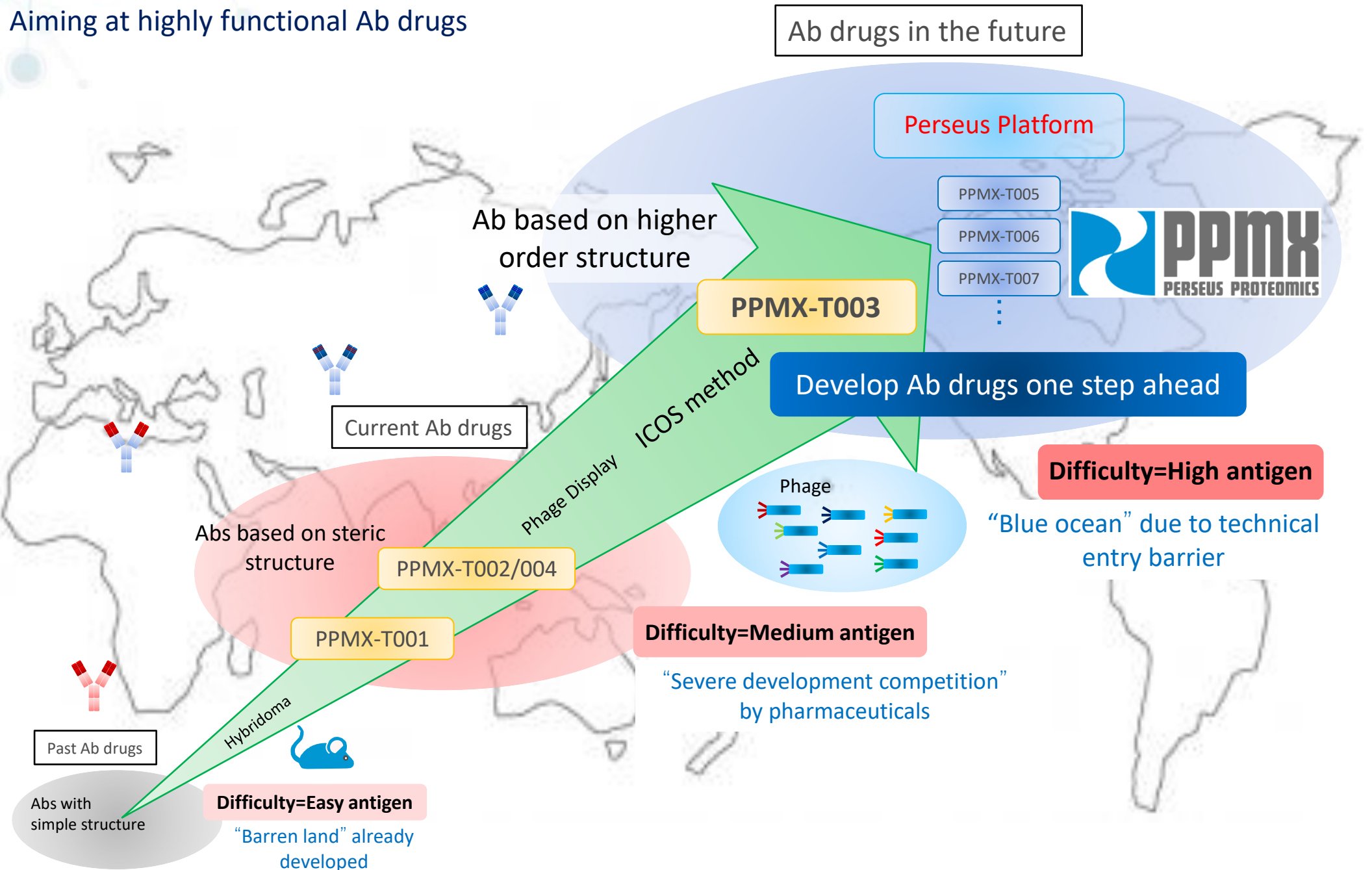
2019.11  
PPMX-T003  
In-house  
P1 starts (JP)

2020.4  
PPMX-T002  
FUJIFILM  
P1 starts(JP)

2021.6  
Listed in Mothers TSE

# Bring more Ab drugs to patients

Aiming at highly functional Ab drugs



Perseus Proteomics Inc.

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