

# Corporate Presentation

May 2020

NASDAQ: ABUS

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

# Forward-Looking Statements

This presentation contains forward-looking statements within the meaning of the U.S. Private Securities Litigation Reform Act of 1995 and Canadian securities laws. All statements that are not historical facts are hereby identified as forward-looking statements for this purpose and include, among others, statements relating to: the potential for HBV to have a larger market opportunity than HCV; our ability to meet a significant unmet medical need; the sufficiency of our cash and cash equivalents to extend into mid 2021; our expectation for multiple 60 mg dose and 90 mg single-dose data in the second half of 2020; the potential for an oral HBsAg reducing agent and potential all oral combination therapy; our objective to complete IND enabling studies for AB-836 in the second half of 2020; the potential for AB-836 to be low dose with a greater therapeutic window and to address known capsid resistant variants T33N and I105T; the potential for AB-836 to be once daily dosing; our expectations regarding the timing and clinical development of our product candidates; the timeline to a combination cure for HBV; our coronavirus strategy; and other statements relating to our future operations, future financial performance, future financial condition, prospects or other future events.

With respect to the forward-looking statements contained in this presentation, Arbutus has made numerous assumptions regarding, among other things: the timely receipt of expected payments; the effectiveness and timeliness of preclinical studies and clinical trials, and the usefulness of the data; the timeliness of regulatory approvals; the continued demand for Arbutus' assets; and the stability of economic and market conditions. While Arbutus considers these assumptions to be reasonable, these assumptions are inherently subject to significant business, economic, competitive, market and social uncertainties and contingencies including uncertainties and contingencies related to the ongoing COVID-19 pandemic. Forward-looking statements herein involve known and unknown risks, uncertainties and other factors that may cause the actual results, events or developments to be materially different from any future results, events or developments expressed or implied by such forward-looking statements. Such factors include, among others: anticipated pre-clinical and clinical trials may be more costly or take longer to complete than anticipated, and may never be initiated or completed, or may not generate results that warrant future development of the tested drug candidate; changes in Arbutus' strategy regarding its product candidates and clinical development activities; Arbutus may not receive the necessary regulatory approvals for the clinical development of Arbutus' products; economic and market conditions may worsen; and market shifts may require a change in strategic focus; and the ongoing COVID-19 pandemic could significantly disrupt our clinical development programs. A more complete discussion of the risks and uncertainties facing Arbutus appears in Arbutus' Annual Report on Form 10-K, Quarterly Report on Form 10-Q and Arbutus' periodic disclosure filings which are available at [www.sec.gov](http://www.sec.gov) and at [www.sedar.com](http://www.sedar.com).

The forward-looking statements made in connection with this presentation represent our views only as of the date of this presentation (or any earlier date indicated in such statement). While we may update certain forward-looking statements from time to time, we specifically disclaim any obligation to do so, even if new information becomes available in the future.

COVID-19. In December 2019 an outbreak of a novel strain of coronavirus (COVID-19) was identified in Wuhan, China. This virus continues to spread globally, has been declared a pandemic by the World Health Organization and has spread to nearly every country in the world. The impact of this pandemic has been, and will likely continue to be, extensive in many aspects of society. The pandemic has resulted in and will likely continue to result in significant disruptions to businesses. A number of countries and other jurisdictions around the world have implemented extreme measures to try and slow the spread of the virus. These measures include the closing of businesses and requiring people to stay in their homes, the latter of which raises uncertainty regarding the ability to travel to hospitals in order to participate in clinical trials. Additional measures that have had, and will likely continue to have, a major impact on clinical development, at least in the near-term, include shortages and delays in the supply chain, and prohibitions in certain countries on enrolling subjects in new clinical trials (e.g. in Australia). It is not possible to predict if the COVID-19 pandemic will negatively impact our plans and timelines.

# Investment Highlights

Therapeutic focus – curing chronic Hepatitis B Virus (HBV) Infection

Significant  
Unmet Medical  
Need in HBV

Global HBV  
prevalence double  
that of HCV,  
potential for  
larger market  
opportunity

Goal of HBV  
Functional  
Cure

Undetectable HBV  
DNA and HBsAg  
delivered through  
finite duration  
treatment with a  
combination of  
drugs with different  
modes of action

Broad  
HBV  
Portfolio

HBV assets include:  
RNAi  
Capsid Inhibitors  
PD-L1  
HBV RNA  
Destabilizers

Coronavirus  
Research  
Initiative

Focused on direct  
acting antivirals  
targeting the viral  
polymerase and  
protease

Team with  
Antiviral  
Expertise &  
Proven Track  
Record

Applying  
knowledge gained from  
HIV and HCV success to  
HBV  
and Coronaviruses

# Proven Leadership Team

Successful track records in the discovery, development, and commercialization of multiple antivirals including sofosbuvir, etravirine, rilpivirine, telaprevir and simeprevir



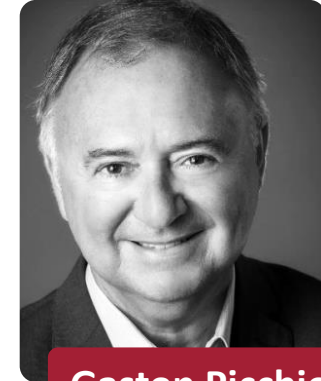
**William H. Collier**

President and CEO



**Michael J. Sofia, PhD**

Chief Scientific Officer



**Gaston Picchio, PhD**

Chief Development Officer



**David C. Hastings**

Chief Financial Officer



**Elizabeth Howard, PhD, JD**

EVP, General Counsel and  
Chief Compliance Officer



**Michael J. McElhaugh**

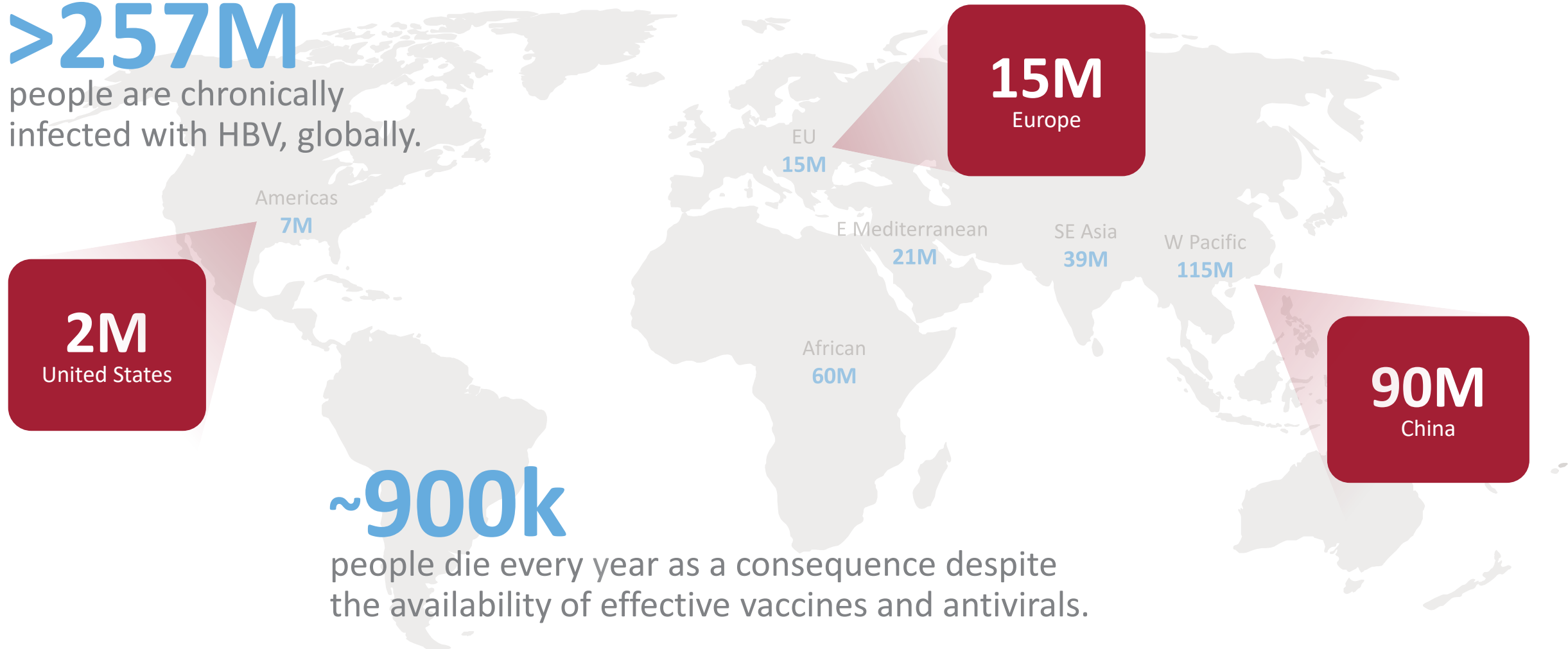
Chief Business Officer



# HBV Presents a Significant Unmet Medical Need

>257M

people are chronically infected with HBV, globally.



~900k

people die every year as a consequence despite the availability of effective vaccines and antivirals.

# Significant Opportunity to Improve HBV Cure Rates

HBV cures are achievable with today's SOC in **<5% of patients**. Sustained HBsAg and HBV DNA loss after end-of-treatment\* is rare.

\*undetectable HBsAg and HBV DNA 6 months after end-of-treatment accepted as a functional cure..

## STANDARD OF CARE THERAPIES FOR CHRONIC HBV

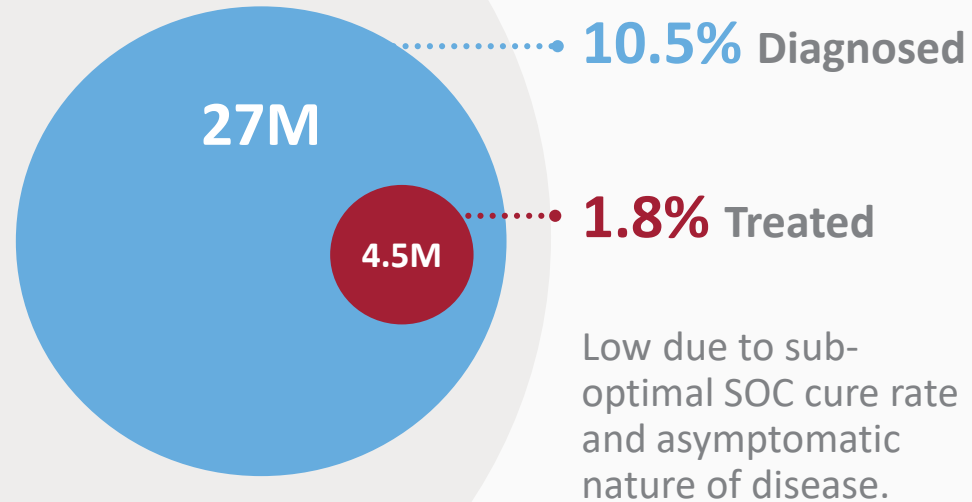
|                                     | PegIFN   | Entecavir | Tenofovir | New HBV Therapies            |
|-------------------------------------|----------|-----------|-----------|------------------------------|
| Dosing Duration                     | 48-weeks | Chronic   | Chronic   | rate of Undetectable HBV DNA |
| HBV DNA Undetectable (<60-80 IU/ml) | 7-19%    | 67-90%    | 76-93%    | +                            |
| HBsAg Loss                          | ~3-7%    | ~1-2%     | ~1-3%     | rate of HBsAg Loss           |
|                                     |          |           |           | =                            |
|                                     |          |           |           | HIGHER CURES RATES           |

Achievable **HBV Cure Rates** with Current SOC

SOC: Standard Of Care | HBsAg: HBV Surface Antigen | PegIFN: Pegylated Interferon  
Source: EASL HBV Clinical Practice Guidelines, 2017 - Pegasys, PEG-Intron, Baraclude and Viread Package Inserts

# Compelling Growth Opportunity in the **HBV Market**

**257M**  
chronic HBV



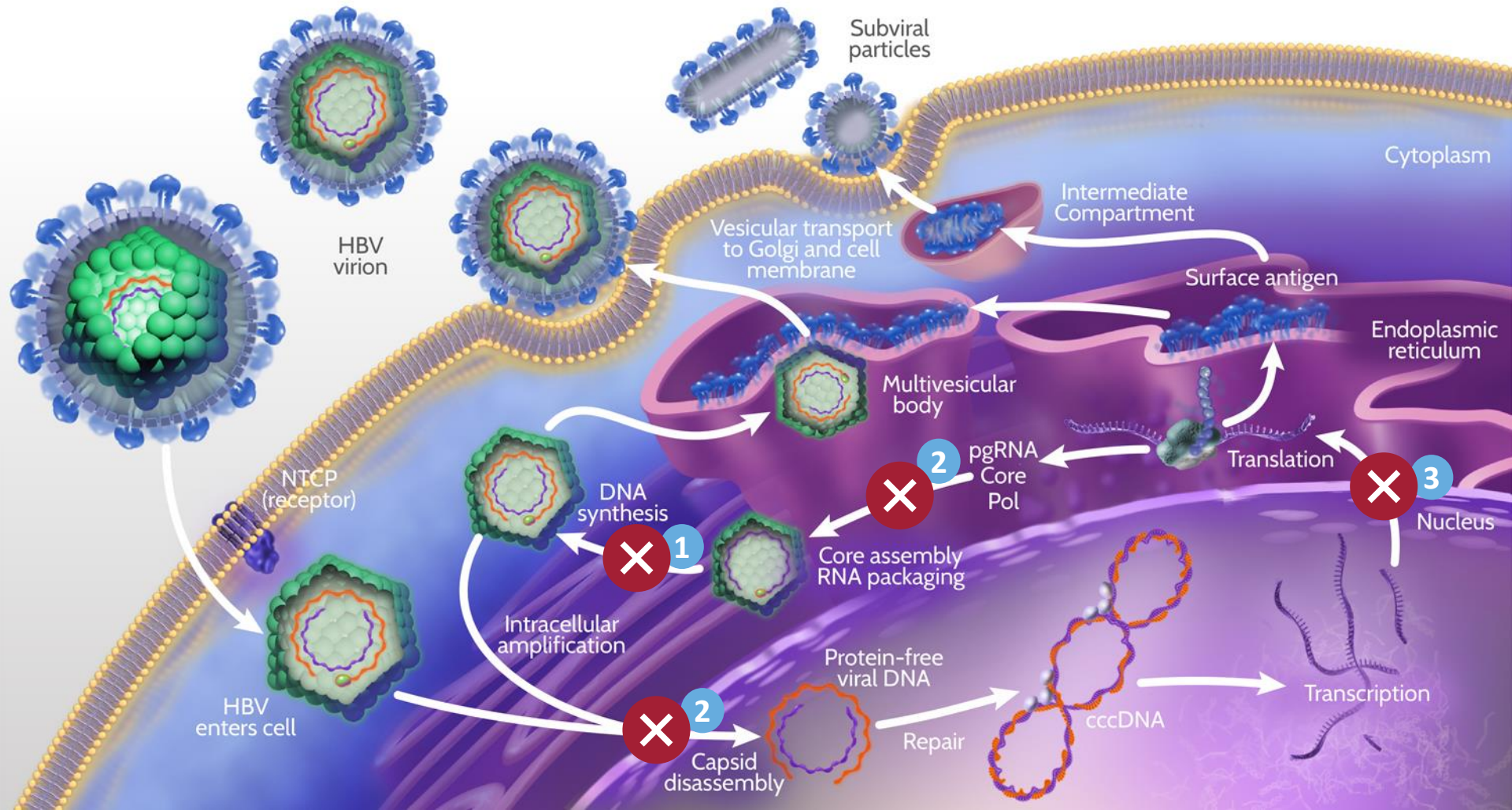
**An HBV curative regimen** would substantially increase **diagnosis** and **treatment** rates to unlock significant **market growth opportunities.**



# HBV Lifecycle Illustrates Key Points for Intervention

A combination of agents with complementary MOA **is needed to cure HBV**

1. Nucleoside Analogue
2. Capsid Inhibitor
3. RNAi & RNA Destabilizer



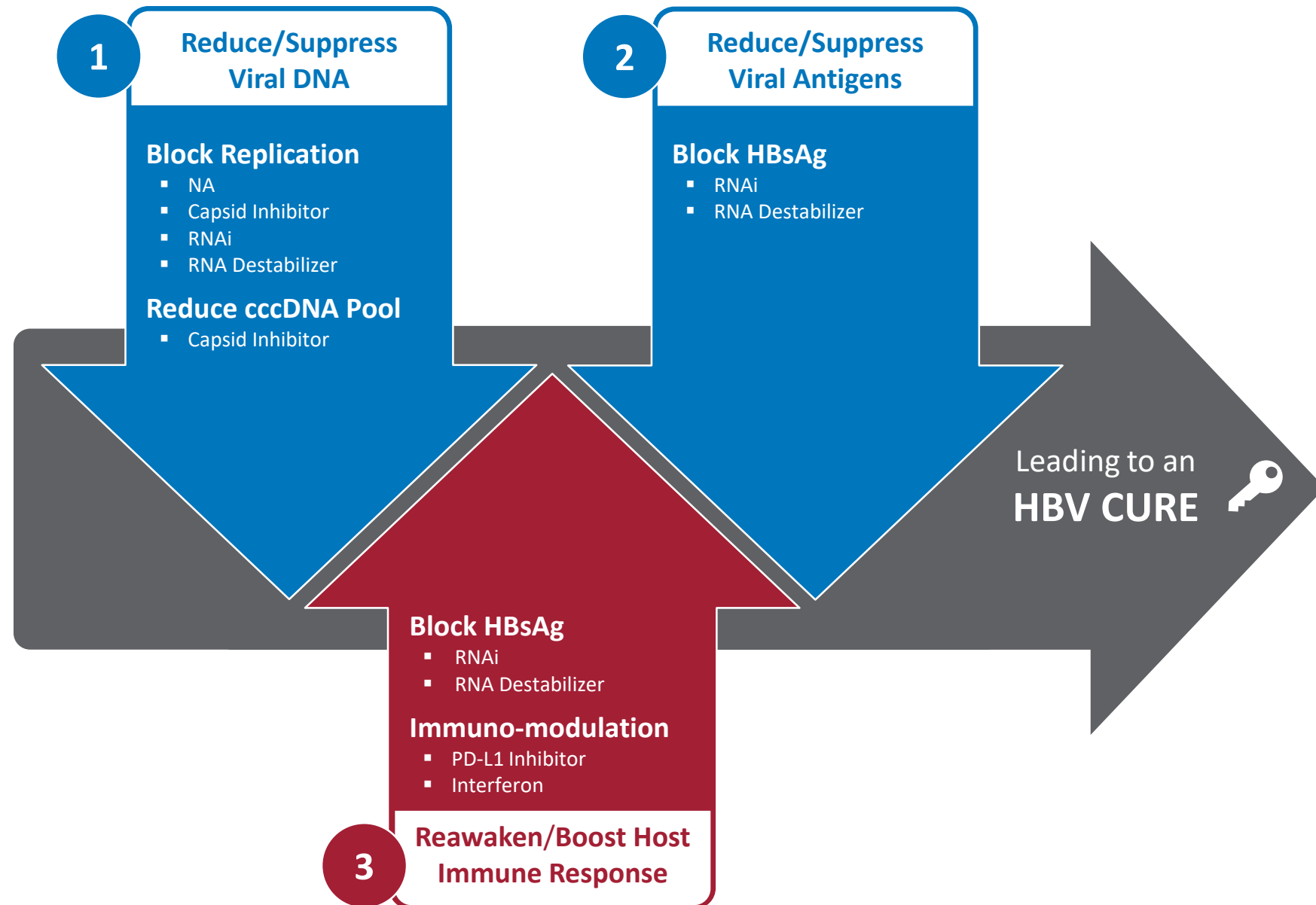


# Keys to Therapeutic Success

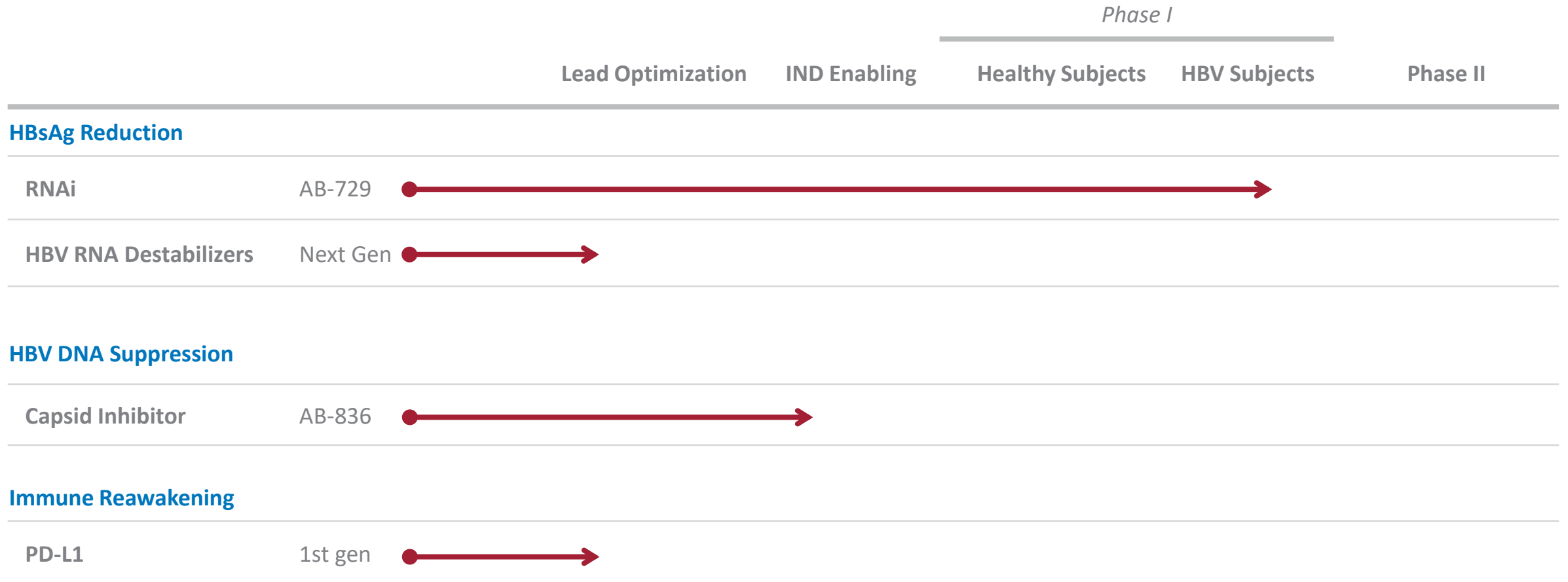
Suppress HBV DNA and viral antigens

Reawaken host immune response

Therapeutic success will require a combination of agents with complementary MOAs



# Arbutus HBV Pipeline



# AB-729

## RNAi Therapeutic

**Proprietary GalNAc-conjugate delivery technology** provides liver targeting and enables **subcutaneous dosing**



Single trigger RNAi agent targeting all HBV transcripts

Inhibits HBV replication and lowers all HBV antigens

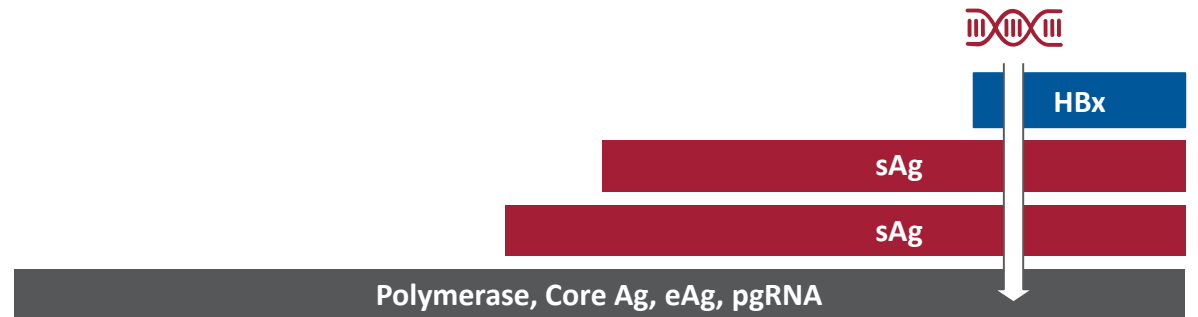
Pan-genotypic activity across HBV genotypes

Demonstrated complementarity with capsid inhibitors

Actively targets the liver

Active against cccDNA derived and integrated HBsAg transcripts

Clean profile in long term preclinical safety studies



# AB-729

## RNAi Therapeutic

**In May 2020,**  
Arbutus announced  
additional positive single-  
dose Phase 1a/1b clinical  
trial results for AB-729

**Continuous HBsAg decline** with a single 60 mg dose through week 12 with mean HBsAg decline of approximately 1.0 log matching the 180 mg cohort at week 12.

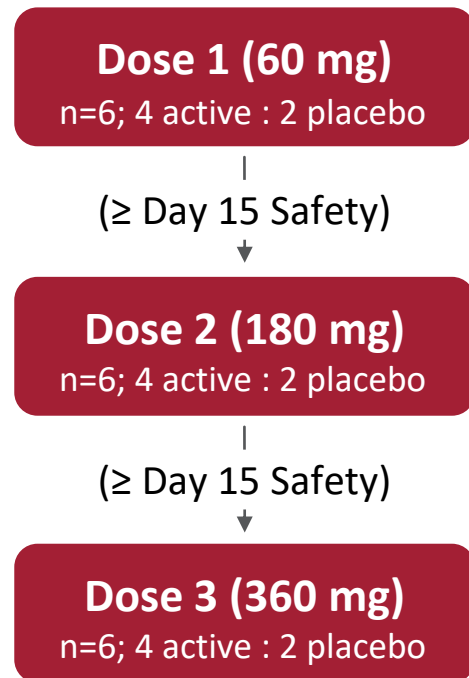
**All subjects had normal ALTs/ASTs** throughout the 12 week follow up period.

**All subjects responded to therapy** with everyone achieving at least a -0.62 log reduction in HBsAg at week 12 in the 60 mg dose group with a maximum decline of -2.14 log.

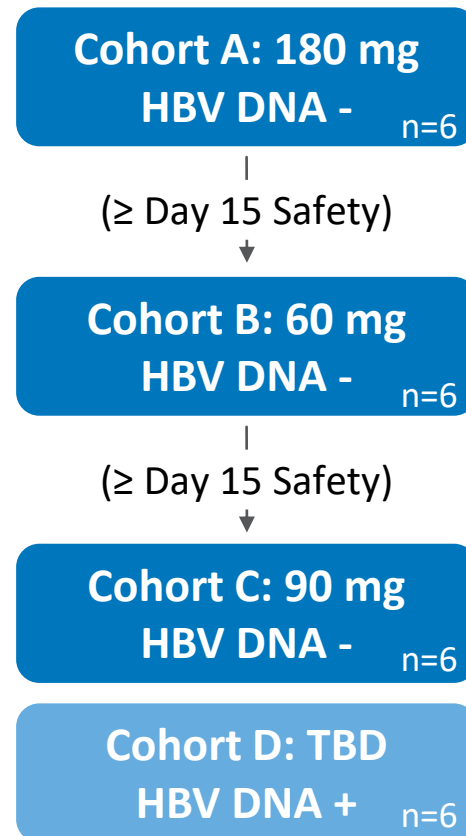
**AB-729 may provide a competitive advantage** through low dose and reduced frequency of injections.

# AB-729-001 Study Design

## Part 1: Single Ascending Dose In Healthy Subjects

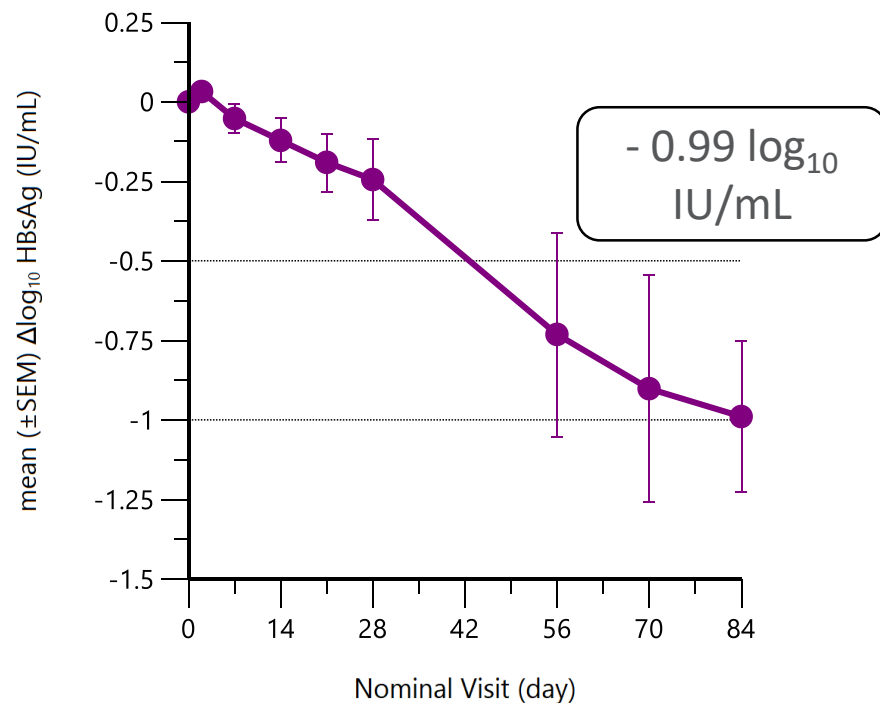


## Part 2: Single Doses In Chronic Hepatitis B Subjects

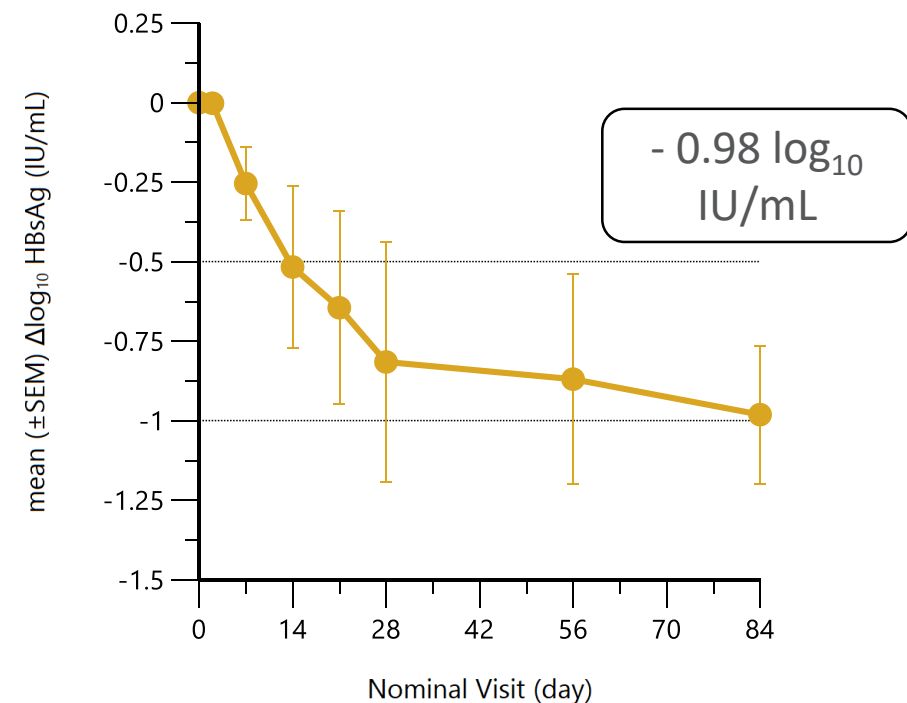


# Continuous Mean HBsAg Decline of $\sim 1 \log_{10}$ with a Single 60 mg Dose Matching HBsAg Decline of 180 mg at Week 12

**AB-729 60 mg (N=6)**



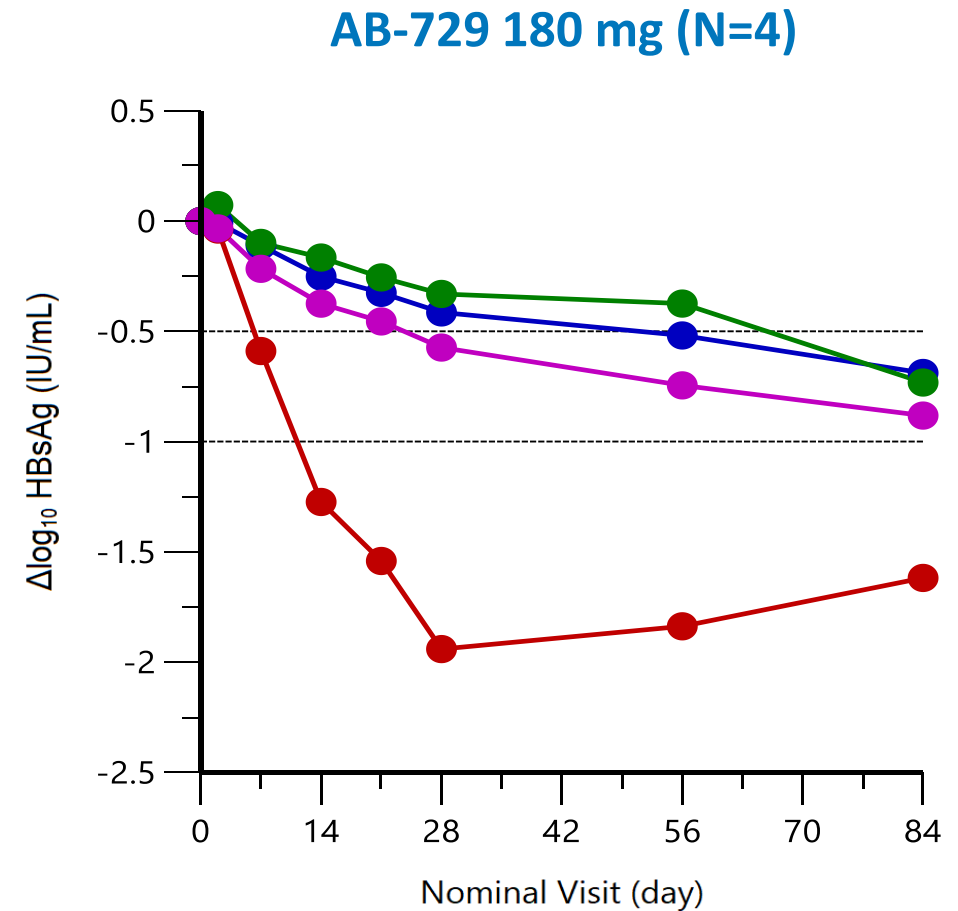
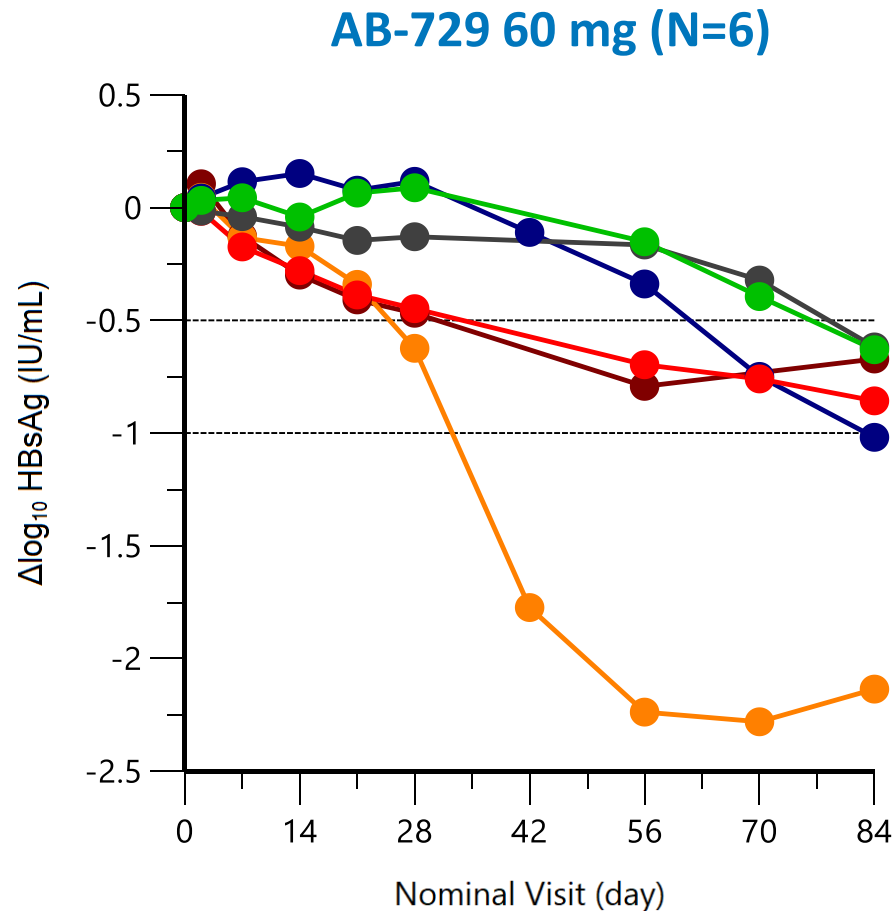
**AB-729 180 mg (N=4)**





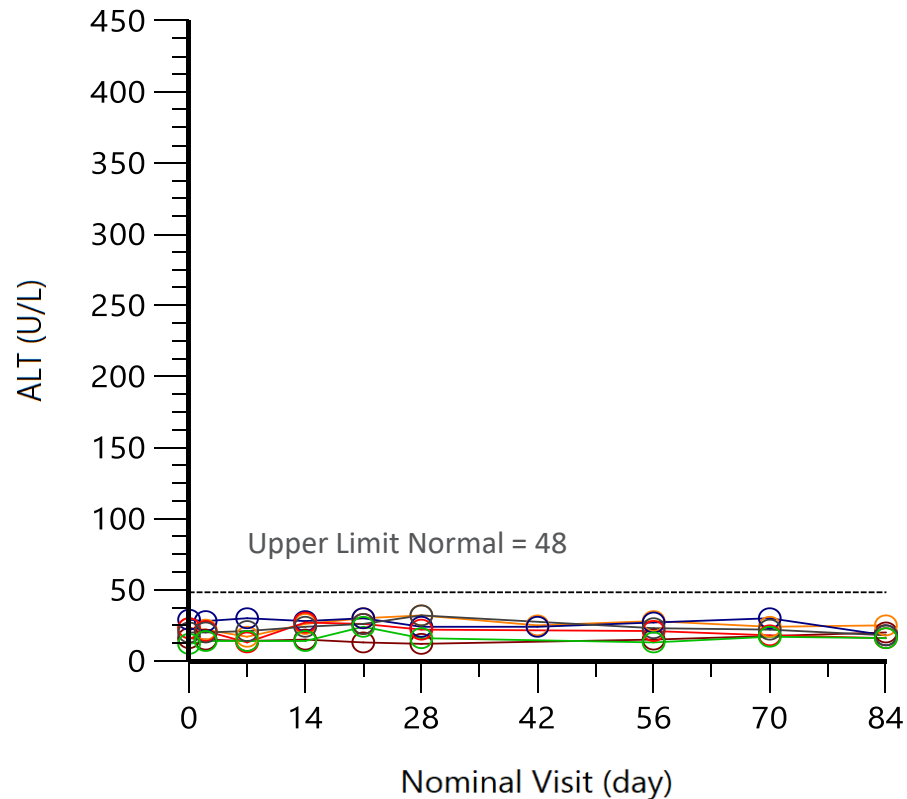
# All Subjects Responded in the 60 mg Single-Dose Cohort

Minimum HBsAg decline of -0.62 log<sub>10</sub> and maximum HBsAg decline of -2.14 log<sub>10</sub> at week 12

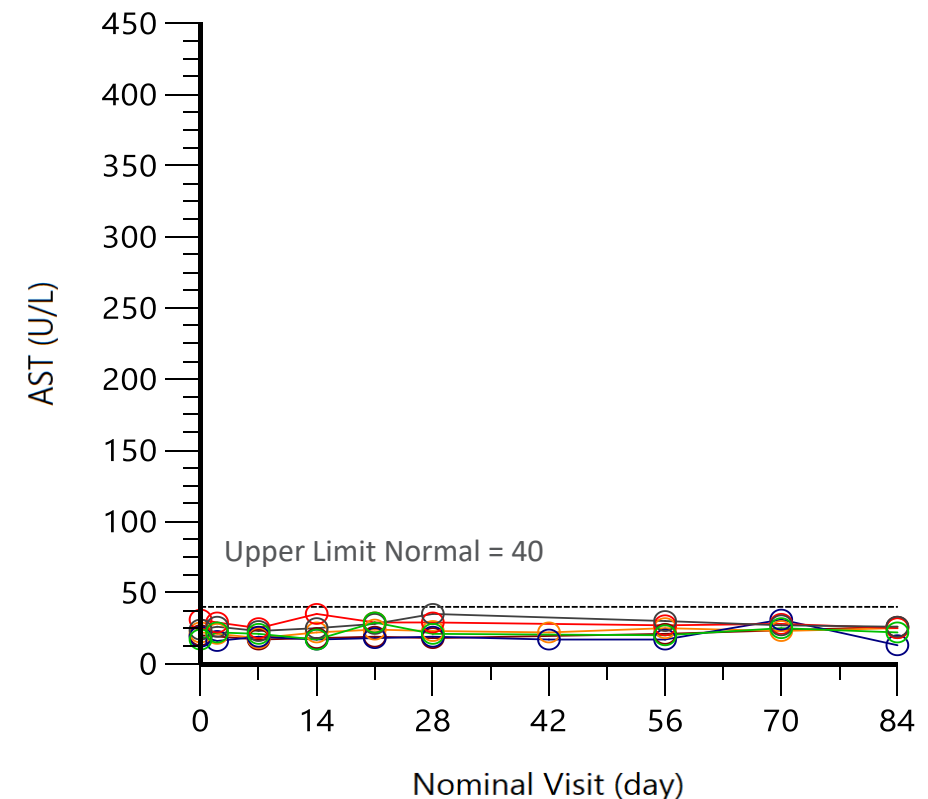


# AB-729 60 mg Single-Dose Generally Safe and Well Tolerated with Normal ALT/AST Through 12 Weeks

## Alanine Aminotransferase

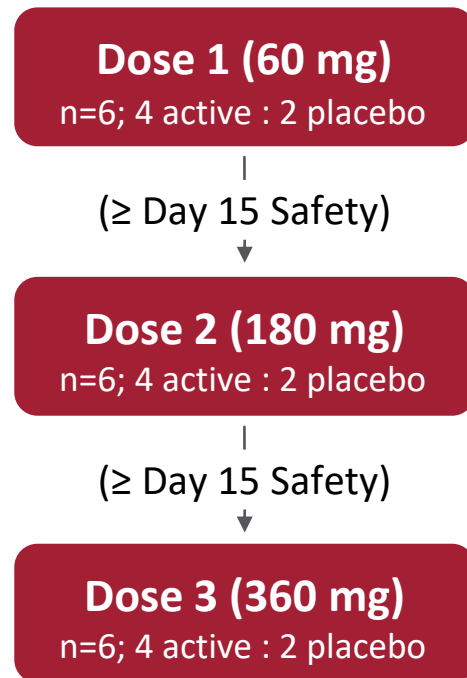


## Aspartate Aminotransferase

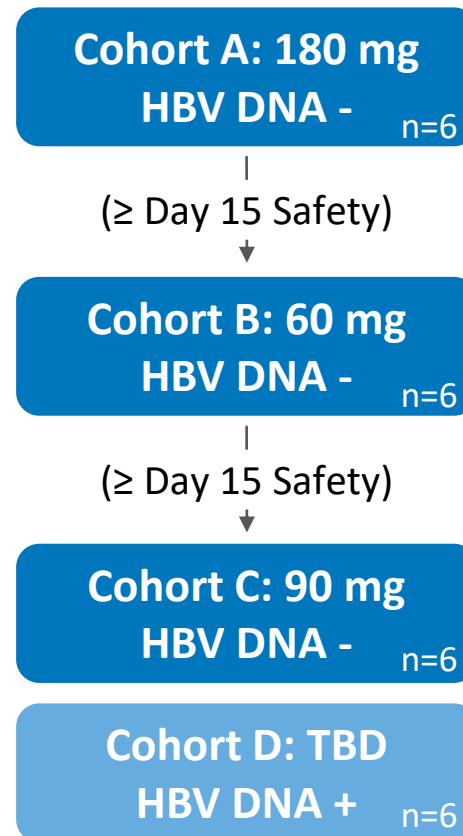


# AB-729-001 Study – Next Steps

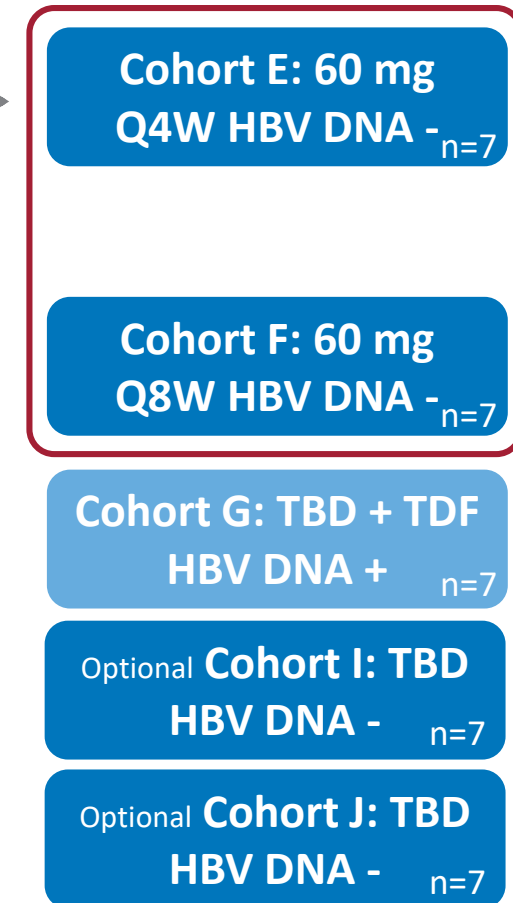
## Part 1: Single Ascending Dose In Healthy Subjects



## Part 2: Single Doses In Chronic Hepatitis B Subjects



## Part 3: Multiple Doses In Chronic Hepatitis B Subjects



# AB-836

## Capsid Inhibitor

### IND enabling studies ongoing

Potential for increased potency and enhanced resistance profile

Novel chemical series differentiated from AB-506 and other competitor compounds in the Class II capsid inhibitor space

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Leverages a novel binding site within the core protein dimer-dimer interface

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Improved intrinsic potency with  $EC_{50} \leq 10$  nM

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Active against NA resistant variants

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Potential to address known capsid resistant variants T33N and I105T

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Provides the potential for low dose and wide therapeutic window

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Projected to be once daily dosing

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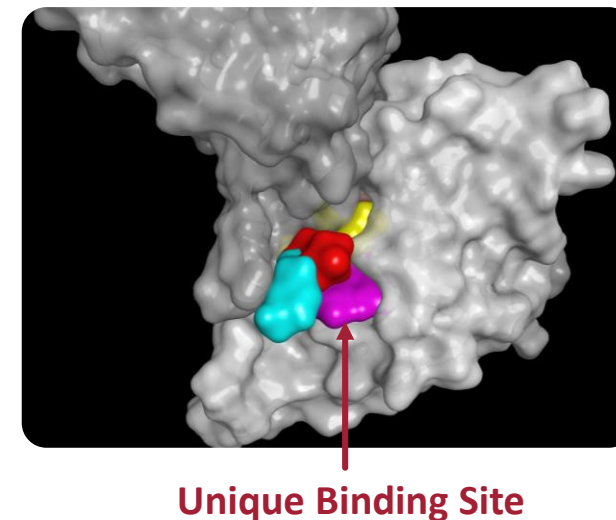
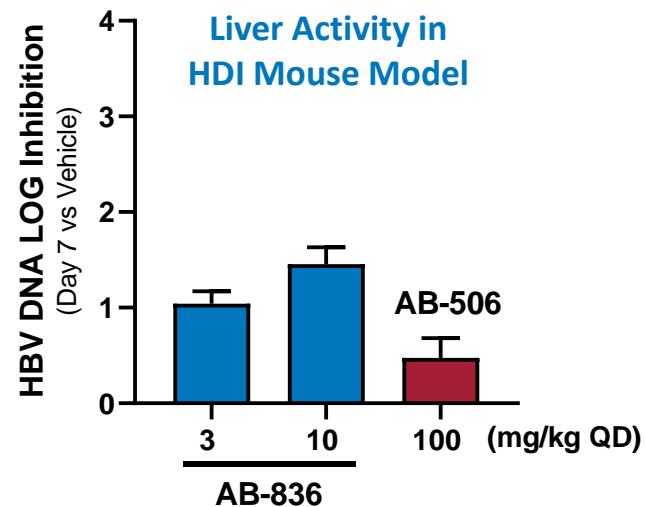
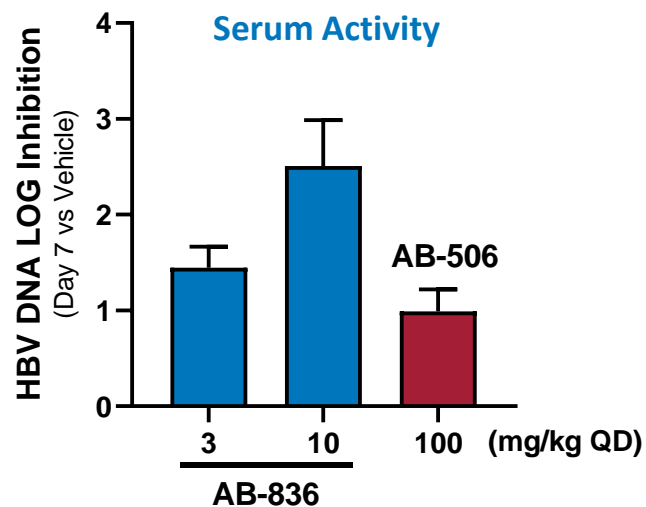
Pangenotypic

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Combinable with other MOA agents

# AB-836: A Next Generation Capsid Inhibitor

| Compound | HBV DNA / 1 <sup>o</sup> Mechanism |  |  |   | cccDNA Formation / 2 <sup>o</sup> Mechanism            | Human Serum Shift                           |
|----------|------------------------------------|--|--|---|--|---|
|          | HepDE19 (EC <sub>50</sub> μM)      | HBV infected PHH (EC <sub>50</sub> μM) | HBV infected HepG2-NTCP-C4 (EC <sub>50</sub> μM) | Core I105T Mutation (EC <sub>50</sub> mM) | HBV infected HepG2-NTCP-C4 (HBsAg EC <sub>50</sub> μM) | (FC in EC <sub>50</sub> in 40% Human Serum) |
| AB-506   | 0.077                              | 0.032                                  | 0.101  | 1.26                                      | 1.430  | 6x  |
| AB-836   | 0.010                              | 0.002                                  | 0.012  | 0.118                                     | 0.196  | 2x  |



# Next Gen RNA Destabilizer Program

**Offers a novel mechanism of action** to reduce HBsAg and other viral proteins and viral RNA

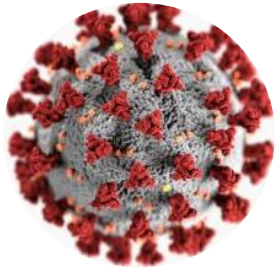
**Continuing active research** and development of a next generation small molecule

**We believe this approach offers potential** for an oral HBsAg reducing agent and all oral combination therapy



# Coronavirus Strategy

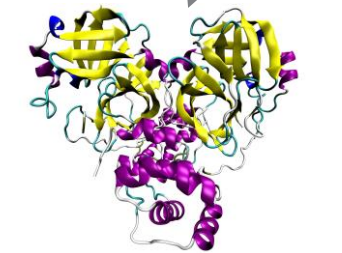
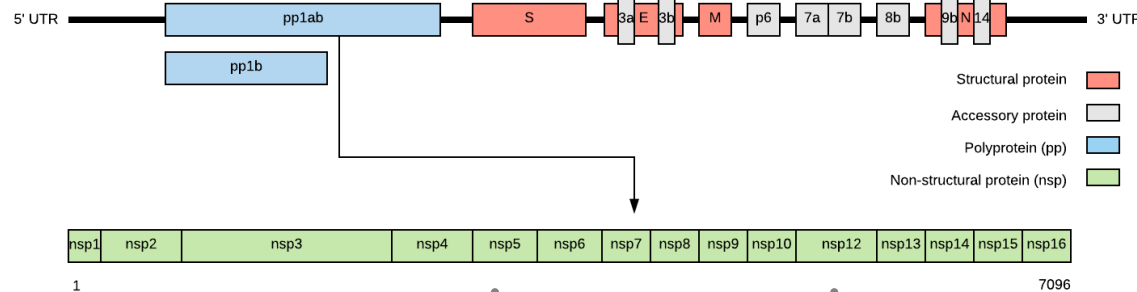
Leveraging our proven expertise and capabilities in antiviral drug discovery and development



**+RNA Virus**

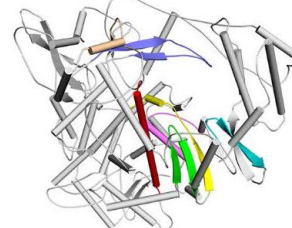
**31 kb Genome**

**nsp5 protease & nsp12 polymerase**  
essential enzymes  
for replication



COVID-19 Virus nsp5 / 3CLpro

**Viral Protease**



COVID-19 Virus nsp12

**Viral Polymerase**

**Long term commitment**

**Pan-coronavirus focused**

**Small Molecule Direct-Acting Antivirals**

Directed Effort

- nsp12 Viral Polymerase - nucleosid(t)es
- nsp5 Main Viral Protease - denovo design

**Screening Effort**

- Proprietary library screening through COVID R&D consortium

# Key Objectives for 2020

Cash balance of \$88.1M as March 31, 2020, cash runway into mid-2021

| Objective  | Anticipated Timing |
|--|--------------------|
| AB-729 preliminary phase 1a/1b single-dose data            | ✓ Late 1Q 2020     |
| AB-729 additional week 12 60 mg single-dose data           | ✓ May 2020         |
| AB-729 multi-dose 60 mg data (4 and 8 wk dosing intervals) | 2H 2020            |
| AB-729 week 12 single-dose 90 mg data                      | 2H 2020            |
| AB-836 complete IND enabling studies                       | 2H 2020            |