



# Corporate Presentation

---

*Investor Events*

---

*SEPTEMBER 12, 2023*

# Disclaimer & Forward-Looking Statement

IMPORTANT NOTICE – YOU MUST READ THE FOLLOWING BEFORE CONTINUING. THIS PRESENTATION HAS BEEN PREPARED BY GENFIT AND IS FOR INFORMATION PURPOSES ONLY.

CERTAIN OF THE INFORMATION CONTAINED HEREIN CONCERNING ECONOMIC TRENDS AND PERFORMANCE IS BASED UPON OR DERIVED FROM INFORMATION PROVIDED BY THIRD-PARTY CONSULTANTS AND OTHER INDUSTRY SOURCES. WHILE GENFIT BELIEVES THAT SUCH INFORMATION IS ACCURATE AND THAT THE SOURCES FROM WHICH IT HAS BEEN OBTAINED ARE RELIABLE, GENFIT HAS NOT INDEPENDENTLY VERIFIED THE ASSUMPTIONS ON WHICH PROJECTIONS OF FUTURE TRENDS AND PERFORMANCE ARE BASED. IT MAKES NO GUARANTEE, EXPRESS OR IMPLIED, AS TO THE ACCURACY AND COMPLETENESS OF SUCH INFORMATION.

This presentation contains certain forward-looking statements, including those within the meaning of the Private Securities Litigation Reform Act of 1995 with respect to GENFIT, including, but not limited to statements about GENFIT's corporate strategy and objectives, the potential sizes of the markets for PBC, cholangiocarcinoma, ACLF, hepatic encephalopathy (HE) and urea cycle disorder (UCD), commercial certainty within these markets and the potential of elafibranor to receive marketing authorization and be commercialized in PBC by Ipsen, development plans for our pipeline programs and expected timing for potential regulatory approvals and clinical milestones, as well as projections regarding our cash runway. The use of certain words, including "believe," "potential," "expect" and "will" and similar expressions, is intended to identify forward-looking statements. Although the Company believes its expectations are based on the current expectations and reasonable assumptions of the Company's management, these forward-looking statements are subject to numerous known and unknown risks and uncertainties, which could cause actual results to differ materially from those expressed in, or implied or projected by, the forward-looking statements. These risks and uncertainties include, among other things, the uncertainties inherent in research and development, including in relation to safety, biomarkers, cost of, progression of, and results from, its ongoing and planned clinical trials, review and approvals by regulatory authorities in the United States, Europe and worldwide, of our drug and diagnostic candidates, exchange rate fluctuations, potential synergies related to the acquisition of Versantis, our capacity to integrate its assets, develop its programs and our continued ability to raise capital to fund our development, as well as those risks and uncertainties discussed or identified in the Company's public filings with the AMF, including those listed in Chapter 2 "Main Risks and Uncertainties" of the Company's 2022 Universal Registration Document filed with the AMF on April 18, 2023, which is available on the Company's website ([www.genfit.com](http://www.genfit.com)) and on the website of the AMF ([www.amf-france.org](http://www.amf-france.org)) and public filings and reports filed with the U.S. Securities and Exchange Commission ("SEC") including the Company's 2022 Annual Report on Form 20-F filed with the SEC on April 18, 2023.

In addition, even if the Company's results, performance, financial condition and liquidity, and the development of the industry in which it operates are consistent with such forward-looking statements, they may not be predictive of results or developments in future periods. These forward-looking statements speak only as of the date of publication of this document. Other than as required by applicable law, the Company does not undertake any obligation to update or revise any forward-looking information or statements, whether as a result of new information, future events or otherwise.

CERTAIN OF THE INFORMATION CONTAINED HEREIN CONCERNING ECONOMIC TRENDS AND PERFORMANCE IS BASED UPON OR DERIVED FROM INFORMATION PROVIDED BY THIRD-PARTY CONSULTANTS AND OTHER INDUSTRY SOURCES. WHILE GENFIT BELIEVES THAT SUCH INFORMATION IS ACCURATE AND THAT THE SOURCES FROM WHICH IT HAS BEEN OBTAINED ARE RELIABLE, GENFIT HAS NOT INDEPENDENTLY VERIFIED THE ASSUMPTIONS ON WHICH PROJECTIONS OF FUTURE TRENDS AND PERFORMANCE ARE BASED. IT MAKES NO GUARANTEE, EXPRESS OR IMPLIED, AS TO THE ACCURACY AND COMPLETENESS OF SUCH INFORMATION.

# GENFIT Highlights

## Who we are



French **biopharmaceutical** company  
Dual-listed on Euronext & Nasdaq (GNFT)



Improving the life of patients  
with **liver diseases**




Specific focus on **rare** and **severe** liver  
diseases with **high unmet medical need**




Expertise bringing **early-stage assets**  
to **commercial readiness**



 Lille & Paris

 Zurich

 Cambridge, MA



United Nations  
Global Compact

PAQTE



Prime



## Our pipeline

ELAFIBRANOR in **PBC** (Positive Phase 3)

5 assets in **ACLF** and its complications

**VS-01** (Phase 2)

NTZ (Phase 2 ready)

SRT-015 (preclinical)

CLM-022 (preclinical)

VS-02-HE (preclinical)

**GNS561 CCA** (Phase 2)

VS-01 **UCD/OA** (preclinical)

**Diagnostic programs**

NIS2+ ('at risk' NASH)

TS-01 (ammonia)

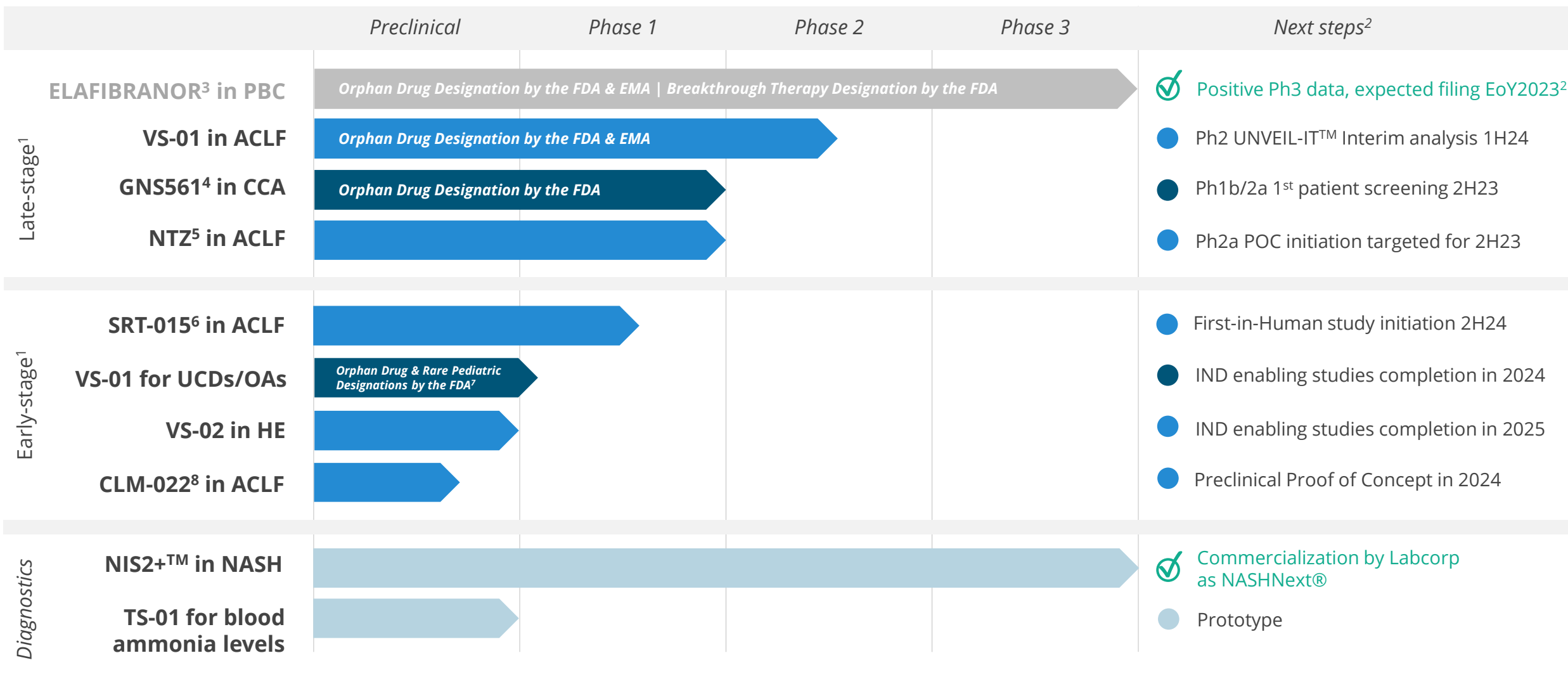
## Cash position

**€128.6M** as of March 31, 2023  
Cash until **3Q24\***

Potential **milestones** related to  
elafibranor in 2023 and 2024  
extending cash runway  
**Royalties** will help finance operations

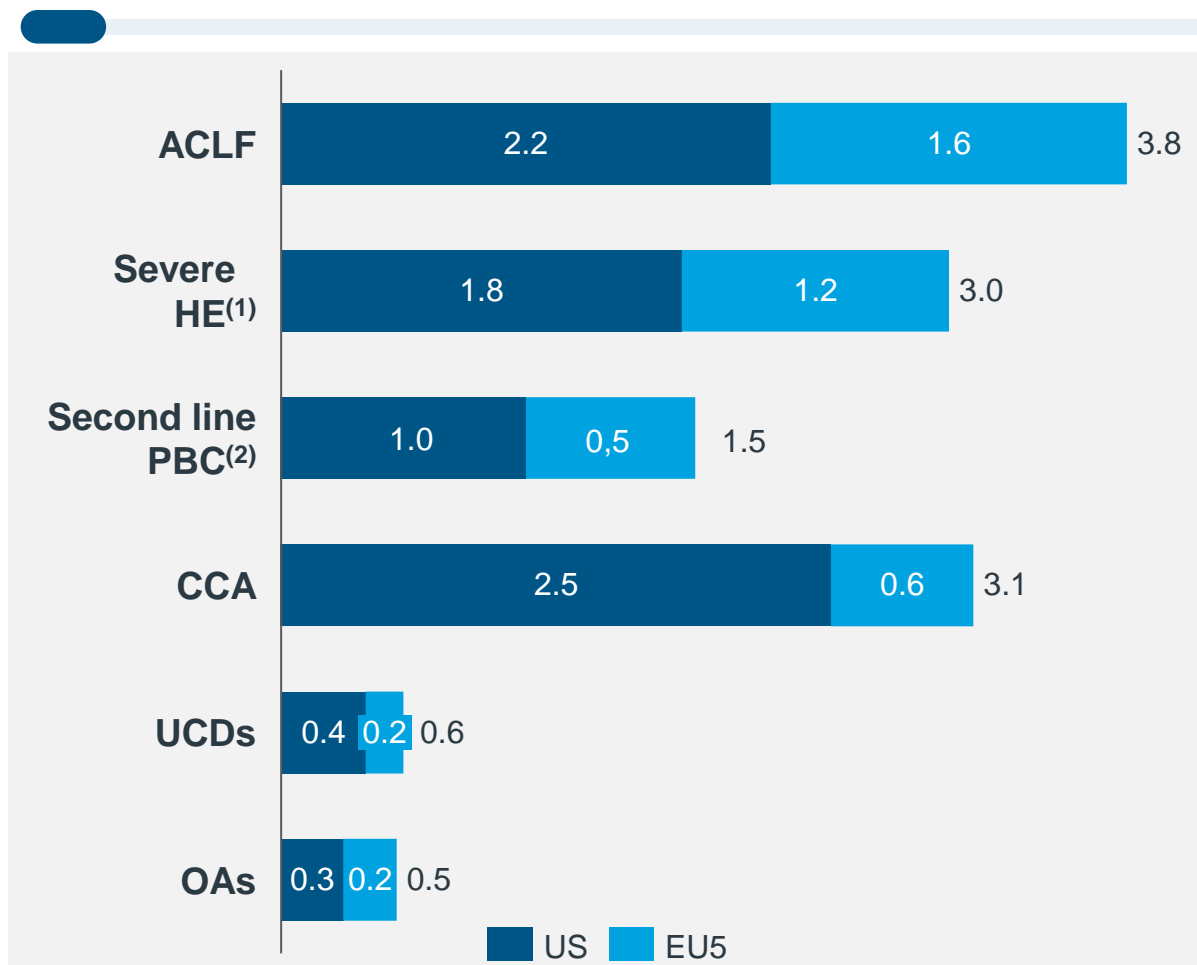
\* We expect that our existing cash, cash equivalents and current financial assets will enable us to fund our operating expenses and capital expenditure requirements until approximately the fourth quarter of 2024. This is based on current assumptions and without taking exceptional events into account, including potential milestone payments should the ELATIVE® study be successful.

# Pipeline: 10 Programs from Early-stage to Commercialization



# Market Potential: an Overall ~12.5bn USD Opportunity

## Estimated Overall Market Size (US+UE5) by 2030, bnUSD



## Assumptions<sup>(3)</sup>

- Prevalence: 155k (EU5) / 80k (US) for grade 1 / 2 ACLF patients
- Drug price could amount to \$30-40k per patient in US in secondary prevention for ACLF1/2. With restricted subpopulation in ACLF2 for acute life-threatening event, drug price could amount up to ~\$50-150k<sup>(4)</sup>

- Hospitalizations per year: 195k (EU5) / 200k (US)
- Drug price ranges: analogues in acute ICU costs would potentially range from \$15-20k in US and \$7-15k in EU5 based on economic burden of hospitalizations

- Prevalence: 52k (EU5) / 54k (US) for 40% of patients moving to 2L
- Drug gross price ranges per year: ~\$30k in EU5 in 2022 and ~\$84k in US expected to slightly evolve as competition will arise in second line

- Prevalence: 15k (EU5) / 15k (US)
- Drug price ranges per month: [\$500 - \$9k] in EU5 and [\$30k] in US

- Prevalence: 1k (EU5) / 1.3k (US)
- Drug price ranges per year: [\$500k - \$700k] in US and [\$300k - \$500k] in EU5

- Incidence in newborns: 129 (US), 198 (EU5)
- Drug price ranges per year: [\$96 - \$81k] in EU5 and [\$200 - \$300k] in US

(1) Only acute HE considered in estimations (2) Addressable market for second Line post UDCA (3) Estimation calculations include duration of treatment, potential eligibility to drug treatment, compliance rates based on analogues in rare diseases, gross-to-net price estimate depending on therapeutic area & disease (4) Acquired aplastic anemia could be a relevant analogue, treatments that include blood transfusions, stem cell transplant, immunosuppressants and bone marrow stimulants cost: approx. \$72k/patient per year

# Several Catalysts over the Next Few Years

## Elafibranor in PBC\*

- **Expected Filing in US and Europe\*\***

- **Potential US and European approval & commercialization\***

2023

2024

2025

## Pipeline

- **VS-01-ACLF**  
*Ph2 1<sup>st</sup> patient screening*
- **GNS561 CCA**  
*Ph1b/2a 1<sup>st</sup> patient screening*
- **NTZ ACLF**  
*Ph2a POC initiation*

- **VS-01-ACLF**  
*Ph2 clinical efficacy data*
- **GNS561 CCA**  
*Ph1b/2a biomarker data*
- **SRT-015 ACLF**  
*Initiation of First-in- Human study*
- **VS-01-UCD/OA**  
*IND enabling studies completion*
- **CLM-022 ACLF**  
*Preclinical POC*

- **VS-02 HE**  
*IND enabling studies completion*
- **GNS561 CCA**  
*Ph1b/2a clinical efficacy data*
- **NTZ ACLF**  
*Ph2a clinical data*

# Acute-on-Chronic Liver Failure (ACLF), a Growing Health Burden

## Definition, Epidemiology & Costs

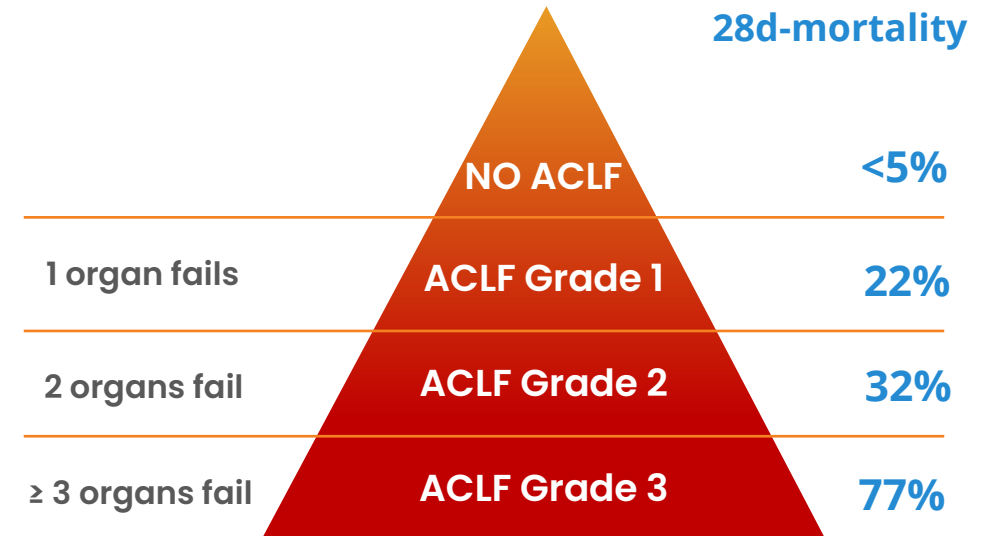
**ACLF = Abrupt life-threatening worsening of pre-existing chronic liver disease (CLD)/cirrhosis**

- Characterised by **pronounced systemic inflammation, hepatic and extrahepatic organ dysfunction & failure** (brain, kidneys, coagulation, cardiovascular and respiratory)
- High short-term mortality** (22% to 74% mortality at 28 days, depending on severity grade<sup>1</sup>)

### Relationship between CLD, cirrhosis & ACLF

- 1.5 billion people** suffer from chronic liver disease worldwide<sup>2</sup>
- Cirrhosis is the **11<sup>th</sup> cause of death** and **15<sup>th</sup> cause of morbidity** worldwide<sup>2</sup>
- Number of **cases of decompensated cirrhosis are projected to rise** in the next decade<sup>3</sup>
- ~1/3 patients** admitted with decompensated cirrhosis present with ACLF<sup>3</sup>
- The **cost per hospitalization for ACLF is 3.5-fold higher** than that for cirrhosis<sup>4</sup> (\$53,570 versus \$15,193)

**The Higher the ACLF Grade, the Higher the Mortality**



**No approved treatment in this indication = high unmet medical need**

# ACLF, a Multifactorial Syndrome

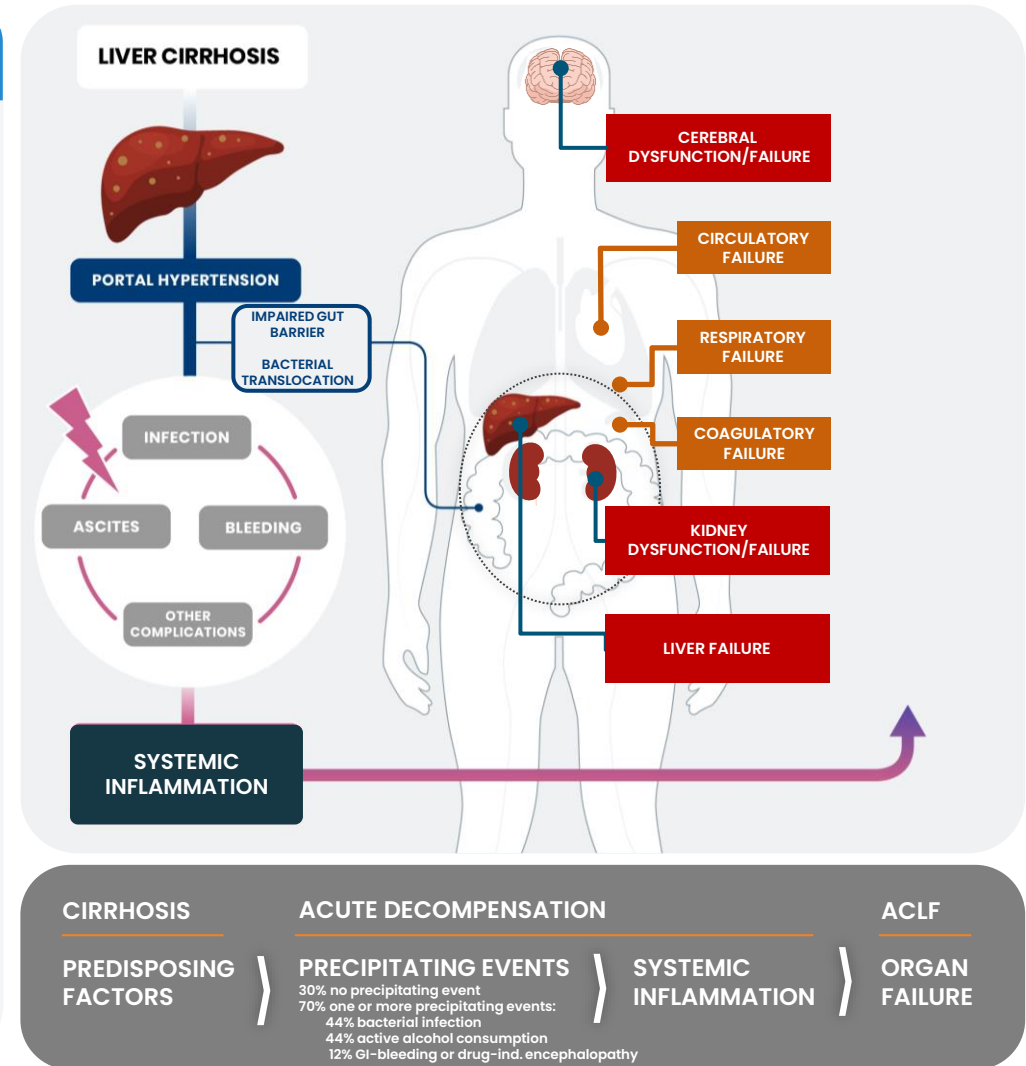
## Liver Disease Etiology & Pathogenesis

### Liver Disease Etiology<sup>1</sup>

- › ALD
- › DILI
- › Viral Hepatitis
- › MASLD
- › Autoimmune & cholestatic diseases
- › Hereditary diseases

### Mechanisms of pathogenesis<sup>2</sup>

- › **Portal hypertension** and splanchnic vasodilation cause **ascites**, and may lead to systemic hypotension & **organ hypoperfusion**
- › Dysbiosis, impaired gut barrier, and **bacterial translocation** are drivers of systemic inflammation
- › **Pronounced systemic inflammation** mediated via **PAMPs, DAMPs** leads to a cascade of downstream **cytokine signaling & immunoparesis**
- › **Mitochondrial ATP-depletion**, systemic **organ cell injury & hepatocellular cell death** are major contributing factors
- › Organ dysfunction and failure promote the **accumulation of ACLF-related metabolites incl. ammonia**





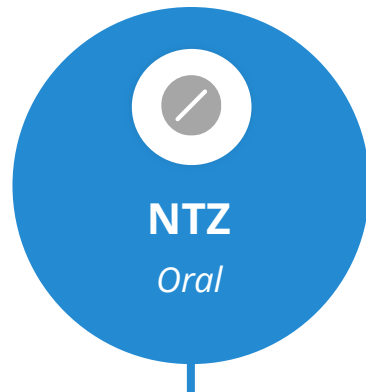
# Our Scientific Approach: A Diverse Portfolio with Complementary MoAs

We are developing a **diverse drug portfolio** with **complementary mechanisms of action**, to better address the **complexities of ACLF** and improve **treatment outcomes**.



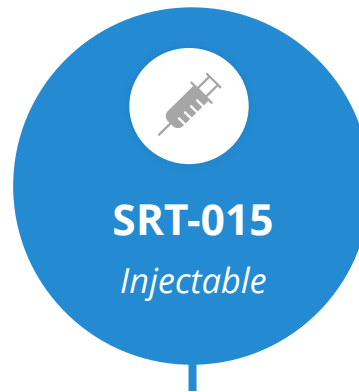
*Liposomal-based  
technology*

To remove **ACLF-related toxins** from the blood incl. **ammonia**



*Anti-inflammatory  
and anti-bacteria*

To reduce **systemic inflammation**, impede **PAMPs release** and **bacterial translocation**



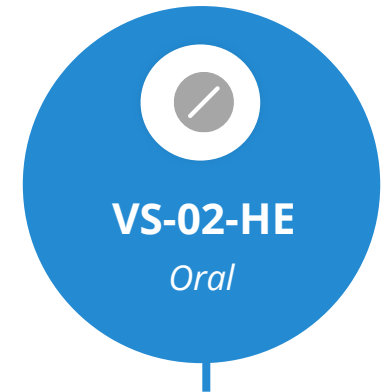
*ASK1  
inhibitor*

To inhibit **hepatocellular death**, **inflammation** and **fibrosis**



*NLRP3 inflammasome  
inhibitor*

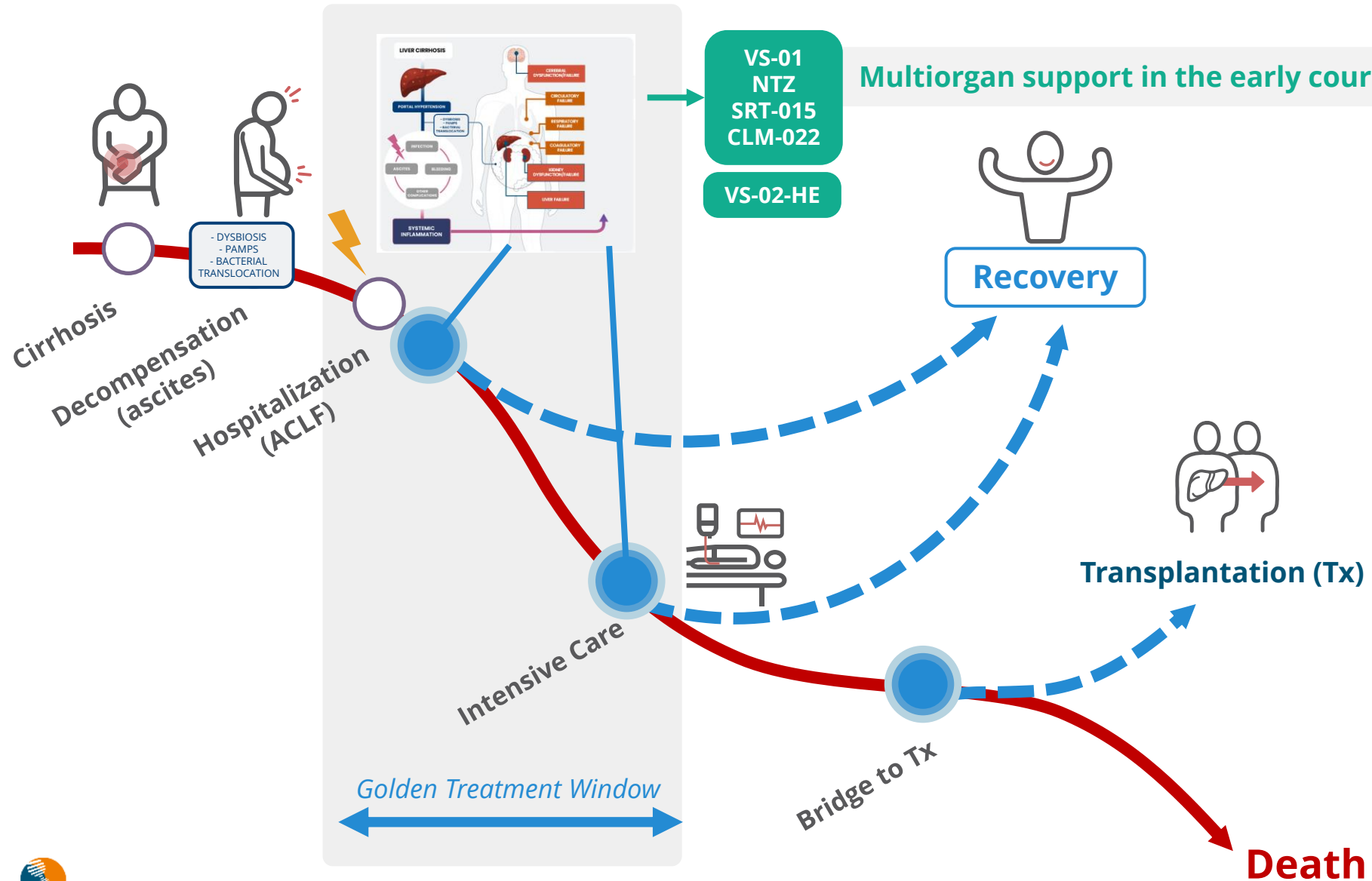
To inhibit **inflammation** and **cell death (pyroptosis)**



*Urease  
inhibitor*

To reduce **hyperammonemia**, stabilize blood ammonia and **prevent HE**

# Our Scientific Approach: Targeting the Golden Treatment Window

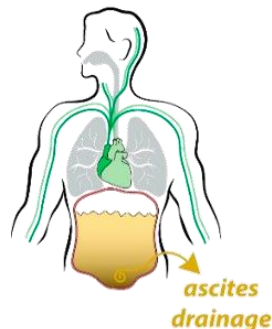


- ### TREATMENTS GOALS
- **R**esolve ACLF
  - **I**mprove survival
  - **C**hance of liver transplant increased
  - **H**ealthcare costs reduced

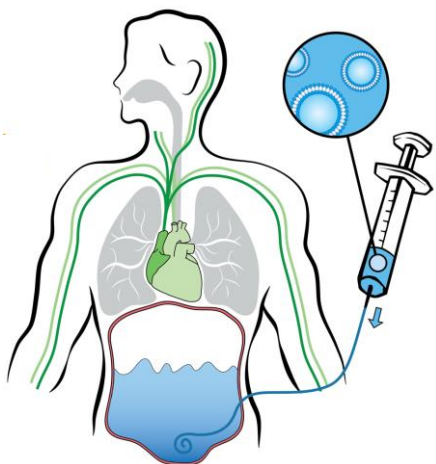
# VS-01, our Liposomal-based Technology for Ammonia Clearance in Blood

## Standard of care

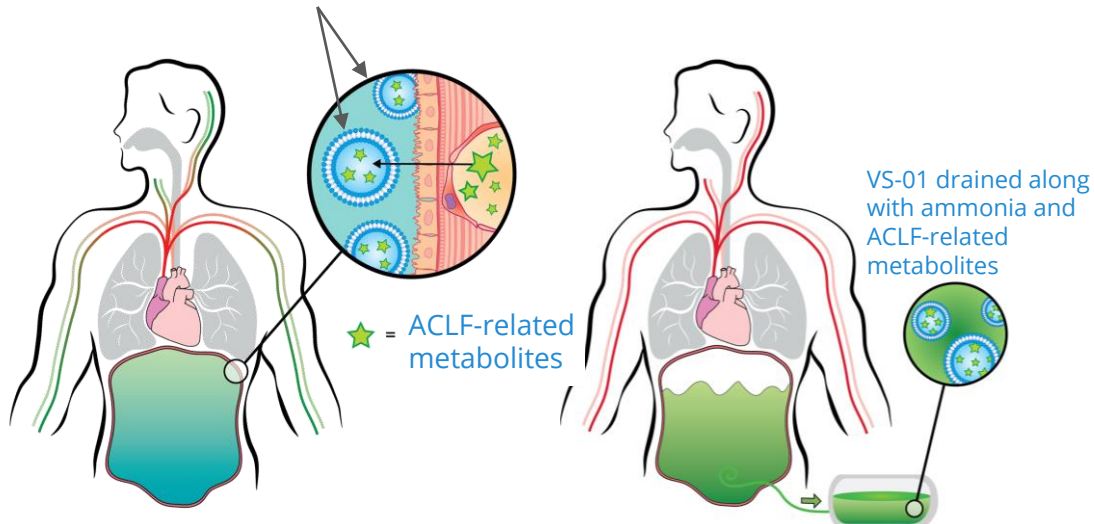
Ascites drainage



## VS-01



## VS-01 scavenging liposomes



Intraperitoneal route of administration following paracentesis

## VS-01 in brief

Targets **first-line treatment for ACLF** to **reverse** the disease

Delivered via **in-place peritoneal access** catheter

**Targets multiorgan support:** brain, liver, and kidney

## Supporting Evidence

**Phase 1b FIH study<sup>2</sup> in patients with decompensated liver cirrhosis, ascites and covert HE**

- › Generally **Safe and well-tolerated**
- › **Improvement** on overall **liver disease severity**
- › Dose-dependent **ammonia removal** from the body
- › **Improvement in psychometric tests**
- › Reduction of **ACLF-related metabolites**
- › Reduction of **infection-related metabolites**

Upcoming milestone

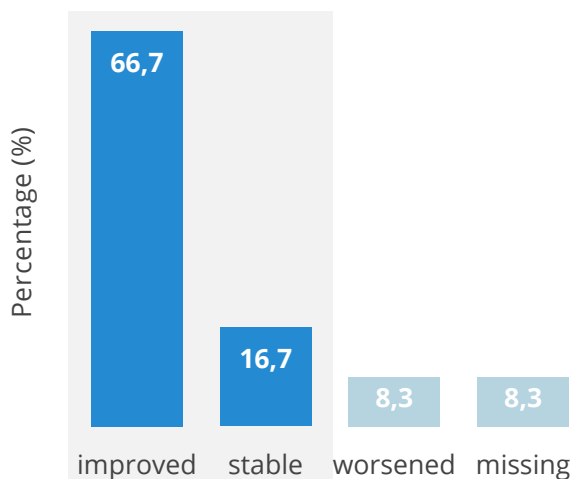
**Ph2 Interim Analysis 1H24**

# VS-01-ACLF: FIH Ph1b Study Efficacy Results on Liver & Brain Function<sup>1</sup>

## IMPROVEMENT OF OVERALL LIVER DISEASE SEVERITY

e.g., assessed by Child-Pugh Score (CPS)

Improved or stable disease:  
**83.4%**

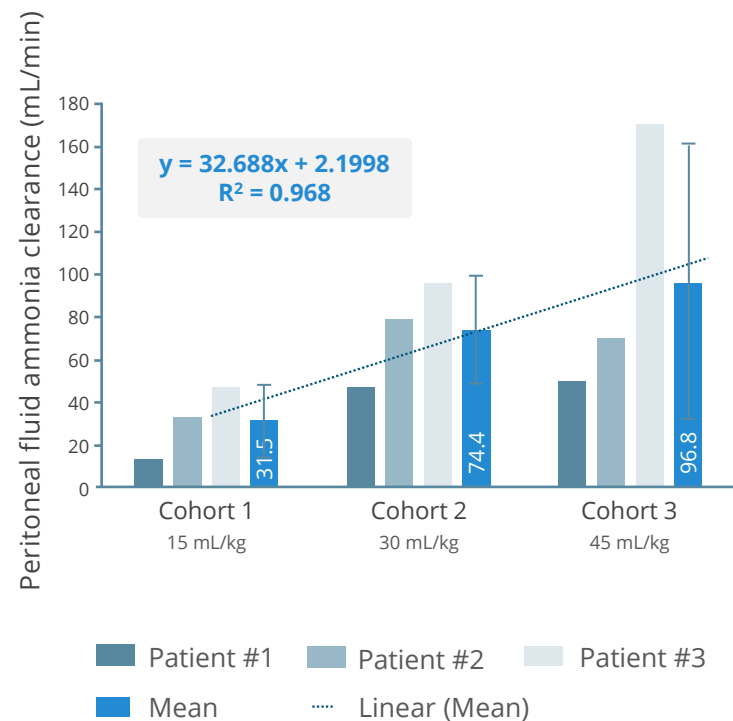


NO PATIENTS PROGRESSED TO ACLF



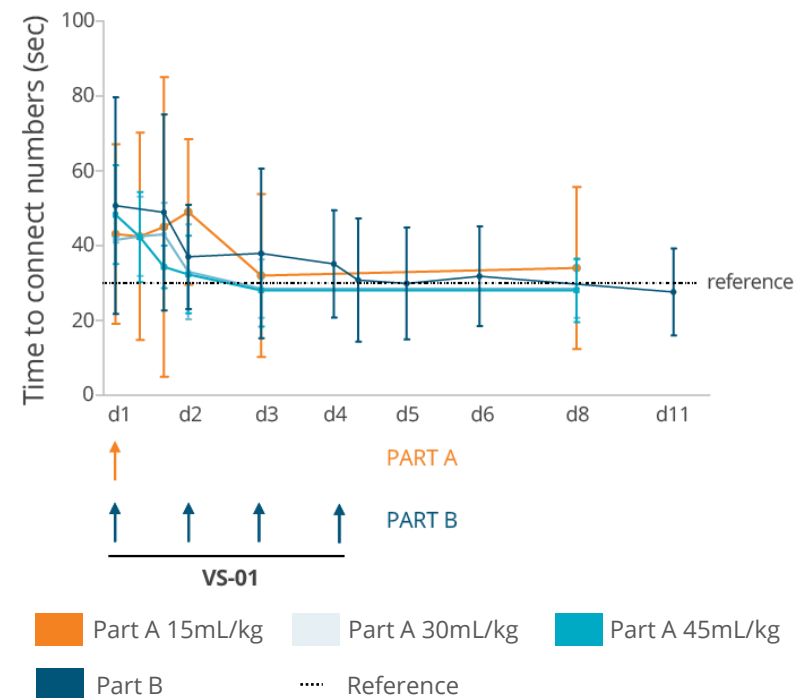
## DOSE-DEPENDENT AMMONIA REMOVAL FROM THE BODY

Ammonia clearance increased with VS-01 dosage in peritoneal fluid



## IMPROVEMENT IN PSYCHOMETRIC TESTS FOR HE

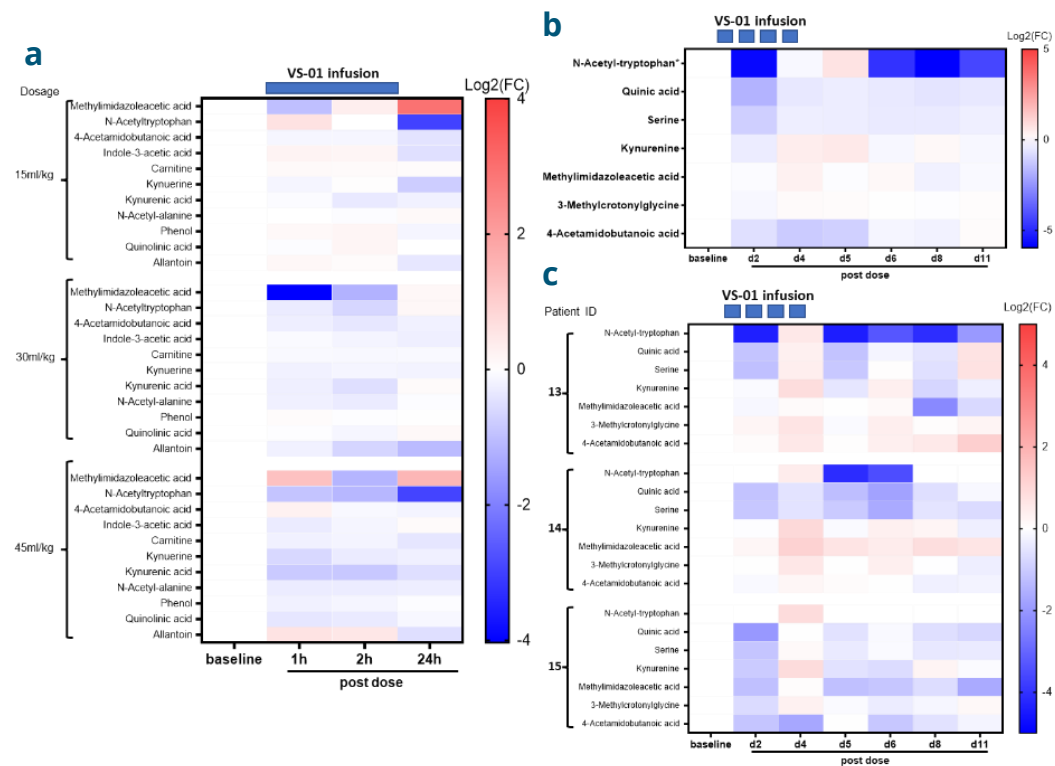
e.g., number connection test was performed faster



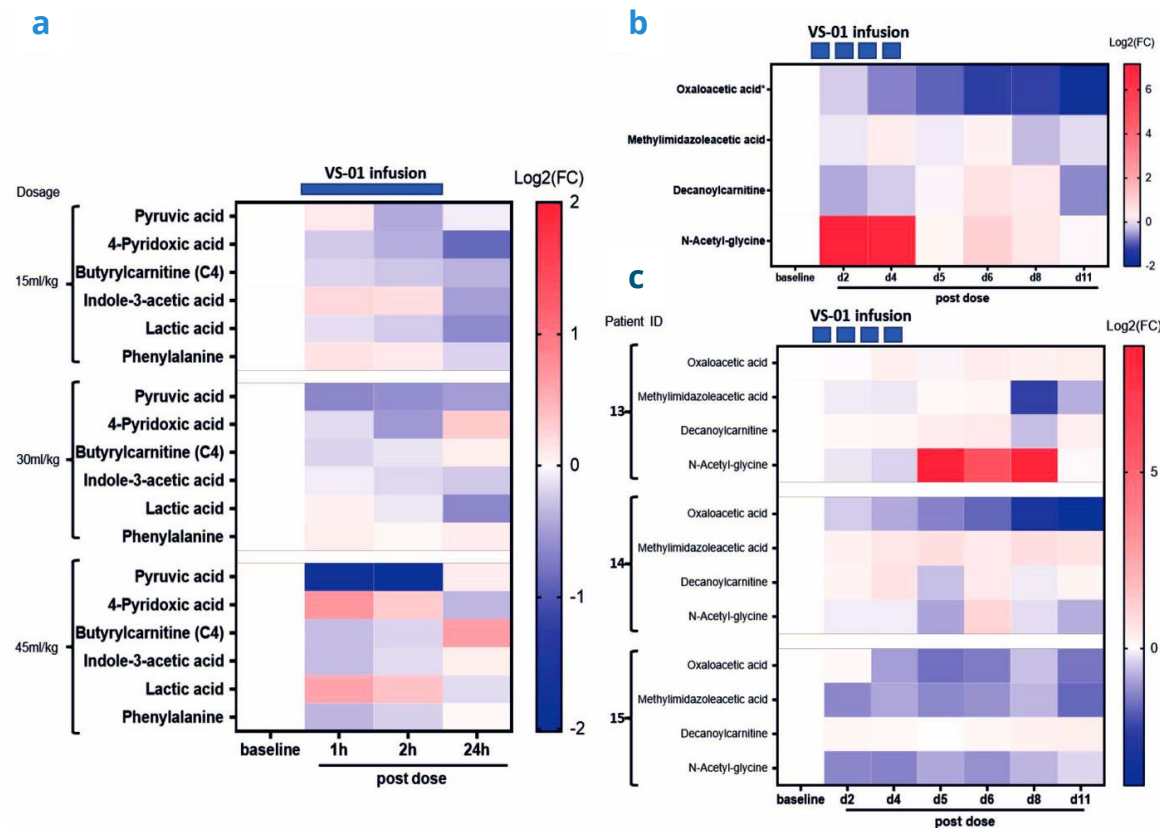
# VS-01-ACLF: FIH Ph1b Study Efficacy Results on Metabolites & Inflammation<sup>1</sup>

## REDUCTION OF ACLF-RELATED METABOLITES<sup>2</sup>

VS-01 reduced plasma metabolites associated with organ failures<sup>3</sup>



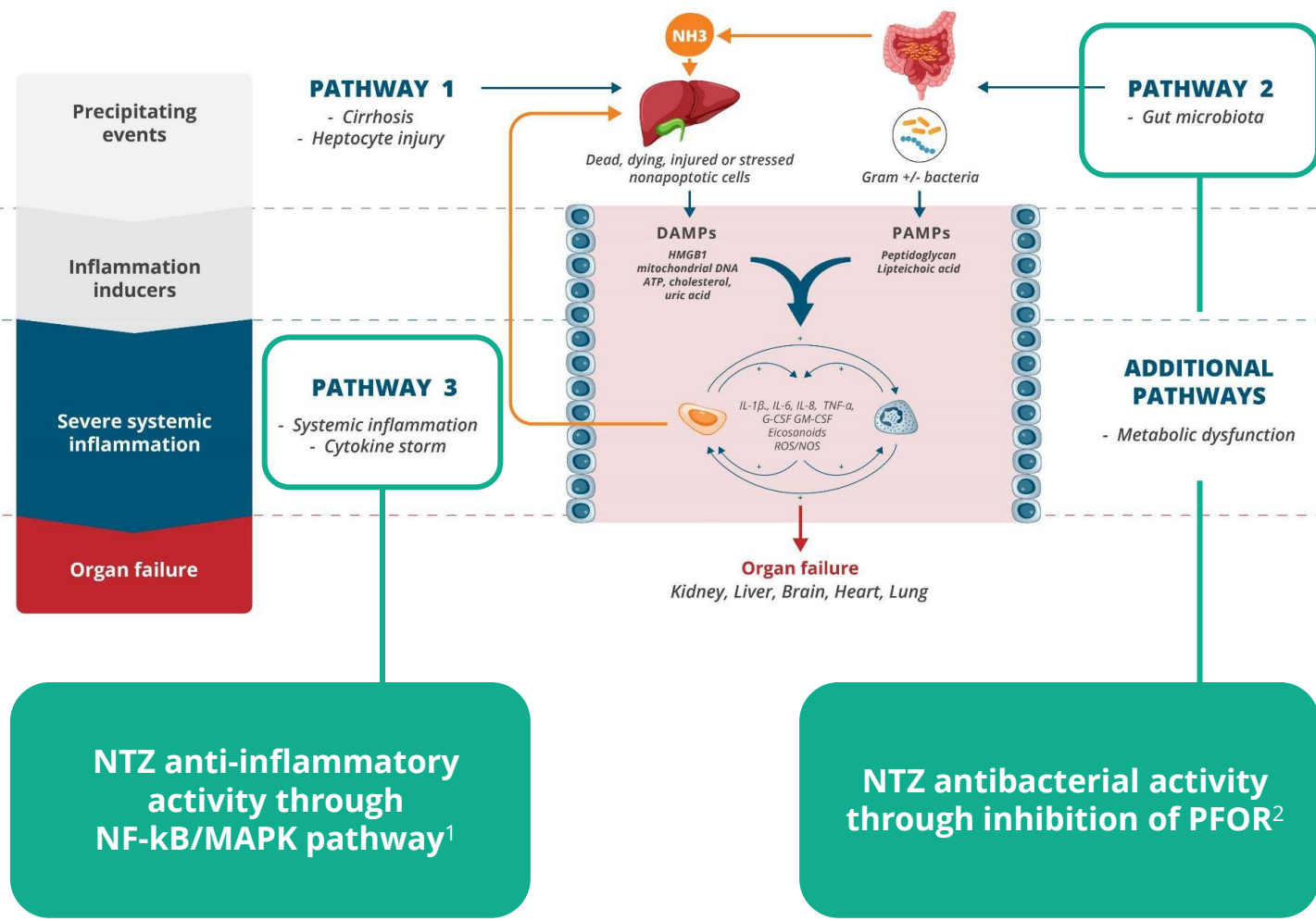
VS-01 reduced plasma metabolites associated with bacterial infection<sup>4</sup>



Two abstracts presented at EASL-ILC 06/2022

Abstract selected for 2022 EASL 'Best of International Liver Congress Summit' resource

# NTZ, Antiparasitic with Anti-Inflammatory and Antibacterial Effects



## NTZ in brief

Antiparasitic medication used to treat infections  
Several clinical trials and observational studies<sup>1,2</sup> reported **anti-inflammatory** and **anti-bacterial effects** of NTZ in human

## Supporting Evidence

### Preclinical

- › Reduces **LPS-induced inflammation** in healthy rats
- › Beneficial effects on **liver function markers** (bil, alb) in models of cirrhosis
- › Reduces **brain edema** in models of ACLF (BDL)
- › Reduces **inflammation markers** in models of ACLF (BDL)
- › Improves **survival** in treatment models of Sepsis (CLP)

### Clinical (phase 1 studies)

- › Was generally **well tolerated**, with a **favorable safety profile**, in subjects with HI and RI
- › In patients with severe HI, demonstrated **trends for improvement in inflammatory and liver markers**<sup>3</sup>

Upcoming milestone

**Ph2a POC init. targeted for 2H23**

# NTZ, Activity in Disease Models Support ACLF Clinical Development



NTZ **reduces LPS-induced** inflammation in healthy rats\*



NTZ has **beneficial effects on liver function** markers (bil, alb) in models of cirrhosis\*



NTZ **reduces brain edema** in models of ACLF (BDL)

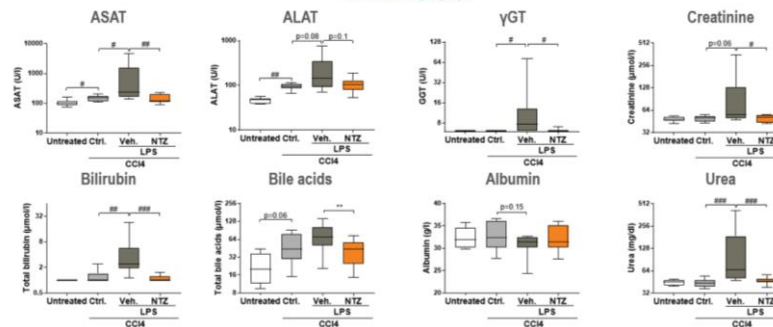
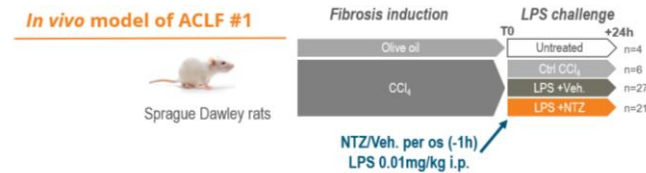


NTZ **reduces inflammation markers** in models of **ACLF** (BDL)

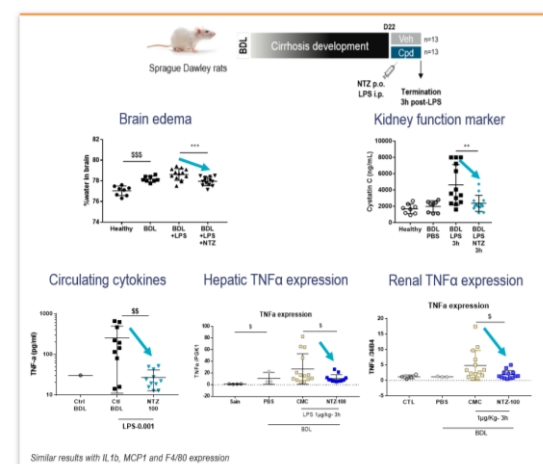


NTZ **improves survival** in **treatment** models of **Sepsis** (CLP)

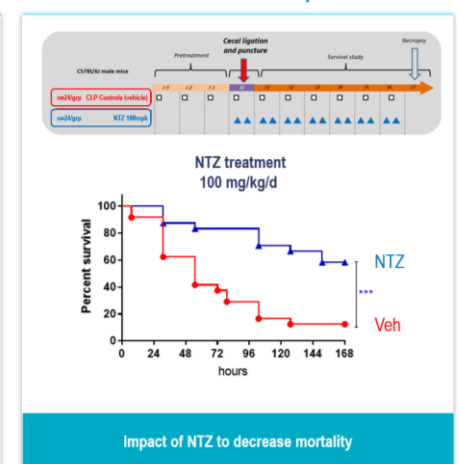
+ NTZ completed 2 Ph1 studies in subjects with hepatic impairment (HI) and renal impairment (RI), showing a favorable safety and tolerability profile in both studies



**In vivo model of ACLF #2**



**In vivo model of sepsis**



# 3 Early-stage Programs for the Treatment of ACLF and its Complications

## SRT-015<sup>1</sup> in ACLF



ASK1  
inhibitor

### Supporting evidence

#### Preclinical and clinical<sup>2</sup>

- › In kidney diseases, limits renal inflammation, apoptosis and fibrosis
- › In liver diseases, prevents hepatocyte death, inflammation and fibrosis
- › In brain disorders, limits neurodegeneration
- › In inflammatory diseases, limits damaging immune responses
- › In cardiopulmonary disease, slows the onset of heart failure



First-in-Human  
study initiation  
**2H24**

## CLM-022<sup>3</sup> in ACLF



NLRP3  
inflammasome  
inhibitor

### Supporting evidence

#### Preclinical studies<sup>4</sup>

- › primarily in animal models of liver injury and inflammation, have shown promise for NLRP3 inflammasome inhibitors in reducing liver damage and inflammation



Preclinical Proof  
of Concept  
in **2024**

## VS-02 in HE



Urease  
inhibitor

### Supporting evidence

#### Preclinical proof of concept<sup>7</sup>

- › Superior urease inhibitory activity in vitro
- › Cytotoxicity and mutagenicity assessment
- › In vivo efficacy to significantly reduce plasmatic ammonia and brain glutamine in bile duct-ligated rats



IND enabling  
studies  
completion  
in **2025**

### About HE

- Major complications of **advanced liver disease** and **portal hypertension**<sup>6</sup>

- **30-40% of patients with cirrhosis** will experience at least 1 episode<sup>5</sup>

- **Independent risk factor of mortality** in ACLF<sup>6</sup>



# Other Therapeutic Areas: GNS561 for the Treatment of KRAS-mutated CCA

## About CCA

- › **Malignancy** of the bile ducts => as cancer grows, leads to **damage to the liver** and other organs
- › Approx. **15% of all primary liver tumors** and **3% of gastrointestinal cancers**<sup>2</sup>
- › Without treatment **<20% of patients survive 5 years** from diagnosis<sup>1</sup>
- › KRAS mutation (20 to 40% of CCA) **not addressed by current treatments**

Rare and severe disease with **high unmet medical needs**



## Total CCA market estimates

**\$3.1 bn\***



GNS561, a **clinical-stage candidate** that localizes in lysosome where it **binds and inhibits PPT1**, resulting in:

- › lysosomal unbound Zn<sup>2+</sup> accumulation
- › impairment of cathepsin activity
- › blockage of autophagic flux
- › altered location of MTOR
- › lysosomal membrane permeabilization

### GNS561\*\* in CCA

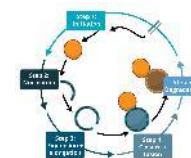


*PPT1 inhibitor in combination with a MEK inhibitor (MEKi)*

### Supporting evidence

#### Preclinical and PHASE 1b data<sup>3</sup>

- Antitumor activity in human cell lines (HCC, ICCA)
- Decreases tumor number and size in transgenic HCC mouse model
- First-in-human effects of PPT1 inhibition using GNS561/ Ezurpimtrostat in patients with primary/secondary liver cancers



- Autophagy promotes cancer cell survival, tumor growth and treatment resistance



**Ph1b/2a 1<sup>st</sup> patient**  
**Randomized 3Q23<sup>4</sup>**

# Other Therapeutic Areas: VS-01, ammonia clearance in HAC

## About Urea Cycle Disorders (UCD) and Organic Acidemias (OA)

- › Groups of **congenital metabolic diseases** => abnormalities in the metabolism of ammonia leading to **hyperammonemic crises (HAC)**
- › **Ultra rare** (1900 HAC/year in US+EU5)<sup>1,2,3</sup>
- › **Very high mortality** (75% after 5 years)<sup>1</sup> + survivors often have severe brain injuries
- › **No acute treatment** available for early onset crises

**Urgent unmet needs in the treatment of acute hyperammonemic crises (HAC)**



## Total UCD/OA market estimates

**\$0.6 bn\***



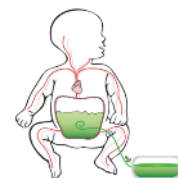
## VS-01 in UCD/OA, an **existing asset leveraged** in another indication

- › Peritoneal route well adapted to pediatric patients
- › Rapid treatment onset in all hospitals
- › Complementary to other therapeutical approaches

### VS-01 in UCD/OA



*Potential first-line peritoneal route treatment*



### Supporting evidence from ACLF program

#### Preclinical proof of concept

- VS-01 demonstrated superior ammonia clearance than commercial peritoneal dialysis in-vivo<sup>4,5,6,7</sup>
- Ammonia clearance in adult patients with decompensated cirrhosis at least comparable with hemodialysis<sup>8</sup>



IND enabling studies completion in **2024**

# Diagnostic Program: NIS2+™ for the Identification of at-risk NASH

## NASH diagnosis in brief

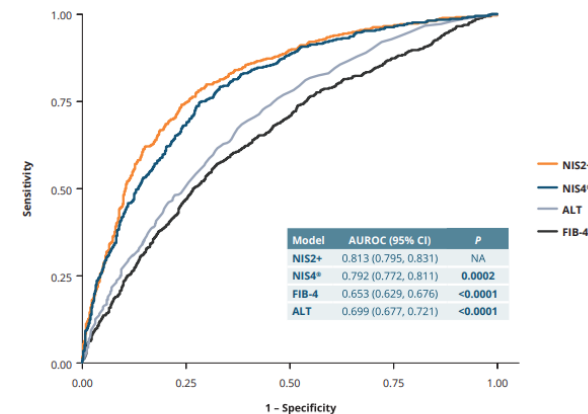
- › 6.7M patients have **NASH and significant fibrosis** (F $\geq$ 2) in the US<sup>1</sup> but only **900,000 are diagnosed**
- › **Poor disease awareness** among patients with NAFLD due to nonspecific symptoms<sup>2,3</sup>
- › Liver biopsy, the reference standard for NASH, poses **risks for patients** and has **technical limitations**<sup>4</sup>
- › Patients with NASH and significant fibrosis (F $\geq$ 2) (ie. at-risk NASH), are **at increased risk of progression** and are designated as **eligible to potential treatments**<sup>5</sup>

- › There are **no non-invasive blood-based diagnostic tests** specifically developed to identify at-risk NASH, impeding **large scale patient's diagnostic**
- › Potential **availability of anti-NASH drugs** in the near future could **bring incentive to diagnose** and for **large scale clinical use**

We **developed and validated NIS2+™**, an optimization of the NIS4® technology for the **diagnosis of at-risk NASH**

- › Data demonstrate **robust and improved clinical performance for efficient identification of at-risk NASH**, irrespective of patient characteristics such as age, sex and T2D<sup>6,\*\*</sup>
- › This NIT\* has the potential to be widely used for both **ruling out and ruling in at-risk NASH** thus be implemented at **large-scale in clinical practice**

## ROC curves and AUROC values for NIS2+™, NIS4®, FIB-4 and ALT for the detection of at-risk NASH



Test cohort: N=2035 (929 [45.65%] patients with at-risk NASH). AUROCs were compared using paired Delong tests. ALT, alanine aminotransferase; AUROC, area under the receiver operating characteristic curve; FIB-4, fibrosis-4; NA, not applicable; ROC, receiver operating characteristic.

- › NIS2+™ returned the highest AUROC value (0.813) among NIS4®, FIB-4, and ALT despite the removal of Hemoglobin A1c (HbA1c), and Alpha2-macroglobulin (A2M)

**NIS2+™ data published in peer-reviewed *Journal of Hepatology* in May 2023, and in *Hepatology Communications* in August 2023**

At EASL Congress in June 2023, and after several years of discussion among the relevant stakeholders, it was announced that nonalcoholic steatohepatitis (NASH) would now be referred to as Metabolic dysfunction-associated steatohepatitis (MASH). In addition, Nonalcoholic fatty liver disease (NAFLD) will now be referred to as metabolic dysfunction-associated steatotic liver disease (MASLD). GENFIT is progressively transitioning its documentation over to this new nomenclature and both NASH and MASH terms may appear in our documents during this period

# Conclusion

Potential future revenues from PBC program



**Potential future milestones and royalty revenues** expected to further strengthen GENFIT's financial position, and **accelerate pipeline development**

Large and diversified ACLF pipeline



**5 programs** under development to address ACLF, a severe disease with high unmet medical need, and a **>\$4bn market potential**

- From preclinical to **Phase 2**
- With **complementary MoAs**

Frequent newsflow as pipeline advances



Up to **7 data readouts** expected in the coming **2 years** related to biomarker or clinical efficacy

- in **ACLF**
- in **CCA**



We thank you for  
your attention

---

SEPTEMBER 12, 2023