

ACTIVATING THE POWER WITHIN

VBI-2901: Multivalent Coronavirus Vaccine

Broadly Reactive Preclinical Immunity Supporting Development World Vaccine Congress



Forward-Looking & Safe Harbor Statements

Certain statements in this presentation that are forward-looking and not statements of historical fact are forward-looking statements within the meaning of the safe harbor provisions of the Private Securities Litigation Reform Act of 1995 and are forward-looking information within the meaning of Canadian securities laws (collectively "forward-looking statements").

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All such forward-looking statements made herein are based on our current expectations and we undertake no duty or obligation to update or revise any forward-looking statements for any reason, except as required by law.





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Enveloped Virus Like Particle (eVLP) Platform

Ideal Platform to Leverage: eVLPs have Structural Homology with SARS-CoV2

eVLPs *Enveloped Virus-Like Particles*

- VBI's Proprietary Platform Technology -

eVLPs expand the list of potentially-viable target indications by providing a stable core (Gag Protein) and lipid bilayer

Flexible and customizable

Highly immunogenic with demonstrated safety profile

A perfect match for enveloped viruses (such as coronaviruses)





Multiple eVLP Candidates have Clinical Experience

VBI Pipeline: Balanced Portfolio of Therapeutic & Preventative Vaccines

The eVLP Platform is a Foundation for the Majority of VBI's Clinical Programs

Disease	Name/Program	Technology	Preclinical	Phase 1	Phase 2	Phase 3	Registration/ Commercial	
Approved Vaccines								
Hepatitis B (HBV)	PreHevbrio ^{1,2} Hepatitis B Vaccine (Recombinant)	VLP						
Therapeutic Candidates								
Hepatitis B (HBV)	VBI-2601 (BRII-179)	VLP						
Glioblastoma (GBM)	VBI-1901	eVLP						
Other Virus+ Tumors	Undisclosed	eVLP						
Prophylactic Candidates								
Cytomegalovirus (CMV)	VBI-1501	eVLP						
COVID-19 (Ancestral)	VBI-2902 (monovalent)	eVLP						
COVID-19 (Beta Variant)	VBI-2905 (monovalent)	eVLP						
Coronaviruses	VBI-2901 (multivalent)	eVLP						
Coronaviruses	Undisclosed (multivalent)	eVLP						
Zika	VBI-2501	eVLP						



¹Approved for use in the U.S. for the prevention of infection caused by all known subtypes of hepatitis B virus in adults 18 years of age and older ²Approved for use in Israel, under the brand name Sci-B-Vac[®], for active immunization against hepatitis B virus (HBV) infection

VBI-2900 Program is Focused on Long-Term Protection Against Coronaviruses

Program Goals: Broad protection, exceptional safety, and durability of immune response

	VBI-2902	VBI-2905	VBI-2901				
	Monovalent COVID-19	Monovalent COVID-19 B.1.351 Variant	Trivalent Pan-Coronavirus				
Schematic	eVLP	evlp	eVLP				
Construct Design	SARS-CoV-2 Ancestral/Wuhan spike antigen	SARS-CoV-2 Beta (B.1.351) spike antigen	SARS-CoV-1, MERS, and SARS-CoV-2 spike antigens				
Adjuvant & Presentation	Alum adsorbed, liquid @ 4-8C with stability > 2 years (based on platform data)						
TPP/Development Rationale	 Human proof-of-concept vs. ancestral strain 	 Demonstration of strain specific immunity as booster 	 Broad protection Safe & well tolerated Ideal for VoCs and potential for emerging variants/future zoonotic 				



VBI's coronavirus pipeline is supported by partnerships with :



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▼ VBI-2902 : Human Proof-of-Concept vs. ▼ SARS-CoV-2 Ancestral/Wuhan Strain

Preclinical Data from VBI-2902 Candidate Confirmed Value of eVLP Presentation

Preclinical data demonstrated superior immunogenicity of matched-Spike protein on eVLPs vs. standard recombinant protein (both Alum adjuvanted)







VBI-2902a Clinical Data Update

PNA50 neutralizing antibody titers project a high degree of efficacy (> 90%)



Pseudoparticle Neutralization Titers

WHO normalized, validated PNA assay, sera tested 28 days after a 2nd dose in seronegative subjects (Gilbert PB, 2021):

VBI

Moderna: AZ:

VBI:

83 IU50/ml → 90% efficacy 140 IU50/ml → 90% efficacy

176 IU50/ml

T-Cell Immunity is Induced Against Both S1 and S2 Domains of Spike Protein after 2 Doses of VBI-2902





VBI-2902 Demonstrates Clean Safety & Tolerability : No Grade 3 or 4 Events Observed

Local Reactogenicity:

Dose #2	Group Gl VBI-2902a + Placebo n=19	Group 2 VBI-2902a + VBI-2902a n=18	Group 3 Placebo n=18
Any, n (%)	6 (31.6)	12 (66.7)	8 (44.4)
Pain, n (%) Grade 1 Grade 2 Grade 3 Grade 4	4 (21.1) 4 (21.1) 0 0 0	11 (61.1) 10 (55.6) 1 (5.6) 0 0	8 (44.4) 8 (44.4) 0 0 0
Tenderness, n (%) Grade 1 Grade 2 Grade 3 Grade 4	5 (26.3) 5 (26.3) 0 0 0	10 (55.6) 7 (38.9) 3 (16.7) 0 0	5 (27.8) 3 (16.7) 2 (11.1) 0 0
Redness, n (%)	0	0	0
Swelling, n (%)	0	0	0
Itchiness, n (%)	0	0	0

- VBI-2902 was well-tolerated with no safety signals observed
- Reactogenicity was mild-to-moderate and lasted for 1-2 days
- There was no increase in reactogenicity with subsequent doses

Systemic Reactogenicity:

Dose #2	Group Gl VBI-2902a + Placebo n=19	Group 2 VBI-2902a + VBI-2902a n=18	Group 3 Placebo n=18
Any, n (%)	7 (36.8)	10 (55.6)	10 (55.6)
Fatigue, n (%)	7 (36.8)	4 (22.2)	5 (27.8)
Grade 1	4 (21.1)	2 (11.1)	3 (16.7)
Grade 2	3 (15.8)	2 (11.1)	2 (11.1)
Grade 3/4	0	0	0
Fever, n (%) Grade 1 Grade 2 Grade 3/4	0	0	1 (5.6) 1 (5.6) 0 0
Headache, n (%)		7 (38.9)	5 (27.8)
Grade 1	0	6 (33.3)	4 (22.2)
Grade 2	U	1 (5.6)	1 (5.6)
Grade 3/4		0	0
Myalgia, n (%)	1 (5.3)	4 (22.2)	4 (22.2)
Grade 1	1 (5.3)	3 (16.7)	2 (11.1)
Grade 2	0	1 (5.6)	2 (11.1)
Grade 3/4	0	0	0
Nausea / Vomiting, n (%)			1 (5.6)
Grade 1	0	0	1 (5.6)
Grade 2	0	0	0
Grade 3/4			0
Diarrhea, n (%)	1 (5.3)		1 (5.6)
Grade 1	1 (5.3)	Ο	1 (5.6)
Grade 2	0	U	0
Grade 3/4	0		0

₹ Trivalent eVLP Candidate (VBI-2901) vs. Ancestral (VBI-2902) and Beta (VBI-2905) Monovalent Candidates

Despite Current Challenges: Data Suggests Broadly Reactive, Durable Immunity is Possible – Answer May Lie in the Past

- Sauer et al : Isolated broadly cross-reactive mAbs after immunizing with MERS, then SARS-CoV-2 vaccines
- Tan et al : Broad immunity found in SARS-1 survivors who had been vaccinated with SARS-CoV-2 vaccine



Sequence Homology of VBI-2900 Vaccine Candidates

Spike sequence homology for VBI-2901 (Wul, MERS, SARS), VBI-2902 (Wul), and VBI-2905 (Beta) as they relate to not only previous & currently circulating variants, but also bat (RaTG13) and pangolin (Pan-GX, Pan-GD) coronaviruses distant to circulating human strains

	MERS	Pan-GX	RaTG13	Pan-GD	Delta	Kappa	C.1.2	Beta	Lambda	Wu1	Alpha	Rs4231	SARS1
MERS	100.00	20.45	20.45	20.45	20.45	20.45	20.45	20.45	20.91	20.45	20.45	20.55	21.10
Pan-GX	20.45	100.00	87.89	87.89	85.65	86.10	86.10	86.55	86.10	86.55	86.55	76.13	75.11
RaTG13	20.45	87.89	100.00	89.69	90.13	89.69	90.13	89.69	89.69	90.13	90.13	76.13	75.11
Pan-GD	20.45	87.89	89.69	100.00	95.96	95.96	95.52	95.96	95.96	96.86	96.41	77.48	74.21
Delta	20.45	85.65	90.13	95.96	100.00	99.10	97.76	97.76	98.65	99.10	98.65	75.68	73.76
Kappa	20.45	86.10	89.69	95.96	99.10	100.00	98.21	98.21	98.65	99.10	98.65	75.68	73.30
C.1.2	20.45	86.10	90.13	95.52	97.76	98.21	100.00	99.10	97.76	98.65	99.10	75.68	72.85
Beta	20.45	86.55	89.69	95.96	97.76	98.21	99.10	100.00	97.76	98.65	99.10	76.13	73.30
Lambda	20.91	86.10	89.69	95.96	98.65	98.65	97.76	97.76	100.00	99.10	98.65	75.68	73.30
Wul	20.45	86.55	90.13	96.86	99.10	99.10	98.65	98.65	99.10	100.00	99.55	76.13	73.30
Alpha	20.45	86.55	90.13	96.41	98.65	98.65	99.10	99.10	98.65	99.55	100.00	76.13	73.30
Rs4231	20.55	76.13	76.13	77.48	75.68	75.68	75.68	76.13	75.68	76.13	76.13	100.00	81.90
SARS1	21.10	75.11	75.11	74.21	73.76	73.30	72.85	73.30	73.30	73.30	73.30	81.90	100.00



Key Point: VBI-2902 & VBI-2901 share the same Wu1/ancestral SARS-CoV-2 Spike sequence. Addition of MERS & SARS-1 are highly divergent from currently circulating strains

VBI-2901a (Trivalent) PreClinical PoC Data

Trivalent 2901 candidate offers superior breadth relative to monovalent 2902 –trivalent consistently out-performs 2902 on its own turf (strain-matched)



VBI-2901 offered 2-9X higher RBD binding antibodies than VBI-2902

Trivalent VBI-2901 Induced Robust Neutralizing Titers Against an Extended Panel of Variants in Mice

Neutralizing responses elicited by VBI-2901 are 2-5X higher than VBI's ancestral (2902) and Beta (2905) monovalent candidates

Immunogenicity of trivalent VBI-2901a: Three groups of 10 mice were immunized with 2 doses of VBI-2901a, VBI-2902a, or VBI-2905a 3 weeks apart. Blood was collected at day 14 after the last injection for monitoring of humoral responses. Neutralization of EPT measured by PRNT90, neutralization of pseudoparticles expressing S from Wu-1, Delta, and Kappa variants are represented as half-maximum inhibitory dilutions (neutralization ID50). Due to technical limitations, only 8 sera per group were tested against Wu-1 and Kappa pseudoparticles and 4 sera against Delta pseudoparticles.

VBI-2901 Golden Hamster Challenge Data vs. Ancestral SARS-CoV-2

VBI-2901 offered strong protection with highly significant reduction in viremia in lungs & nasal shedding

- Despite dose and strain matching, VBI-2901 (trivalent) offered stronger protection in ancestral challenge vs VBI-2902 (monovalent, ancestral)
- Breadth appears to have clinical benefit in animals
- Reduction in nasal viremia may offer transmission benefit

Figure 12. Infectious virus in nasal washes. Infectious virus was detected using a cell culture method. All nasal wash samples collected from 1, 2 and 3 dpc were examined (10a; n=12). For 5, 7, 9, 11 and 14 dpc, infectious virus was examined on qRT-PCR positive samples plus selected negative controls (10b; n=6). All animals were challenged on Day 0. Each dot represents one animal. Horizontal bars represent medians. Group A=Saline control; Group B=VBI-2902a; Group C=VBI-2901e; LOD=limit of detection. Zero values are included as 1 values for graphing purposes.

VBI-2900 Program Summary & Next Steps

eVLP Benefits

Proof-of-Concept Data Obtained

- Strong immune responses when compared to standard recombinants Multivalency allows for broadly reactive vaccines
- Highly potent immunogenicity, comparable to licensed vaccines Low clinical doses (5ug) & no requirement for novel adjuvants Exceptional safety/tolerability profile typical of alum-adjuvanted subunit vaccines
 - Preclinical data demonstrated improvement of VBI-2901 over monovalent eVLPs

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Future Development

Continued testing of breadth vs. emerging variants

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Summer 2022 : First clinical study of VBI-2901 expected to initiate

Acknowledgements :

VBI Vaccines Inc. | www.vbivaccines.com