JÆTERNA VZENTARIS In Pursuit of Medical Innovations

June 2022 Corporate Presentation

Forward-Looking Statements

This document contains forward looking statements (as defined by applicable securities legislation) made pursuant to the safe harbor provision of the US Securities Litigation Reform Act of 1995 and forward-looking information (as defined under applicable Canadian securities laws), which reflect the current expectations regarding future events of Aeterna Zentaris Inc. (the "Company", "we", "our"). Forward looking statements and forward-looking information may include, but are not limited to statements preceded by, followed by, or that include the words " will," "expects," "believes," "intends," "would," "could," "plans," or ""planned" and similar terms that relate to future events, performance, or our results.

Such statements include, but are not limited to, the Company's ability to deliver multiple development and regulatory milestones with respect to diagnostics, therapeutics and vaccines, including developing manufacturing processes for immunomodulating therapeutics and selection of development candidates, the potential of oral Coronavirus vaccines to induce mucosal immunity to prevent infection and avoid transmission, the Company's expectations regarding the development and manufacturing of oral Coronavirus vaccines, the potential use of MacrilenTM macimorelin as a therapeutic, including its use as a potential treatment for Amyotrophic Lateral Sclerosis (ALS, Lou Gehrig's disease), the size, timing and scope of our commercial and development pipeline for AIM biologicals, AEZS-150, macimorelin as a therapeutic and oral Coronavirus vaccines, the Company's expectations regarding its cash runway and its ability to fund operations beyond 2023 and the expected timing of future key milestones, studies, agreements and approvals.

Forward looking statements and forward-looking information contained in this presentation are based on our opinions, estimates and assumptions in light of our experience and perception of historical trends, current conditions and expected future developments, as well as other factors that we currently believe are appropriate and reasonable in the circumstances. There can be no assurance that the underlying opinions, estimates and assumptions will prove to be correct.

Forward looking statements involve known and unknown risks and uncertainties, including those discussed in this presentation and in our Annual Report on Form 20F, under the caption "Key Information Risk Factors" filed with the relevant Canadian securities regulatory authorities in lieu of an annual information form and with the US Securities and Exchange Commission. Known and unknown risks and uncertainties could cause our actual results to differ materially from those in forward looking statements and forward-looking information. Such risks and uncertainties include, among others, our heavy dependence on the success of MacrilenTM macimorelin and related out licensing arrangements and the continued availability of funds and resources to successfully develop and commercialize MacrilenTM and our in licensed products and technologies, the ability of the Company to enter into licensing, development, manufacturing and marketing and distribution agreements with other pharmaceutical companies, universities or others and keep such agreements in effect (including that the Company the Company may be unable to successfully negotiate a license agreement for any technology or products for which it has an option), the Company's ability to identify therapeutic uses for MacrilenTM macimorelin or to in license product candidates, the Company's reliance on third parties for the manufacturing and commercialization of MacrilenTM macimorelin, potential delay or termination or lack of success of resulting in significant litigation or arbitration, and, more generally, uncertainties related to the regulatory process, the degree of market acceptance of MacrilenTM macimorelin, the impact of securities class action litigation, shareholder lawsuits or other litigation on our cash flow, results of operations and financial position, our ability to protect our intellectual property, general changes in economic conditions and the impact of the COVID-19 pandemic on our operations, plans and prospects, including to the initiation and

Readers of this presentation should consult our quarterly and annual filings with the Canadian and US securities commissions for additional information on risks and uncertainties Given these uncertainties and risk factors, readers are cautioned not to place undue reliance on these forward looking statements and forward looking information. The forward looking statements and information in this presentation are made as of the date hereof and we disclaim any obligation to update any such factors or to publicly announce any revisions to any of the forward looking statements or forward looking information contained herein to reflect future results, events or developments, unless required to do so by a governmental authority or applicable law.

Certain Other Matters

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Investment Highlights

Diversified portfolio focused on areas of significant unmet medical need

Leveraging network of large pharmaceutical and university partners and collaborators

Executing strategy to deliver multiple development and regulatory milestones

Strong financial position with runway expected to fund operations through 2023¹

Key Focus Areas:

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Diagnostics

• Rare endocrine disorders

Therapeutics

- Auto-immune diseases
- Neurodegenerative disease
- Endocrine disorders

Salmonella-Based Vaccine Platform

- SARS-CoV-2 (Covid-19)
- Chlamydia



Diagnostic Commercial and Development Pipeline

	Program	Indication	Preclinical	Phase 1	Phase 2	Phase 3	Commercial	Co-Development Partner
ostics	Macimorelin	Adult Growth Hormone Deficiency (AGHD)						novo nordisk
Diagn	Macimorelin	Childhood-Onset Growth Hormone Deficiency (CGHD)						novo nordisk

Commercial Rights





Therapeutic and Vaccine Development Pipeline

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	Program	Indication	Preclinical	Phase 1	Phase 2	Phase 3	Program Highlights	
Therapeutics	AIM Biologicals	Neuromyelitis Optica Spectrum Disorder (NMOSD) Parkinson's Disease (PD)					Abstract accepted at 13 th Intl. Congress on Autoimmunity (Athens, June 2022) Presented positive preclinical results at IMMUNOLOGY2022 [™]	Julius-Maximilians- UNIVERSITÄT WÜRZBURG
	Macimorelin	Amyotrophic Lateral Sclerosis (ALS, Lou Gehrig's disease)					Entered material transfer agreement and option to in-license in January 2021	THE UNIVERSITY OF QUEENSLAND
	AEZS-150 (Delayed clearance parathyroid hormone)	Chronic Hypoparathyroidism					In-licensed program in March 2021	The University Of Sheffield.
							Program Highlights	
Vaccine	Salmonella-Based Vaccine Platform	COVID-19 (SARS-CoV-2) Chlamydia Trachomatis					In-licensed program in March 2021 In-licensed program in September 2021	Julius-Maximilians- UNIVERSITÄT WÜRZBURG



Development Programs

Advancing Therapeutics and Vaccines to Secure Long-Term Growth



AIM Biologicals

Targeted Immunomodulating Therapeutics



Technology platform enabling highly specific treatment for auto-immune diseases with well-defined target antigens



Modulation of immune response towards autoantigens by induction of antigen-specific regulatory T cells and by elimination of cognate antigenspecific cytotoxic T cells



Entered exclusive license and R&D agreements with Julius-Maximilians-University of Wuerzburg in January and September 2021



Targeted, highly specific, autoimmunity modifying proteins

Technology is based on "**feto-maternaltolerance**" – It is derived from the body's natural process that protects a fetus against the mother's immune system while the mother is protected against pathogens



Fetal cells persist in mothers for decades after delivery and vice versa → lasting, selective immune tolerance which cannot be explained by barrier function¹



The Promise of Targeted, Physiological, Antigen-Specific Immunomodulation







Autoimmune Diseases

Enhanced by autoreactive effector (cytotoxic) T cells Ameliorated by protective regulatory T cells

Immunosuppressive Therapeutics

Reduce disease symptoms Often cause severe side effects

Antigen-specific Immunomodulation

Various antigens may be used for highly selective de-sensibilization Relevant side effects should be avoided



AIM Biologicals Concept¹

- Adaptable platform of immunomodulating molecules
- Soluble HLA-G ß2m fusion proteins presenting a peptide (= antigen)* which is coupled to the proteins via a linker
- The proteins interacts with the target cells via the T-cell receptor (TCR) and ILT-2 to induce immune tolerance



* exemplified by NMOSD-specific antigen



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The Promise of Targeted, Physiological, Antigen-Specific Immunomodulation



Potential selective treatment option for neuromyelitis optica spectrum disorder (NMOSD)



Well-defined antigen: Aquaporin-4 (AQP4)



NMOSD is an auto-antibody mediated inflammatory CNS orphan disorder with significant unmet medical need affecting 1 per 100,000 people¹

Next Steps:

- *In-vitro* and *in-vivo* assessments to select development candidate
- Manufacturing process development for selected candidate



Potential treatment option for Parkinson's disease (PD)

 α -Synuclein (α -Syn) described as target



Parkinson's disease is a neurodegenerative movement disorder affecting over 7 million people worldwide, growing evidence suggests that PD could be considered as an autoimmune disease²

Recent Highlights:

- Design and production of antigen-specific AIM biologics molecules
- ✓ *In-vitro* and *in-vivo* assessments in relevant disease models



1: Hor et al.; Frontiers in Neurology 2020

AIM Biologicals Demonstrated Improvements in Preclinical Parkinson's Disease Model

Significant induction of regulatory T cells, improvement in motor function and rescue of substantia nigra neurons





AIM Bios induce regulatory CD8+CD122+ T cells in spleen and lymph nodes



EV+PBS A53T+0.5µg/g aSy68 DbG A53T+2µg/g aSy68_ DbG

> AIM Bios show a trend towards improvement of motor function in the Rotarod test

Significantly improves survival of SN neurons

Number of dopaminergic neurons in SN





AEZS-150: Delayed Clearance Parathyroid Hormone (DC-PTH) Fusion Polypeptide



PTH is a key regulating hormone essential for calcium homeostasis and renal phosphate clearance



Potential to be a self-administered pen to help maintain normal serum calcium and phosphate levels in a once weekly treatment versus current daily injections

Hypoparathyroidism

Body produces abnormally low levels of PTH



Orphan indication ~23-37 per 100,000¹





AEZS-150: Delayed Clearance Parathyroid Hormone (DC-PTH) Fusion Polypeptide

AEZS-150: Fusion-Protein of PTH [1-34] coupled via a linker to a modified growth-hormone binding protein (GHBP)¹

Delayed clearance in comparison to PTH[1-34]

Potential to control serum calcium levels by once weekly treatment PTH [1-34]





Next Steps:

- In depth characterization of development candidate (*in-vitro* and *in-vivo*)
- Manufacturing process development ongoing



Salmonella-Based Vaccine Platform



Live-attenuated bacterial vaccine based on the Salmonella typhi Ty21a carrier strain currently used as a typhoid vaccine



This new, alternative and unprecedented approach with *Salmonella* based vaccines should have the potential to induce mucosal immunity, not only in respiratory, but also in urogenital tract, in order to form a barrier for pathogens entering the body

- \rightarrow Potential to evade infection
- \rightarrow Potential to avoid transmission

UNIVERSITÄT WÜRZBURG Entered exclusive license and R&D agreements with Julius-Maximilians-University of Wuerzburg in 2021 Potential for temperature stable supply chain: ~2°C - 8°C¹

Potential to induce systemic and mucosal immunity to prevent infection → COVID-19 vaccine → Chlamydia Trachomatis vaccine

Adaptable antigen expression

Salmonella Typhi Ty21a carrier strain has been safely used worldwide in more than 150 million administered doses¹



Oral Coronavirus (SARS-CoV-2) Vaccine

Currently undergoing pre-clinical studies for the prevention of coronavirus diseases, including COVID-19 (SARS-CoV-2)



Induction of Immunity

Potential to induce mucosal immunity to prevent infection and avoid transmission

Multiple-Antigens

Higher likelihood for improved defense against mutated virus variants if not only spike protein is used as antigen



Chlamydia Trachomatis Vaccine

- Sexually transmitted gram-negative bacterium infecting over 130 million subjects annually
- Whereas symptomatic infection can be treated by antibiotics, asymptomatic disease can spread to the reproductive tract inducing infertility, miscarriage, or ectopic pregnancy, which is a life-threatening condition
- Ocular infections can lead to inclusion conjunctivitis or trachoma, which is the primary source of visual impairment or infectious blindness
- High medical need to develop a vaccine providing mucosal immunity preventing infection
- In-depth expertise and well-established advanced disease models at University of Wuerzburg²



² <u>https://www.biozentrum.uni-wuerzburg.de/en/mikrobio/forschungsschwerpunkte/chlamydiales</u> ³ Paavonen and Eggert-Kruse, Hum Reprod Update Sep-Oct 1999;5(5):433-47.





Chlamydia trachomatis (magenta) in human cells (green)



Infection with *Chlamydia trachomatis* leads to visual impairment or infectious blindness of about 1.9 million people¹



Approximately 4% of women with chlamydial lower genital tract infection will develop chronic pelvic pain, 3% infertility, and 2% adverse pregnancy outcome.³

Next Steps:

- Design and preparation of candidate vaccine strains
- In-vivo immunology experiments and challenge studies

Macimorelin For the Treatment of ALS







Exclusive rights for AEZS to negotiate a license agreement for the commercial use of the results

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Investigation as a potential treatment for Amyotrophic Lateral Sclerosis (ALS, Lou Gehrig's disease) University researchers to secure grants and conduct pre-clinical and clinical studies

ALS is a neurodegenerative motoneuron disease with progressive loss of motor neurons that control voluntary muscles

Disease symptoms of ALS include muscle twitches, cramps and stiffness, gradual increasing weakness and muscle atrophy, finally leading to respiratory failure in most patients

Treatment approaches consist mainly of supportive therapy, including physical therapy, nutritional and breathing support



The Science Behind Macimorelin in ALS

Importance of Ghrelin and the GH/IGF-1 axes in ALS¹

- Ghrelin is a natural endogenous hormone that stimulates appetite and lowers metabolic rate to promote weight gain
- Ghrelin also regulates the release of the neuroprotective and anabolic hormones growth hormone (GH) and insulin-like growth factor-1 (IGF-1)
- Ghrelin acts via the GHSR1a, which is expressed throughout the body, including the cerebral cortex, spinal cord and muscle (i.e. tissues directly impacted in ALS)
- Majority of motor neuron disease (MND) patients have a moderate to marked GH deficiency²
- Treatment with ghrelin or the ghrelin mimetic GHRP3 was shown to slow weight loss, improvement of muscle strength, and extension of survival in the SOD1G93A mouse model of ALS³. However, clinical use of ghrelin is limited due to its rapid clearance from circulation.

1: Bianchi et al., International Journal of Molecular Sciences, 2017, 18.





The Science Behind Macimorelin in ALS

Macimorelin is a Ghrelin mimetic and modulates various disease-relevant processes



Accelerated disease progression

Next Steps:

- Assessment of alternative formulations
- Proof of concept with macimorelin in disease specific SOD1 mouse model (ongoing)
- TDP-43 transgenic mouse model
- Formalized pre-clinical development

Macimorelin has the potential to benefit patients and to slow disease progression



Roadmap for Advancing Development Programs Towards IND Filing



Macimorelin / Macrilen™

A Disruptive Oral Diagnostic Test Solution for Growth Hormone Deficiency

First and only AGHD test approved by US FDA and European Commission



Growth Hormone is Critical to Lifelong Health



Produced by the pituitary gland (located at the base of the brain)

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Children Promotes growth



Reduction in auxological parameters:

- Short stature
- Low growth velocity (speed) for age
- · Increased fat around the waist
- Delayed tooth development

Adults

Maintains normal body stature and regulates metabolism



No clear signs or symptoms, but recognized by:

- Metabolic syndrome
- Osteoporosis
- Muscle wasting
- Impaired quality of life
- Increased risk of:
- Cardiovascular (CV) issues
- Bone fractures



Macimorelin

Only Approved Oral Diagnostic for GHD

No Other FDA or EC Approved Oral Test

Insulin tolerance test (ITT) considered the "Gold Standard" in GHD detection procedures^{1,2}

Not FDA or EC approved or regulated

*"[ITT Test] is increasingly used less frequently in the U.S. because of safety concerns."*²

"Because the **macimorelin** test is simple, well tolerated with minimal side effects, and of shorter duration with only 3 to 4 blood draws compared to other GH–stimulation tests, it is anticipated that its use will increase over time."²

"Very promising test that is easy to conduct with high reproducibility, safety, and diagnostic accuracy comparable to the ITT...test" ²

1: Molitch et al. J Clin Endocrinol Metab. 2011; 1587-1609 2: AACE 2019 Guidelines: American Association Of Clinical Endocrinologists And American College Of Endocrinology Guidelines For Management Of Growth Hormone Deficiency In Adults And Patients Transitioning From Pediatric To Adult Care, 2019



Pivotal Phase 3 DETECT¹ Study for Diagnosis of CGHD

Currently Enrolling Subjects and Dosing is Underway

• Open-label, single dose, multicenter, multinational

US, Czech Republic, Germany, Poland, Georgia, Italy, Serbia, Romania, Russia, Slovenia and Ukraine

- Macimorelin GHST will be performed twice (for repeatability data)
- Two standard GHSTs as controls: arginine (i.v.), clonidine (p.o.)
- Design suitable to support claim for potential of macimorelin as stand-alone test
- The impact of delays due to the COVID-19 pandemic and the Russian invasion of Ukraine two countries where we planned to recruit patients will extend the recruitment phase into H2 2023.

Children and adolescents from 2 to less than 18 years of age with suspected GHD to be enrolled (Years) <18 2 ≥ 100 subjects worldwide \geq 40 pre-pubertal and 2222 <u>ж</u>шшш ШШШШ 40 pubertal subjects ≥ 25 subjects expected Th, to be enrolled in the U.S.



Macimorelin Commercial Rights

Actively seeking commercial partners in ROW

novo nordisk

License and Assignment Agreement

- Territories: United States and Canada
- Royalties on sales
- Sales milestones
- Aeterna Zentaris controls API supply chain and provides API
- Co-development for expansion into CGHD
 - Novo Nordisk to fund 100% of budgeted Study P02 trial expenses up to €9 million
 - Potential additional expenses to be shared



Corporate Overview



Financial Snapshot NASDAQ: AEZS / TSX: AEZS

Cash runway expected to fund operations through 2023¹

~\$64M Cash on Hand As of March 31, 2022

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~\$25M Market Cap²

~121M Shares Outstanding³ **~601K** 3 month Avg. Volume⁴

Based on Management's current expectations and planned development activities
Based on May 31, 2022 closing price of \$0.2050 per share on NASDAQ and the number of issued and outstanding AEZS shares on that date
Information as of February 22, 2021
Based on information as of May 31, 2022 for the 3-month average daily trading volume on NASDAQ



Management



Klaus Paulini, PhD President and Chief Executive Officer; Managing Director, Aeterna Zentaris GmbH



Eckhard Guenther, PhD

SVP Business Development and Alliance Management Managing Director, Aeterna Zentaris GmbH





Nicola Ammer, MD *SVP Clinical Development, Chief Medical Officer*



Michael Teifel, PhD SVP Non-Clinical Development, Chief Scientific Officer





Investment Summary

Advancing diversified pipeline across multiple high-value therapeutic areas

High-Value Focus Areas:

- Endocrine disorders
- Auto-immune diseases
- Neurodegenerative disease
- COVID-19 oral vaccine
- Chlamydia oral vaccine

- Only oral drug indicated for diagnosis of adult growth hormone deficiency
- Expanding into childhood growth hormone deficiency

Leveraging network of large pharmaceutical and university partners and collaborators

Strong Financial Position with Runway Expected to Fund Operations Through 2023¹



JÆTERNA VZENTARIS In Pursuit of Medical Innovations

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