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

FORM 6-K

REPORT OF FOREIGN PRIVATE ISSUER  
PURSUANT TO RULE 13a-16 OR 15d-16  
UNDER THE SECURITIES EXCHANGE ACT OF 1934

For the month of June 2020

Commission File Number: 001-38764

APTORUM GROUP LIMITED

Unit 232, 2/F, Building 12W  
Phase Three, Hong Kong Science Park, Pak Shek Kok, N.T.,  
Hong Kong  
(Address of principal executive offices)

Indicate by check mark whether the registrant files or will file annual reports under cover of Form 20-F or Form 40-F: Form 20-F  Form 40-F

Indicate by check mark if the registrant is submitting the Form 6-K in paper as permitted by Regulation S-T Rule 101(b)(1):

Indicate by check mark if the registrant is submitting the Form 6-K in paper as permitted by Regulation S-T Rule 101(b)(7):

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We are filing this report to disclose the revised power point presentation the Company will use during corporate presentations; such power point presentation is incorporated herein by reference.

Neither this report nor the power point presentation attached as an exhibit hereto constitutes an offer to sell, or the solicitation of an offer to buy our securities, nor shall there be any sale of our securities in any state or jurisdiction in which such offer, solicitation or sale would be unlawful prior to the registration or qualification under the securities laws of any such state or jurisdiction.

The information in this Form 6-K, including the exhibit shall not be deemed to be "filed" for the purposes of Section 18 of the Securities Exchange Act of 1934, as amended, and shall not be incorporated by reference into any filing under the Securities Act of 1933, as amended, except as shall be expressly set forth by specific reference in such filing.

This Form 6-K is hereby incorporated by reference into the registration statements of the Company on Form S-8 (Registration Number 333-232591) and Form F-3 (Registration Number 333-235819) and into each prospectus outstanding under the foregoing registration statements, to the extent not superseded by documents or reports subsequently filed or furnished by the Company under the Securities Act of 1933, as amended, or the Securities Exchange Act of 1934, as amended.

#### EXHIBIT INDEX

<b>Exhibit No.</b>	<b>Description</b>
99.1	<a href="#">Power Point Presentation</a>

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2

**SIGNATURES**

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, thereunto duly authorized.

**Aptorum Group Limited**

Date: June 29, 2020

By: /s/ Sabrina Khan

Name: Sabrina Khan

Title: Chief Financial Officer

EXHIBIT INDEX

**Exhibit No.**

**Description**

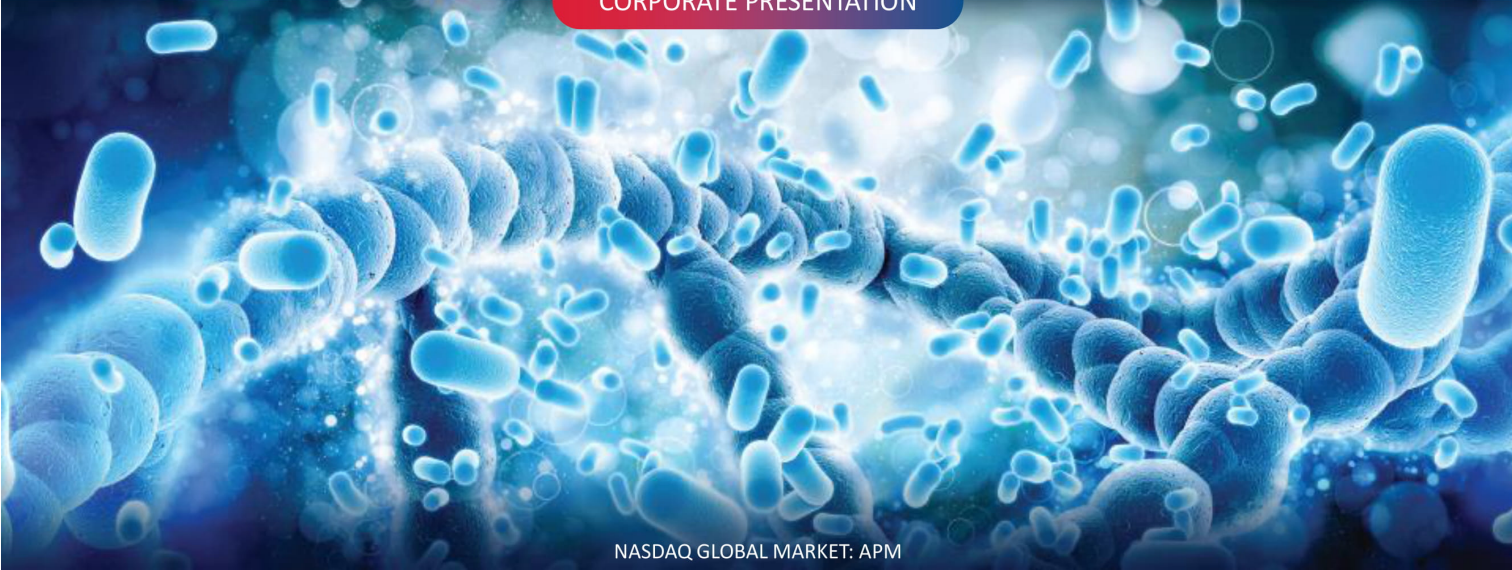
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99.1 [Power Point Presentation](#)



Facilitating Life Science Innovations to Serve Unmet Medical Needs

CORPORATE PRESENTATION



NASDAQ GLOBAL MARKET: APM

# Disclaimer

This document includes statements concerning Aptorum Group Limited and its future expectations, plans and prospects that constitute “forward-looking statements” within the meaning of the Private Securities Litigation Reform Act of 1995. For this purpose, any statements contained herein that are not statements of historical fact may be deemed to be forward-looking statements. In some cases, you can identify forward-looking statements by terms such as “may,” “should,” “expects,” “plans,” “anticipates,” “could,” “intends,” “target,” “projects,” “contemplates,” “believes,” “estimates,” “predicts,” “potential,” or “continue,” or the negative of these terms or other similar expressions. Aptorum Group has based these forward-looking statements, which include statements regarding projected timelines for application submissions, trials and commercialization and market potential of related products, largely on its current expectations and projections about future events and trends that it believes may affect its business, financial condition and results of operations. These forward-looking statements speak only as of the date of this document and are subject to a number of risks, uncertainties and assumptions including, without limitation, risks related to its announced management and organizational changes, the continued service and availability of key personnel, its ability to expand its product assortments by offering additional products for additional consumer segments, development results, the company’s anticipated growth strategies, anticipated trends and challenges in its business, and its expectations regarding, and the stability of, its supply chain, and the risks more fully described in Aptorum Group’s Form 20-F and other filings that Aptorum Group may make with the SEC in the future. As a result, the projections included in such forward-looking statements are subject to change. Aptorum Group assumes no obligation to update any forward-looking statements contained in this document as a result of new information, future events or otherwise.

# About Aptorum Group

## Company information

- Established in 2010, focused on current unmet medical needs, including orphan diseases, infectious diseases, metabolic diseases and women's health, over 15 therapeutic candidates
- **Business Strategy:** From Discovery to Ph2 Proof-of-Concept (PoC)
- **Markets and Regulatory:** Targeted for US FDA clinical, China NMPA and Europe EMA approval and other major countries
- **IPO:** Listed on NASDAQ Global Market (ticker symbol: APM) on December 18, 2018
- **2 Development Sites:** Hong Kong Science and Technology Park (non-GLP, formulation, preclinical work) and Toronto (GLP, GMP, clinical trial coordination)
- ~40 full time staff and ~45 scientists, advisors and consultants with vast experience in drug development and clinical studies



# Aptorum Management and Directors

## Leadership



**MR. IAN HUEN**

*Founder, Chief Executive Officer and Executive Director*

- Over 15 years in global asset management;
- US healthcare equity research analyst at Janus Henderson Group;
- Trustee board member of Dr. Stanley Ho Medical Development Foundation;
- CFA, Princeton University, U.S. (Econ)



**MR. DARREN LUI**

*President and Executive Director*

- Over 13 years in global capital market;
- Extensive experience in Investment in UK, Singapore, US, etc.;
- ICAS, CFA & Associate of Chartered Institute of Securities & Investments (UK);
- First-Class Honors from Imperial College (Biochemistry)



**DR. CLARK CHENG**

*Chief Medical Officer and Executive Director*

- Almost 10 years working in Raffles Medical Group as Operations Director and Deputy General Manager;
- Received medical training at the University College London in 2005 & obtained membership of the Royal College of Surgeons of Edinburgh in 2009;
- MBA, University of Iowa, U.S.



**MISS SABRINA KHAN**

*Chief Financial Officer*

- Almost 10 years serving US & Asian healthcare companies;
- Extensive experience in business development, restructuring, US & Asian IPO, and M&A deals;
- Solid accounting experience gained from Big 4;
- Advanced China Certified Taxation Consultant;
- CPA, University of Hong Kong (BBA(Acc. & Fin))



**DR. THOMAS LEE WAI YIP**

*Head of Research and Development*

- Former Assistant Professor at The Chinese University of Hong Kong (CUHK) specialized in drug delivery and formulation development;
- 10 years from Novartis & Celgene;
- B.Pharm.(Hons), CUHK; Ph.D. in Pharmaceutical Sciences (Drug Delivery), the University of Wisconsin-Madison



**DR. ANGEL NG SIU YAN**

*Chief Operating Officer*

- Research Officer cum Project Manager at The University of Hong Kong (HKU) towards cadaveric trial for a novel soft robotics medical device;
- Former Project Manager at Hong Kong Science & Technology Parks Corporation and CUHK;
- B.Sc (Hons), HKU; M.Sc in Composite Materials, Imperial College London; Ph.D. in Mechanical Engineering, HKU

## Independent Non-Executive Directors



**PROFESSOR DOUGLAS ARNER**

Kerry Holdings Professor in Law, HKU



**DR. JUSTIN WU**

COO of CUHK Medical Centre



**DR. MIRKO SCHERER**

CEO of CoFes China



**MR. CHARLES BATHURST**

Founder of Summerhill Advisors Limited



# Aptorum Team

## Clinical Advisor



**DR. NISHANT AGRAWAL**  
*Senior Clinical Advisor - ATL*

- Professor of Surgery, School of Medicine, University of Chicago



**DR. HENRY CHAN LIK YUEN**  
*Senior Clinical Advisor - ATL*

- Associate Dean, Faculty of Medicine, CUHK



**DR. PHILIP W.Y. CHIU**  
*Senior Advisor - ATL*

- Professor, Department of Surgery, Institute of Digestive Disease, CUHK



**DR. VINCENT MOK CHUN TONG**  
*Senior Advisor - ATL*

- Head of Division of Neurology, Dept of Medicine & Therapeutics, CUHK

## Consultants, Advisors and Principal Investigators



**DR. KEITH CHAN**  
*Consultant*

- Adjunct professor and advisor at the Research Center for Drug Discovery, National Yang Ming University in Taipei;
- Former Division Director of Office of Generic Drugs, US FDA;
- Co-founder of Globomax LLC;
- Formerly employed at Ciba-Geigy



**DR. LAWRENCE BAUM**  
*Senior Scientific Advisor - VIL*

- Hon. Asso. Professor, Department of Psychiatry, The University of Hong Kong



**DR. FRANCIS SZELE**  
*Senior Scientific Advisor - VIL*

- Asso. Professor, Department of Physiology, Anatomy & Genetics, University of Oxford;
- Asst. Professor, Subventricular Zone, Northwestern University;
- Ph.D. in Biology, The University of Pennsylvania, U.S.



**MR. WILLIAM WEISS**  
*Consultant*

- Currently Director of Preclinical Service and Instructor of Pharmaceutical Sciences, College of Pharmacy, University of North Texas;
- 38 years of experience in drug discovery and development of antimicrobials including antibiotics, antivirals and antifungals;
- Former Director of Cumbre Pharmaceuticals Inc;
- Former Group Leader at Wyeth for 17 years;
- Formerly employed at Schering-Plough for 7 years;
- BSc in Microbiology from Rutgers University; MSc in Microbiology from Penn State University and Fairleigh Dickinson University



**DR. KIRA SHEINERMAN**  
*Senior Strategic Consultant*

- Co-Founder, CEO and Executive Director of Diamir Biosciences;
- Serves as a Managing Director, Healthcare Investment Banking at H.C. Wainwright & Co.;
- Ph.D. in Biomedical Sciences from Mount Sinai School of Medicine in New York
- Honors MBA from Zicklin School of Business, Baruch College, City University of New York



**PROFESSOR RICHARD KAO**  
*Principal Investigator*

- Asso. Professor, Department of Microbiology, The University of Hong Kong;
- Ph.D. in Microbiology, the University of British Columbia;
- The recipient of the Innovation Academy Award from the International Consortium of Prevention and Control of Infection (ICPIC) in Geneva, Switzerland in 2017

# Current progress of leading pipeline programs and discovery

→ Lead Projects → Other Candidates → Non-therapeutics Candidates

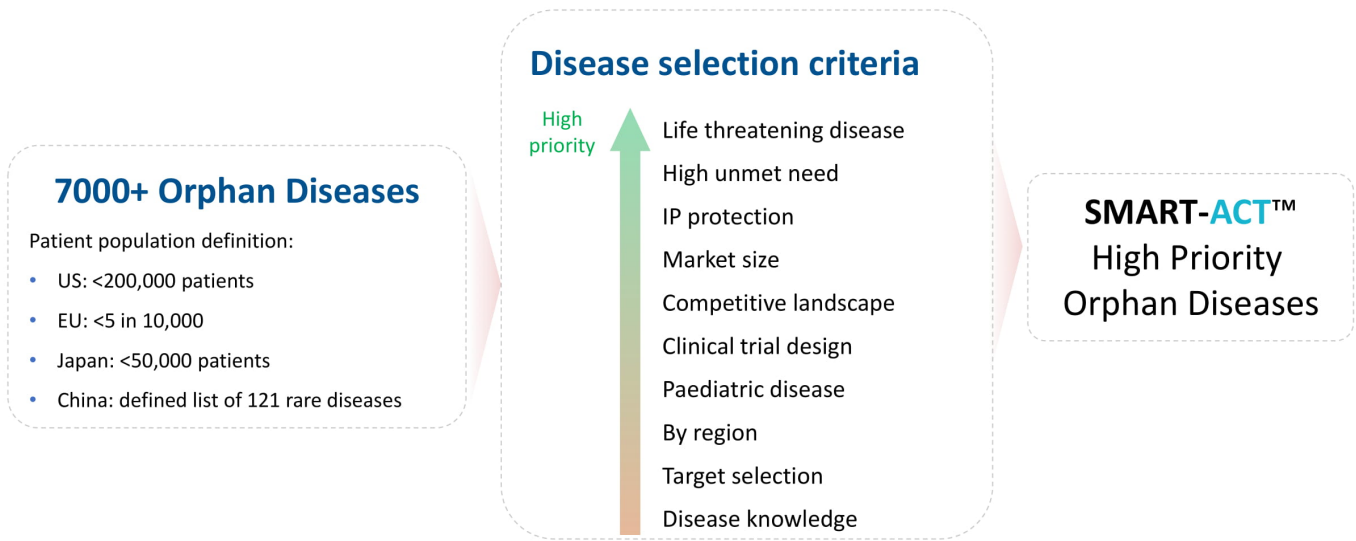
Projects	Candidate / Modality	Indication	Computational Discovery	In vitro validation	Existing Ph/III clinical safety data <sup>1</sup>	In vivo validation	IND preparation & submission	Ph/III w/ limited population <sup>2</sup>	
<b>SACT's Series</b>									
SACT-1	Repurposed Drug Molecule	Neuroblastoma	→						
SACT-2	Repurposed Drug Molecule	To be disclosed	→						
SACT-3	Repurposed Drug Molecule	To be disclosed	→						
SACT-COV19	Repurposed Drug Molecule	Coronavirus Disease 2019 (COVID-19)	→						

Projects	Candidate / Modality	Indication	Development Stage						
			Target Identification & Selection	Lead Discovery	Lead Optimization	IND-Enabling	Phase 1	Phase 2	Phase 3
<b>Acticle's Series</b>									
ALS-4	Small molecule	Treatment of bacterial infections caused by Staphylococcus aureus including MRSA	→						
ALS-1	Small molecule	Treatment of viral infections caused by Influenza virus A	→						
ALS-2	Small molecule	Treatment of bacterial infections caused by Staphylococcus aureus including MRSA	→						
ALS-3	Small molecule	Reviving existing antibiotics to overcome drug Resistance	→						
<b>Claves' Series</b>									
CLS-1	Macromolecule	Treatment of Obesity	→						
CLS-2	To be disclosed	To be disclosed	→						
CLS-3	To be disclosed	To be disclosed	→						

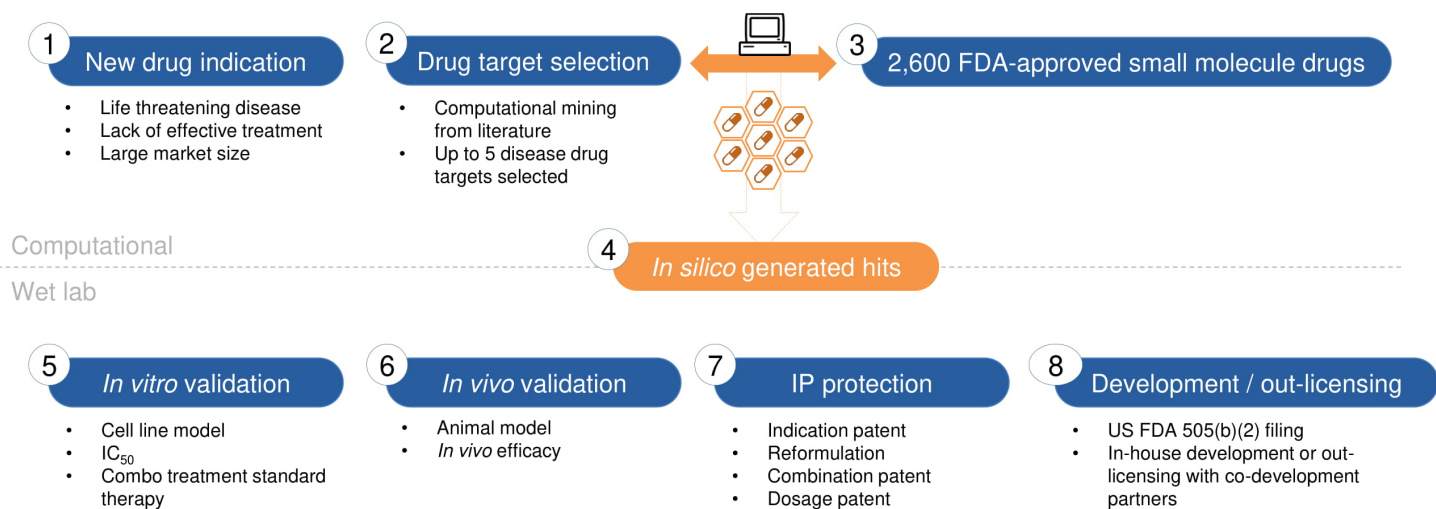
1. Refers to the drug's existing Phase I/II safety data previously conducted by a third party. Does not refer to clinical trials conducted by Aptorum 2. Subject to FDA's approval

## Current progress of leading pipeline programs and discovery

Projects	Candidate / Modality	Indication	Development Stage						
			Target Identification & Selection	Lead Discovery	Lead Optimization	IND-Enabling	Phase 1	Phase 2	Phase 3
<b>Nativus' Series</b>									
NLS-1	Small molecule	Treatment of Endometriosis							
<b>Scipio's Series</b>									
SPLS-1	83b-1 Novel Quinoline erivative	Treatment of Liver Cancer							
<b>Videns' Series</b>									
VLS-1	Curcumin-MNP (Medical Imaging Agent for MRI Diagnosis)	Diagnosis of Alzheimer's Disease							
VLS-2	MITA	Treatment of Alzheimer's & Parkinson's Disease							
VLS-4	Imaging Agent for MRI Diagnosis	Diagnosis of Alzheimer's Disease							
Projects	Modality	Target Customer	Formulation			Commercialisation			
NativusWell™ DOI (NLS-2)	Supplement	Women undergoing menopause							
<b>Signate's Series</b>									
Projects	Candidate / Modality	Indication	Development Stage						
			Lab-based Phantom Trial	Animal Trial	IDE Application Approval	Safety/ Feasibility Clinical Study	Pivotal Clinical Study	Process of obtaining PMA	
SLS-1	Robotic Catheter Platform for Intra-operative MRI-Guided Cardiac Catheterization	Heart Rhythm Disorders by Cardiac Electrophysiology Intervention							



# SMART-ACT™: Pipeline Workflow



# SACT-1 (neuroblastoma): market overview

SACT-1 targets neuroblastoma, a cancer that develops from nerve cells

## MARKET SIZE



## Prevalence

- ~700 cases of high risk neuroblastoma (NB) patients each year in the US<sup>3</sup> and we estimated EU has 1.5x those cases, c. 1050 high risk NB patients per year.
- Accounts for ~15% of all cancer-related deaths in the pediatric population<sup>4</sup>

## Orphan drug designation<sup>5</sup>

- Neuroblastoma is a rare disease and drugs are qualified for orphan designation by the FDA
- Designated orphan drugs receive 7 years of market exclusivity in US and 10 years of marketing exclusivity in EU
- Patents on reformulation, if granted, will provide up to 20 years of patent exclusivity from the application date in parallel to the market exclusivity

1. *Pediatr Rev.* 2018 Feb;39(2):57-67; 2. "Neuroblastoma Market Global Industry Perspective, Comprehensive Analysis, Size, Share, Growth, Trends, and Forecast 2019 – 2023" (2019). MRR Research. 3. *Curr Oncol Rep.* 2009 Nov;11(6):431-8 4. *Paediatr Drugs.* 2011 Aug 1;13(4):245-55 5. <https://www.fda.gov/about-fda/office-special-medical-programs/office-orphan-products-development>

All conclusory statements on this slide are based on Aptorum's internal tests/experimentation and has not yet been verified by clinical trials or third party testing.

# In vitro drug activity against neuroblastoma cell lines

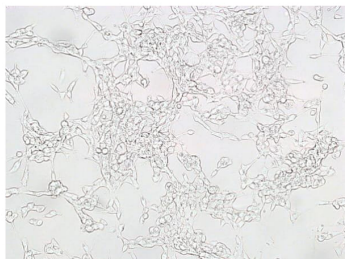
- SACT-1's potential action against neuroblastoma might be patentable
- We find that its action against neuroblastoma could be patentable

## Timeline

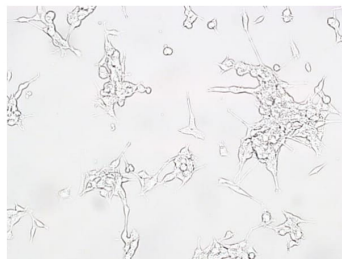
→ Lead Projects → Other Candidates → Non-therapeutics Candidates

Projects	Candidate / Modality	Indication	Computational Discovery	In vitro validation	Existing Ph/III clinical safety data	In vivo validation	IND preparation & submission	Ph/III w/ limited population
SACT's Series								
SACT-1	Repurposed Drug Molecule	Neuroblastoma		→				

Control treatment on neuroblastoma cells



SACT-1 treatment on neuroblastoma cells

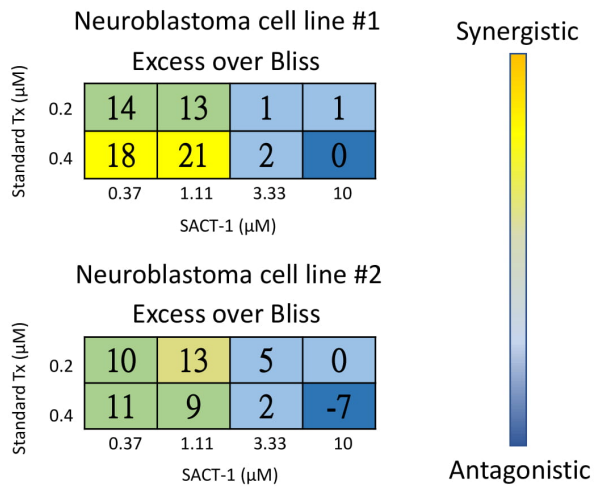
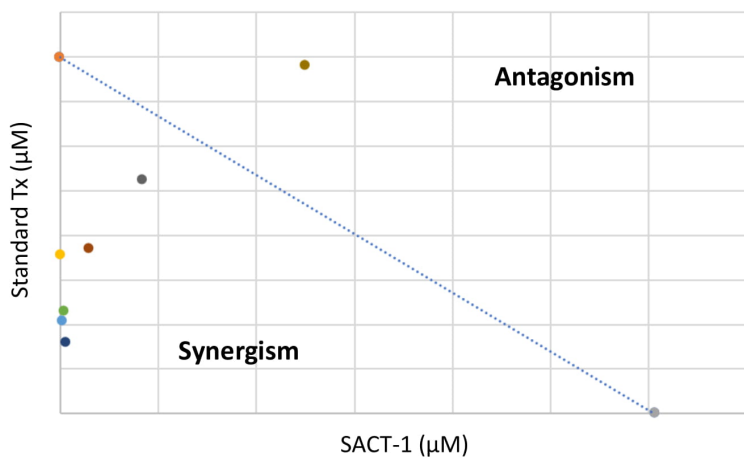


	IC <sub>50</sub> [μM] For IMR-32	IC <sub>50</sub> [μM] For SK-N-BE(2)	IC <sub>50</sub> [μM] For SK-N-SH	IC <sub>50</sub> [μM] For SH-SY5Y
SACT-1	2.97	3.37	2.75	3.12

The results shown in this slide are based on Aptorum's internal (in vitro/in vivo) tests/experiments that have not been verified in clinical trials and/or third party testing.

# Synergistic effect of SACT-1 in combination with standard treatment

- Synergistic effect observed for SACT-1 in combination with standard treatment in 2 different neuroblastoma cell lines, as seen in the isobologram (left) and the Excess over Bliss (right)



The results shown in this slide are based on Aptorum's internal (in vitro/in vivo) tests/experiments that have not been verified in clinical trials and/or third party testing



## SACT-1: safety & tolerability

### Well-established Safety profile based on a FDA approved product

- Did not show genotoxic potential even at the highest feasible concentration dose (*in vitro* and *in vivo*)
- In a phase IIb study over 2 years, all SACT-1 doses were safe and well tolerated
- No dose relationship between RPV and adverse events (AE)

RPV	25mg/day (N=93)	75mg/day (N=95)	150mg/day (N=91)
Median treatment duration, weeks	101	100	100
Adverse events (AE)			
Any grade 2-4 AE at least possibly related to RPV	20%	20%	21%
AEs leading to discontinuation	9%	12%	14%
Any serious AE	13%	14%	10%
Deaths	0%	2%	0%

### FDA approved pharmacokinetics profile

- Data package can be potentially accepted by the FDA in our 505(b)(2) new drug application
- Relatively long half-life ( $t_{1/2} = 43-55h$ ). Frequent dosing may not be required

Ref: doi: 10.1089/AID.2011.0050

## Executive summary: Acticle projects

### ALS-4

- Aptorum's lead program ALS-4 is an anti-virulent, non-bactericidal drug candidate for *Staphylococcus aureus* infections including MRSA<sup>1</sup>
- Unlike all major treatments on the market, ALS-4 is an orally administered anti-virulent molecule using a non-bactericidal approach<sup>1</sup>, potentially reducing significant risks of developing *S. aureus* resistance
- Targeting IND submission by H2 2020
- Upon IND approval, a Phase I clinical study to commence in H2 2020 in North America
- Targeting to submit written request for approval under the newly established LPAD regulatory pathway (Limited Population Pathway for Antibacterial and Antifungal Drugs), to expedite marketing approval and commercialization

### ALS-1

- A unique antiviral therapeutic against Influenza A that has a more upstream target than Tamiflu<sup>®</sup> which is shown to be more effective in vitro<sup>1</sup>
- Viral resistance to Tamiflu and other neuraminidase inhibitors has risen rapidly in recent years<sup>2</sup>
- ALS-1 has a distinct mechanism of action compared with Tamiflu<sup>®</sup> and Xofluza<sup>TM1,3</sup>

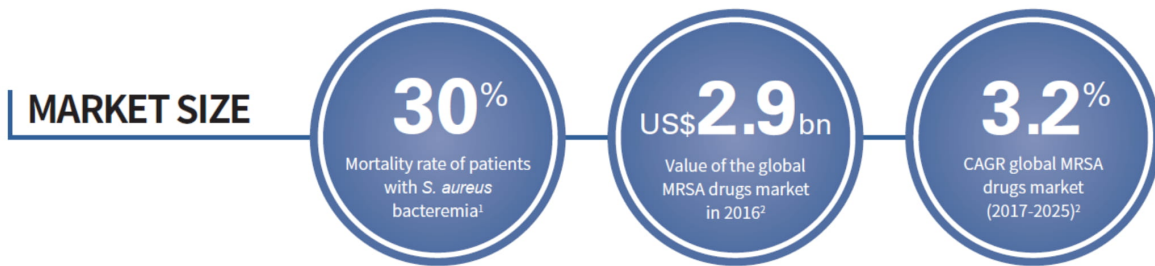
### ALS-2/ALS-3

- Additional novel anti-virulent, non-bactericidal approach therapeutics targeting Gram-positive bacteria<sup>1</sup>
- In discovery/lead optimization stage and generating good traction towards doing IND-enabling studies<sup>1</sup>

1. Based on Aptorum's internal tests/experimentation and has not yet been verified by clinical trials or third party testing; 2. Influenza Antiviral Medications: Summary for Clinicians. CDC. <https://www.cdc.gov/flu/professionals/antivirals/summary-clinicians.htm>; 3. Nat Biotechnol. 2010 Jun;28(6):600-5

## ALS-4: Market Overview

ALS-4 is an anti-virulent, non-bactericidal drug candidate for *Staphylococcus aureus* infections including MRSA



### Third-party infectious disease drugs or company-related mergers and acquisitions

- In 2014, Merck's acquisition of Cubist Pharmaceuticals, a large developer of antibiotics, for USD 8.4bn<sup>3</sup>
- In 2018, Roivant's licensing of Intron's Phase II asset for USD 667.5m in upfront and milestone payments<sup>4</sup>

1. Clin Microbiol Rev. 2012 Apr;25(2):362-86; 2. "Methicillin-resistant Staphylococcus Aureus (MRSA) Drugs Market - Global Industry Analysis, Size, Share, Growth, Trends, and Forecast, 2017-2025" (2018). Transparency market research; 3. <https://dealbook.nytimes.com/2014/12/08/merck-agrees-to-acquire-drug-maker-cubist-for-9-5-billion/>; 4. <https://www.prnewswire.com/news-releases/roivant-sciences-and-intron-bio-sign-licensing-deal-for-novel-anti-superbugs-biologic-sal200-300753307.html>

# Market Approved Drugs for MRSA Infections

## Frequently prescribed antibiotics for MRSA infections<sup>1</sup>

Product (Company)	Antibiotic Class	Indication(s)	RoA	Dose	Cost of Treatment (duration)	Notes
Vancomycin (Generic)	Glycopeptide	Severe infections caused by MRSA	IV / oral*	2g/day	USD 101-144 (7-10 days)	<ul style="list-style-type: none"> <li>Currently, the most frequently prescribed antibiotic for MRSA suspected infections<sup>1,2</sup></li> <li>In clinical use for &gt;60 years<sup>3</sup>, vancomycin-resistant <i>S. aureus</i> (VRSA) was first discovered in 2002<sup>4</sup></li> </ul>
Daptomycin (Merck)	Lipopeptide	ABSSSI, <i>S. aureus</i> bacteremia	IV	4-6mg/kg/day	USD 6,736-23,710 <sup>5</sup> (14-42 days)	<ul style="list-style-type: none"> <li>In clinical use since 2003<sup>6</sup></li> <li>Daptomycin resistance described in <i>S. aureus</i> as early as 2006<sup>7</sup></li> </ul>
Linezolid (Pfizer)	Oxazolidinone	ABSSSI, CABP, HABP, uSSSI	IV / oral	0.8-1.2g/day	IV: USD 1,920-5,376 Oral: USD 2,978-11,429 (10-14 days)	<ul style="list-style-type: none"> <li>In clinical use since 2003<sup>8</sup>. Entirely synthetic, not expected to develop clinical resistance<sup>9</sup>, however</li> <li>Linezolid resistance encountered clinically since 2010<sup>9</sup></li> </ul>
Ceftaroline fosamil (Actavis)	Cephalosporin	ABSSSI, CABP	IV	1.2g/day	USD 1,831-5,127 (5-14 days)	<ul style="list-style-type: none"> <li>In clinical use since 2010<sup>10</sup></li> <li>Ceftaroline resistance encountered clinically since 2016<sup>11</sup></li> </ul>
Tigecycline (Pfizer)	Glycycycline	ABSSSI, CABP, CIAI	IV	0.1-0.2mg/day	USD 1,888-4,977 (5-14 days)	<ul style="list-style-type: none"> <li>In clinical use since 2005<sup>12</sup></li> <li>Tigecycline resistance encountered clinically in developing countries since 2017<sup>13,14</sup></li> </ul>
Telavancin (Theravance Biopharma)	Lipoglycopeptide	ABSSSI, HABP, VABP	IV	10mg/kg/day	USD 3,002-10,568 (7-21 days)	<ul style="list-style-type: none"> <li>In clinical use since 2009<sup>15</sup></li> <li>Vancomycin resistance leads to a 4-8x increase in telavancin MIC (minimum inhibitory concentration)<sup>16</sup></li> </ul>

ABSSSI: acute bacterial skin and skin structure infection; CABP: community-acquired bacterial pneumonia; HABP: hospital-acquired bacterial pneumonia; CIAI: complicated intra-abdominal infection; VABP: ventilator-associated bacterial pneumonia; \* Only for intestinal infections; 1. Reproduced from "Companies Take Aim at MRSA Infections" P T, 2016 Feb; 41(2): 126-128; 2. Clin Infect Dis. 2011 Feb 1;52(3):e18-55; 3. Clin Infect Dis. 2006 Jan 1;42 Suppl 1:S5-12; 4. Centers for Disease Control and Prevention. [https://www.cdc.gov/hai/settings/lab/vrsa\\_lab\\_search\\_containment.html](https://www.cdc.gov/hai/settings/lab/vrsa_lab_search_containment.html); 5. Cost of treatment of Daptomycin for *S. aureus* bacteremia at a dosage of 6mg/kg; 6. FDA. [https://www.accessdata.fda.gov/drugsatfda\\_docs/nda/2003/21-572\\_Cubicin.cfm](https://www.accessdata.fda.gov/drugsatfda_docs/nda/2003/21-572_Cubicin.cfm); 7. Int J Antimicrob Agents. 2006 Oct;28(4):280-7; 8. FDA. [https://www.accessdata.fda.gov/drugsatfda\\_docs/nda/2002/21-1308003\\_211315003\\_211325003\\_ZyvoTOC.cfm](https://www.accessdata.fda.gov/drugsatfda_docs/nda/2002/21-1308003_211315003_211325003_ZyvoTOC.cfm); 9. Pharmaceuticals (Basel). 2010 Jul; 3(7):1988-2006; 10. FDA. [https://www.accessdata.fda.gov/drugsatfda\\_docs/nda/2010/200327orig1s000toc.cfm](https://www.accessdata.fda.gov/drugsatfda_docs/nda/2010/200327orig1s000toc.cfm); 11. J Antimicrob Chemother. 2016 Jun; 71(6):1736-1738; 12. FDA. [https://www.accessdata.fda.gov/drugsatfda\\_docs/nda/2005/21-821\\_Tygaicil.cfm](https://www.accessdata.fda.gov/drugsatfda_docs/nda/2005/21-821_Tygaicil.cfm); 13. New Microbes New Infect. 2017 Sep; 19: 8-12; 14. Journal of Microbiology and Infectious Diseases 2017; 7 (4):173-177; 15. FDA. [https://www.accessdata.fda.gov/drugsatfda\\_docs/nda/2009/022110s000toc.cfm](https://www.accessdata.fda.gov/drugsatfda_docs/nda/2009/022110s000toc.cfm); 16. Clin Infect Dis. 2015 Sep 15;61 Suppl 2:S58-68.

# ALS-4: Addressing the Shortfall of Vancomycin

## Vancomycin

- Generic antibiotic that is the most frequently prescribed for MRSA-suspected infections<sup>1,2</sup>
- After >60 years<sup>3</sup> of clinical use, its use against *S. aureus* is becoming limited. Vancomycin has been shown to have slow bactericidal activity, poor anti-staphylococcal activity, poor tissue penetration, and high rates of infection relapse<sup>4,5,6,7,8,9</sup>
- The shortcomings of Vancomycin has been compounded since the discovery of vancomycin-resistant *S. aureus* (VRSA) in 2002<sup>10</sup>
- Vancomycin is not orally bioavailable and must be administered intravenously in order to treat systemic infections<sup>11,12</sup>. Oral vancomycin is only effective for treating local intestinal infections<sup>13</sup>. Therefore, for MRSA-suspected infections oral vancomycin is only indicated for the treatment of pseudomembranous colitis<sup>14</sup>

## ALS-4: Stand Alone or as Combination Therapy with Antibiotics (e.g. Vancomycin)

- As a combination therapy believed to overcome the shortcomings of vancomycin.<sup>15</sup>
- ALS-4 can potentially complement other bactericidal antibiotics as well, therefore ALS-4 is not a direct competitor of antibiotics
- Synergistic effects of other drugs with vancomycin against MRSA has been demonstrated previously with  $\beta$ -lactam antibiotics and vancomycin<sup>16</sup>

1<sup>st</sup> place, Innovation Academy Category, ICPIC 2017



- Awarded to the Company's Hong Kong team, led by Dr. Richard KAO
- For the revolutionary concept of applying chemical genetics to tackle MRSA infection, which forms the scientific basis of ALS-2, ALS-3 and ALS-4

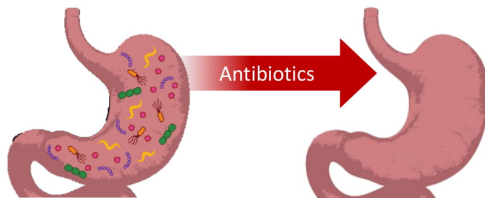
1. "Companies Take Aim at MRSA Infections" P T. 2016 Feb; 41(2): 126-128; 2. Clin Infect Dis. 2011 Feb 1;52(3):e18-55; 3. Clin Infect Dis. 2006 Jan 1;42 Suppl 1:S5-12; 4. Antimicrob Agents Chemother. 2008 Jan;52(1):192-7; 5. Clin Infect Dis. 2007 Jan 15;44(2):190-6; 6. Clin Infect Dis. 2007 Sep 1;45(5):601-8; 7. J Clin Microbiol. 2011 Oct;49(10):3669-72; 8. Clin Infect Dis. 2007 Sep 15;45 Suppl 3:S191-5; 9. J Clin Microbiol. 2004 Jun;42(6):2398-402; 10. Centers for Disease Control and Prevention. [https://www.cdc.gov/hai/settings/lab/vrsa\\_lab\\_search\\_containment.html](https://www.cdc.gov/hai/settings/lab/vrsa_lab_search_containment.html); 11. J Infect. 2018 Dec;77(6):489-495; 12. StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019-2018 Nov 18; 13. HealthJade, <https://healthjade.net/vancomycin/>; 14. Medscape, <https://reference.medscape.com/drug/firvanq-vancocin-vancomycin-342573>; 15. Combination Antibiotic Treatment of Serious Methicillin-Resistant Staphylococcus aureus Infections, <https://www.thieme-connect.com/products/ejournals/pdf/10.1055/s-0034-1396906.pdf>; 16. J Clin Microbiol. 2016 Mar; 54(3): 565-568

# ALS-4: Value Proposition

## Antibiotic

- Antibiotic resistance in *S. aureus* has been discovered in most prescribed antibiotics for MRSA<sup>1</sup>
- Broad spectrum and indiscriminate<sup>2</sup>
- Commonly affect normal flora, may lead to superinfection in case of drug resistance<sup>3</sup>

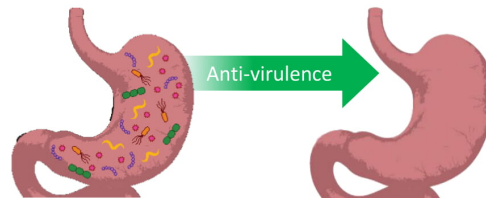
### Indiscriminate clearance



## Anti-virulence (ALS-4)

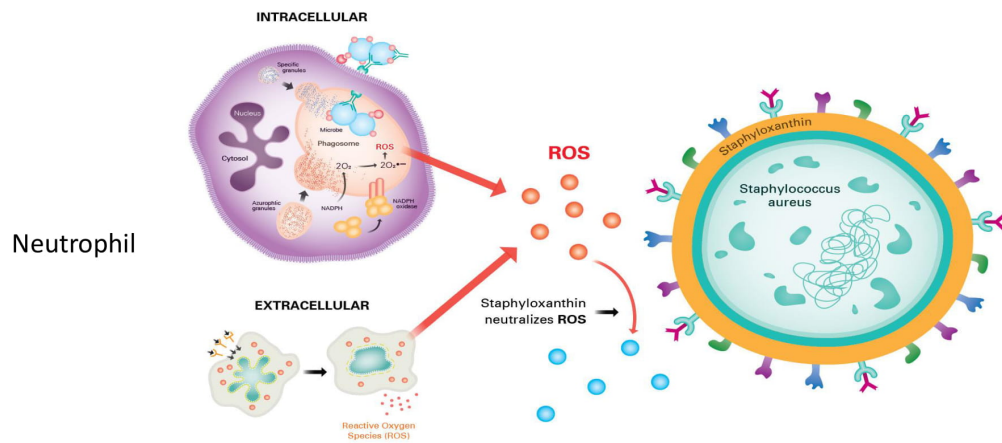
- ✓ Not bactericidal, potentially less selective pressure and much less likely for bacteria to develop resistance<sup>4,5</sup>
- ✓ "Disarms" the bacteria by reducing pathogenicity<sup>4,5,6</sup>
- ✓ Bacterial clearing is mediated by host immunity<sup>4,5</sup>

### Directed against pathogen



1. Refer to "ALS-4: Approved Drugs for MRSA Infections" for complete set of sources; 2. P T. 2016 Feb; 41(2): 126-128; 3. J Infect Dis. 2018 Jan 30;217(4):628-636; 4. Based on Aptorum's internal tests/experimentation and has not yet been verified by clinical trials or third party testing; 5. MBio. 2017 Sep 5;8(5). pii: e01224-17; 6. J Exp Med. 2005 Jul 18;202(2):209-15.

# Mechanism of Action: Staphyloxanthin of *Staphylococcus aureus*

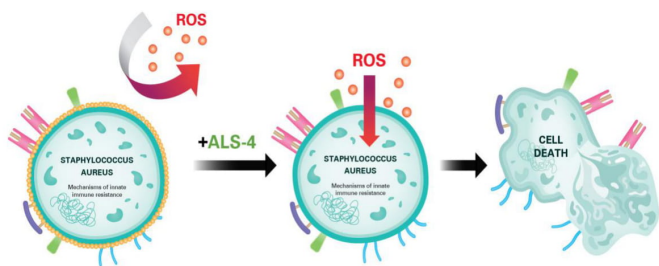


The above diagram summarizes the mechanism of action by Staphyloxanthin of *Staphylococcus aureus*:

- Neutrophils kill bacteria including *Staphylococcus aureus* intracellularly or extracellularly via Reactive Oxygen Species “ROS-oxygen radicals released by neutrophils trigger the subsequent bacterial damage processes”<sup>1</sup>.
- To counteract, Staphyloxanthin protects the bacteria by serving as an anti-oxidant to neutralize the ROS secreted by neutrophils<sup>2</sup>.

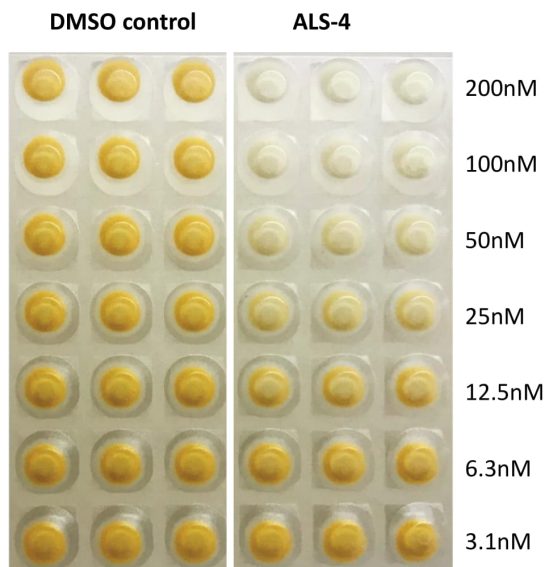
<sup>1</sup>Annu Rev Immunol. 2005;23:197-223; <sup>2</sup>mBio. 2017 Sep 5;8(5). pii: e01224-17

# Mechanism of Action-ALS-4 on Staphyloxanthin Synthesis



The above diagram summarizes our findings about how ALS-4 inhibits Staphyloxanthin synthesis:

- ALS-4 inhibits a key enzyme in the biosynthesis of Staphyloxanthin with an  $IC_{50} = 20nM$ .
- In the absence of Staphyloxanthin, the bacteria become susceptible to damage by ROS, triggering the usual series of mechanisms by neutrophils that ultimately leads bacterial cell death.



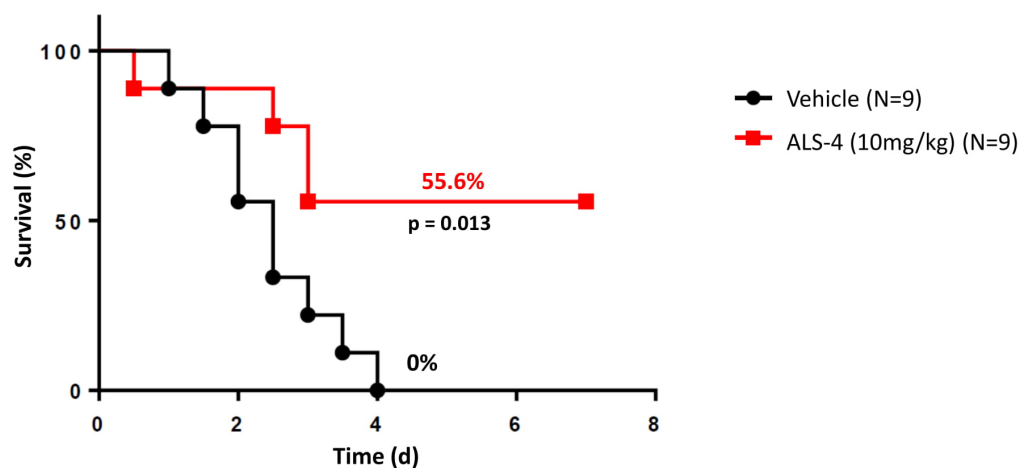
The description of ALS-4 and related conclusory statements on ALS-4 on this slide are based on Aptorum's internal tests/experimentation and has not yet been verified by clinical trials or third party testing.



## ALS-4: oral formulation treatment in an MRSA survival study

The combination of ALS-4's anti-virulence properties together with host immune system, efficacy is still superior.

The below in-vivo data includes rats infected with a lethal dose of MRSA USA300 in a bacteremia model.

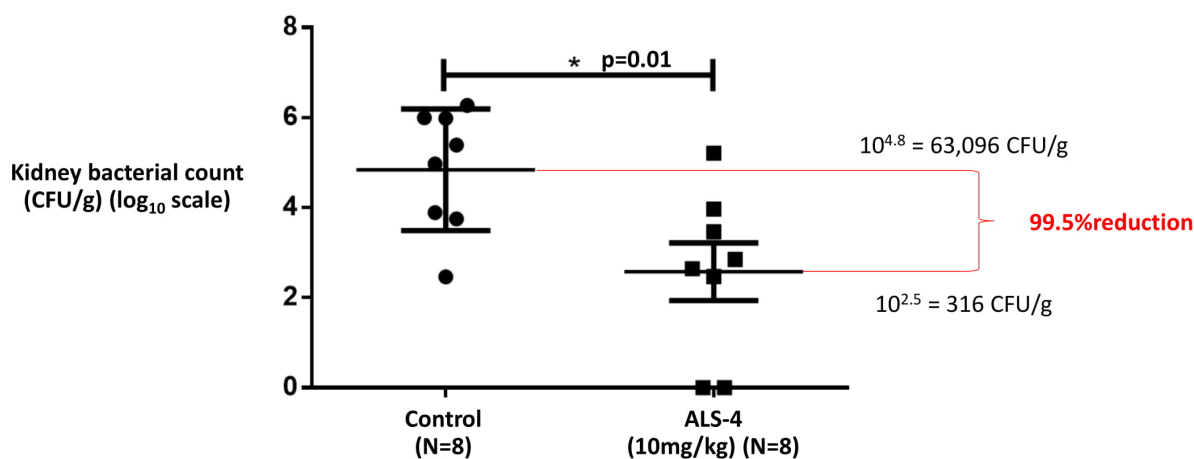


- A lethal dose ( $10^9$  CFU) of MRSA was introduced through the tail vein
- ALS-4 was administered **orally** 30 minutes after infection for twice a day thereafter

The results shown in this slide are based on Aptorum's internal (in vitro/in vivo) tests/experiments that have not been verified in clinical trials and/or third party testing

## ALS-4: oral formulation treatment in a non-lethal bacteremia model

ALS-4 is shown to greatly reduce organ bacterial count in a bacteremia animal model



- Rats were challenged with a non-lethal dose ( $10^7$  CFU) of MRSA through the tail vein
- In order to simulate a more realistic clinical scenario, treatment was introduced 14-days after infection, where ALS-4 was administered orally twice a day at 10mg/kg per animal

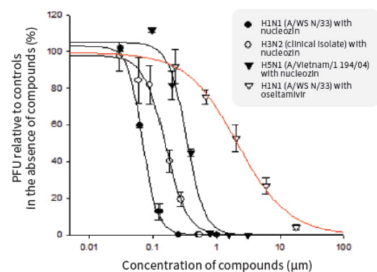
The results shown in this slide are based on Aptorum's internal (in vitro/in vivo) tests/experiments that have not been verified in clinical trials and/or third party testing

# ALS-1: Targeting a Novel Druggable Target for Influenza A

## ALS-1 inhibits influenza A nucleoprotein (NP)

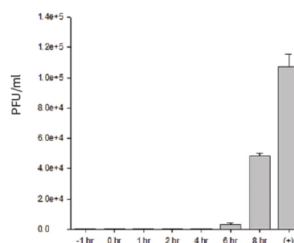
- NP is the most abundantly expressed protein during the course of an infection<sup>1</sup>. Its primary function is to encapsidate the virus genome for RNA transcription, replication and packaging. It is also a key adapter molecule between virus and host processes<sup>1</sup>
- ALS-1, by targeting NPs, acts upstream of Neuraminidase inhibitors such as Tamiflu, which target the last stage (budding) of the viral life cycle<sup>2</sup>. This novel mechanism distinguishes ALS-1 from all other currently marketed antiviral drugs<sup>3</sup>

### ALS-1 outperforms Tamiflu® (oseltamivir, in red) in vitro with a lower IC<sub>50</sub><sup>2</sup>



This figure shows the concentration dependence of ALS-1 in reducing the plaque-forming unit (pfu, a measure of number of infectious virus particulates) of human H1N1, H3N2 and H5N1 influenza viruses. The IC<sub>50</sub> for these viruses is between 0.1-1µM

### ALS-1 inhibited viral growth up to 6 hours after infection, indicating antiviral activities reside on post-entry and post-nuclear events<sup>2</sup>



This figure shows that MDCK cells were infected and ALS-1 (1 µM) was added before infection (- 1 h), at the time of infection (0 h) and at 1, 2, 4, 6 and 8 hour after infection as indicated. (+) control without ALS-1

1. J Gen Virol. 2002 Apr;83(Pt 4):723-34; 2. Nat Biotechnol. 2010 Jun;28(6):600-5; 3. Refer to the next slide "ALS-1: A Unique Antiviral Therapeutic Against Influenza A".

# Claves Executive Summary

## Human Microbiota

- We live in constant symbiosis with our gut bacteria, and dysbiosis can be the cause to numerous diseases<sup>1</sup>

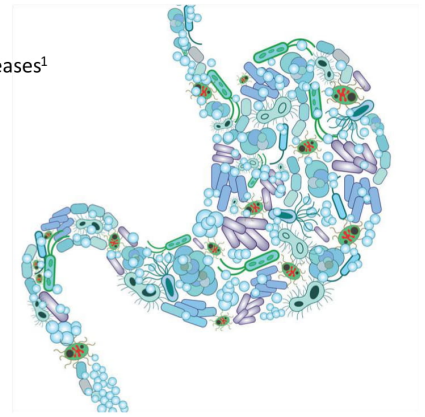
## Claves Technology

- The Claves technology is designed to physically modulate the chemical signaling of diseases-causing microbiota<sup>2</sup>
- Highly scalable large molecule technology with over 70 potential therapeutic targets possible for development<sup>2</sup>
- Claves therapeutics bind target chemicals with high affinity and specificity, they are non-absorbable and expected to be free from any systemic toxicity<sup>2,3</sup>
- Multiple candidates under development for various indications<sup>2</sup>

## CLS-1: Lead Program Targeting Obesity

- CLS-1 is the lead program in the Claves projects, intended to target metabolites secreted by the microbiota linked to obesity<sup>2</sup>
- CLS-1 is also shown to modulate gut microbiota population linked to obesity<sup>2,3</sup>
- CLS-1 achieves significant weight loss in a mouse model without affecting the gut mucosa, inflammation, and the functions of the liver and kidneys<sup>2,3</sup>
- Non-absorbable nature of the Claves therapeutics may expedite traditional toxicological studies<sup>2</sup>

1. Lancet. 2003 Feb 8;361(9356):512-9; 2. Based on Aptorum's internal tests/experimentation and has not yet been verified by clinical trials or third party testing; 3. Data available in this presentation; 4. Current understanding of dysbiosis in disease in human and animal models, <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4838534/>



- Contains **100s of species of microbes**
- Constantly producing **1000s of active metabolites**
- Some metabolites provides immunological and metabolic benefits
- **Dysbiosis (microbial imbalance) is a significant factor in disease<sup>4</sup>**

# Claves Platform and CLS-1: Value Proposition

## CLS-1

- Identified specific microbiota metabolite linked to obesity
- Novel therapeutic that physically modulates microbiota metabolite
- Acts locally in the gut with high affinity and specificity
- Non-absorbable and is expected to be free from any systemic toxicity
- Significant weight loss in an animal study

## Claves Platform

- Novel platform technology that can be customized to bind a wide variety of microbiota metabolites with high affinity and specificity
- Sustainable pipeline of drug candidates for treatment of multiple indications (see next page)

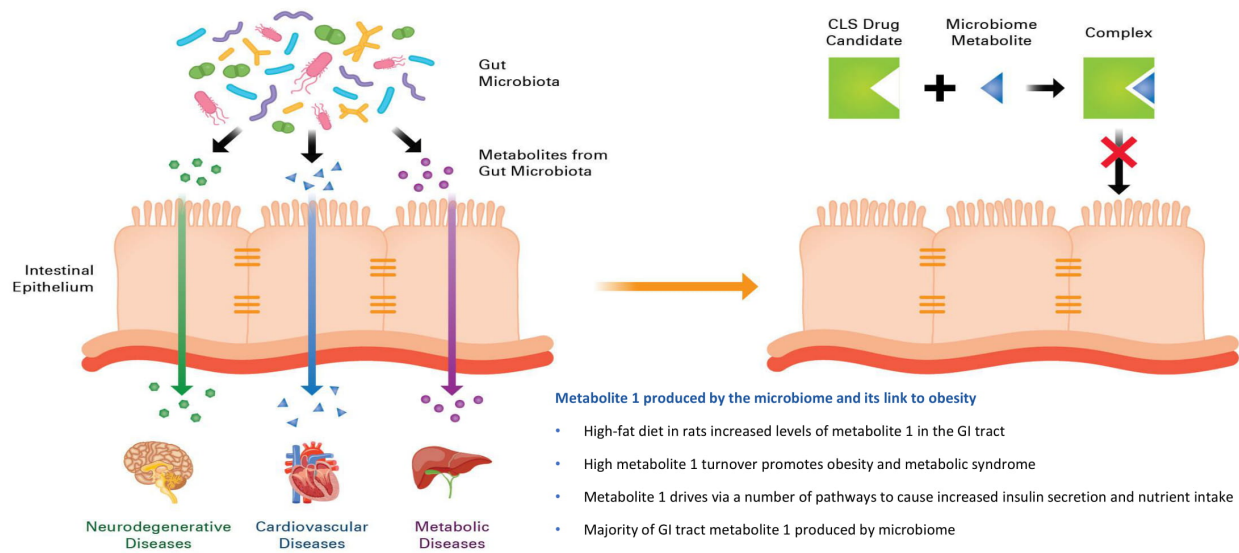


## POSSIBLE INDICATIONS

	SYSTEMIC DISEASES	DIGESTIVE DISEASES
	Obesity	C. difficile infection
	Diabetes	Colorectal cancer
	Fatty liver	Inflammatory bowel disease
	Cardiovascular diseases	Irritable bowel syndrome
	Renal failure	
	Depression	
	Parkinsonism	
	Autistic spectrum disorder	

All conclusory statements on this slide are based on Aptorum's internal tests/experimentation and has not yet been verified by clinical trials or third party testing.

# Mechanism of Action



## Metabolite 1 produced by the microbiome and its link to obesity

- High-fat diet in rats increased levels of metabolite 1 in the GI tract
- High metabolite 1 turnover promotes obesity and metabolic syndrome
- Metabolite 1 drives via a number of pathways to cause increased insulin secretion and nutrient intake
- Majority of GI tract metabolite 1 produced by microbiome
- **CLS-1 specifically binds to and removes metabolite 1 from the body**
- CLS-1 is a macromolecule that, due to its particle size (over 20um), cannot be absorbed into the systemic circulation (both on its own and in combination with the metabolite 1)
- The combined form passes out through the digestive system and therefore, removing metabolite 1 from the gut.

All conclusory statements on this slide are based on Aptorum's internal tests/experimentation and has not yet been verified by clinical trials or third party testing.

# CLS-1: Market Overview

CLS-1: the lead program in the Claves projects, targeting obesity



## Recent Deals in Obesity Treatment

- Boehringer Ingelheim committed up to USD 300m to work with Gubra on obesity treatments

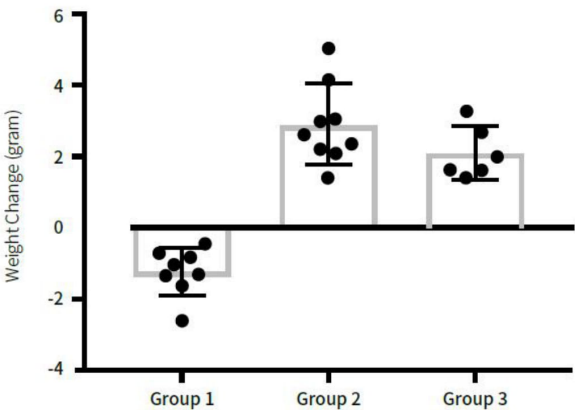
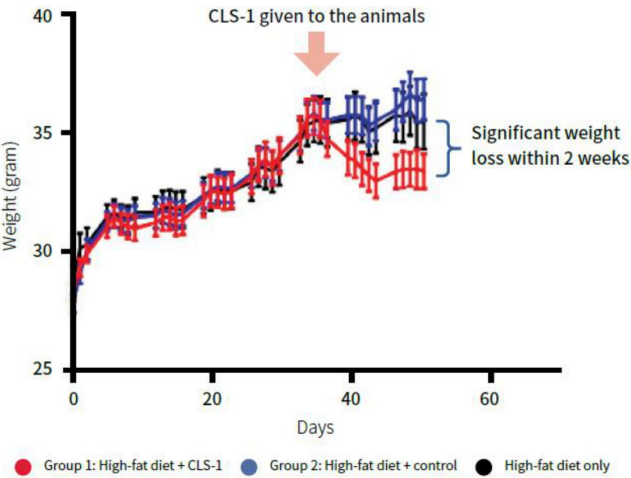
## Competing Drugs

- CLS-1 is a drug candidate for obesity treatment that achieves its effect by modulating the chemical signaling of gut microbiota. There are no obesity treatment drugs on the market using similar mechanism<sup>3</sup>.

1. World Health Organization. Obesity and overweight fact sheet. <https://www.who.int/news-room/fact-sheets/detail/obesity-and-overweight>; 2. "Obesity Treatment Market To Reach USD 19.90 Billion By 2026 " (2019). Reports And Data. <https://www.globenewswire.com/news-release/2019/06/06/1865530/0/en/Obesity-Treatment-Market-To-Reach-USD-19-90-Billion-By-2026-Reports-And-Data.html>; 3. To the extent of our knowledge at the time of writing

# CLS-1: Efficacy in a Mouse Model

CLS-1 treatment significantly reduces body weight in mice



The above data are based on Aptorum's internal tests and has not yet been verified by clinical trials or third party testing.



# NLS-2: Executive Summary

## NLS-2<sup>1</sup>

- NLS-2 is a dietary supplement for the relief of menopausal symptoms.
- The bioactive component of NLS-2 is DOI, a novel non-hormonal compound extracted from Chinese Yam
- DOI significantly increased estradiol biosynthesis and aromatase expression in granulosa cells *in vitro* and *in vivo* (rat animal model)
- Osteoporosis is frequently associated with menopause. DOI increases the apparent bone mineral density, bone volume fraction and trabecular thickness in an *in vivo* rat model
- DOI acts in a tissue-specific manner. Upregulation of aromatase, an enzyme involved in the production of estrogen, by DOI was found in ovary but not in other tissue
- DOI does not cause toxicity *in vitro* based on cell viability in the MTT assay
- Targeting to launch as a dietary supplement in H2 2020



## Timeline<sup>2</sup>

→ Lead Projects → Other Candidates → Projected timeline

Projects	Modality	Target Customer	Formulation	Commercialisation
NativusWell™ DOI (NLS-2)	Supplement	Women undergoing menopause	Targeted to launch in HK, UK, Europe in 2020 (registration ongoing)	

1. Lancet. 2003 Feb 8;361(9356):512-9; 2. Data available in this presentation

All conclusory statements on this slide are based on Aptorum's internal tests/experimentation and has not yet been verified by clinical trials or third party testing.

The background of the page is a repeating pattern of blue, semi-transparent molecular models. These models consist of spheres of varying sizes connected by thin rods, representing chemical structures. The spheres are rendered with a glass-like effect, showing highlights and shadows. The overall color palette is a light, airy blue.

# APPENDIX

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## Income Statement Summary (U.S. GAAP)<sup>1</sup>

	Year ended December 31, 2019	Year ended December 31, 2018
	US\$	US\$
Revenue	535,166	383,450
Research and development expenses	(6,939,051)	(3,101,432)
General and administrative fees	(7,373,425)	(4,919,626)
Legal and professional fees	(3,405,705)	(1,811,770)
Net loss attributable to Aptorum Group Limited	(18,686,762)	(14,831,723)
Net loss per share – basic and diluted	(0.64)	(0.53)
Interest (expense) income, net <sup>2</sup>	(3,699,672)	(4,458,191)
Depreciation and amortization	(1,299,618)	(682,902)
Share based compensation expenses	(1,612,832)	-

Notes:

1. The following slide contains selected information for the Company's income statement. Please see the Company's most recently filed Form 20-F for the Company's complete financial statements.

2. During the year ended December 31, 2019 and 2018, the net interest expenses included USD 3.1 M and USD 2.4 M, respectively, amortization of beneficial conversion feature which are non-cash items.

## Selected Balance Sheet Items (U.S. GAAP)<sup>1</sup>

	December 31, 2019	December 31, 2018
	US\$	US\$
Cash, restricted cash and marketable securities	6,356,284	27,121,576
<b>Total current assets</b>	<b>8,032,881</b>	<b>28,722,941</b>
Property, plant and equipment, net	7,093,035	4,260,602
<b>Total assets</b>	<b>23,954,218</b>	<b>45,074,640</b>
Convertible debts	-	(10,107,306)
Warrant liabilities	-	(753,118)
<b>Total current liabilities</b>	<b>(2,674,675)</b>	<b>(12,184,865)</b>
<b>Total liabilities</b>	<b>(9,102,466)</b>	<b>(12,328,738)</b>
<b>Total equity attributable to the shareholders of Aptorum Group Limited</b>	<b>16,361,208</b>	<b>33,114,435</b>
<b>Working Capital<sup>2,3</sup></b>	<b>5,358,206</b>	<b>16,538,076</b>

Notes:

1. The following slide contains selected information for the Company's balance sheets. Please see the Company's most recently filed form 20-F for the Company's complete financial statements.

2. Current assets less current liabilities.

3. Since September 2019, Aptorum Group has access to over USD15m in working capital from shareholder support



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