



Corporate Overview

September 2024



FORWARD LOOKING STATEMENTS

This presentation includes forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995. In some cases forward-looking statements can be identified by terminology such as "may," "should," "potential," "continue," "expects," "anticipates," "intends," "plans," "believes," "estimates," and similar expressions, and include statements regarding oncolytic viruses (OVs) being promising cancer therapeutics; the potential of VCN-01 and Theriva OVs and their ability to overcome key OV challenges; clinical advancement of VCN-01 (including completion of enrollment into the pancreatic ductal adenocarcinoma (PDAC) Phase 2 clinical trial in Q3 2024); potential expansion of VCN-01 use in different lines of PDAC treatment; potential synergistic antitumor effects of VCN-01 with topoisomerase inhibitors; the potential of VCN-01 to enable immuno-oncology therapies in refractory tumors; the potential to obtain expedited status from the FDA; and the potential of the albumin shield to enhance OV systemic delivery. These forward-looking statements are based on management's expectations and assumptions as of the date of this press release and are subject to a number of risks and uncertainties, many of which are difficult to predict that could cause actual results to differ materially from current expectations and assumptions from those set forth or implied by any forward-looking statements. Important factors that could cause actual results to differ materially from current expectations include, among others, Theriva Biologics' product candidates demonstrating safety and effectiveness, as well as results that are consistent with prior results, the ability to initiate and complete clinical trials on time and achieve the desired results and benefits, continuing clinical trial enrollment as expected, the ability to obtain regulatory approval for commercialization of product candidates or to comply with ongoing regulatory requirements, regulatory limitations relating to Theriva Biologics' ability to promote or commercialize their product candidates for the specific indications, acceptance of product candidates in the marketplace and the successful development, marketing or sale of Theriva Biologics' products, developments by competitors that render such products obsolete or non-competitive, Theriva Biologics' ability to maintain license agreements, the continued maintenance and growth of Theriva Biologics' patent estate, the ability to continue to remain well financed, and other factors described in Theriva Biologics' Annual Report on Form 10-K for the year ended December 31, 2023 and its other filings with the SEC, including subsequent periodic reports on Forms 10-Q and 8-K. The information in this presentation is provided only as of the date of this release, and Theriva Biologics undertakes no obligation to update any forward-looking statements contained in this release on account of new information, future events, or otherwise, except as required by law.

OVERVIEW

- **Theriva Biologics** is developing unique oncolytic viruses optimized for systemic administration
- **VCN-01** is undergoing a Phase 2b clinical trial in first-line metastatic pancreatic cancer in combination with SoC chemotherapy
- **VCN-01** Phase 1 clinical trials support multiple additional indications (retinoblastoma, HNSCC, CRC) and combinations (immunotherapies)
- **Albumin Shield™** platform and innovative **VCN-X** oncolytic virus discovery engine enable development of a distinct product pipeline

Financial Snapshot

Exchange	NYSE American
Ticker	TOVX
Cash (06/30/2024)	\$16.6M
Projected cash runway	Q2 2025
Average Daily Volume (3M Ave)	715,000
Locations	Rockville, MD Barcelona, Spain



CRC colorectal cancer. HNSCC head and neck squamous cell carcinoma. SoC standard of care. Immunotherapies include immune checkpoint inhibitors and CAR-T cells.

THERIVA PIPELINE

Candidate	Target	Pre-IND	Phase 1	Phase 2	Phase 3	Collaborators	Status*
VCN-01 Selective, Stroma Degrading OV	Pancreatic Cancer (IV) with gemcitabine/nab-paclitaxel						Phase 2b Study On-going Orphan Drug Designation US, EU Fast Track Designation US
	Retinoblastoma (IVit)						Phase 1 Complete, CSR in preparation Orphan Drug Designation US Rare Pediatric Disease Designation US
	HNSCC (IV) + durvalumab						Phase 1 Complete, CSR in preparation
	Solid Tumors – Brain, Ovarian, PDAC (IV)						Phase 1 Studies On-going
VCN-X and Albumin Shield OVs	Solid tumors (IV)						Preclinical Studies On-going
SYN-004 ^[1,2] Oral β -lactamase	Prevention of aGVHD in allo-HCT						Phase 1b/2a On-going
SYN-020 Oral IAP	Potential indications include NAFLD/NASH, celiac, radiation enteritis						Phase 1 Studies Complete



*Based on Management's current beliefs and expectations. aGVHD acute graft-vs-host disease; allo-HCT allogeneic hematopoietic cell transplant. CSR clinical study report. IAP recombinant bovine intestinal alkaline phosphatase. HNSCC head and neck squamous cell carcinoma. IV intravenous. IVit intravitreal.

VCN-01 IS A UNIQUELY ENGINEERED HUMAN ADENOVIRUS 5

Systemic

Access primary and **metastatic** lesions
High dose, highly replicating

Selective

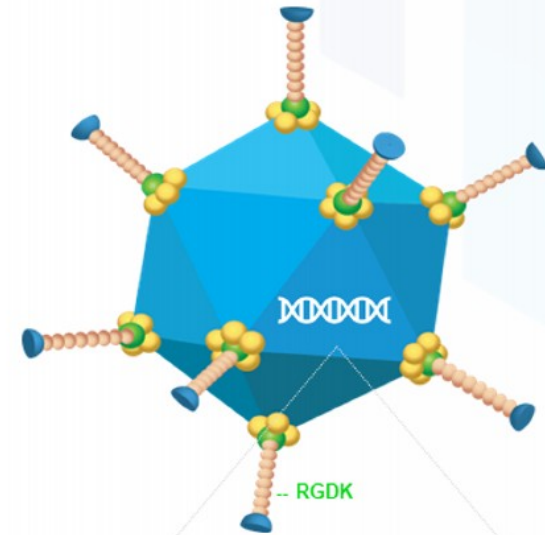
Replicates only in **tumor** cells
Liver detargeted

Stroma Degrading

Expresses **PH20** (hyaluronidase)
after viral replication cycle

Self Reporting

PH20 in blood is a potential **biomarker**
for virus replication in tumors

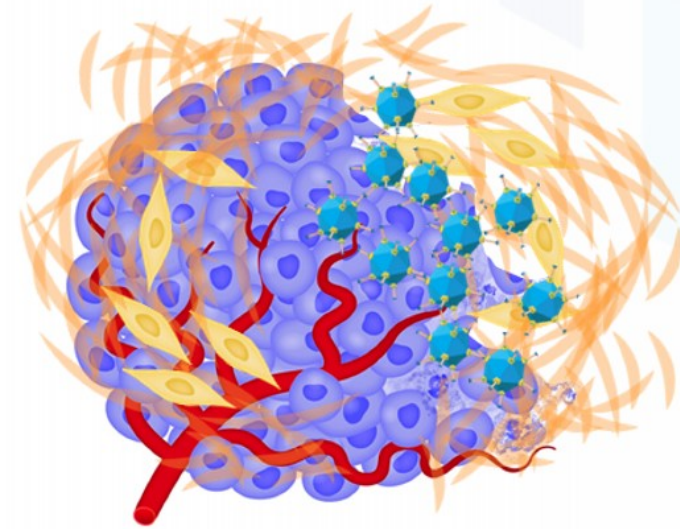
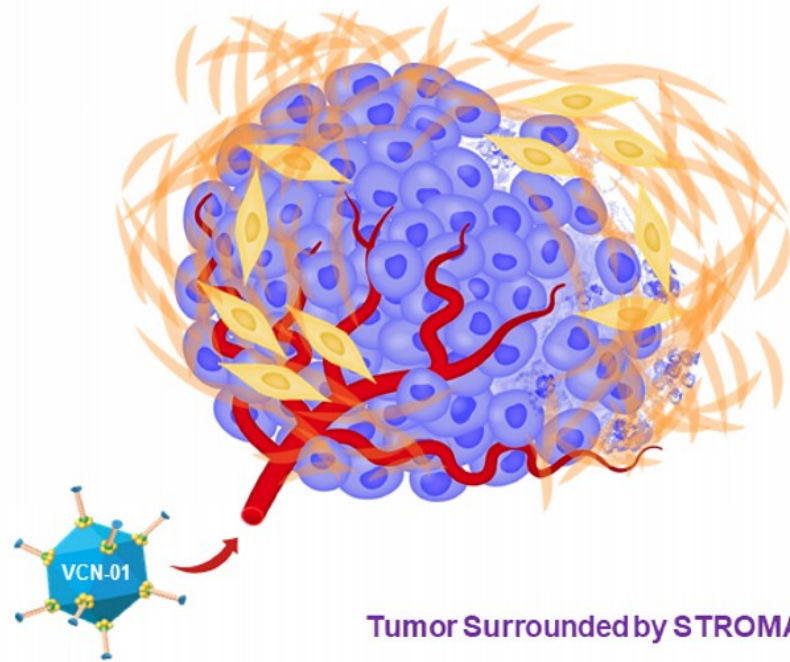


E2F binding +++ → E1a-Δ24 → MLP → PH20

VCN-01 IS A UNIQUELY ENGINEERED HUMAN ADENOVIRUS 5

1 **SYSTEMIC** administration enables VCN-01 access to primary tumor and metastases and detargets the liver

2 **SELECTIVE** replication at very high levels lyses tumor cells directly without harming healthy tissues

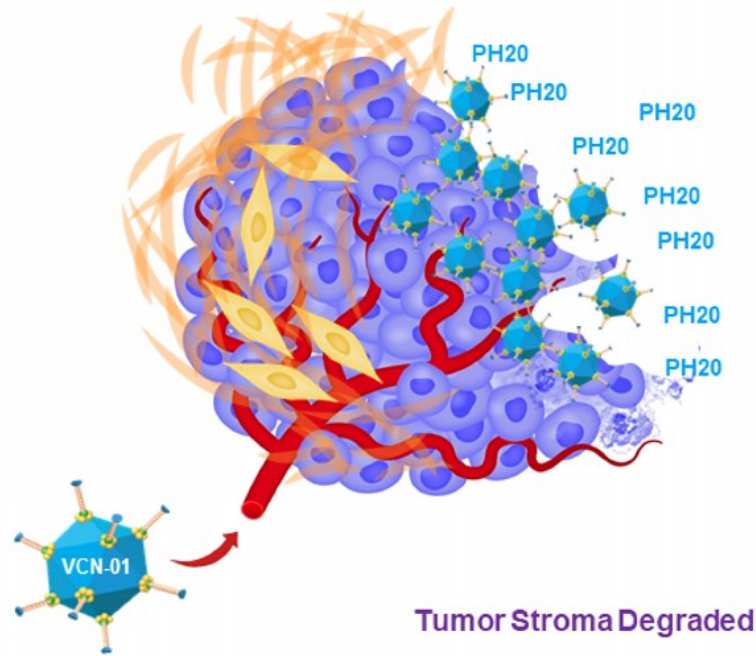


Tumor Surrounded by STROMA

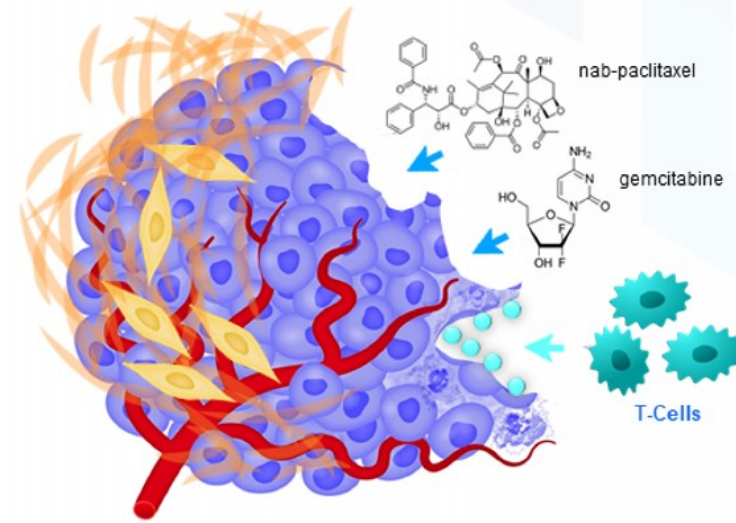


VCN-01 DESIGNED TO HAVE MULTIPLE ANTI-TUMOR ACTIONS

3 **STROMA** degradation by PH20 facilitates tumor access and destruction by coadministered cancer therapies



4 **IMMUNOGENIC** actions of VCN-01 turn "cold" tumors "hot" and elicit an anti-tumor immune response



VCN-01 COMPARED TO OTHER ONCOLYTIC VIRUSES IN DEVELOPMENT

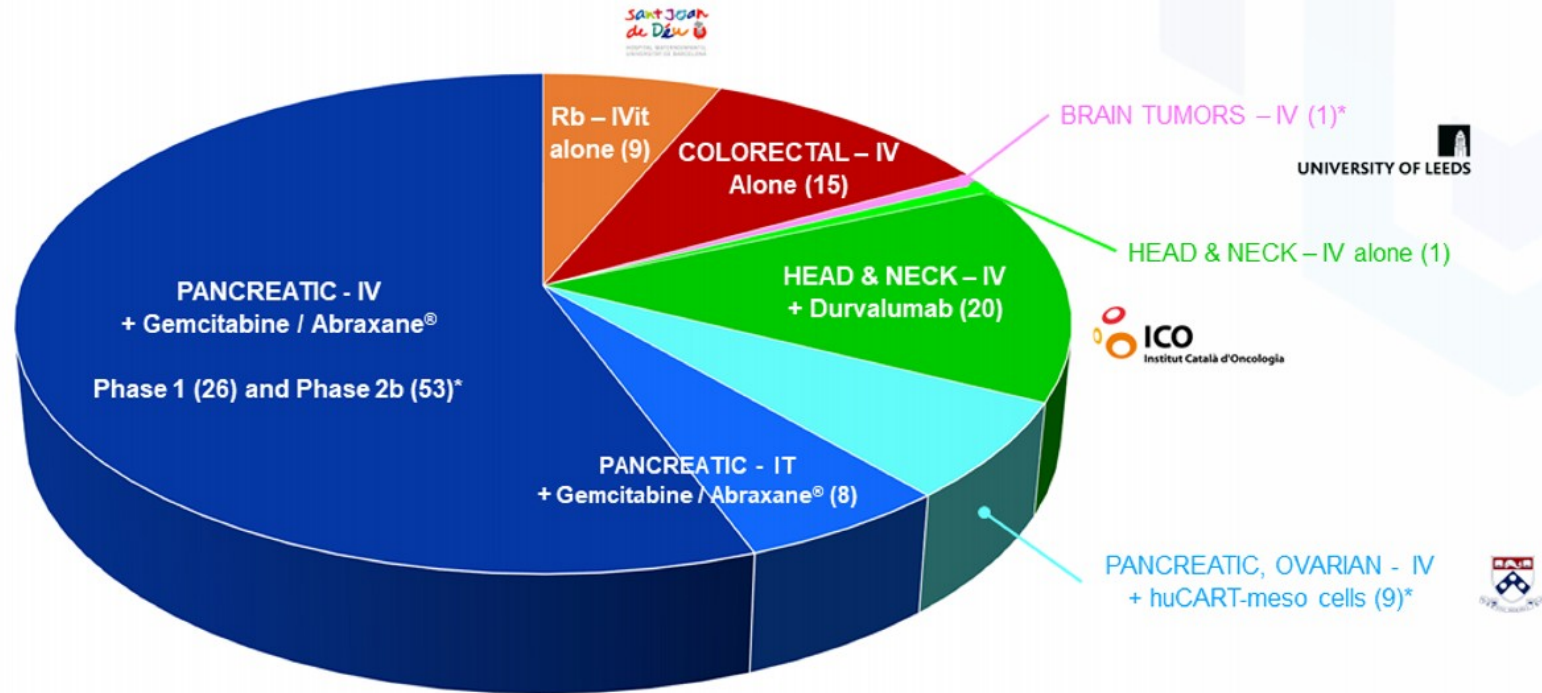
COMPANY	THERIVA BIOLOGICS	CG ONCOLOGY	GENELUX	ONCOLYTICS BIOTECH	REPLIMMUNE
Ticker	NYSE MKT: TOVX	NASDAQ: CGON	NASDAQ: GNLX	NASDAQ: ONCY	NASDAQ: REPL
Market Cap ¹	\$4M	\$2.5B	\$85M	\$71M	\$757M
Product	VCN-01	CG0070	Olvi-Vec	Pelareorep	RP1, RP2
Virus	Adenovirus 5	Adenovirus 5	Vaccinia	Reovirus	Herpes Simplex
Type	DNA	DNA	DNA (enveloped)	RNA	DNA (enveloped)
Tumor selectivity mechanism	Selective replication (Rb-E2F dysfunction)	Selective replication (Rb-E2F dysfunction)	Low tumor IFN TK deletion	Low tumor IFN Ras activation	Low tumor IFN ICP34.5 deletion
Therapeutic Transgene	PH20	GM-CSF	GM-CSF, GALV-GP R(-), anti-CTLA-4
Lead Indication (Ph)	Pancreatic (2b)	Bladder (3)	Ovarian (3)	Pancreatic, GI (2b)	Melanoma (3)
Route	IV	IVESIC	IP	IV	IT
Dose	1x10 ¹³ vp ²	1x10 ¹² vp	3x10 ⁹ pfu	4.5x10 ¹⁰ TCID ₅₀	1x10 ⁷ pfu/mL
Stroma Degrading	Yes	No	No	No	No
Biomarker	PH20	..	β-GAL, β-GLU, GFP



¹At 18Sep2024. ²Approximately 7x10¹¹ TCID₅₀. References describing the different OV product candidates are provided in the appendix; information is also available on each company's website and clinicaltrials.gov

VCN-01 EXTENSIVE CLINICAL PROGRAM

142 patients treated with VCN-01 to date in multiple indications and combinations



(Number of VCN-01 Patients Treated in Parentheses)

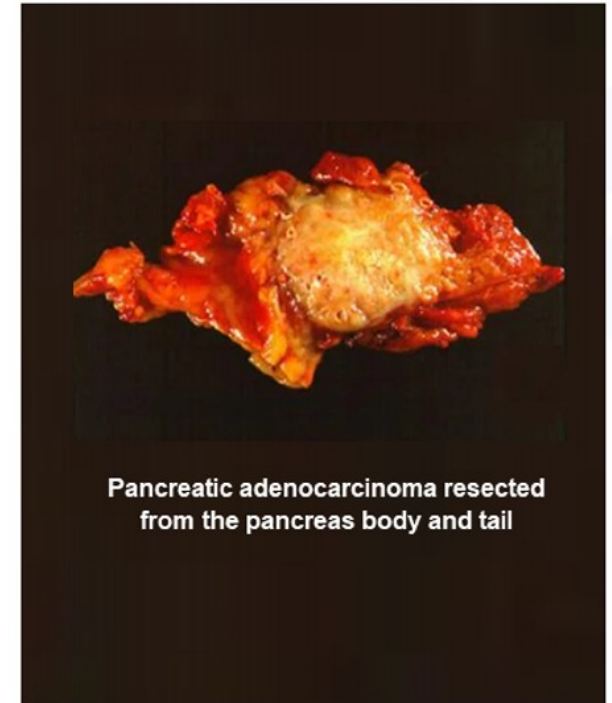


*On-going study. Abraxane® - nab-paclitaxel. Durvalumab (IMFINZI®, AstraZeneca) is an anti-PD-L1 mAb immune checkpoint inhibitor. huCART-meso are autologous T cells engineered to express an extracellular single chain variable fragment (scFv) with mesothelin specificity. IT - intratumoral. IV - intravenous. IVit - intravitreal. Rb - retinoblastoma. See Appendix for study registry numbers and publications.

VCN-01 LEAD INDICATION PANCREATIC CANCER

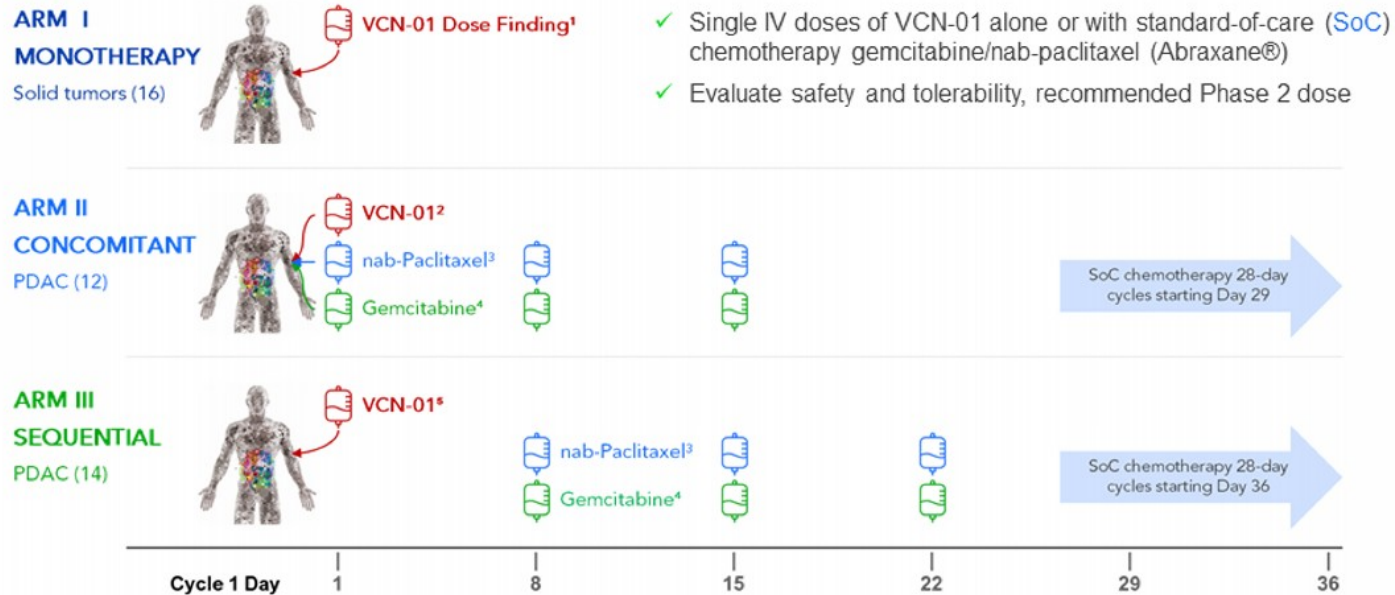
Highly fatal cancer protected by dense tumor stroma

- Orphan disease with the highest mortality of all solid tumors
 - Median survival 8-11 months for metastatic disease^{1,2}
 - USA est. 66,440 new cases and 51,750 deaths in 2024³
- **Hyaluronic acid** in stroma is associated with reduced treatment efficacy and poor prognosis⁴
 - VCN-01 designed to degrade hyaluronic acid
- Incidence is growing worldwide
 - Est. treatment market ~\$2.5B (2022) ~\$7.0B (2030)⁵



VCN-01 FAVORABLE OUTCOMES IN PHASE 1 TRIAL

Multicenter, open-label, dose escalation study (NCT02045602)



¹Single dose of VCN-01 (1×10^{11} to 1×10^{12} vp/dose) administered by 10 min IV infusion. ²VCN-01 doses 3.3×10^{12} vp (n=6) and 1×10^{12} vp (6). ³nab-Paclitaxel (Abraxane®; 30 min infusion) administered at least 4 hours after VCN-01. ⁴Gemcitabine (30 min infusion) administered immediately after Abraxane® infusion. ⁵VCN-01 doses 3.3×10^{12} vp (8) 1×10^{12} vp (6). Garcia-Carbonero (2022) J Immunother Cancer 10:e003255.

VCN-01 FAVORABLE OUTCOMES IN PHASE 1 TRIAL

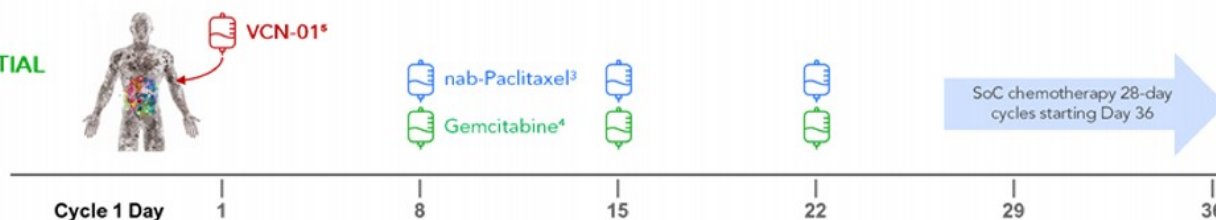
Multicenter, open-label, dose escalation study (NCT02045602)

OUTCOME	VCN-01 DOSE, virus particles (n) ¹			SoC ALONE ²
	3.3x10 ¹² (6)	1.0x10 ¹³ (6)	Combined (12)	Phase 3 (431)
Responders, %	16.7%	83.3%	50.0%	22.9%
Median OS, months	13.1	20.8	13.5	8.5
Median PFS, months	9.9	6.3	6.7	5.5
Survival ≥12 months	.	.	67%	35%

KOLs advise that Hazard Ratio <0.7 is a significant patient outcome

RELATED AEs IN ≥1 PATIENT ¹	CTCAE SEVERITY	
	VCN-01 Combined, Sequential Regimen	Grade 1-2
Pyrexia/Influenza-like Illness	12 (85.7%)	-
Nausea	3 (21.4%)	-
Vomiting	3 (21.4%)	-
Asthenia/Fatigue	3 (21.4%)	-
Transaminase increases (ALT, AST)	2 (14.3%)	2 (14.3%)
Thrombocytopenia	2 (14.3%)	-

ARM III
SEQUENTIAL
PDAC (14)

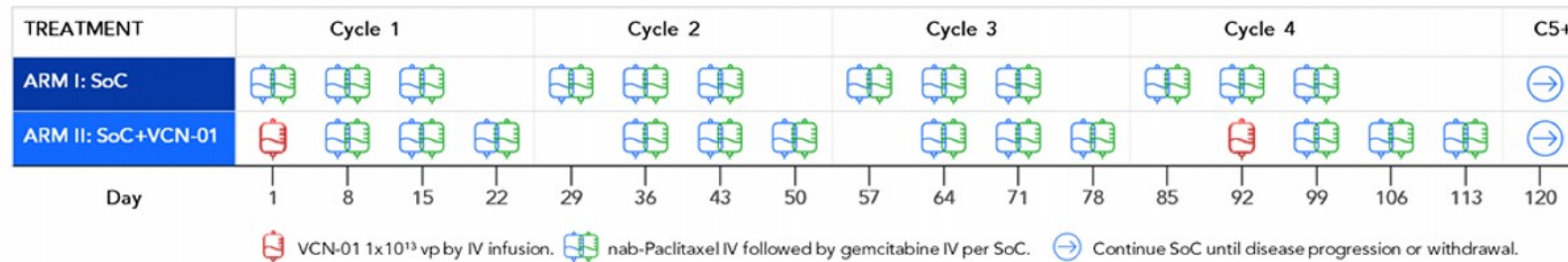


¹Single dose of VCN-01 (1x10¹¹ to 1x10¹³ vp/dose) administered by 10 min IV infusion. ²VCN-01 doses 3.3x10¹² vp (n=6) and 1x10¹³ vp (6). ³nab-Paclitaxel (Abraxane®; 30 min infusion) administered at least 4 hours after VCN-01. ⁴Gemcitabine (30 min infusion) administered immediately after Abraxane® infusion. ⁵VCN-01 doses 3.3x10¹² vp (8) 1x10¹³ vp (6). Garcia-Carbonero (2022) J Immunother Cancer 10:e003255.

VIRAGE PHASE 2B CLINICAL TRIAL in PANCREATIC CANCER

Multicenter, open-label, randomized, controlled trial (NCT05673811)

- Study on-going in patients with **first-line** metastatic pancreatic ductal adenocarcinoma (PDAC)
- Achieved target of 92 patients (46 in each arm) enrolled at sites in Spain and the USA
- Randomized 1:1 to receive gemcitabine/nab-paclitaxel SoC or up to **two doses** of VCN-01 plus SoC
- Primary endpoints **overall survival**, VCN-01 safety and tolerability
- Secondary endpoints include **response rates**, progression free survival, landmark survival



VCN-01 STRATEGY AND PRIMARY CATALYSTS

- Strategy

- Seek regulatory agency agreement to convert the VIRAGE Phase 2b study to a Phase 3 registration trial¹
 - Reduce potential Phase 3 size and costs; expected to shorten time to marketing application
 - Leverage Phase 3 status (if agreed) to engage potential partners

- Primary Catalysts and Use of Funds

- Achieved target enrollment into the VIRAGE Study Q3 2024
- Initiate commercial scale manufacturing process development² Q4 2024
- Regulatory feedback on Phase 3 from FDA, AEMPS (Spain) and EMA Q1 2025
- Conversion to Phase 3 trial (subject to regulatory agreement)² Q2 2025

ACHIEVEMENTS AND PROJECTED MILESTONES

• VCN-01 PDAC

- Ph2 enrollment achieved ✓
- Spanish Government Public-Private Loan-Grant ✓
- Meeting with AEMPS (potential pivotal trial design)

• VCN-01 RETINOBLASTOMA

- RPDD granted ✓

• SYN-004 aGVHD

- Cohort 2 DSMC Outcomes

• VCN-01 PDAC

- Meeting with FDA (potential pivotal trial design)
- Initiate commercial scale manufacturing process development

• VCN-01 RETINOBLASTOMA

- ODD from EMA

• VCN-01 + CAR-T

- U. Penn ASGCT poster

• VCN-01 PDAC

- EMA Scientific Advice (potential pivotal trial design)

• VCN-01 RETINOBLASTOMA

- Finalize Phase 2 study design¹

• VCN OV DISCOVERY

- VCN-12 candidate selection²

• VCN-01 PDAC

- Conversion to Phase 3 trial (if regulatory agreement)¹
- Establish feasibility of commercial scale VCN-01 manufacture

• THERICEL

- Commercial availability of proprietary suspension cell line for manufacturing viral products

Q3 2024

Q4 2024

Q1 2025

H2 2025



¹Contingent on Theriva obtaining required funding for a clinical trial and subject to regulatory agreement. ²VCN-12 is an armed version of VCN-11 designed to express an additional functional payload (VCN-11 is the first clinical candidate using the Albumin Binding Domain™ technology). ISS investigator sponsored Phase 1 study. AEMPS Agencia Española de Medicamentos y Productos Sanitarios. ASGCT American Society of Gene and Cell Therapy. EMA European Medicines Agency. ODD Orphan Drug Designation. RPDD rare pediatric disease designation. SIOP International Society of Paediatric Oncology

TOVX CAPITALIZATION TABLE

Common Shares Issued at September 6, 2024			1,382,672
Less: Treasury Shares			(28,809)
Common Shares Issued and Outstanding at September 6, 2024			<u>1,353,862</u>
Stock Options Issued (2007, 2010, 2020 Plans)	175,191	12.9%	
Common Shares Available for Future Grants 2020 Plans	<u>112,640</u>	8.3%	
	<u>287,831</u>		
Total Reserved Common Shares			287,831
Remaining Master Common Stock Reserve			<u>12,358,307</u>
Total Common Shares Authorized			<u>14,000,000</u>
Treasury Shares			28,809

Note: Numbers are post-split values, rounded for fractional shares



APPENDIX



SEASONED LEADERSHIP TEAM



Steven Shallcross

Chief Executive Officer, Chief Financial Officer

Served as the Company's CEO since 2018 and CFO since joining the Company in 2015

Deep operational, financial and international biotech industry experience and proven track record of leading the financial development and strategy in the public sector

Senseonics

VANDA
PHARMACEUTICALS INC.

Innocoll

nuo
THERAPEUTICS

Theriva
BIOLOGICS



Manel Cascalló PhD

General Director, EU Subsidiary

Expertise in oncolytic adenovirus clinical development, received several patents for the use of adenovirus as antitumoral agents and authored many peer-reviewed scientific publications

Deep regulatory experience and serves as an independent expert for the European Medicines Agencies (EMA)

VCN
BIOSCIENCES



Vince Wachter PhD

Head Corporate Development

Nearly 30 years leading corporate strategy, partnering, research, clinical development, and intellectual property programs for start-ups, small companies, and new business units within large companies

Development experience across oncology, infection, GI, metabolic diseases, transplantation, and drug delivery

EASTMAN

Verva
Pharmaceuticals

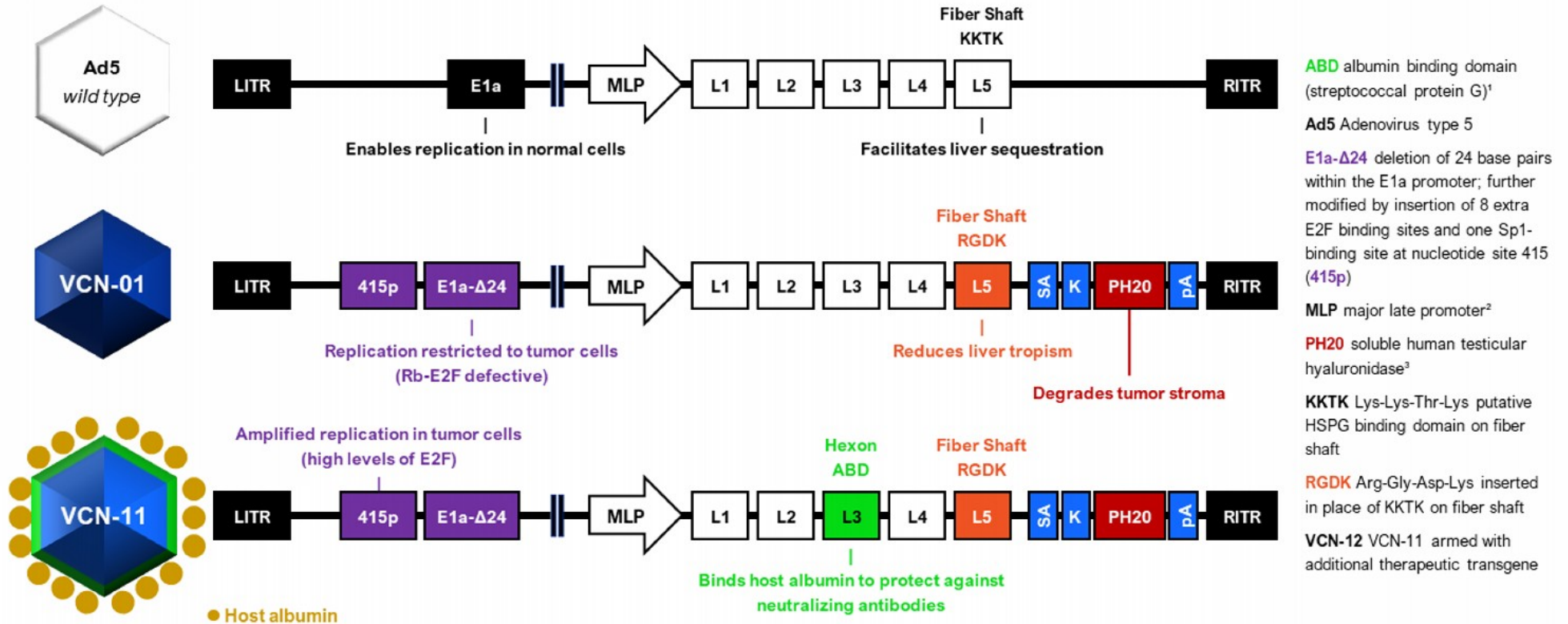
INTELLECTUAL PROPERTY

Hyaluronidase OV	Albumin Shield™	Oral β-Lactamase	Oral IAP
VCN-01, VCN-11	VCN-11, Discovery	SYN-004, -006, -007	SYN-020
Composition of Matter (exp 2030)	Composition of Matter (exp 2034)	Composition of Matter (exp 2031-5)	Manufacturing Know-how (Trade Secret)
Methods of Use and Novel Formulations (examination)	Methods of Use and Novel Formulations (examination)	Methods of Use and Novel Formulations (exp 2035-6)	Methods of Use and Novel Formulations (applications filed)
Use in Rb (exp 2036)			Option to additional IP from MGH
ODD EU (PDAC)			
ODD US (PDAC & Rb)			

VCN-01 IN PANCREATIC CANCER



VCN ONCOLYTIC VIRUS GENETIC MODIFICATIONS



EXTENSIVE VCN-01 PHASE 1 CLINICAL EXPERIENCE

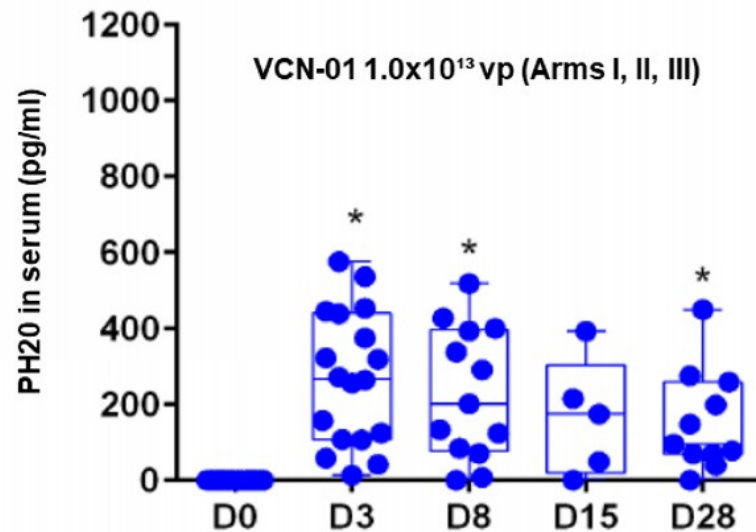
88 Patients Treated in Diverse Cancer Indications

Location	Phase	Indication	Co-therapy	Route	Status	Registry
Multicenter (ESP)	1	Part I: Solid tumor Part II, III: PDAC	Part I: None Parts II,III: Gemcitabine + nab-Paclitaxel	IV	Complete ¹	NCT02045602
Multicenter (ESP)	1	PDAC	Gemcitabine + nab-Paclitaxel	IT	Complete ²	NCT02045589
Hospital Sant Joan de Déu, Barcelona	1	Retinoblastoma	None	IVit	Complete; CSR in prep	NCT03284268
Institut Català d'Oncologia (ICO)	1	HNSCC	Durvalumab	IV	Complete; CSR in prep	NCT03799744
U. Leeds	1	Adult brain tumors	None	IV	Ongoing	ISRCTN 517624869
U. Pennsylvania	1	PDAC, Ovarian	huCART-meso ³	IV	Ongoing	NCT05057715

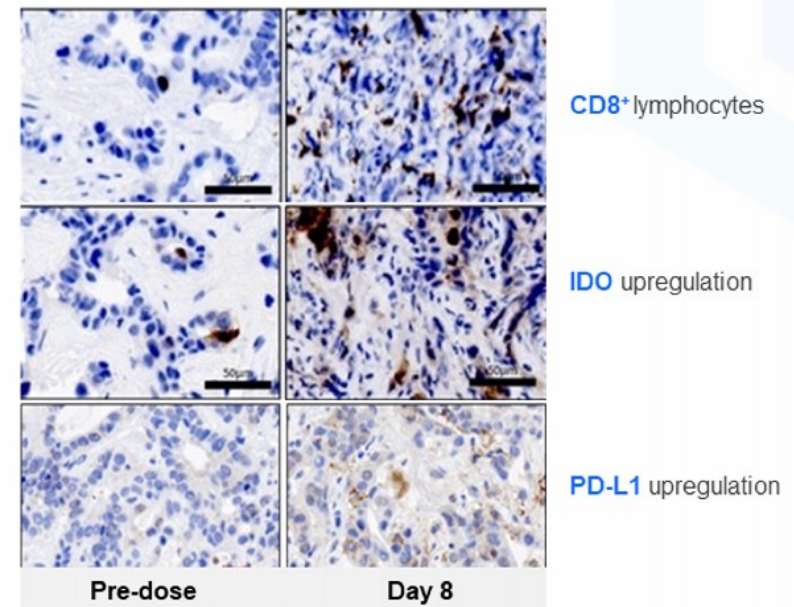
CLINICAL DATA SUPPORT VCN-01 MODE-OF-ACTION

Remodels the tumor matrix and turns “cold” tumors “hot”

Potential biomarker: PH20 levels in patient sera indicate sustained VCN-01 activity in tumors



Immune markers upregulated in biopsies of **hepatic metastases**



MOST COMMON IV VCN-01 RELATED AEs [NCT02045602]

ADVERSE EVENTS ¹	Part I (Alone, n=16)		Part II (Concomitant, 12) ²		Part III (Sequential, 14) ³	
	Grade 1-2	Grade ≥3	Grade 1-2	Grade ≥3*	Grade 1-2	Grade ≥3
Febrile neutropenia	-	-	-	2 (16.7%)	-	-
Neutropenia/Neutrophil count decreased	2 (12.5%)	-	1 (8.3%)	1 (8.3%)	-	-
Thrombocytopenia/Platelet count decreased	1 (6.3%)	1 (6.3%)	3 (25.0%)	4 (33.3%)	2 (14.3%)	-
Diarrhea	3 (18.8%)	-	1 (8.3%)	-	-	1 (7.1%)
Nausea	2 (12.5%)	-	3 (25.0%)	-	3 (21.4%)	-
Vomiting	3 (18.8%)	-	2 (16.7%)	-	3 (21.4%)	-
Asthenia/Fatigue	2 (12.5%)	-	5 (41.7%)	1 (8.3%)	3 (21.4%)	-
Pyrexia/Influenza-like illness	12 (75.0%)	1 (6.3%)	8 (66.7%)	-	12 (85.7%)	-
Transaminase increases (ALT, AST)	2 (12.5%)	3 (18.8%)	1 (8.3%)	1 (8.3%)	2 (14.3%)	2 (14.3%)
Pancreatic enzyme increase (lipase, amylase)	1 (6.3%)	3 (18.8%)	-	-	-	-
Decreased appetite	3 (18.8%)	-	3 (25.0%)	-	-	-
Arthralgia	2 (12.5%)	-	-	-	-	-
Musculoskeletal pain	3 (18.8%)	-	4 (33.3%)	-	-	-
Dizziness	1 (6.3%)	-	1 (8.3%)	-	-	-
Headache	1 (6.3%)	-	1 (8.3%)	-	1 (7.1%)	-
Dyspnea	2 (12.5%)	-	-	-	-	-
Hypotension	2 (12.5%)	-	1 (8.3%)	-	-	-

*Part II: one patient at the highest dose (1x10¹⁵ vp) died from a combination of thrombocytopenia (Grade 4) and enterocolitis (Grade 5)



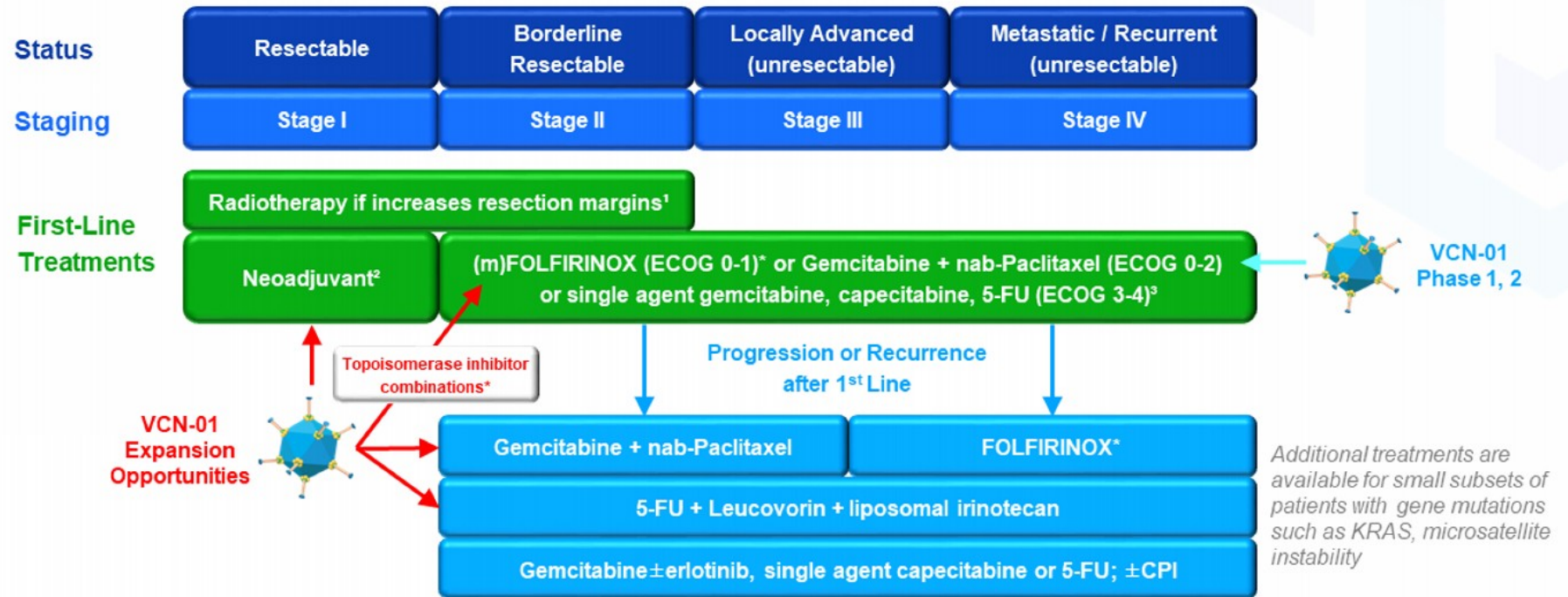
¹Garcia-Carbonero (2022) J Immunother Cancer 10:e003255. ²Concomitant IV VCN-01 3.3x10¹² or 1.0x10¹³ vp/patient administered same day as first dose of SoC IV gemcitabine/nab-paclitaxel. ³Sequential IV VCN-01 3.3x10¹² or 1.0x10¹³ vp/patient administered 7-days prior to first dose of SoC.

VIRAGE PHASE 2 CLINICAL TRIAL DIFFERENTIATORS

- ✓ **First-line** treatment of metastatic PDAC patients
- ✓ **Direct** comparison with standard-of-care chemotherapy in the same trial
- ✓ **Repeated** dosing of VCN-01 may improve treatment outcomes
- ✓ **Open label** provides real-time opportunity to review emerging VCN-01 treatment effects
- ✓ **Orphan Drug Designation** to facilitate regulatory interactions and provide market exclusivity
- ✓ **Fast Track Designation** for more frequent communication with FDA and eligibility for Accelerated Approval and Priority Review

EXPANSION OPPORTUNITIES for VCN-01 in PDAC

Alternate treatment lines and new chemotherapy combinations



VCN-01 WITH GEMCITABINE/ NAB PACLITAXEL

Potential survival benefit compared to all first-line chemotherapy

COMPANY	THERIVA BIOLOGICS (Phase 1)		PDAC FIRST LINE CHEMOTHERAPY			
Virus	VCN-01	VCN-01	
Dose	3.3x10 ¹² vp x1 1x10 ¹³ vp x 1*	1x10 ¹³ vp x 1*
Chemotherapy	Gemcitabine + Nab Paclitaxel	Gemcitabine + Nab Paclitaxel	Gemcitabine + Nab Paclitaxel		FOLFIRINOX	NALIRIFOX
No. Patients	12 (6/dose)	6	431	387	171	383
Response Rate, %	50% [21, 79]	83% [36, 99.6]	29% [25%, 34%]	36.2% [31.4, 41.2]	31.6% [24.7, 39.1]	41.8% [36.8, 46.9]
Progression Free Survival, mos	6.7 [4.5, 11.7]	6.3 [5.7, NE]	5.5 [4.5, 5.9]	5.6 [5.3, 5.8]	6.4 [5.5, 7.2]	7.4 [6.0, 7.7]
12-Mo. Survival, %	66.7%	83.3%	35%	39.5%	48.4%	45.6%
Overall Survival, mos	13.5 [7.1, 29.0]	20.8 [12.2, NE]	8.5 [7.9, 9.5]	9.2 [8.3, 10.6]	11.1 [9.0, 13.1]	11.1 [10.0, 12.1]
	Garcia-Carbonero JTC 10:e003255	Garcia-Carbonero JTC 10:e003255	Von Hoff NEJM 369:1691	Wainberg Lancet 402:1272	Conroy NEJM 364:1817	Wainberg Lancet 402:1272

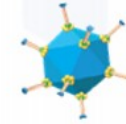


*Approximately 2.2x10¹¹ TCID₅₀ and 7x10¹¹ TCID₅₀ respectively.

THERIVA OV PORTFOLIO HIGHLIGHTS

Unique MOA enables multiple indications and combinations

- **Highly differentiated OV designed to have multiple antitumor effects**
 - Systemic administration, selective tumor replication, stroma degradation
 - Designed to increase cell lysis, tumor immunogenicity, and tumor access by co-administered therapies
- **Multiple potential value opportunities for VCN-01**
 - Encouraging clinical data in PDAC, HNSCC, and retinoblastoma support VCN-01 MOA and safety profile
 - Phase 1 clinical data suggest potential to improve/enable use of immune CPIs in refractory patients
 - Phase 1 clinical data support the feasibility of combining VCN-01 with CAR-T cells in solid tumor patients
- **Regulatory status expected to facilitate VCN-01 development**
 - PDAC: Orphan Drug Designation (FDA, EMA), Fast Track designation (FDA)
 - Retinoblastoma: Orphan Drug Designation (FDA); Rare Pediatric Disease Designation (FDA: access to priority review voucher)
- **Leading OV discovery engine advancing diverse new product candidates**
 - Potent tumor killing with potential single agent efficacy



CPI immune checkpoint inhibitor. PDAC pancreatic ductal adenocarcinoma. R/M HNSCC refractory/metastatic head and neck squamous cell carcinoma. SoC standard of care (gemcitabine + nab-paclitaxel in PDAC). Alternate indications include retinoblastoma and brain tumors; phase 1 data support additional evaluation in colorectal cancer. Topoisomerase inhibitors include topotecan and irinotecan.

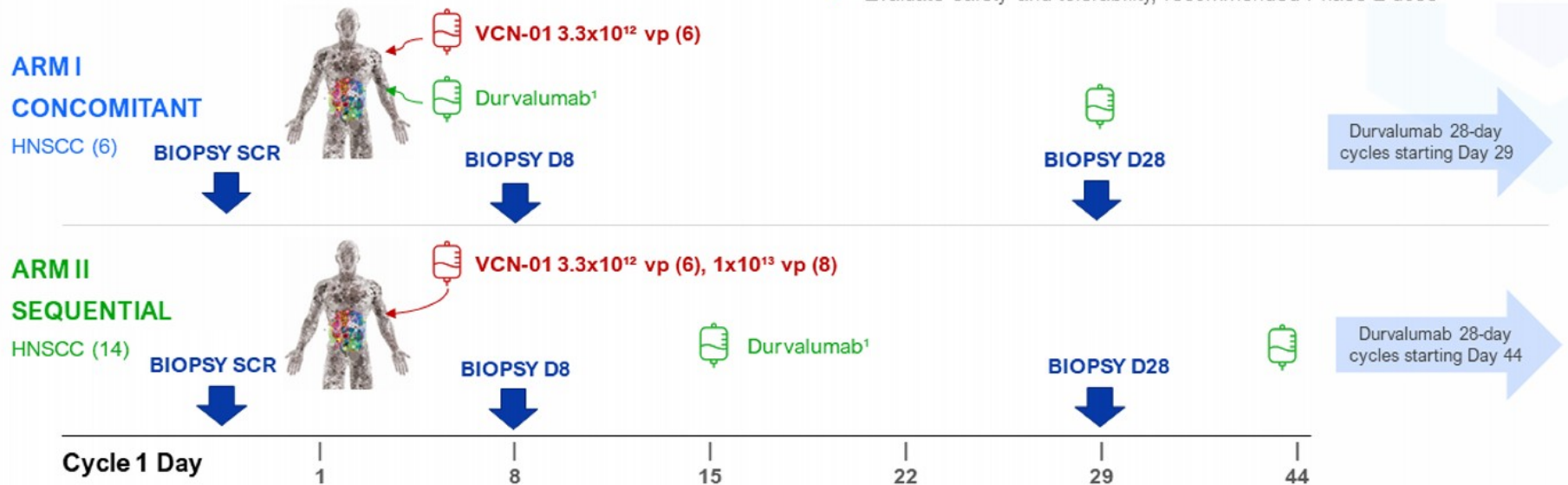
VCN-01 IN HEAD & NECK CANCER



VCN-01 IV + ANTI-PD-L1 PHASE 1 TRIAL in HNSCC

Multicenter, open-label, dose escalation study (NCT03799744)

- ✓ Single IV doses of VCN-01 combined with anti-PD-L1
- ✓ Patients with metastatic squamous cell carcinoma of the head & neck previously **REFRACTORY** to anti-PD(L)1 treatment (R/M HNSCC)
- ✓ Evaluate safety and tolerability, recommended Phase 2 dose



EXTENDED SURVIVAL with VCN-01+DURVALUMAB

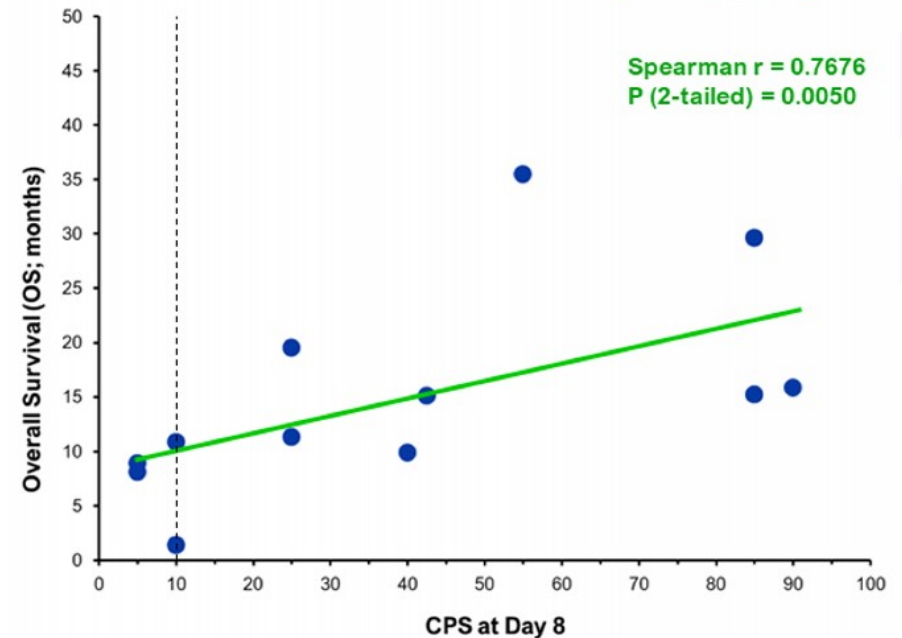
Survival correlated with PD-L1 upregulation after VCN-01 treatment

- Higher than expected survival (OS) despite previous anti-PD(L)1 failure

Regimen	Median OS (95% CI), mos	
	3.3x10 ¹² vp	1.0x10 ¹³ vp
Concomitant	10.4 (8.9-NE)	..
Sequential	15.5 (15.1-NE)	17.3 (11.3-NE)

- No correlation of survival with baseline tumor PD-L1 expression (CPS) BUT significant correlation of survival with CPS 8-days after VCN-01 treatment

Overall Survival vs CPS in Biopsies at Day 8



VCN-01 MAY SENSITIZE PATIENTS TO SUBSEQUENT THERAPY

Patients responded to subsequent chemotherapy after progressing with VCN-01 + durvalumab

ARM	ICI Treatment Progression (Pre-trial)		Current Trial			1st Line after Current Trial	2nd Line after Current Trial
	Median OS post-1st ICI		ORR	Median PFS	Median OS	ORR	ORR
Concomitant Low (3.3E12vp)	21.6 (19.2-NE)		0/6	1.7 (1.6-NE)	10.4 (8.9-NE)	3/5	1/2
Sequential Low (3.3E12vp)	23.9 (16.6-NE)		1/6	3.7 (2.2-NE)	15.5 (15.1-NE)	3/6	1/6
Sequential High (1E13vp)	21.8 (12.9-NE)		0/6	2.1 (1.4-NE)	17.3 (11.3-NE)	2*/5	1/4

*Complete Responses

AE PROFILE FOR THE COMBINATION OF VCN-01 AND DURVALUMAB

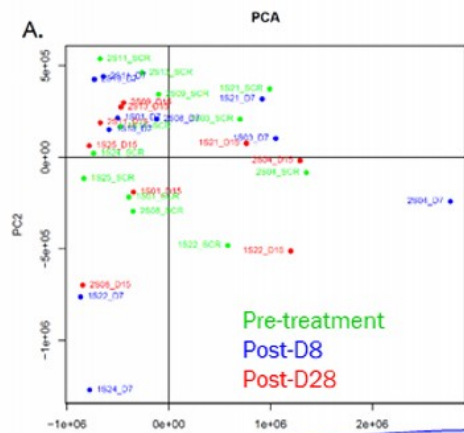
Most common AEs related to IV VCN-01 [NCT03799744]

Adverse Reactions	Arm I - Concomitant (Dose 3,3E12 , n=6) ²		Arm II - Sequential (Dose 3,3E12 , n=6) ³		Arm II - Sequential (Dose 1E13 , n=8) ³		
	CTCAE Grade	Grade 1-2	Grade ≥3	Grade 1-2	Grade ≥3	Grade 1-2	Grade ≥3
Pyrexia		2 (33,0%)	-	5 (62,5%)	-	3 (50%)	-
Influenza like illness		3 (50,0%)	-	3 (37,5%)	2(25%)	2 (33,3%)	-
Asthenia/Fatigue		2 (33,0%)	-	2 (25%)	1 (12,5%)	4 (66,6%)	-
AST increased		4 (66,7%)	1 (16,6%)	2 (25%)	-	-	2 (33,3%)
ALT increased		3 (50,0%)	1 (16,6%)	1 (12,5%)	-	-	2 (33,3%)
Decreased Appetite		1 (16,6%)	-	3(37,5%)	-	2 (33,3%)	-
Lymphocyte count decreased		1 (16,6%)	-	-	3 (37,5%)	-	-
Myalgia		-	-	3(37,5%)	-	1 (16,6%)	-
Hypotension		-	-	2 (25%)	-	1 (16,6%)	-
Chills		1 (16,6%)	-	1(12,5%)	-	1 (16,6%)	-
Vomiting		1 (16,6%)	-	1(12,5%)	-	1 (16,6%)	-
Anemia		2 (33,0%)	-	1(12,5%)	-	1 (16,6%)	-
Nausea		-	-	1(12,5%)	-	1 (16,6%)	-
Headache		-	-	1(12,5%)	-	1 (16,6%)	-
Erythema		1 (16,6%)	-	1(12,5%)	-	-	-
Guillain-Barre Syndrome		-	1 (16,6%)	-	1 (12,5%)	-	-
Hepatic enzymes increased		-	-	-	1 (12,5%)	-	-
GGT Increased		-	-	-	-	-	1 (12,5%)

VCN-01 INDUCES TRANSCRIPTOMIC CHANGES in TUMOR MICROENVIRONMENT

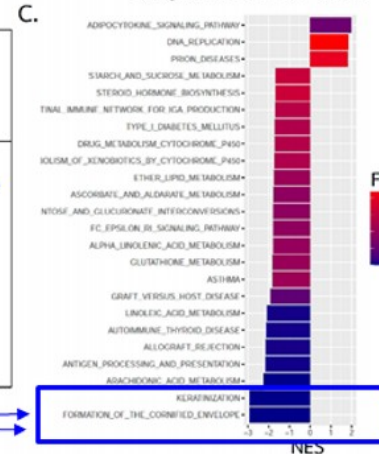
RNAseq Analysis in Clinical Samples from HNSCC Patients [NCT03799744]

Principal Component Analysis¹
including all the Pre- and Post-treatment samples



Most significant Reactome and KEGG pathways in GSEA (Gene Set Enrichment Analysis)¹

analysis D8 vs Pre-treat.



analysis D28 vs Pre-treat.



B.

Gene	Gene product	log2FoldChange	padj	
CIDEA	Cell Death Inducing DFFA Like Effector C	5.524	0.047	Up D7
TAGLN3	Transgelin 3	1.615	0.037	
GPR3	G Protein-Coupled Receptor 3	1.157	0.039	
ABP1	Auxin-binding protein 1	-1.636	0.013	
CD207	CD207 Molecule	-1.960	0.044	Down D7
MEGF10	Multiple EGF Like Domains 10	-2.008	0.043	
OSTalpha	Organic solute transporter alpha	-2.179	0.039	
CD1E	CD1E Molecule	-2.316	0.010	
ATP10B	ATPase Phospholipid Transporting 10B	-2.556	0.044	
FCER1A	Fc Epsilon Receptor 1a	-2.687	0.006	
LOC285629	-2.818	0.001	
LCE1B	Late Corneal Envelope 1B	-7.921	0.006	

Sustained differential gene expression profiles associated with downregulation of matrix-related pathways

VCN-01 FINDINGS in R/M HNSCC

Data support VCN-01 MOA and immune enhancing effects

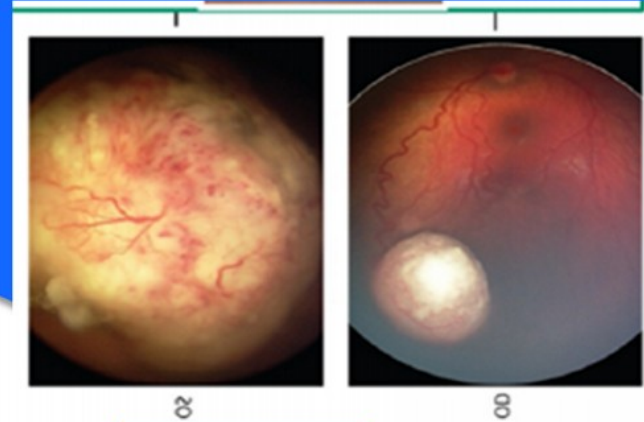
- VCN-01 has an acceptable adverse event profile when administered prior to durvalumab (Imfinzi®)
- VCN-01 reaches tumors, has sustained replication and PH20 expression
- VCN-01 treatment led to downregulation of tumor matrix genes and increased levels of immune markers in tumor biopsies (CD8, PD-L1, IDO)
- VCN-01-treated patients showed **increased response** to subsequent chemotherapy treatment lines after progressing on this trial

VCN-01 IN RETINOBLASTOMA



RETINOBLASTOMA, A RARE PEDIATRIC MALIGNANCY

- Retinoblastoma (Rb) is an orphan indication that accounts for ~2-3% of all childhood cancers¹
- 200-300 cases each year in the USA, EU²⁻⁴
- VCN-01 selectivity enables development as an intravitreal treatment for Rb patients
- VCN-01 could be used in combination with chemotherapy to potentially improve outcomes
- VCN-01 could be used as a rescue therapy for patients who fail standard therapy

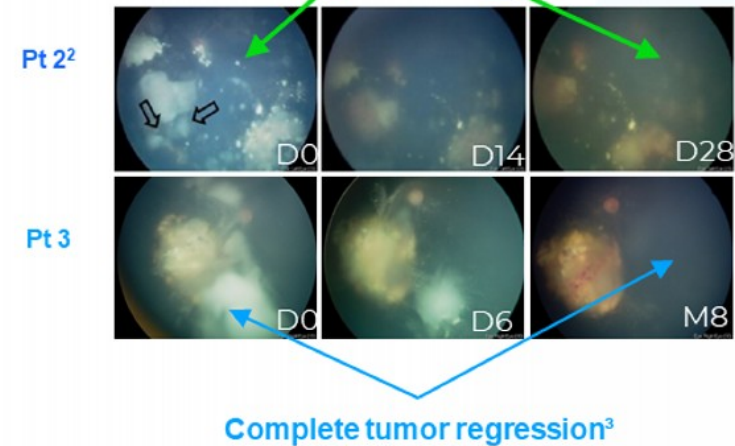


VCN-01 IN RETINOBLASTOMA

- Single center, open-label, dose escalation study of intravitreal (IVit) VCN-01¹⁻³
 - Children aged 1-12 years (n=9)
 - Retinoblastoma that is recurrent or refractory to chemotherapy and for whom enucleation is the best treatment option
 - VCN-01 doses of 2.0×10^9 vp per eye (n=1) or 2.0×10^{10} vp per eye (n=8) on days 1 and 15
- Promising antitumor activity and appropriate adverse event profile and tolerability at RP2D
 - Reduction of vitreous seeds in 4 patients of 9 evaluable patients
 - Enucleation avoided in 3 patients; low VCN-01 dose and/or damage from prior chemotherapy meant the eye could not be saved in 6 patients
- Earlier VCN-01 intervention anticipated to have better outcomes

Promising Results in Patients Treated with High Dose VCN-01

Reduced number and size of tumor vitreous seeds following VCN-01 administration²



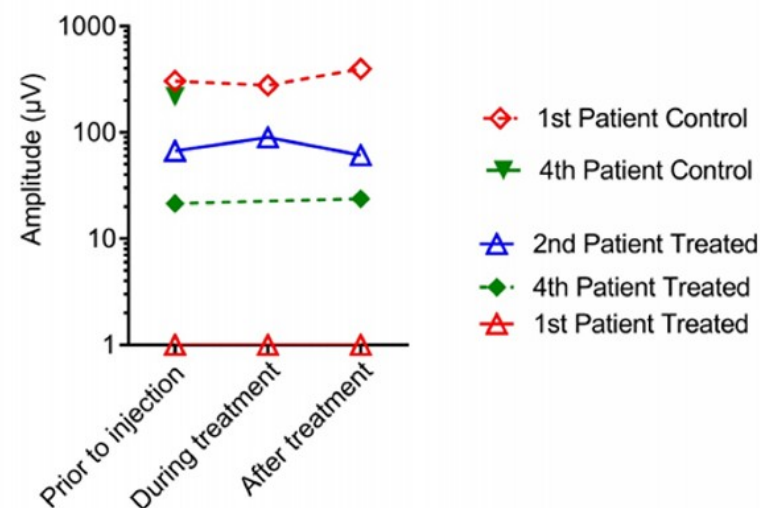
INTERIM ADVERSE EVENT DATA FOR INTRAVITREAL VCN-01

Two Intravitreal VCN-01 Doses of 2.0×10^9 or 2.0×10^{10} vp per eye¹

Adverse Reaction	Pts	All Grades		Grade ≥ 3	
CTCAE grade	N	n	%	n	%
Uveitis	6	2	33%	2	33%
Eye oedema	6	1	17%	0	0%
Conjunctival hyperemia	6	1	17%	0	0%
Eye inflammation	6	1	17%	0	0%

- VCN-01 was reasonably well tolerated after intravitreal administration², although some turbidity and vitritis associated with intravitreal inflammation was observed
- Intravitreal inflammation was managed with local and systemic administration of anti-inflammatory drugs
- VCN-01 did not appear to change retinal function (electroretinographic signaling)
- VCN-01 does not replicate in healthy retinal tissue of patients with either somatic or germline Rb mutation³

Stable Electroretinographic Signals



VCN-01 DEVELOPMENT IN RETINOBLASTOMA

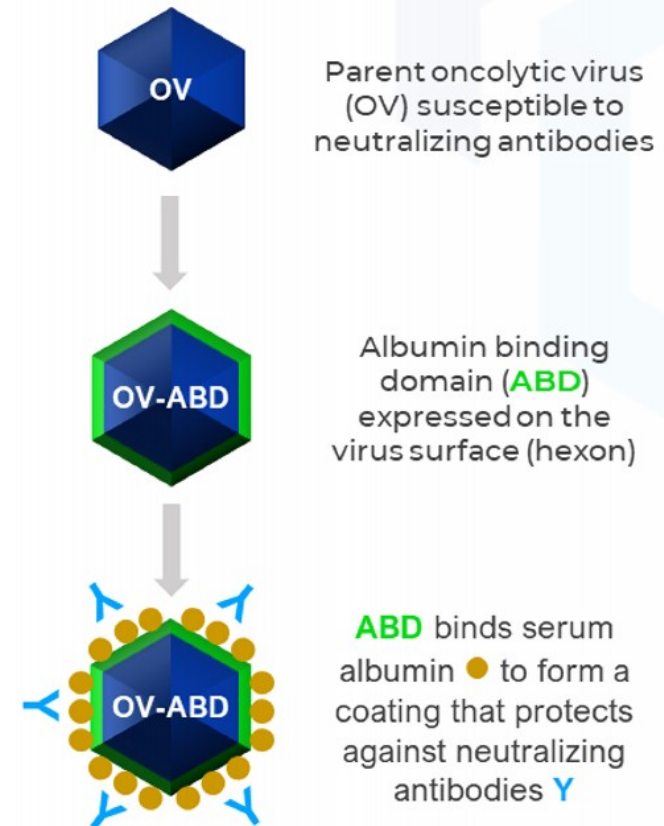
- Phase 1 ISS Completed H1 2024
 - Initial data demonstrate acceptable adverse event profile and one durable complete response
- Developing a clinical protocol for an open-label, multinational study
 - Retinoblastoma patients with vitreous seeds
 - IVit VCN-01 in combination with chemotherapy (no defined SoC)
 - PI Dr. Guillermo Chantada, MD PhD¹
- Status
 - US Orphan Drug Designation (EU application in process)
 - Pre-IND meeting with FDA completed Q4 2023
 - Rare Pediatric Disease Designation (potential eligibility for Priority Review Voucher)

VCN-X NEXT GENERATION
OV DISCOVERY PLATFORM



ALBUMIN SHIELD™ to ENHANCE OV SYSTEMIC DELIVERY

- Albumin Shield technology protects OVs as they travel to tumors after systemic administration^{1,2}
- Albumin Shield modified OVs bind albumin in the patient's blood to form a protective coating
- Albumin Shield genetic modification allows parent and progeny virus to be albumin coated
- Albumin Shield may enable [multiple IV administrations](#) for hard-to-treat patients
- Albumin Shield first candidate VCN-11 is being prepared for a potential Phase 1 clinical trial



THERIVA OV PIPELINE DISCOVERY AND DEVELOPMENT

Advancing founders' decade of world leading OV innovation

Common Features

Clinically-tested Adenovirus Expressing PH20
Hyaluronidase to Degrade Stroma

+

Albumin Shield™ To Prevent Neutralization by
anti-viral Antibodies and Facilitate IV Multidosing

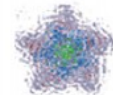
+

Unique Multifunctional Proteins to Turn Cold Tumors
Hot and Enhance Anti-tumor Immune Response

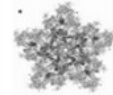
Product Specific Features



VCN-11 Hyaluronidase alone



VCN-12 Hyaluronidase + Toxins



VCN-13 Fusion Hyaluronidase + scPD-L1



VCN-XX Hyaluronidase + other payloads

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