



Corporate Overview

April 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 H1 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 oncolytic virus discovery engine enable development of a distinct product pipeline

Financial Snapshot

Exchange	NYSE American
Ticker	TOVX
Cash (12/31/2023)	\$23.2M
Projected cash runway	Q1 2025
Average Daily Volume (3M Ave)	100.7K
Locations	Rockville, MD Barcelona, Spain

THERIVA PIPELINE

Candidate	Target	Pre-IND	Phase 1	Phase 2	Collaborators	Status*
VCN-01 Selective, Stroma Degrading OV	Pancreatic Cancer (IV) with gemcitabine/nab-paclitaxel					Phase 2 Study On-going (ODD EU, US)
	Retinoblastoma (IVit)					Phase1 Study On-going (ODD US)
	HNSCC (IV) + durvalumab					Phase 1 Treatment Complete Survival Follow-up On-going
	Solid Tumors – Brain, Ovarian, PDAC (IV)					Phase 1 Studies On-going
VCN-11 Albumin Shield OV	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 [3] Oral IAP	Potential indications include NAFLD/NASH, celiac, radiation enteritis					Phase 1 Studies Complete

VCN-01 IS A UNIQUELY ENGINEERED HUMAN ADENOVIRUS 5

Systemic

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

Selective

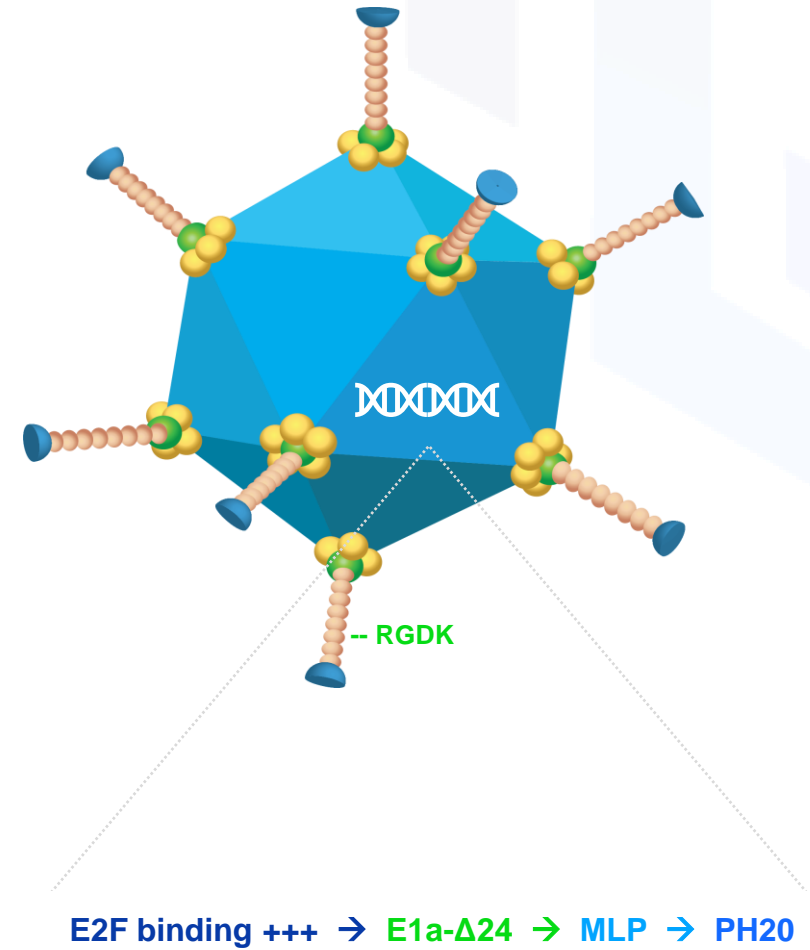
Replicates only in **tumor** cells
Liver detargeted

Stroma Degrading

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

Self Reporting

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



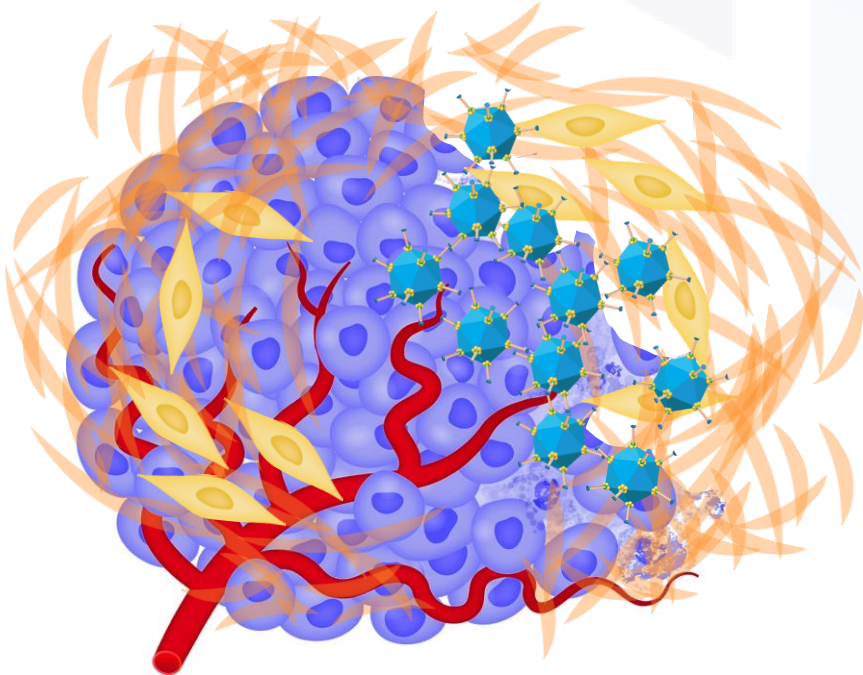
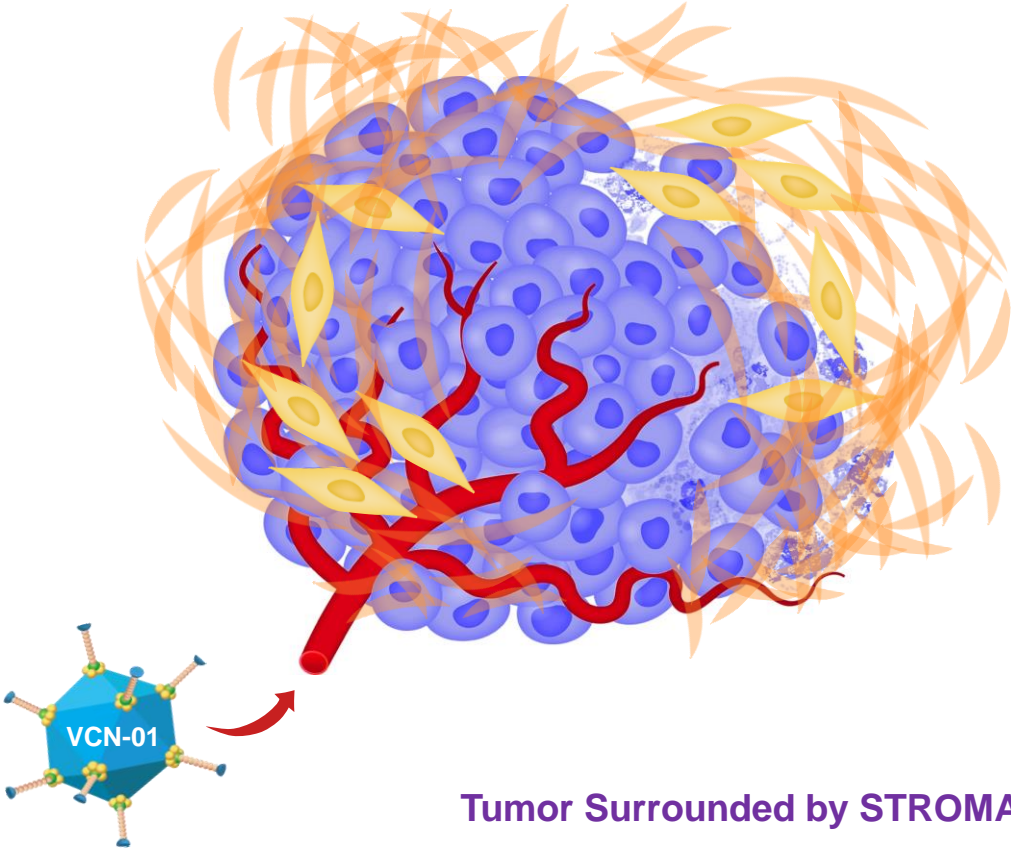
VCN-01 IS A UNIQUELY ENGINEERED HUMAN ADENOVIRUS 5

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



Stroma



Cancer Associated Fibroblast

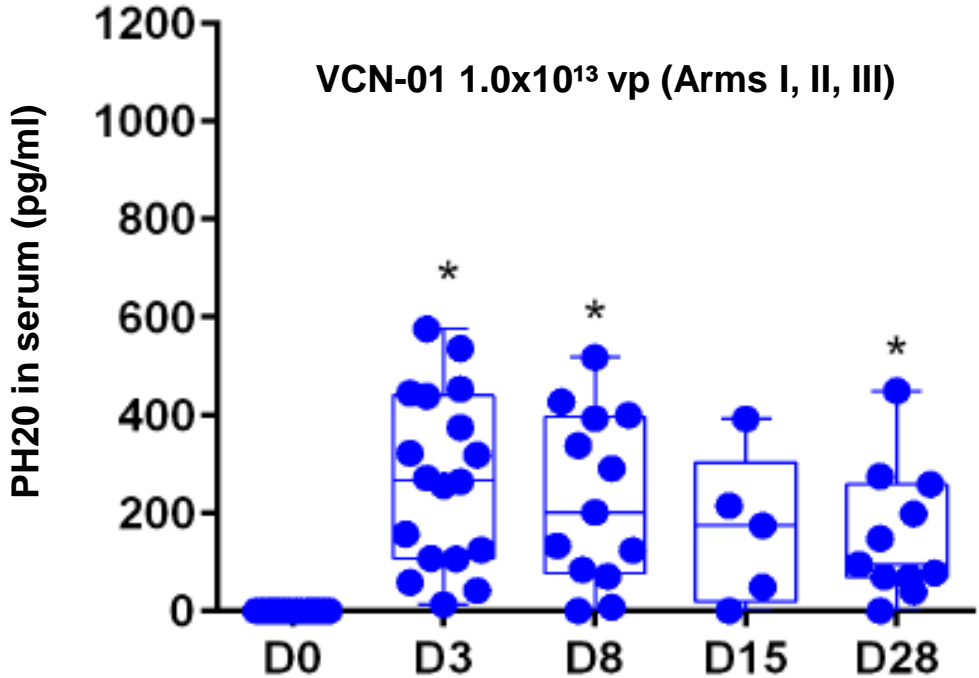
VCN-01 COMPARED TO OTHER ONCOLYTIC VIRUSES

COMPANY	THERIVA BIOLOGICS	CG ONCOLOGY	GENELUX	ONCOLYTICS BIOTECH	REPLIMMUNE
Ticker	NYSE MKT: TOVX	NASDAQ: CGON	NASDAQ: GNLX	NASDAQ: ONCY	NASDAQ: REPL
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 (2/3)
Route	IV	IVESIC	IP	IV	IT
Dose	1x10 ¹³ vp*	1x10 ¹² vp	3x10 ⁹ pfu	4.5x10 ¹⁰ TCID ₅₀	1x10 ⁷ pfu/mL
Stroma Degrading	Yes	No	No	No	No
Biomarker	PH20	..	Beta-GAL, beta-GLU, GFP

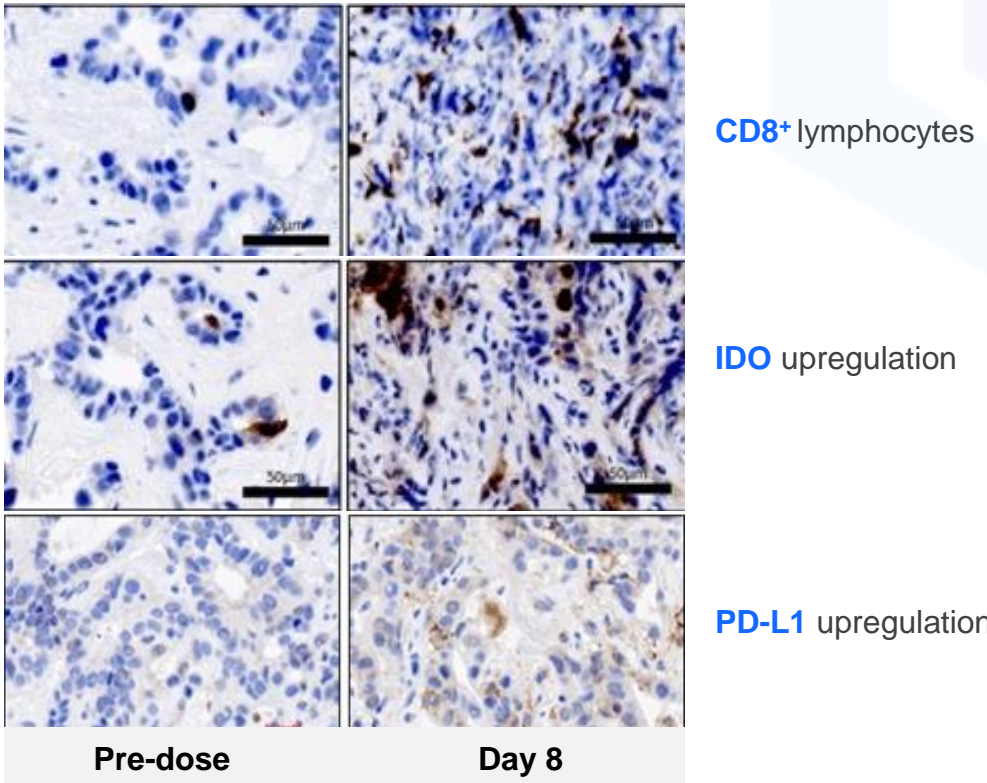
CLINICAL DATA SUPPORT VCN-01 MODE-OF-ACTION

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

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

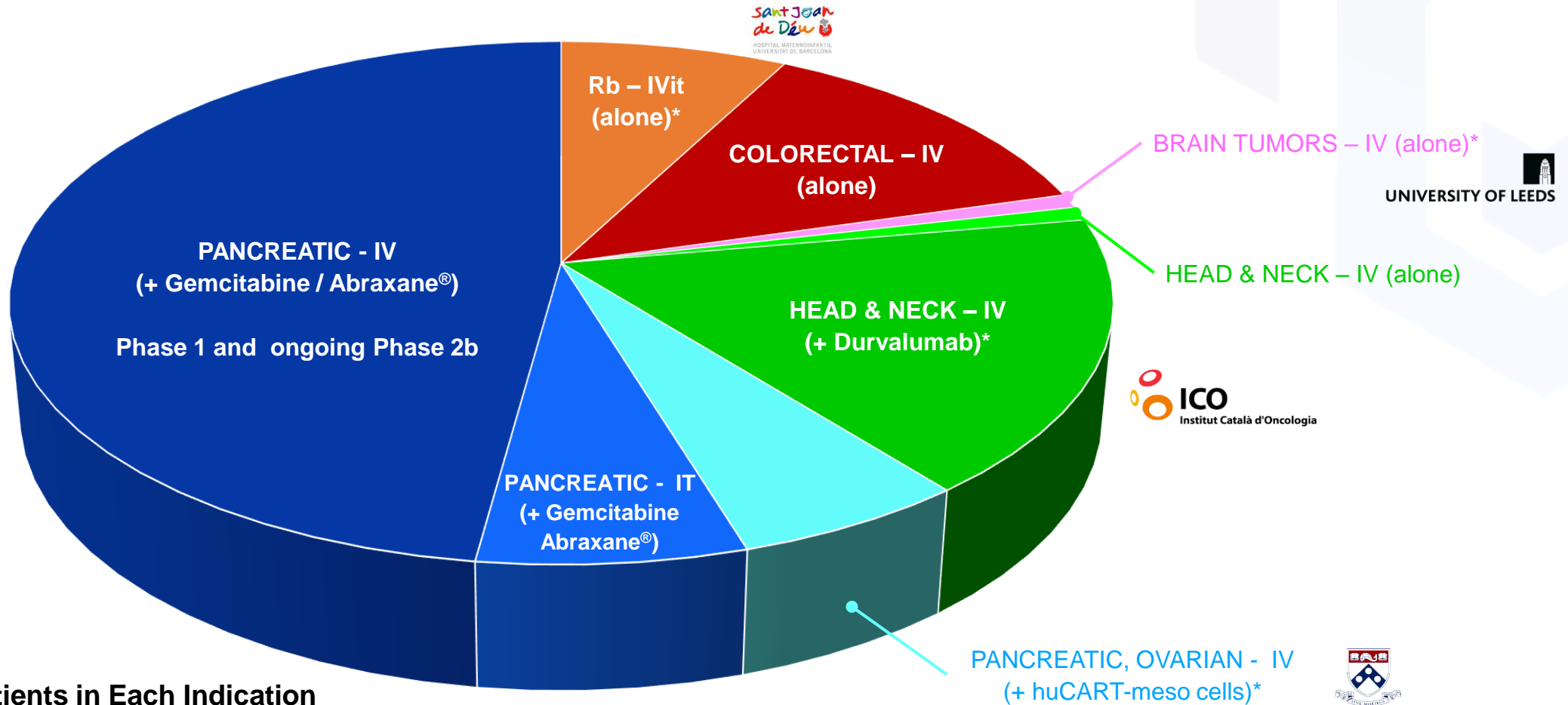


Immune markers upregulated in biopsies of **hepatic metastases**



VCN-01 EXTENSIVE CLINICAL PROGRAM

116 patients treated to date in multiple indications and combinations

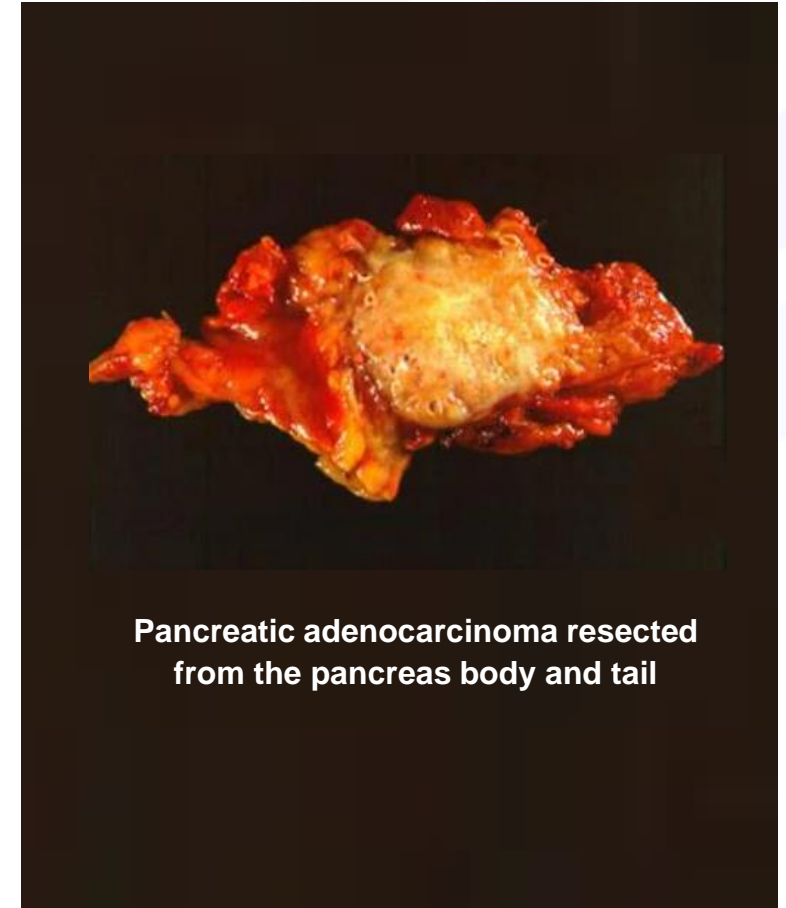


Number of Patients in Each Indication

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 disease¹
 - USA est. 62,210 new cases and 49,830 deaths in 2022^{2,3}
- **Hyaluronic acid** in stroma is associated with reduced treatment efficacy and poor prognosis⁴
 - VCN-01 designed to degrade hyaluronic acid
- Incidence is growing worldwide
 - Estimated treatment market ~\$2.5B (2022) ~\$7.0B (2030)⁵

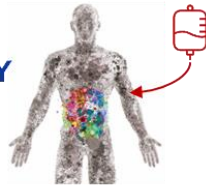


VCN-01 FAVORABLE OUTCOMES IN PHASE 1 TRIAL

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

ARM I MONOTHERAPY

Solid tumors (16)

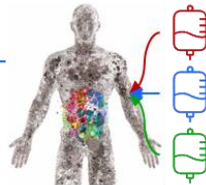


VCN-01 Dose Finding¹

- ✓ Single IV doses of VCN-01 alone or with standard-of-care (SoC) chemotherapy gemcitabine/nab-paclitaxel (Abraxane®)
- ✓ Evaluate safety and tolerability, recommended Phase 2 dose

ARM II CONCOMITANT

PDAC (12)



VCN-01²

nab-Paclitaxel³

Gemcitabine⁴



SoC chemotherapy 28-day cycles starting Day 29

ARM III SEQUENTIAL

PDAC (14)



VCN-01⁵

nab-Paclitaxel³

Gemcitabine⁴



SoC chemotherapy 28-day cycles starting Day 36

Cycle 1 Day

1

8

15

22

29

36

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.

VCN-01 FAVORABLE OUTCOMES IN PHASE 1 TRIAL

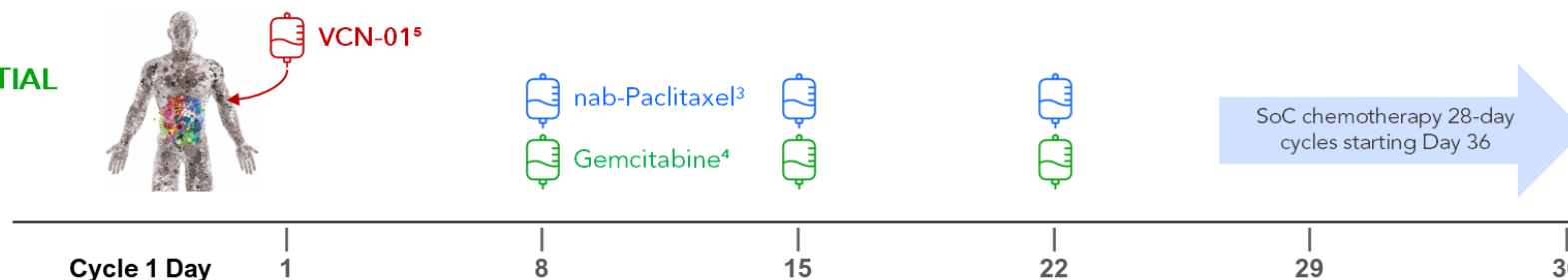
Multicenter, open-label, dose escalation study (NCT02045602)

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	6.7	7.2	5.5
Survival ≥12 months	.	.	67%	35%

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%)	-

KOLs advise that OS ≥15 months is a significant patient outcome

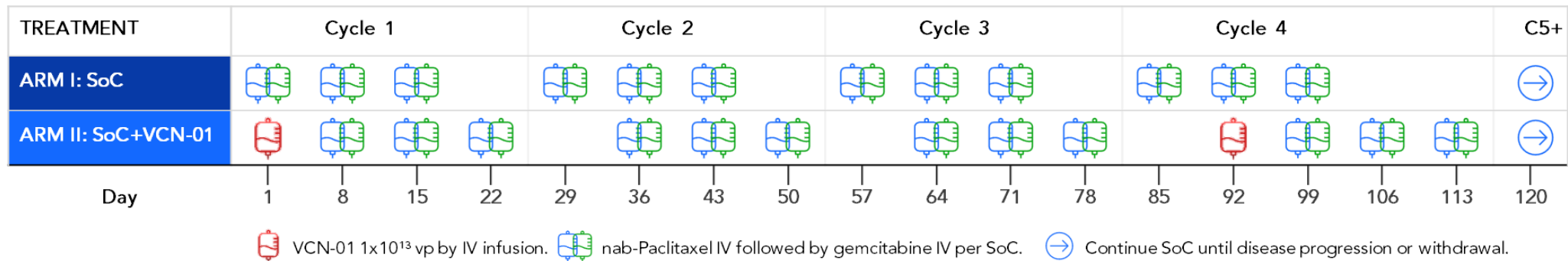
ARM III
SEQUENTIAL
PDAC (14)



VIRAGE PHASE 2B CLINICAL TRIAL in PDAC

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

- Study on-going in patients with **first-line** metastatic pancreatic ductal adenocarcinoma (PDAC)
- Up to 92 patients to be 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



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

VCN-01 DEVELOPMENT IN PANCREATIC CANCER

- Project complete enrollment into the VIRAGE Study H1 2024
 - First DMC safety review completed Q1 2024
 - No safety concerns were raised and no protocol amendments were requested
- Potential interim data analysis H2 2024
 - Opportunity to discuss pivotal study design and potential expedited status with FDA and EMA
- Evaluating potential expansion opportunities in pancreatic cancer¹
 - Combination with FOLFIRINOX or NALIRIFOX²

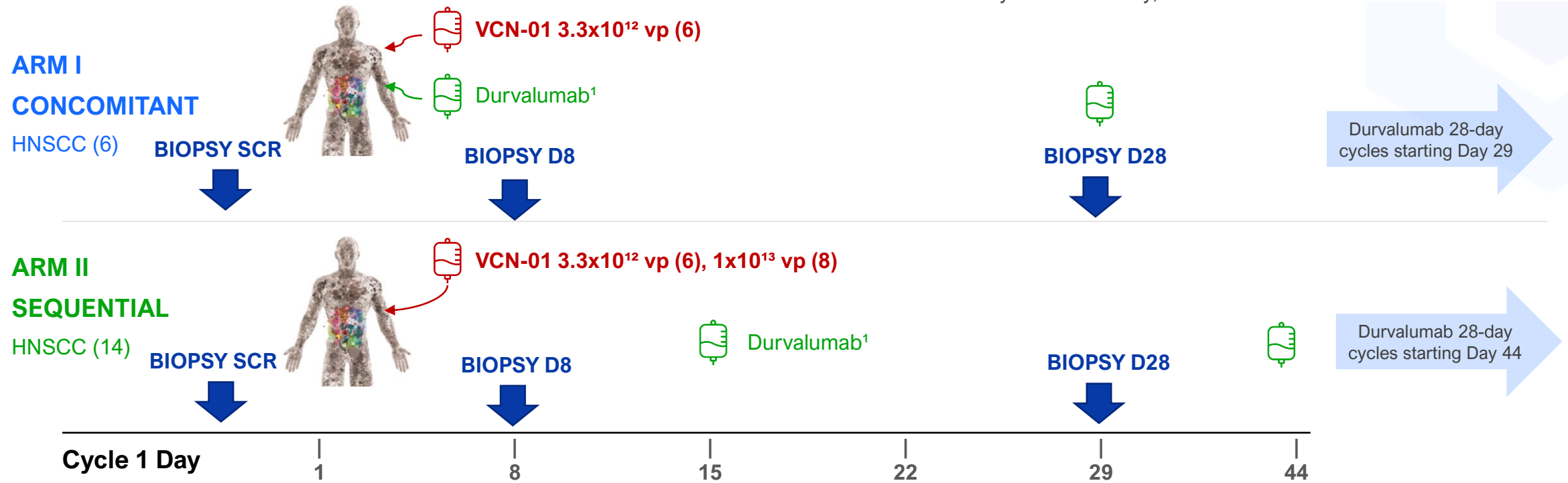
VCN-01 IN HEAD & NECK CANCER



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

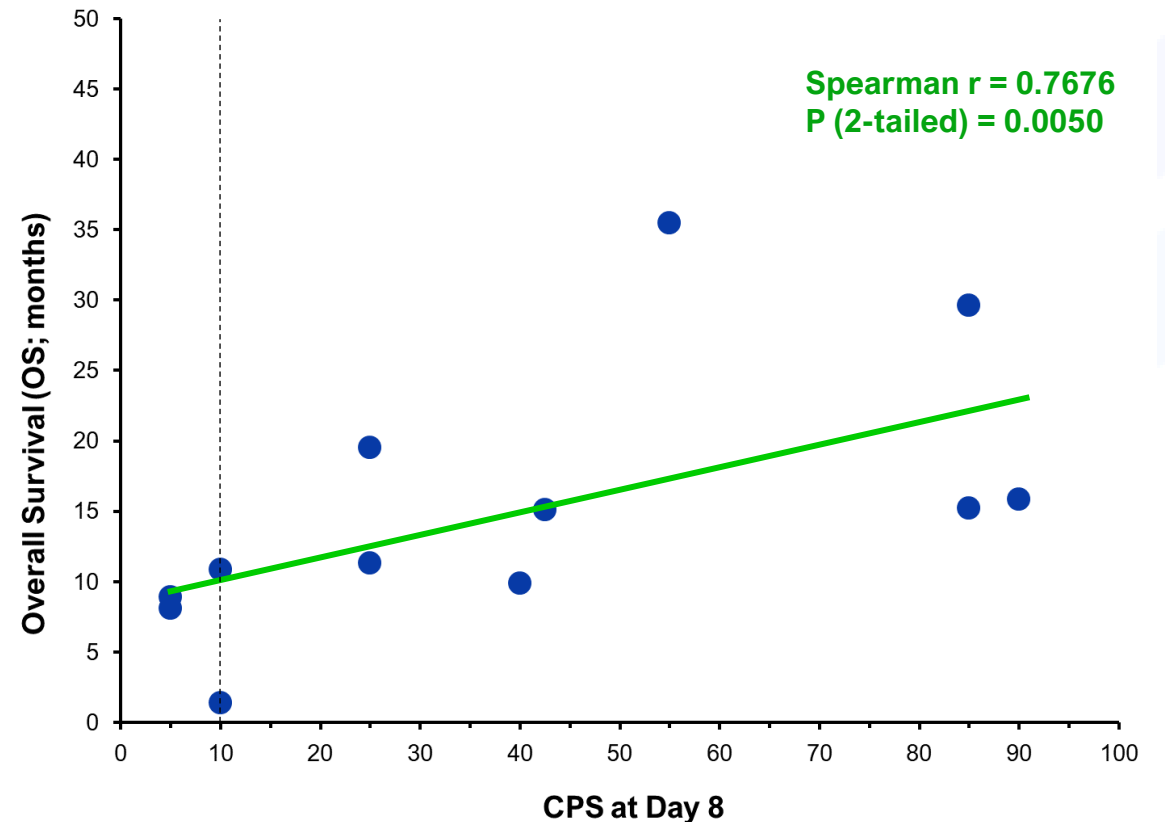
Survival correlated with PD-L1 upregulation after VCN-01 treatment

- 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) Median OS post-1st ICI	<u>Current Trial</u>			1st Line after Current Trial	2nd Line after Current Trial
		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

VCN-01 FINDINGS in R/M HNSCC

Data support VCN-01 MOA and immune enhancing effects

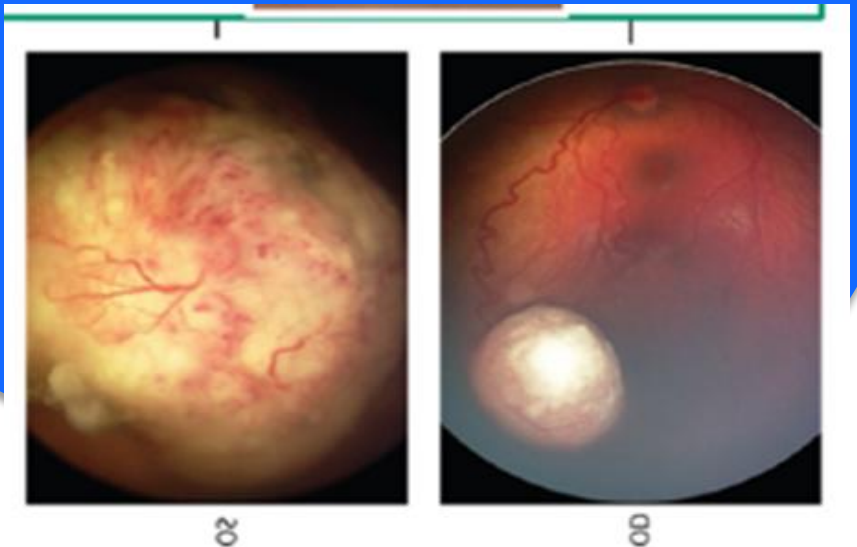
- VCN-01 has an acceptable safety profile when administered prior to durvalumab (Imfinzi®)
- VCN-01 reaches tumors, has sustained replication and PH20 expression
- VCN-01 treatment led to downregulation of tumor matrix genes and increased levels of immune markers in tumor biopsies (CD8, PD-L1, IDO)
- 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

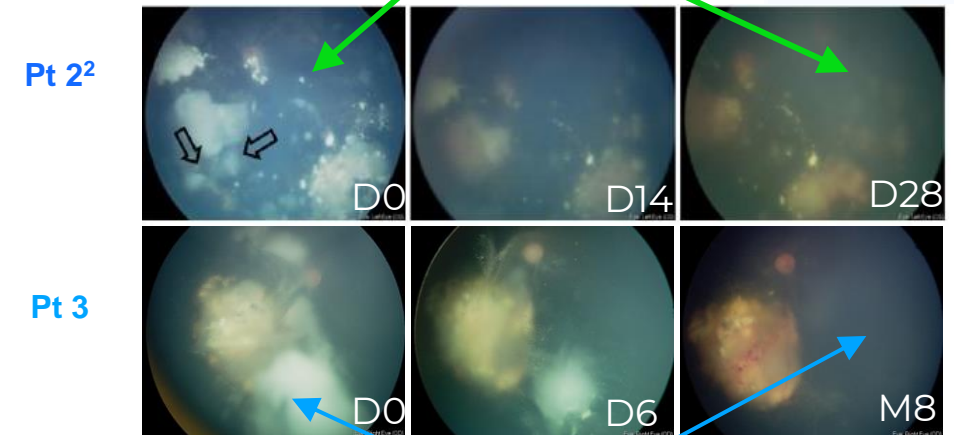


VCN-01 IN RETINOBLASTOMA

- On-going 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 safety 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



Complete tumor regression³

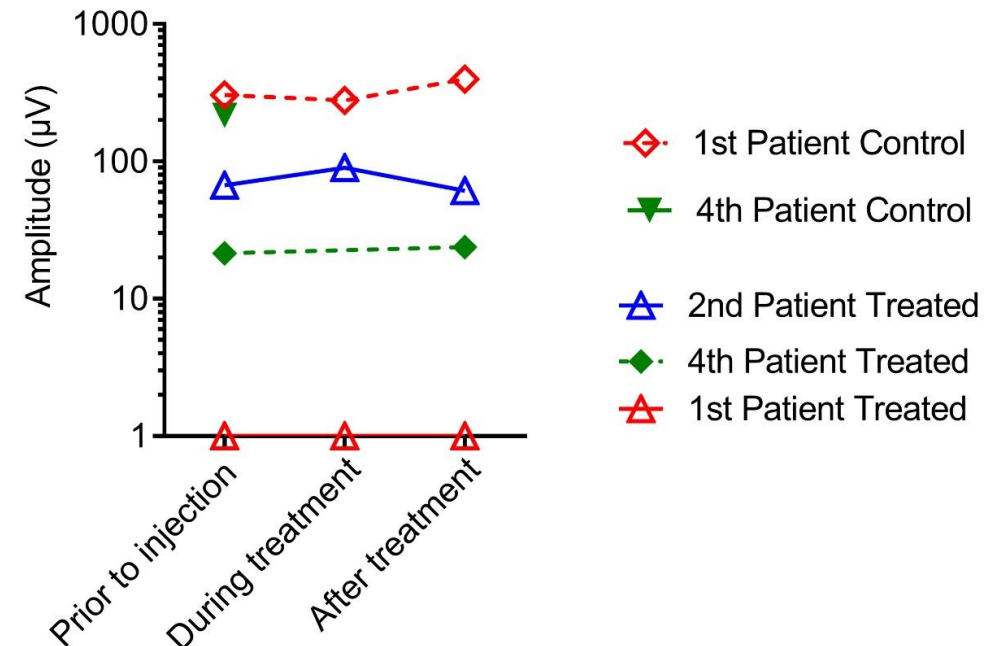
INTERIM SAFETY DATA FOR INTRAVITREAL VCN-01

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

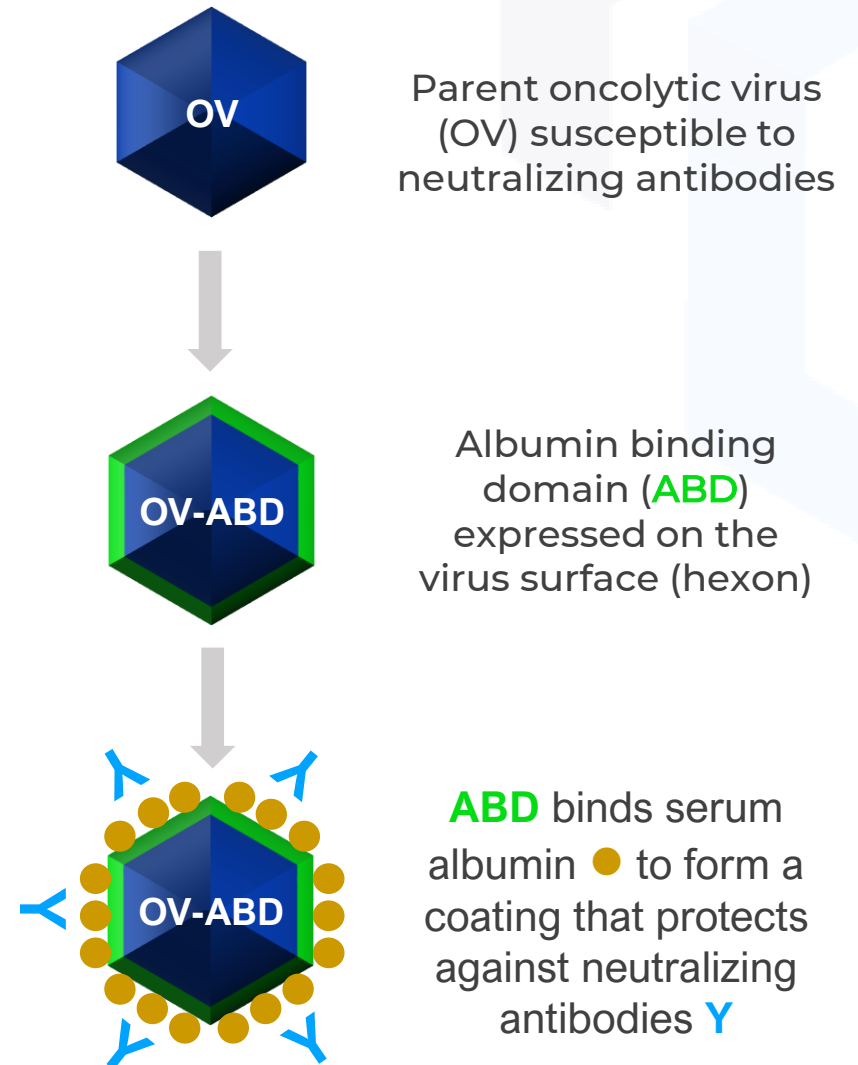
- Phase 1 ISS to Complete in H1 2024
 - Enrollment completed and last patients in the follow-up period
 - Initial data demonstrate acceptable safety and one durable complete response
- Developing a clinical protocol for an open-label, multinational study
 - Retinoblastoma patients with vitreous seeds
 - 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
 - Potential to apply for expedited status and Rare Pediatric Disease Designation

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

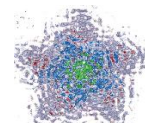
+

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

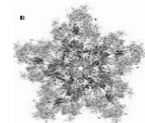
Product Specific Features



VCN-11 Hyaluronidase alone



VCN-12 Hyaluronidase + Toxins



VCN-13 Fusion Hyaluronidase + scPD-L1



VCN-XX Hyaluronidase + other payloads

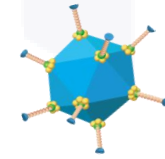


SUMMARY



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
- Clinical data in different indications highlight 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
 - Orphan Drug Designation in PDAC and retinoblastoma
 - Opportunity to apply for expedited status and/or Rare Pediatric Disease Designation (access to priority review voucher)
- Leading OV discovery engine advancing diverse new product candidates
 - Potent tumor killing with potential single agent efficacy

ACHIEVEMENTS AND PROJECTED MILESTONES

- **VCN-01 PDAC**
Positive DMC safety review ✓
- **VCN-01 RETINOBLASTOMA**
Patient dosing completed ✓
- **VCN-01 PDAC**
Phase 2 enrollment complete
ASGCT poster (preclinical)¹
ASCO poster (trial design)
- **VCN-01 RETINOBLASTOMA**
Patient follow-up complete
- **SYN-004 aGVHD**
Phase 1b/2a data 2nd cohort
- **VCN-01 PDAC**
Potential interim data analysis (Q3/Q4)
- **VCN OV DISCOVERY**
VCN-12 candidate selection²
- **SYN-004 aGVHD**
Phase 1b/2a potential initiation of 3rd cohort (data est. H1 2025)
- **VCN-01 PDAC**
Potential FDA and EMA discussions of pivotal design
- **VCN-01 RETINOBLASTOMA**
SIOP presentation (Phase 1)³
Finalize Phase 2 study design
- **VCN-01 + CAR-T**
Anticipate additional data presentations by U. Penn

Q1 2024

Q2 2024

Q3 2024

Q4 2024

FINANCIAL AND INVESTMENT GOALS

- Immediate Goals
 - Augment share register with additional sophisticated institutional investors
 - Establish a realistic share price/valuation to facilitate financial options
 - External validation through a funded discovery or development partnership
- Mid- to Longer-Term Goals
 - Institutional investment to enable registration studies in lead indication(s)
 - Financing or partnering to advance OV pipeline and explore additional indications

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



APPENDIX



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)			

EXTENSIVE VCN-01 PHASE 1 CLINICAL EXPERIENCE

87 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	Ongoing; partial data available	NCT03284268
Institut Català d'Oncologia (ICO)	1	HNSCC	Durvalumab	IV	Treatment complete; Initial data H2'22	NCT03799744
U. Leeds	1	Adult brain tumors	None	IV	Ongoing	ISRCTN 517624869
U. Pennsylvania	1	PDAC, Ovarian	huCART-meso ³	IV	Ongoing	NCT05057715

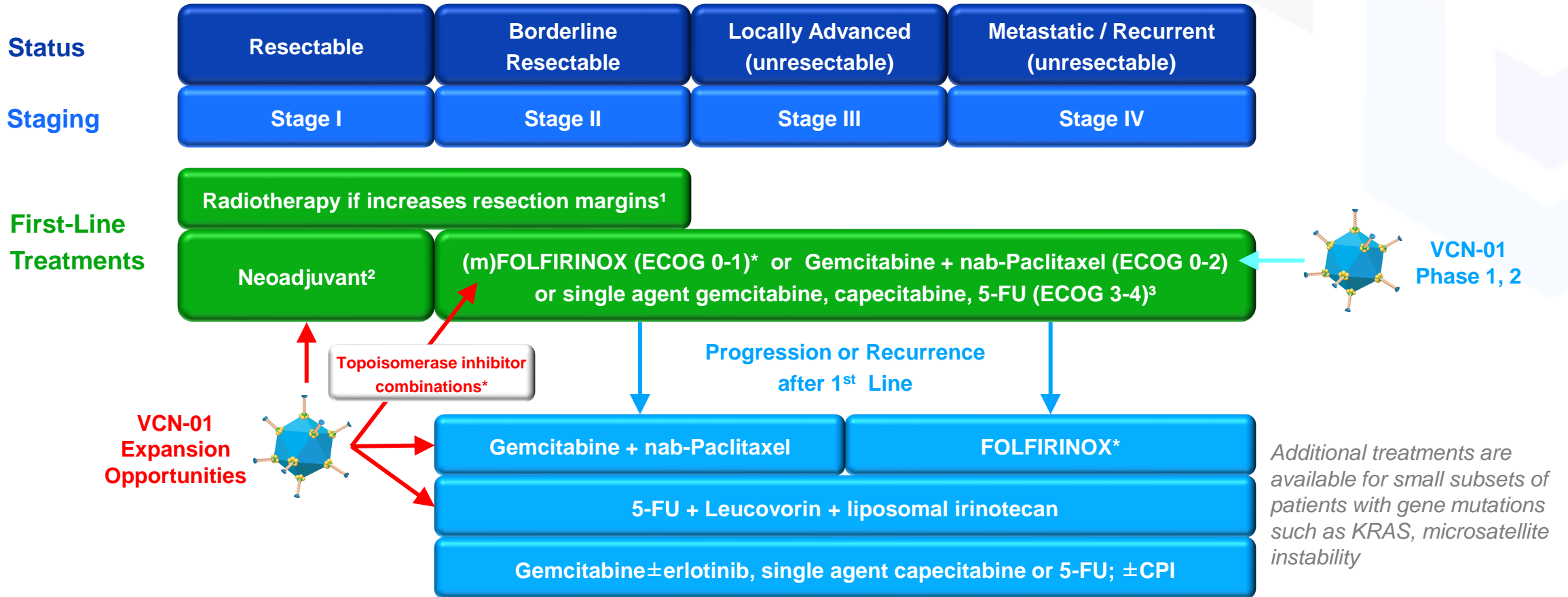
MOST COMMON IV VCN-01 RELATED AEs [NCT02045602]

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 (1x10¹³ vp) died from a combination of thrombocytopenia (Grade 4) and enterocolitis (Grade 5)

EXPANSION OPPORTUNITIES for VCN-01 in PDAC

Alternate treatment lines and new chemotherapy combinations



AE PROFILE FOR THE COMBINATION OF VCN-01 AND DURVALUMAB

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

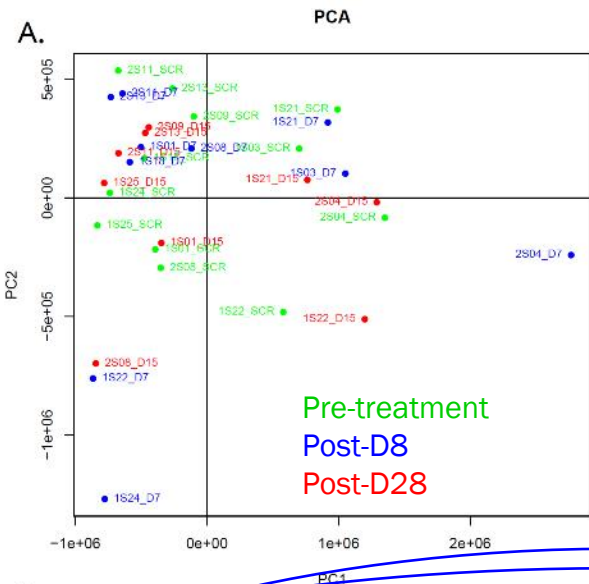
Adverse Reactions	Arm I - Concomitant (Dose 3,3E12 , n=6) ²		Arm II - Sequential (Dose 3,3E12 , n=6) ³		Arm II - Sequential (Dose 1E13 , n=8) ³		
	CTCAE Grade	Grade 1-2	Grade ≥3	Grade 1-2	Grade ≥3	Grade 1-2	Grade ≥3
Pyrexia		2 (33,0%)	-	5 (62,5%)	-	3 (50%)	-
Influenza like illness		3 (50,0%)	-	3 (37,5%)	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]

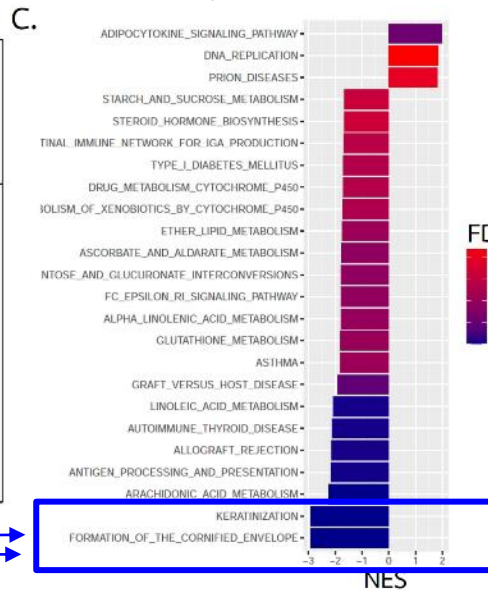
Principal Component Analysis¹

including all the Pre- and Post-treatment samples

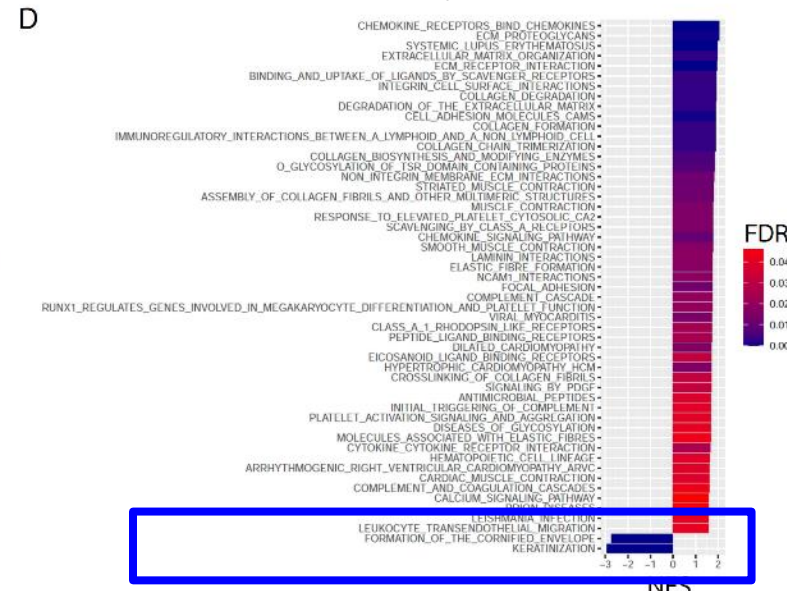


Most significant Reactome and KEGG pathways in GSEA (Gene Set Enrichment Analysis)¹

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 Cornified Envelope 1B	-7.921	0.006	

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

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