

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

**FORM 8-K**

**CURRENT REPORT**

**Pursuant to Section 13 or 15(d)  
of the Securities Exchange Act of 1934**

**Date of Report (Date of earliest event reported): August 3, 2023**

**Arbutus Biopharma Corporation**

(Exact name of registrant as specified in its charter)

**British Columbia, Canada**

(State or Other Jurisdiction of Incorporation)

**001-34949**

(Commission File Number)

**98-0597776**

(I.R.S. Employer Identification No.)

**701 Veterans Circle**

**Warminster, Pennsylvania 18974**

(Address of Principal Executive Offices) (Zip Code)

**(267) 469-0914**

(Registrant's telephone number, including area code)

(Former name or former address, if changed since last report)

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:

- Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)
- Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)
- Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))
- Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

Securities registered pursuant to Section 12(b) of the Act:

Title of each class	Trading Symbol(s)	Name of each exchange on which registered
Common Shares, without par value	ABUS	The Nasdaq Stock Market LLC

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

Emerging growth company

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act.

**Item 2.02. Results of Operations and Financial Condition.**

On August 3, 2023, Arbutus Biopharma Corporation (the “Company”) issued a press release announcing its financial results for the second quarter ended June 30, 2023 and certain other information. A copy of the press release is furnished as Exhibit 99.1 hereto and is incorporated by reference herein.

**Item 8.01. Other Events.**

On August 3, 2023, the Company posted an updated corporate presentation on its website at [www.arbutusbio.com](http://www.arbutusbio.com). A copy of the presentation is filed herewith as Exhibit 99.2 and is incorporated by reference herein.

**Item 9.01. Financial Statements and Exhibits.****(d) Exhibits.**

<u>Exhibit Number</u>	<u>Description</u>
<a href="#">99.1</a>	<a href="#">Press Release dated August 3, 2023</a>
<a href="#">99.2</a>	<a href="#">Corporate Presentation dated August 3 2023</a>
104	Cover page interactive data file (formatted as inline XBRL).

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**SIGNATURE**

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

**Arbutus Biopharma Corporation**

Date: August 3, 2023

By: /s/ David C. Hastings  
David C. Hastings  
Chief Financial Officer

## Arbutus Reports Second Quarter 2023 Financial Results and Corporate Update

**Regulatory approval received in New Zealand to advance AB-101, our oral PD-L1 inhibitor, into a Phase 1 clinical trial with dosing to begin this quarter**

**AB-729 (imdsiran), in combination with pegylated interferon alfa-2a, in a Phase 2a clinical trial, was generally well tolerated and appears to result in continued HBsAg declines in some patients**

**First patient dosed in additional arm of Phase 2a clinical trial combining imdsiran, VTP-300, NA therapy and nivolumab – advancing towards goal of further stimulating host HBV-associated immunity**

**Cash runway into the first quarter of 2025**

**Conference Call and Webcast Today at 8:45 AM ET**

WARMINSTER, Pa., Aug. 03, 2023 (GLOBE NEWSWIRE) -- Arbutus Biopharma Corporation (Nasdaq: ABUS) (“Arbutus” or the “Company”), a clinical-stage biopharmaceutical company leveraging its extensive virology expertise to develop novel therapeutics that target specific viral diseases, today reported second quarter 2023 financial results and provided a corporate update.

“In the second quarter of 2023, we achieved two important milestones in our Phase 2a clinical trials that support our efforts in developing AB-729 (imdsiran), our lead RNAi therapeutic, as a cornerstone therapy in a functional cure treatment regimen for HBV,” said William Collier, Arbutus President and Chief Executive Officer. “First, we reported data from our Phase 2a clinical trial showing that imdsiran in combination with interferon, is well tolerated and appears to result in continued HBsAg declines in some patients. Second, we made solid progress towards our goal of further stimulating host HBV-associated immunity, as we dosed the first patient in the additional treatment arm of the ongoing Phase 2a trial assessing the addition of low-dose nivolumab, a PD-1 monoclonal antibody, to VTP-300 and imdsiran.”

Mr. Collier continued, “Regarding our early-stage HBV assets, we are now prepared to move AB-101 forward into a Phase 1 clinical trial in New Zealand, which we expect to initiate this quarter, and AB-161, our oral RNA destabilizer, is in an on-going Phase 1 clinical trial. Additionally, we are on-track to complete IND-enabling studies with our coronavirus M<sup>Pro</sup> inhibitor candidate, AB-343, as well as initiate IND-enabling studies for a coronavirus nsp12 inhibitor candidate in the second half of this year.”

### Pipeline Updates and Key Milestones

#### Imdsiran (AB-729, RNAi Therapeutic)

- At the European Association for the Study of the Liver (EASL) Congress, we presented data from our on-going Phase 2a clinical trial (AB-729-201), evaluating the safety, tolerability and antiviral activity of the combination of imdsiran and pegylated interferon alfa-2a (IFN) in patients with chronic hepatitis B virus (cHBV). Preliminary data suggests that the addition of IFN to imdsiran was generally well tolerated and appears to result in continued HBsAg declines in some patients. The mean HBsAg decline from baseline during the imdsiran lead-in phase was 1.6 log<sub>10</sub> at week 24 of treatment which is comparable to what was previously seen in other imdsiran clinical trials. Four patients reached HBsAg below the lower limit of quantitation (LLOQ) at some point during IFN treatment. We plan to provide a further update on this clinical trial when we have additional meaningful patient data.
- We have completed enrollment in the first group of our Phase 2a clinical trial (AB-729-202) that is evaluating imdsiran, nucleos(t)ide analogue (NA) therapy and Vaccitech’s HBV antigen-specific immunotherapeutic, VTP-300. Preliminary data from patients in the clinical trial are expected in the second half of 2023.

We recently expanded the AB-729-202 clinical trial to enroll 20 patients who will receive imdsiran (60mg every 8 weeks) plus NA therapy for 24 weeks followed by VTP-300 plus up to two doses of low-dose nivolumab (Opdivo<sup>®</sup>). In June 2023, we announced that the first patient received the first dose of imdsiran in this additional arm. Preliminary data from this additional treatment arm are expected in 2024.

#### AB-161 (Oral RNA destabilizer)

- The Phase 1 clinical trial with AB-161 is on-going with single-ascending dose data expected in the second half of 2023. AB-161 is our next-generation oral HBV-specific RNA destabilizer, which is being developed as part of a potential all-oral treatment regimen to functionally cure HBV. Recently reported preclinical data showed that AB-161 provides robust anti-HBV activity including suppression of HBV RNA and HBsAg production *in vitro* and *in vivo*.

#### AB-101 (Oral PD-L1 Inhibitor)

- In April 2023, AB-101 was placed on clinical hold by the U.S. Food and Drug Administration (FDA) during the Investigational New Drug (IND) application review process prior to dosing subjects. In July 2023, the New Zealand Medicine and Medical Device Safety Authority (Medsafe) approved our CTA application for a Phase 1 clinical trial in New Zealand for AB-101, and we believe the protocol approved by Medsafe adequately addresses the clinical trial design and safety monitoring issues raised by the FDA. We are planning to initiate the Phase 1 clinical trial this quarter. We are developing AB-101 to reawaken and boost the immune system of patients with cHBV. Preclinical data generated thus far indicates that AB-101 is highly potent and mediates activation and reinvigoration of HBV-specific T-cells from cHBV patients.

### COVID-19 and Pan-Coronavirus Programs

- We are continuing to conduct IND-enabling studies with AB-343 and are on track to complete those studies in the second half of 2023.

- We are continuing to direct our research efforts to identifying an nsp12 viral polymerase inhibitor clinical candidate. Such a candidate could potentially be combined with AB-343 to achieve better patient treatment outcomes and for use in prophylactic settings. We expect to nominate an nsp12 inhibitor clinical candidate and initiate IND-enabling studies in the second half of 2023.

## Corporate Updates

- In July 2023, we announced that Melissa V. Rewolinski, PhD was appointed to the Board of Directors. Melissa brings to the Board more than 20 years of strategic, operational and drug development experience within the pharmaceutical industry.
- In July 2023, we announced the promotion of Karen Sims, MD, PhD to Chief Medical Officer. Karen is a board-certified infectious disease physician with more than 12 years of industry experience in conducting and overseeing early stage through global Phase 2 clinical trials. She joined Arbutus in April 2017 and has held positions of increasing seniority, including most recently as Vice President, Clinical Development, before being promoted to Chief Medical Officer.
- In July 2023, we also announced the appointment of Christopher Naftzger as General Counsel and Chief Compliance Officer. Chris succeeds Dr. Elizabeth Howard who will continue in an advisory role with respect to the on-going patent infringement litigations. Chris brings more than 25 years of legal experience, including over a decade of experience serving in senior in-house counsel positions with life science companies.

## Financial Results

### Cash, Cash Equivalents and Investments

As of June 30, 2023, we had cash, cash equivalents and investments in marketable securities of \$163.5 million compared to \$184.3 million as of December 31, 2022. During the six months ended June 30, 2023, we used \$46.9 million in operating activities, which was partially offset by \$24.6 million of net proceeds from the issuance of common shares under our “at-the-market” offering program. We expect our 2023 net cash burn to range from between \$90 to \$95 million, excluding any proceeds received from our “at the market program”. We believe our cash runway will be sufficient to fund our operations into the first quarter of 2025.

### Revenue

Total revenue was \$4.7 million for the three months ended June 30, 2023 compared to \$14.2 million for the same period in 2022. The decrease of \$9.5 million for the 2023 period was due primarily to lower revenue recognition from our license agreement with Qilu compared to the 2022 period based on lower employee labor hours expended by us in the 2023 period compared to the 2022 period to perform our manufacturing obligations under the license agreement.

### Operating Expenses

Research and development expenses were \$17.7 million for the three months ended June 30, 2023 compared to \$22.9 million for the same period in 2022. The decrease of \$5.2 million was due primarily to a decrease in expenses for drug supply manufacturing for our imdusiran, AB-101 and AB-161 clinical trials, as well as a decrease in expenses related to our AB-836 Phase 1a/1b clinical trial, which was discontinued in the fourth quarter of 2022. These were partially offset by an increase in expenses for our coronavirus program, including drug supply manufacturing. General and administrative expenses were \$6.0 million for the three months ended June 30, 2023, compared to \$5.2 million for the same period in 2022. This increase was due primarily to increases in non-cash stock-based compensation expense and professional fees.

### Net Loss

For the three months ended June 30, 2023, our net loss was \$17.1 million, or a loss of \$0.10 per basic and diluted common share, as compared to a net loss of \$14.2 million, or a loss of \$0.10 per basic and diluted common share, for the three months ended June 30, 2022.

### Outstanding Shares

As of June 30, 2023, we had approximately 166.9 million common shares issued and outstanding, as well as approximately 20.2 million stock options and unvested restricted stock units outstanding. Roivant Sciences Ltd. owned approximately 23% of our outstanding common shares as of June 30, 2023.

## UNAUDITED CONDENSED CONSOLIDATED STATEMENTS OF LOSS (in thousands, except share and per share data)

	Three Months Ended June 30,		Six Months Ended June 30,	
	2023	2022	2023	2022
Revenue				
Collaborations and licenses	\$ 3,885	\$ 12,556	\$ 9,394	\$ 23,774
Non-cash royalty revenue	766	1,685	1,944	3,048
<b>Total revenue</b>	<b>4,651</b>	<b>14,241</b>	<b>11,338</b>	<b>26,822</b>
Operating expenses				
Research and development	17,692	22,942	35,967	41,404
General and administrative	5,980	5,200	11,532	10,092
Change in fair value of contingent consideration	(636)	208	(363)	409
<b>Total operating expenses</b>	<b>23,036</b>	<b>28,350</b>	<b>47,136</b>	<b>51,905</b>
Loss from operations	(18,385)	(14,109)	(35,798)	(25,083)
Other income (loss)				
Interest income	1,461	396	2,729	555

Interest expense	(171)	(482)	(369)	(988)
Foreign exchange gain	1	3	5	3
Total other income (loss)	1,291	(83)	2,365	(430)
Loss before income taxes	(17,094)	(14,192)	(33,433)	(25,513)
Income tax expense	—	—	—	(4,444)
<b>Net loss</b>	<b>\$ (17,094)</b>	<b>\$ (14,192)</b>	<b>\$ (33,433)</b>	<b>\$ (29,957)</b>
Net loss per common share				
Basic and diluted	\$ (0.10)	\$ (0.10)	\$ (0.20)	\$ (0.20)
Weighted average number of common shares				
Basic and diluted	166,063,284	148,750,048	163,855,661	148,589,711

**UNAUDITED CONDENSED CONSOLIDATED BALANCE SHEETS**  
(in thousands)

	<u>June 30, 2023</u>	<u>December 31, 2022</u>
Cash, cash equivalents and marketable securities, current	\$ 152,484	\$ 146,913
Accounts receivable and other current assets	6,316	4,226
Total current assets	158,800	151,139
Property and equipment, net of accumulated depreciation	5,370	5,070
Investments in marketable securities, non-current	11,057	37,363
Right of use asset	1,585	1,744
Other non-current assets	11	103
<b>Total assets</b>	<b>\$ 176,823</b>	<b>\$ 195,419</b>
Accounts payable and accrued liabilities	\$ 8,805	\$ 16,029
Deferred license revenue, current	15,327	16,456
Lease liability, current	397	372
Total current liabilities	24,529	32,857
Liability related to sale of future royalties	8,787	10,365
Deferred license revenue, non-current	—	5,999
Contingent consideration	7,168	7,531
Lease liability, non-current	1,646	1,815
Total stockholders' equity	134,693	136,852
<b>Total liabilities and stockholders' equity</b>	<b>\$ 176,823</b>	<b>\$ 195,419</b>

**UNAUDITED CONDENSED CONSOLIDATED STATEMENTS OF CASH FLOWS**  
(in thousands)

	<u>Six Months Ended June 30,</u>	
	<u>2023</u>	<u>2022</u>
Net loss	\$ (33,433)	\$ (29,957)
Non-cash items	2,911	3,154
Change in deferred license revenue	(7,128)	27,815
Other changes in working capital	(9,210)	(686)
<b>Net cash (used in) provided by operating activities</b>	<b>(46,860)</b>	<b>326</b>
Net cash provided by (used in) investing activities	18,119	(73,886)
Issuance of common shares pursuant to Share Purchase Agreement	—	10,973
Issuance of common shares pursuant to the Open Market Sale Agreement	24,604	268
Cash provided by other financing activities	555	357
Net cash provided by financing activities	25,159	11,598
Effect of foreign exchange rate changes on cash and cash equivalents	3	—
Decrease in cash and cash equivalents	(3,579)	(61,962)
Cash and cash equivalents, beginning of period	30,776	109,282
Cash and cash equivalents, end of period	27,197	47,320
Investments in marketable securities	136,344	153,329
<b>Cash, cash equivalents and marketable securities, end of period</b>	<b>\$ 163,541</b>	<b>\$ 200,649</b>

**Conference Call and Webcast Today**

Arbutus will hold a conference call and webcast today, Thursday, August 3, 2023, at 8:45 AM Eastern Time to provide a corporate update. To dial-in for the conference call by phone, please register using the following link: [Registration Link](#). A live webcast of the conference call can be accessed through the Investors section of Arbutus' website at [www.arbutusbio.com](http://www.arbutusbio.com).

An archived webcast will be available on the Arbutus website after the event.

### **About imdusiran (AB-729)**

Imdusiran is an RNA interference (RNAi) therapeutic specifically designed to reduce all HBV viral proteins and antigens including hepatitis B surface antigen which is thought to be a key prerequisite to enable reawakening of a patient's immune system to respond to the virus. Imdusiran targets hepatocytes using Arbutus' novel covalently conjugated N-Acetylgalactosamine (GalNAc) delivery technology enabling subcutaneous delivery. Clinical data generated thus far has shown single- and multi-doses of imdusiran to be generally safe and well-tolerated, while also providing meaningful reductions in hepatitis B surface antigen and hepatitis B DNA. Imdusiran is currently in multiple Phase 2a clinical trials.

### **About AB-101**

AB-101 is our lead oral PD-L1 inhibitor candidate that we believe will allow for controlled checkpoint blockade and enable oral dosing, while minimizing the systemic safety issues typically seen with checkpoint antibody therapies. Immune checkpoints such as PD-1/PD-L1 play an important role in the induction and maintenance of immune tolerance and in T-cell activation. Preclinical data generated thus far indicates that AB-101 mediates activation and reinvigoration of HBV-specific T-cells from cHBV patients. We believe AB-101, when used in combination with other approved and investigational agents, could potentially lead to a functional cure in HBV chronically infected patients. We are also exploring oncology applications for our internal PD-L1 portfolio.

### **About AB-161**

AB-161 is our next generation oral small molecule RNA destabilizer, specifically designed to target the liver. Mechanistically, RNA destabilizers target the host proteins PAPD5/7, which are involved in regulating the stability of HBV RNA transcripts. In doing so, RNA destabilizers lead to the selective degradation of HBV RNAs, thus reducing HBsAg levels and inhibiting viral replication. To provide a proprietary all-oral treatment regimen for patients with cHBV, we believe inclusion of a small molecule RNA destabilizer is key.

### **About AB-343**

AB-343 is our lead coronavirus drug candidate that inhibits the SARS-CoV-2 main protease (M<sup>Pro</sup>), a validated target for the treatment of COVID-19 and potential future coronavirus outbreaks. In our pre-clinical research conducted to date, AB-343 has shown pan-coronavirus antiviral activity, no reduction in potency against known SARS-CoV-2 variants, robust activity against SARS-CoV-2 M<sup>Pro</sup> resistant strains, and a favorable drug-drug interaction profile with no need for ritonavir boosting. We see an opportunity to pursue a potential combination therapeutic strategy focusing on M<sup>Pro</sup> and nsp12 viral polymerase targets to reduce hospitalizations, achieve better patient treatment outcomes and provide pre-exposure prophylactic therapy.

### **About HBV**

Hepatitis B is a potentially life-threatening liver infection caused by the hepatitis B virus (HBV). HBV can cause chronic infection which leads to a higher risk of death from cirrhosis and liver cancer. Chronic HBV infection represents a significant unmet medical need. The World Health Organization estimates that over 290 million people worldwide suffer from chronic HBV infection, while other estimates indicate that approximately 2.4 million people in the United States suffer from chronic HBV infection. Approximately 820,000 people die every year from complications related to chronic HBV infection despite the availability of effective vaccines and current treatment options.

### **About Coronaviruses**

Coronaviruses are a large family of viruses that range from the common cold to more severe diseases such as severe acute respiratory syndrome (SARS), Middle East respiratory syndrome (MERS), and COVID-19. COVID-19 has caused approximately 7.2 million deaths globally according to an analysis by the Institute for Health Metrics and Evaluation (IHME). As we strive to identify and develop new antiviral small molecules to treat COVID-19 and future coronavirus outbreaks, we have focused our research efforts on two essential targets critical for replication across all coronaviruses – nsp5 protease and nsp12 polymerase.

### **About Arbutus**

Arbutus Biopharma Corporation (Nasdaq: ABUS) is a clinical-stage biopharmaceutical company leveraging its extensive virology expertise to develop novel therapeutics that target specific viral diseases. Our current focus areas include Hepatitis B virus (HBV), SARS-CoV-2, and other coronaviruses. To address HBV, we are developing a RNAi therapeutic, an oral PD-L1 inhibitor, and an oral RNA destabilizer to potentially identify a combination regimen with the aim of providing a functional cure for patients with chronic HBV by suppressing viral replication, reducing surface antigen and reawakening the immune system. We believe our lead compound, imdusiran (AB-729), is the only RNAi therapeutic with evidence of immune re-awakening. Imdusiran is currently being evaluated in multiple phase 2 clinical trials. We also have an ongoing drug discovery and development program directed to identifying novel, orally active agents for treating coronaviruses, (including SARS-CoV-2), for which we have nominated a compound and have begun IND-enabling pre-clinical studies. In addition, we are also exploring oncology applications for our internal PD-L1 portfolio. For more information, visit [www.arbutusbio.com](http://www.arbutusbio.com).

### **Forward-Looking Statements and Information**

This press release contains forward-looking statements within the meaning of the Section 27A of the Securities Act of 1933 and Section 21E of the Securities Exchange Act of 1934, and forward-looking information within the meaning of Canadian securities laws (collectively, forward-looking statements). Forward-looking statements in this press release include statements about our future development plans for our product candidates; the expected cost, timing and results of our clinical development plans and clinical trials with respect to our product candidates; our expectations with respect to the release of data from our clinical trials and the expected timing thereof; our expectations and goals for our collaborations with third parties and any potential benefits related thereto; statements regarding our plans for AB-101 in light of the FDA's clinical hold; the potential for our product candidates to achieve success in clinical trials; and our expected financial condition, including our anticipated net cash burn, the anticipated duration of cash runways and timing regarding needs for additional capital.

With respect to the forward-looking statements contained in this press release, Arbutus has made numerous assumptions regarding, among other things: 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 patent litigation matters.

Additionally, there are known and unknown risk factors which could cause Arbutus' actual results, performance or achievements to be materially different from any future results, performance or achievements expressed or implied by the forward-looking statements contained herein. Known risk factors include, among others: anticipated pre-clinical studies 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 product candidate; Arbutus may elect to change its 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; uncertainties associated with litigation generally and patent litigation specifically; it may take considerable time and expense to resolve the clinical hold that has been placed on AB-101 by the FDA, and no assurance can be given that the FDA will remove the clinical hold; Arbutus and its collaborators may never realize the expected benefits of the collaborations; and market shifts may require a change in strategic focus.

A more complete discussion of the risks and uncertainties facing Arbutus appears in Arbutus' Annual Report on Form 10-K, Arbutus' Quarterly Reports on Form 10-Q and Arbutus' continuous and periodic disclosure filings, which are available at [www.sedar.com](http://www.sedar.com) and at [www.sec.gov](http://www.sec.gov). All forward-looking statements herein are qualified in their entirety by this cautionary statement, and Arbutus disclaims any obligation to revise or update any such forward-looking statements or to publicly announce the result of any revisions to any of the forward-looking statements contained herein to reflect future results, events or developments, except as required by law.

### **Contact Information**

#### **Investors and Media**

Lisa M. Caperelli  
Vice President, Investor Relations  
Phone: 215-206-1822  
Email: [lcaperelli@arbutusbio.com](mailto:lcaperelli@arbutusbio.com)





# Corporate Presentation

NASDAQ: ABUS

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

August 3, 2023



# 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 market opportunity for HBV; Arbutus' ability to meet a significant unmet medical need; the sufficiency of Arbutus' cash and cash equivalents for the anticipated durations; the expected cost, timing and results of Arbutus' clinical development plans and clinical trials, including its clinical collaborations with third parties; the potential for Arbutus' product candidates to achieve their desired or anticipated outcomes; Arbutus' expectations regarding the timing and clinical development of Arbutus' product candidates, including its articulated clinical objectives; the timeline to a combination cure for HBV; Arbutus' coronavirus strategy; Arbutus' expectations regarding its technology licensed to third parties; the expected timing and payments associated with strategic and/or licensing agreements; the patent infringement lawsuit against Moderna; and other statements relating to Arbutus' 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 pre-clinical 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 and patent litigation matters. 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; uncertainties associated with litigation generally and patent litigation specifically; market shifts may require a change in strategic focus; the parties may never realize the expected benefits of the collaborations; and the ongoing COVID-19 pandemic could significantly disrupt Arbutus' 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). All forward-looking statements herein are qualified in their entirety by this cautionary statement, and Arbutus disclaims any obligation to revise or update any such forward-looking statements or to publicly announce the result of any revisions to any of the forward-looking statements contained herein to reflect future results, events or developments, except as required by law.

# Our Strategy

Leverage the proven track record of success established with our team's expertise in understanding and treating viral infections by discovering and developing a broad, differentiated pipeline of therapies targeting chronic HBV, COVID-19, and future coronavirus outbreaks.



Develop a **combination therapy that includes antivirals and immunologics** to provide a finite duration treatment for people with cHBV that results in >20% functional cure rate.



Develop **novel oral pan-coronavirus antivirals targeting essential viral proteins** with the goal of reducing hospitalizations and providing pre-exposure prophylactic therapy.

# Investment Highlights



Indications with significant unmet medical need & large market opportunities

Focused on developing functional cure for HBV and oral pan-coronavirus therapeutics



Team with virology expertise and proven track record

Discovered, developed & commercialized multiple drugs



Broad portfolio of internally discovered assets with distinct MOAs

RNAi therapeutic  
PD-L1 inhibitor  
RNA destabilizer  
M<sup>pro</sup> inhibitor  
Nsp12 polymerase inhibitor



Lead HBV compound – imdusiran (AB-729) RNAi therapeutic in multiple Phase 2a combination clinical trials

Data shows imdusiran is generally safe and well-tolerated and has shown meaningful suppression of HBsAg while on- or off-treatment



Strong financial position

Cash runway into Q1 2025



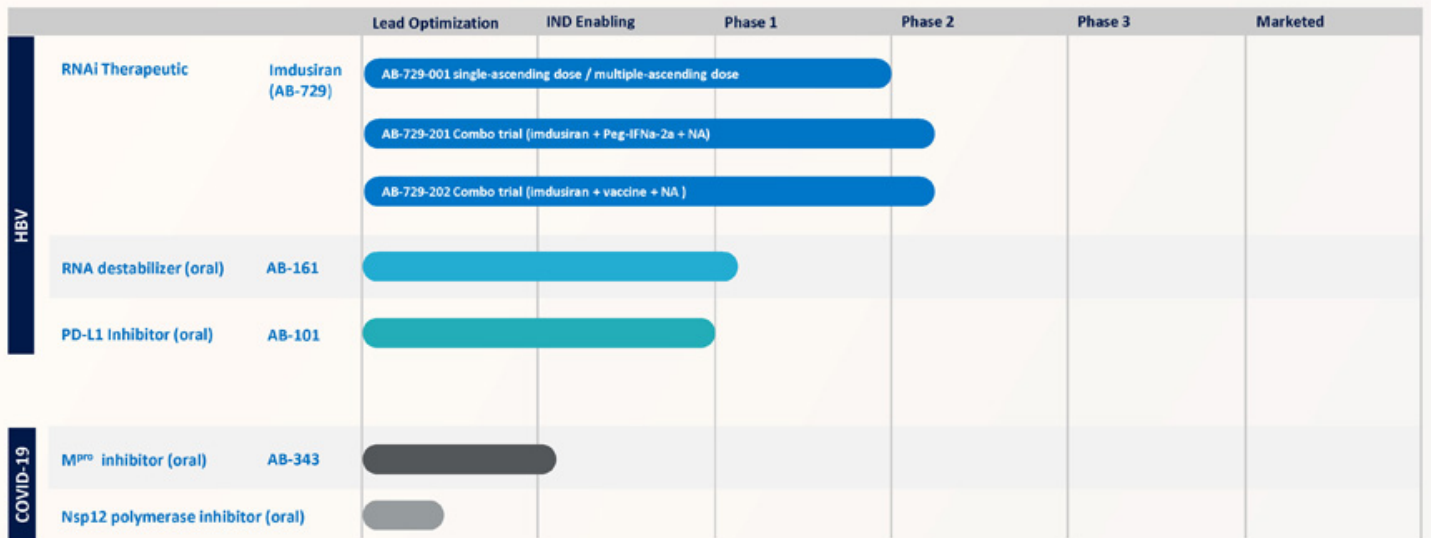
Patented LNP technology

Receiving licensing royalties arising from Alnylam's Onpattro® and seeking damages for Moderna & Pfizer/BioNTech COVID-19 vaccine sales



MOA: Mechanism of Action | PD-L1: Programmed death-ligand 1 | M<sup>pro</sup>: Main protease  
NSP12: Non-structural protein | HBsAg: Hepatitis B surface antigen

# Broad Pipeline



NA: Nucleoside Analogue

# HBV Overview



## Cause & Symptoms

- Life-threatening liver infection caused by hepatitis B virus (HBV)
- Transmitted through body fluids and from mother to child
- Long-term chronic infection (cHBV) leads to higher risk of cirrhosis and/or liver cancer



## Diagnosis

- HBsAg detection
- Additional biomarkers necessary to determine stage of disease



## Treatments

- NA therapy – lifelong daily therapy, aimed at reducing HBV DNA and risk of cirrhosis and/or HCC
- Peg-IFN $\alpha$  – administered weekly; poorly tolerated
- <5% of patients achieve functional cure



## Rationale

- Need for finite and more efficacious HBV treatments that further improve long-term outcomes and increase functional cure rate
- Combination therapy with different MOAs will be required to reduce HBsAg, suppress HBV DNA, and boost immune system

Sources for all data on slide:

1 Hepatitis B Fact Sheet, WHO <https://www.who.int/news-room/fact-sheets/detail/hepatitis-b>; Hep B Foundation link <https://www.hepb.org/what-is-hepatitis-b/what-is-hepb/facts-and-figures/>; Kowdley et al. Hepatology (2012) Prevalence of Chronic Hepatitis B Among Foreign-Born Persons Living in the US by Country of Origin

2 Pegasis, PEG-Intron, Baraclude and Viread Package Inserts

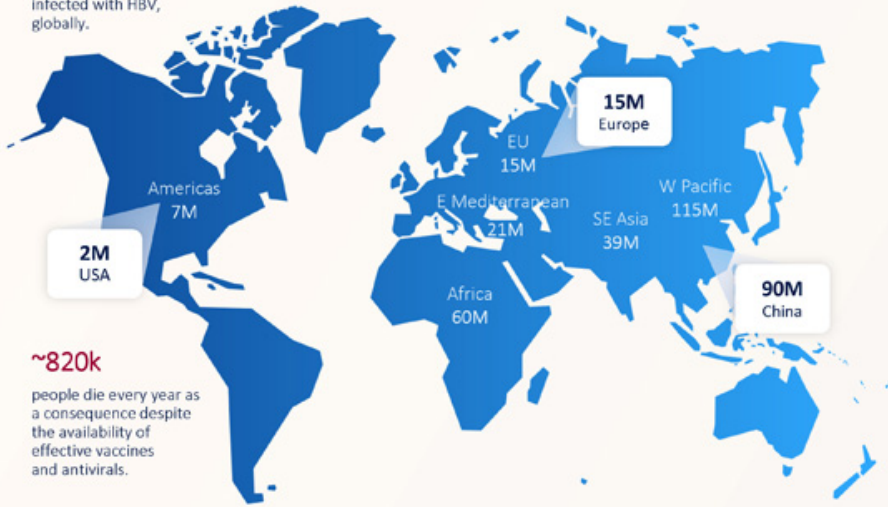


HBsAg: HBV Surface Antigen | HCC: Hepatocellular carcinoma

# HBV Presents a Significant Unmet Medical Need

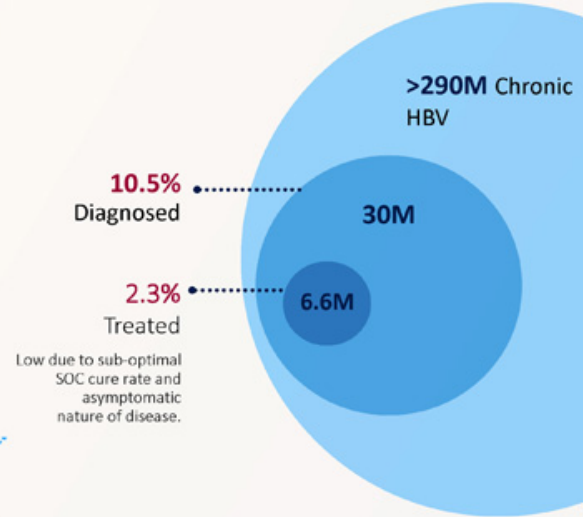
>290M

people are chronically infected with HBV, globally.



~820k

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



SOC: Standard of Care

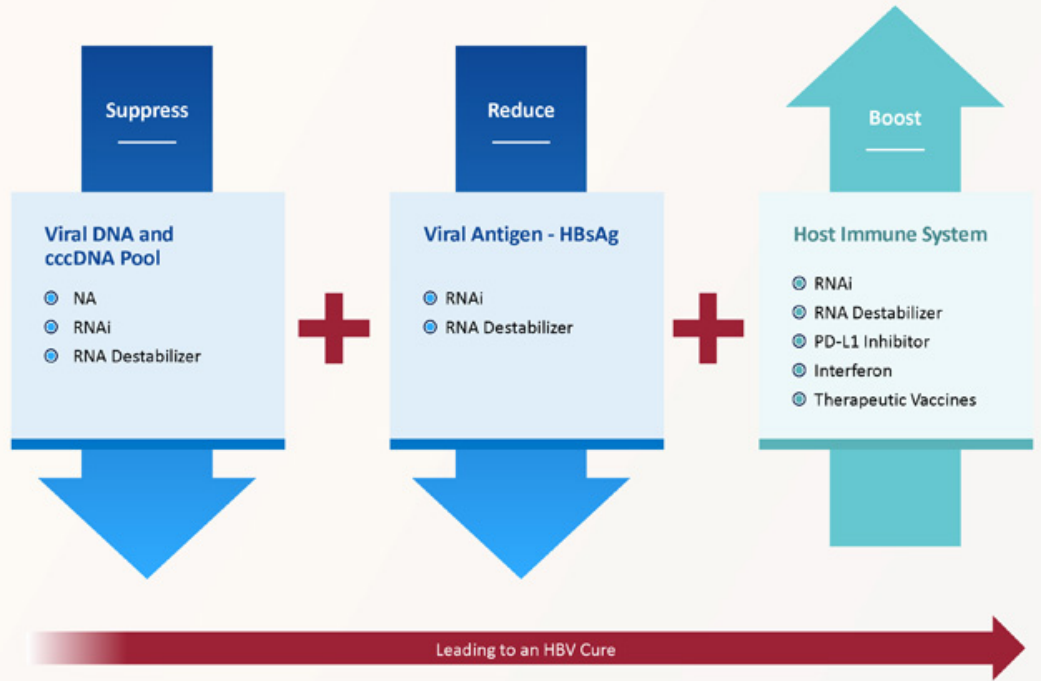
Sources: <https://www.who.int/news-room/fact-sheets/detail/hepatitis-b>  
<https://www.hepb.org/what-is-hepatitis-b/what-is-hepb/facts-and-figures/>

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# 3-Pronged Approach to Therapeutic Success

- Suppress HBV DNA
- Reduce viral antigens
- + Boost host immune response

Therapeutic success will require a combination of agents with complementary MOAs.





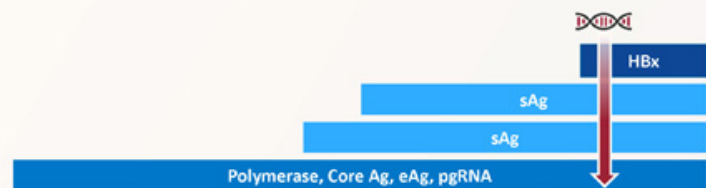
## Imdusiran

# 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 other agents
- Actively targets the liver
- Active against cccDNA derived and integrated HBsAg transcripts
- Clean profile in long term preclinical safety studies



# AB-729-001 Phase 1a/1b Clinical Trial

## Part 1 & 2:

### Single-ascending dose

#### Imdusiran monotherapy conclusions:

- Robust HBsAg declines across all cohorts
- HBV DNA declines in HBV DNA+ patients

## Part 3: Multiple Ascending Dose in cHBV Patients

E: 60mg Q4W  
HBV DNA-

F: 60mg Q8W  
HBV DNA-

G: 90mg Q8W + TDF  
HBV DNA+

I: 90mg Q8W  
HBV DNA-

J: 90mg Q12W  
HBV DNA-

K: 90mg Q8W HBV DNA-,  
HBeAg+ only

HBeAg: HBV E antigen | TDF: tenofovir disoproxil fumarate  
Data presented at EASL 2022

# Robust HBsAg Declines Irrespective of Dose, Dosing Schedule, HBeAg or HBV DNA Status

Mean (SE) Baseline and  $\Delta \log_{10}$  HBsAg by Visit

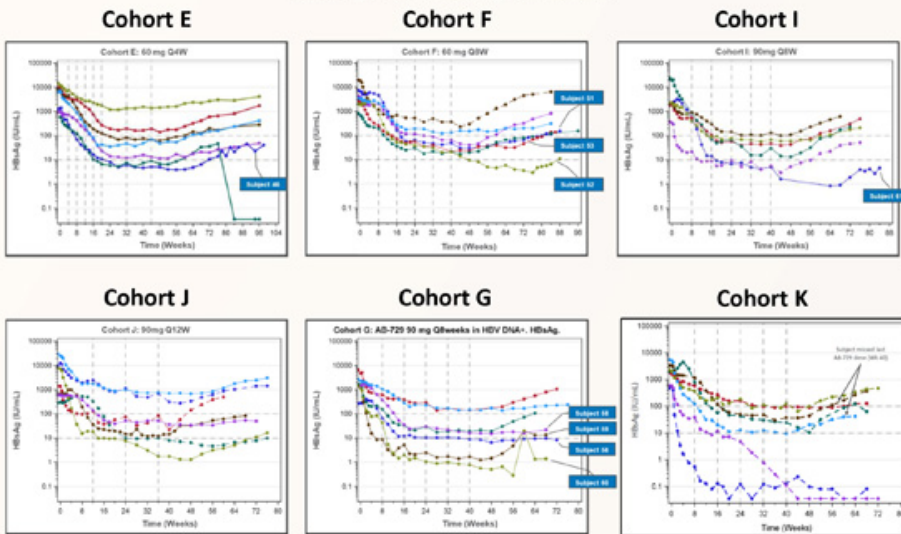
Visit	HBV DNA-					HBV DNA+
	Cohort E (N=7)	Cohort F (N=7)	Cohort I (N=6)	Cohort J (N=7)	Cohort K (N=7)	Cohort G (N=7)
Baseline	3.51 (0.20)	3.53 (0.17)	3.36 (0.23)	3.37 (0.28)	3.23 (0.14)	3.14 (0.14)
Treatment Week 12	-1.10 (0.15)	-1.02 (0.11)	-1.30 (0.19)	-1.06 (0.31)	-1.63 (0.39)	-1.56 (0.32)
Treatment Week 24	-1.84 (0.16)	-1.57 (0.09)	-1.79 (0.22)	-1.56 (0.25)	-1.99 (0.35)	-1.82 (0.29)
Treatment Week 48	-1.89 (0.18)	-1.90 (0.14)	-1.91 (0.32)	-1.80 (0.41)	-2.57 (0.61)	-2.05 (0.31)
Follow Up Week 12	-1.74 (0.20)	-1.59 (0.23)	-1.42 (0.26)	-1.52 (0.40)	-2.38 (0.75)	-1.50 (0.13)
Follow Up Week 24	-1.43 (0.18)	-1.26 (0.21)	-1.37 (0.39)	-1.49 (0.35)	-1.82 (0.63)	-1.53 (0.29)
Follow Up Week 48	-1.55 (0.56)	-1.01 (0.24)	-0.88 (0.33)	-1.04 (0.20)	-1.86 (0.70)	-1.10 (0.27)

Data shown as mean (SE)  $\log_{10}$  IU/mL; minimum of 5 subjects/timepoint. Last Indusiran (A8-729) dose Cohort E: Week 44, Cohorts F, I, G, K: Week 40, Cohort J: Week 36; HBsAg Assay LLOQ = 0.07 IU/mL; \*N=6; \*N=5

- All Cohorts achieved at least a  $-1.8 \log_{10}$  decline in mean HBsAg at the end of the treatment period (Week 48)
- Mean HBsAg levels remained below baseline values at Follow Up Week 48
- There were no significant differences in mean HBsAg declines between the 60 mg and 90 mg doses or between different dosing intervals

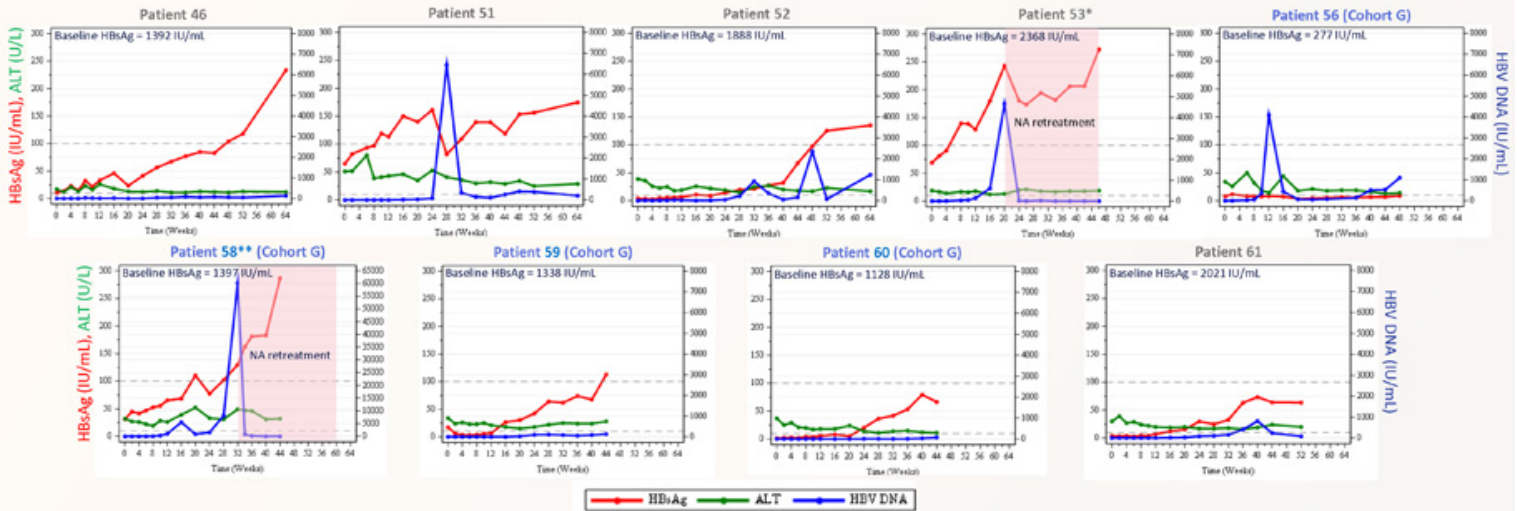
# AB-729-001: Robust & Sustained HBsAg Declines **While On- or Off-Treatment with Imdusiran**

Change in HBsAg vs time



- 33 of 41 patients had HBsAg < 100 IU/mL at some point during the trial
- 1 patient in Cohort E (baseline HBsAg = 583.5 IU/mL) who qualified but declined to participate in NA discontinuation seroconverted at Week 84 (HBsAg < LLOQ and HBsAb = 189 IU/mL at last visit); liver enzymes remained within normal limits.
- 2 patients in Cohort K reached HBsAg < LLOQ on multiple visits with detectable HBsAb levels

# AB-729-001: Imdusiran Shows Low Levels of HBV Biomarkers Persisting in cHBV Patients **While Off-Treatment**



- 7 of 9 (78%) subjects remain off NA therapy for 44-64 weeks and all completed imdusiran treatment over 1½ years ago
- Most subjects have maintained low HBV DNA levels off treatment
- HBsAg remains between  $-0.8$  and  $-1.6 \log_{10}$  IU/mL below baseline values
- NA discontinuation post-imdusiran treatment appears well tolerated with no ALT flares

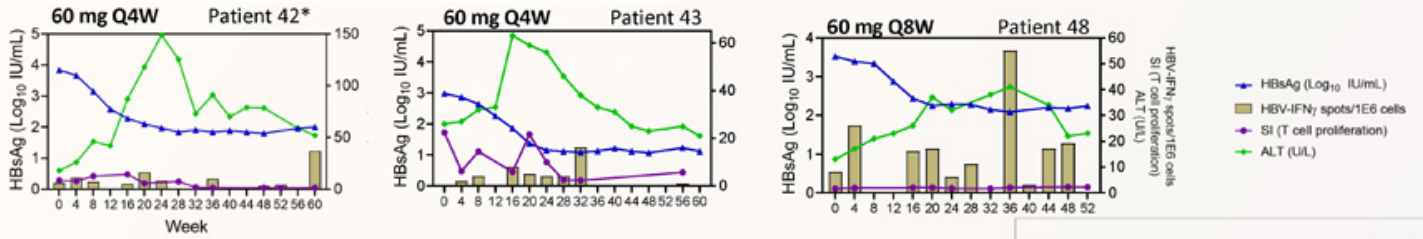


Data presented at GHS 2023

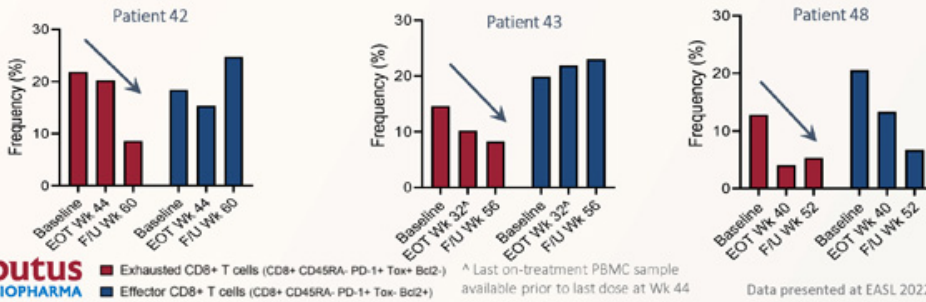
\* Patient 53 restarted NA therapy at Investigator's request after the NA d/c FU W20 visit (pink shaded area).  
 \*\* Patient 58 restarted therapy after the NA d/c FU W36 visit (pink shaded area).

# AB-729-001: Treatment with Imdusiran Reactivates HBV Specific Immunity in Some Patients

## Imdusiran Increased HBV-Specific T-Cell Activation



## Imdusiran Decreased Exhausted T-Cells



- Upregulation of HBV-specific T-cell activation markers observed in all 7 patients assessed to date
- Two profiles of HBV-specific T cell IFN- $\gamma$  responses observed
  - Elevation between Wk 16-28 which coincides with nadir of HBsAg reduction
  - \*Elevation after imdusiran dosing completed, between Wk 48-60



Exhausted CD8+ T cells (CD8+ CD45RA- PD-1+ Tbx+ Bcl2-)  
Effector CD8+ T cells (CD8+ CD45RA- PD-1+ Tbx+ Bcl2+)

<sup>^</sup> Last on-treatment PBMC sample available prior to last dose at Wk 44

Data presented at EASL 2022

## AB-729-001 Safety Summary

- Indusiran is generally safe and well-tolerated after repeat dosing for up to 48 weeks
- No treatment-related SAEs or discontinuations due to AEs
- No treatment-related Grade 3 or 4 AEs
- No treatment-related Grade 3 or 4 laboratory abnormalities
  - Grade 1 and Grade 2 ALT elevations have improved or stabilized with continued treatment
- Injection site AEs were all Grade 1 (erythema, pain, bruising)
- No clinically meaningful changes in ECGs or vital signs
- After NA treatment discontinuation, no ALT flares have been observed

# AB-729-001 Clinical Trial **Key Takeaways**

**Imdusiran provided robust and comparable HBsAg declines regardless of dose, dosing interval, HBeAg or DNA status**

**Discontinuation of both imdusiran and NA-therapy results in sustained reduction in HBsAg and HBV DNA in 7 of 9 patients**

**Imdusiran results in HBV-specific T-cell immune restoration and decrease of exhausted T-cells in some patients**

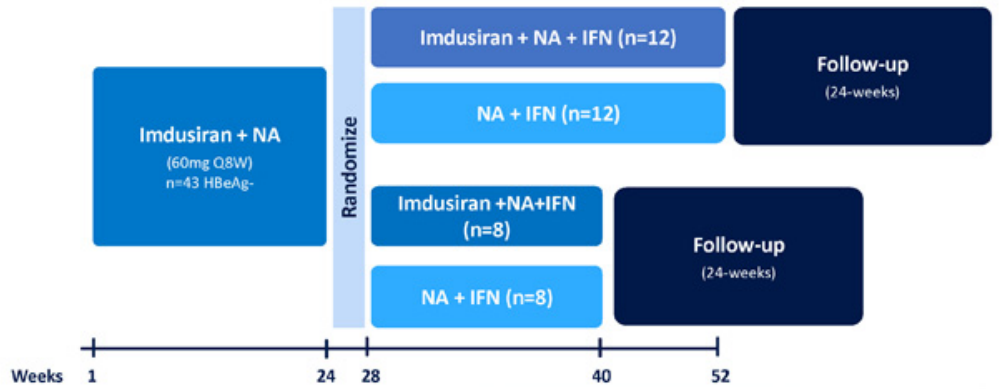
**Imdusiran was generally safe and well-tolerated after completing dosing in 41 patients**



## AB-729-201:

# Phase 2a POC Clinical Trial

Imdusiran in combination with ongoing NA therapy and short courses of Peg-IFN $\alpha$ -2a in CHBV patients



### Multi-center, open-label Phase 2a

**Primary objective:** evaluate safety and tolerability of imdusiran in combination with Peg-IFN $\alpha$ -2a in patients with NA-suppressed CHBV

**Preliminary results\*:** treatment was generally well tolerated with continued HBsAg declines in some patients during the IFN treatment period

- Mean HBsAg decline during lead-in phase was 1.6 log<sub>10</sub> at week 24 of treatment
- 93% of patients (38 of 41 randomized) had HBsAg levels <100 IU/mL during treatment period
- 4 patients reached HBsAg levels <LLOQ during IFN treatment

After 24-weeks of follow-up, patients are assessed to stop NA therapy. Those patients that stop NA therapy will be followed for an additional 48 weeks.

POC: Proof of Concept  
\*Data presented at EASL 2023

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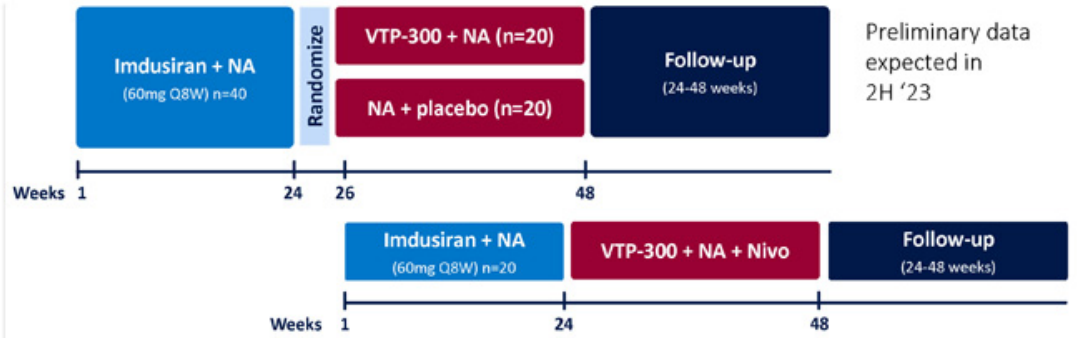
## AB-729-202:

# Phase 2a POC Clinical Trial



### POC Phase 2a clinical

**trial** evaluating imdusiran in combination with Vaccitech's immunotherapeutic, VTP-300, with or without low dose nivolumab, and a NA



**Primary objective:** evaluate safety and reactogenicity of imdusiran followed by VTP-300 or placebo

At week 48 all participants who are eligible to discontinue NA therapy will be followed for 48-weeks

Expanded the clinical trial to include an additional arm with nivolumab (Opdivo®), and dosed first patient in this arm in the first half of 2023

*Full rights retained by the Companies of their respective product candidates and all costs split equally*

# Imdusiran Strategic Collaboration



## Exclusive Licensing\* and Strategic Partnership

Develop, manufacture and commercialize imdusiran in mainland China, Hong Kong, Macau and Taiwan

\*ABUS retains the non-exclusive right to develop and manufacture in the Qilu territory for exploiting AB-729 in the rest of the world



## Deal economics for Arbutus:

<b>\$40M</b>	Upfront payment (received in 2022)
<b>\$15M</b>	Equity investment (received in 2022)
<b>Up to \$245M</b>	Commercialization and milestone payments
<b>Double-digit up to low twenties %</b>	Tiered royalties on annual sales

### Qilu Pharmaceutical:

One of the leading pharmaceutical companies in China, provides development, manufacturing, and commercialization expertise to this partnership



# AB-161: Next Generation Oral RNA Destabilizer

## Safety

**Next generation small molecule overcomes peripheral neuropathy** non-clinical safety findings with first generation molecule

## Novelty

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

## Convenience

Potential for an **oral HBsAg reducing agent** and all oral combination therapy

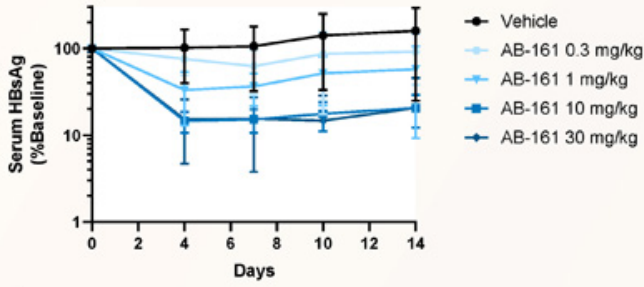
AB-161 is currently in a Phase 1 clinical trial

# AB-161 Reduces HBsAg in AAV-HBV Mouse Model

## Compound concentration in liver drives efficacy

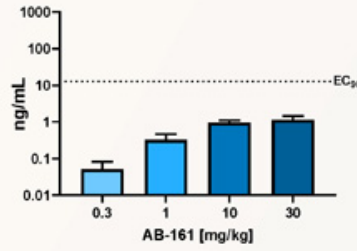
- AB-161 effective as a once-daily dose in AAV-HBV mouse model (0.3, 1, 10, 30 mg/kg QD)
  - Dose-dependent reduction of HBsAg, also observed with BID dosing (0.3 and 1 mg/kg BID)
- HBsAg reduction achieved when fraction unbound  $C_{24h} > EC_{90}$  in liver

AAV-HBV mouse  
AB-161 QD for 14 days

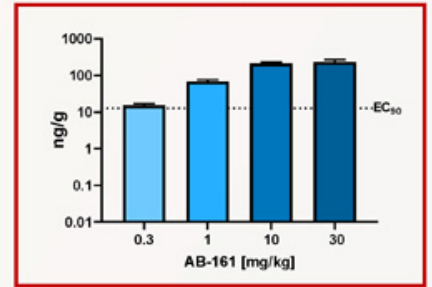


Fraction Unbound Concentrations ( $C_{24h}$ )

$[AB-161]_{free}$ : plasma



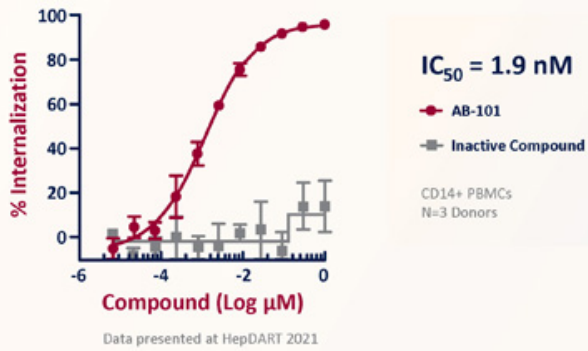
$[AB-161]_{free}$ : liver



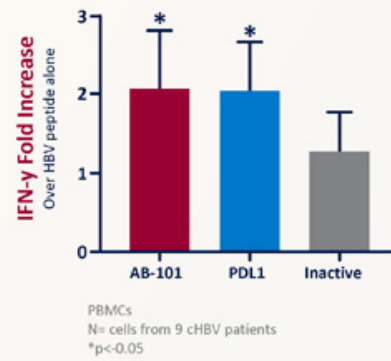
# AB-101: Small-Molecule Oral PD-L1 Inhibitor for HBV

AB-101 is highly potent with demonstrated activity against PD-L1 in cells from chronic HBV patients

AB-101 reduces PD-L1 on the surface of human primary myeloid cells



AB-101 reinvigorates HBV-specific cHBV patient T-cells



# AB-101: Oral PD-L1 Inhibitor for HBV Immune Reactivation

## Rationale

- HBV immune tolerance is a critical driver of cHBV infection
- PD-1:PD-L1 checkpoint axis plays a key role in immune tolerization in cHBV
- PD-L1 expression upregulated during HBV infection
- PD-1 upregulated on HBV-specific T- and B-cells
- Inhibition associated with HBsAg loss in some cHBV patients

## Small-Molecule Inhibitor Approach

- Allows controlled checkpoint blockade
- Enables oral dosing
- Designed to reduce systemic safety issues seen with Abs

## AB-101

- Blocks PD-L1/PD-1 interaction at sub-nM concentrations
- Activates HBV-specific immune responses in T-cells from cHBV patients *in vitro*
- Novel MOA identified
- Demonstrates a robust checkpoint mediated *in vivo* effect
- Improves HBV-specific T- and B-cell responses *ex vivo*

Regulatory approval received in New Zealand to advance AB-101 into a Phase 1 clinical trial; dosing expected to begin Q3 2023

# Coronavirus Program Overview



## Cause & Symptoms

- Coronavirus Infections, such as COVID-19 caused by SARS-CoV-2
- Spreads through breathing out droplets and small particles that contain the virus
- Older adults and people with severe underlying conditions at higher risk of developing serious complications
- Virus continues to mutate with variant strains developing



## Population

- ~6.9M deaths globally<sup>1</sup>
- In US: ~80M cases; 1M deaths (as of March 2022)



## Treatments

### Vaccines

- Durability of effect uncertain, boosters required, limited efficacy on variant strains

### Therapies

- Sub-optimal



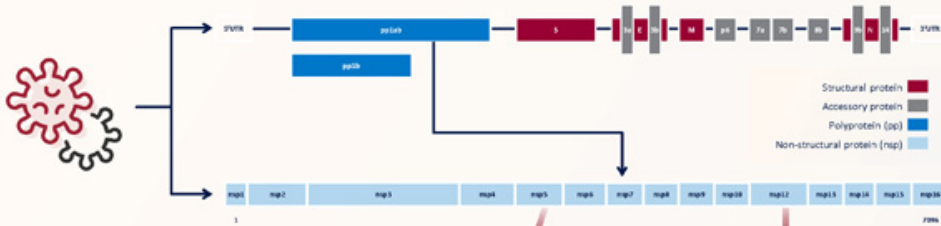
## Rationale

- Pan-coronavirus focused: need for effective and safe therapies to combat COVID-19 and future coronavirus outbreaks
- Address essential viral targets – nsp12 viral polymerase and nsp5 viral protease
- Potential for combo therapy to enhance efficacy and reduce symptomology



# Coronavirus Strategy

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



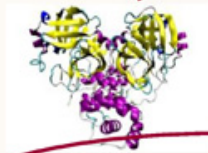
**X-CHEM**  
TAKE THE LEAD. DRIVE DISCOVERY.

**PROTEROS**  
REACH RIGHT FASTER

## Collaboration

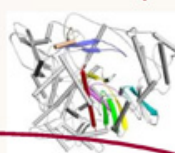
- Proprietary DEL library screening and structural biology for M<sup>pro</sup> inhibitor discovery
- First milestone reached; several unique compound series that inhibit nsp5 protease identified
- Advancing to lead optimization stage

nsp5 protease & nsp12 polymerase essential enzymes for replication



COVID-19 Virus nsp5 / M<sup>pro</sup>

**Viral Protease**  
de novo design



COVID-19 Virus nsp12

**Viral Polymerase**  
nucleos(t)ides

## Arbutus Strategy

- Pan-coronavirus focused
- Combination therapy approach
- Expected to complete IND-enabling studies for AB-343 in 2H 2023

# AB-343: M<sup>PRO</sup> Coronavirus Candidate

## Activity

- Highly potent (IC<sub>50</sub> < 8nM)
- Equipotent against all known COVID-19 variants
- Robust activity against M<sup>PRO</sup> resistant variants

## Safety

- Highly selective for coronavirus M<sup>PRO</sup> vs human proteases
- Clean cell toxicity profile
- Off-target assessment results unremarkable

## Convenience

- Preclinical PK supports ritonavir-free dosing
- No anticipated drug-drug interactions
- Data supports combination strategy

AB-343 is currently in IND-enabling studies

# 2023 Key Milestones

Cash balance\* of \$164M as of June 30, 2023, cash runway into Q1 2025; 2023 net cash burn of between \$90M and \$95M

Milestone	Anticipated Timing 2023
Imdusiran: Dose first patient in the imdusiran+VTP-300+Nivo arm of the ongoing Phase 2a Vaccitech trial	1H <input checked="" type="checkbox"/>
Imdusiran: Preliminary IFN data from patients in the AB-729-201 clinical trial	1H <input checked="" type="checkbox"/>
Imdusiran: Follow-up off-treatment data from AB-729-001 clinical trial	1H <input checked="" type="checkbox"/>
Imdusiran: Preliminary data from Phase 2a POC clinical trial with imdusiran+VTP-300+NA therapy	2H
AB-161: Initial data from Phase 1 single-ascending dose clinical trial in healthy subjects	2H
AB-101: Initiate single-ascending dose portion of Phase 1 clinical trial in healthy subjects	2H
AB-343, COVID M <sup>pro</sup> : Complete IND-enabling studies	2H
COVID Nsp12: Nominate a clinical candidate and initiate IND-enabling studies	2H

Thank You

