

# IMUGENE

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November 2019



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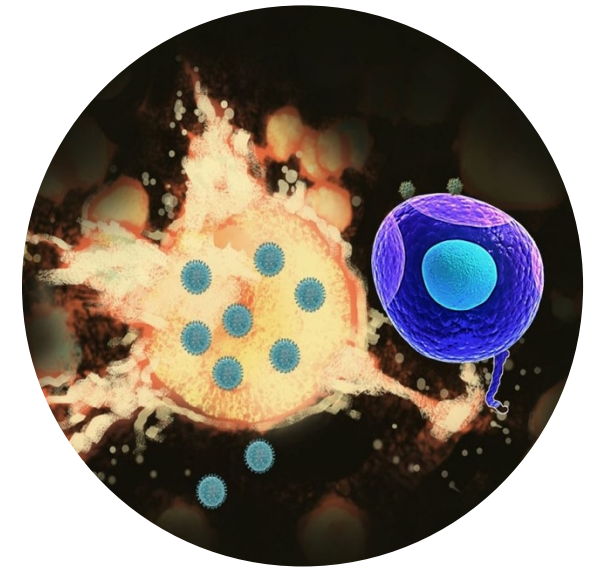
# INTRODUCTION TO IMUGENE

- Imugene is a biotech company headquartered in Australia and publicly traded on the Australian Securities Exchange (ASX:IMU)
- Technology originated from the Medical University of Vienna, invented by Prof Christoph Zielinski and Prof Ursula Wiedermann
- Late 2013, Paul Hopper built Imugene around this technology
- 2017: HER-Vaxx, our HER2 targeted B-Cell Immunotherapy, entered the clinic
- 2018: Licensed extensive B cell portfolio from OSU and Mayo Clinic comprising of HER1, HER2, HER3, VEGF, IGF-1R, CD28, combinations thereof and the PD-1 B-Cell Immunotherapy
- July, 2019: Licensed a prolific oncolytic virus from City of Hope



# INVESTMENT HIGHLIGHTS

- Novel technology in one of the most sought-after areas of cancer immunotherapy today: oncolytic viruses a.k.a. cancer killing viruses that stimulate immune recognition of cancer and our B-cell Immunotherapy pipeline
- CF33 poised to enter into 2 x Phase 1 clinical trials in 2020; PD1-Vaxx to enter into phase 1 study in 2020
- Robust intellectual property: long patent life & composition of matter for CF33 to 2037 and HER-Vaxx to 2036
- Highly experienced immunotherapy developers including the oncolytic virus team (all ex-viralytics)
- Potential applications across many cancers, including combination with CTLA4/PD-1/PD-L1 checkpoint inhibitors or with engineered immune cells
- Outstanding scientific provenance across all our technology: B-cell immunotherapies from Medical University of Vienna and OSU & CF33 from one of the US leading cancer centres, City of Hope in Los Angeles with Inventor, Professor Yuman Fong who is an internationally recognized oncolytic virus and cancer expert
- Attractive license terms - worldwide exclusive rights to the technology



# International leadership team with extensive commercialisation expertise in the sector



**Leslie Chong**  
SYDNEY, AU  
Managing Director & CEO

- 21+ years of oncology experience across Phase I – III clinical development programs
- Ex Senior Clinical Program Lead at Genentech, one of the world's most successful biotech businesses which sold the best selling breast cancer drug Herceptin
- Also worked at global majors GSK and Exelixis



**Paul Hopper**  
SYDNEY, AU  
Executive Chairman

- Founder of Imugene
- Former Chairman of Viralytics
- Founder & Director of Prescient
- Chairman of SUDA Pharmaceutical
- Extensive international & ASX biotech capital markets experience particularly in immuno-oncology & vaccines



**Dr Axel Hoos**  
PHILADELPHIA, USA  
Non-Executive Director

- Senior Vice President and Head of Oncology at GSK
- Former Medical Lead for Yervoy, the first immuno-oncology treatment to improve first survival
- Chairman of the Sabin Vaccine Institute
- Co-Chair of the Cancer Immunotherapy Consortium Think-Tank



**Mr Charles Walker**  
BRISBANE, AU  
Non-Executive Director

- Experienced listed biotech CEO and CFO (ASX:ACL and ASX:IMU)
- Extensive financial markets experience having executed 50+ cross border transactions
- Clinical experience includes managing pipeline of drugs in all stages from discovery, through to Phase III to product launch



**Dr Jens Eckstein**  
CAMBRIDGE, USA  
Non-Executive Director

- Managing Partner of Apollo Ventures
- Former president of SR One Ltd., the VC arm of GSK
- 15+ years in VC experience funding early to clinical stage biopharmaceutical companies
- Extensive experience as chairman, board director and founder of several biotechnology and venture capital companies.
- Creator of OneStart, the world's largest life science accelerator



**Dr Lesley Russell**  
PHILADELPHIA, USA  
Non-Executive Director

- 25+ years of senior international operational and leadership experience having worked at Amgen, Eli Lilly, Teva, and Cephalon
- Extensive knowledge and experience with new drug development

Imugene has a team with oncology drug development experience

# Imugene's Scientific Advisory Board consists of world leading oncologists, researchers and developers



**Prof Pravin Kaumaya**  
OHIO STATE UNIVERSITY, USA

- Prof of Medicine Department of Obstetric Gynecology at Ohio State University
- Research focus in tumour immunology, mechanisms of tumour cell-immune cell interactions, and immune mechanisms
- Research focus on fields of vaccine with emphasis on peptide vaccines for cancer



**Dr. Michael Galigiuri**  
CITY OF HOPE, USA

- President of City of Hope National Medical Center and holds the Deana and Steve Campbell Physician-in-Chief.
- Recent President of the American Association for Cancer Research (AACR) in 2017



**Prof. Josep Taberero**  
VALL D'HEBRON, BARCELONA, SPAIN

- President of European Society for Medical Oncology (ESMO)
- President of the Medical Oncology Department at the Vall d'Hebron
- Director of the Vall d'Hebron Institute of Oncology (VHIO)



**Prof Tanius Bekail Saab**  
MAYO CLINIC, USA

- Professor of College of Medicine and Science
- Program Co-Leader, GI Cancer, Mayo Clinic Cancer Center
- Medical Director, Cancer Clinical Research Office (CCRO)
- Senior Associate Consultant, Mayo Clinic AZ



**Prof Peter Schmid**  
BARTS CANCER INSTITUTE, UK

- Medical Oncologist
- Expertise in breast and lung cancer, cancer immunotherapy and early drug development
- Leads the Centre of Experimental Medicine at Barts Cancer Institute



**Prof. Ursula Wiedermann-Schmidt**  
UNIVERSITY OF VIENNA, AUSTRIA

- Co-inventor of HER-Vaxx
- Professor of Vaccinology at Medical University of Vienna



**Dr Neil Segal**  
MEMORIAL SLOAN KETTERING CANCER CENTER, USA

- Medical Oncologist
- Expertise in GI, Colon, Pancreatic cancers
- Active clinical immunology researcher
- Clinical lead in several trials using PD-L1 inhibitors



**Dr Yelina Janjigian**  
MEMORIAL SLOAN KETTERING CANCER CENTER, USA

- Medical Oncologist
- Expertise in esophageal and stomach (gastric) cancer
- Active in GI clinical trials testing combinations of Her-2 and checkpoint inhibitor therapies

Imugene has a world renowned advisory board of scientists and oncologists

# ONCOLYTIC VIRUS SCIENTIFIC ADVISORS



Professor Yuman Fong, OV SAB Chair



Prasad S. Adusumilli

Deputy Chief, Thoracic Service; Co-Director, Mesothelioma Program; Head, Solid Tumors Cell Therapy, Cellular Therapeutics Center. Memorial Sloan Kettering Cancer Centre. These therapies include immunotherapy (enhancing patients' own immune systems using genetic and cell engineering) and oncolytic viral therapy (killing cancer cells using genetically engineered viruses).



Memorial Sloan Kettering  
Cancer Center



Dr Rebecca Auer

Associate Scientist Cancer Therapeutics Program, The Ottawa Hospital Research Institute and Cross-Appointed Member, Associate Professor Department of Surgery and Department Biochemistry, Microbiology and Immunology University of Ottawa. Director of Cancer Research Ottawa Hospital.



uOttawa



Prof James Market

James Garber Galbraith Endowed Chair of Neurosurgery, University of Alabama at Birmingham. His major interest remains the use of herpes simplex virus and other viruses as oncolytic and gene therapy vectors for the treatment of malignant brain tumors and other cancers







Imugene is acquiring the worldwide exclusive license to a promising **oncolytic virus technology** developed at the City of Hope Cancer Centre in Los Angeles.



The virus, known as CF33, is a chimeric poxvirus, and is poised to enter Phase 1 clinical trials in 2020.

# LANDSCAPE: RECENT ONCOLYTIC VIRUS TRANSACTIONS

Oncolytic viruses are attracting the serious attention of big pharma companies such as Merck, Boehringer and Janssen which have made three acquisitions in 2018 alone totalling **over \$1.0 billion**, including Viralytics.

\$340m



\$200m



\$502m



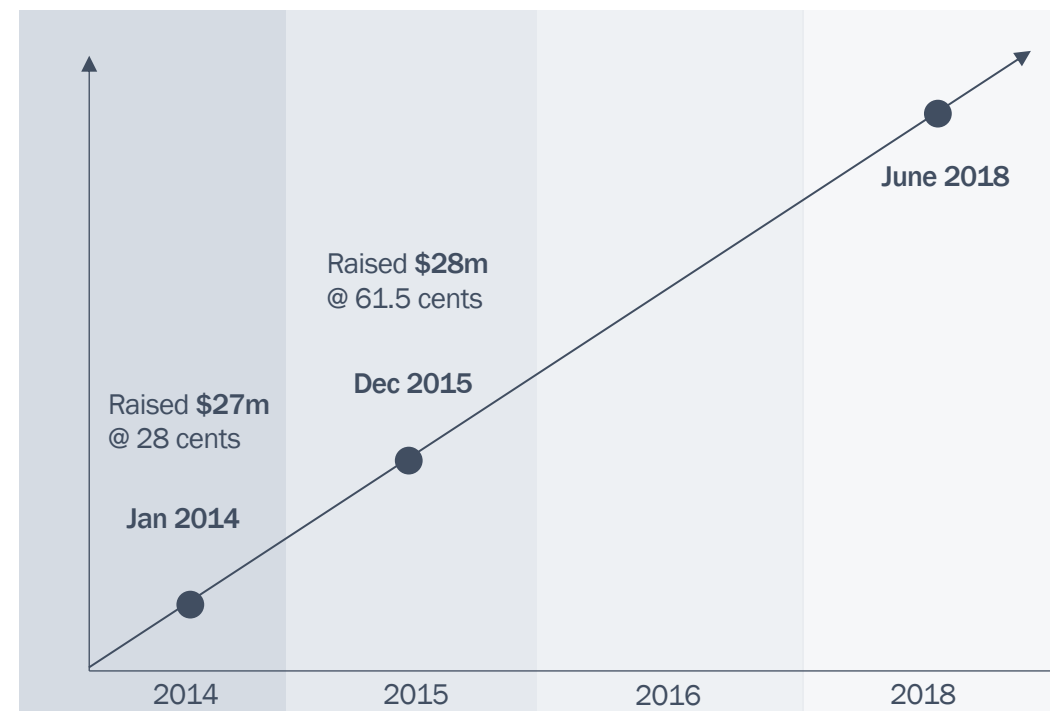
# VIRALYTICS CASE STUDY

## ACQUIRED BY MERCK FOR \$502M

**\$502M** Acquired by  
**MERCK** @ \$1.75



<b>Virus</b>	Picornovirus/coxsackie
<b>Stage of Development</b>	Phase 2
<b>Disease types</b>	Melanoma, bladder, colorectal, non small cell lung
<b>Industry collaboration</b>	Checkpoint combination trial with Merck
<b>Investors</b>	Orbimed, Abbingworth, Baker Bros, BVF, Quest
<b>Team</b>	Paul Hopper (Chair), McColl, Prof Darren Shafren, Turvey, Post



# THE INVENTOR & CITY OF HOPE



Professor  
Yuman  
Fong



A pioneer, both in the operating room and in the laboratory, Prof Yuman Fong, M.D., The Sangiacomo Family Chair in Surgical Oncology and chair of The City of Hope Dept of Surgery is an *internationally recognized expert* in liver and pancreatic cancer. He has developed many new surgical techniques and instruments. He has also led research efforts to use genetically modified viruses to destroy cancer cells.

Prof Fong joined City of Hope in 2014 after more than two decades at the renowned Memorial Sloan-Kettering Cancer Center in New York City.

Prof Fong is both an *author and innovator*. He has written and edited over 700 scholarly articles as well as 14 textbooks. He is currently the Editor-in-Chief of *Molecular Therapy Oncolytics* (Cell Press).

Prof Fong has had leadership roles in regulatory aspects of gene therapy, including serving as Chair or the Recombinant DNA Advisory Committee of the National Institutes of Health of the United States.

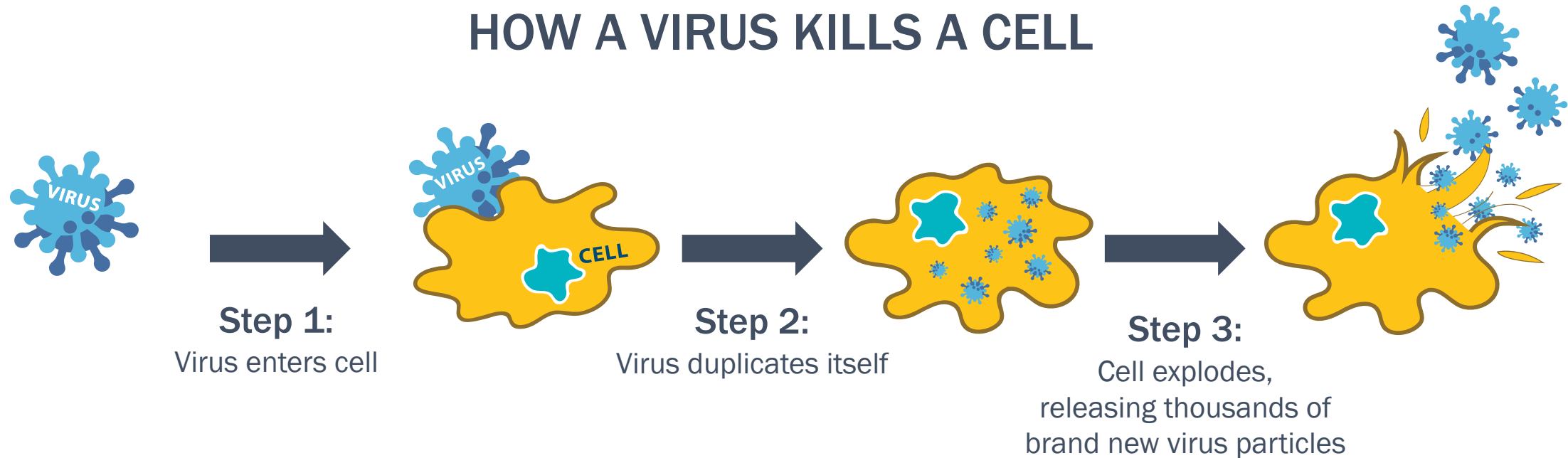
City of Hope, in Los Angeles, is *a leading research and treatment center* for cancer, diabetes and other life-threatening diseases. Founded in 1913, it is designated as a comprehensive cancer center, the highest recognition bestowed by the National Cancer Institute. City of Hope is also a founding member of the National Comprehensive Cancer Network, with research and treatment protocols that advance care throughout the US.

City of Hope has been ranked as one of the nation's "Best Hospitals" in cancer by *U.S. News & World Report* for over 10 years.

City of Hope has GMP facilities that produces clinical trials materials for many academic centers and is the alpha clinic trials site for CIRM

# ONCOLYTIC VIRUS (CF33) MECHANISM OF ACTION

## HOW A VIRUS KILLS A CELL



- **Direct infection**, replication within and cancer cell killing
- **Viral infection** increases local checkpoint targets (PD-1, PD-L1, CTLA4 etc), stimulating the immune system to recognise tumors
- **Cell death** is immunogenic [surface expression of calreticulin, release of adenosine triphosphate (ATP) and release of high mobility group box 1 (HMGB1)]
- **Human sodium** iodine symporter (hNIS) expression allows additional use of  $^{131}\text{I}$ iodine or  $^{188}\text{Re}$ Rhenium killing of infected cells and adjacent cells

# MAJOR ADVANTAGES OF CF33



Preclinical data has demonstrated that CF33 is more **efficacious** than all parental viruses and some viruses in clinical trials.



Especially impressive is that CF33 can **shrink multiple types** of cancer at an extremely low dose (1000 PFU). Also the potential to be more **synergistic** with standard-of-care therapies and emerging novel therapies

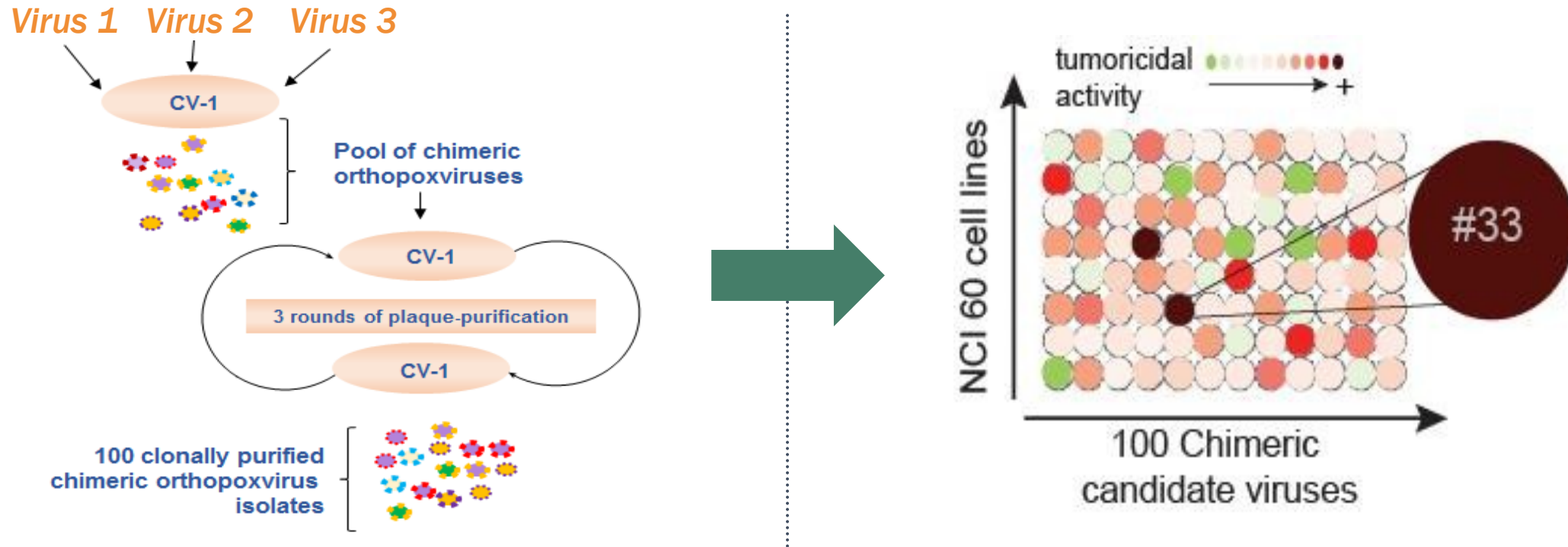


Importantly, CF33 **shrinks** not only injected tumors, but also non-injected **distant tumors** (abscopal effect).

## KEY DIFFERENTIATION

1. **DNA virus** - Much easier to manipulate and vectorize to carry foreign gene as therapeutic payloads
2. **CF33** - more potent in terms of;
  - a) Range of cancer cell types infectible,
  - b) Low doses necessary for cancer killing in vitro and in vivo, and
  - c) Therapeutic window (dose for toxicity minus dose for efficacy)
3. **CF33** can be made in high titres
4. **CF33** can be used in multiple doses without complete neutralization by host immune system

# HOW WAS CF33 DERIVED?



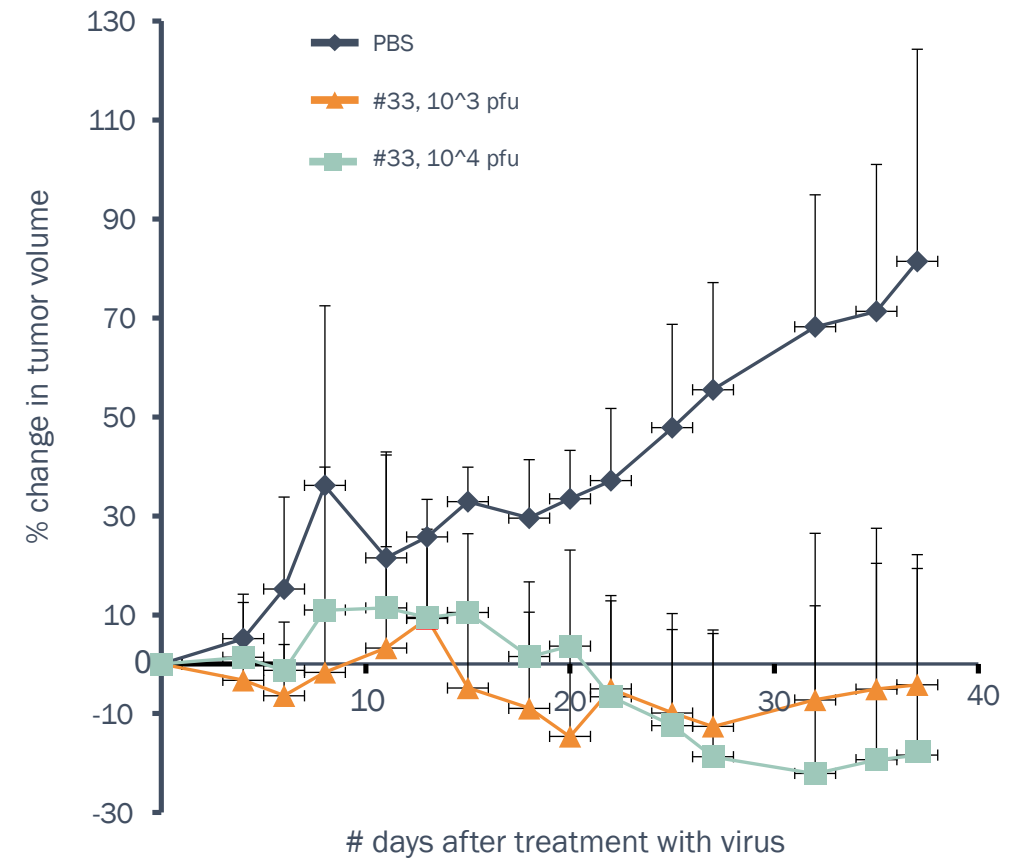
1. 100 chimeric orthopoxviruses and 100 chimeric parapoxviruses were generated
2. Several orthopoxvirus and parapoxvirus chimeras showed superior cancer cell killing in the NCI-60 cell lines
3. CF33 is the chimeric orthopoxvirus chosen for further evaluation *in vivo* and clinical development

## CF33 SHRINKS TRIPLE-NEGATIVE BREAST CANCER

Mice treated with both  
Intratumoral (IT) & IV virus

The viral dose used was **2-5 orders of magnitude** lower than doses used for oncolytic viruses under clinical testing

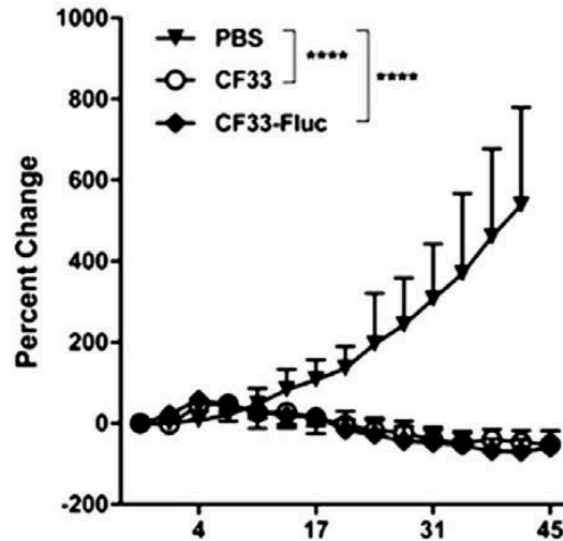
*Mol Ther Oncolytics*. 2018 Jun 29;9





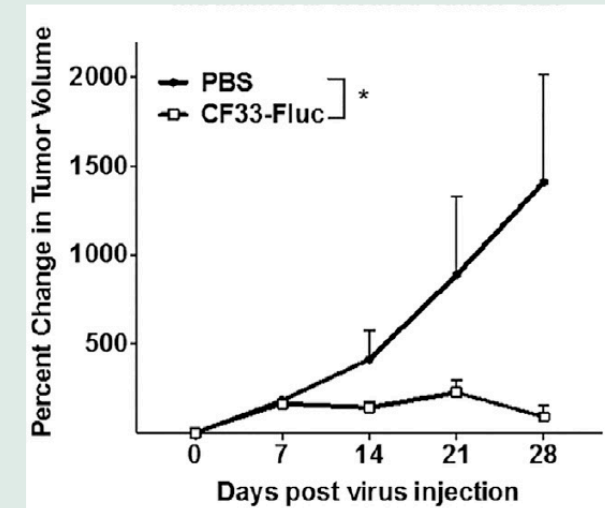
# COMPELLING TUMOUR INHIBITION IN MULTIPLE CANCERS

## PANCREATIC



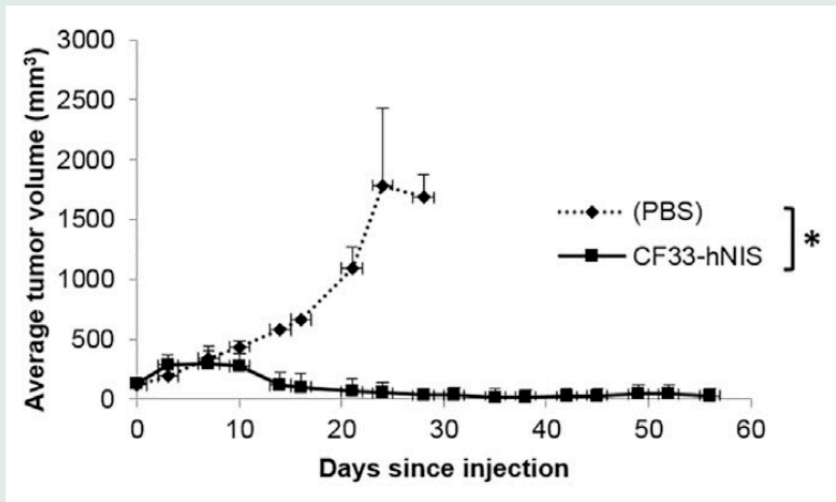
*J Transl Med.* 2018, 16, 110

## COLORECTAL



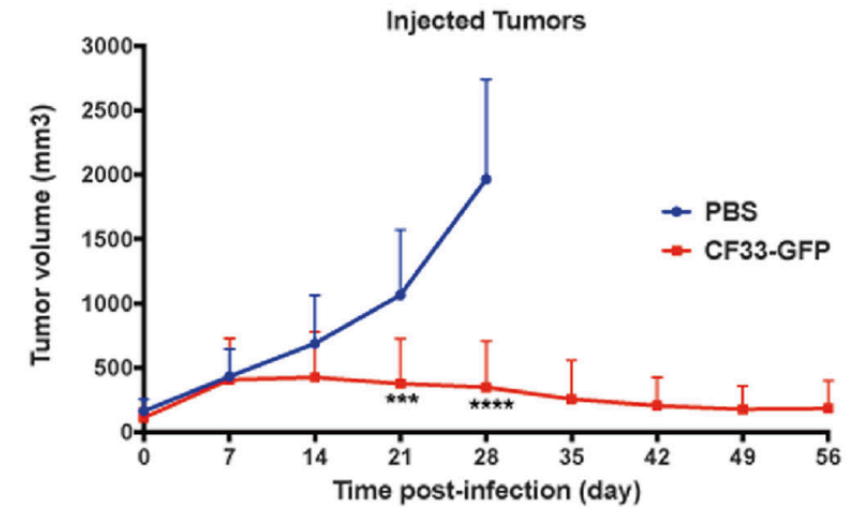
*Mol Ther Oncolytics.* 2018, 9, 13

## COLON



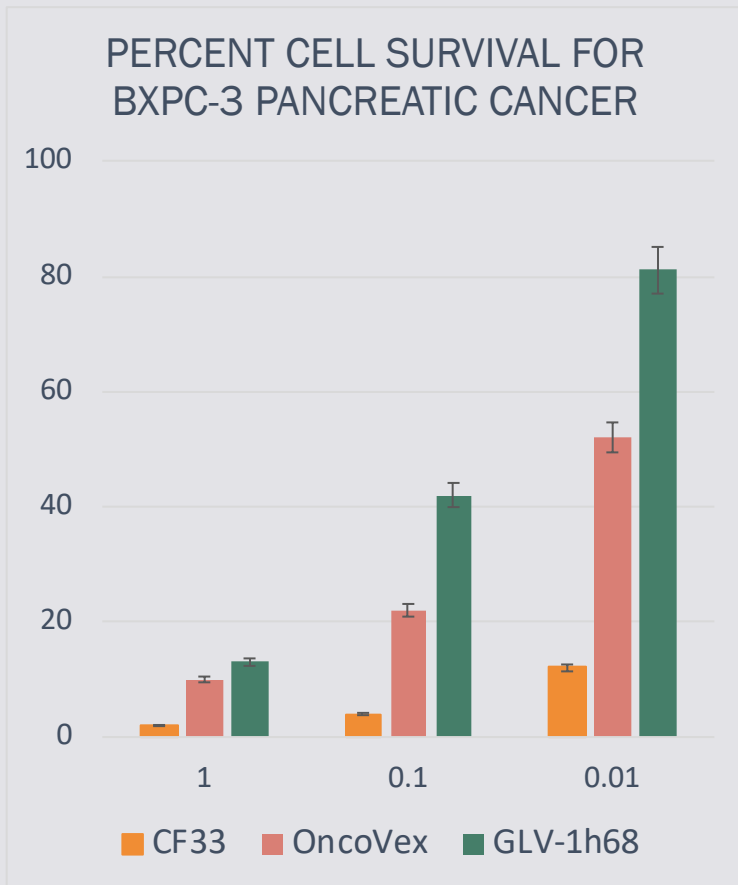
*Mol Ther Oncolytics.* 2019, 13, 82

## LUNG

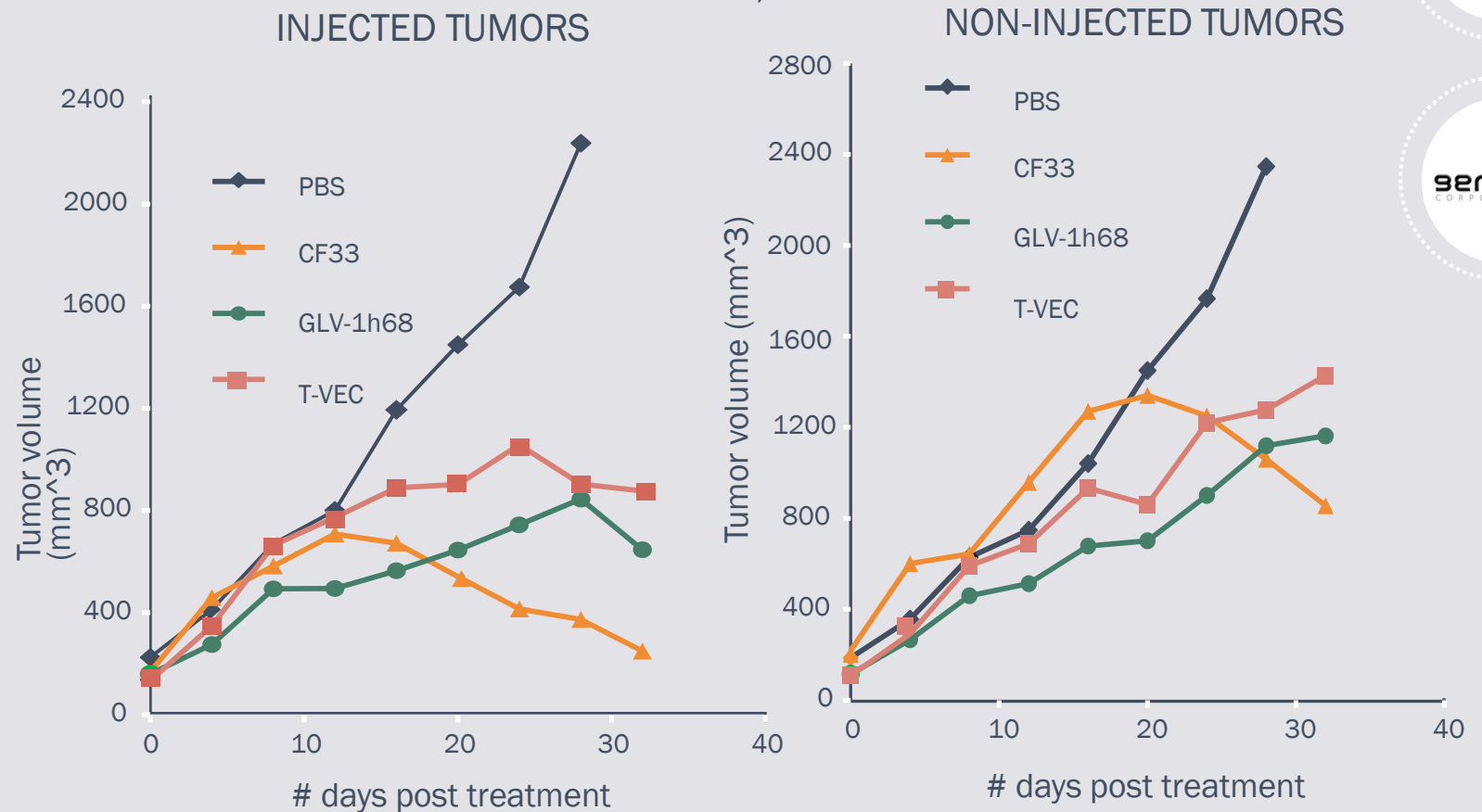


*Cancer Gene Ther.* 2019

# CF33 OUTPERFORMS AMGEN & GENELUX VIRUSES



MICE BEARING THE **A549 XENOGRAPTS** WERE TREATED WITH INDICATED VIRUSES AT A DOSE OF  $10^3$  PFU/MOUSE



# CF33 GMP MANUFACTURING AT CITY OF HOPE



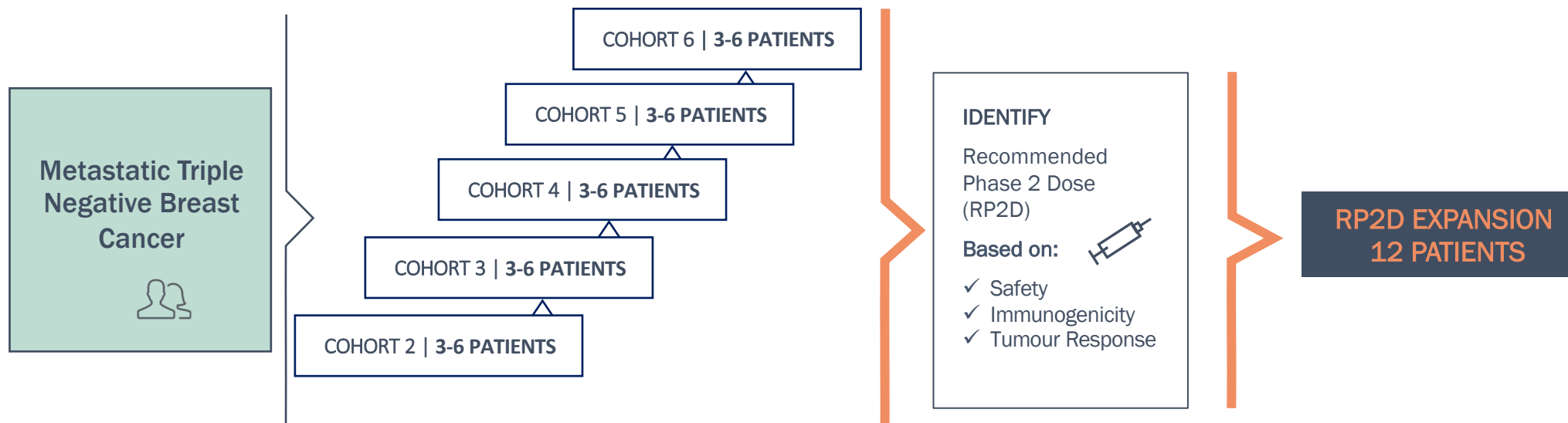
## Center for biomedicine and genetics (CBG)

The Center for Biomedicine & Genetics (CBG) is a California-licensed, 20 000 square foot, multi-product biologics manufacturing facility within City of Hope. With twelve ISO7 production rooms in three product type “zones”, a dedicated aseptic fill suite and a staff with extensive biopharmaceutical experience, the CBG is capable of producing virtually any type of biologic at scales suitable for Phase I through Phase II clinical trials.

- ✓ GMP Phase 1 CHECKvacc (CF33 + hNIS + PD-L1) virus material completed






## Proposed Phase 1 Triple Negative Breast Cancer Study



- ❑ Disease of need
  - 8-13 month survival for metastatic disease with few treatments
- ❑ Potential target for immunotherapy
  - Expresses PD1, PD-L1
- ❑ Treatment responses to Atezolizumab (JAMA Oncology, 5:74, 2019)
  - 1<sup>st</sup> line: 24%; 2<sup>nd</sup> line: 6%
  - Approved by FDA 8-March, 2019
- ❑ Potential for registration in well-designed, randomized P2 study

Indication	TNBC
 FDA IND	CHECKvacc: CF33-hNIS-aPDL1
 N	Part 1=18-24 ; Part 2=12
Location	Single Center: COH
 Admin Route	Intratumoral (IT)
PI	Dr Yuan Yuan

## Proposed Phase 1 & Phase 2 MAST (MIXED ADVANCED SOLID TUMOURS) STUDY

MAST Study Phase 1 Dose Seeking/Signal Finding		MAST Study Phase 2 Simon 2 Stage Design		
	<b>Indication</b>	Lung, TNBC, Melanoma, Bladder, Gastric, Colorectal	<b>Indication</b>	Select tumors from Phase 1
	<b>FDA IND Study Design</b>	1.Vaxinia: CF33-hNIS monotherapy 2.Vaxinia + Immune Checkpoint Inhibitor (ICI) Combination	<b>FDA IND</b>	Vaxinia + Immune Checkpoint Inhibitor (ICI) Combination
	<b>N</b>	Monotherapy: 6 cohorts of 3-6 patients Combination: 2 cohorts of 3-6 patients	<b>N</b>	Depends on the number of Indications
	<b>Location</b>	Multi Centre	<b>Location</b>	Multi Centre
	<b>Admin Route</b>	IT or IV	<b>Admin Route</b>	IT or IV

# INTELLECTUAL PROPERTY

## FOUNDATION PATENT (2037)

<b>PCT</b>	<b>US2017/046163</b>
<b>Title</b>	Chimeric poxvirus compositions & use thereof
<b>Inventor</b>	Yuman Fong
<b>Assignee</b>	City of Hope
<b>Primary Date</b>	9 August 2016
<b>International Publication</b>	18 February 2018

PCT application filing date was 8/9/2017, and estimated expiration date is in late 2037. The patent application includes both composition of matter and method of use. It is currently pending with the opportunity to secure worldwide rights. International search report was favorable.



(43) International Publication Date  
15 February 2018 (15.02.2018)

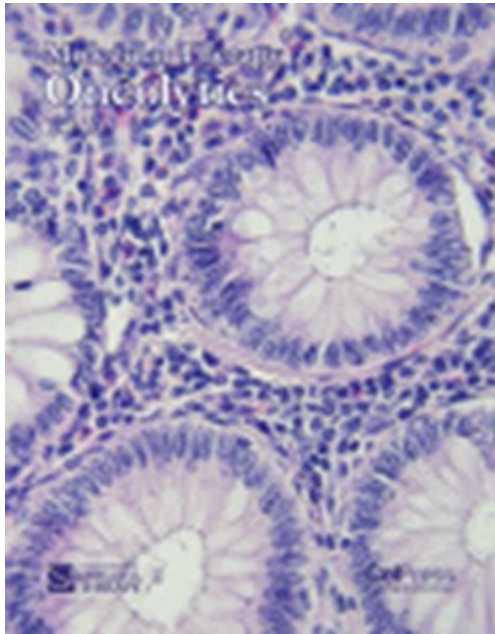


**WO 2018/031694 A1**

- (51) **International Patent Classification:**  
*C12N 7/01* (2006.01)      *C07K 16/28* (2006.01)  
*C12N 15/863* (2006.01)    *A61K 31/7088* (2006.01)  
*C07K 14/47* (2006.01)    *A61K 35/76* (2015.01)
- (74) **Agent:** HETZER-EGGER, Claudia et al; Minitz Levin Cohn Ferris Glovsky And Popeo, P.C., One Financial Center, Boston, MA 021 11 (US).
- (81) **Designated States** (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BN, BR, BW, BY, BZ, CA, CH, CL, CN, CO, CR, CU, CZ, DE, DJ, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IR, IS, JO, JP, KE, KG, KH, KN, KP, KR, KW, KZ, LA, LC, LK, LR, LS, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SA, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW.
- (84) **Designated States** (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LR, LS, MW, MZ, NA, RW, SD, SL, ST, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (AL, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MK, NL, PL, PT, RO, SI, SK, SM, TR), Industrial Property Rights (AR, AU, BR, CA, CL, CO, EC, EG, ES, FI, FR, GB, GR, HU, IE, IL, IN, JP, KR, KZ, LS, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SA, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW).
- (21) **International Application Number:** PCT/US20 17/046 163
- (22) **International Filing Date:** 09 August 2017 (09.08.2017)
- (25) **Filing Language:** English
- (26) **Publication Language:** English
- (30) **Priority Data:**  
 62/372,408      09 August 2016 (09.08.2016)      US  
 62/5 19,010      13 June 2017 (13.06.2017)      US
- (71) **Applicant:** CITY OF HOPE [US/US]; 1500 E. Duarte Road, Duarte, CA 91010 (US).
- (72) **Inventors:** FONG, Yuman; 5219 La Canada Boulevard, La Canada, CA 9101 1 (US). CHEN, Nanhai; 9167 Buck-

# CORE SCIENCE PUBLISHED IN LEADING PEER PUBLICATIONS

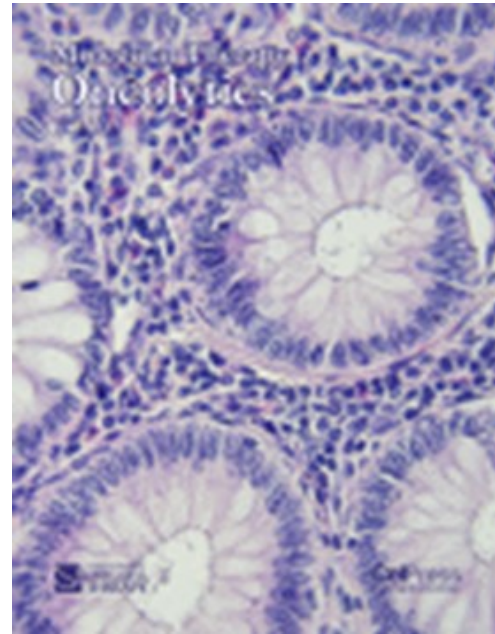
2018



*Mol Ther Oncolytics*. 2018 Jun 29;9

A Novel Oncolytic Chimeric Orthopoxvirus Encoding Luciferase Enables Real-Time View of **Colorectal Cancer** Cell Infection

2018



*Mol Ther Oncolytics*. 2018 Jun 29;9

Endogenous AKT Activity Promotes Virus Entry and Predicts Efficacy of Novel Chimeric Orthopoxvirus in **Triple-Negative Breast Cancer**

2018



*J Transl Med*. 2018 Apr 26;16:110

Novel Oncolytic Chimeric Orthopoxvirus Causes Regression of **Pancreatic Cancer** Xenografts and Exhibits Abscopal Effect at a Single Low Dose

2019



SPRINGER NATURE

*Cancer Gene Ther*. 2019 17 June

A chimeric poxvirus with J2R (thymidine kinase) deletion shows safety and anti-tumor activity in **lung cancer** models

# SELECTED ONCOLYTIC VIRUS DEALS

Date	Source	Buyer	Deal type	Up-front (\$m)	Note
May 2019	Transgene	Astrazeneca	Licensing	10	Five research candidates
Sep 2018	Viratherapeutics	Boehringer Ingelheim	Acquisition	245	VSV-GP project, preclinical
<b>Feb 2018</b>	<b>Viralytics</b>	<b>Merck &amp; Co</b>	<b>Acquisition</b>	<b>394</b>	<b>Cavatak, phase II asset</b>
Nov 2017	Oncolytics	Adlai Norte	Licensing	5	Far East development of Reolysin
Oct 2017	Turnstone Biologics	Abbvie	Licensing	Undisclosed	Ad-MG1-MAGEA3, phase I/II asset
Dec 2016	Ignite Immunotherapy	Pfizer	Acquisition	Undisclosed	50% stake
Dec 2016	Psioxus	Bristol-Myers Squibb	Licensing	Undisclosed	NG-348, preclinical asset
Dec 2016	Takara Bio	Otsuka	Licensing	Undisclosed	Japan rights to HF10
Nov 2016	Virttu Biologics	Sorrento	Acquisition	25 (equity)	Seprehvir, phase II asset
Jun 2016	Psioxus	Bristol-Myers Squibb	Licensing	10	Enadenotucirev, phase I collaboration
Jun 2015	Oncos	Targovax	Acquisition	Undisclosed	Structured as a 50/50 merger
Jan 2015	Omnis	Astrazeneca	Licensing	Undisclosed	VSV project, phase II
Nov 2013	Jennerex	Sillajen	Acquisition	Undisclosed	\$150m biodollar value
Jan 2011	Biovex	Amgen	Acquisition	424	Imlygic, approved for melanoma in 2015

Source: <https://www.evaluate.com/vantage/articles/news/snippets/astrazeneca-doubles-down-oncolytic-viruses> Company statements.

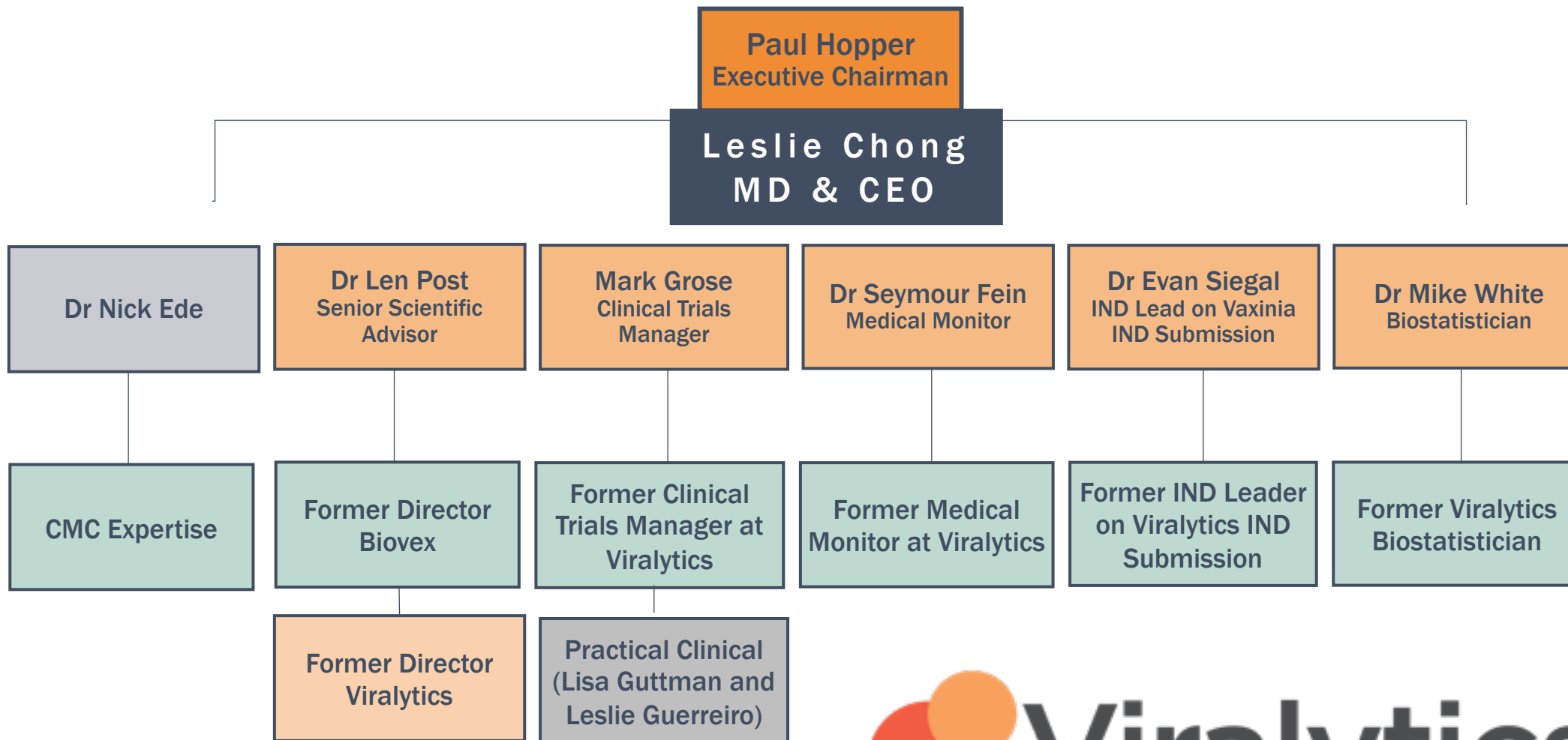


# KEY CURRENT & EMERGING COMPETITION

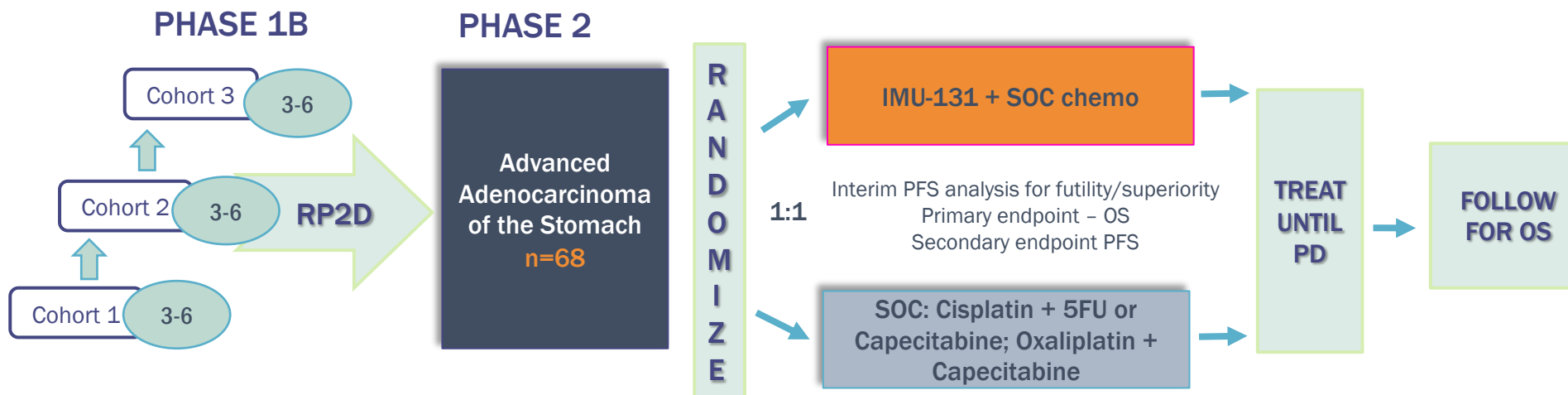
Product	Target/Virus	Comparator	Development Phase & Key Results
Oncorine	Squamous cell carcinoma of the head and neck	Opdivo	Approved in China
Talimogene laherparepvec	Melanoma	Opdivo	Approved in USA
Pexa-Vec (JX-594)	Melanoma	Opdivo	Phase III
REOLYSIN	Squamous cell carcinoma of the head and neck	Opdivo	Phase III
DNX-2401	Melanoma	Opdivo	Phase II
CAVATAK™	Melanoma	Opdivo	Phase II
ColoAd1	Colorectal cancer	Opdivo	Phase I/II
SEPREHVIR	Melanoma	VIRTTU	Phase I/IIa
GL-ONC1	Ovarian cancer	Genelux	Phase I

➤ Running out of IP  
 ➤ Too expensive to deliver  
 ➤ Poor efficacy

# CF33 MANAGEMENT TEAM



# HER-Vaxx PHASE 1B/2 STUDY DESIGN



Phase	Phase 1B	Phase 2
Indication	Newly diagnosed HER2+ gastric cancer	Newly diagnosed HER2+ gastric cancer
Endpoint	Safety & Tolerability, Immunogenicity, RP2D	Primary: OS, Secondary: PFS, Safety & Tolerability, Immune Response
No of Patients	up to 18	68
Site location	Asia, Eastern Europe, India	Asia, Eastern Europe, India

# HER-Vaxx PHASE 1B: DESIGN & RESULTS



## Trial

- HER2 Gastric or GEJ cancer
- Phase 1b
- Open label
- Dose escalation
- 14 sites in Asia and Eastern Europe



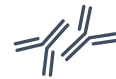
## Patients

- Advanced stage IIIb or IV
- 7 HER2+++, 3 HER2++ (FISH positive), 4 HER2++ expressing tumors
- Age 57yo (21 - 79)
- ECOG 1(7) and 0(7)
- 9 Asian, 5 Caucasian
- 5 female, 9 male



## Study

- 14 patients in 3 cohorts (10µg (3), 30µg (6) and 50µg (5))
- Dosed on D0, D14, D35
- IMU-131 in combination with chemo: cisplatin and 5FU or capecitabine



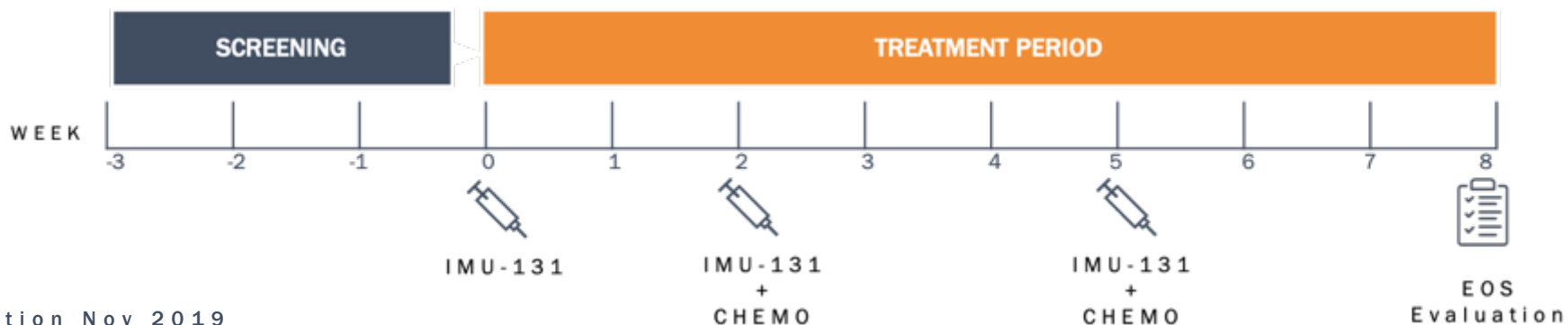
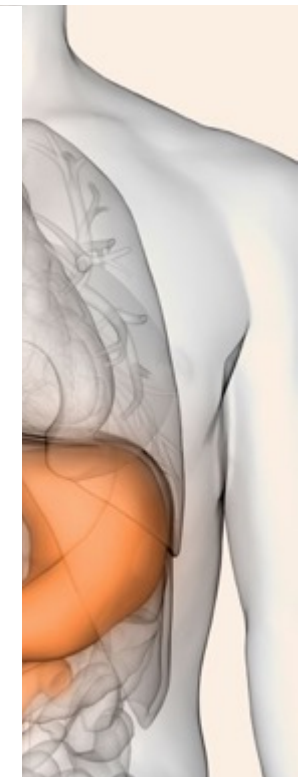
## Endpoints

- Recommended Phase 2 Dose of IMU-131
- Safety and Toxicity
- Immunogenicity (anti-peptide (P467) and anti-HER-2 antibody titres)



## Study Results

- No safety or toxicity issues
- All patients had increased antibody response
- 1 Complete Response
- 5 Partial Response
- 4 Stable Disease
- 1 Progressive Disease
- 50 µg selected as RP2D



# GOING FORWARD: HER-V<sub>axx</sub> PHASE 2 COMMENCED



## Trial

- Phase 2
- Open label
- Asia
- Eastern Europe
- India



## Patients

- HER-2+++
- HER-2++ FISH/CISH +ve
- Advance or metastatic Gastric Cancer
- Stage IIIb/IV
- 68 patients in two arms



## Study

### Randomized

HER-V<sub>axx</sub> in combination with standard of care chemotherapy

### Or

Standard of care chemo: Cisplatin and 5FU or capecitabine or oxaliplatin

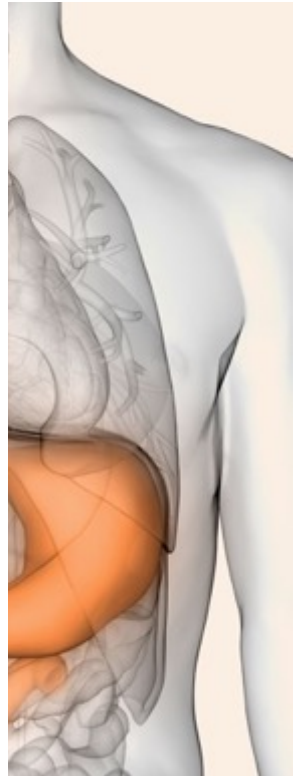


## Primary Endpoints

- Overall survival

## Secondary Endpoints

- Progression-free survival
- Safety and Tolerability
- Immune response



First patient dosed March 2019

# HER-Vaxx PUBLICATIONS

## AACR

**IMUGENE**  
 A Phase Ib open label multicenter study with a HER2/neu peptide vaccine administered with cisplatin and 5-fluorouracil or capecitabine chemotherapy shows safety, immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

**Abstract** HER2/neu overexpression is a key prognostic factor in gastric cancer. Immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

**Background:** HER2/neu overexpression is a key prognostic factor in gastric cancer. Immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

**Methods:** A Phase Ib open label multicenter study with a HER2/neu peptide vaccine administered with cisplatin and 5-fluorouracil or capecitabine chemotherapy shows safety, immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

**Results:** The study shows safety, immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

**Conclusion:** The study shows safety, immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

**Figure 1:** Kaplan-Meier plot showing overall survival (OS) for patients with HER2/neu overexpressing advanced cancer of the stomach. The plot compares two treatment groups: cisplatin and 5-fluorouracil (n=15) and capecitabine chemotherapy (n=15). The OS curves are similar, indicating no significant difference in survival between the two groups.

## AACR 2019

A Phase Ib open label multicenter study with a HER2/neu peptide vaccine administered with cisplatin and 5-fluorouracil or capecitabine chemotherapy shows safety, immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

## ESMO-GI

**IMUGENE**  
 A Phase Ib Study of IMU-131 HER2/neu Peptide Vaccine plus Chemotherapy in Patients with HER2/neu Overexpressing Metastatic or Advanced Adenocarcinoma of the Stomach or Gastroesophageal Junction

**Abstract** HER2/neu overexpression is a key prognostic factor in gastric cancer. Immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

**Background:** HER2/neu overexpression is a key prognostic factor in gastric cancer. Immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

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**Results:** The study shows safety, immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

**Conclusion:** The study shows safety, immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

**Figure 2:** Kaplan-Meier plot showing overall survival (OS) for patients with HER2/neu overexpressing advanced cancer of the stomach. The plot compares two treatment groups: cisplatin and 5-fluorouracil (n=15) and capecitabine chemotherapy (n=15). The OS curves are similar, indicating no significant difference in survival between the two groups.

## ESMO-GI 2019

A Phase 1B study of IMU-131 HER2/NEU peptide vaccine plus chemotherapy in patients with HER2/NEU overexpressing metastatic or advanced adenocarcinoma of the stomach or gastroesophageal junction

## ASCO

**IMUGENE**  
 A Phase Ib Study of IMU-131 HER2/neu Peptide Vaccine plus Chemotherapy in Patients with HER2/neu Overexpressing Metastatic or Advanced Adenocarcinoma of the Stomach or Gastroesophageal Junction

**Abstract** HER2/neu overexpression is a key prognostic factor in gastric cancer. Immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

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**Figure 3:** Kaplan-Meier plot showing overall survival (OS) for patients with HER2/neu overexpressing advanced cancer of the stomach. The plot compares two treatment groups: cisplatin and 5-fluorouracil (n=15) and capecitabine chemotherapy (n=15). The OS curves are similar, indicating no significant difference in survival between the two groups.

## ASCO 2019

A Phase 1b study of IMU-131 HER2/NEU peptide vaccine plus chemotherapy in patients with HER2/NEU overexpressing metastatic or advanced adenocarcinoma of the stomach or gastroesophageal junction

## ESMO

**IMUGENE**  
 Comprehensive results of a Phase Ib study with a HER2/neu B-cell peptide vaccine administered with cisplatin and 5-fluorouracil or capecitabine chemotherapy shows safety, immunogenicity and clinical response in patients with HER2/neu overexpressing advanced gastric cancer

**Abstract** HER2/neu overexpression is a key prognostic factor in gastric cancer. Immunogenicity and clinical response in patients with HER2/neu overexpressing advanced cancer of the stomach

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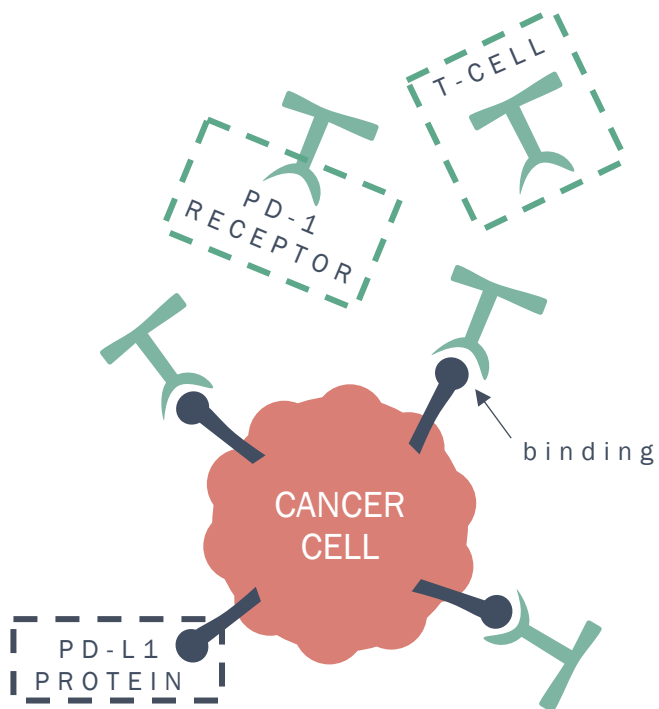
**Figure 4:** Kaplan-Meier plot showing overall survival (OS) for patients with HER2/neu overexpressing advanced cancer of the stomach. The plot compares two treatment groups: cisplatin and 5-fluorouracil (n=15) and capecitabine chemotherapy (n=15). The OS curves are similar, indicating no significant difference in survival between the two groups.

## ESMO-2019

Comprehensive results of a Phase Ib study with a HER2/neu B-cell peptide vaccine administered with cisplatin and 5-fluorouracil or capecitabine chemotherapy shows safety, immunogenicity and clinical response in patients with HER2/neu overexpressing advanced gastric cancer

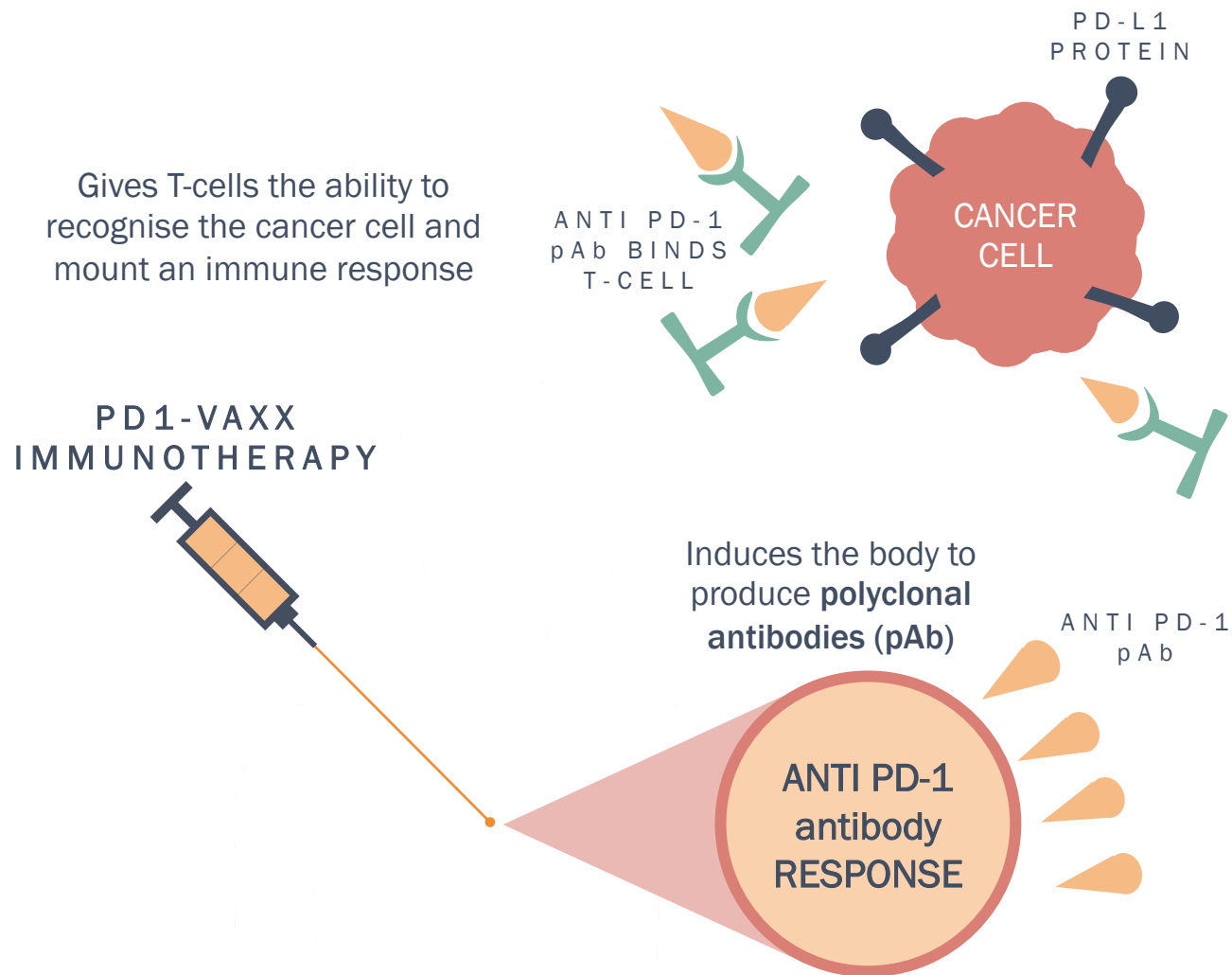
# HOW DOES PD1-Vaxx WORK?

## HOW CANCER STAYS UNDETECTED BY THE IMMUNE SYSTEM



The PD-L1 protein binds to the PD-1 receptor and stops the T-Cell from recognising the cancer cell, allowing the cancer cell to survive and spread

## PD1-VAXX STOPS THE CANCER CELL FROM AVOIDING T-CELL RECOGNITION AND KILLING



# PD1-Vaxx PUBLICATIONS

## AACR

**Abstract 1453: Development of a novel PD-1 vaccine and in combination with two Chimeric HER-2 peptide vaccine provides synergistic inhibition of tumor growth in a syngeneic Balb/c model challenged with CT26/HER-2 carcinoma cell line**

Tania Bokil Gadd, Jay Chhabra, Ashish Bhat, Manish Patel, John Guo, Prash T Rajasekar, Mayo Clin Cancer Center, Phoenix, AZ; The Ohio State University Wexner Medical Center and The James Comprehensive Cancer Center, Columbus, OH; UCLA, Los Angeles, CA

**Background:** Despite blockade of the signaling axis between PD-1 and BTLA-1 with monoclonal antibodies to have remarkable clinical success in the treatment of cancer and increased response activity across a broad set of cancer histologies. Similarly to treatment with monoclonal antibodies, novel B-cell epitopes have been discovered by utilizing a specific immune response that an antibody can induce using B-1 of response, while utilizing immune modulator to suppress. We have a B-cell epitope vaccine approach that is based on the protein-protein interface that binds B-1 epitopes and is based on a chimeric HER-2 peptide vaccine. We have a B-cell epitope vaccine approach that is based on the protein-protein interface that binds B-1 epitopes and is based on a chimeric HER-2 peptide vaccine. We have a B-cell epitope vaccine approach that is based on the protein-protein interface that binds B-1 epitopes and is based on a chimeric HER-2 peptide vaccine.

**Methods:** We have a B-cell epitope vaccine approach that is based on the protein-protein interface that binds B-1 epitopes and is based on a chimeric HER-2 peptide vaccine. We have a B-cell epitope vaccine approach that is based on the protein-protein interface that binds B-1 epitopes and is based on a chimeric HER-2 peptide vaccine.

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## AACR 2019

Development of a novel PD-1 vaccine and in combination with two Chimeric HER-2 peptide vaccine provides synergistic inhibition of tumor growth in a syngeneic Balb/c model challenged with CT26/HER-2 carcinoma cell line

## ESMO

**Abstract 1218P: Antitumor activity and safety of a novel PD-1 vaccine (PD1-Vaxx) alone and in combination with two chimeric HER-2 peptide vaccine (B-Vaxx) in syngeneic Balb/c mice and canines**

Prash T Rajasekar, Jay Chhabra, Ashish Bhat, Manish Patel, Nicholas King, Linda Chung, Tania Bokil Gadd, The Ohio State University Wexner Medical Center, The James Comprehensive Cancer Center, Columbus, OH, UCLA, Los Angeles, CA; Imugene Inc, San Diego, CA; Mayo Clinic Cancer Center, Phoenix, AZ

**Background:** Monoclonal antibodies (mAbs) targeting PD-1 with a high-affinity and high specificity appear to have remarkable clinical success in the treatment of cancer. Similarly to treatment with monoclonal antibodies, novel B-cell epitopes have been discovered by utilizing a specific immune response that an antibody can induce using B-1 of response, while utilizing immune modulator to suppress. We have a B-cell epitope vaccine approach that is based on the protein-protein interface that binds B-1 epitopes and is based on a chimeric HER-2 peptide vaccine.

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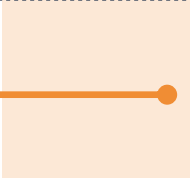
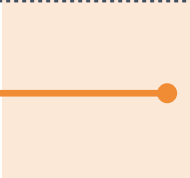
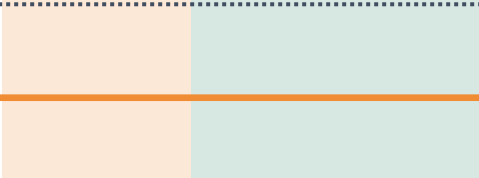
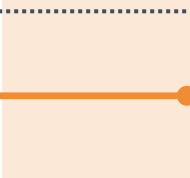

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## ESMO-2019

Antitumor activity and safety of a novel PD-1 vaccine (PD1-Vaxx) alone and in combination with two chimeric HER-2 peptide vaccine (B-Vaxx) in syngeneic Balb/c mice and canines



# IMUGENE HAS A DEVELOPING PIPELINE

	Pre-Clinical	Clinical development Phase 1	Clinical development Phase 2	Key Data / Results	Key IP patents
<b>Vaxinia (CF33)</b>				<ul style="list-style-type: none"> <li>CF33 has shown strong anti tumour responses in preclinical studies</li> <li>Inhibition of tumour growth in nearly all NCI60 models in TNBC, Lung, Pancreatic etc.</li> <li>Signs of increased tumour growth inhibition with CF33 + anti PD-L1</li> </ul>	Intellectual property patents expiring 2037
<b>CheckVacc (CF33 &amp; aPD-L1)</b>				<ul style="list-style-type: none"> <li>Pre-clinical studies showed cancer growth inhibition was better than compared to Amgen or Genelux oncolytic virus.</li> <li>Potentially solves the industry problem of additive toxicity of combined checkpoint inhibitors if safety of CF33 is maintained in combination</li> </ul>	Intellectual property patents expiring 2037
<b>HER-Vaxx (HER-2)</b>				<ul style="list-style-type: none"> <li>Successful completion of Phase 1b trials</li> <li>Strong trial results with no safety or toxicity issues</li> <li>All patients had increased antibody response</li> <li>11/14 evaluable patients with encouraging clinical responses</li> </ul>	Intellectual property patents going out to 2036
<b>PD1-Vaxx</b>				<ul style="list-style-type: none"> <li>PD1-Vaxx has shown encouraging response in preclinical studies</li> <li>Strong inhibition of tumour growth in mouse models of colorectal cancer (outperformed industry standard mouse PD-1 mAb)</li> <li>Signs of increased tumour growth inhibition when co-administered with B-Vaxx</li> </ul>	Intellectual property patents expiring March 2037 & February 2038
<b>B-Vaxx (HER-2)</b>				<ul style="list-style-type: none"> <li>Positive Phase 1 results and now currently in phase 2</li> <li>B-Vaxx is fully funded by OSU grant</li> <li>14/24 evaluable late stage patients with encouraging clinical response</li> </ul>	Intellectual property patents expiring April 2027 & August 2030

# MULTIPLE VALUE REALISATION PATHWAYS



**SALE OF COMPANY**

OR



**PARTNER WITH BIG  
PHARMA**

OR



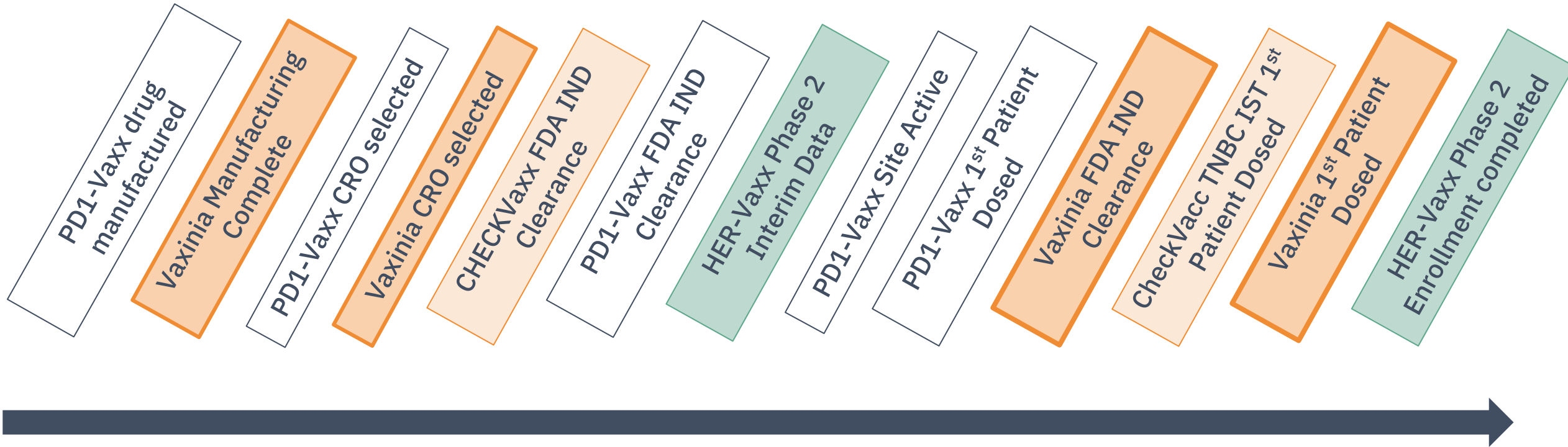
**LICENSE A TARGET  
DISEASE**

OR



**DEVELOP  
INDEPENDENTLY**

# MULTIPLE NEAR & MEDIUM TERM VALUE INFLECTION POINTS



Next 12 months

# Financial Summary

## Public Market Overview

Share Price <sup>1</sup>	A\$0.023
Market Capitalisation <sup>2</sup>	A\$83.1M
Cash equivalents (30 Sep 19 + R&D rebate)	A\$17.8M
Enterprise Value	A\$65.3M

## Top 5 Shareholders (as at October 2019)

National Nominees Limited	5.6%
Dr. Nicholas Smith	3.3%
Paul Hopper	2.1%
HSBC Custody Nominees (Australia)	1.8%
Sarah Cameron	1.7%

Note:

1. As of 29 October 2019

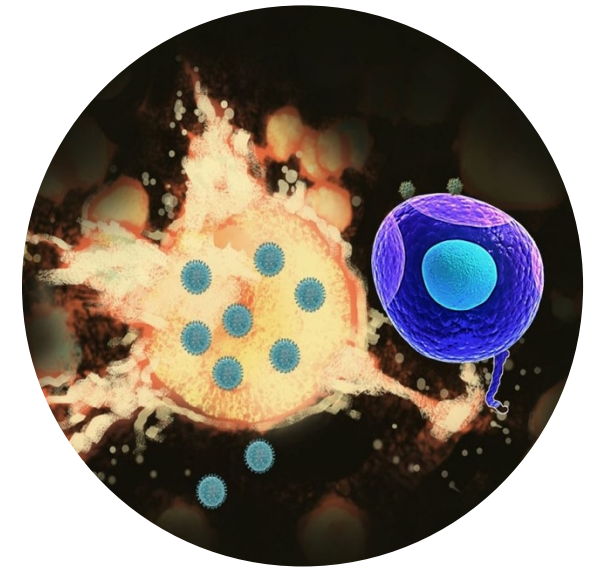
2. Market capitalization calculations based on ordinary shares (3.61bn) only and excludes the dilutive impact of options outstanding (652m)

## Share Price Performance (last 6 months)



# INVESTMENT HIGHLIGHTS

- Novel technology in one of the most sought-after areas of cancer immunotherapy today: oncolytic viruses a.k.a. cancer killing viruses that stimulate immune recognition of cancer & our B-cell Immunotherapy pipeline
- CF33 poised to enter into 2 x Phase 1 clinical trials in 2020; PD1-Vaxx to enter into phase 1 study in 2020
- Robust intellectual property: long patent life & composition of matter for CF33 to 2037 and HER-Vaxx to 2036
- Highly experienced immunotherapy developers including the oncolytic virus team (all ex-viralytics)
- Potential applications across many cancers, including combination with CTLA4/PD-1/PD-L1 checkpoint inhibitors or with engineered immune cells
- Outstanding scientific provenance across all our technology: B-cell immunotherapies from Medical University of Vienna and OSU & CF33 from one of the US leading cancer centres, City of Hope in Los Angeles with Inventor, Professor Yuman Fong who is an internationally recognized oncolytic virus and cancer expert
- Attractive license terms - worldwide exclusive rights to the technology









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**Managing Director & CEO**

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