

**sequana**medical

Needham Healthcare Conference, April 2022

Ian Crosbie, CEO

Euronext: SEQUA.BR



## Innovators in the treatment of diuretic-resistant fluid overload

liver disease  malignant ascites  heart failure

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## Regulatory disclaimer:

- The **alfapump**<sup>®</sup> system has not yet received regulatory approval in the United States and Canada. Any statement in this presentation about safety and efficacy of the **alfapump**<sup>®</sup> system does not apply to the United States and Canada. In the United States and Canada, the **alfapump**<sup>®</sup> system is currently under clinical investigation (POSEIDON Study) and is being studied in adult patients with refractory or recurrent ascites due to cirrhosis. For more information regarding the POSEIDON clinical study visit [www.poseidonstudy.com](http://www.poseidonstudy.com).
- DSR<sup>®</sup> therapy is still under development and it should be noted that any statements regarding safety and efficacy arise from ongoing pre-clinical and clinical investigations which have yet to be completed. DSR<sup>®</sup> therapy is currently not approved for clinical research in the United States or Canada. There is no link between DSR<sup>®</sup> therapy and ongoing investigations with the **alfapump**<sup>®</sup> system in Europe, the United States or Canada.

## COVID-19 disclaimer:

- Sequana Medical is closely following the evolution of the COVID-19 global health crisis and is in constant dialogue with its partners to assess the impact and adapt operations accordingly.
- Sequana Medical has put in place mitigation plans to minimise delays. The impact of increased demands on the healthcare systems, limitations on non-essential hospital visits and procedures, social-distancing and travel restrictions may result in further delays to execution of clinical studies and impact sales.
- Sequana Medical will continue to update the market as needed and whenever possible.

## Note:

- **alfapump**<sup>®</sup> is a registered trademark. DSR<sup>®</sup> and **alfapump DSR**<sup>®</sup> are registered trademarks in the Benelux, China, the EU, United Kingdom, and Hong Kong.

# Uniquely positioned in two large markets



- **alfapump<sup>®</sup> in liver disease – over €3 Bn / year <sup>(1)</sup>**
  - NASH is changing liver cirrhosis market and driving growth
  - Approved in EU / FDA breakthrough designation in US
  - North American pivotal study de-risked / primary endpoint Q4 '22
  - Direct commercialisation in US

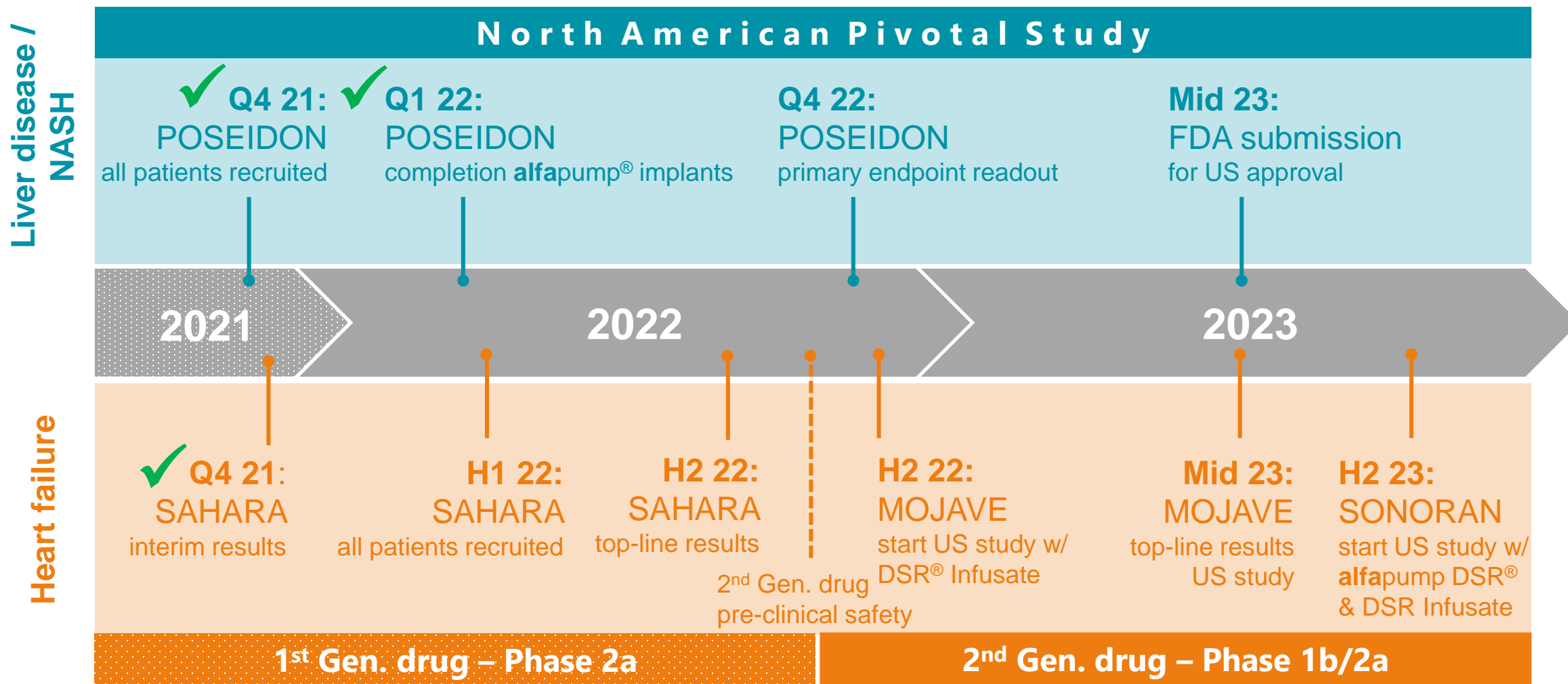


- **DSR<sup>®</sup> in heart failure – over €5 Bn / year <sup>(2)</sup>**
  - Congestion is a key driver of heart failure and key clinical challenge
  - Ph. 2a 1<sup>st</sup> Gen. drug – clinical proof-of-concept
  - Low-risk proprietary 2<sup>nd</sup> Gen. drug – on track for Q4 US clinical study
  - Partnering after US efficacy study



- **Proprietary technologies treating diuretic-resistant fluid overload**
  - Key clinical problem in liver disease, heart failure, renal failure and cancer
  - Diuretic-resistance is common – alternatives have significant disadvantages
- **Strong granted IP portfolio**

# Strong outlook for value drivers



Note: Presented timelines are subject to further developments related to the COVID-19 pandemic








**alfapump®**

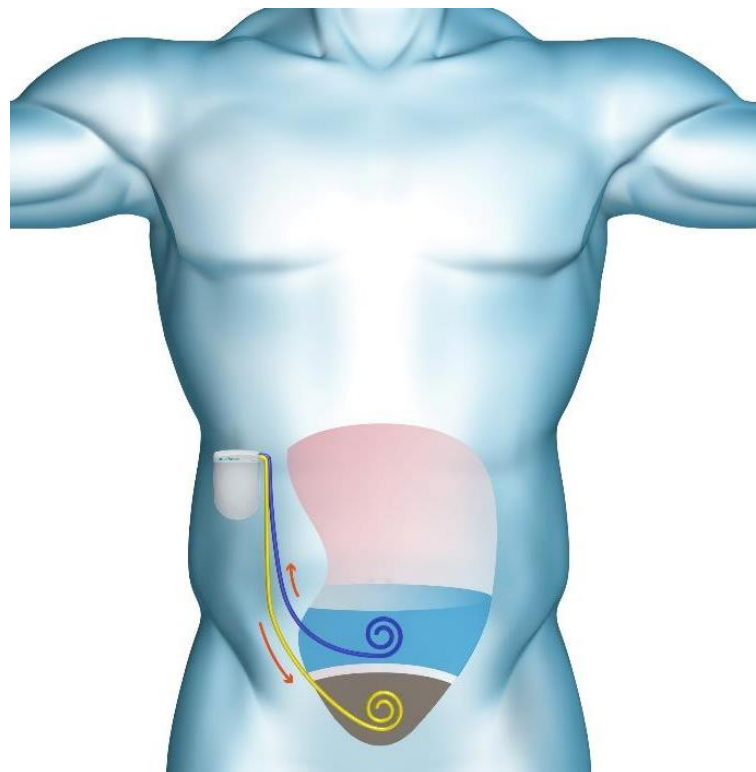
Proven step change in the  
treatment of liver refractory  
ascites





# alfapump®

Eliminating fluid from the peritoneal cavity – working in partnership with the bladder



-  Fully implanted
-  Automatic operation
-  Wireless battery charging
-  Settings wirelessly adjusted
-  Remote data monitoring

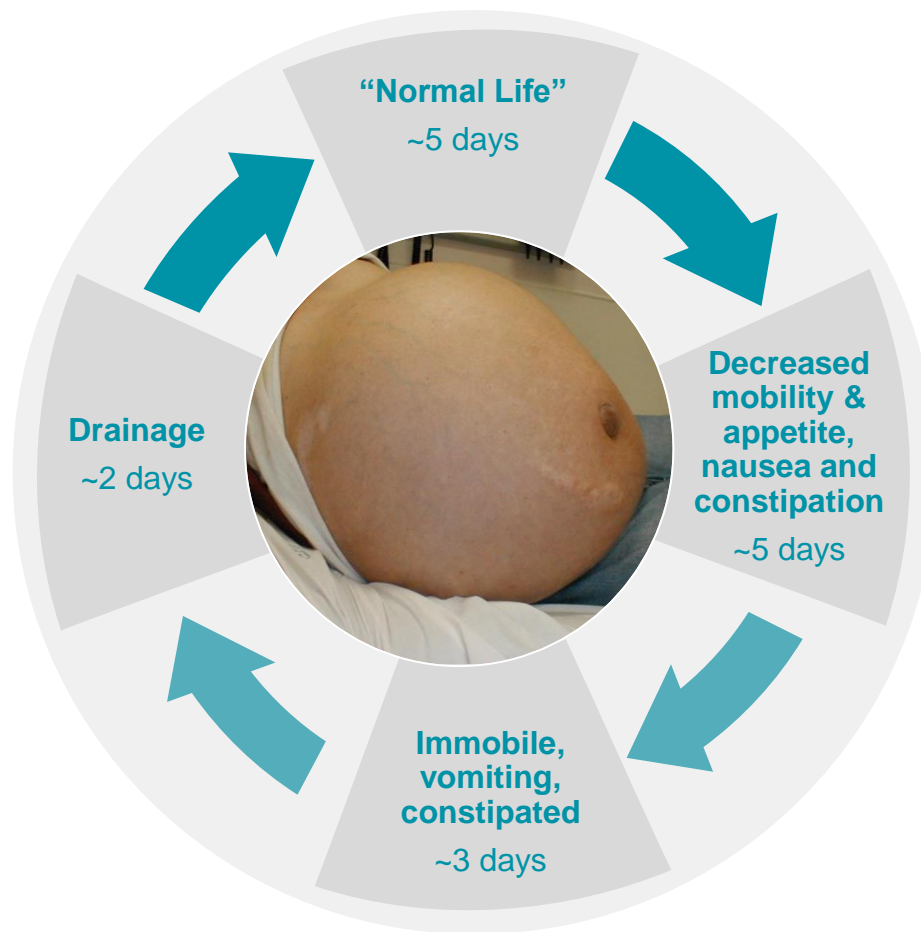


-  Easy implantation
-  Long-term implantation & catheter patency
-  Moves up to 4 litres / day
-  Virtually non-clogging
-  No significant heating during charging and operation

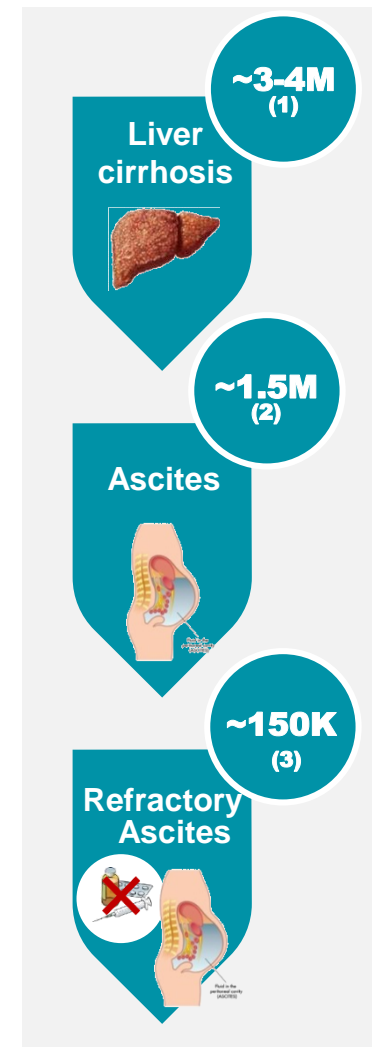
***Proven capabilities – over 900 systems implanted***  
***Strong IP barriers through extensive patent portfolio & know-how***

# Refractory ascites – key complication of liver cirrhosis

Fatty liver disease / NASH is driving dramatic growth and change in attitudes to liver cirrhosis patients



Typical patient life<sup>(4)</sup>



Note : Prevalence of NASH in US is expected to increase by 63% between 2015-2030; Estes et al., 2018

Source 1 Management estimate in US based on Estes et al; GlobalData Nash Epidemiology Forecast to 2026; Nouredin et al., 2013

Source 2: Runyon 2009: approximately 50% of cirrhotic patients develop ascites within 10 years of diagnosis of cirrhosis

Source 3: Ginès et al., NEJM 2004: refractory ascites occurs in 5-10% patients with ascites

Source 4: Presentation of Dr. Rajiv Jalan at EASL in 2018, Large Volume Paracentesis (LVP) treatment cycle for refractory ascites

# NASH transforming the face of liver cirrhosis

In US, liver cirrhosis is transitioning to a mainstream disease requiring modern treatment options





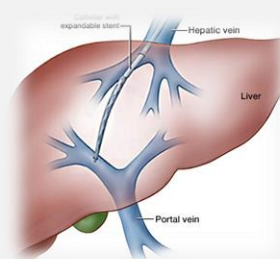
# Limitations of existing therapies

## Drainage (“Large Volume Paracentesis / LVP”)



Painful, Poor Quality of Life, Short Term Benefit

## Transjugular Intrahepatic Portosystemic Shunt (TIPS)



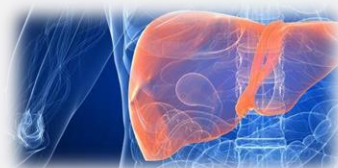
Complications, Contraindications

## Permanent Catheter System



External Catheter, Risk for Infections / Blockage

## Liver transplantation



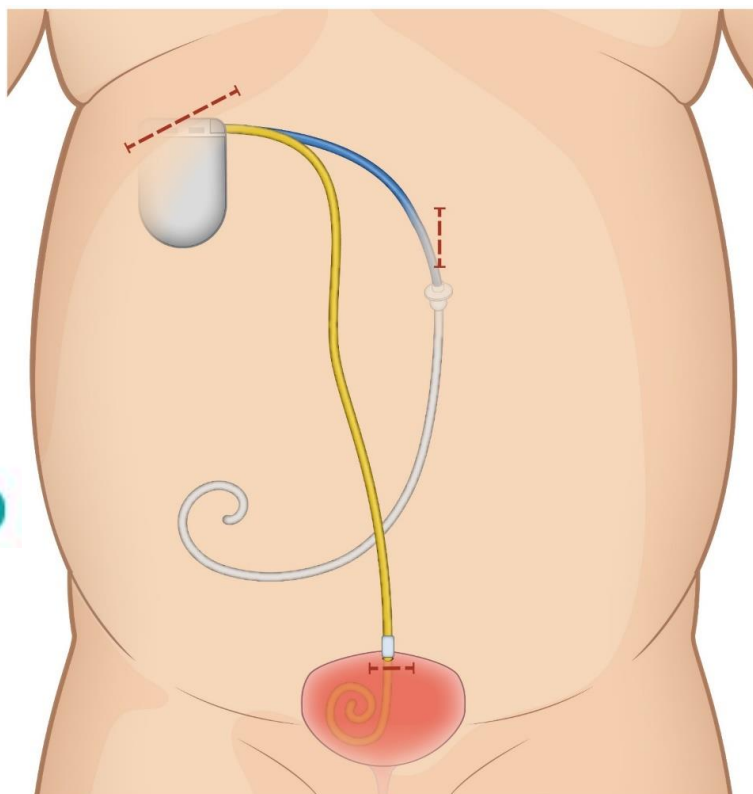
High Cost, Limited Availability

**alfapump®**



# alfapump® strong clinical and economic rationale

Over 900 implants and hundreds of years of patient experience



- ✔ Reduced burden of disease
- ✔ Improved patient QoL
- ✔ Cost savings for hospitals and payers

Estimated treatment cost / patient\*:

**LVP: ~\$54K** ↔ **alfapump®: ~\$35K**

~\$1.8K / LVP<sup>(1)</sup>      ~\$25K / alfapump  
 2 LVP / month      ~\$10K / implantation  
 15 months

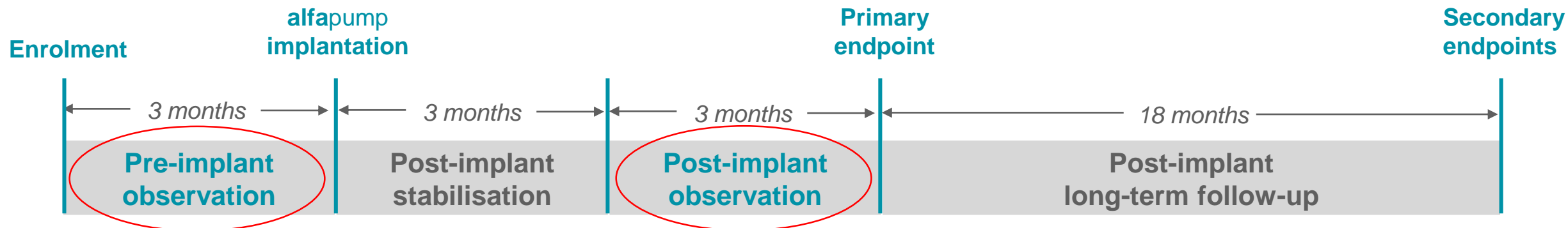
\* Management estimate of US treatment costs, assuming no complications

QoL: Quality of Life; LVP: Large Volume Paracentesis



# North American Pivotal Study (POSEIDON) underway

Pivotal Cohort of 40 implanted patients; Roll-In (“training”) cohort of 29 implanted patients



## POSEIDON Study Endpoints

**Primary efficacy:** 1) 50% reduction in average monthly frequency of Therapeutic Paracentesis (“TP”) post-implant vs. pre-implant  
2) 50% of patients achieve a 50% reduction in the requirement for TP post-implant vs. pre-implant

**Primary safety:** Rate of **alfapump** related re-interventions adjudicated by the Clinical Events Committee (CEC)

**Secondary:** QoL (SF36, Ascites-Q), nutritional status, health economics, safety (device and/or procedure-related AEs), survival

# Interim POSEIDON: Positive for primary endpoints

Data from first 26 Roll-In patients

## EFFICACY

- ✓ Over 90% reduction in mean Therapeutic Paracentesis (TP) frequency (primary endpoint >50% reduction)
- ✓ 100% patients with > 50% reduction in mean TP frequency per month (primary endpoint >50% of patients)

## SAFETY

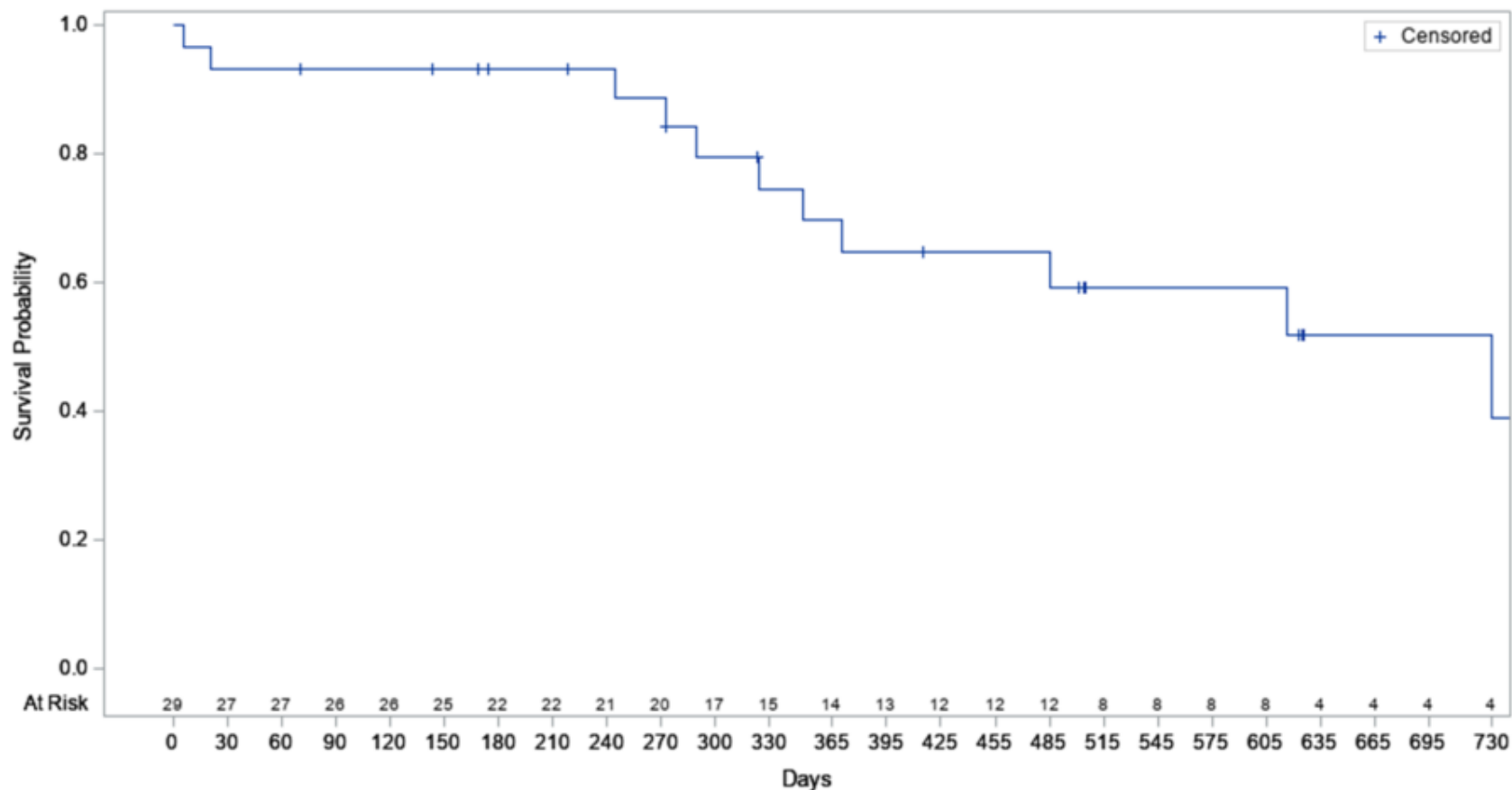
- ✓ In line with expectations – 3 composite primary safety events

## QUALITY OF LIFE

- ✓ Clinically important improvement maintained for up to 12 months post-implantation

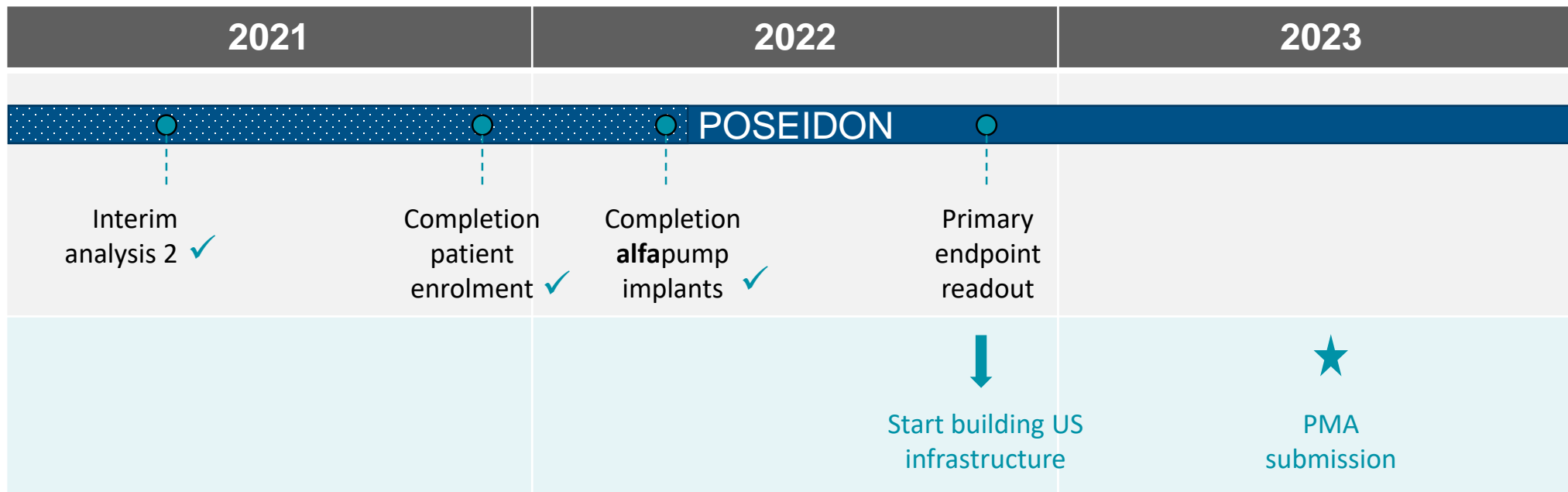
# Interim POSEIDON: 70% survival at 12 months

Preliminary survival rate analysis of Roll-In Cohort (25 March 2022)



**Mean survival probability of 70% at 12 months compares favourably to published literature reporting a survival rate for refractory ascites patients of only 50% at 12 months<sup>1</sup>**

# North American alfapump® approval on track for 2024



*NTAP for breakthrough devices de-risks reimbursement in key Medicare population\**



*\*On the basis of existing ICD-10 codes issued for the **alfapump**, the likely DRG coding will be 423, 424 and 425 "OTHER HEPATOBILIARY OR PANCREAS O.R. PROCEDURES"*

*PMA: Pre-Market Approval; NTAP: New Technology Add-On Payment*

# US – Go direct to 140 liver transplant centres

Highly efficient approach to target doctors and patients – driven by treatment guidelines



140 liver transplant centres



~50 person team initially

- 35 sales reps
- 10 clinical specialists
- 5 corporate



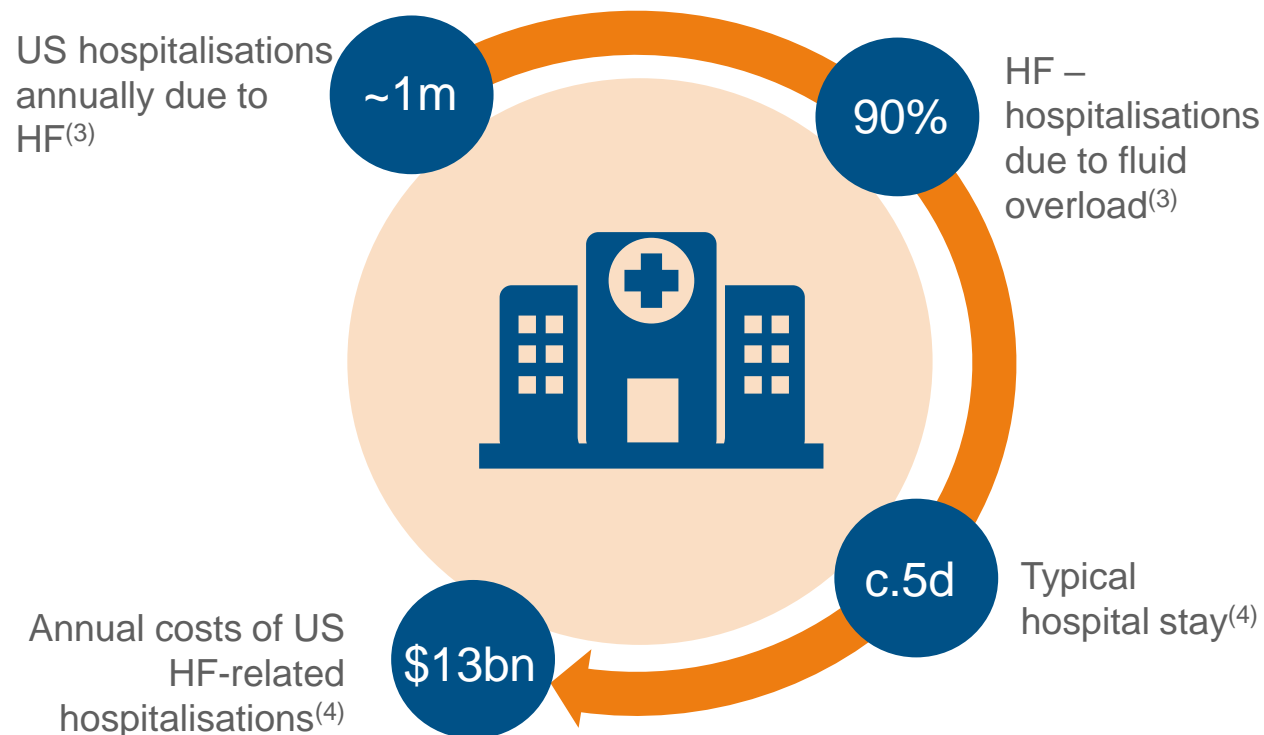
**DSR<sup>®</sup>**

Breakthrough approach to  
**persistent congestion in heart  
failure**



# Diuretic-resistant congestion in heart failure

Removal of congestion is a key therapeutic target and maintaining renal function is a clinical challenge



- 40% of heart failure patients on IV loop diuretics have a poor response<sup>(1)</sup>
- 24% re-admission rate at 30 days<sup>(2)</sup>

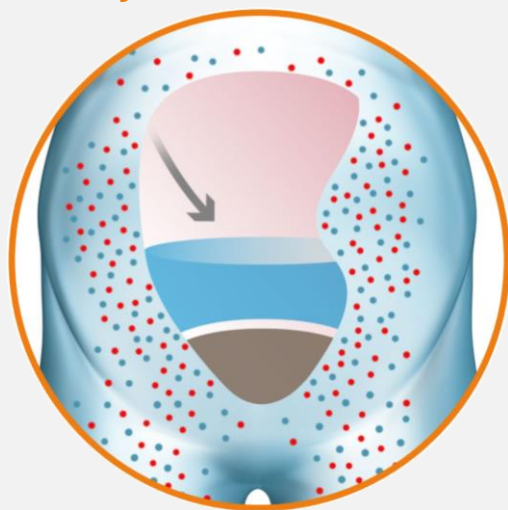


# Direct Sodium Removal (DSR<sup>®</sup>) platform

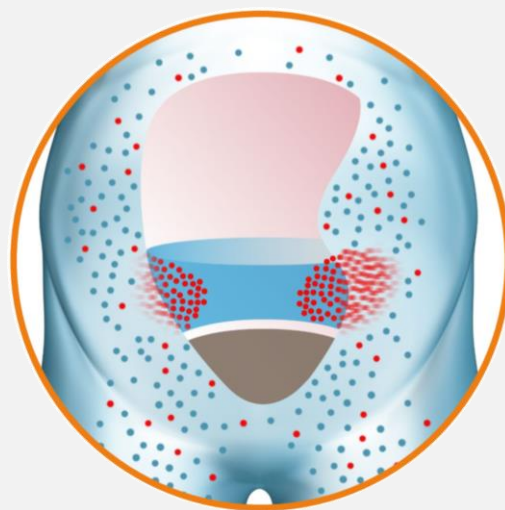
Eliminating fluid spread across the body – working in partnership with the kidneys



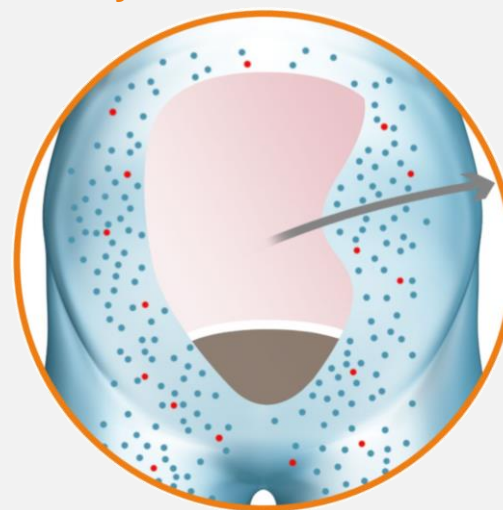
1 Sodium-free DSR infusate administered to peritoneal cavity



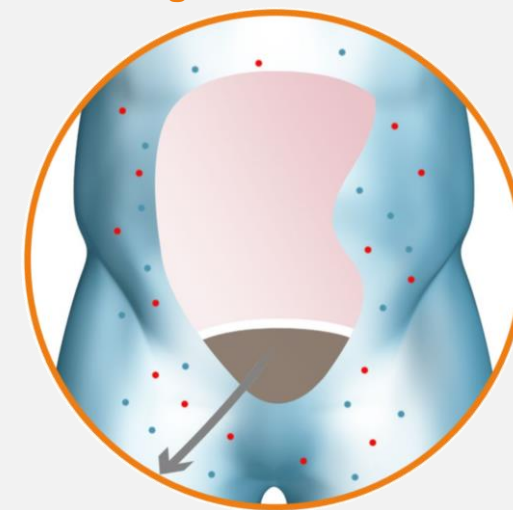
2 Sodium diffuses from body into DSR infusate



3 DSR infusate + extracted sodium removed from the body



4 Body eliminates free water to restore sodium balance, reducing the fluid overload

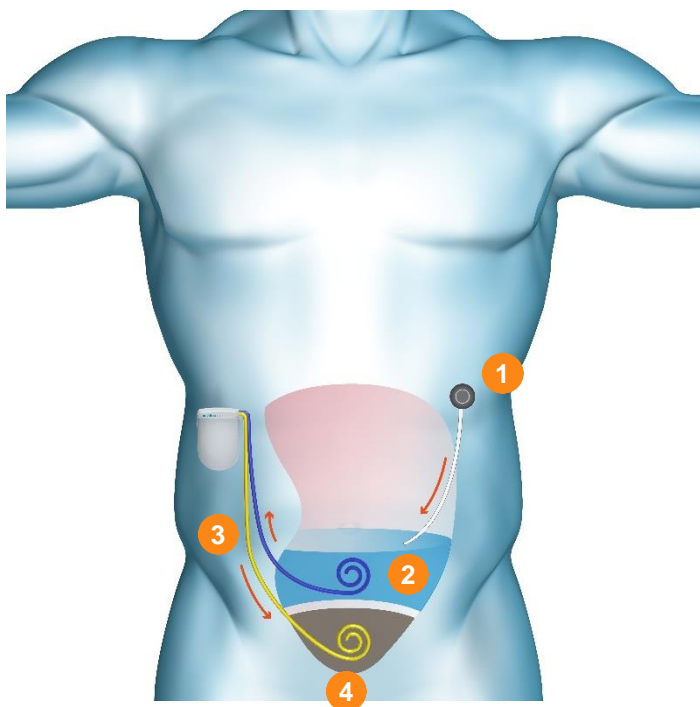


- water
- sodium

*Fundamental patents to reduce fluid overload in heart failure patients granted in the US and Europe*

# alfapump DSR<sup>®</sup> leveraging proven alfapump<sup>®</sup> platform

Fully implanted system for long-term DSR<sup>®</sup> therapy – keeping patients out of the hospital



- 1 Sodium-free DSR infusate administered to peritoneal cavity via implanted subcutaneous port
- 2 Sodium diffuses into DSR infusate
- 3 **alfapump** pumps sodium-rich DSR infusate into the bladder
- 4 Body eliminates excess fluid through osmotic ultrafiltration and urination

# DSR<sup>®</sup> – Encouraging phase 2a heart failure data

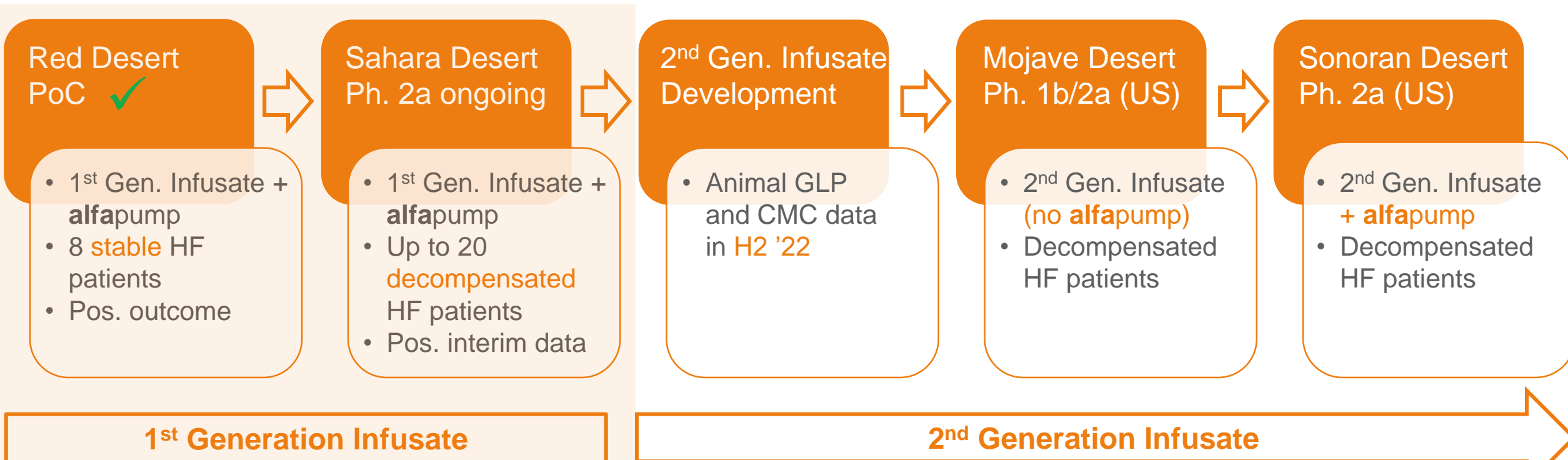
Clearing congestion while preserving renal function is a key objective of heart failure therapy

| RED DESERT – Completed  | SAHARA DESERT – Ongoing (Interim data)   |
|---|--|
| 8 Euvolemic heart failure patients  | 6 Decompensated heart failure patients   |
| <p>✓ <b>Clinical proof-of-concept</b></p> <p>✓ <b>Clear improvement in cardio-renal status</b></p> <ul style="list-style-type: none"> <li>• 30% decrease in NT-proBNP*</li> <li>• 22% increase in eGFR* / decrease in creatinine*</li> </ul> <p>✓ <b>Dramatic and durable improvement in diuretic response</b></p> <ul style="list-style-type: none"> <li>• 40-96% reduction in diuretic dose 9-19 months after study completion</li> </ul> | <p>✓ <b>Safe, effective &amp; rapid decongestion, &amp; restore euvolemia</b></p> <p>✓ <b>Clear improvement in cardio-renal status</b></p> <ul style="list-style-type: none"> <li>• &gt;30% decrease in NT-proBNP*</li> <li>• Stable eGFR* and creatinine*</li> </ul> <p>✓ <b>Dramatic and durable improvement in diuretic response</b></p> <ul style="list-style-type: none"> <li>• &gt;90% reduction in diuretic dose 3 months* after intensive DSR therapy</li> </ul> |

\* Mean value

**NT-proBNP:** N-terminal-pro hormone B-type Natriuretic Peptide (analysed in local lab); **eGFR:** estimated glomerular filtration rate

# Proprietary heart failure drug development programme



## 1<sup>st</sup> Generation Infusate

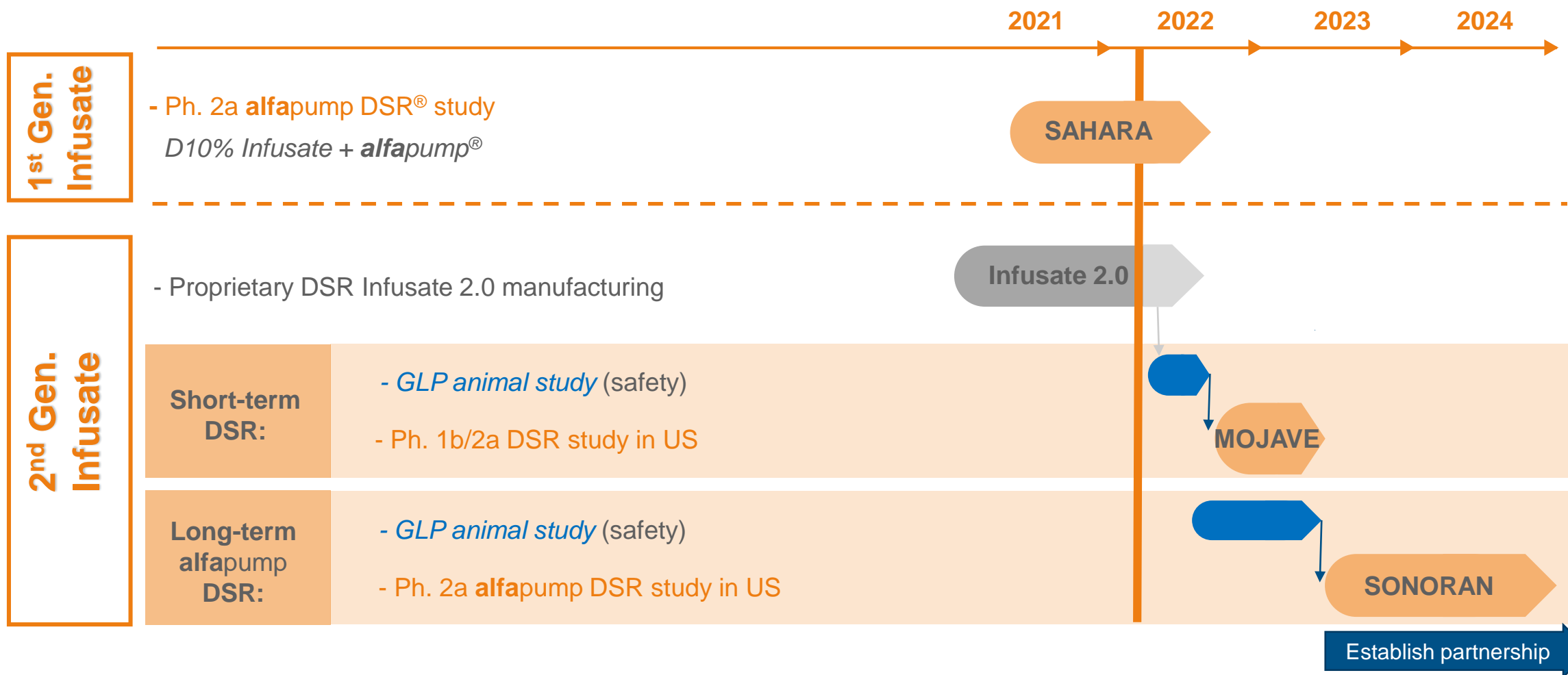
- ✓ Sodium-free D10% (off-the-shelf)
- ✓ Clinical Proof-of-Concept
- ✓ Rapid clinical path

## 2<sup>nd</sup> Generation Infusate

- ✓ Sodium-free dextrose / icodextrin (proprietary)
- ✓ Improved therapeutic profile
- ✓ Favorable safety profile
- ✓ IP protection drives recurring revenue from high gross margin consumable

# DSR<sup>®</sup> – plan to partner after US efficacy study

Step-by-step approach to introduction of breakthrough heart failure therapy



Timelines subject to further developments related to the ongoing COVID-19 pandemic  
 Description and timing of these studies are subject to change and/or feedback from applicable regulatory authorities



# Outlook

Strong **near term value drivers**  
with clear **long term potential**



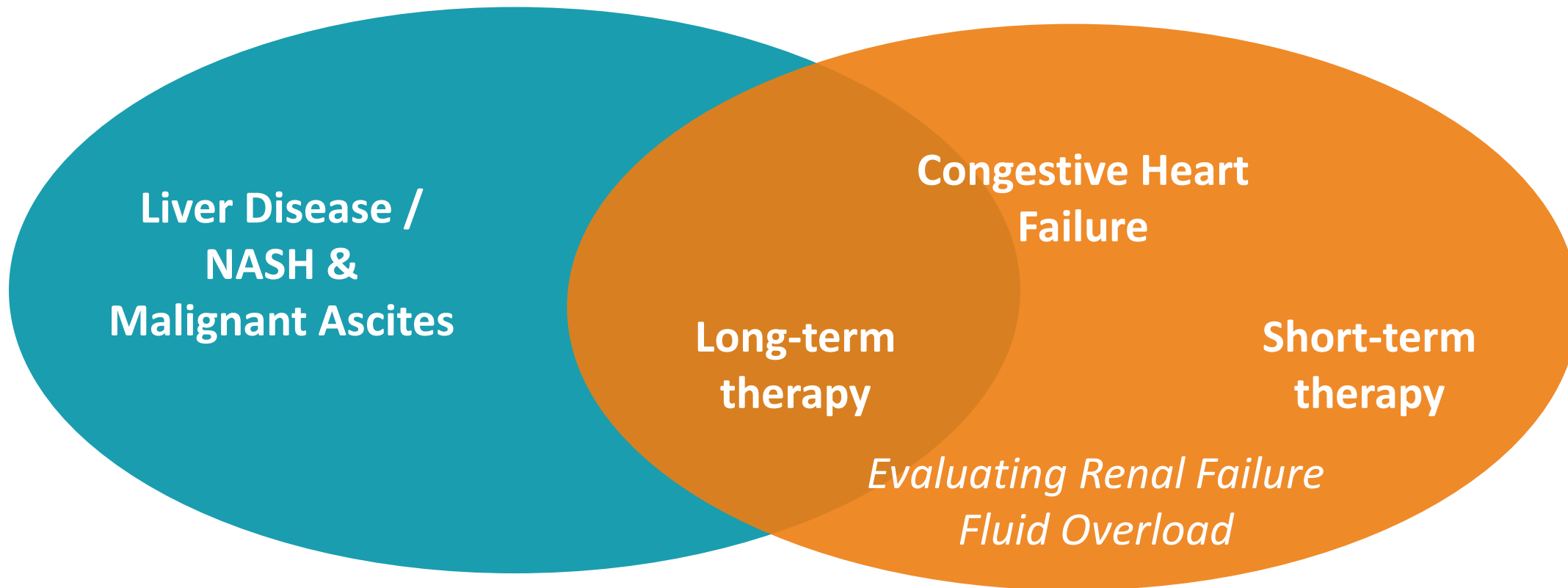
# Building on our two proprietary platforms

Complementary approaches to diuretic-resistant fluid overload

alfapump® 

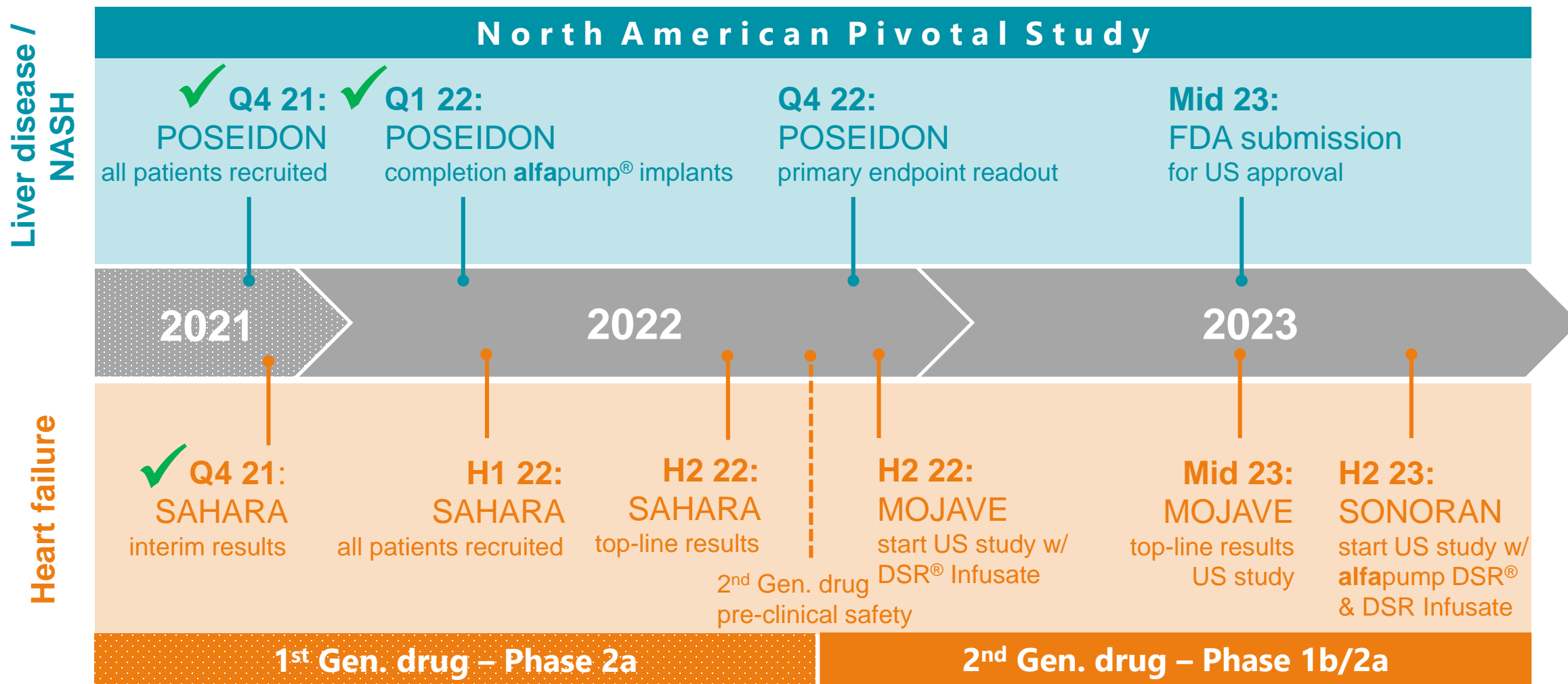
alfapump DSR®

DSR® 





# Strong outlook for value drivers



Note: Presented timelines are subject to further developments related to the COVID-19 pandemic

# Strongly positioned for growth in both our markets



- **alfapump<sup>®</sup> in liver disease / NASH – over €3 Bn / year <sup>(1)</sup>**
  - NASH is changing liver cirrhosis market and driving growth
  - FDA breakthrough device status / Strong IP portfolio
  - North American pivotal study de-risked – Fully implanted / Positive interim data
  - North American approval on track for 2024 / Go direct to 140 liver transplant centres



- **DSR<sup>®</sup> in heart failure – over €5 Bn / year <sup>(2)</sup>**
  - Clearing congestion while preserving renal function is a key objective of heart failure therapy
  - Clinical proof-of-concept with 1<sup>st</sup> Gen. drug – Encouraging phase 2a data
  - Development of proprietary 2<sup>nd</sup> Gen. drug – Strong IP / Driver of high margin recurring revenue
  - Establish partnership after US efficacy study mid-2023



# Contact info



[IR@sequanamedical.com](mailto:IR@sequanamedical.com)



+32 498 053579

[www.sequanamedical.com](http://www.sequanamedical.com)