

# Ascentage Pharma Group

Advancing Therapies That  
Restore Apoptosis

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# Ascentage: Innovative Science

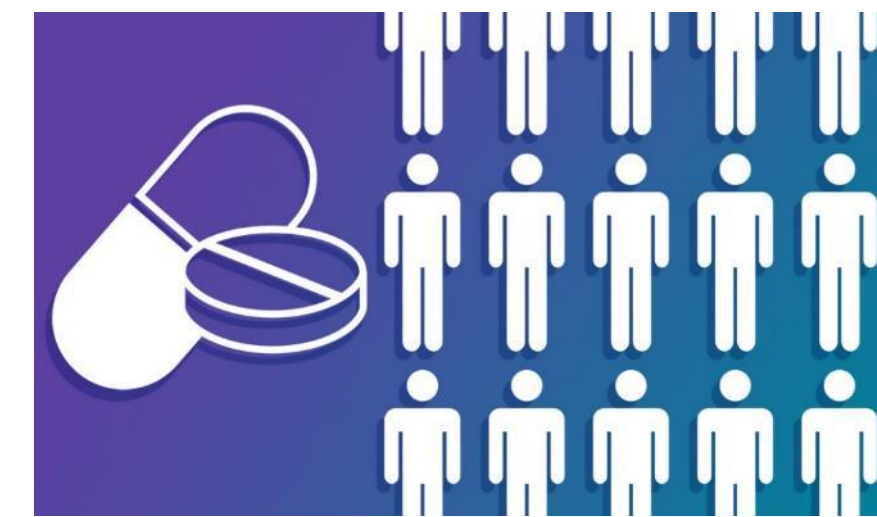
Proprietary PPI Platform delivering first-and/or best-in-class potential drugs

## BREAKTHROUGH SCIENCE



**80** ISSUED PATENTS  
**300+** PENDING APPLICATIONS  
**90+** PUBLICATIONS

## STRONG PIPELINE



**12** NOVEL COMPOUNDS  
**24** INDS  
**40+** CLINICAL TRIALS  
**10+** INDICATIONS

## DEDICATED TEAM



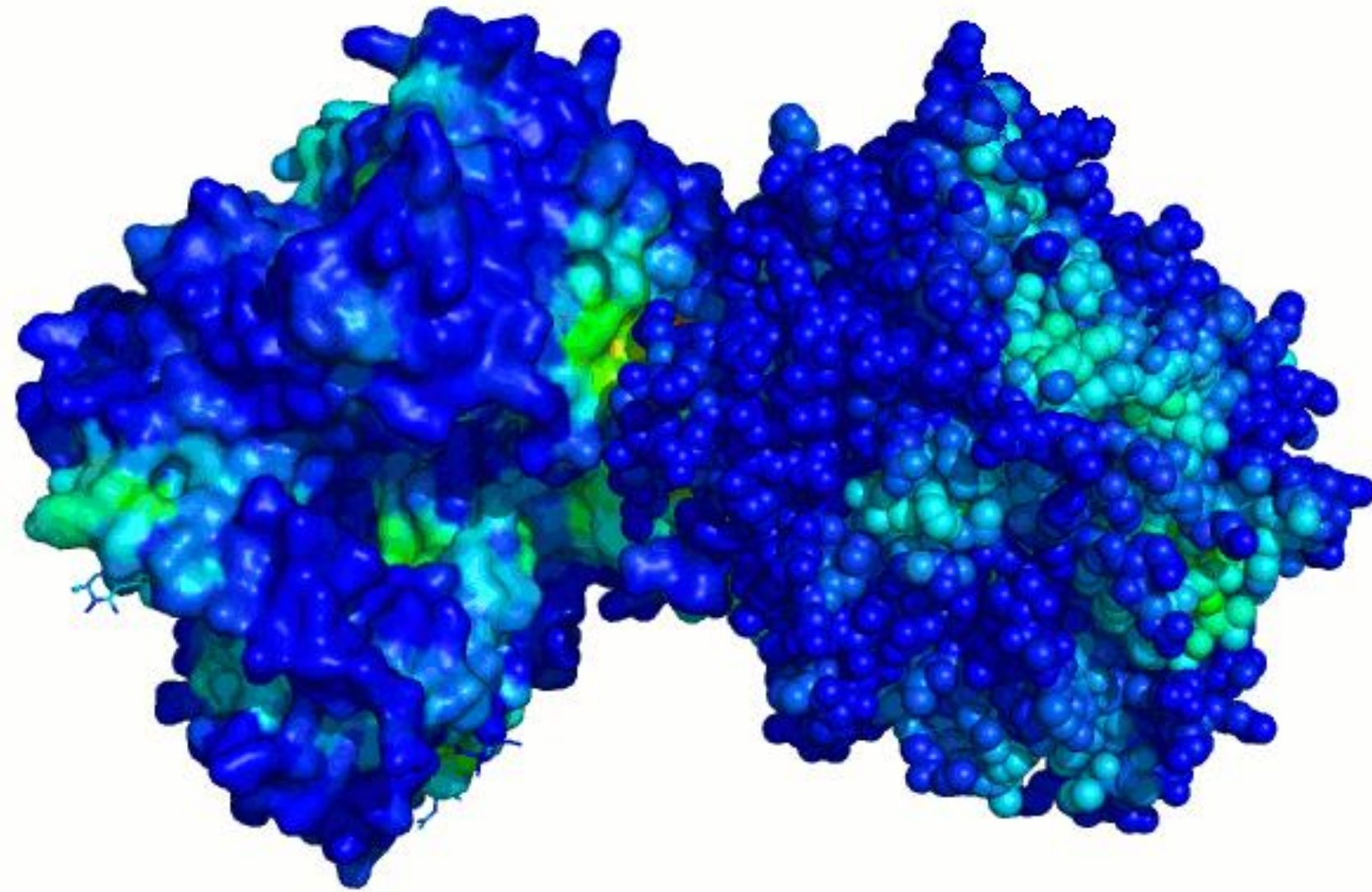
**1** VISION: BUILDING A GLOBAL BIOTECH COMPANY  
**20+** YEARS' COMMITMENT OF EXECUTIVE TEAM  
**400+** EMPLOYEES

## GLOBAL DEVELOPMENT



INTEGRATED ORGANIZATION IN **CHINA, UNITED STATES** AND **AUSTRALIA**

# Global Leader Developing Protein-Protein Interactions Drugs

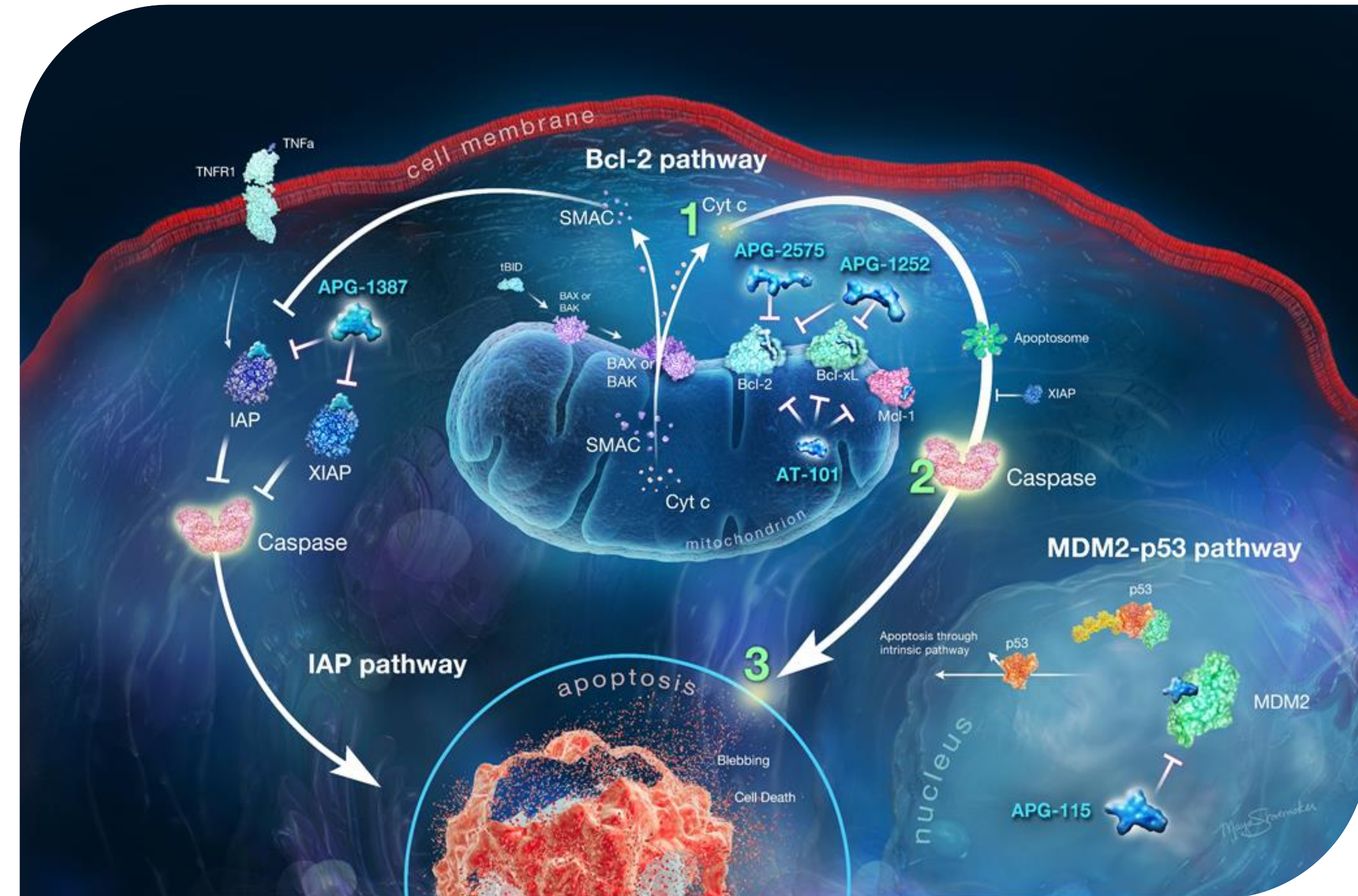


- **Protein-protein interactions** (PPIs) play a crucial role in cellular processes, and are implicated in many diseases, from cancer to viral infections
- **PPI** targets can't be penetrated by large molecules, leaving small molecules the only viable choice for drug development
- **PPIs** have broad, shallow, relatively featureless binding sites, hence historically “**difficult to drug**”. There is only one PPI-targeting drug approved in oncology, venetoclax

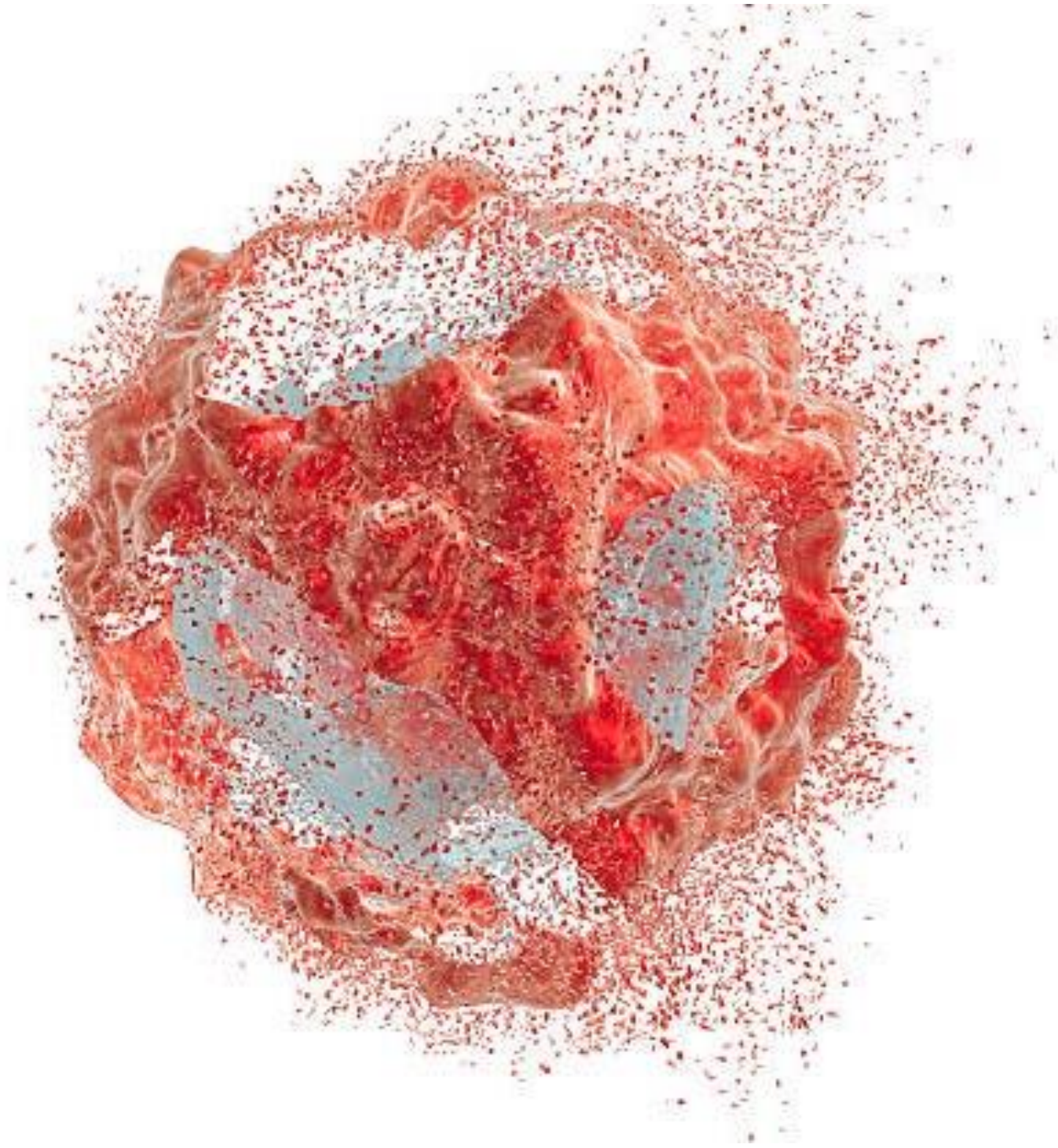
# Focused on Apoptosis

**Apoptosis** (programmed cell death) is an essential biological process. The average adult human loses between **50 to 70 billion** cells each day due to apoptosis.

- Apoptosis plays a **crucial role** in developing and maintaining the health of the body by eliminating old and unhealthy cells.
- When apoptosis doesn't function correctly, cells that should be eliminated persist or become immortal causing **cancer** and **leukemia**.
- Ascentage has discovered **four** potentially **first- or best-in-class** candidates targeting three distinct classes of PPIs.



# 2020 H1 Key Achievements



- Submitted NDA for **HQP1351** in patients with T315I-mutant CP-CML and AP-CML in China in June 2020
- Started **4** new Phase Ib/II studies of APG-2575 both in China and U.S. APG-115, APG-1387 and APG-1252 entered into phase Ib/II studies respectively.
- Obtained **2** orphan drug designation from U.S FDA for HQP1351 in TKI resistant CML and APG-2575 in WM. Obtained **1** fast track designation from U.S FDA for HQP1351
- Entered **2** global clinical collaborations: APG-2575 with CALQUENCE® (acalabrutinib) in r/r CLL/SLL and APG-115 with KEYTRUDA® (pembrolizumab) in advanced solid tumors

# 12 Month Clinical Milestones

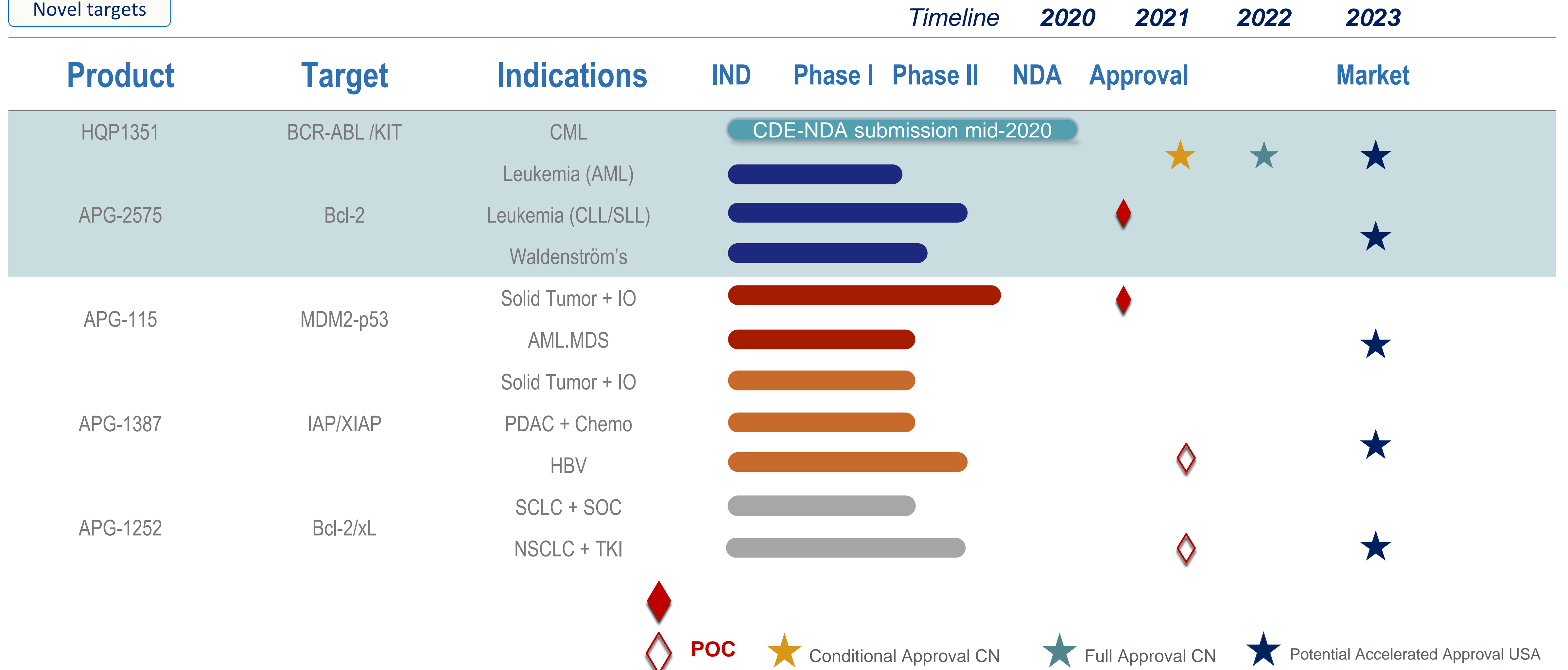
- Launch of 3rd Bcr-Abl inhibitor **HQP1351** in China
- Bcl-2 selective inhibitor **APG-2575** in Ph II r/rCLL reaches POC
- MDM2-p53 **APG-115** + Keytruda® reaches POC in targeting checkpoint resistant/relapsed NSCLC or melanoma patients
- IAP/XIAP Dimer **APG-1387** + Keytruda® reaches Ph II POC study targeting checkpoint resistant/relapsed NSCLC patients
- IAP/XIAP Dimer **APG-1387** reaches Ph II POC targeting Chronic Hepatitis B



# High Value Portfolio Opportunities

Validated targets

Novel targets



POC



Conditional Approval CN



Full Approval CN

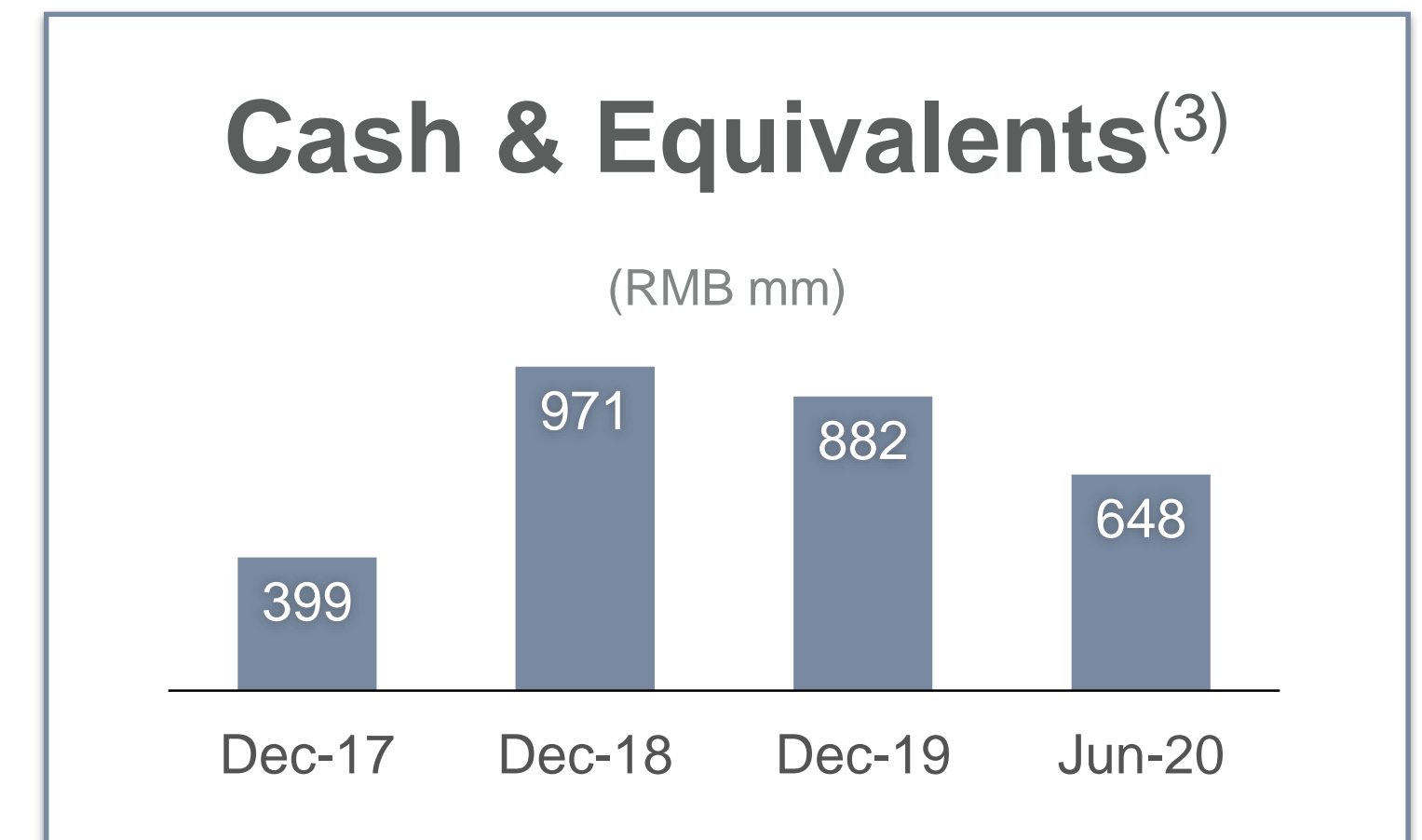
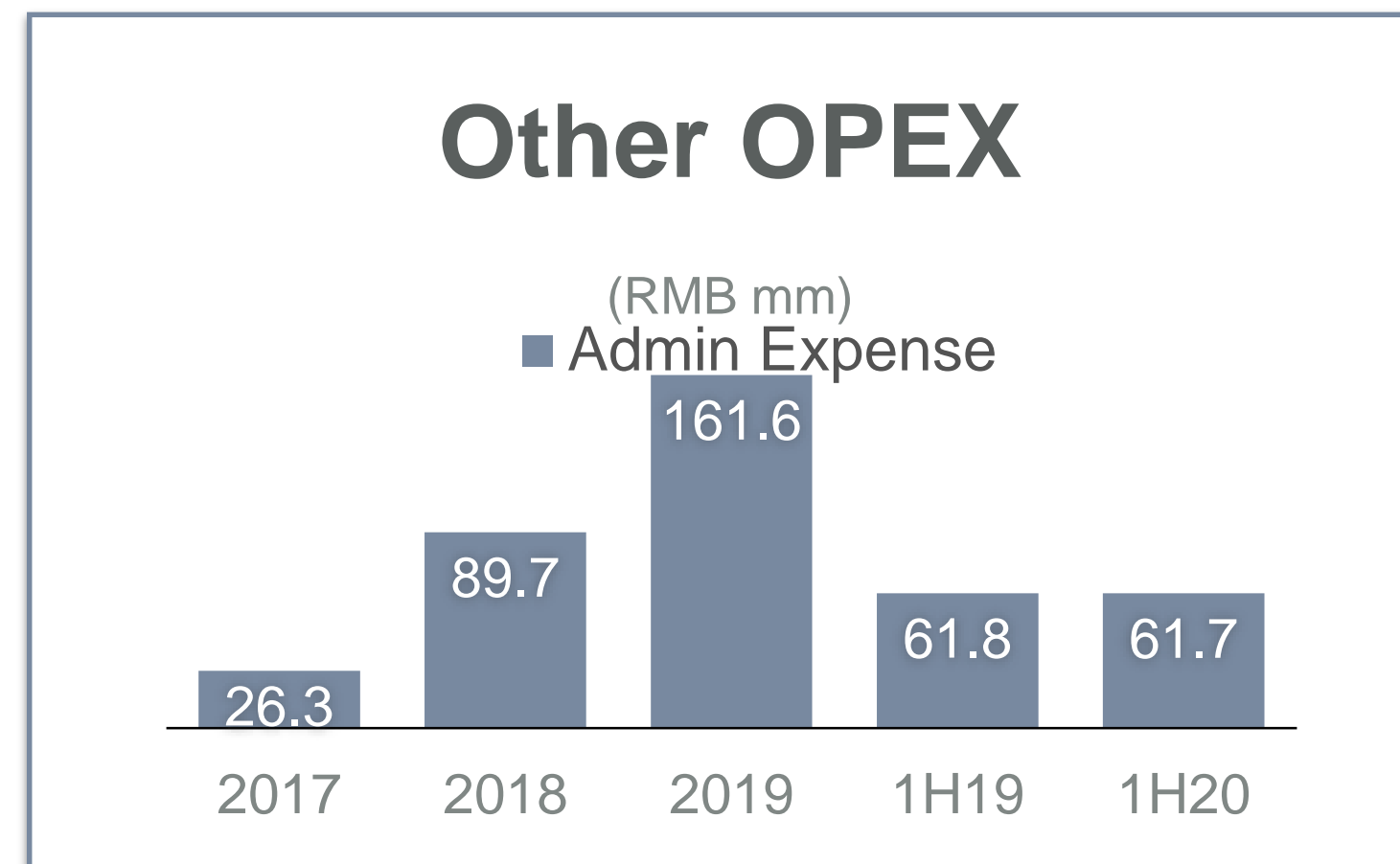
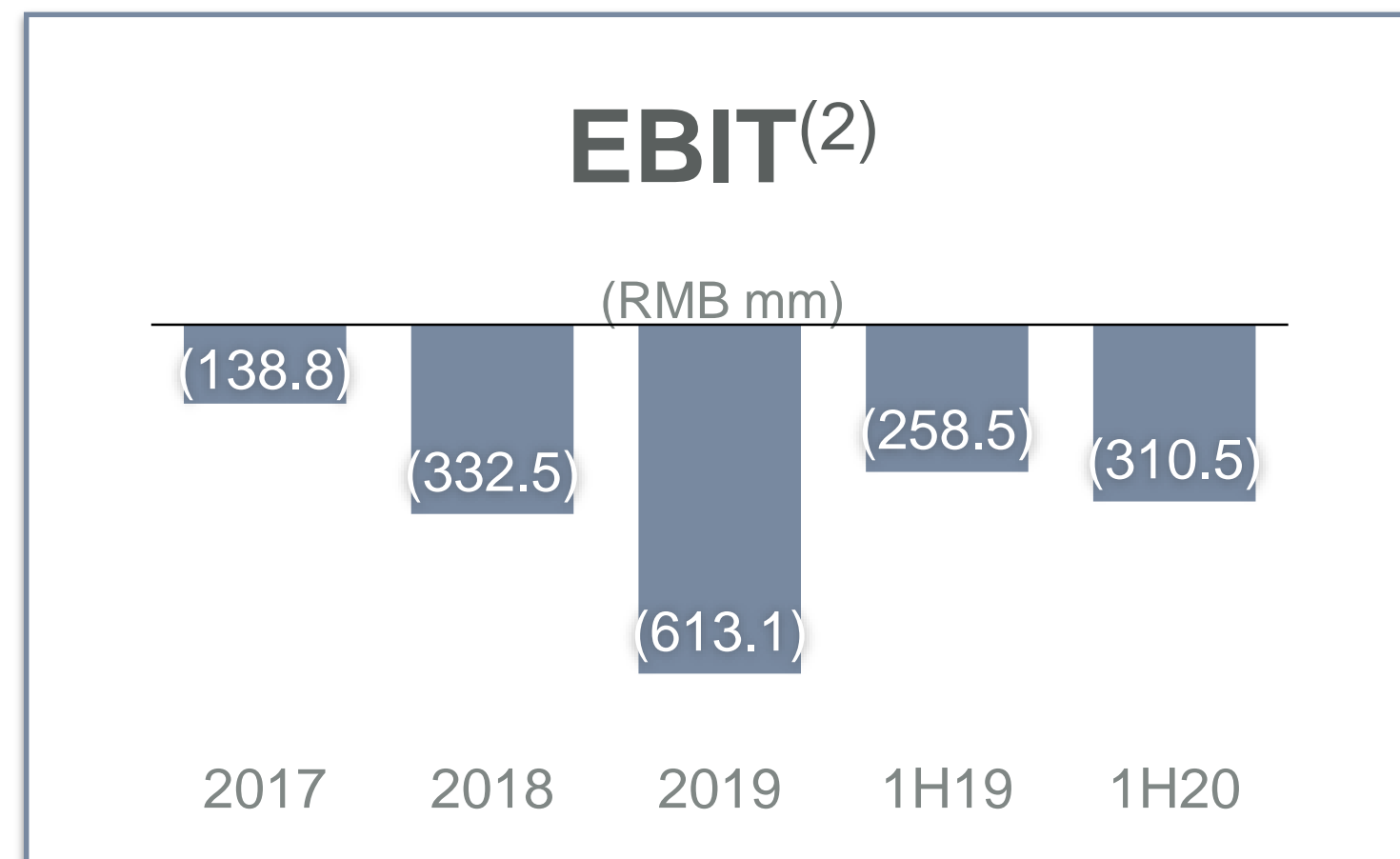
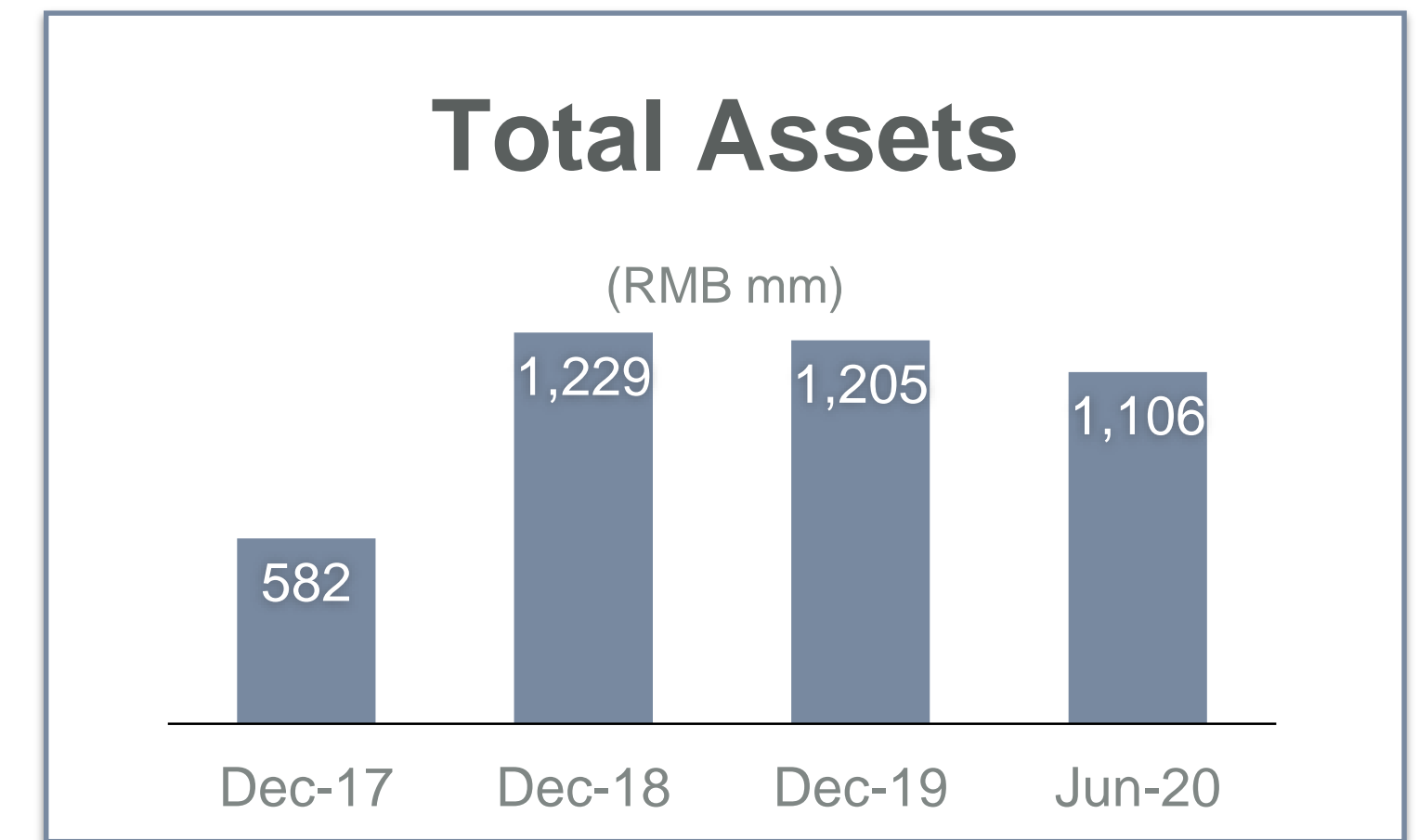
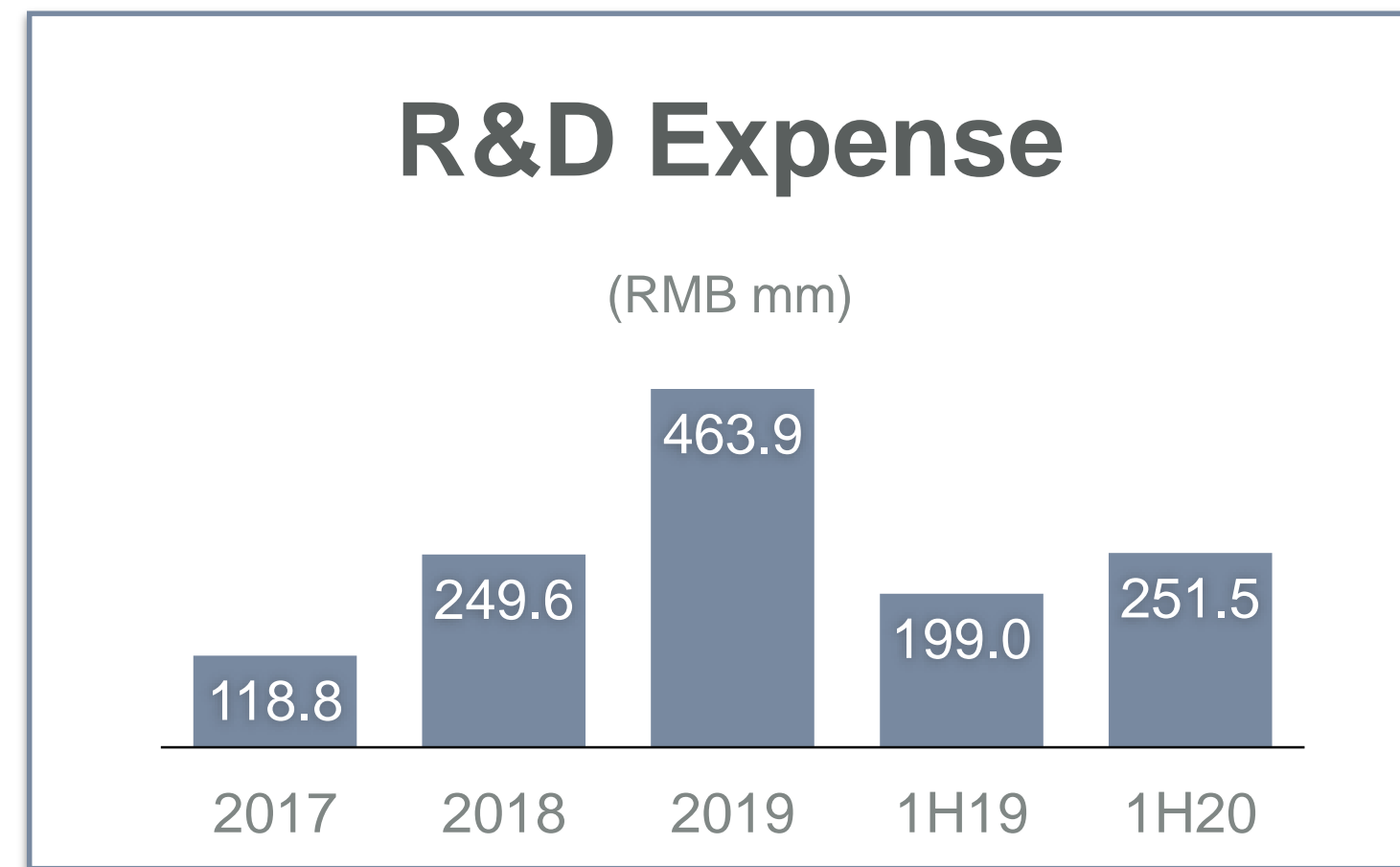
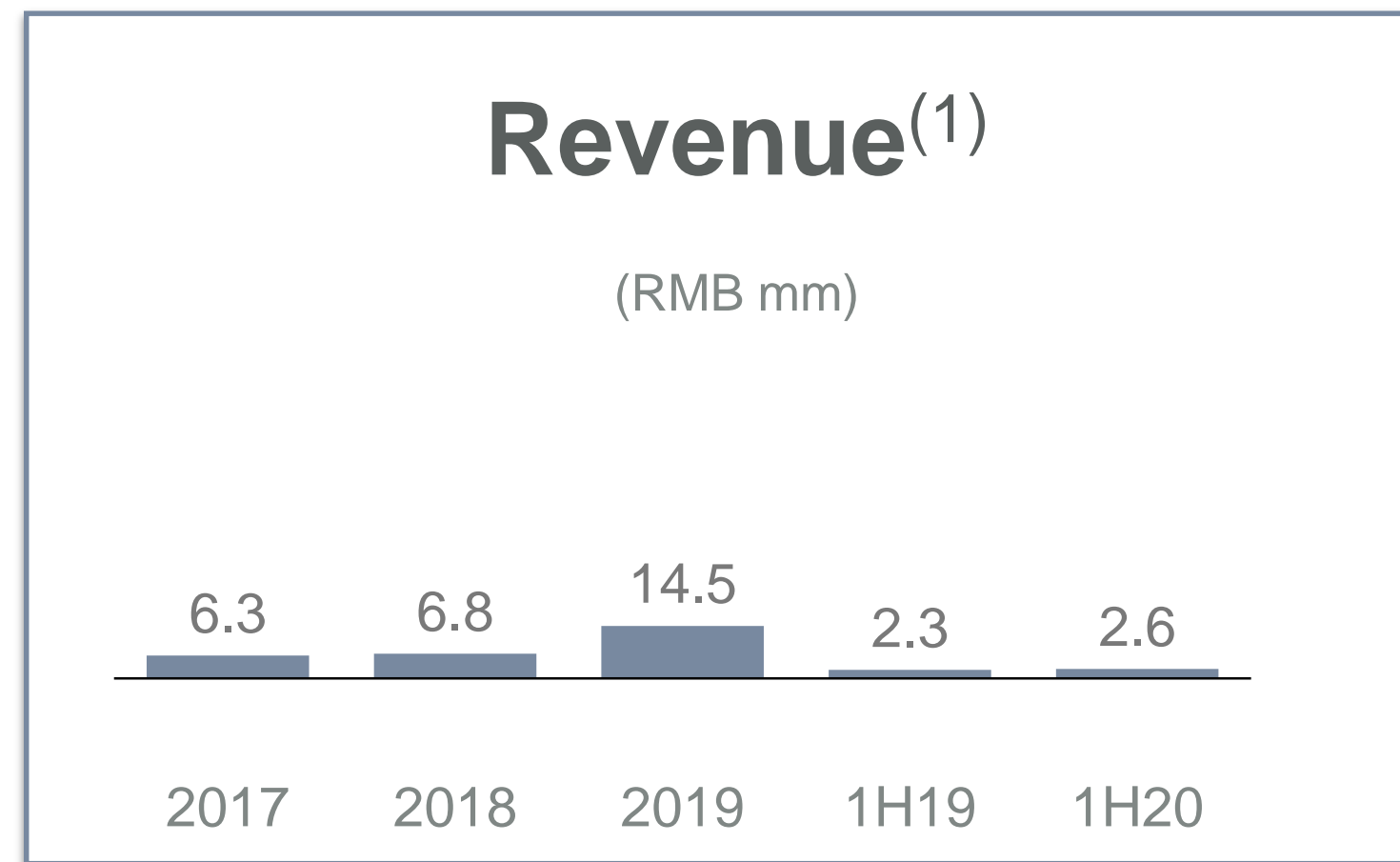


Potential Accelerated Approval USA

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Source: Company data Note: All data as of December 31, 2019

# Key Financial Highlights

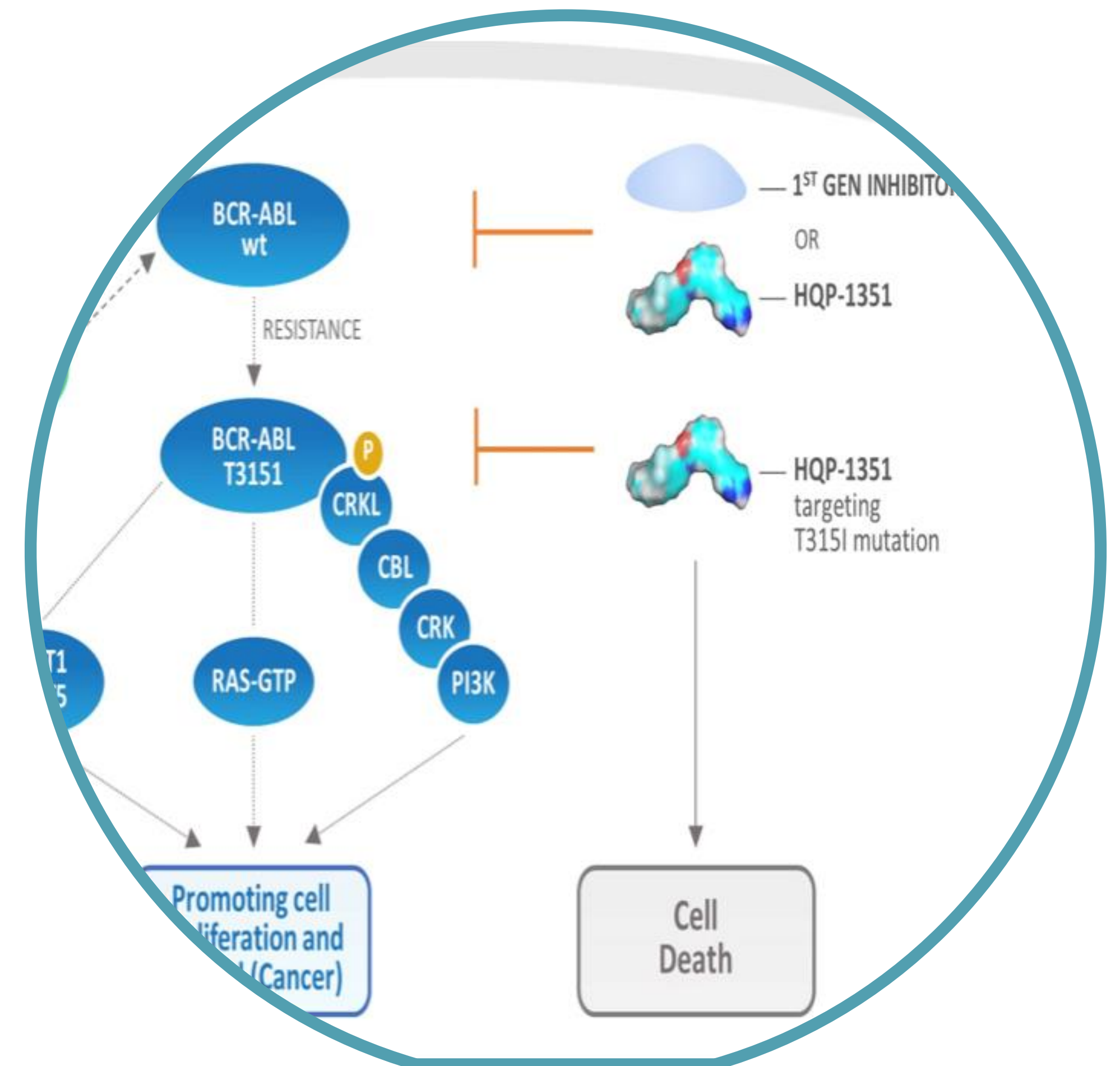


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1) its revenue from provision of research and development services, and compounds library and intellectual property license fee income; 2) EBIT = Gross Profit – R&D Expense – Other OPEX 3) Cash & Equivalents include cash and bank balances, and other financial assets, which represent mainly investment in short-term financial productsThe group derives

# HQP1351 Olverembatinib Overview

3<sup>rd</sup> Gen BCR-ABL/KIT  
Multi-kinase Inhibitor

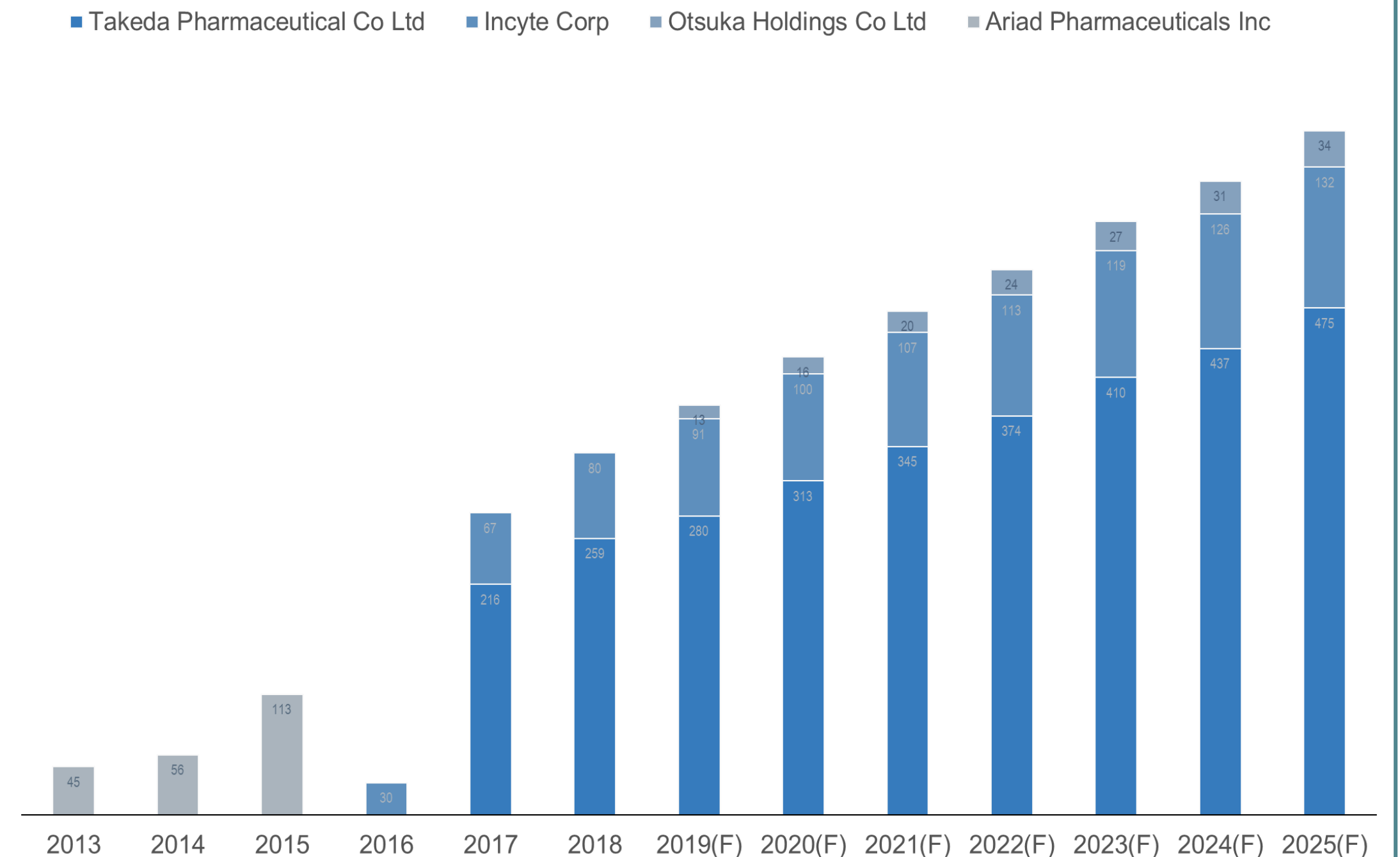


# BCR-ABL a Validated Target

## Unmet Needs for Targeting BCR-ABL

- One of the most frequent BCR-ABL mutations is T315I, ranging from 5 to 25% of CML cases
- Amongst multiple BCR-ABL mutations T315I is also the deadliest mutation; it is resistant to second generation TKIs too (i.e. dasatinib, nilotinib)
- Until now, only ponatinib has been able to overcome TKI-resistance

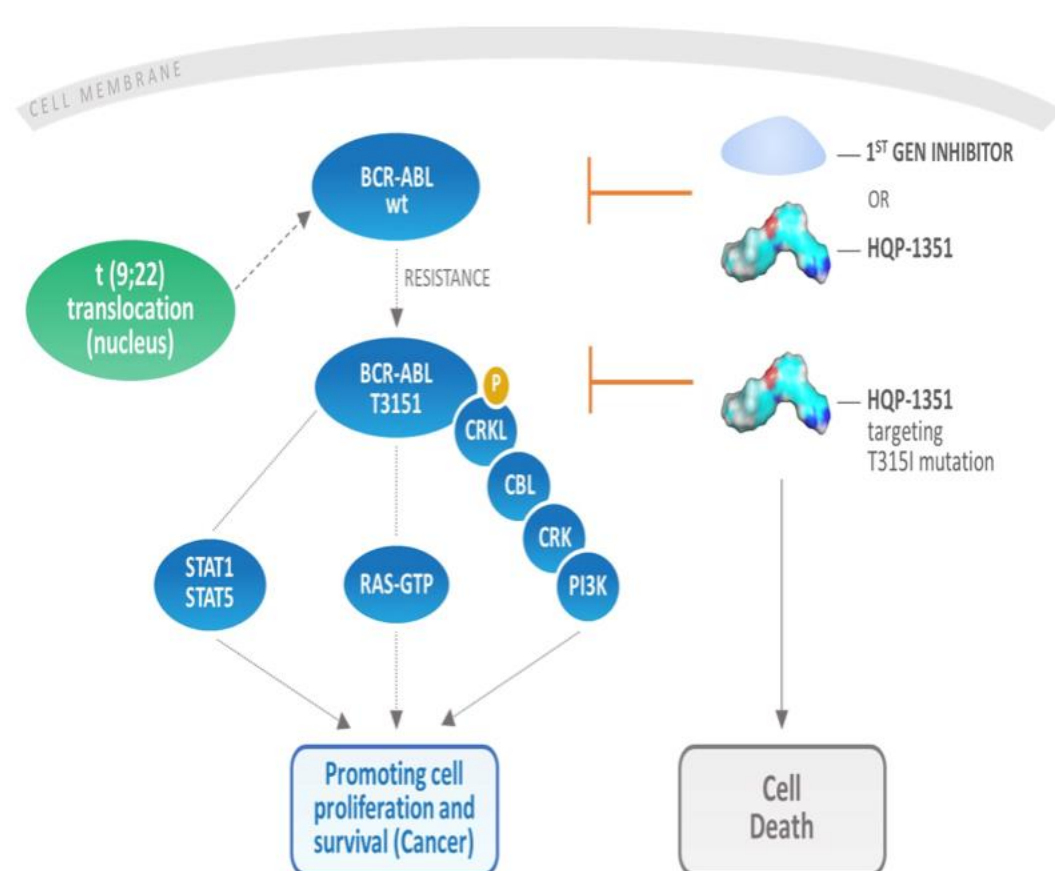
## Global sales of ponatinib forecasted at \$641M



# HQP1351 Olverembatinib

3<sup>rd</sup> Gen BCR-ABL/KIT  
Multi-kinase Inhibitor  
Targets TKI resistant mutations

## BCR-ABL/KIT TKI for mT315I

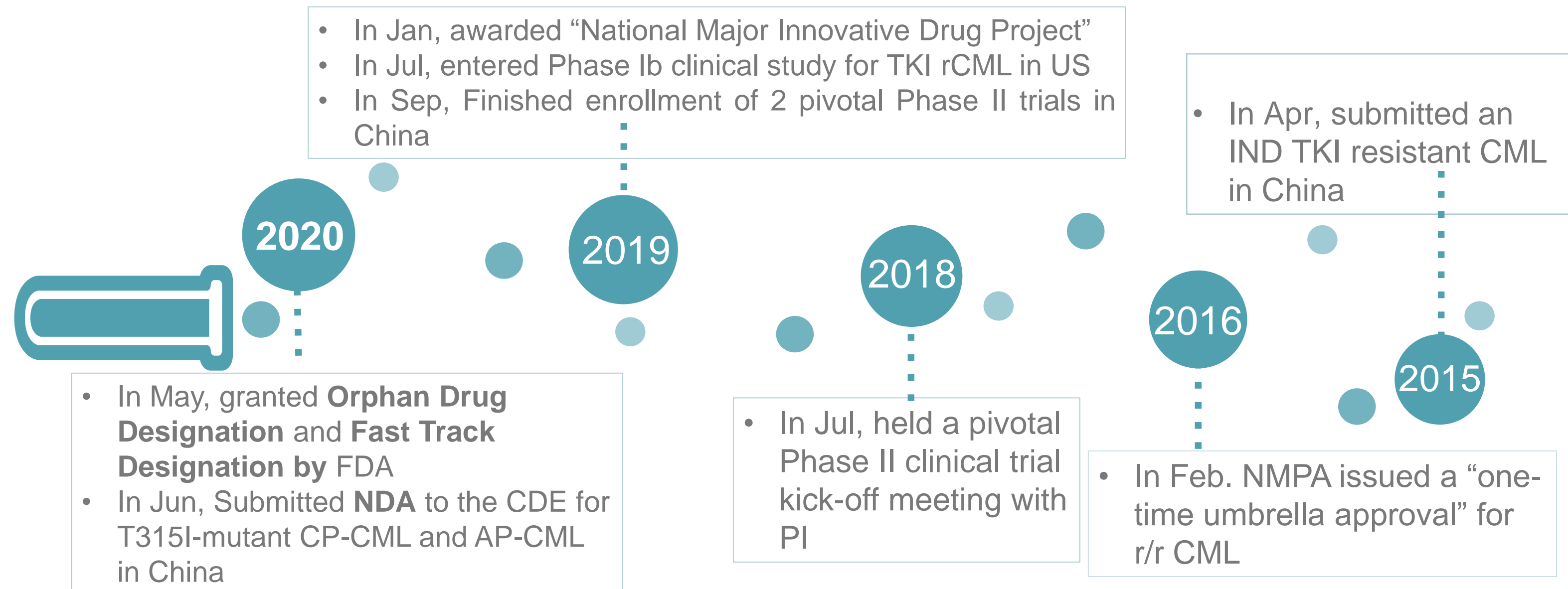


Source: Company data

## Development Milestones

- Submitted NDA to the CDE in China in July 2020
- **Fast Track Designation** approved by FDA for CML in April 2020
- **Orphan Drug Designation** approved for CML in April 2020
- **Ph Ib bridging trial** in US enrolling patients at MD Anderson Cancer Center
- Results of Ph I trial of HQP1351 in CP/AP TKI resistant / intolerant CML were orally presented on ASH 2018 and 2019; nominated as “Best of ASH” in 2019

## Milestones & Developments



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# Well-tolerated with minimal dose interruptions

## Ph I: HQP1351 is well-tolerated

- 92 out of 101 patients have finished three cycles of treatment:
  - Longest duration of treatment is 45 months
  - The average observation period for the Ph I clinical trial is more than 1 year
  - 2 out of 101 patients has discontinued treatment due to AEs
- Most treatment-related AEs were mild or moderate**
- Grade 3 or 4 thrombocytopenia reported in HQP1351 treated patients, consistent with other TKIs
- No** cardiovascular, cerebrovascular, or peripheral vascular thrombosis, fatal myocardial infarction or stroke was reported, compared to serious arterial occlusion cases observed in 35% of ponatinib treated patients in clinical trials
- The liver toxicity was rarely reported and was mild or moderate, compared to ALT or AST elevation observed in 56% (all grade) and 8% (grade 3 or 4) of patients treated with ponatinib

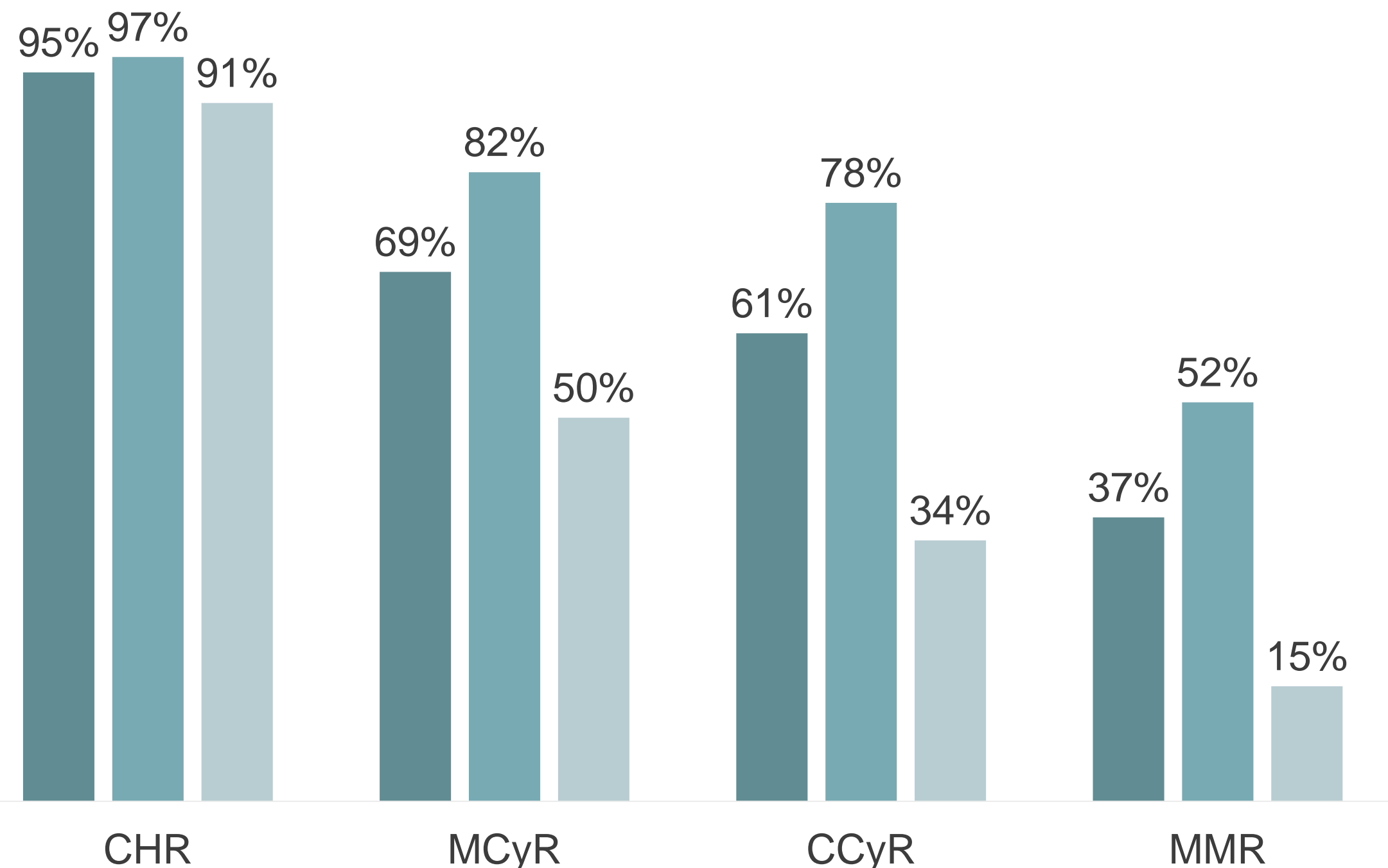
### Summary of all Grade 3 | 4 AEs and SAEs in overall subjects

AE (>10% of Patients)	Grade 3, 4 n(%)	SAE n (%)
Thrombocytopenia	50 (49.5)	6 (5.9)
Leukopenia	20 (19.8)	0 (0)
Anemia	12 (11.9)	2 (2)
Hypertriglyceridemia	8 (7.9)	0 (0)
ALT elevation	2 (2)	0 (0)
AST elevation	3 (3)	0 (0)
Hyperbilirubinemia	1 (1)	0 (0)
Proteinuria	5 (5)	0 (0)
CPK elevation	2 (2)	0 (0)
Pyrexia	7 (6.9)	1 (1)
Rash	2 (2)	0 (0)
Skin Mass	1(1)	0 (0)

# Responses in Total Patients

CP

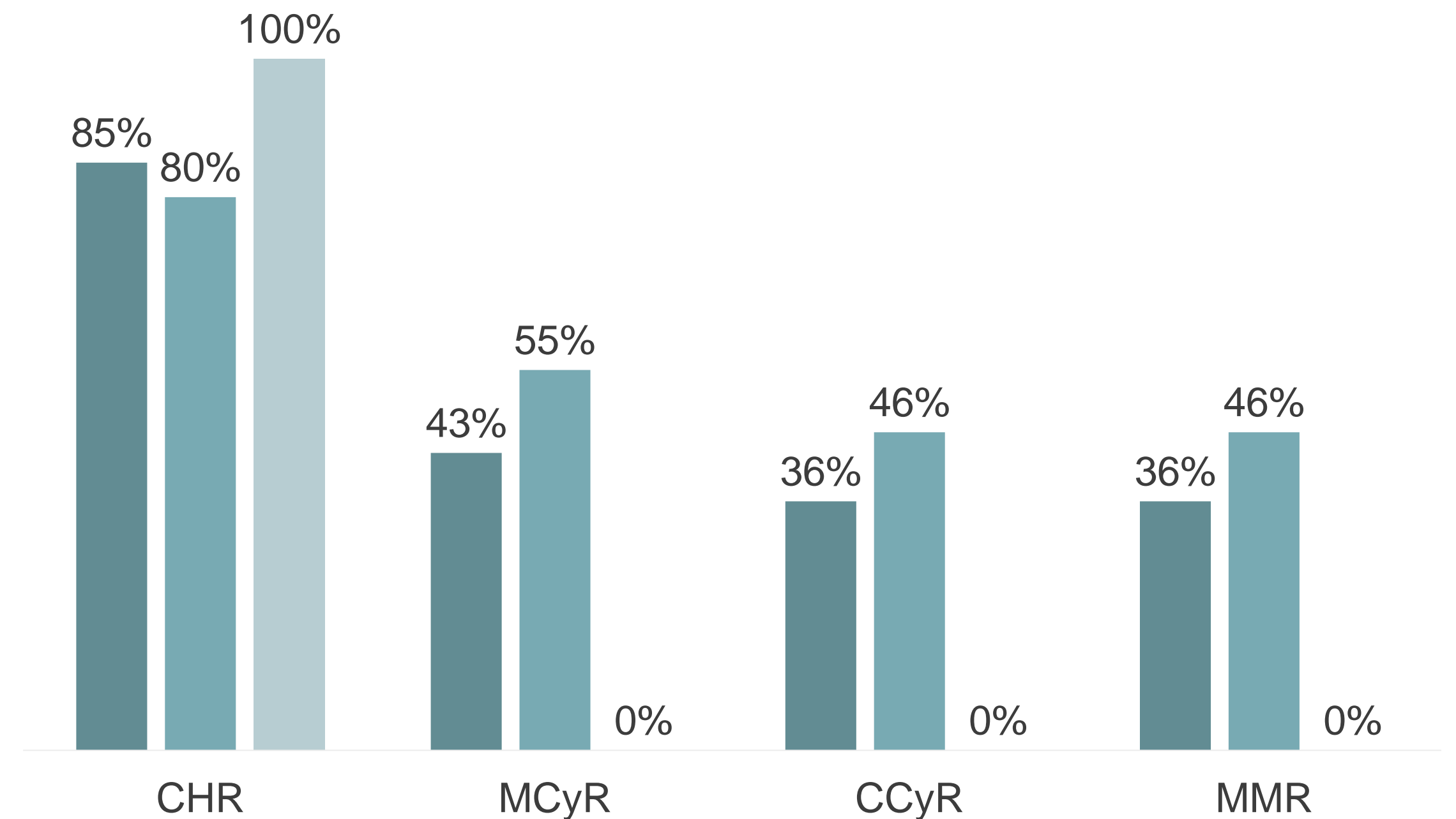
■ Total n=87 ■ T315I+ n=52 ■ T315I- n=35



CML Response Criteria: Complete Hematological Response(CHR),  
Bone Marrow; Major Cytogenic Response (MCyR\*) Complete Cytogenic Response (CCyR),  
 Major Molecular Response (MMR<sup>^</sup>) | \* MCyR is a validated End Point, ^ MMR defined by PCR (<1/1000)

AP

■ Total n=14 ■ T315I+ n=11 ■ T315I- n=3

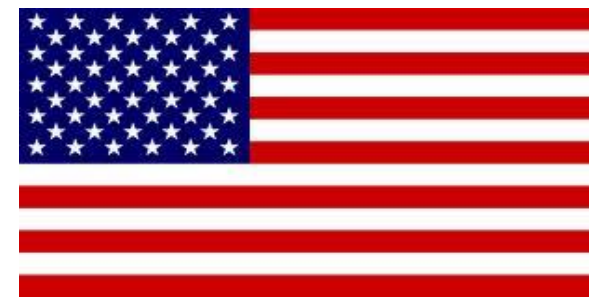


CML Response Criteria: Complete Hematological Response(CHR),  
Bone Marrow; Major Cytogenic Response (MCyR\*) Complete Cytogenic Response (CCyR),  
 Major Molecular Response (MMR<sup>^</sup>) | \* MCyR is a validated End Point, ^ MMR defined by PCR (<1/1000)

# CML Patient Numbers

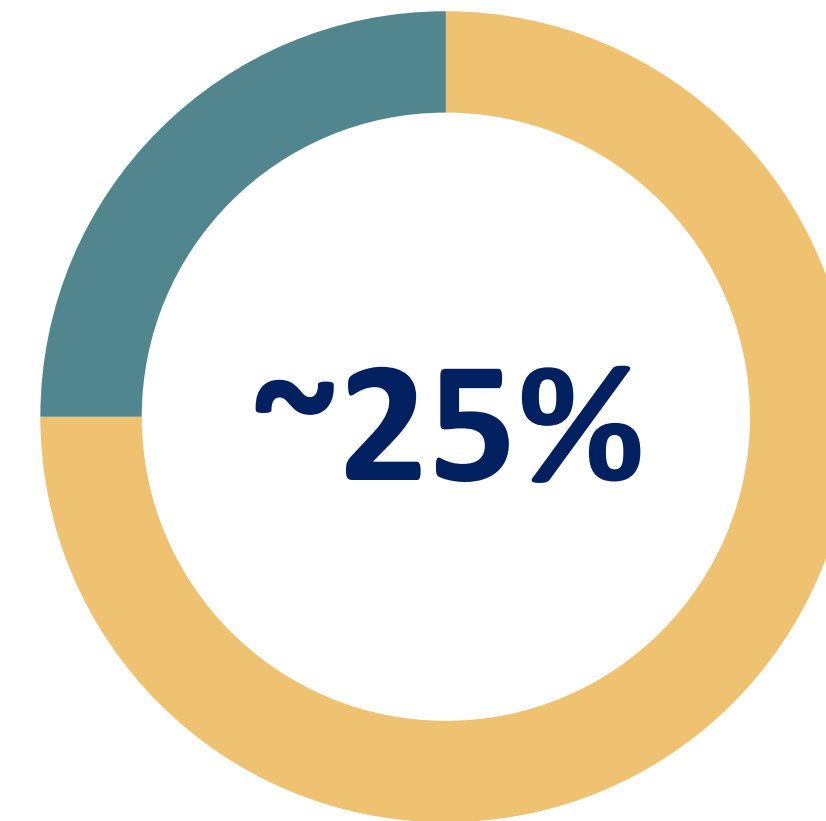
**51,000+**

CML patients in US



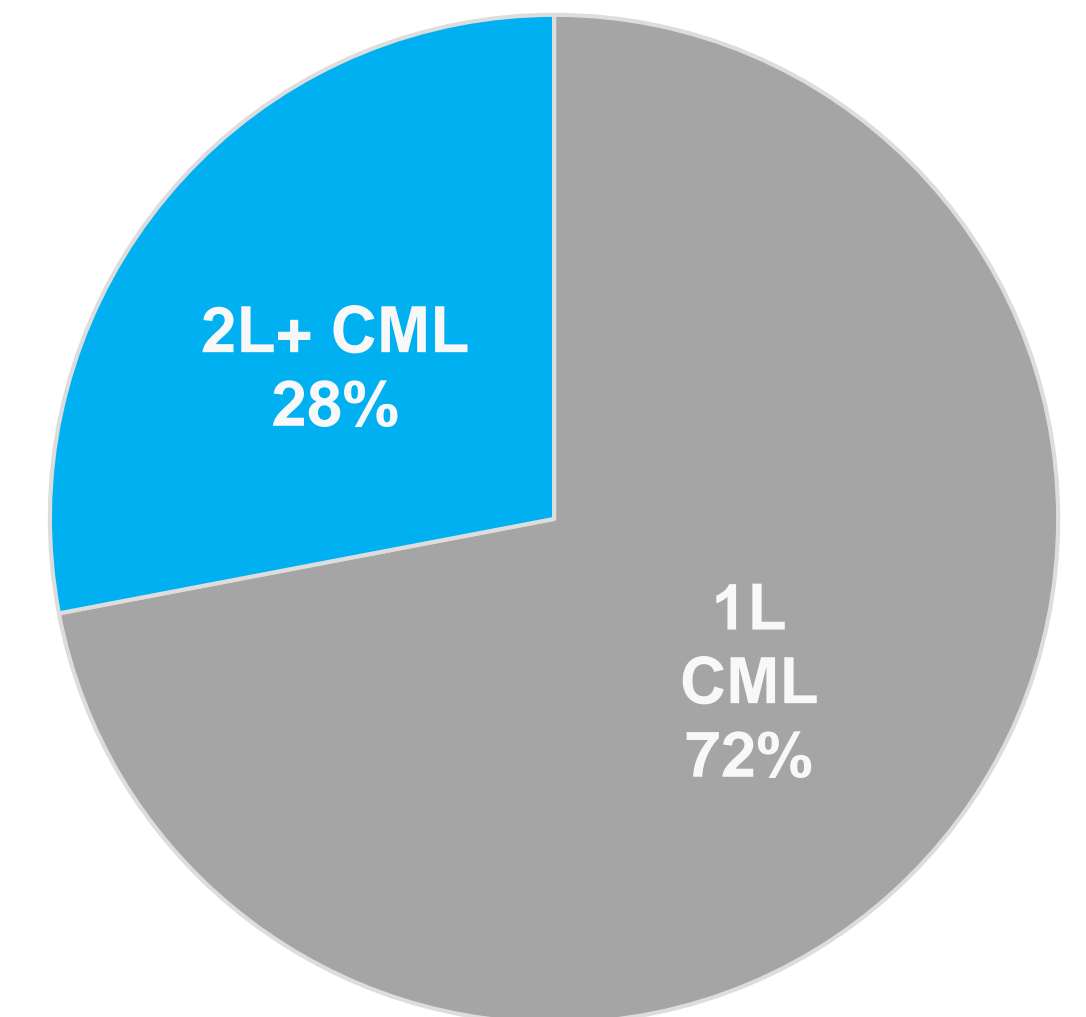
**75,000+**

CML patients in China



Over 25% of patients with **BCR-ABL-mutated CML** have the T315I mutation<sup>2</sup>, which has been associated with **resistance to treatment and poor outcomes**<sup>3</sup>

## China's CML patient by lines of treatment



CML

**\$5.5B**  
Market

**33.3K**  
Incidence

**135K**  
Prevalence

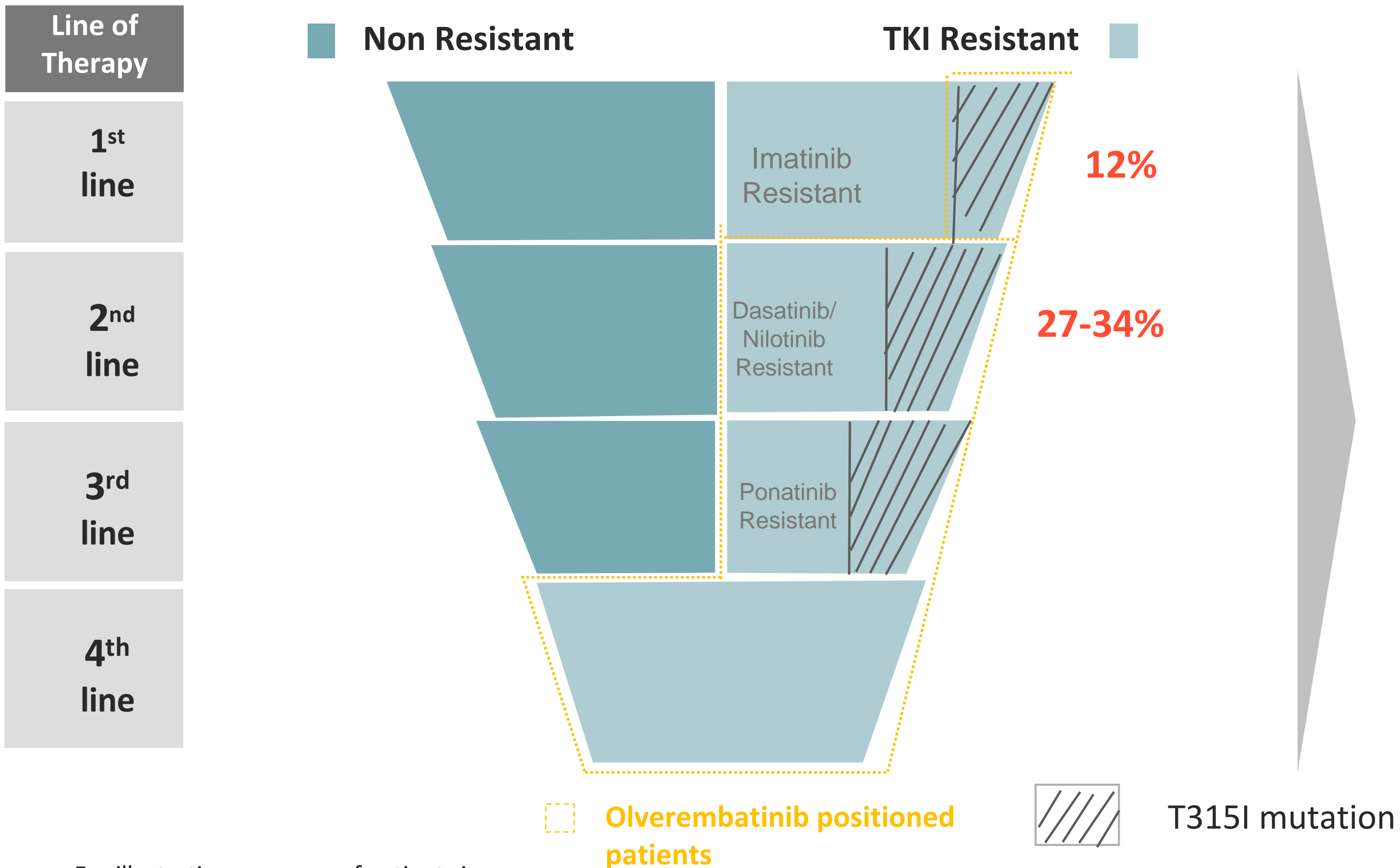
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Source: 1. Frost & Sullivan 2. My Cancer Genome 2014. 3. Nicolini, et al. Leukemia 2006;20:1061-6, Global Data, DRG.

# China CML TKI-Resistance Patient Pool

CML patients treated with TKI may have resistance

TKI resistance & T315I mut share

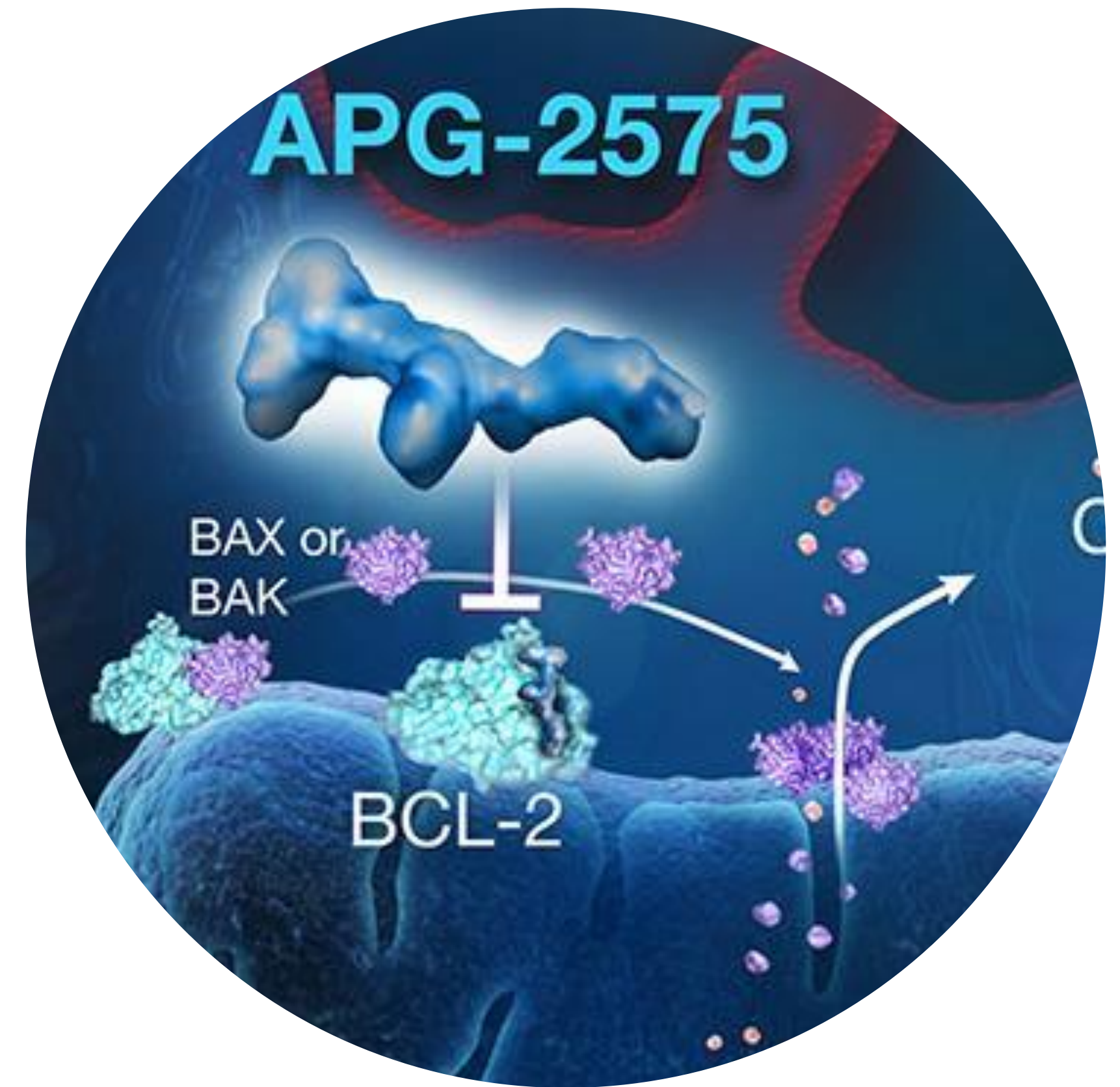


For illustrative purposes of patient size

- In a Chinese review analysis, 52.7%, 21.8%, 25.5% cases experienced resistance to imatinib, nilotinib and dasatinib<sup>1</sup>
- Over **half** of imatinib-, nilotinib-, and dasatinib-resistant cases developed BCR-ABL mutation<sup>1</sup>
- **T315I mutation** was the most frequent mutation detected in imatinib-, nilotinib-, and dasatinib-resistant cases, accounting for 12.3%, 27.3%, and 34.1%<sup>1</sup>

# APG-2575 Overview

BCL-2 Selective Inhibitors





# BCL-2 is a Validated Target

## BCL-2 inhibitor



### Bcl-2 Selective Inhibitors

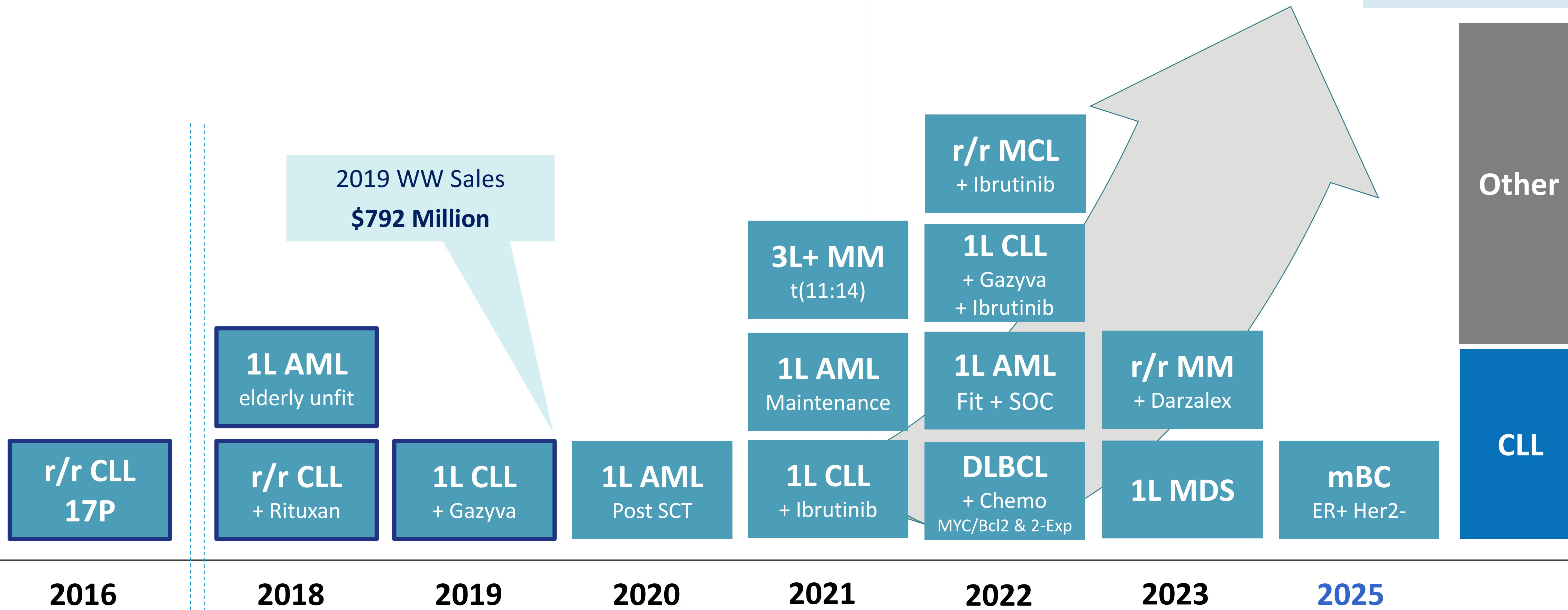
		
Compound	APG-2575	Venetoclax (ABT-199)
MOA	Orally available and Bcl-2 selective inhibitor	Orally available and Bcl-2 selective inhibitor
Clinical stage	Ph Ib/II	Marketed (CLL, AML)
Indication	CLL, AML, WM, MM, T-PLL	CLL, AML, MM, MCL, MDS, NHL, ALL, Breast cancer, Prostate cancer
Combo agents	BTK, CD20, MDM2, BCR-ABL TKI	BTK, CD20, CDK9, Pi3K, MDM2, JAK, PD-(L)1, FLT-3, IDH, CD33, CD38, etc.
Comments	<ul style="list-style-type: none"> <li>• Patient-friendly daily dose-ramp-up</li> <li>• No or Low TLS</li> <li>• Less risk DDI</li> <li>• Less neutropenia likely</li> <li>• Strong synergy with in-house MDM2-p53 inhibitor APG-115</li> <li>• Plan to focus on the China market</li> </ul>	<ul style="list-style-type: none"> <li>• NDA approved in April 2016</li> <li>• First-in-class Bcl-2 inhibitor</li> <li>• 5 FDA Breakthrough Therapy designations</li> <li>• 4 approved indications across CLL and AML populations</li> <li>• 250+ trials across US, China, EU, Japan, etc.</li> <li>• Enrolled 10,000+ patients</li> </ul>

- Tumor cells may become dependent on Bcl-2 for survival
- Inhibiting Bcl-2 releases pro-apoptotic proteins, which trigger apoptosis through the apoptosome

# Venetoclax is projected to generate ~\$6 Bn WW sales in 2026

250+ Active Trials | 89 Sponsored | 13 Potential New Indications

Venetoclax ~ \$6 Bn Worldwide sales Forecasted 2026



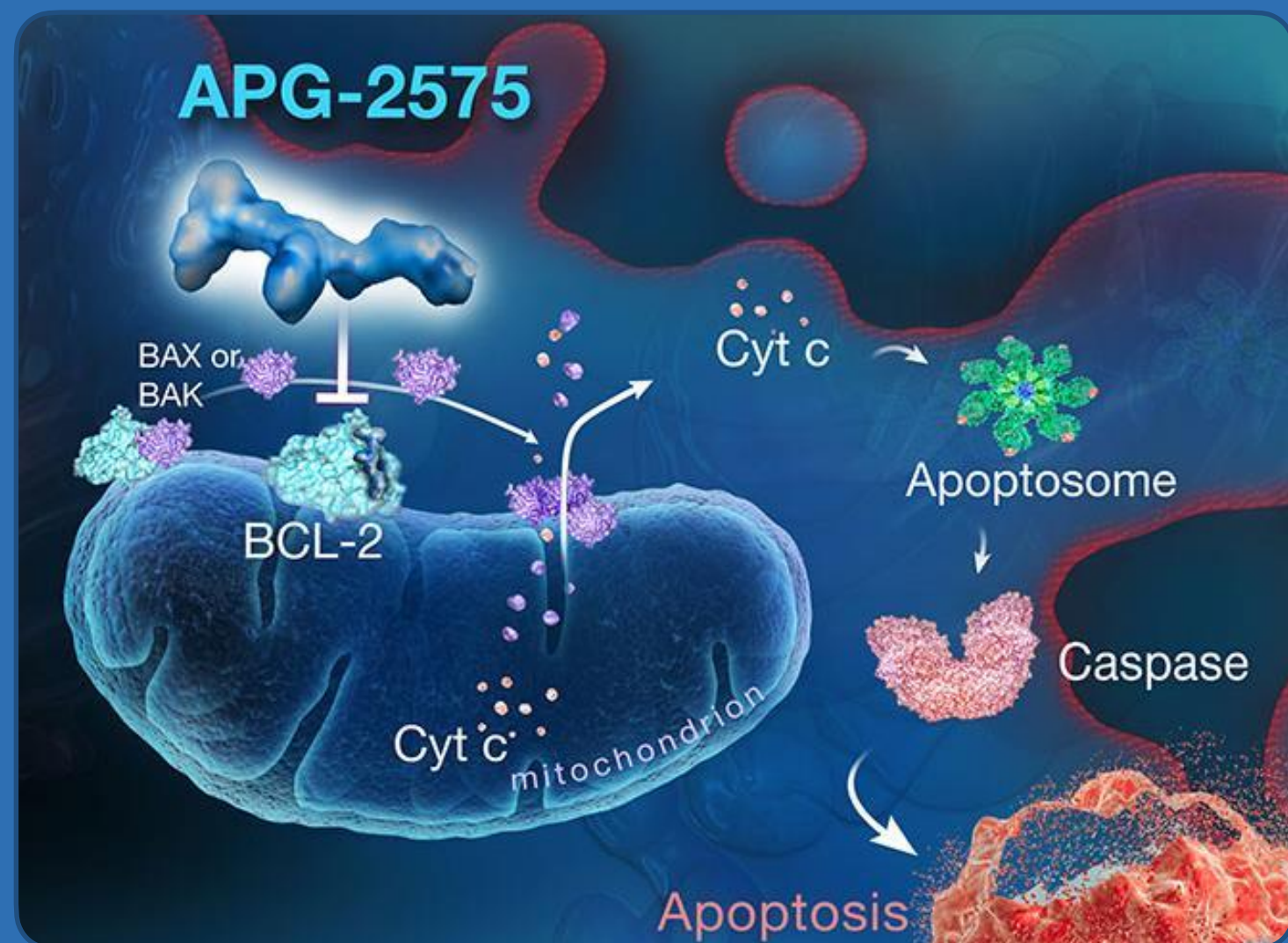
Approved indications

Reference for APG 2575: 2nd BCL-2 inhibitor vs. 1st BCL-2 inhibitor

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# APG-2575

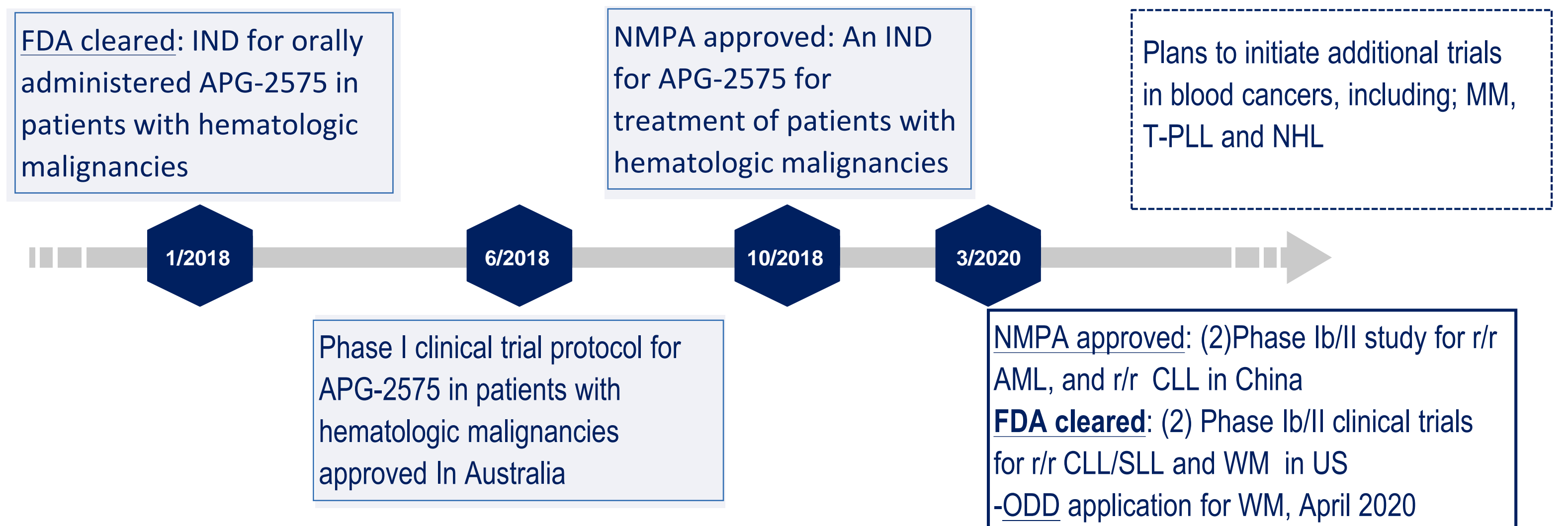
BCL-2 Selective Inhibitor  
 Novel, orally administered Bcl-2 selective inhibitor, follow to Venclexta®



## Summary of Key Results

- Ph I trial of APG-2575 in hematologic malignancies enrolling US & Australia
  - 21 patients enrolled up to 800mg (7 dose cohorts), all with daily dose ramp-up
  - CLL (n=8) completed daily dose ramp-up with no clinical TLS
    - 6/8 patients reached criteria for hematologic CR or PR (nodes & ALC)
  - Interim safety data shows APG-2575 is well-tolerated, No DLTs, only lab TLS, and MTD has not been reached
- Phase I trial in China has reached third dose cohort, No DLTs

## Milestones & Developments



# APG-2575

## Clinical Development

### Progress to Proof of Concept

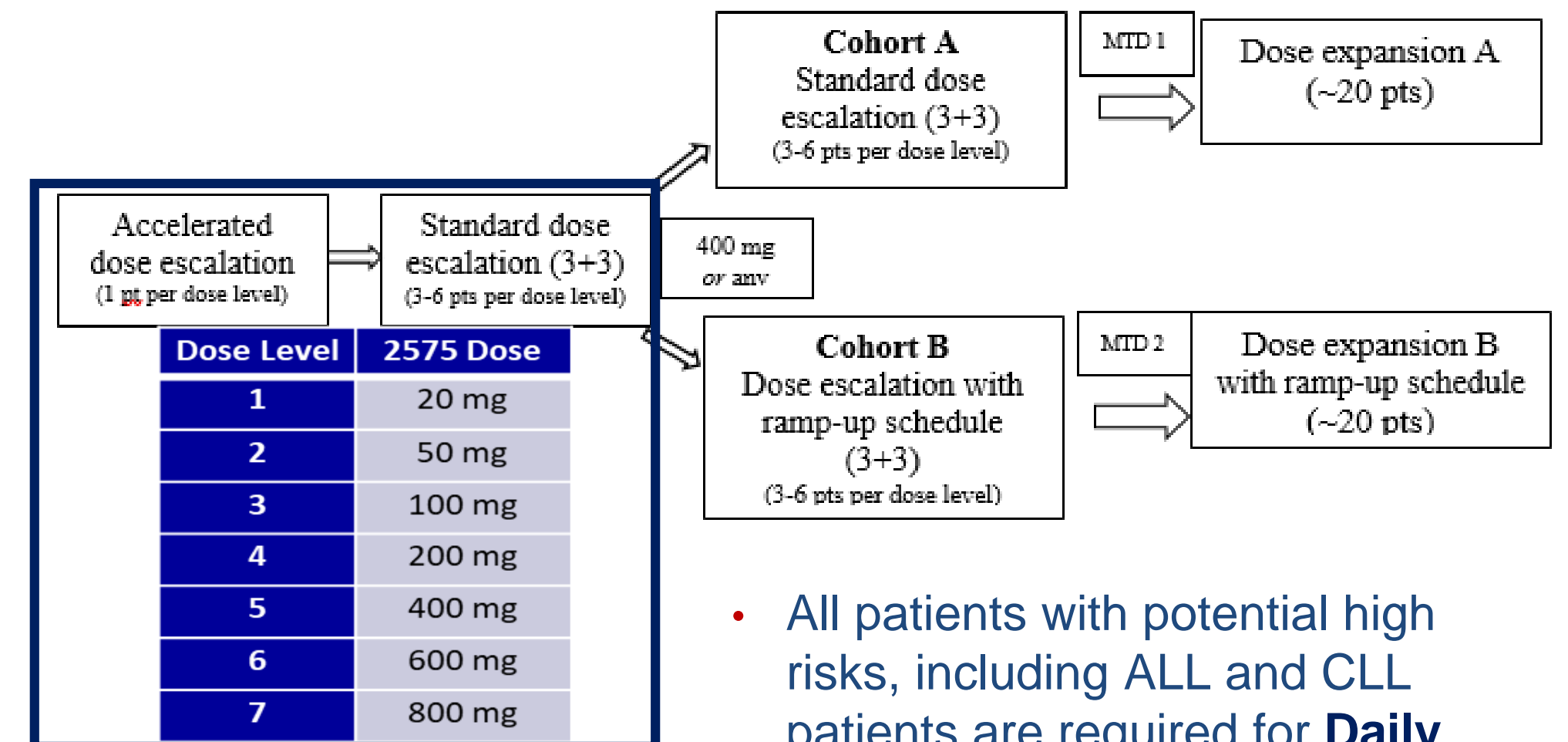
#### Trial 1 - U.S. & Australia

- 21 patients with hematologic malignancies have been treated with APG-2575 up to 800mg (6 dose levels)
  - All 8 CLL patients completed the daily dose ramp-up without TLS.
  - 4 CLL patients have reached a criteria for hematological CR (ALC)
  - 2 CLL patients have reached PR (lymph node & ALC)
- Interim data shows APG-2575 is well-tolerated
  - No DLTs, No Clinical TLS and the MTD has not been reached

#### Trial 2 - China

- 4 patients have completed the first cycle of treatment
- No Serious Adverse Reaction
- **NMPA approved Ph Ib / II studies for r/r CLL, AML in China**
- **FDA cleared two Ph Ib/II clinical trial for r/r CLL/SLL & r/r WM**

### Study Design of APG-2575 Ph I Trials



- All patients with potential high risks, including ALL and CLL patients are required for **Daily Ramp-up** (NOT weekly ramp-up like venetoclax) prior to receiving daily treatment at a desired dose cohort.

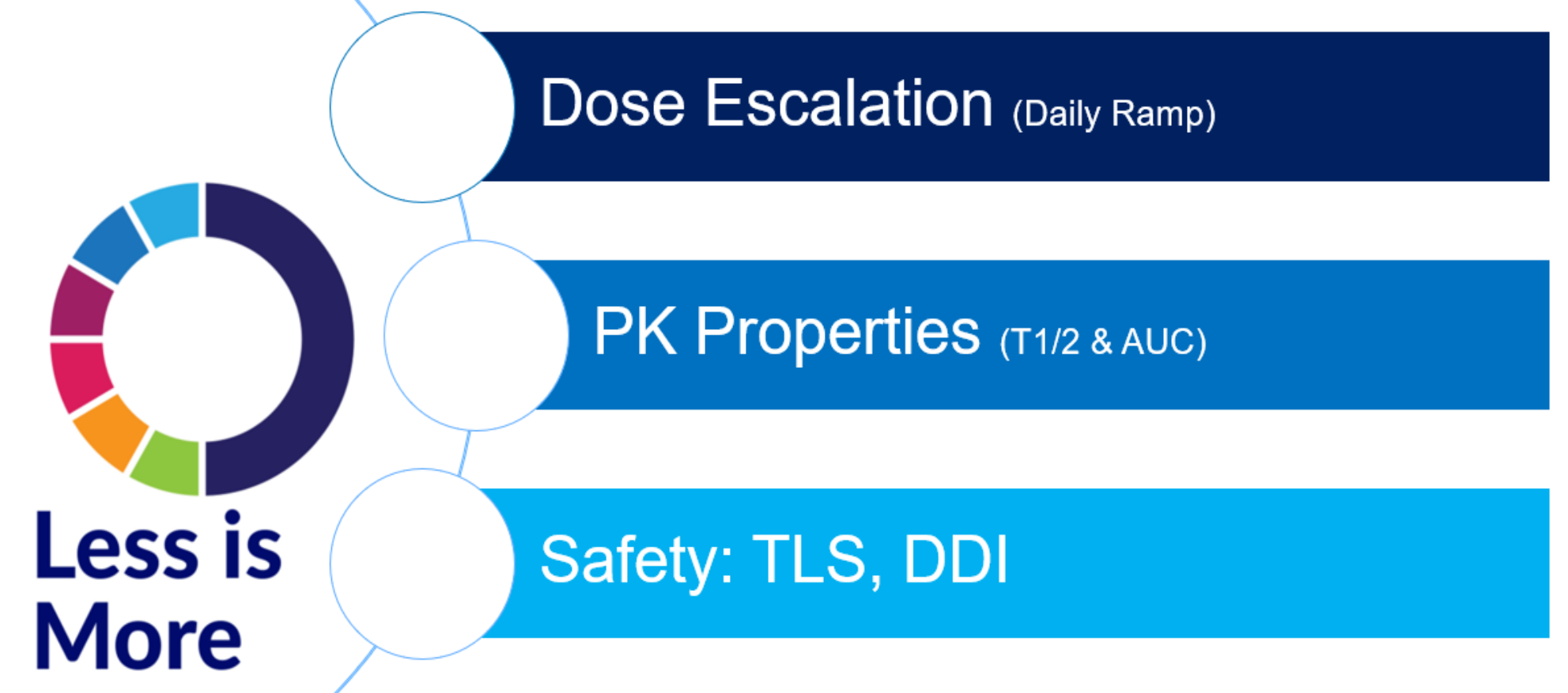
# APG-2575 and Venetoclax

- Venetoclax was the first Oncology PPI drug approved by the FDA (AbbVie 20+ years)
- Venetoclax is the third transformative therapy for lymphoma, after Rituxan and Imbruvica
- >250 Venetoclax trials are being conducted, potentially expanding to >13 indications

## Differences Compared to Venetoclax:

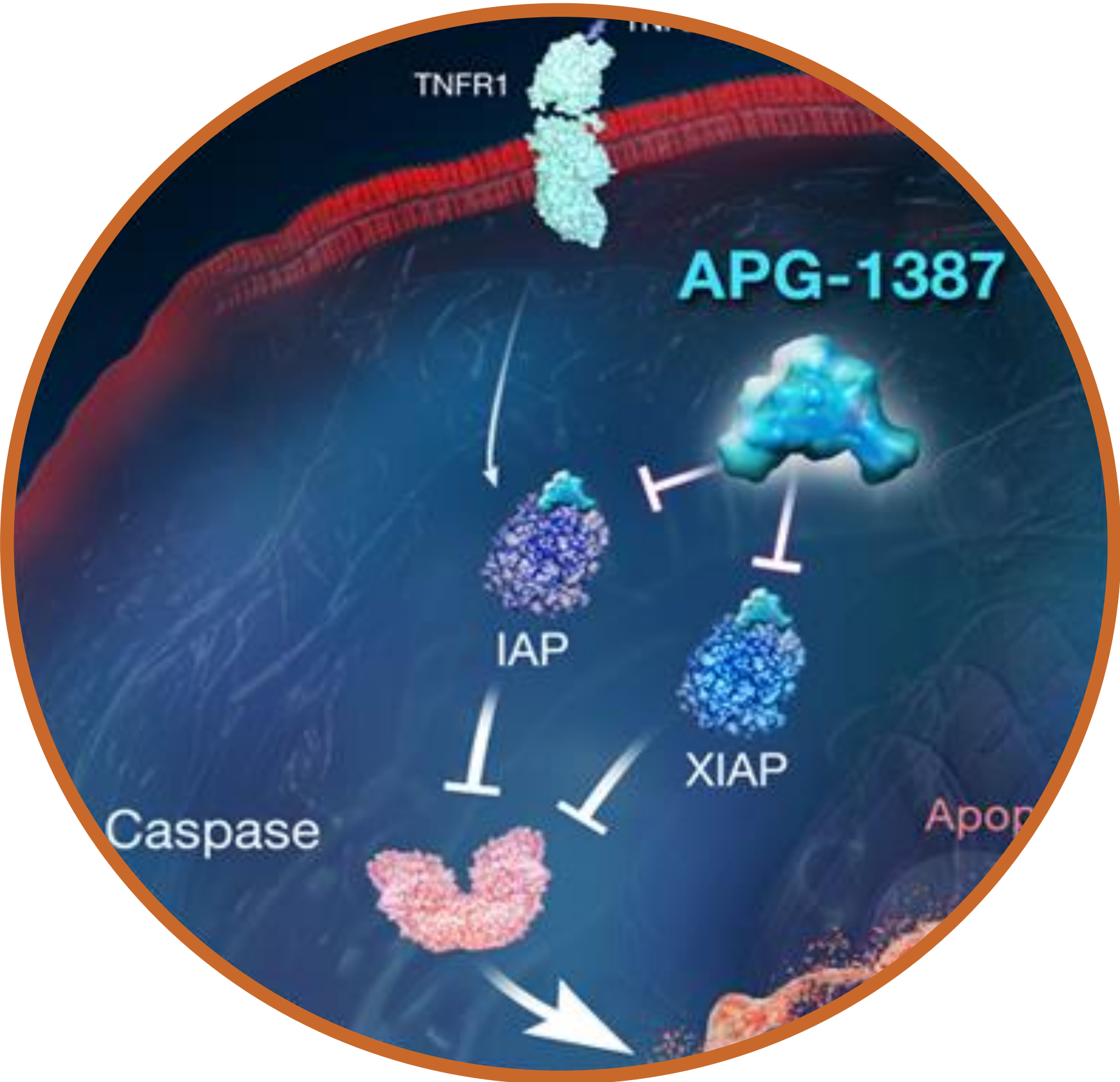
- No Clinical TLS, Lab TLS
- **Daily Ramp-up** verse weekly ramp up
- **Short  $T_{1/2}$  & AUC**--potentially lower risk of TLS with better safety profile

## When Selectively Targeting BCL-2



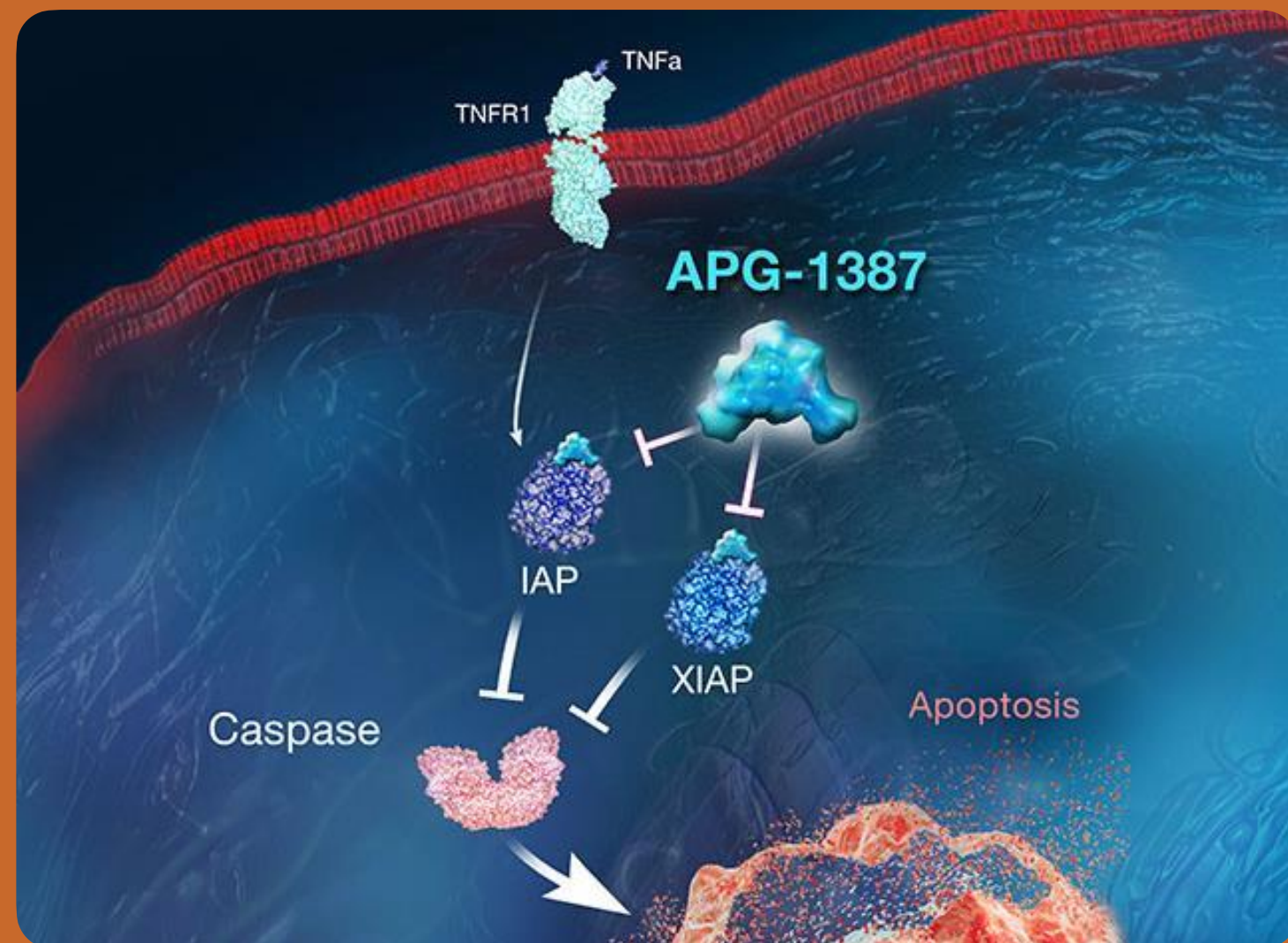
# APG-1387 Overview

An Antagonist of IAP/XIAP  
(SMAC Mimetic) Dimmer



# APG-1387

An Antagonist of IAP/XIAP  
(SMAC Mimetic) Dimmer



## Immuno-Oncology Development

- The only IAP-targeting drug to enter clinical trials in China and Completed the Ph I monotherapy clinical trials in solid tumors in US and China
- A Phase Ib clinical trial in combination with pembrolizumab (“Keytruda”) in solid tumors ongoing
- In 2020, two Phase Ib/II clinical trials of APG-1387 combined with immuno-checkpoint inhibitor or chemotherapy in advance solid tumors have been approved

## CHB Developments

- A Phase Ib trial in naive Chronic Hepatitis B (CHB) patients completed the enrollment and the Phase Ib trial is ongoing
- A Phase II trial combo with NAs in CHB patients is ongoing globally

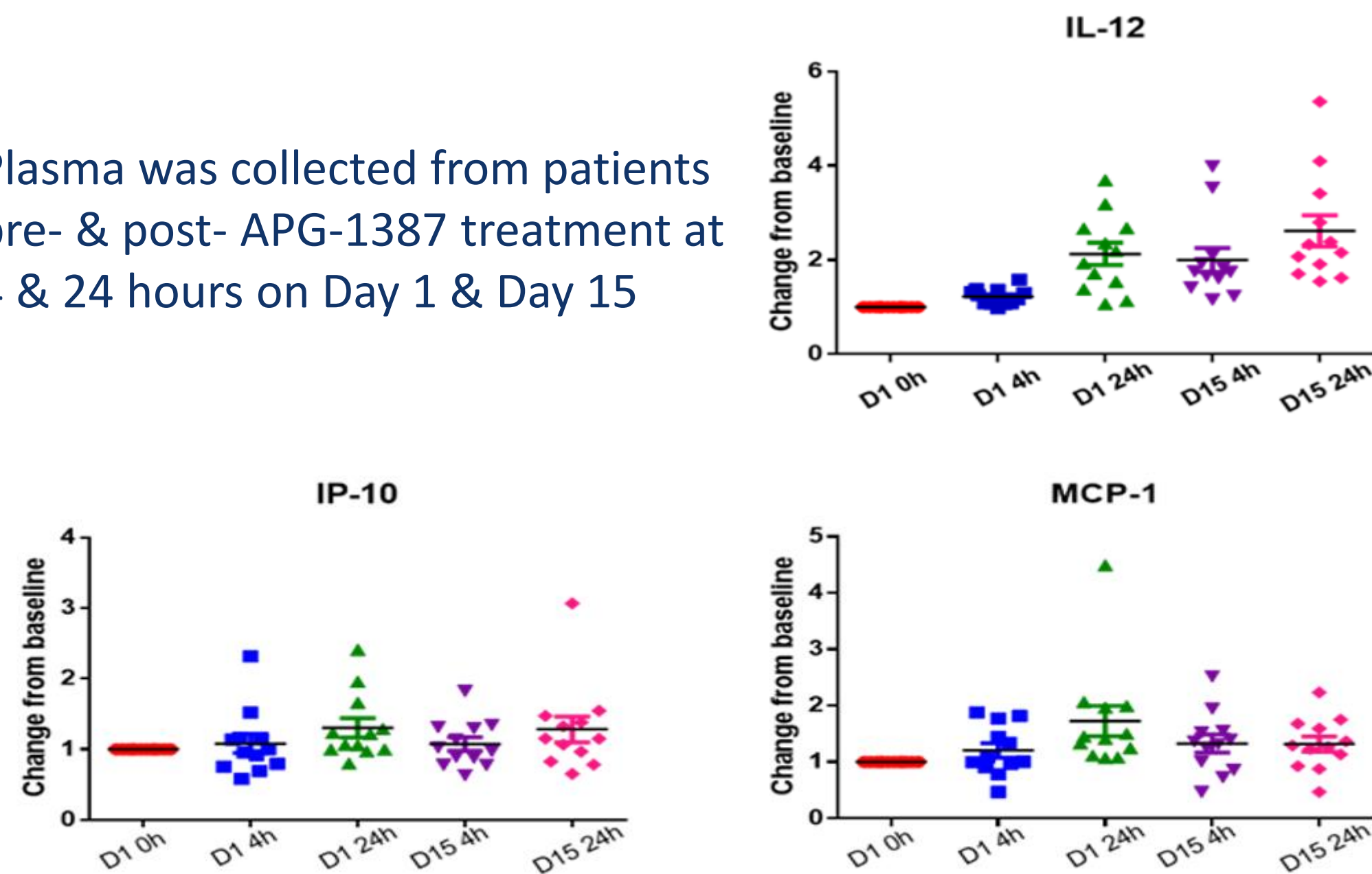
# APG-1387 Clinical Development

## Ph Ib | Immune Modulation and Activity

Ph Ib IO resistant/relapsed patients | Combination with pembrolizumab

### A potential host immune modulator

Plasma was collected from patients pre- & post- APG-1387 treatment at 4 & 24 hours on Day 1 & Day 15



- Human Cytokine 30-Plex analyses showed that IL-12, IP-10, and MCP-1 were increased in the plasma 24 hours post treatment with APG-1387.
- IL-12 elevation was observed in a time- and dose-dependent manner.

### Antitumor Activity

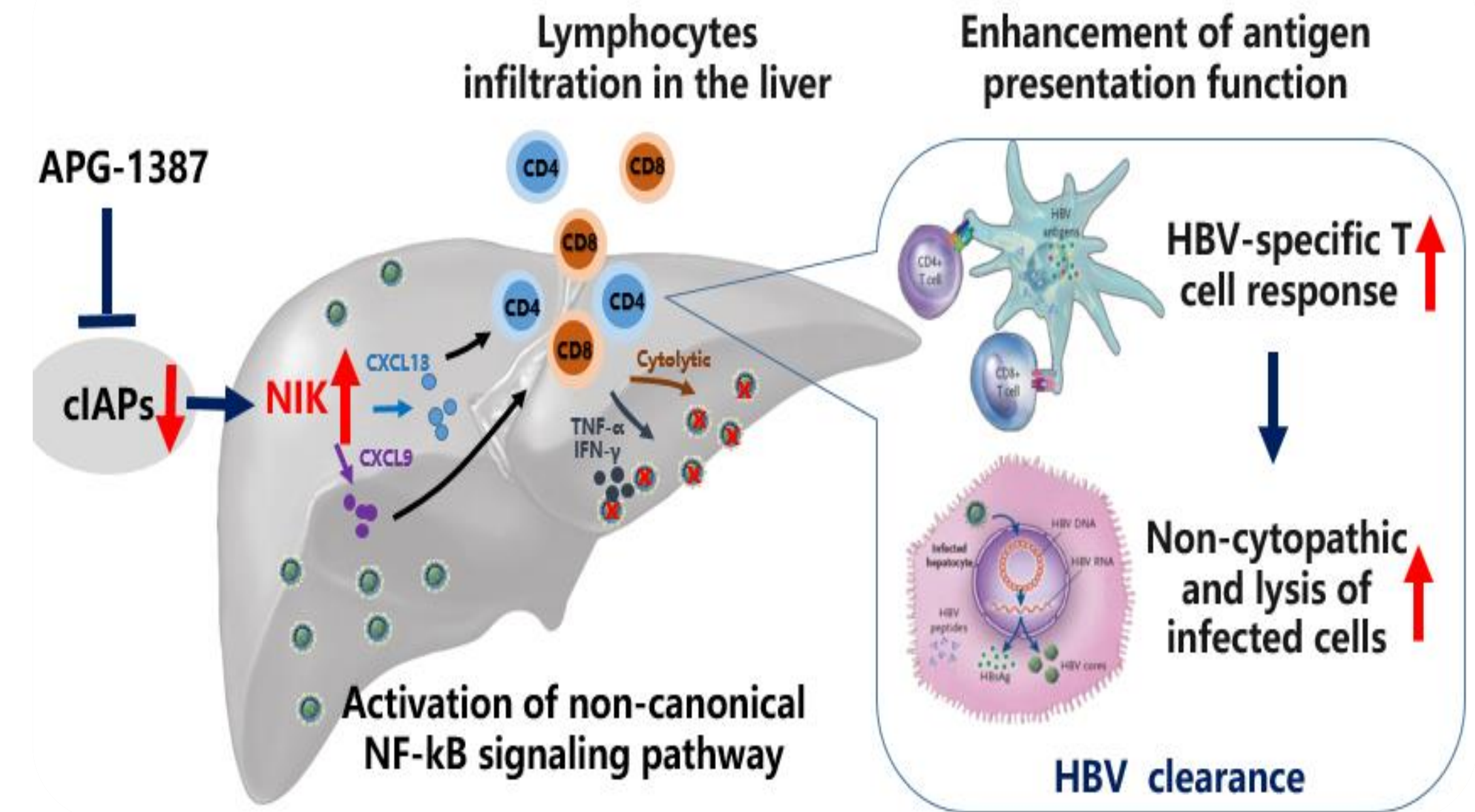
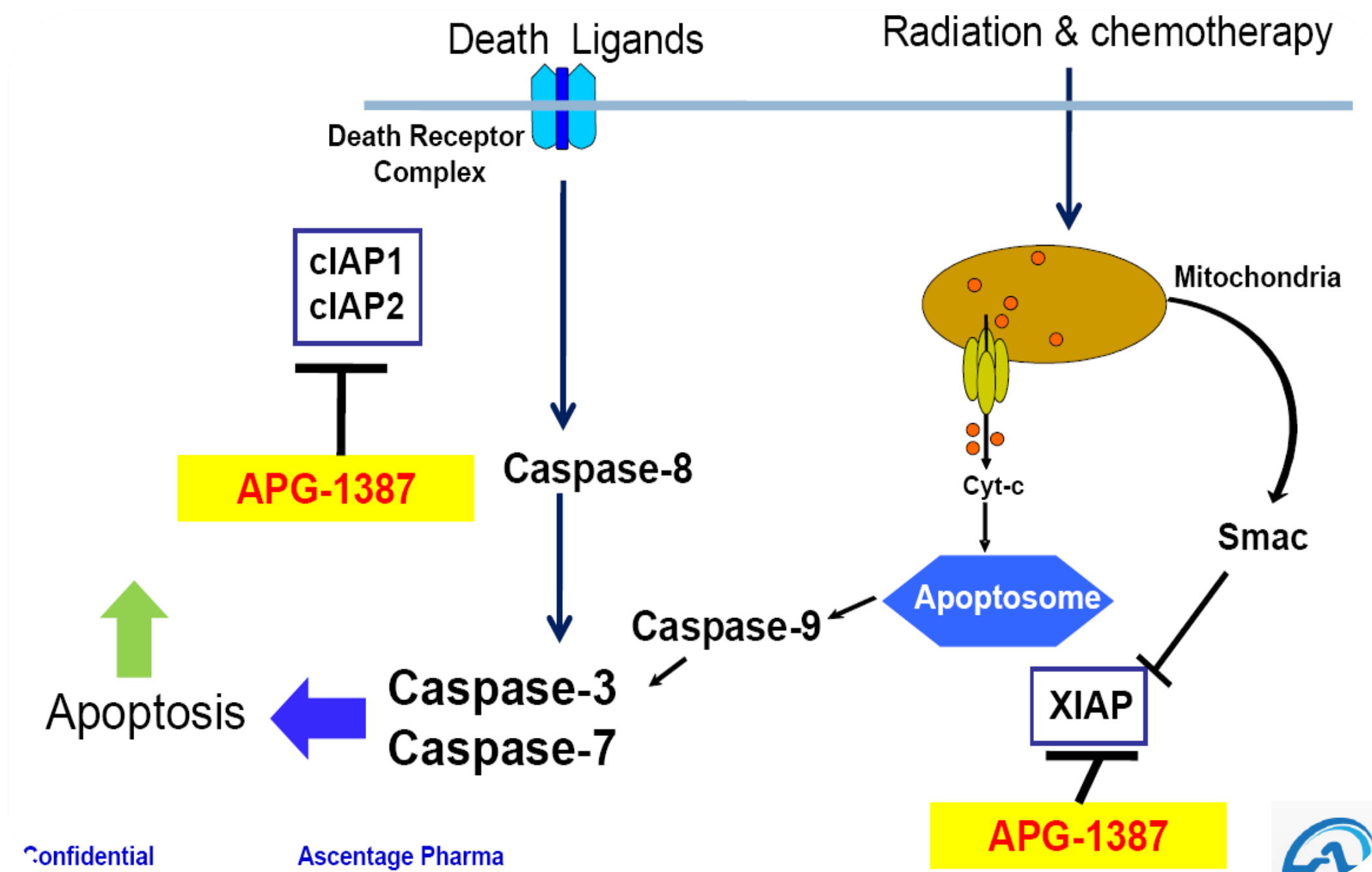
Response	All Cancers (N=41)	NSCLC (n=4)	Colorectal cancer (n=8)	Breast cancer (n=9)
ORR (CR+PR) Objective responses	10.8% (4/37)	50% (2/4)	12.5% (1/8)	11.1% (1/9)
DCR (SD + ORR) Disease control	43.2% (16/37)	100% (4/4)	50% (4/8)	33.3% (3/9)
Best overall response, n				
CR	0	0	0	0
PR	4	2	1	1
SD	12	2	3	2
PD	21	0	4	6
Non-evaluable	4	0	0	1

- Among 37 efficacy evaluable patients;
  - 4-PR (2 NSCLC | 1 CRC | 1 BC)
  - 12- SD | NSCLC cohort; 50% ORR | 100% DCR

# APG-1387

## A Novel Pan-IAP Antagonist (SMAC Mimetic) Dimmer

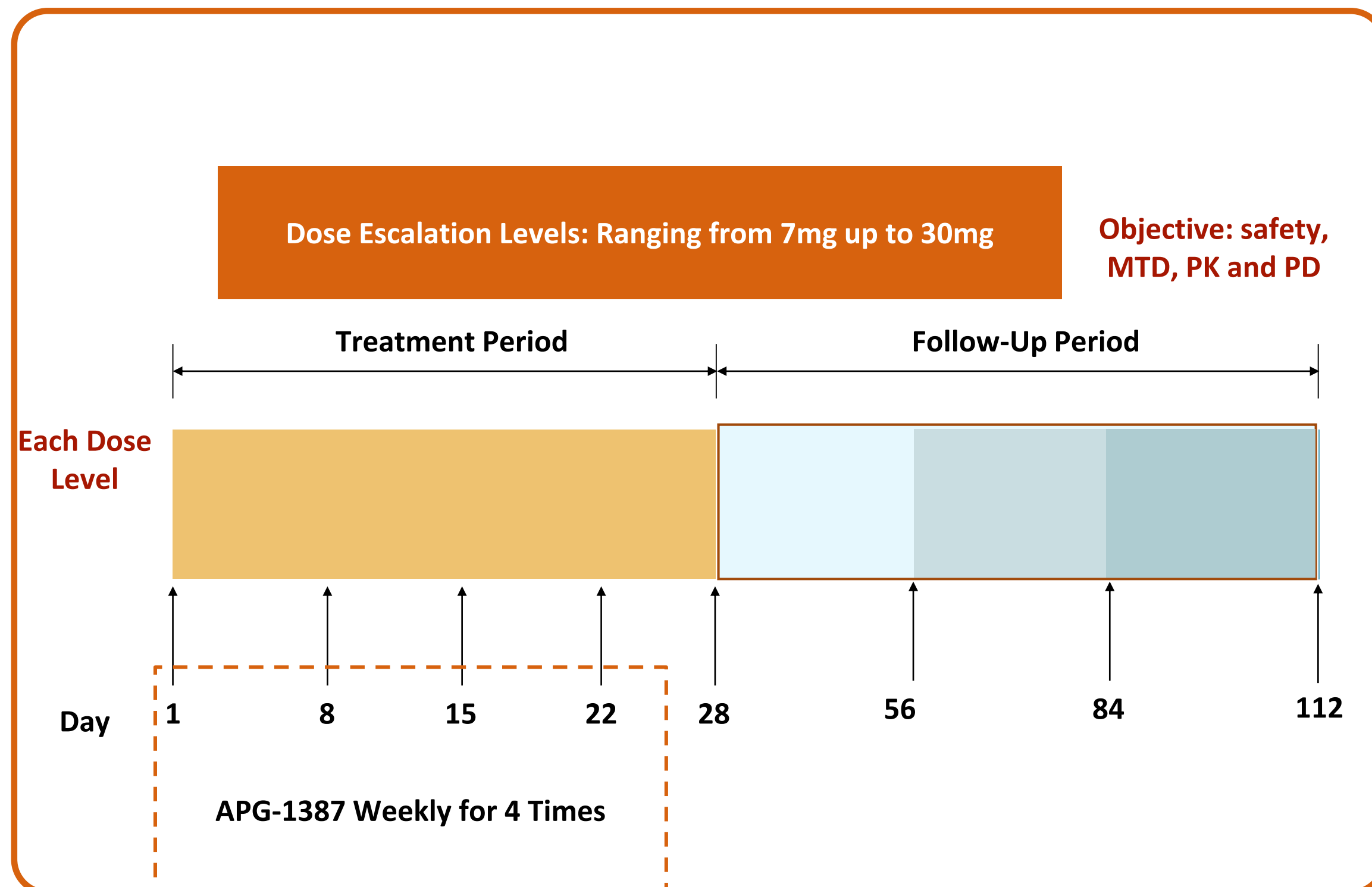
- Class I original innovation drugs, multiple small molecule IAPs antagonists, which can block the activity of IAPs family proteins (XIAP, cIAP-1, cIAP-2 and ML-IAP) and induce apoptosis.
- Preclinical studies suggest that it may be a new way to obtain functional cure for chronic hepatitis B.



# APG-1387

## Chronic Hepatitis B Clinical Development

### Study Design of APG-1387 Monotherapy in CHB



- As of latest evaluable day, a total of 30 patients were involved in studies.
- The MTD has not yet been determined. No DLT was observed. All AEs were mild to moderate in severity (Gr 1 or 2)
- After just 4 doses & compared to baseline
  - HBV DNA levels declined in 23 out of 30 patients <sup>1</sup>
  - HBsAg levels declined in 17 out of 30 patients
  - Some patients' HBV DNA and HBsAg levels continued to decline during the follow-up without further treatments

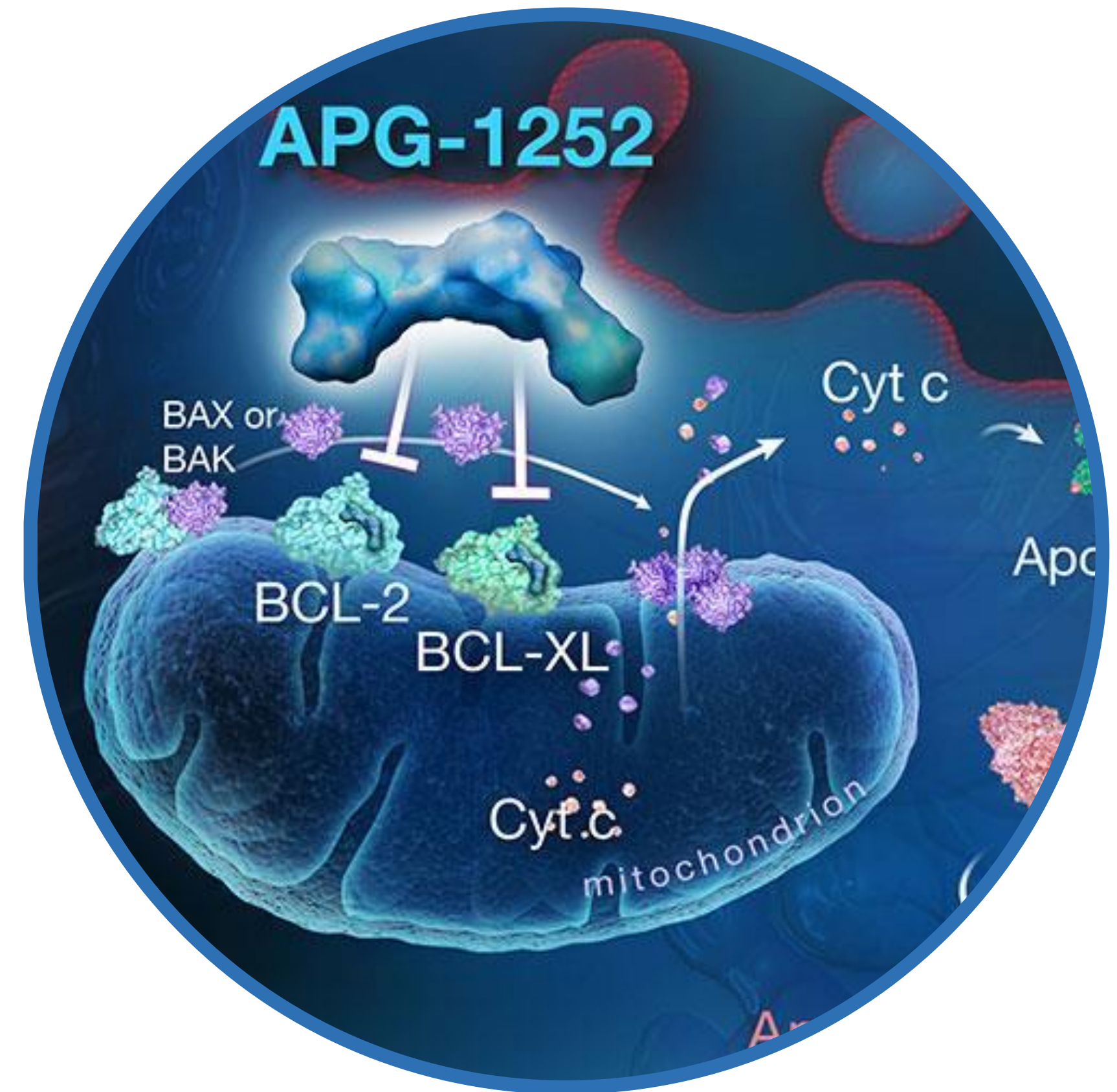
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Source: Company data Note: Study design for illustrative purpose only : actual clinical trial design may deviate from this illustrative chart

<sup>1</sup>.Among 33 treated patients, 3 patients are enrolled in June,2020 in the 2nd extension part. Their efficacy results (including HBV DNA and HBsAg changes) haven't been analyzed due to the short duration.

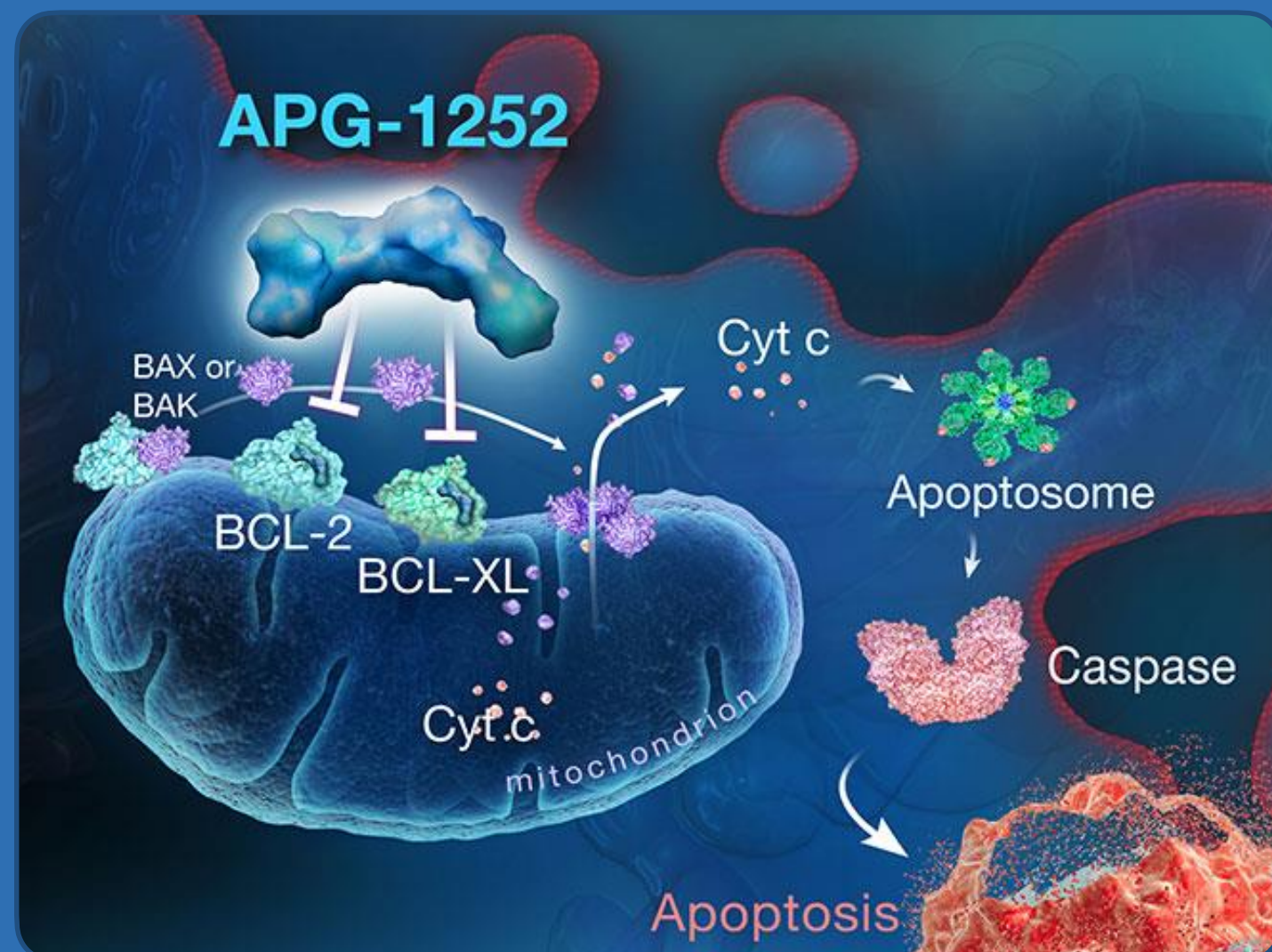
# APG-1252 Overview

Bcl-2/Bcl-xL Inhibitor



# APG-1252 pelcitoclax

BCL-2/BCL-xL Inhibitor



## Clinical Development

- Two Phase I dose-escalation trials in patients with advanced cancers in the United States and Australia ongoing
- A Phase I dose-escalation/expansion trial as a monotherapy in patients with SCLC in China ongoing
- 65 Patients are involved in the dose escalation trials

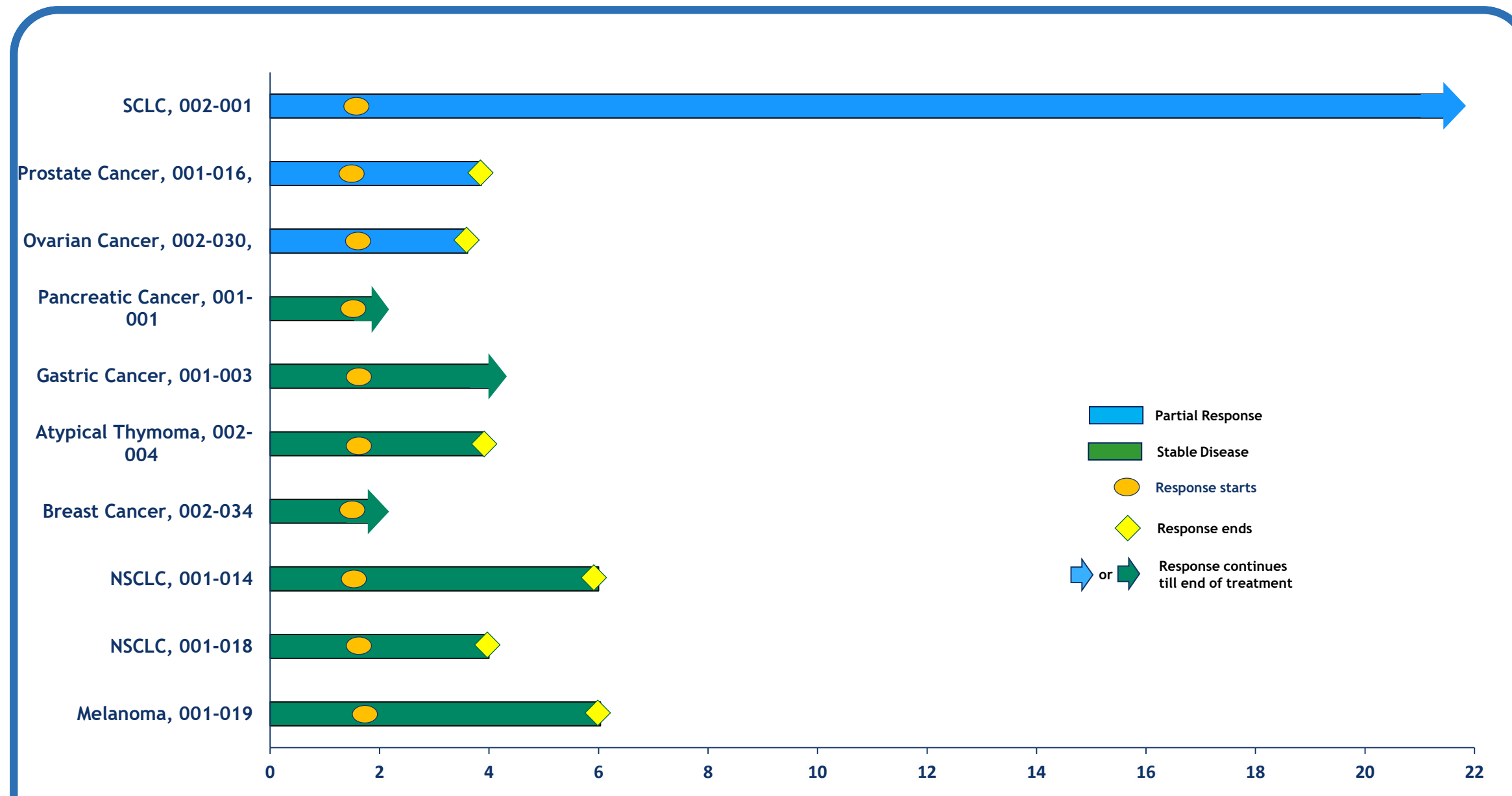
## Milestone

- New IND submitted to FDA in Dec 2019 : APG-1252 in combination with Paclitaxel for patients with SCLC
- Pending Phase I results, planning a Phase II trial in relapsed/refractory NSCLC, or r/r NSCLC, in the United States and China.

# Palcitoclax (APG-1252)

Ph I Interim Efficacy Data | n=42

## Single agent activity in advances solid tumors



- A total of 7 patients achieved SD, 4 of them were at 10mg, BIW; 20mg, BIW, 40mg, BIW and 240mg, BIW (patient #001-001, 001-003, 002-004 and 002-034).
- Three patients achieved SD at 320mg, BIW or QW cohort.(patient #001-014, 001-018 and 001-019).
- Five patients had SD lasted for ≥4 cycles, among them 2 patients had SD lasted for ≥ 6 cycles.

### Durable PR in a patient with SCLC

Before APG-1252

After APG-1252

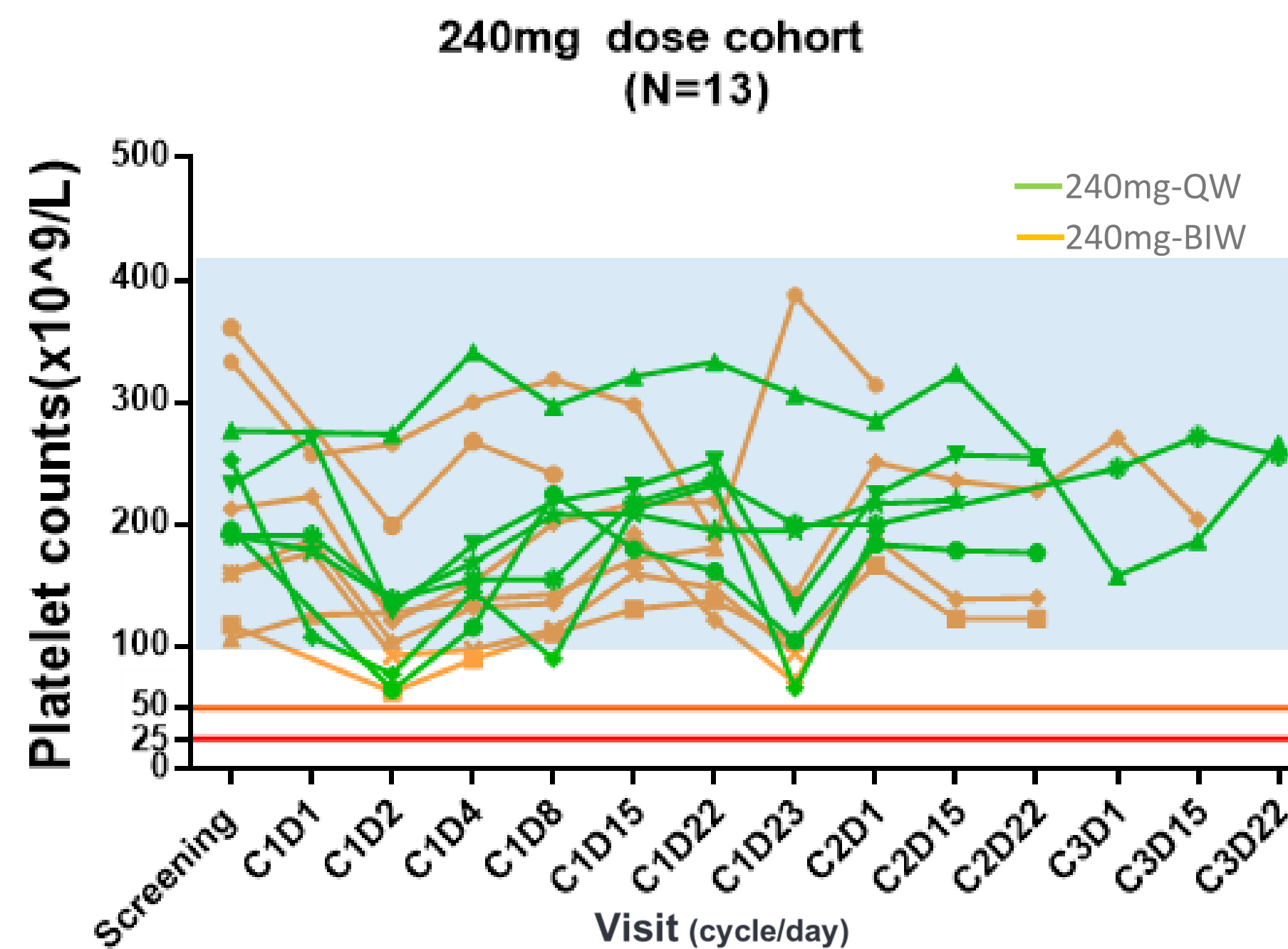
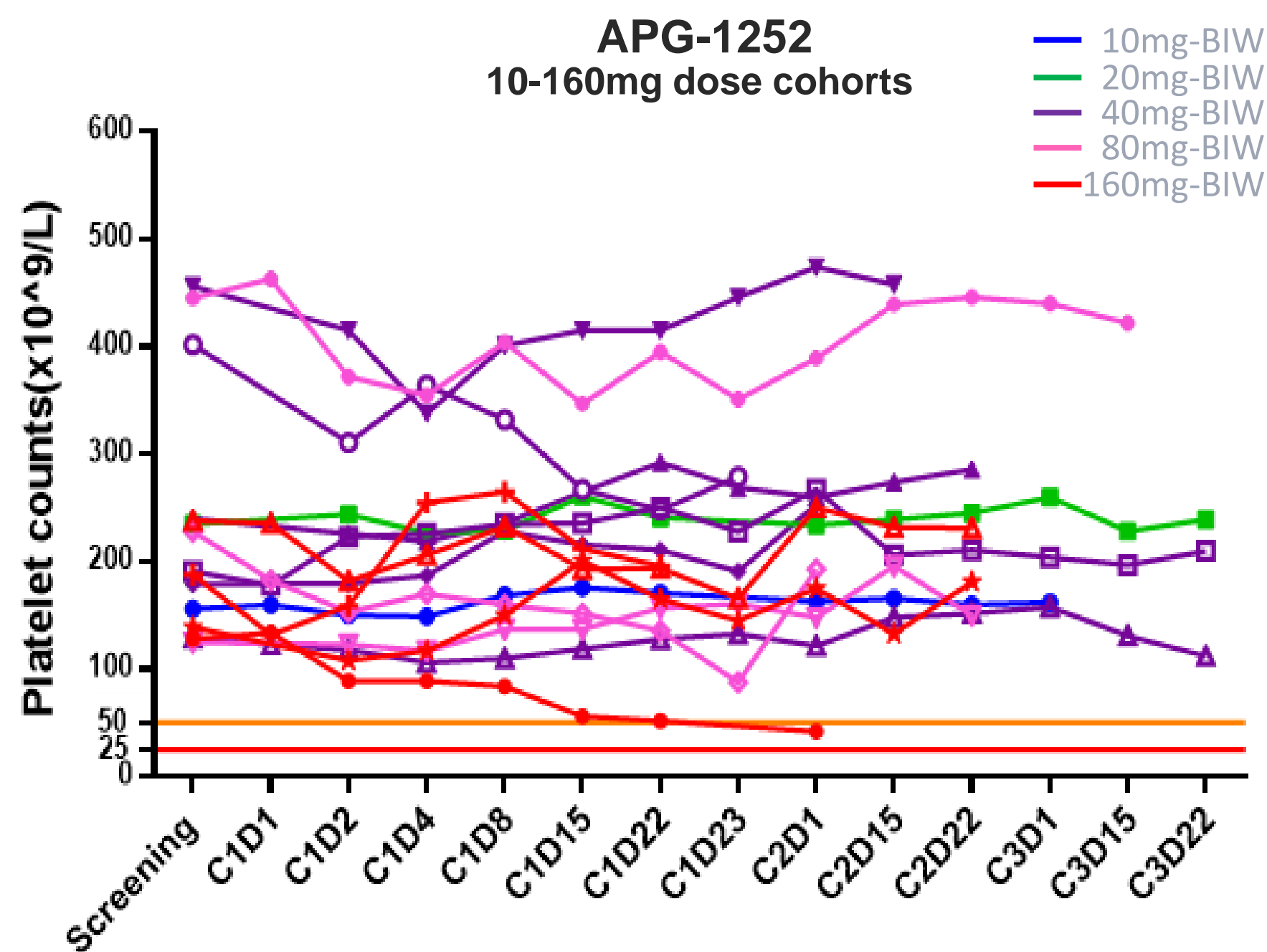


Hepatic tumor size decreases 44%  
Response maintained > 20 cycles

# Palcitoclax (APG-1252)

Ph I Safety Data | Platelet Toxicity

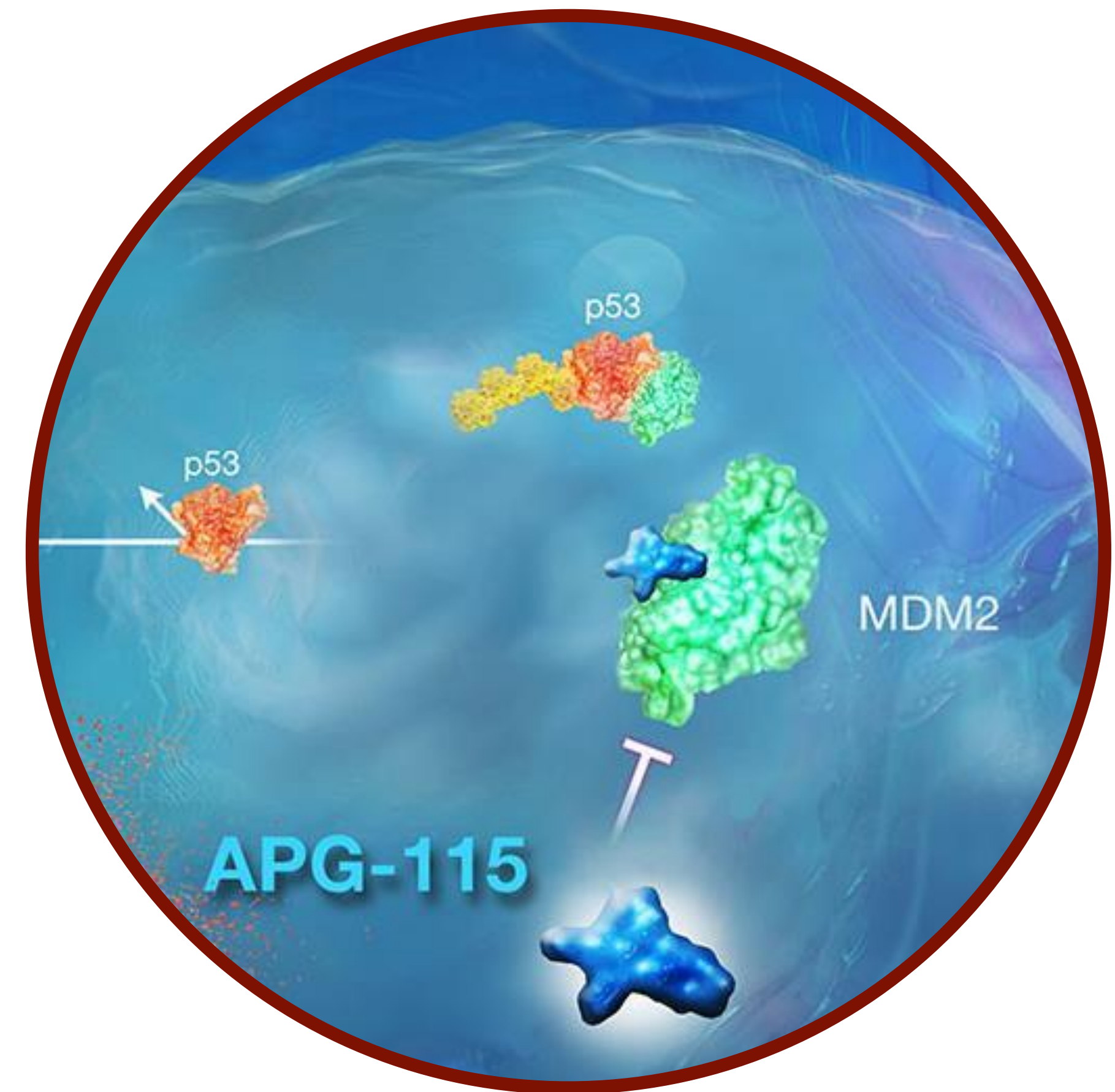
## APG-1252 Solves Platelet Toxicity by Design; 240mg QW RP2D



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# MDM2-p53

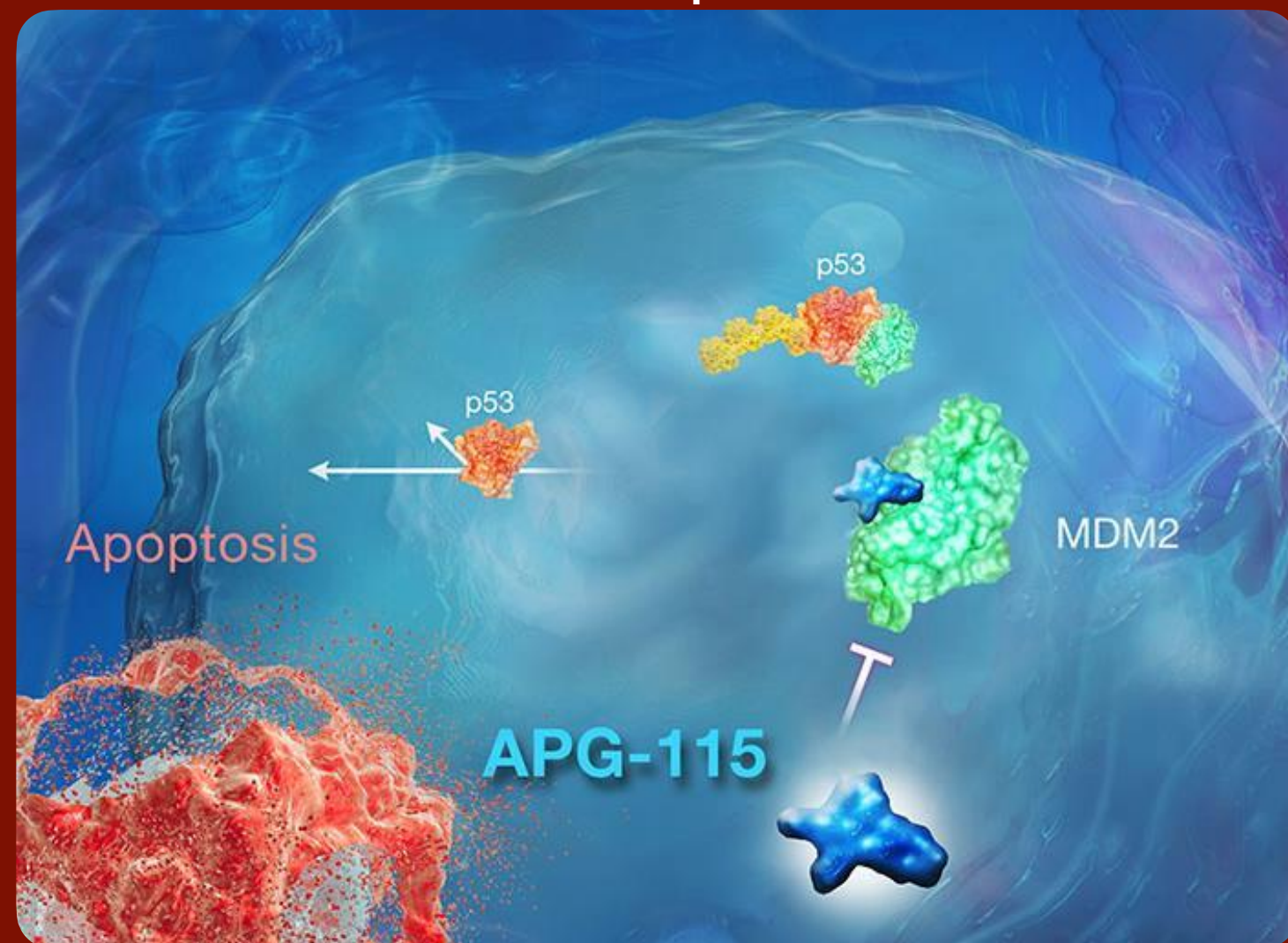
Activates p53 tumor suppression via  
MDM2-p53 PPI



# APG-115 Overview

MDM2-p53 Inhibitor

Activates p53 tumor suppression  
via MDM2-p53 PPI



## Clinical Development

- Completed Two Phase I trials in the U.S. and China, respectively in advanced solid tumors or lymphoma
- Completed enrollment of the Ph Ib clinical trial (19 patients were treated) treated in combination with Keytruda® with 4 dose-escalation cohorts in the U.S.
- A Ph II trial in combination with Keytruda® in patients with advanced solid tumors is ongoing, focus on the r/r IO melanoma, NSCLC and others.
- 1 patient confirmed CR, 2 PRs were observed, 7 patients had SD as the best response; the total DCR is 55.5% with an ORR of 16.9% (among 18 efficacy evaluable subjects)

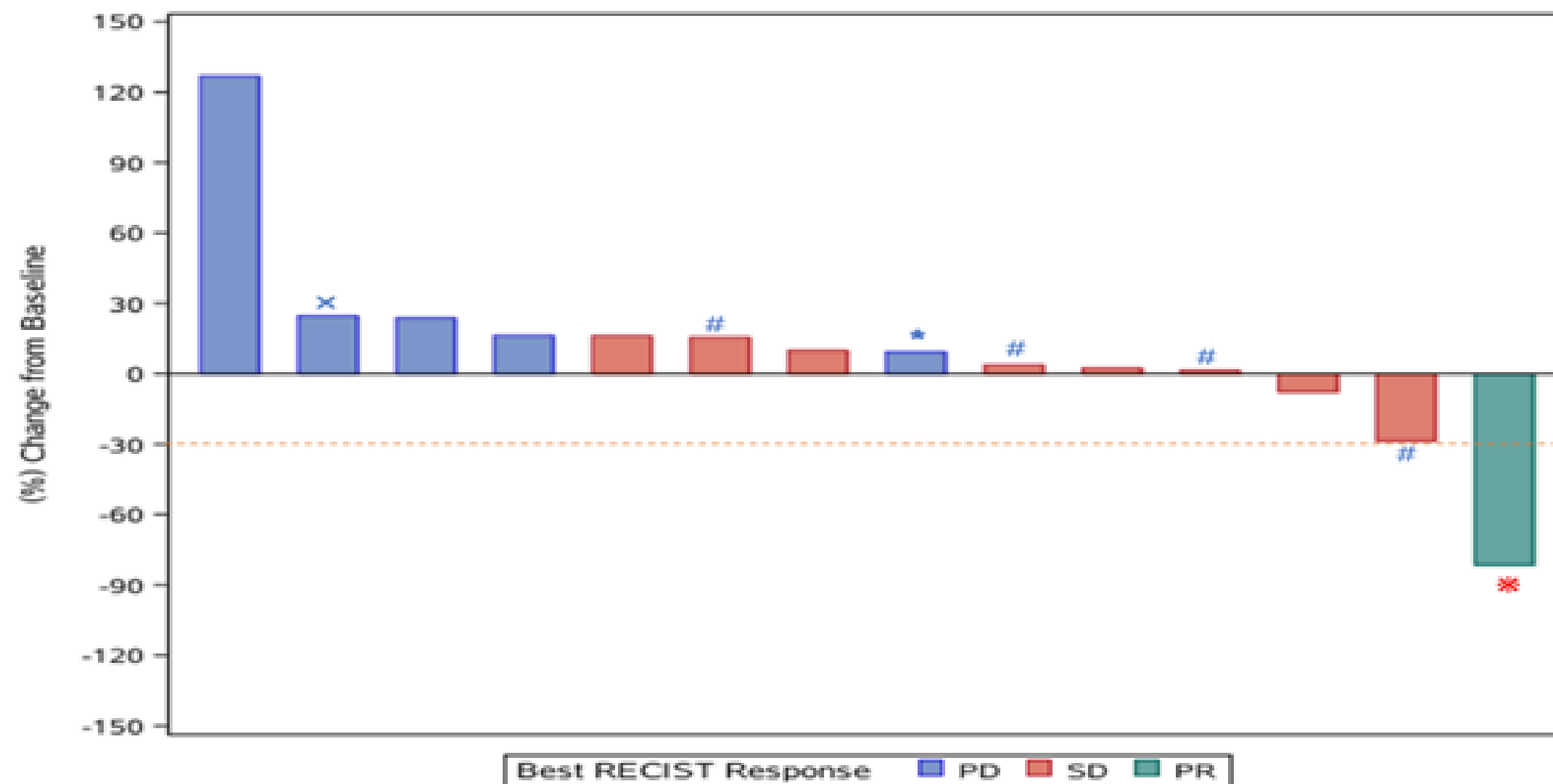
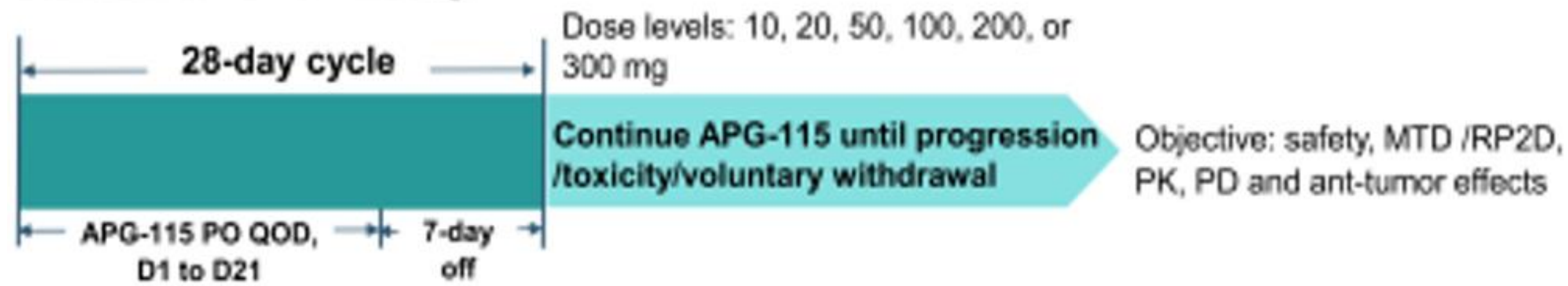
## Milestones

- Made an oral presentation on the preliminary results at the International Congress on Targeted Anticancer Therapies by European Society for Medical Oncology in February 2019
- Phase Ib/II clinical trial for APG-115 in combination with chemotherapeutic or other targeted agents for the treatment of patients with hematologic malignancies was approved by the NMPA in China in July 2019
- We plan to submit additional INDs for combination trials in China and U.S.
- We have completed dosing of the first patient in its Phase Ib clinical study treating patients with hematologic malignancies in China in July 2020

# APG-115 US-101

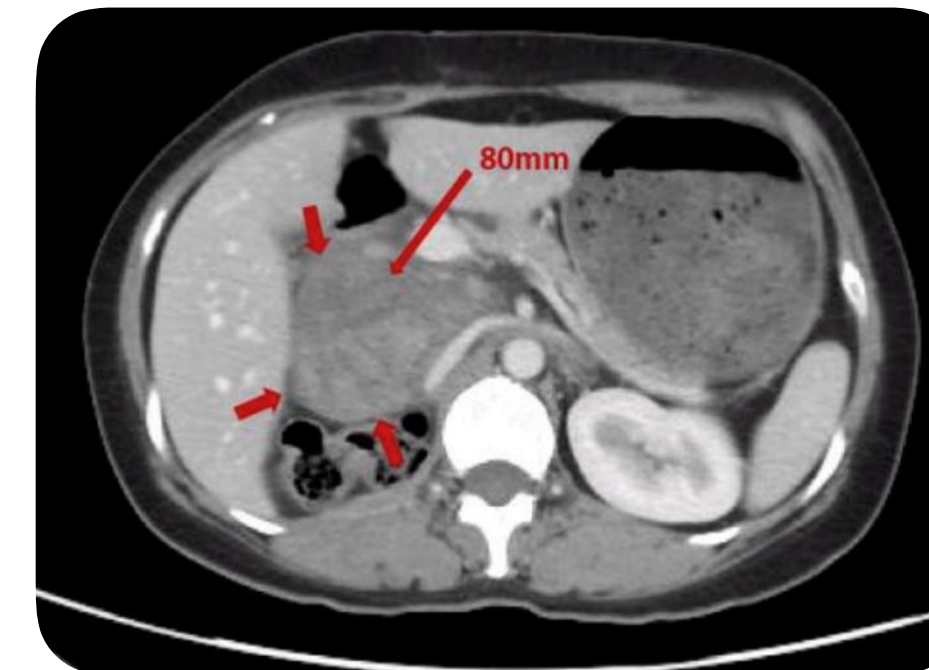
## Single Agent Ph I Study Results

Phase I dose escalation study - accelerated dose escalation, then standard "3+3" design



## Single Agent Activity

80mm  
Baseline



36 mm  
Cycle 2  
~55% Decrease



- 39 yr old Female, with Lipomatoid Liposarcoma, lymphatic metastasis (T2N1M0 )
- 5 cycles of AD Chemo (Adriamycin+ Dacarbazine)
- APG-115: 150mg QOD

# APG-115 US-002

## Ph Ib | Overview and Treatment

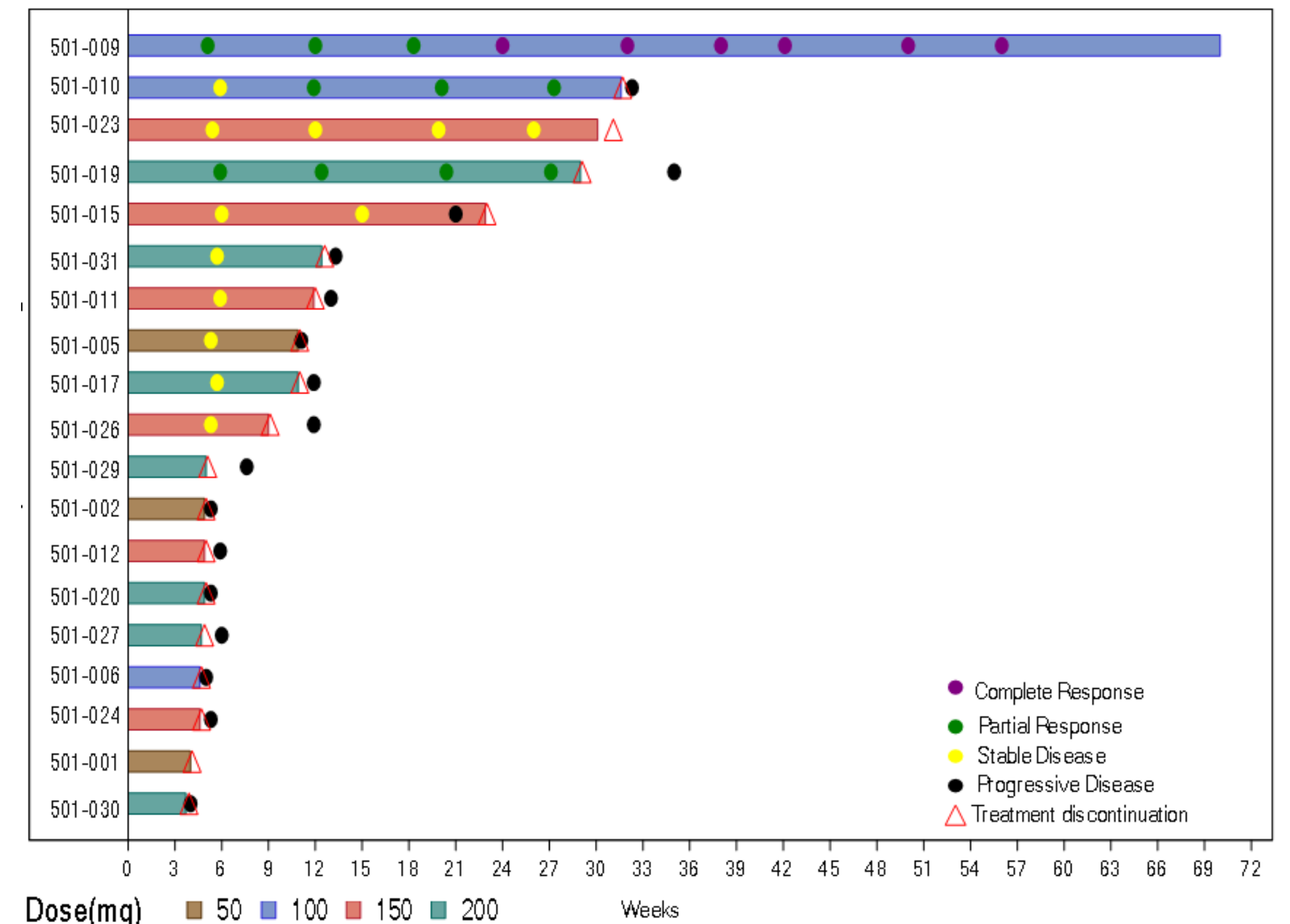
### Ph Ib IO resistant/relapsed patients| Combination with Keytruda®

#### Overview & Safety

Treatment Related AEs (at least Grade 3) by Preferred Term					
	50 mg (n=3)	100 mg (n=3)	150 mg (n=6)	200 mg (n=7)	Overall (n=19)
Any drug-related AEs with Severity Grade at least 3, n(%)	0	1 (33.3)	2 (33.3)	3 (42.9)	6 (31.6)
Platelet count decreased	0 (0.0)	0 (0.0)	2 (33.3)	2 (28.6)	4 (21.1)
Neutrophil count decreased	0 (0.0)	1 (33.3)	1 (16.7)	1 (14.3)	3 (15.8)
Adrenal insufficiency	0 (0.0)	0 (0.0)	0 (0.0)	1 (14.3)	1 (5.3)
Anemia	0 (0.0)	0 (0.0)	0 (0.0)	1 (14.3)	1 (5.3)
Febrile neutropenia	0 (0.0)	0 (0.0)	0 (0.0)	1 (14.3)	1 (5.3)
Lymphocyte count decreased	0 (0.0)	0 (0.0)	1 (16.7)	0 (0.0)	1 (5.3)
White blood cell count decreased	0 (0.0)	0 (0.0)	0 (0.0)	1 (14.3)	1 (5.3)

- MTD not reached, No DLT observed
- **RP2D** is determined as **150mg QOD**
- No new safety finding when combined with pembrolizumab
- PK: AUC & Cmax generally increase dose proportionally over the dose range of 50-200 mg.
- PD: MIC-1(biomarker of TP53 activation) serum increase was exposure dependent within the dose range.
- Efficacy: **ORR: 16.7%** (1 CR|2PR) + 7SD = **DCR: 55.5%**
  - Resp: CR-Ovarian | PR-NSCLC, Appen. Adeno. | 7SD | 8PD

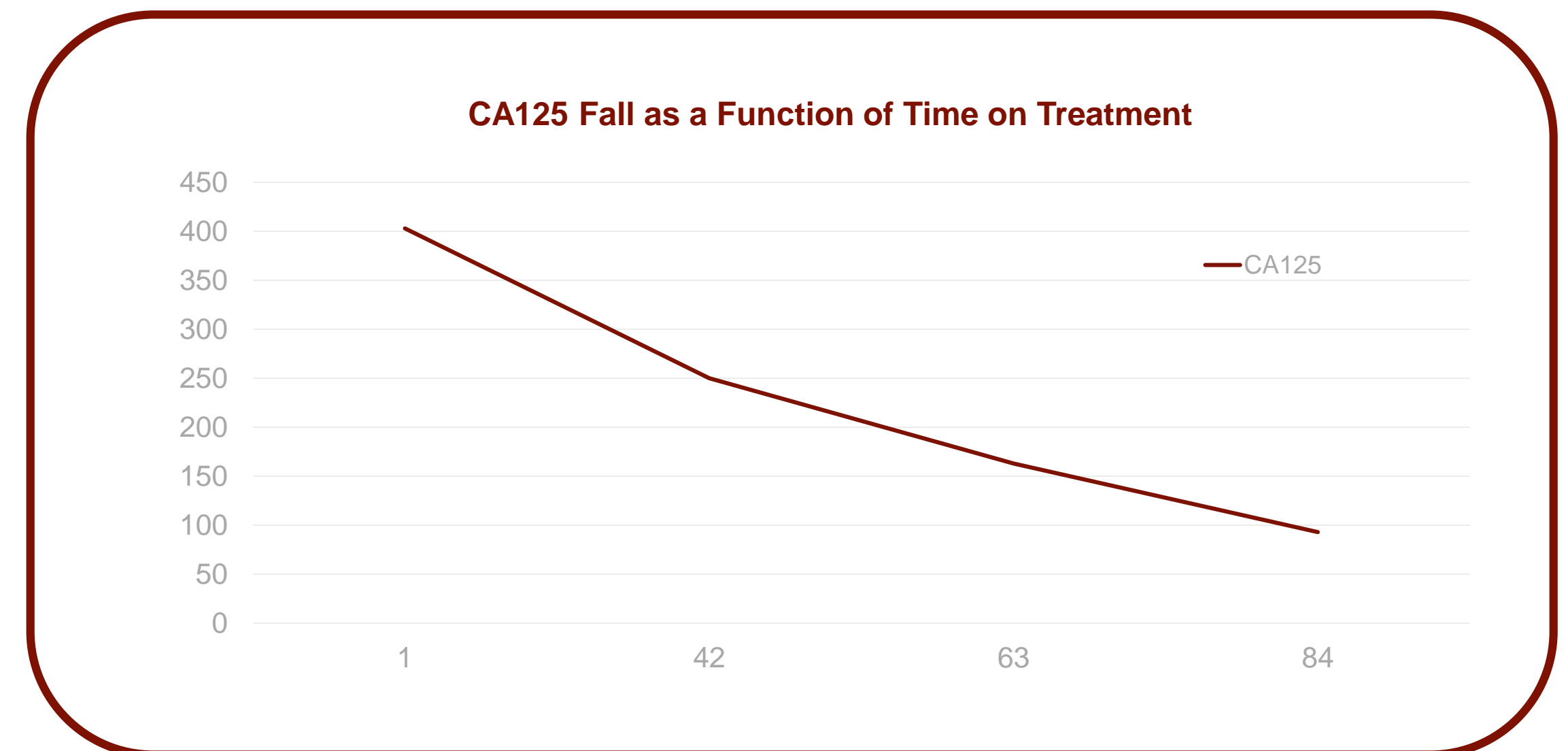
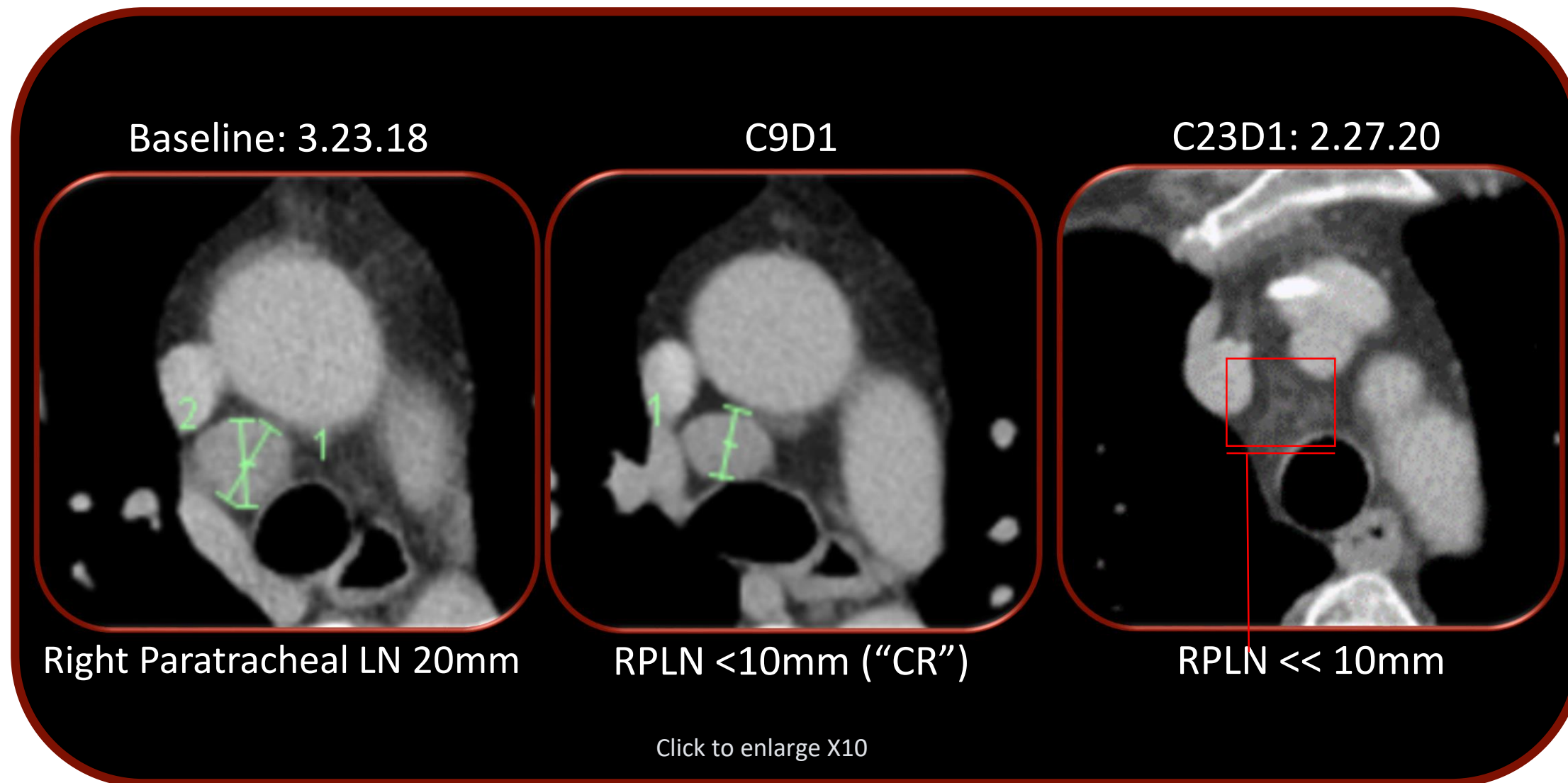
#### Overview & Treatment Duration



# APG-115: Promising Efficacy

Ph Ib | Combined with Keytruda

APG-115 and Keytruda achieves a CR in heavily pre-treated, ATM-mutated Ovarian Cancer



Treatment History			Clinical Trial
Initial Tx	Tx		
Neoadjuvant • Paclitaxel • Carboplatin • TAH BSO	Adjuvant • Carboplatin • Docetaxel	Relapse < 6mo. • <u>Doxil</u> • Topotecan • Bevacizumab • PD XMT1536	APG-115 (150mg) & Keytruda (200mg)

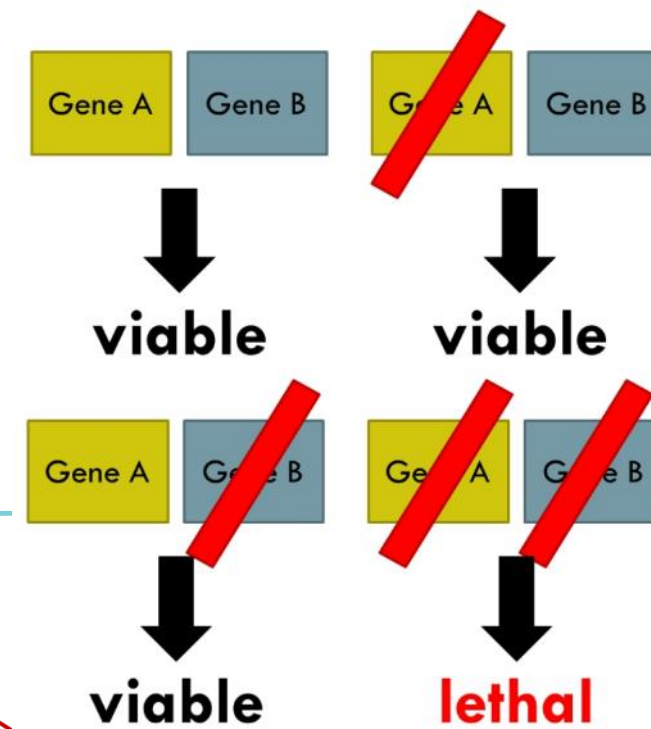
Trial to date(N=19) ; 3 dosing cohorts: 50mg | 100mg | 150 mg

Efficacy(N=18) ;	Safety
1 CR   2PR   7SD ORR = 16.9%   DCR= 55.5%	The combination is well-tolerated No DLTs, No additive AEs

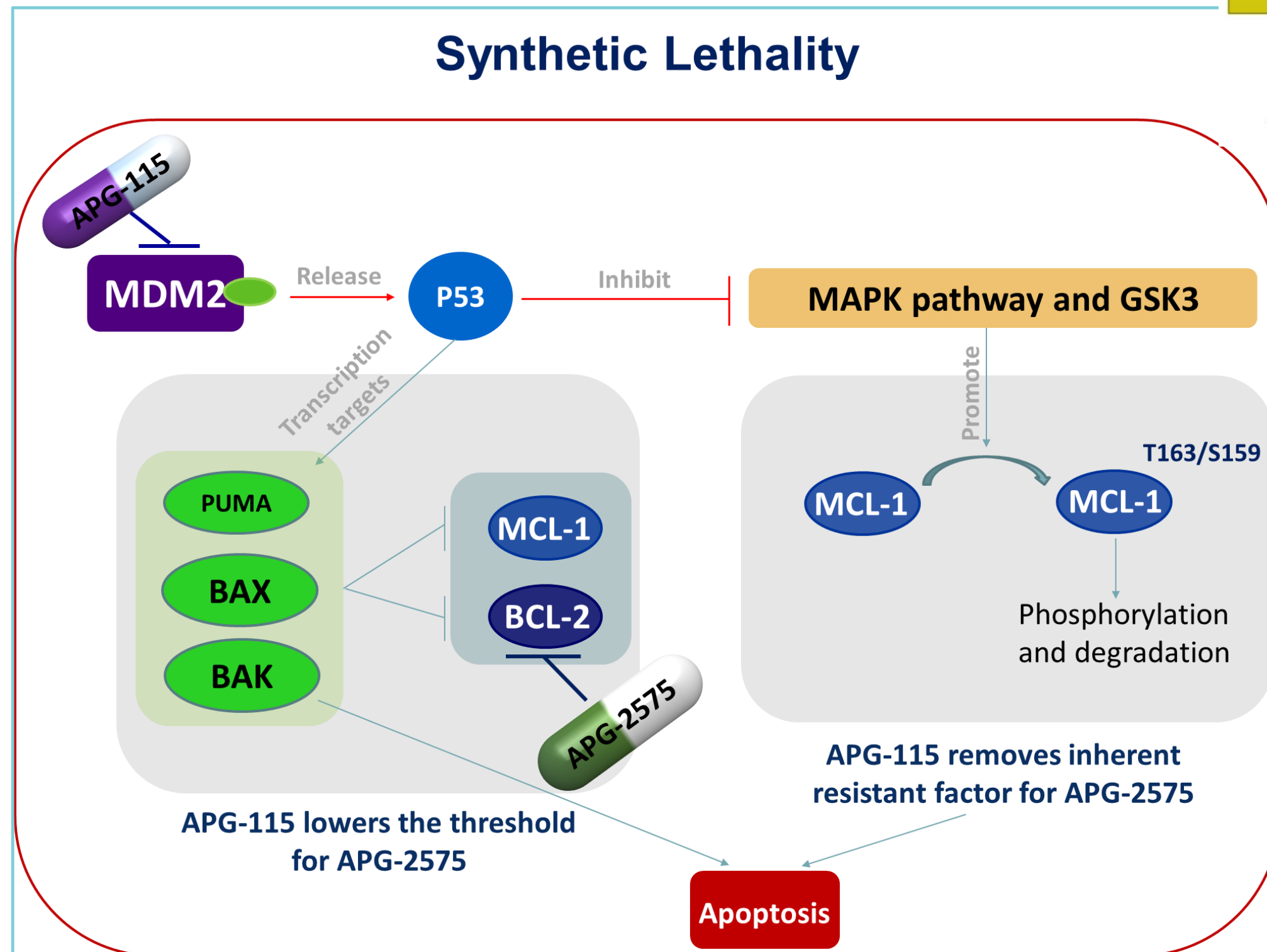
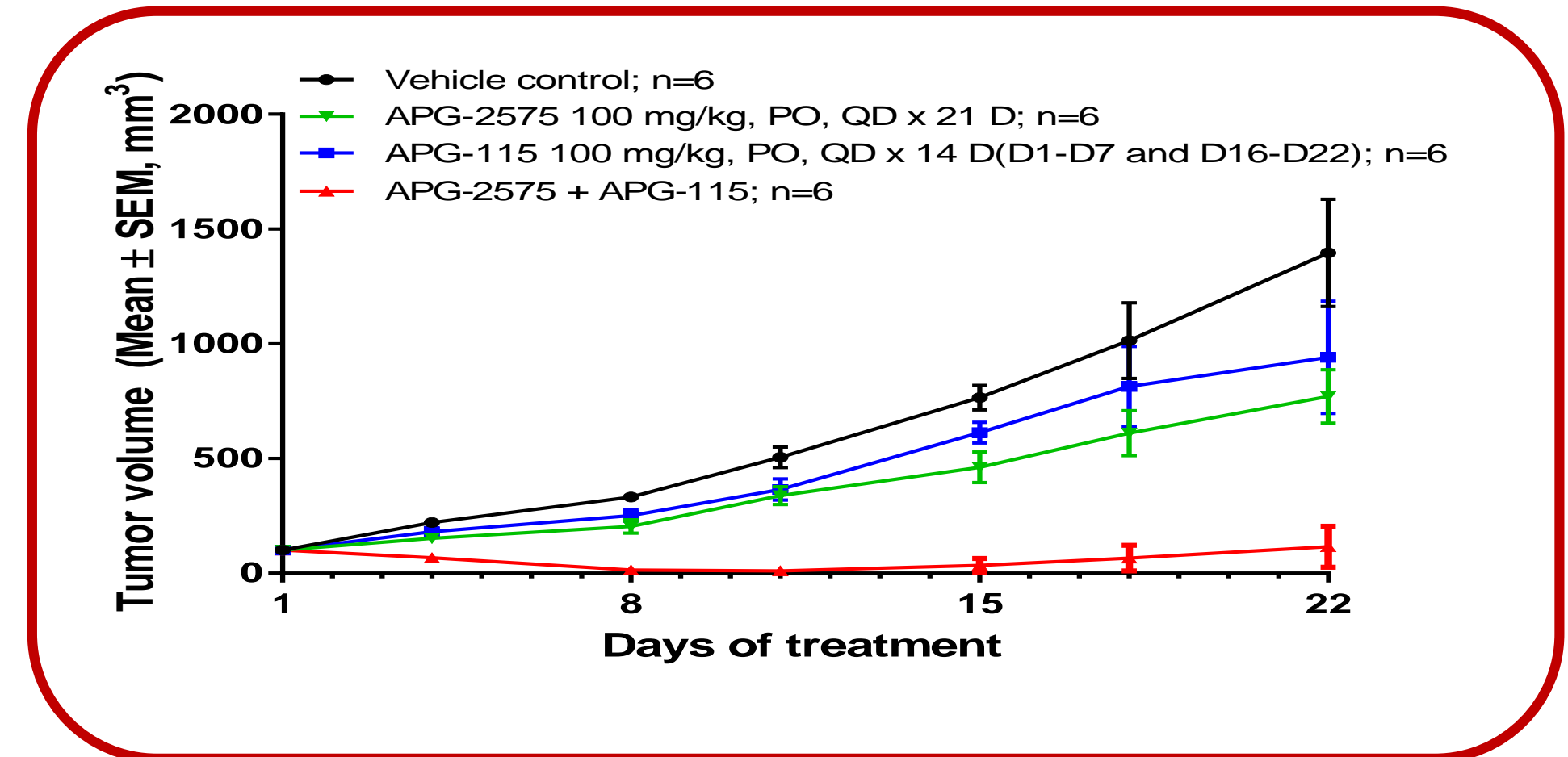
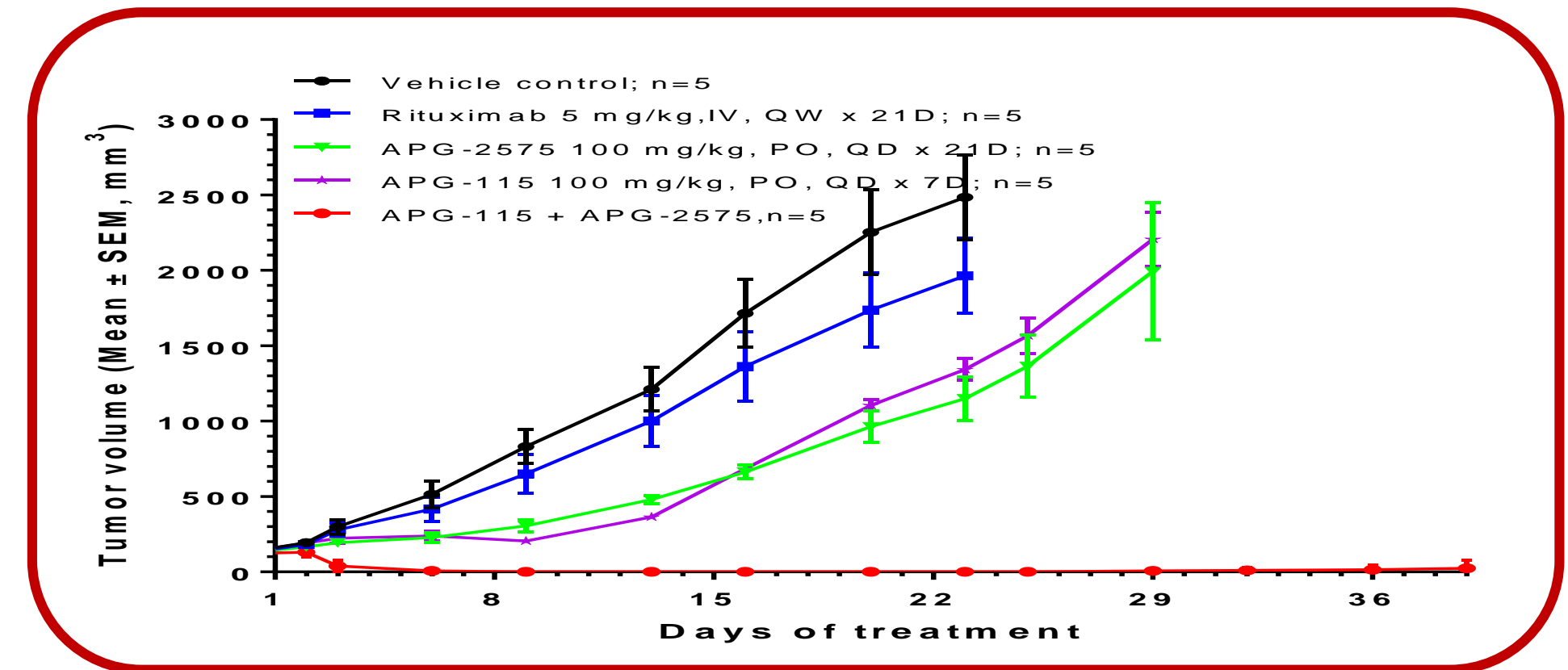
# Synthetic Lethality

## Combination of APG-115 + APG-2575

- “Synthetic lethality” describes a strategy where blocking two mutations result in cell death, but the cancerous cells only has one mutations. By artificially inducing a second mutation the medicine can induce cancerous cell death.

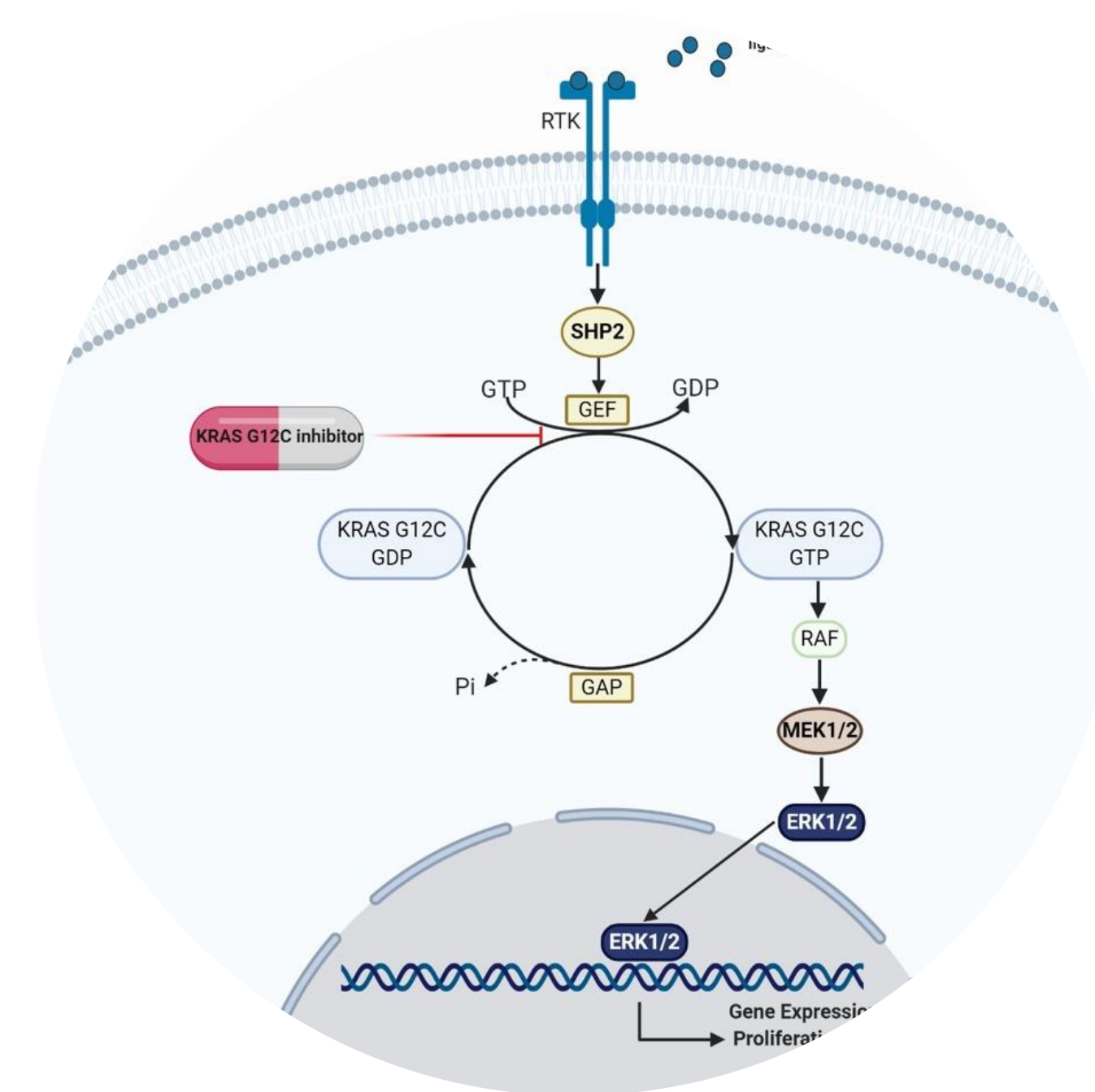


### Complete Response in Animal Tumor Models



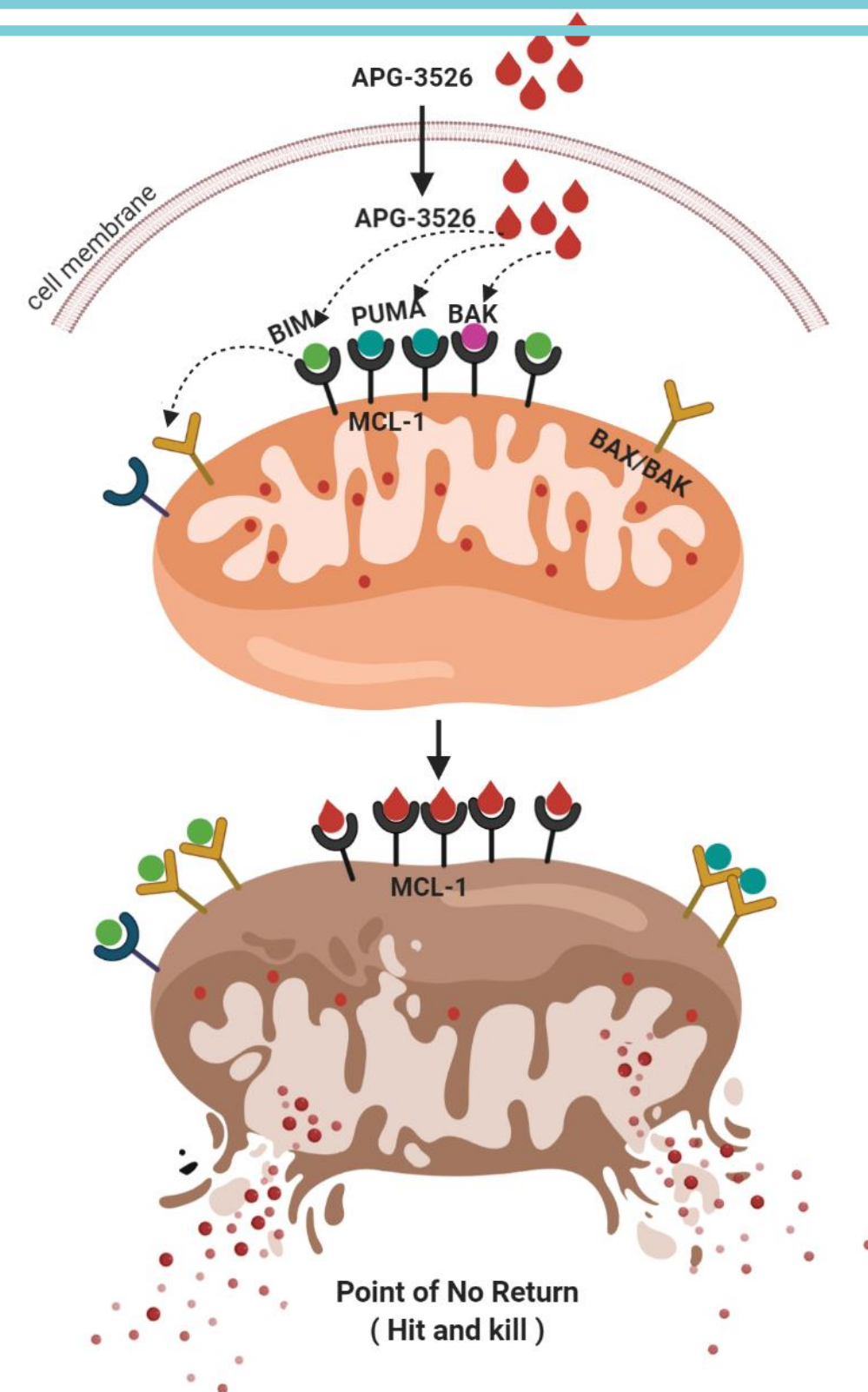
# Pre-Clinical Asset

MCL-1 inhibitor/ EED Selective/ BCR-ABL

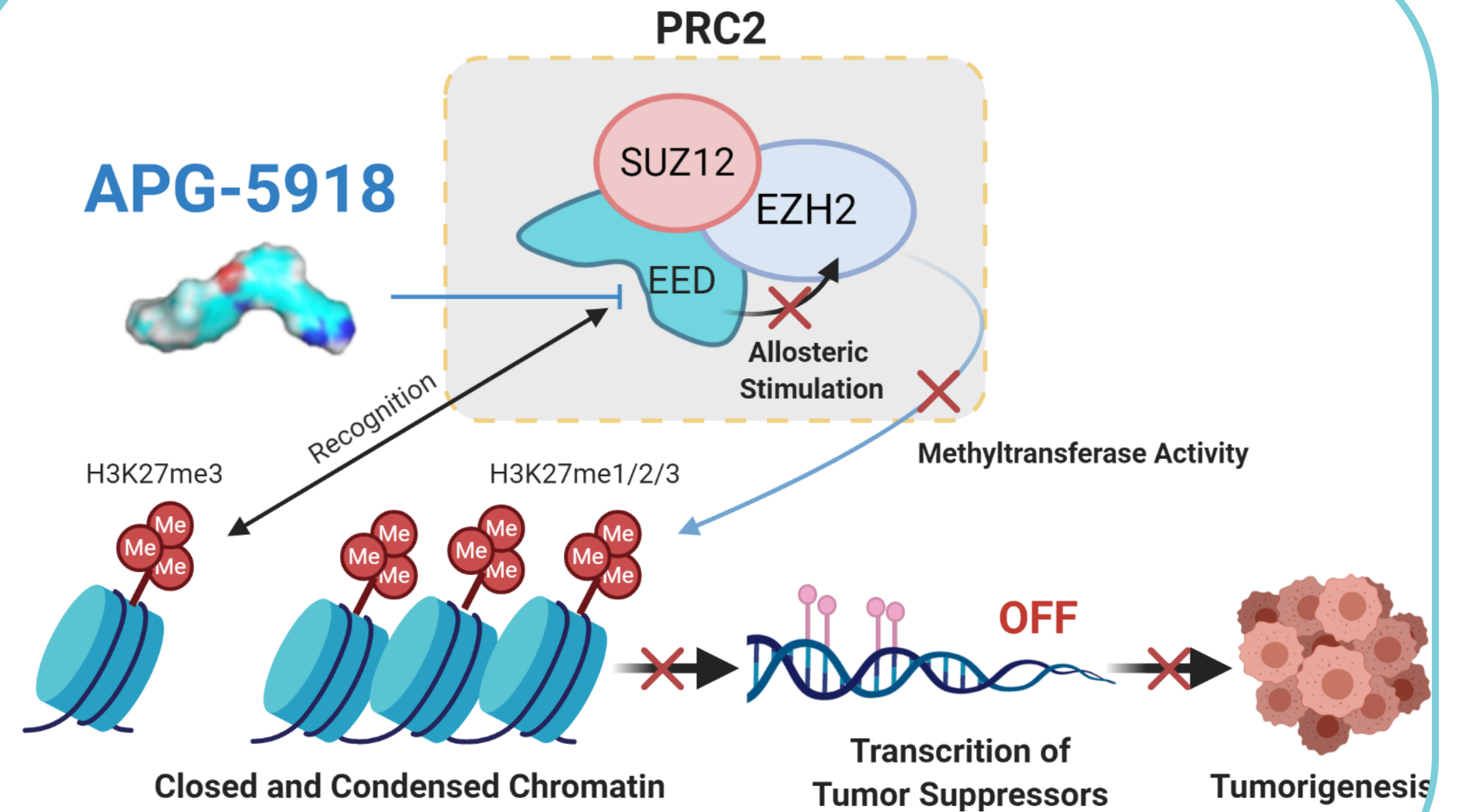


# MoA of Mcl-1 Inhibitor and EED Inhibitor

## Mcl-1 Inhibitor

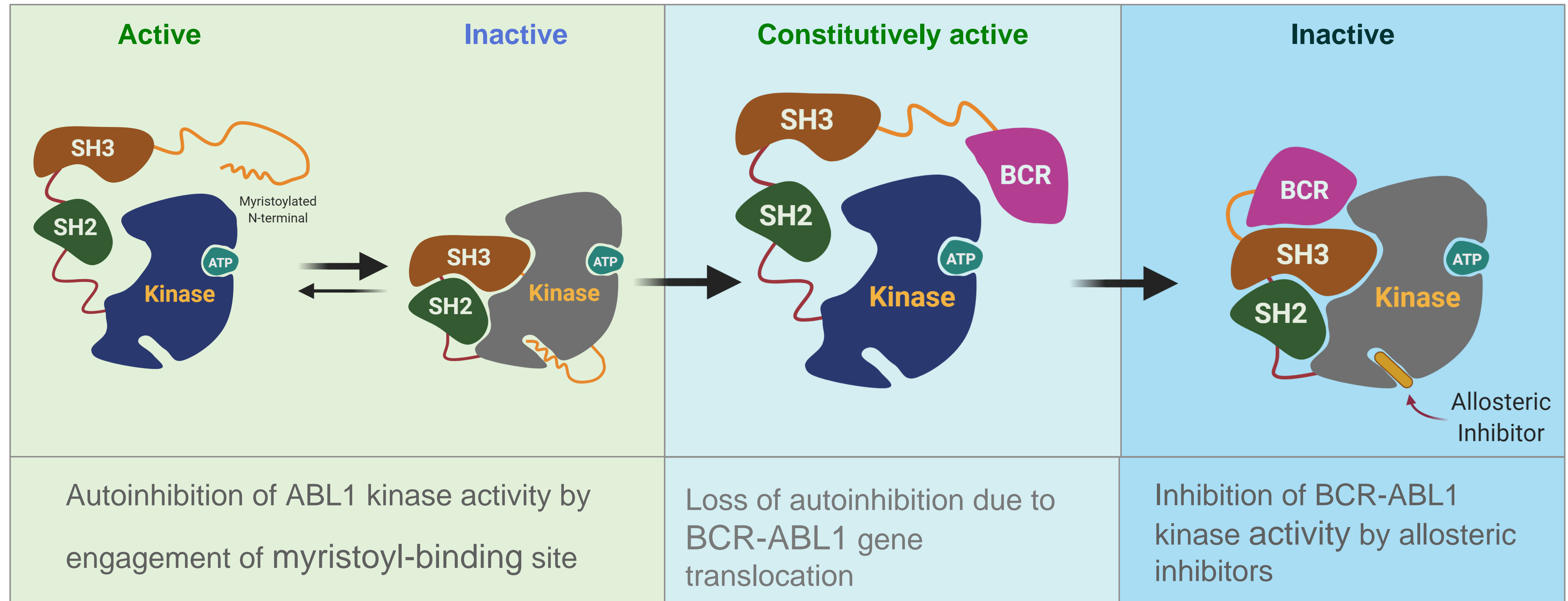


## EED Inhibitor



# MoA of BCR-ABL1 Allosteric Inhibitor

Inhibition of BCR-ABL1 kinase activity by allosteric inhibitors

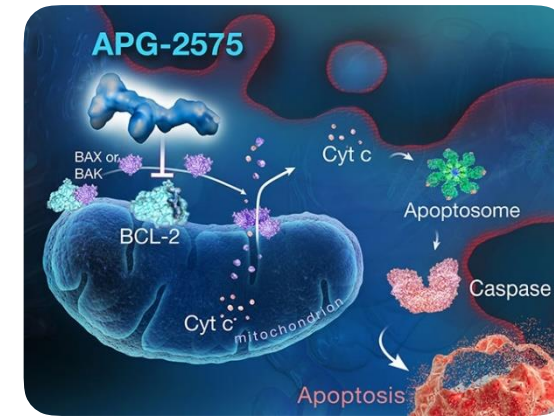


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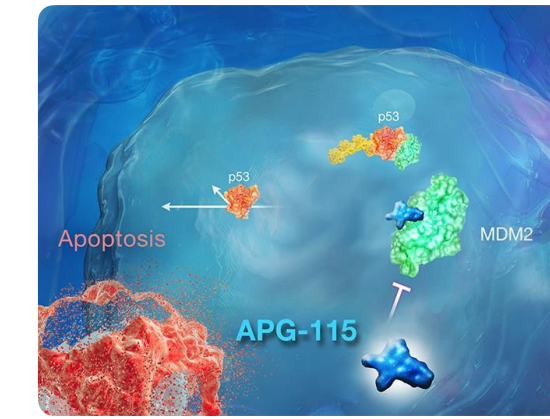
Source: N Engl J Med. 2019 Dec 12;381(24):2315-2326. J Med Chem. 2018 Sep 27;61(18):8120-8135. Cancer Res. 2012 Oct 1;72(19):4890-5.



## Strategic Alliances



BCL2

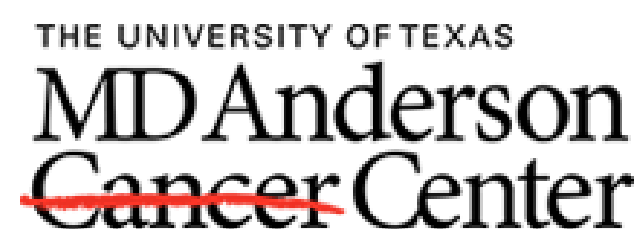


MDM2-p53



- Entered a global clinical collaboration with Acerta Pharma, the hematology research and development center of excellence of AstraZeneca;
- Ascentage Pharma will sponsor a clinical trial to study the combination of Ascentage Pharma's APG-2575, Acerta Pharma's CALQUENCE® (acalabrutinib), evaluating the efficacy and safety of this combination therapy in patients with r/r CLL/SLL;
- The study has already initiated in US with the dosing of first patient, and planned to expand in Europe, and Australia.

- Entered a global clinical collaboration with MSD;
- We will sponsor an open-label, multicenter, phase Ib/II study (NCT03611868) to evaluate the safety and efficacy of APG-115 with KEYTRUDA® (pembrolizumab) in multiple cohorts of advanced solid tumors (i, e., NSCLC, melanoma);
- The Phase II portion of the study has initiated and is expected to enroll 80 patients at multiple sites in the United States.



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Ascentage Pharma

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**Ming Guo, Ph.D.**  
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**Jeff Kmetz**  
CHIEF BUSINESS OFFICER




**Thomas Knapp**  
SVP, GENERAL COUNSEL




**Su Zhang**  
CHIEF FINANCIAL OFFICER




**James (Jim) Tripp**  
SVP, PORTFOLIO MANAGEMENT  
AND HEAD OF US OPERATIONS



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- Editor-in-chief, Journal of Medicinal Chemistry



Journal of  
**Medicinal Chemistry**



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- Dean of the University of Michigan Medical School from 1998-2006
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M.D., PhD, FASCO

- Chief of the Immuno-Oncology Service, The Lloyd J. Old Chair in Clinical Investigation at Memorial Sloan Kettering Cancer Center
- Director, Parker Institute for Cancer Immunotherapy at MSK
- Professor of Medicine, Weill Medical College of Cornell University



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- Member, Board of Directors, Tesaro



**Arul Chinnaiyan**

M.D., PH.D.

- Howard Hughes Medical Institute Investigator
- S.P. Hicks Endowed Professor at the University of Michigan Medical School



# IP Portfolio for Major Clinical Compounds

Core Compound	Patent Type	Year Patent Expires
APG-1252	Product (Core compound structure)Process; Formulation; Combination; Use	2034
APG-2575	Product (Core compound structure); Combination; Process; Use	2037
APG-115	Product (Core compound structure); Process; Combination; Use	2035
APG-1387	Product (Core compound structure); New indication; Combination; Use	2033
HQP1351	Product (Core compound structure); Process; Combination; Use; Formulation	2031



# Investment Highlights



Global leader  
in apoptosis targeting  
therapy development



Product pipeline with  
first- and best-in-class  
potential



Compelling combination  
opportunities with  
significant upside potential



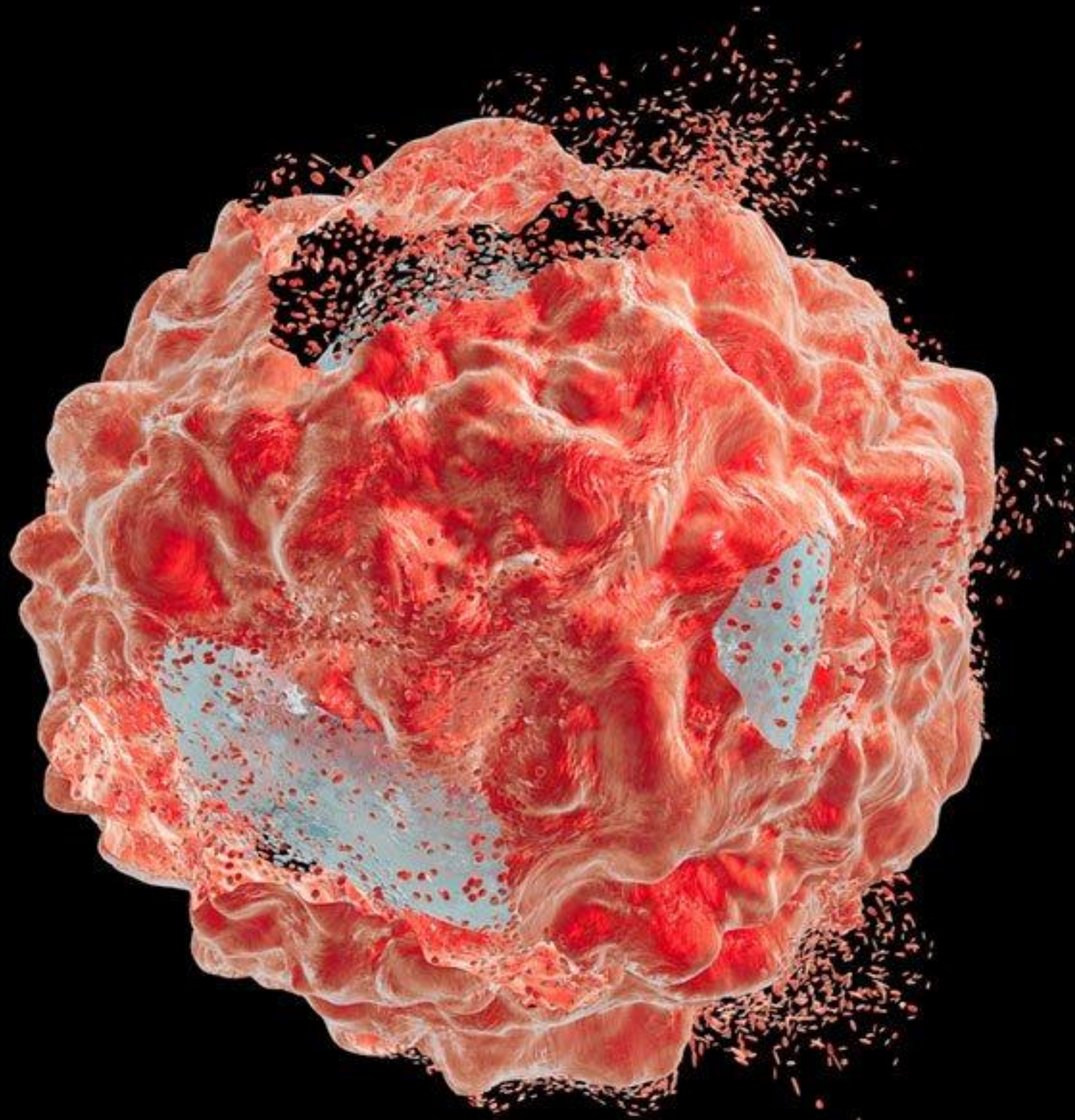
Strong global  
intellectual property  
portfolio



Experienced and  
visionary management  
team and talents



Global Collaboration with  
Leading Companies and  
Institutions



# Ascentage Pharma Group

*Advancing Therapies That  
Restore Apoptosis*