

Singularly Focused on HBV

March 2020

NASDAQ: ABUS 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 expected cash burn rate for 2020; our expectation for AB-729 for preliminary results from our Phase I trial to be available late in the first quarter 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; 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. 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. A more complete discussion of the risks and uncertainties facing Arbutus appears in Arbutus' Annual Report on Form 10-K and Arbutus' periodic disclosure filings which are available at www.sec.gov and at www.sedar.com.

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HBV Presents a Significant Unmet Medical Need



Sources: Global Hepatitis Report and Hepatitis B Fact Sheet, WHO (2017) <u>http://www.who.int/mediacentre/factsheets/fs204/en/</u> Kowdley et al. Hepatology (2012) Prevalence of Chronic Hepatitis B Among Foreign-Born Persons Living in the US by Country of Origin

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BIOPHARMA

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



Achievable HBV Cure Rates with Current SOC

Compelling Growth Opportunity in the HBV Market



An HBV curative regimen

would substantially increase **diagnosis** and **treatment** rates to unlock significant **market growth opportunities.**



Investment Highlights

Singular therapeutic focus - curing chronic Hepatitis B Virus (HBV) Infection



Proven Leadership Team

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







Chief Development Officer



Michael J. McElhaugh



HBV Lifecycle Illustrates Key Points for Intervention

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



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

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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

Duration of HBsAg reduction supports once per month dosing

Demonstrated complementarity with capsid inhibitors

Phase I initiated in July 2019

Preliminary results in healthy volunteer and HBV subjects expected in late Q1 2020



AB-729 In Vivo Single & Multiple Dose Response & Duration

Strong dose response in AAV mouse model

Stepwise reduction of HBsAg with monthly repeat dose administration





AB-729 Phase 1a/b Study Design

Preliminary results anticipated late Q1 2020

<u>Part 1:</u> Blinded SAD in Healthy Volunteers	<u>Part 2:</u> SAD in HBV Subjects	<u>Part 3:</u> 3 and 6 Month Multiple- dose in HBV Subjects
Starting dose 60 mg	Starting dose selected from Part 1	Dose selected from part 2
6 subjects per cohort (4 active, 2 placebo)	6 subjects per cohort	7 subjects per cohort
	CHB on stable NA Rx (HBV DNA neg-), HBeAg pos+ or neg-	CHB on stable NA Rx (HBV DNA neg-), HBeAg pos+ or neg-
	Naïve CHB, HBeAg pos+ or neg-	Rx Naïve CHB, HBeAg pos+ or neg-



AB-452 and Next Gen RNA Destabilizer Program

- AB-452 development discontinued following extensive pre-clinical evaluations
- However, we believe this target offers potential for an oral HBsAg reducing agent and potential all oral combination therapy
- Continuing active research and development of a next generation small molecule



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
- Leverages a novel binding site within the core protein dimer-dimer interface
- Improved intrinsic potency with EC50 < 10 nM</p>
- Active against NA resistant variants
- Potential to address known capsid resistant variants T33N and I105T
- Provides the potential for low dose and wide therapeutic window
- Projected to be once daily dosing
- Pangenotypic
- Combinable with other MOA agents

AB-836: A Next Generation Capsid Inhibitor

	HBV DNA / 1° Mechanism				cccDNA Formation / 2° Mechanism	Human Serum Shift
Compound	HepDE19 (EC ₅₀ μM)	HBV infected PHH (EC₅₀ μM)	HBV infected HepG2-NTCP-C4 (EC ₅₀ μM)	Core I105T Mutation (EC ₅₀ μM)	HBV infected HepG2-NTCP-C4 (HBsAg EC₅₀ μM)	(FC in EC₅₀ in 40% Human Serum)
AB-506	0.077	0.032	0.101	1.26	1.430	бх
AB-836	0.010	0.002	0.012	0.118	0.196	2x



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Unique Binding Site

HAP: Heteroaryldihydropyrimidine | SBA: Sulfamoylbenzamide | PHH: Primary Human Hepatocytes

Key Objectives for 2020

Objective	Anticipated Timing
AB-729 preliminary phase 1a/1b single dose data	Late 1Q 2020
AB-729 multiple dose data	2H 2020
AB-836 complete IND enabling studies	2H 2020



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