

# Precision Oncology for Therapies of Tomorrow

Aptose Biosciences is a science-driven clinical-stage biotechnology company developing first-in-class targeted agents to address the unmet clinical need in chronic lymphocytic leukemia (CLL), non-Hodgkin's lymphoma (NHL), acute myeloid leukemia (AML), myelodysplastic syndrome (MDS), and other hematologic malignancies.

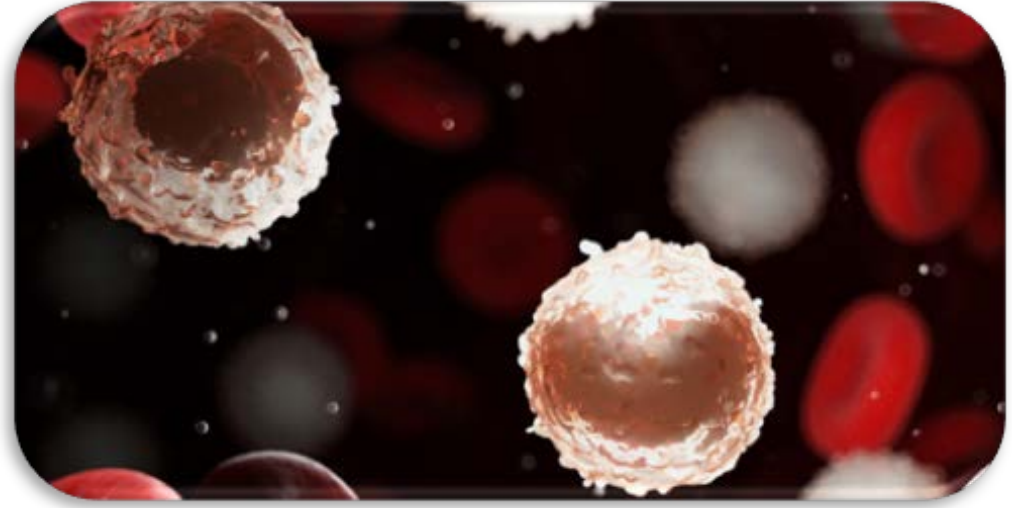
## Aptose Annual Shareholders Meeting

NASDAQ: **APTO**  
TSX: **APS**

[www.apdose.com](http://www.apdose.com)

June 02, 2020





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# During Past Year Aptose Expanded Leadership Team

*...building best possible team at all levels*

**Dr. William G. Rice, PhD**

Chairman, President &  
Chief Executive Officer



**Mr. Gregory Chow**

Executive Vice President &  
Chief Financial Officer



**Dr. Jotin Marango, MD, PhD**

Senior Vice President &  
Chief Business Officer



**Dr. Rafael Bejar, MD, PhD**

Senior Vice President &  
Chief Medical Officer



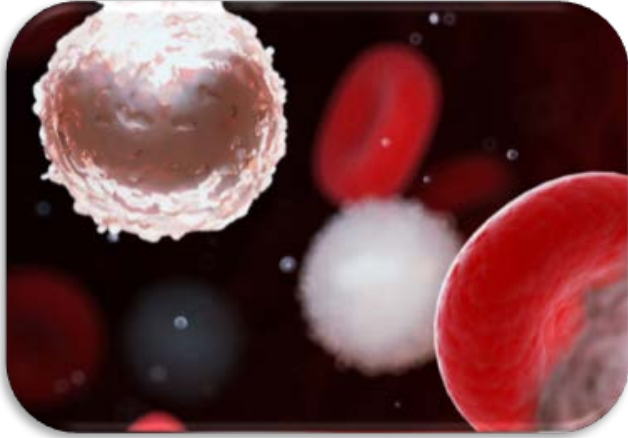
**Mr. Victor Montalvo-Lugo, MS**

Vice President of Clinical Operations

# During Past Year Aptose Established Financial Stability to Support Development of Our Clinical Assets

- Completed **Two Major Financings** During 2019
  - Raised \$21.275 million May 2019
  - Raised \$74.175 million December 2019
  - Built Strong Institutional Participation
- Established New \$200 million Shelf Registration
- Established New \$75 million At-the-Market (ATM) Facility (Q2-2020)
- Maintained and Strengthened Strong Banking Relationships
- Enhanced Research Analyst Coverage





# APTO-253

## Advanced Clinical Development

### Phase 1a/b Ongoing

Small Molecule MYC Inhibitor

For the Treatment of AML

1. [MYC dysregulation](#) is key driver of AML, certain B-cell cancers and solid tumors
2. [MYC gene expression](#) potentially inhibited by APTO-253
3. [Ph1a/b](#) trial for [AML/MDS](#) ongoing with APTO-253
4. APTO-253 first agent to inhibit MYC expression and well tolerated in patients

# APTO-253

## Update for Ongoing Phase 1b Dose Escalating Clinical Trial

- ✓ • Dose Level 1 (20mg/m<sup>2</sup>) Completed 1 AML Patient
- ✓ • Dose Level 2 (40mg/m<sup>2</sup>) Completed 1 MDS Patient
- ✓ • Dose Level 3 (66mg/m<sup>2</sup>) Completed 3 AML Patients
- Dose Level 4 (100mg/m<sup>2</sup>) Ongoing 3 Patients Required
  - 1 Completed 28d Cycle

- To date, well-tolerated & no drug-related SAEs
- Observed suppression of MYC expression in peripheral blood cells at all dose levels and with AML and MDS patients
- Plan to dose escalate to boost exposure between dosing (7 day period)

# **COVID-19 Creates Headwinds to Program:**

## **Risks to APTO-253 Clinical Trial**

- **IV Infusion is Less Desired by Clinical Sites During the COVID-19 High Risk Period**
  - Clinical Sites reluctant to house immunocompromised AML/MDS patients in clinics for hospitalization or extended periods of time
- **COVID-19 May Result in Less Data Available at ASH 2020 Conference**
  - Efforts underway to boost enrollment

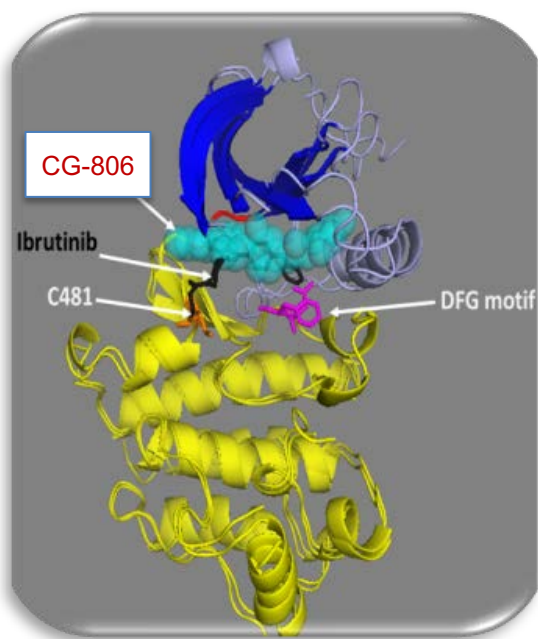
# CG-806

## 1<sup>st</sup>-in-Class

## Oral Kinase Inhibitor

🔗 Mutation Agnostic FLT3 Inhibitor

🔗 Mutation Agnostic rBTK Inhibitor



- ❑ Oral, small molecule “**reversible**” kinase inhibitor
- ❑ Cluster-selective with **highly unique kinase inhibitory profile**
- ❑ Developing across **spectrum of hematologic malignancies**
  - **lymphoid** malignancies (**CLL** & NHL)
  - **myeloid** malignancies (**AML** & MDS)
- ❑ Ongoing trial **Ph1a/b for CLL** and lymphoid malignancies as rBTKi
- ❑ Planning trial **Ph1a/b for AML** and myeloid malignancies as FLT3i



# Properties & Trial Design Minimize the Potential Impact of COVID-19

## *Support Maintenance of Timelines for CG-806 Clinical Programs*

- **“Essential” Treatment** is Required for Critically Sick Cancer Patients
  - CG-806 is being developed for **CLL**/NHL lymphoid cancers and **AML**/MDS myeloid cancers
- **Oral Administration** Places Less Strain on Clinic / Hospital Staff / System / Patients
  - *Oral* administration does not require hospital stays
  - Capsules can be shipped directly to patients and vital signs and blood samples can be collected remotely
- **Safety Profile** Thus Far Reduces Risk to Patients
  - Minimizes number of non-essential clinic visits for additional supportive care
- **Remote Monitoring** Using eDiary for Data Collection Avoids Hospital Visits
- **Study Design** Supports Patient Accrual and Minimizes Risks
  - Scans in B-cell cancer patients only every two cycles
  - Leveraging both Specialty Regional Cancer Clinics and Large Institutional Centers

# CG-806 Phase 1 Clinical Development Plan for Patients with Lymphoid (CLL) and Myeloid (AML) Malignancies

CLL & NHL  
Lymphoid

## 1<sup>st</sup> : Phase 1a/b Ongoing in Patients with R/R CLL & NHL

- Seek to define safety, tolerability, PK and PD properties and RP2D in CLL/NHL patients
- Seek responses in CLL/NHL patients

R/R AML patients are acutely ill, and we did not wish to dose sub-therapeutically  
During CLL trial, identified a dose likely to be “therapeutically active” for AML patients

Pending FDA  
Approval

AML  
Myeloid

## 2<sup>nd</sup> : Perform Phase 1a/b : R/R AML

- Selected starting dose for recommendation and submitted new IND for AML
- Plan to define safety, tolerance, PK, PD and RP2D
- Seek responses in AML patients

## Dose Escalation Phase

- Administered oral capsules
- Twice daily on a 28-day cycle
- Plan to perform 6 dose levels
- Accelerated titration design
- Planned expansion cohorts



# CG-806 Now in Dose Level 4 of Phase 1a/b Clinical Trial in CLL/NHL

## Dose Level 1 (150mg BID for 28d) Completed



Only **One Patient** Required in Dose Level 1

- R/R-CLL/SLL with TP53 mutation ; Heavily pretreated
- Challenging Case with TP53 mutation – No DLTs and in Cycle 10 (now dose escalated)

## Dose Level 2 (300mg BID for 28d) Completed



Only **One Patient** Required in Dose Level 2

- R/R-CLL with unmutated IGHV ; Marrow involvement, neutropenia and thrombocytopenia
- Highly complicated disease to manage – No DLTs and completed Cycle 4

## Dose Level 3 (450mg BID for 28d) Completed



**Three Patients** Required in Dose Level 3 – 3 Patients completed Cycle 1

- Two Follicular Lymphoma and one SLL patients

## Dose Level 4 (600mg BID for 28d) Ongoing

**Three Patients** Required in Dose Level 3

# CG-806 Delivered Clinical Evidence of Safety, Pharmacologic Activity and Favorable Oral Pharmacokinetics

- Patient **Dose Escalation** Advancing Efficiently

- Dose Levels 1, 2 and 3 completed
- Dose Level 4 underway

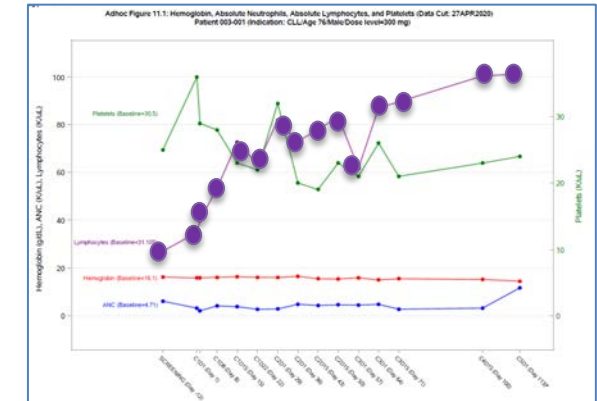


- Evidence of **Safety** and **Tolerability** to Date

- No QTc or atrial fibrillation cardiovascular events to date

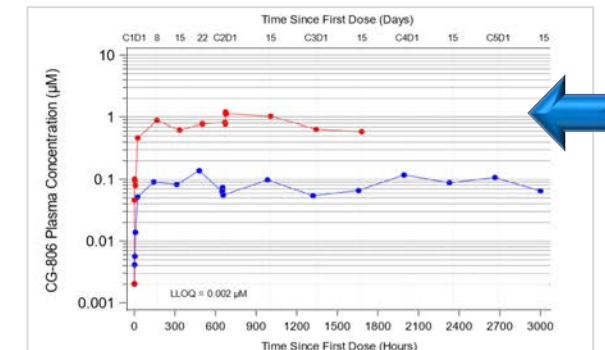
- Evidence of **Pharmacologic Activity** as Early as Dose Level 2

- **Target Engagement:** Plasma inhibits P-BTK, P-SYK, P-ERK, P-PDGFR $\alpha$  in EOL1
- **Lymphocytosis:** BTK inhibition in CLL promotes exfiltration



- Favorable **Steady-State Plasma Exposure Levels**

- Plan to **Continue Dose Escalation** and Seek Clinical POC



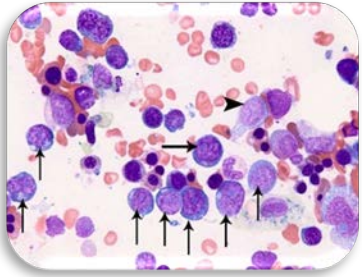




## **CG-806 : A New Class of Drugs**

- **Only BTK Inhibitor that also Inhibits FLT3**
- **Developing for CLL and AML**

# CG-806 for the Treatment of AML & Myeloid Malignancies

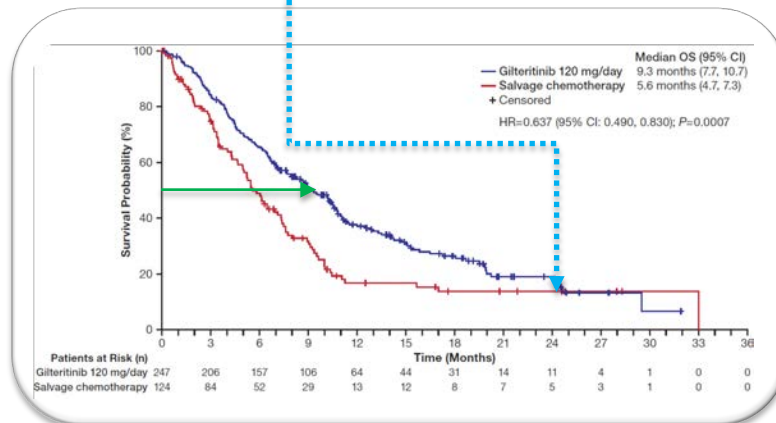
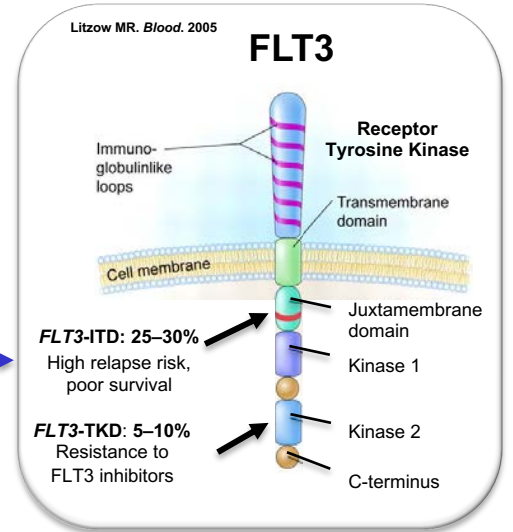


## Deadly Cancer of Blood/Bone Marrow (Orphan Disease)

- ~21,450 diagnosed this year / ~10,920 deaths this year<sup>1</sup>
- The 5-year survival rate for patients with AML approximately 28.3%

## Limitation of Current FLT3 Inhibitors and Other Agents

- **FLT3-ITD** mutation is key driver in **25-35% of AML patients**<sup>2,3</sup>
- Current “Dirty” agents (Midostaurin®, etc.) are limited → **Toxicity**
- Current “Selective” (Gilteritinib®, Quizartinib®) agents not durable → **Resistance**
- Current agents susceptible to mutations in TP53, Ras, FLT3 (ITD/TKD/GK)



## Desperate Need for Improved AML Agents → CG-806

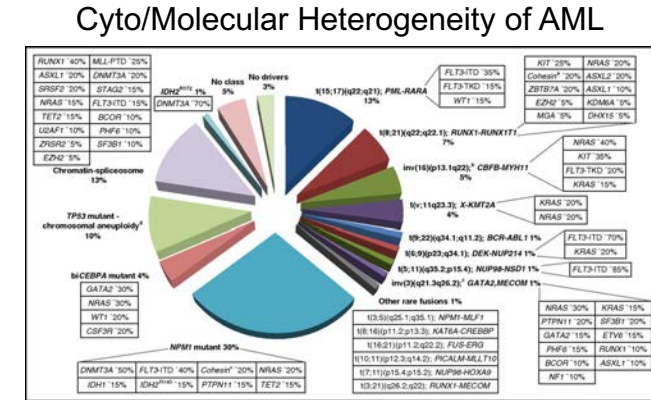
- CG-806 potently inhibits **all WT and mutant forms of FLT3**: ITD/TKD/GK/WT
- CG-806 suppresses **multiple oncogenic signaling pathways** to avoid resistance
- CG-806 retains activity in presence of **diverse mutational background**
- CG-806 **combines effectively** with other therapies, i.e. venetoclax/venclerxa®

<sup>(1)</sup> American Cancer Society : <sup>2</sup> Cancer. 2014 July 15; 120(14): 2142-2149 : <sup>3</sup> Blood 2016;128(5):686-698.

# Developing CG-806 for the Treatment of AML

- **Strong Rationale to Develop for AML with High Potential Value:**

- Broadly potent against AML cells
  - Patients with mutated FLT3, TP53, IDH1, IDH2, SRF2, ASXL1 and RAS
  - Patients with WT-FLT3 (approximately 70% of R/R AML patients)
- More potent than other FLT3 inhibitors on >200 AML patient samples
- Delivers cures in xenograft models of human AML without toxicity



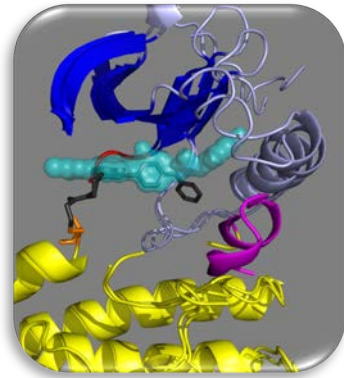
- **Phase 1 Planned : R/R AML Patients with Unmet Needs**

- Patients who failed other FLT3 inhibitors
- Patients who failed IDH-1 inhibitors
- Patients who failed venetoclax
- Patients with mutated p53, mutated RAS
- Patients with wild type-FLT3
- Patients unfit for intensive therapies

- Plan to initiate dosing with an active dose
- &
- Rapidly differentiate CG-806 from other FLT3i's

# Developing CG-806 Broadly Across Hematologic Malignancies

- **Uniquely and Selectively Inhibits Clusters of Kinases**
  - Targets kinases that are drivers of hematologic malignancies (lymphoid and myeloid)
  - Yet, avoids kinases generally associated with toxicity
- **Phase 1 Ongoing in R/R CLL & NHL Lymphoid Cancer Patients**
  - Targeting BTK and multiple survival pathways to treat patients failing other agents
  - Observed safety, pharmacologic activity and predictable PK characteristics
  - Continuing to dose escalate and seek safety, PD responses and efficacy responses
- **Phase 1 Planned in R/R AML Myeloid Cancer Patients**
  - Targeting FLT3 and multiple survival pathways to treat patients failing other agents
  - Plan to initiate dosing at active level; Potential for rapid development and value creation





# Additional Accomplishments

- **Website Upgraded**
- **Generic Naming of CG-806 Ongoing**
- **Scaled CG-806 API Manufacturing**
- **Scaled CG-806 Capsule Manufacturing**
- **Instituted Major Upgrades to Clinical Operations**
- **Aspire to Error-free Execution (Avoid Unforced Errors)**
- **Developing New Oral Formulations for CG-806 and APTO-253**
- **Maintaining Strong Financials – Allow Clinical Data to Emerge & Build Value**
- **Presented Data at Key Medical Conferences: EHA, ESH, ASH, AACR**

DRUG	TARGET	RIGHTS	INDICATIONS	Preclinical Stage	Clinical Proof-of-Concept	Pivotal Stage
<b>CG-806</b>	Pan-BTK	Aptose: WW CG: Korea	CLL NHL	B-Cell Malignancies		
<b>CG-806</b>	Pan-FLT3	Aptose: WW CG: Korea	AML MDS	AML / MDS Planned		
<b>APTO-253</b>	MYC	Aptose: WW	AML MDS	AML / MDS Single Agent		
<b>APL-581</b>	BRD4/JAK	Aptose / Ohm	Hematologic Cancers	AML Single Agent		

## 2020 Anticipated Catalysts

<b>CG-806</b>	1H: Seek FDA allowance for AML trial 2H: Seek clinical activity in AML patients 2H: Seek clinical activity in B-cell cancer patients 1-2H: Presentation of clinical data during EHA (B-cell) and ASH (B-cell & AML)
<b>APTO-253</b>	1-2H: Continue dose escalation in AML/MDS patients 2H: Explore additional cancer indications 2H: Presentation of clinical data during ASH

Patients, Their Families, Care Givers

Dedicated Employees

Shareholders

**We Thank You!**

