



# Corporate Presentation

(NASDAQ: DCTH)

December 2021



## Forward-looking Statements

The Private Securities Litigation Reform Act of 1995 provides a safe harbor for forward-looking statements made by the Company or on its behalf. This presentation contains forward-looking statements, which are subject to certain risks and uncertainties that can cause actual results to differ materially from those described. Factors that may cause such differences include, but are not limited to, uncertainties relating to: the timing and results of the Company's clinical trials, including without limitation the mOM and ICC clinical trial programs, as well as the receipt of additional data and the performance of additional analyses with respect to the mOM clinical trial, our determination whether to continue the ICC clinical trial program or to focus on other alternative indications, and timely monitoring and treatment of patients in the global Phase 3 mOM clinical trial and the impact of the COVID-19 pandemic on the completion of our clinical trials; the impact of the presentations at major medical conferences and future clinical results consistent with the data presented; approval of Individual Funding Requests for reimbursement of the CHEMOSAT procedure; the impact, if any, of ZE reimbursement on potential CHEMOSAT product use and sales in Germany; clinical adoption, use and resulting sales, if any, for the CHEMOSAT system to deliver and filter melphalan in Europe including the key markets of Germany and the UK; the Company's ability to successfully commercialize the HEPZATO KIT/CHEMOSAT system and the potential of the HEPZATO KIT/CHEMOSAT system as a treatment for patients with primary and metastatic disease in the liver; our ability to obtain reimbursement for the CHEMOSAT system in various markets; approval of the current or future HEPZATO KIT/CHEMOSAT system for delivery and filtration of melphalan or other chemotherapeutic agents for various indications in the U.S. and/or in foreign markets; actions by the FDA or foreign regulatory agencies; the Company's ability to successfully enter into strategic partnership and distribution arrangements in foreign markets and the timing and revenue, if any, of the same; uncertainties relating to the timing and results of research and development projects; and uncertainties regarding the Company's ability to obtain financial and other resources for any research, development, clinical trials and commercialization activities. These factors, and others, are discussed from time to time in our filings with the Securities and Exchange Commission. You should not place undue reliance on these forward-looking statements, which speak only as of the date they are made. We undertake no obligation to publicly update or revise these forward-looking statements to reflect events or circumstances after the date they are made.

Delcath



# Executive Summary

Delcath aims to be the leader in targeted, safe and highly-effective minimally-invasive treatments for patients with cancers of the liver.

| UNMET NEED<br>LIVER CANCER   | PERCUTANEOUS HEPATIC<br>PERFUSION (PHP)  | COMPANY &<br>CLINICAL PROGRAM  | LARGE MARKET<br>OPPORTUNITY   |
|--|--|--|---|
| <p>Incidence US/EU</p> <ul style="list-style-type: none"><li>• &gt;200K primary and metastatic liver tumors per year<sup>2-14,29</sup></li></ul> <p>Current local/regional treatments</p> <ul style="list-style-type: none"><li>• Cannot treat the whole liver</li><li>• Targeted to visible and accessible tumors</li><li>• Limited in their ability to retreat</li></ul> | <p>PHP drug-device platform</p> <ul style="list-style-type: none"><li>• Delivers high dose chemotherapy to the entire liver</li><li>• Limits systemic exposure</li><li>• Minimally invasive, repeatable and well-tolerated</li></ul> <p>US: HEPZATO KIT<br/>EU: CHEMOSAT</p> | <p>FOCUS pivotal trial</p> <ul style="list-style-type: none"><li>• Metastatic Ocular Melanoma (mOM)</li><li>• Primary endpoint met*</li><li>• NDA submission mid '22</li></ul> <p><b>Real World Evidence</b></p> <ul style="list-style-type: none"><li>• &gt;1k commercial treatments in EU</li><li>• Multiple single center publications</li></ul> <p>ANTICIPATED FDA APPROVAL: Q4 2022</p> | <p>Near-term (mOM)</p> <ul style="list-style-type: none"><li>• &gt;\$300M TAM in US and EU</li><li>• No effective standard of care</li></ul> <p>Longer Term (CRC, ICC, Pancreatic, etc.)</p> <ul style="list-style-type: none"><li>• &gt;&gt;\$1B TAM</li><li>• Investigator interest in more than 10 other tumor types</li></ul> |

Delcath

Metastatic Ocular Melanoma (mOM)<sup>2,3</sup>, Cholangiocarcinoma (ICC)<sup>4,5</sup>, Liver-dominant Breast Cancer (mBC)<sup>8-11</sup>, Metastatic Neuroendocrine Tumors (mNET)<sup>6,7</sup>, Metastatic Pancreatic Cancer (mPC)<sup>5,16</sup>, Metastatic Colorectal Cancer (mCRC)<sup>12,13</sup>, Hepatocellular carcinoma (HCC)<sup>29</sup>

# Liver-Dominant Cancers

High incidence with poor prognosis

Up to  
**80%**

Many patients with liver metastases are not amenable to surgical resection largely due to extensive tumor burden<sup>1</sup>



**Liver: Common Site of Metastases**



**Limited Effective Systemic Treatments**

- » Systemic therapies - low efficacy
- » Immuno-oncology agents - become less effective in the presence of metastases



**Limited Overall Survival – Unresectable Liver Cancer**

- » Often the life-limiting organ

## US Incidence of Liver Dominant Cancers

(partial set shown)

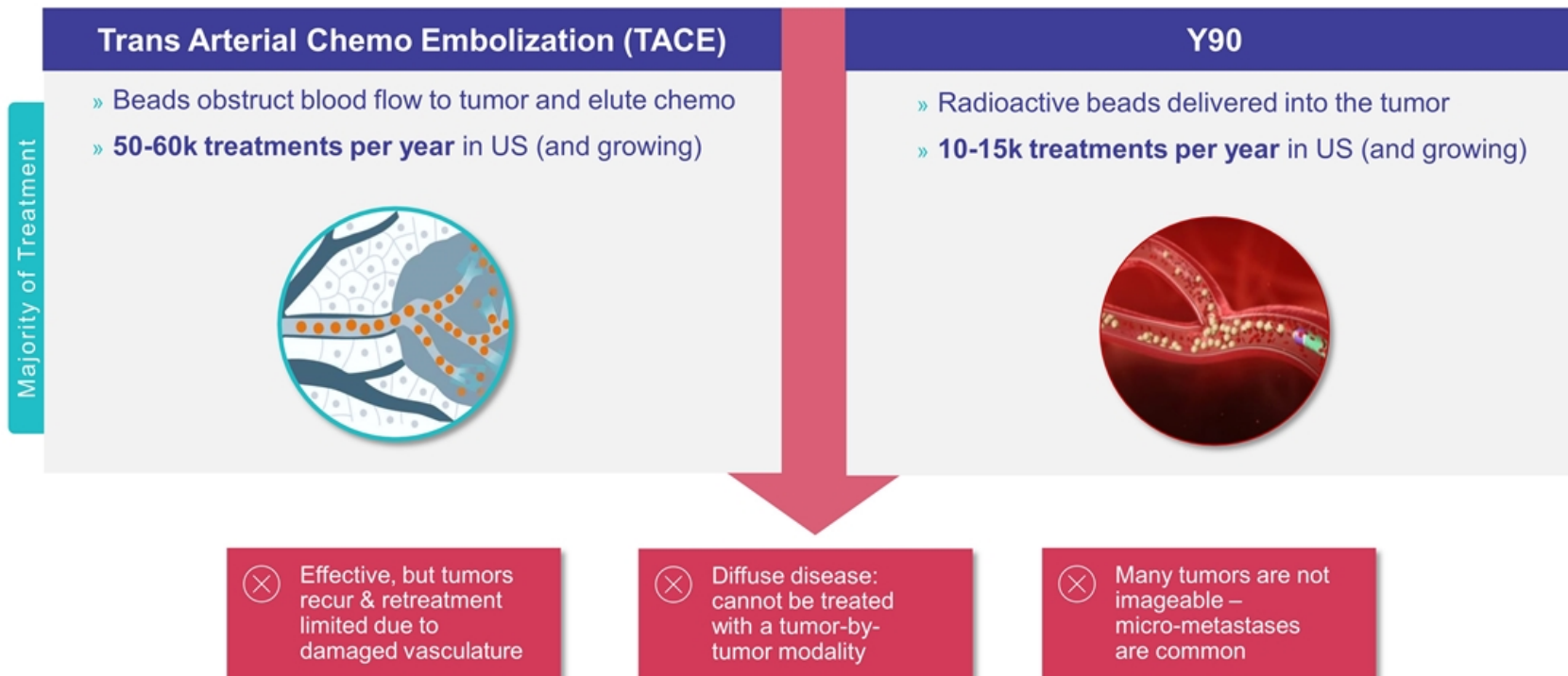


Delcath

Metastatic Ocular Melanoma (mOM)<sup>2,3</sup>, Cholangiocarcinoma (ICC)<sup>4,5</sup>, Liver-dominant Breast Cancer (mBC)<sup>6-11</sup>, Metastatic Neuroendocrine Tumors (mNET)<sup>12,13</sup>, Metastatic Pancreatic Cancer (mPC)<sup>14,15</sup>, Metastatic Colorectal Cancer (mCRC)<sup>16,17</sup>, Hepatocellular carcinoma (HCC)<sup>18</sup>

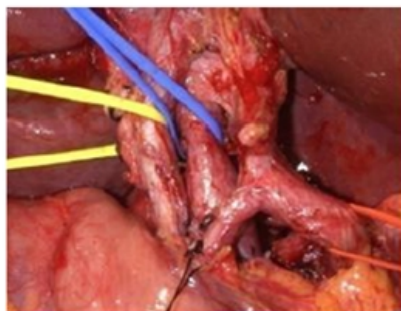


# Limitations of Current Liver-Directed Therapies



# Isolated Hepatic Perfusion (IHP)

## The pathway to developing Percutaneous Hepatic Perfusion



### Benefits

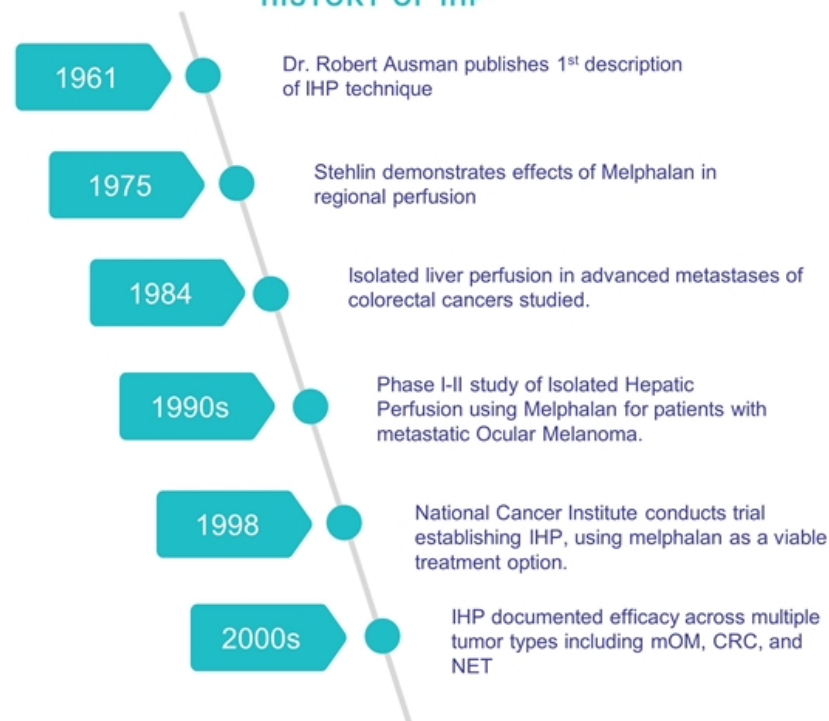
- » Temporarily isolates liver blood supply
- » Delivers substantially higher concentrations of chemotherapy (Melphalan) with limited toxicity



### Limitations

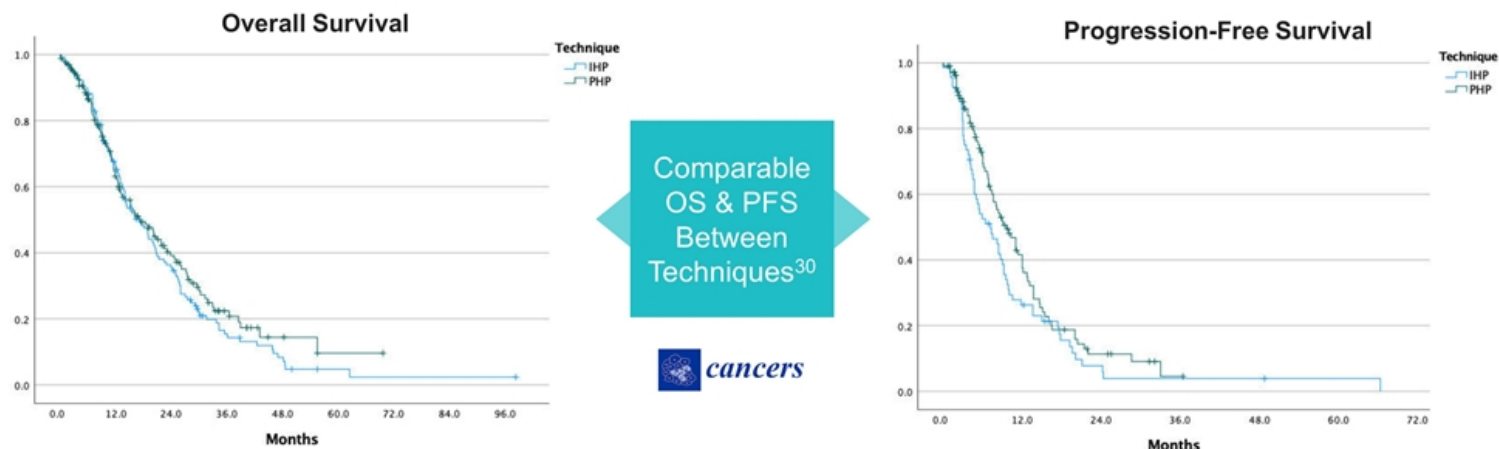
- » High treatment related mortality (>5%)
- » Not repeatable and few patients are eligible

### HISTORY OF IHP<sup>1</sup>



Delcath

# PHP Advances IHP Clinical Benefits

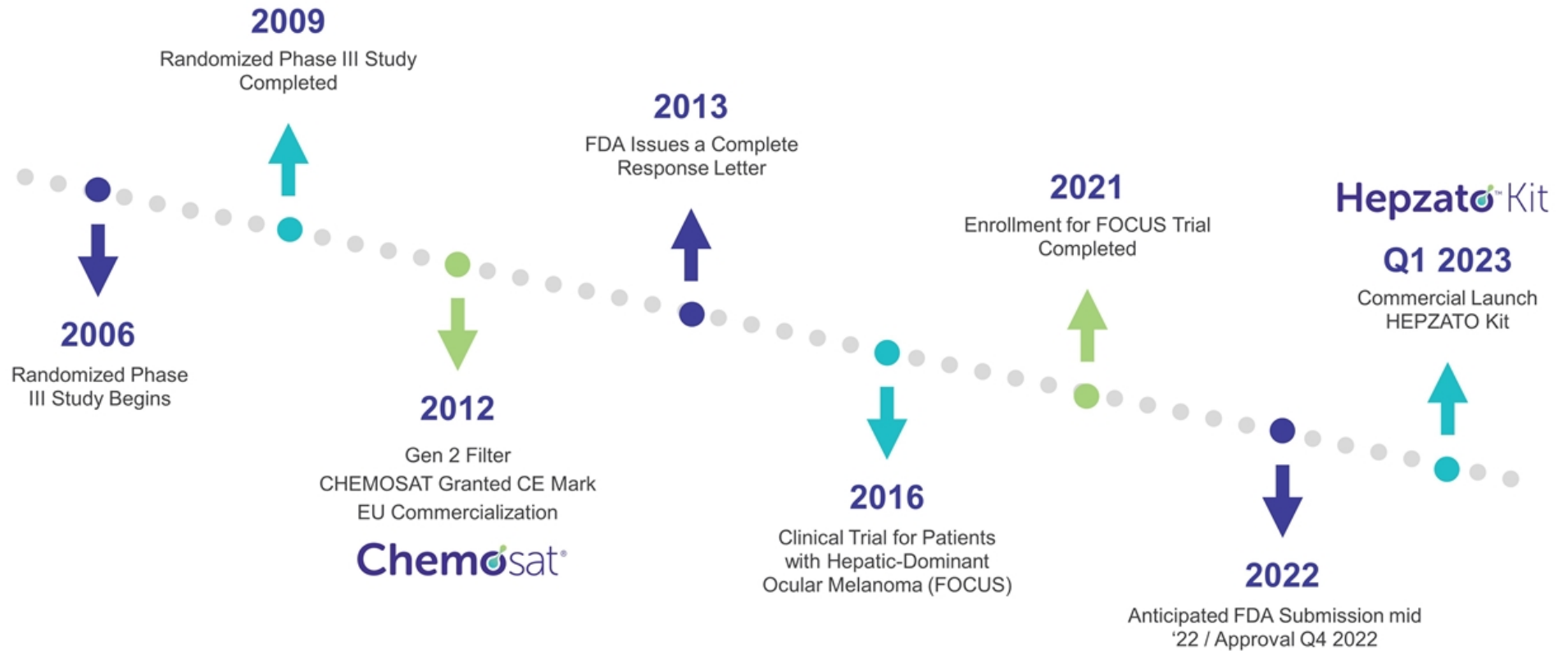


“ There was no difference in overall survival (OS) or progression-free survival (PFS) between IHP and PHP for patients with uveal melanoma liver metastases, but patients have **significantly less of a risk for complications and mortality following PHP.**”

Delcath



# History of HEPZATO Kit Development

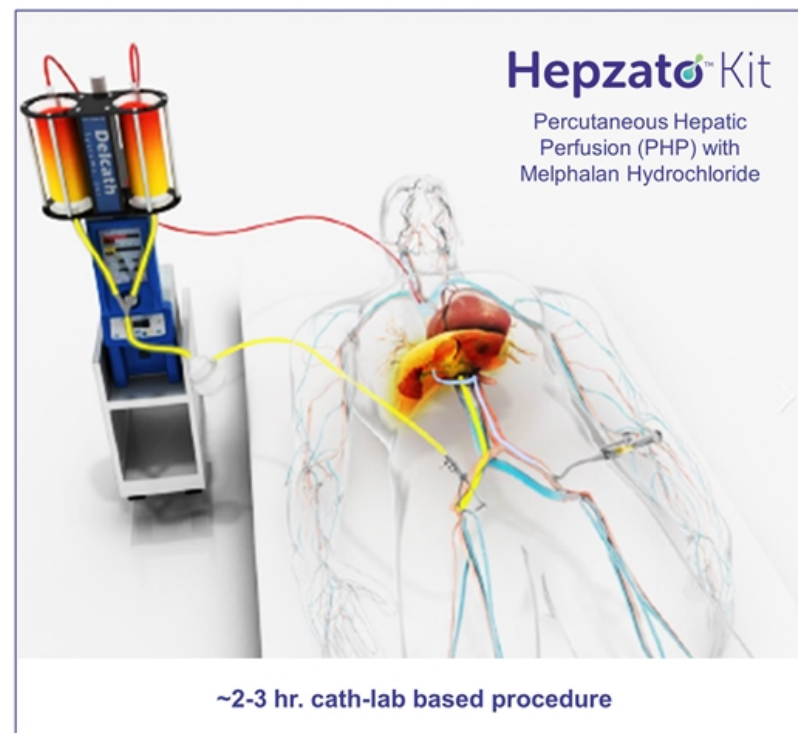


# HEPZATO™ Kit: Percutaneous Hepatic Perfusion (PHP)

Repeatable, safe & effective liver-focused disease control

## Next-Generation, Minimally-Invasive Liver-Directed Treatment

The only minimally invasive cancer treatment that isolates the liver from systemic circulation, allowing for repeated delivery of high-dose chemo to the entire liver while limiting systemic side effects.



Delcath

# Three Steps. Targeted Treatment.

## Hepzato™ Kit

Novel, whole-organ treatment that provides targeted, high-dose liver chemo while minimizing systemic exposure.

1

### ISOLATION

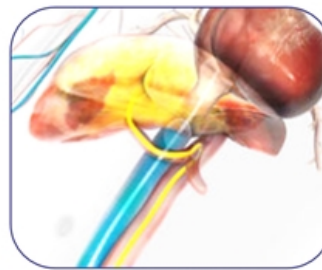
Hepatic venous flow is isolated, enabling 12x increased dose



2

### SATURATION

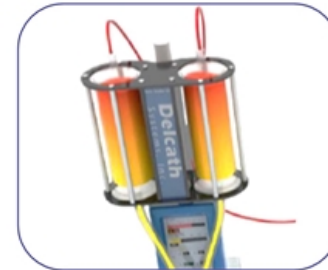
Melphalan (chemo) treats micro and macro lesions simultaneously



3

### FILTRATION

Proprietary filters remove greater than 85% of chemo from the body<sup>1\*</sup>



Delcath

\*In vitro model - Data on file  
1. de Leede E., et al. Cardiovascular Intervent Radiol. 2017 Aug;40(8):1196-1205.



# mOM: Beachhead Market Opportunity

No FDA-approved treatment, no current standard of care



## Unmet Need

- » ~6,000 cases of ocular melanoma per year in the US/EU<sup>13,17</sup>
- » 50% metastasize, 90% to the liver<sup>3,14</sup>
- » Median survival up to 12 months.<sup>15</sup>

## Low Risk Opportunity

- » FOCUS pivotal trial has met primary endpoints to support approval in mOM<sup>19</sup>
- » Significantly improved safety profile over Gen 1 filter technology
- » Real world safety and efficacy demonstrated in EU

## High Barrier to Entry

- » EXCLUSIVE: Granted orphan indication status allows for extended exclusivity
- » HEPZATO is a combination drug device regulated by CDER – no ANDA pathway
- » Melphalan granted orphan indication

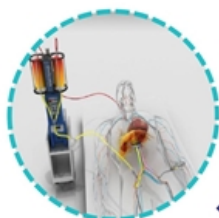
## Favorable Commercial Economics

- » Payer/hospital financial stakeholder interviews suggest expected pricing is on par with immuno-oncological agents ~\$250k annually
- » 20 US treatment centers = ~80% patients

Delcath

# Competitive Landscape for mOM

HEPZATO™ is the only highly-effective, targeted mOM treatment that enables repeat treatments while optimizing QoL



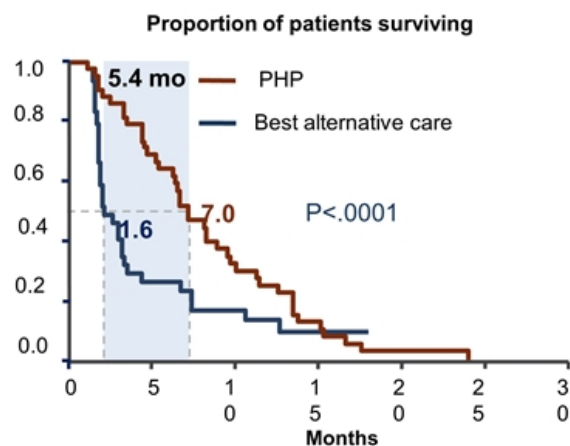
|                                    | Minimally Invasive – Liver Directed |                    |                        | Infusion – Systemic         |                            |
|------------------------------------|-------------------------------------|--------------------|------------------------|-----------------------------|----------------------------|
|                                    | HEPZATO™                            | TACE <sup>23</sup> | Y90/SIRT <sup>21</sup> | Mono/Combo IO <sup>24</sup> | Tebentafusp <sup>22*</sup> |
| High Efficacy<br>ORR %             | 31.4%                               | <21%               | <17%                   | 5.5%                        | Up to 9% <sup>25</sup>     |
| OS at 12 months<br>(% surviving)   | 75%**                               | -                  | -                      | -                           | 73%***                     |
| Repeatable (>3x)                   | ✓                                   | X / ✓              | X                      | ✓                           | ✓                          |
| Preserves QoL                      | ✓                                   | ✓                  | ✓                      | X                           | ✓                          |
| FDA Approved for mOM               | Q4 2022                             | X                  | X                      | Melanoma                    | Pending                    |
| Applicable to most mOM<br>patients | ✓                                   | ✓                  | ✓                      | ✓                           | X                          |

Delcath

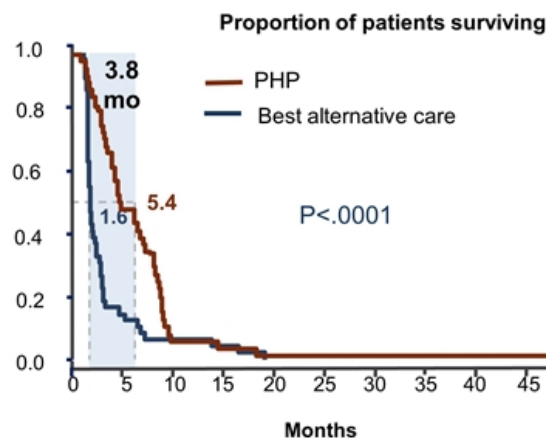
\*HLA A+ patient indication only      \*\*mITT, BAC OS 47%, HR 0.37, 95% CI 0.17, 0.79, p-value 0.01  
\*\*\*Control OS 59%, HR 0.51, 95% CI 0.37, 0.71, p-value <0.001

# First Phase 3 RCT Results\*

**Hepatic Progression Free Survival  
(IRC Assessment)**



**Overall Progression Free Survival  
(INV Assessment)**



**Response Rates  
(ITT population)**

| Cohort | PHP<br>(N=44) | BAC<br>(N=49) | P-<br>Value |
|--------|---------------|---------------|-------------|
| hOR    | 36.4%         | 2.0%          | <0.001      |
| ORR    | 27.3%         | 4.1%          | =0.003      |

**Crossover design confounded overall survival analysis –  
most subjects in BAC arm [57.1%] crossed over to PHP arm**

Delcath

\*Mix of mOM and metastatic melanoma with >90% patients diagnosed with mOM - NDA 201848 Clinical Study Report dated 15 August 2012.



# Safety Issues and Resulting Improvements

## Safety Issue

Hematological toxicities led to 3 patient deaths

| Adverse Event<br>G3/4 | Gen 1<br>Hughes 2016 <sup>20</sup> |    |
|-----------------------|------------------------------------|----|
|                       | %                                  | n  |
| Anemia                | 62.9%                              | 44 |
| Neutropenia           | 85.7%                              | 60 |
| Thrombocytopenia      | 80.0%                              | 56 |

**Inappropriate patient selection and procedural issues** led to 1 patient death and other AE's

- ~90% liver involvement causing tumor lysis syndrome

## Improvement


Gen 2 Filter introduced in 2013

| Adverse Event<br>G3/4 | Gen 2<br>Karydis 2018 <sup>15</sup> |    | % Improvement<br>Gen 1 → 2 |
|-----------------------|-------------------------------------|----|----------------------------|
|                       | %                                   | n  |                            |
| Anemia                | 29.4%                               | 15 | 53% ↓                      |
| Neutropenia           | 31.3%                               | 16 | 64% ↓                      |
| Thrombocytopenia      | 31.3%                               | 16 | 61% ↓                      |

- Protocol amendments were put in place for patient selection
- Training improved

FDA required these issues be addressed prior to the start of the FOCUS trial

Delcath



## FOCUS Trial

### 2<sup>nd</sup> Registration Clinical Trial for Patients with mOM

## FOCUS

### OVERVIEW:

- Multinational, multicenter, single-arm trial
- Endpoints:
  - » Primary: Objective Response Rate compared to historic control
  - » Secondary: Duration of response, disease control rate, overall survival, progression free survival, safety, PK, QoL
- 102 subjects enrolled, 91 completed treatments at 30 centers in the US and EU
- HEPZATO Tx every 6-8 weeks up to a maximum of 6 cycles

# FOCUS Trial Analysis: Prespecified Endpoint Met

Intent to Treat:

| Primary Effectiveness Endpoint <sup>19</sup> | PHP<br>(N=91 treated + 11 untreated) | 95% CI*       |
|--|--------------------------------------|---------------|
| Objective Response Rate                      | 31.4%                                | [22.55-41.31] |

\*A meta-analysis of checkpoint inhibitors (476 patients, 16 publications) calculated a 95% Confidence Interval for ORR of 3.6% - 8.3%\*

Lower bound 22.55% far exceeds  
8.3% upper bound prespecified  
threshold.

PRELIMINARY DATA - SUBJECT TO CHANGE

Delcath

## Hematological Toxicities - Comparison with Previous Trials

| Grade 3 or higher Adverse Events | Focus Trial<br>(n=91) | Hughes 2016<br>(n=70) |
|----------------------------------|-----------------------|-----------------------|
| Anemia                           | 27 (29.7%)            | 44 (62.9%)            |
| Thrombocytopenia                 | 24 (26.4%)            | 56 (80.0%)            |
| Neutropenia                      | 18 (19.8%)            | 60 (85.7%)            |

Hematological AE's consistent with  
European experience

# FOCUS Trial – Safety Comparison with Previous Trials

| Category                                  | FOCUS Trial<br>(N=91) | Pooled Analysis of<br>Prior Studies (N=121) |
|---|-----------------------|---|
| Patients who Withdrew due to an AE or SAE | 20 (22%)              | 46 (38%)                                    |
| Patients who Required a Dose Reduction    | 12 (13.2%)            | 27 (22.3%)                                  |
| Average Number of Cycles                  | 4.1                   | 2.8   |

Improvement in tolerability led to a larger number of treatments



# Recent Initial Approvals Using ORR in Single-Arm Oncology Trials

| Single trial<br>n=50  |   |  |   |  |   |  |  |  |
|---|---|--|---|--|---|--|--|--|
| Danyelza<br>(naxitamab-gqgk)  | Gavreto<br>(pralsetinib)                    | Monjuvi<br>(tafasitamab-cxix)                        | Tazverik<br>(tazemetostat)  | Zepzelca<br>(lurbinectedin)  | Tabrecta<br>(capmatinib)                            | Trodelvy<br>(sacituzumab)  | Pemazyre<br>(pemigatinib)  | Koselugo<br>(selumetinib)  |
| Accelerated   | Accelerated                                 | Accelerated  | Accelerated   | Accelerated  | Accelerated   | Accelerated  | Accelerated  | Accelerated  |
| Relapsed or refractory neuroblastoma in bone or marrow post response or stable disease to prior therapy | Metastatic <i>RET</i> fusion-positive NSCLC | Relapsed or refractory diffuse large B-cell lymphoma | Relapsed or refractory follicular lymphoma positive for EXH2 mutation | Metastatic SMLC with progression on or after platinum chemotherapy | Metastatic NSCLC with mutation MET exon 14 skipping | Metastatic triple-negative breast cancer after at least 2 prior metastatic disease therapies | Previously treated metastatic cholangiocarcinoma with FGFR2 fusion | Neurofibromatosis Type 1 with inoperable plexiform neurofibromas |

| Single trial<br>N=43   |   |   |  | Pooled subgroup<br>analysis n=51   3<br>single arm trials |   |  |  | Pooled subgroup<br>analysis n=72   2<br>single arm trials |
|--|---|---|--|---|---|--|--|---|
| Ayvakit<br>(avapritinib)   | Enhertu<br>(famtrastuzumab deruxtecan)  | Padcev (enfortumab vedotin)   | Brukinsa<br>(zanubrutinib)                           | Rozlytrek<br>(entrectinib)                                | Xpovio (selinexor)  | Balversa<br>(erdafitinib)  | Vitrakvi<br>(larotrectinib)                                    | Libtayo<br>(cemiplimab-rwlc)                              |
| Standard   | Accelerated   | Accelerated   | Accelerated  | Standard  | Accelerated   | Accelerated  | Accelerated  | Standard  |
| Unresectable or metastatic gastrointestinal stromal tumor with PDGFRA exon 18 mutation | Unresectable or metastatic HER2+ breast with two or more prior anti HER2 regimens in metastatic setting | Metastatic urothelial cancer with previously received PD-1 or PD-L1 and platinum chemotherapy | Mantle cell lymphoma with at least one prior therapy | Metastatic NSCLC that is <i>ROS1</i> +                    | Relapsed or refractory multiple myeloma with at least 4 prior therapies | Metastatic urothelial carcinoma with susceptible FGFR 3(2) alterations | Solid tumors with neurotrophic receptor tyrosine kinase fusion | Metastatic squamous cell carcinoma                        |

## Supportive Evidence: Comparison Versus BAC

| Best Alternative Care (BAC) Arm        | Enrolled<br>N=42 | Treated<br>N=32 |
|--|------------------|-----------------|
| Dacarbazine                            | 1                | 0               |
| Ipilimumab                             | 7                | 1               |
| Pembrolizumab                          | 8                | 6               |
| Transarterial Chemoembolization (TACE) | 26               | 25              |

### Amended Study

- » FOCUS was initially a RCT against Best Alternative Care (BAC)
- » Due to enrollment challenges as a result of known limited efficacy of BAC control arm and availability of treatment with PHP (CHEMOSAT), FDA agreed to amend it to single-arm, non-RCT

# FOCUS Trial – Exploratory Analyses vs BAC

## Statistically Significant ORR and DCR Advantage vs. BAC

Intent to Treat:

| Efficacy Endpoint                 | PHP<br>(N=102)  | BAC<br>(N=42)   | P-Value* |
|-----------------------------------|-----------------|-----------------|----------|
| Objective Response Rate - Primary | 32 (31.4%)      | 4 (9.5%)        | 0.0059   |
| 95% CI                            | [22.55 - 41.31] | [2.66 - 22.62]  |          |
| Disease Control Rate              | 67 (65.7%)      | 12 (28.6%)      | <0.0001  |
| 95% CI                            | [55.63 - 74.81] | [15.72 - 44.58] |          |

Modified Intent to Treat\*\*:

| Efficacy Endpoint       | PHP<br>(N=91)   | BAC<br>(N=32)   | P-Value* |
|-------------------------|-----------------|-----------------|----------|
| Objective Response Rate | 32 (35.2%)      | 4 (12.5%)       | 0.0154   |
| 95% CI                  | [25.44 – 45.88] | [3.51 – 28.99]  |          |
| Disease Control Rate    | 67 (73.6%)      | 12 (37.5%)      | 0.0002   |
| 95% CI                  | [63.35 - 82.31] | [21.10 - 56.31] |          |

\*Chi-square

\*\* mITT Population – any patient who received at least one study treatment

Delcath

## FOCUS Trial – Exploratory Analyses vs BAC

ORR Advantage Coupled With Meaningful Duration of Response

|                                       | mITT Population |               |
|---------------------------------------|-----------------|---------------|
|                                       | PHP<br>(N=91)   | BAC<br>(N=32) |
| Duration of Response<br>(DOR, median) | 14.00 mos.      | NC            |
| 95% CI                                | [8.54 - NC]     | [6.93 - NC]   |
| Patients with Confirmed<br>CR or PR   | 32              | 4             |
| Patients with Subsequent PD           | 14 (43.7%)      | 1 (25.0%)     |
| Censored                              | 18 (56.3%)      | 3 (75.0%)     |

# FOCUS Trial - Exploratory Analyses vs BAC

PHP Progression-Free Survival ~3X that of BAC<sup>19</sup>

| Secondary Endpoint               |          | PHP<br>(N=91)*  | BAC<br>(N=32)* | P-Value* |
|----------------------------------|----------|-----------------|----------------|----------|
| Median Progression-Free Survival |          | 9.03 mos.       | 3.12 mos.      | 0.0007   |
| 95% CI                           |          | [6.34 - 11.56]  | [2.89 - 5.65]  |          |
| PFS Status                       | Events   | 64 (70.3%)      | 25 (78.1%)     |          |
|                                  | Censored | 27 (29.7%)      | 7 (21.9%)      |          |
| Hazard Ratio Estimate            |          | 0.39            |                | 0.0002   |
| 95% CI                           |          | [0.237 - 0.643] |                |          |

\* Treated patients only, per the protocol untreated patients were not followed

PRELIMINARY DATA - SUBJECT TO CHANGE

Delcath



## Focus Trial Results – 12 Month Survival\*

Intent to Treat:

| Secondary Endpoint       | PHP<br>(N=102) | BAC<br>(N=42) |
|--------------------------|----------------|---------------|
| % Surviving at 12 months | 68%            | 36%           |
| Hazard Ratio**           | 0.42           |               |
| 95% CI                   | 0.20 - 0.88    |               |
| p-value                  | 0.0215         |               |

Modified Intent to Treat\*\*\*:

| Secondary Endpoint       | PHP<br>(N=91) | BAC<br>(N=32) |
|--------------------------|---------------|---------------|
| % Surviving at 12 months | 75%           | 47%           |
| Hazard Ratio*            | 0.37          |               |
| 95% CI                   | 0.17, 0.79    |               |
| p-value                  | 0.010         |               |

\* Post Hoc analysis

\*\* Log Rank Test

\*\*\* mITT Population – any patient who received at least one study treatment

# Focus Trial Results – Overall Survival

Data still maturing  
PHP enrollment ended  
in May 2020, BAC in  
2018  
OS will be analyzed 24  
months post last  
patient last treatment

## Intent to Treat:

| Secondary Endpoint            |          | PHP<br>(N=102)* | BAC<br>(N=42)* | P-Value* | OS<br>pa |
|-------------------------------|----------|-----------------|----------------|----------|----------|
| Overall Survival (OS, Median) |          | 19.25 mos.      | 14.06 mos.     | 0.2021   |          |
| 95% CI                        |          | [16.30 – 24.35] | [9.99 – 19.78] |          |          |
| OS Status                     | Events   | 66 (64.7%)      | 23 (54.8%)     |          |          |
|                               | Censored | 36 (35.3%)      | 19 (45.2%)     |          |          |
| Hazard Ratio Estimate         |          | 0.739           |                | 0.2308   |          |
| 95% CI                        |          | [0.451 – 1.212] |                |          |          |

## Modified Intent to Treat\*\*:

| Secondary Endpoint            |          | PHP<br>(N=91)*  | BAC<br>(N=32)* | P-Value* |
|-------------------------------|----------|-----------------|----------------|----------|
| Overall Survival (OS, Median) |          | 20.53 mos.      | 14.06 mos.     | 0.1626   |
| 95% CI                        |          | [16.59 – 24.53] | [9.99 – 19.78] |          |
| OS Status                     | Events   | 64 (70.3%)      | 23 (71.9%)     |          |
|                               | Censored | 27 (29.7%)      | 9 (28.1%)      |          |
| Hazard Ratio Estimate         |          | 0.708           |                | 0.1725   |
| 95% CI                        |          | [0.431 – 1.163] |                |          |

\*Chi-square

\*\* mITT Population – any patient who received at least one study treatment

Delcath

# mOM Beachhead Market Strategy

## BEACHHEAD MARKET | mOM

## LIVER DISEASE



### SIGNIFICANT REVENUE OPPORTUNITY:

- Oncologists\* believe ~80% of mOM patients would be HEPZATO candidates - ~800 patients
- Considered a significant advancement over other therapies
- Payer & hospital finance stakeholders suggest pricing expectations in the range of IO agents - ~\$256k per yr.
- May be positioned as a first-line treatment due to limited efficacy of available therapies.

US TAM  
**>\$200M**  
per year

\*Source: Boston Health Associates primary research n=13 physicians

Delcath

# Experienced Interventional Oncology Leadership

- Kevin Muir-VP Commercial

- Formerly Head of Sales for US Therasphere Y90 (BTG/Boston Scientific)
- Led sales revenue growth from \$60M to \$220M
- Built sales team to focus on all members of the MDT

- Michael Ujhelyi - US Medical Director

- Formerly Head of Medical Affairs US Therasphere (BTG/Boston Scientific)
- Built Medical Science Liaison Team
- Responsible for Clinical Trial recruitment and IISs and IITs



# Specialized, Targeted Sales Team Will Leverage Expanded Access Protocol (EAP) and Longitudinal Data

## EAP (FDA Approved)

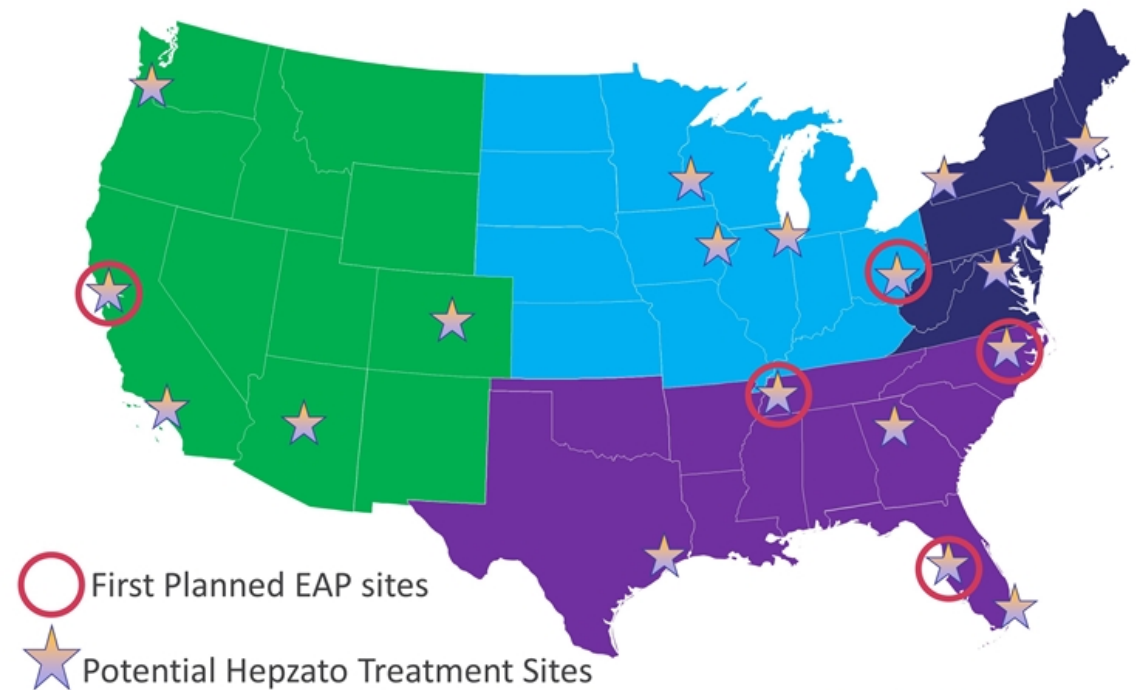
- Provide immediate access to patients
- First Commercial Sites
- Train new medical teams to use Hepzato after launch

## Regional Based Sales Team

- Experienced, Oncology focused
- Upon launch, placed in key geographies
- Supplement with Clinical Support Specialist

## Leverage Longitudinal Data

- Partnered with data provider to access patient level longitudinal data with 3-week refresh
- Accurately map and quantify surveillance, referral and treatment patterns at the patient and MD level



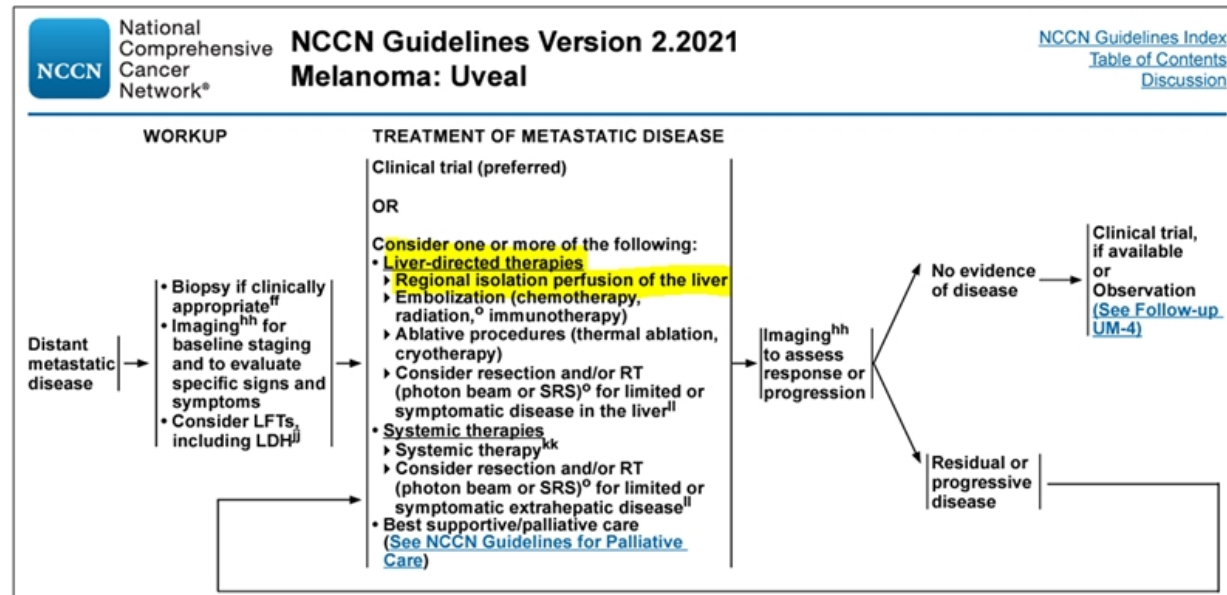
Delcath



# PHP Is Likely Part of Current NCCN Guidelines for mOM

“Regional Isolation Perfusion of the Liver”

PHP- Percutaneous Hepatic Perfusion



# Reimbursement

HEPZATO will be billed as a drug with a J-Code

- Medicare Patients

- Majority of patients will be outpatient (2 midnight rule) with the drug directly covered by Medicare
- For patients which become inpatient patients split billing (inpatient / outpatient) allows the drug to still be directly billed (e.g., not paid under a DRG)

- Private Payer Patients

- Private Payers for rare disease generally follow Medicare guidelines and we expect these patients to be treated as outpatients
- Prior-Authorization of patients might be needed, we are planning to contract out a hub service
- Centers of Excellence (PPS exempt and NCI designated Cancer Centers) have the leverage to negotiate favorable rates and reimbursement terms (our target sites are all either PPS exempt or NCI Cancer Centers)

# EU – Broad Reimbursement Pending Focus Trial Data, But Strong Interest Across Multiple Indications



- » CE Marked - available in ~23 centers in 4 countries
- » Currently distributed by MEDAC Pharma
- » MEDAC has been notified of our intent to terminate – discussions ongoing



- » NICE (UK) upgraded status from “Research” to “Special Status”
- » German reimbursement based on annual hospital special request (“ZE” process)



- » Strong interest to fuel additional indications driven by HCP's



- » 1,343 commercial Chemosat kits shipped to the EU
- » Queensbury facility has been inspected 21 times by the Notified Bodies LRQA and BSI, Health Authorities FDA and ANVISA

## CHEMOSAT Used In 13 Tumor Types

~70%: Metastatic Ocular Melanoma (mOM)

### Other Types Treated:

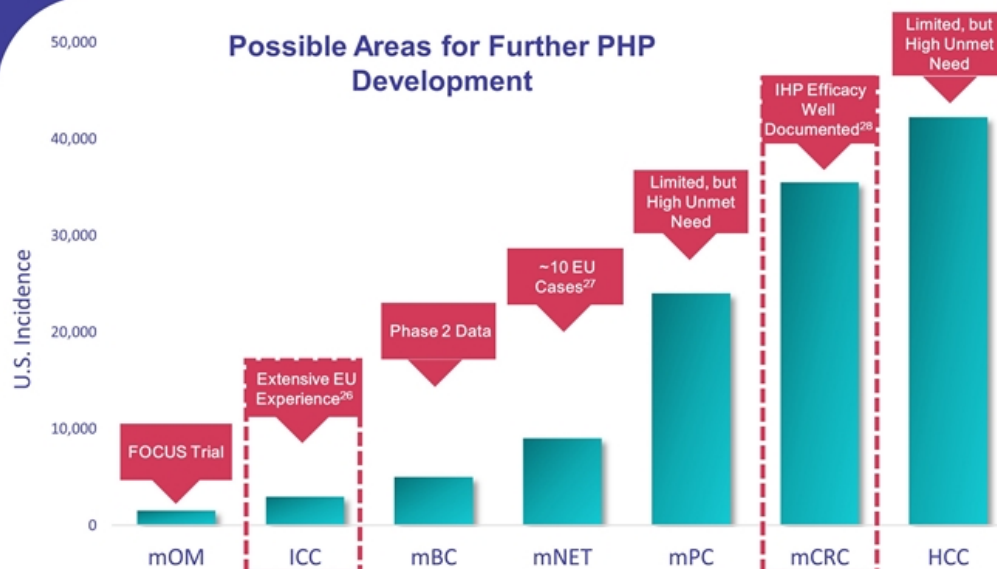
- Intrahepatic Cholangiocarcinoma (ICC)
- Hepatocellular Carcinoma (HCC)
- Metastatic Colorectal Cancer (mCRC)
- Metastatic Breast (mBreast)
- Pancreatic
- Metastatic Neuroendocrine Tumors (mNET)
- Metastatic Cutaneous Melanoma (mCM)

Delcath

# Market Expansion: Liver Disease

BEACHHEAD MARKET | mOM

LIVER DISEASE



US TAM  
**>\$1B**  
Per Year

Delcath

Metastatic Ocular Melanoma (mOM)<sup>2,3</sup>, Cholangiocarcinoma (ICC)<sup>4,5</sup>, Liver-dominant Breast Cancer (mBC)<sup>8-11</sup>, Metastatic Neuroendocrine Tumors (mNET)<sup>6,7</sup>, Metastatic Pancreatic Cancer (mPC)<sup>8,16</sup>, Metastatic Colorectal Cancer (mCRC)<sup>12,13</sup>, Hepatocellular carcinoma (HCC)<sup>29</sup>

## Clinical Rationale for Broad Development Effort

“Broad-spectrum” alkylating agent given at 12X normal systemic doses



- Promising ORR and DCR signals seen across multiple tumor types in Europe and in earlier studies with IHP

Liver mets are often life limiting and reduce I/O efficacy



- When the liver is the life limiting organ, systemic chemotherapy can be paused and HEPZATO added to prolong survival
- Early data supports that combination with I/O agents is safe

PHP treats the entire liver and is not dependent on tumor location



- For patients at high risk of liver mets based on tumor characteristics or ctDNA, adjuvant therapy is logical

Delcath



## Near Term HEPZATO Development Plan

|           | Ongoing                  | 2022 Trial Starts  | 2023+  |
|-----------|--------------------------|--|--|
| Pan Tumor |                          | I/O Combination  |  |
| Other     |                          | LD Pancreatic NET  | LD Breast and/or Pancreatic                                      |
| ICC       |                          | 2 <sup>nd</sup> Line   |  |
| CRC       |                          | LD 2 <sup>nd</sup> Line Additive<br>Stage IV Post Resection ctDNA+ | Hepatic ctDNA Liver MRD Dx<br>Stage II/III Post Resection LD MRD |
| OM        | I/O Combination<br>FOCUS | No Radiological Disease - ctDNA+                                   |  |



Ongoing



Investigator  
Interest  
Confirmed



Pending



Multi-Center  
Sponsored

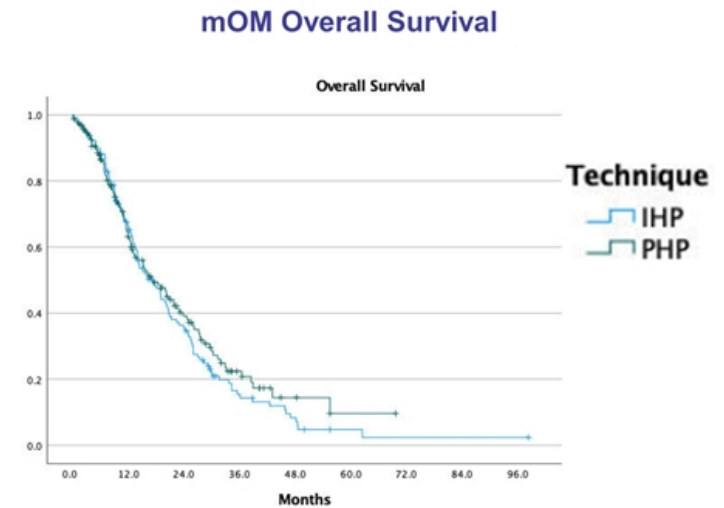
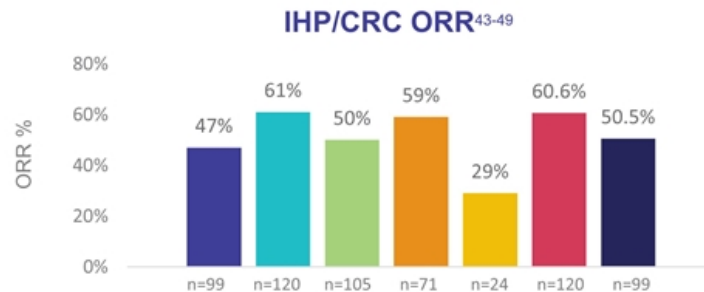
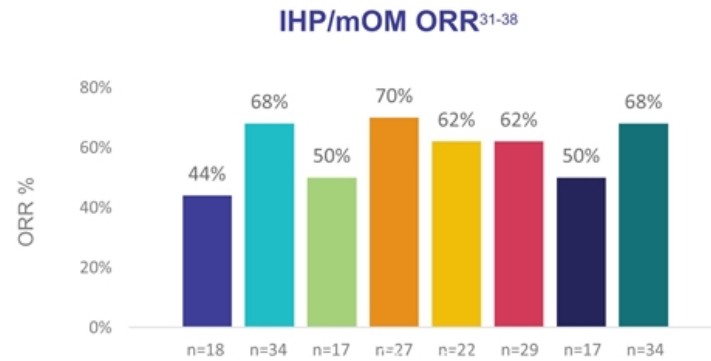


Investigator  
Initiated

Delcath

# Liver Dominant CRC IHP Results Provide Strong Rationale for CRC PHP Trials

mOM Results Similar Between IHP and PHP



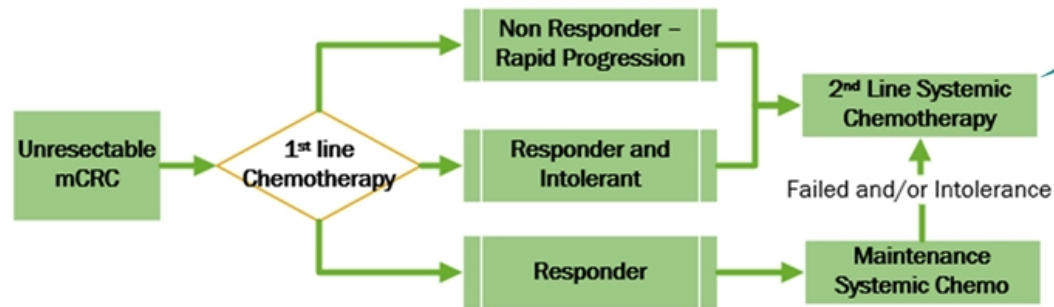
Pending Future Investigation

Delcath

31. Vogl et al (2017). 32. Olofsson et al (2014). 33. Varghese et al (2010). 34. Rizell et al (2008). 35. Alexander et al (2000). 36. Alexander et al (2003). 37. Varghese et al (2010). 38. Olofsson et al (2014). 39. Artzner et al. Cancer Imaging 2019. 40. Bruning et al. Radiology Research and Practice 2020. 41. Karydis et al. Journal of Surgical Oncology 2017. 42. Meijer Annals of Surgical Oncology 2020. 43. van Iersel et al (2010). 44. Alexander et al (2009). 45. van Iersel et al (2008). 46. Rothbarth et al (2003). 47. Vahrmeijer et al (2000). 48. Alexander et al (2009). 49. Van Iersel et al (2010).

## 2<sup>nd</sup> Line Therapy Liver Metastatic CRC

### NCCN Standard of Care



Adding PHP to  
SOC – Not  
Replacing

### Population Base

| US Incidence = 160K new CRC Cases                     | TAM           |
|---|---------------|
| 50% diagnosed metastatic                              | 80K           |
| 50% Liver only metastases                             | 40K           |
| 65-75% are unresectable                               | 26-30K        |
| 85% fail 1 <sup>st</sup> line therapy by 24-36 months | <b>22-25K</b> |

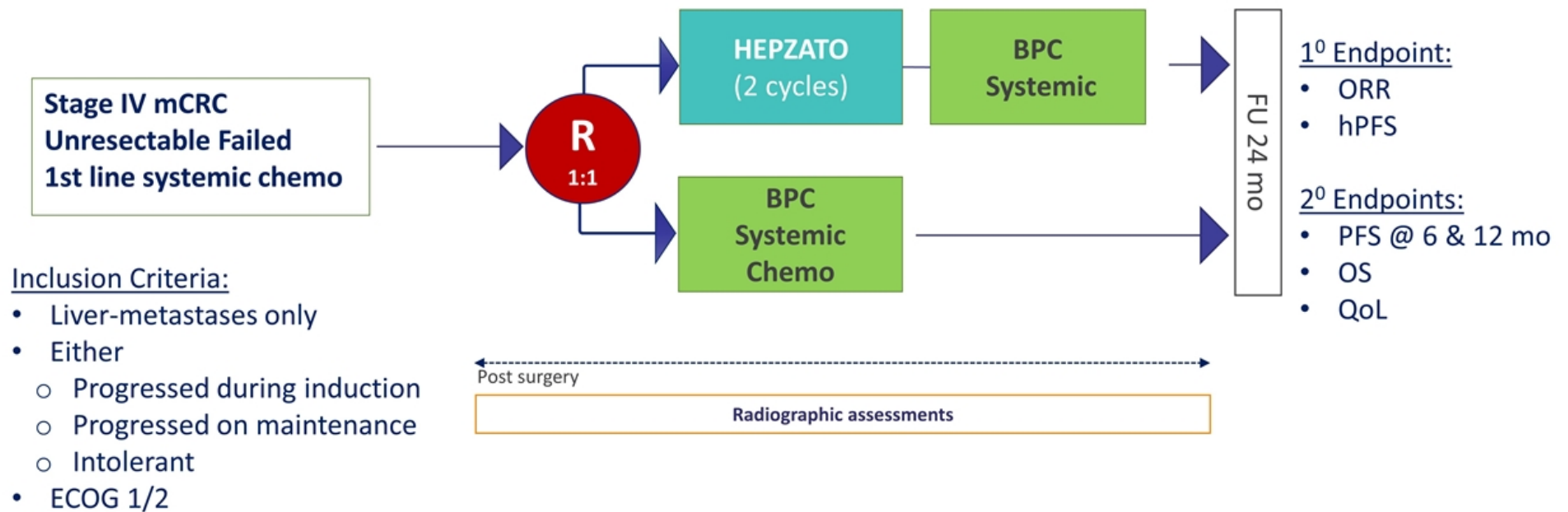
### Current Treatment Options

- Therapy Goal = Disease control
- 1<sup>st</sup> line systemic chemotherapy - 85-90% will have disease progression within 3 yrs

National Cancer Institute. Cancer Stat Facts: Colon and Rectum Cancer. <https://seer.cancer.gov/statfacts/html/colorect.html>  
 Bulut G et al PLoS ONE 16(11): e0259622. <https://doi.org/10.1371/journal.pone.0259622>.

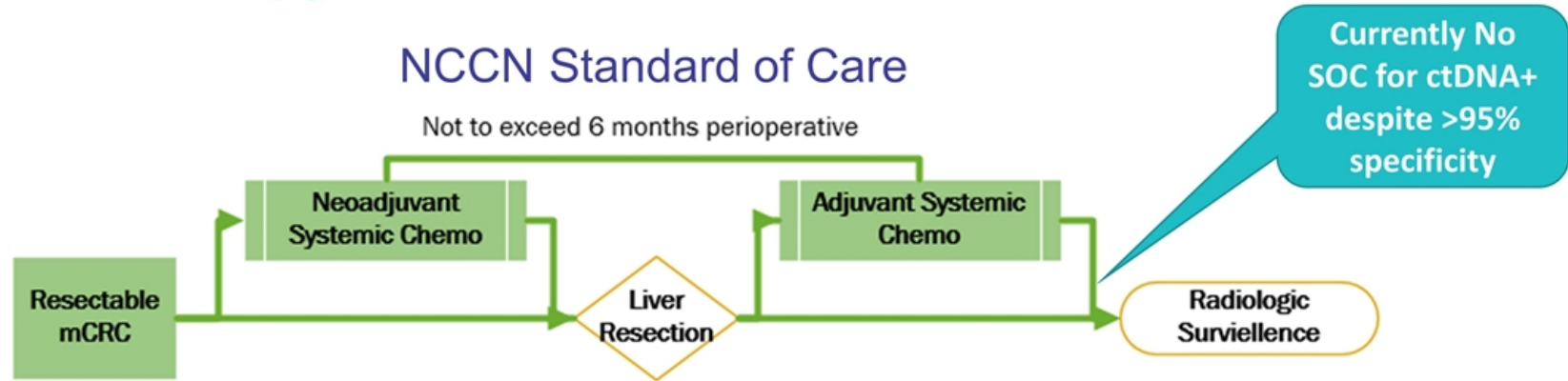
# Hepzato in Stage IV Unresectable mCRC

## Hepzato + Best Physician Choice vs Best Physician Choice



# Adjuvant Therapy : CRC Post Liver Resection

## NCCN Standard of Care



### Population Base

| US Incidence = 160k new CRC Cases                         | TAM          |
|---|--------------|
| 50% diagnosed metastatic                                  | 80K          |
| 50% Liver only metastases                                 | 40K          |
| 25-35% are resectable; initial or converted to resectable | 10-14K       |
| 70% ctDNA positive (based on recurrence)                  | <b>7-10K</b> |

### Current Treatment Options

- Therapy Goal = Prevent recurrence
  - 50-70% recurrence rate within 24 months
  - 70% recur in the liver
- Current adjuvant treatment is +/- chemo up to 6 months perioperative treatment duration

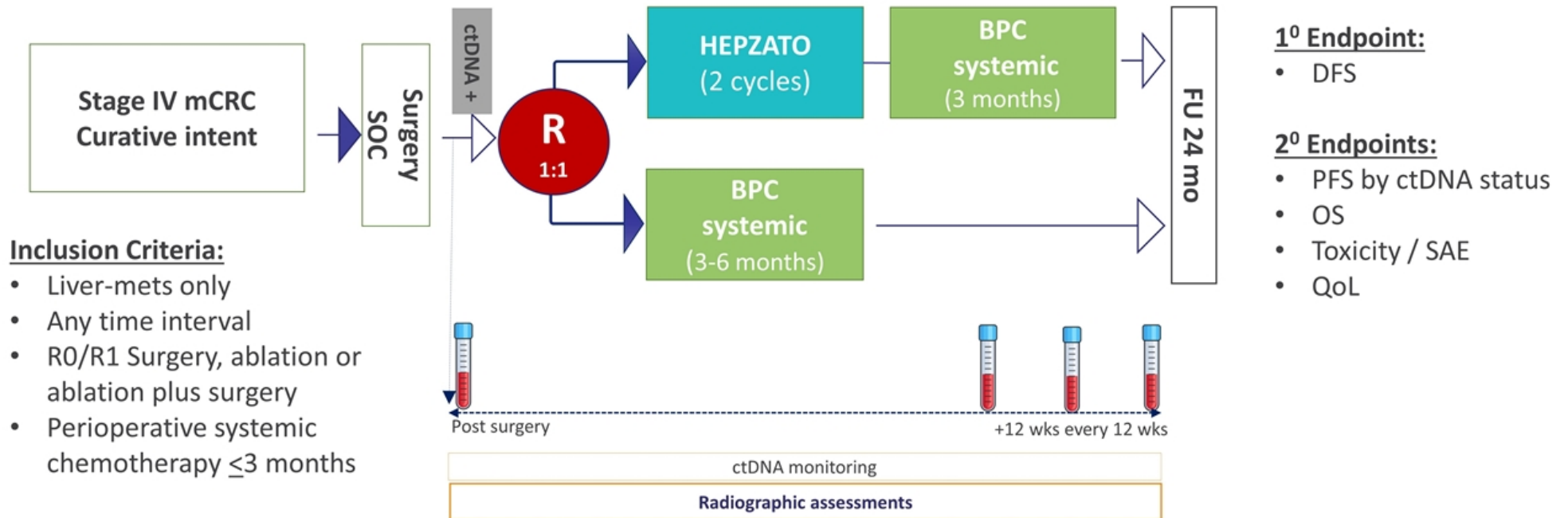
Siegel et al CA CANCER J CLIN 2020;70:145–164

National Cancer Institute. Cancer Stat Facts: Colon and Rectum Cancer. <https://seer.cancer.gov/statfacts/html/colorect.html>.

Holch et al Visc Med 2017;33:70–75 DOI: 10.1159/000454687

# HEPZATO in Post-resection Stage IV ctDNA Positive Patients

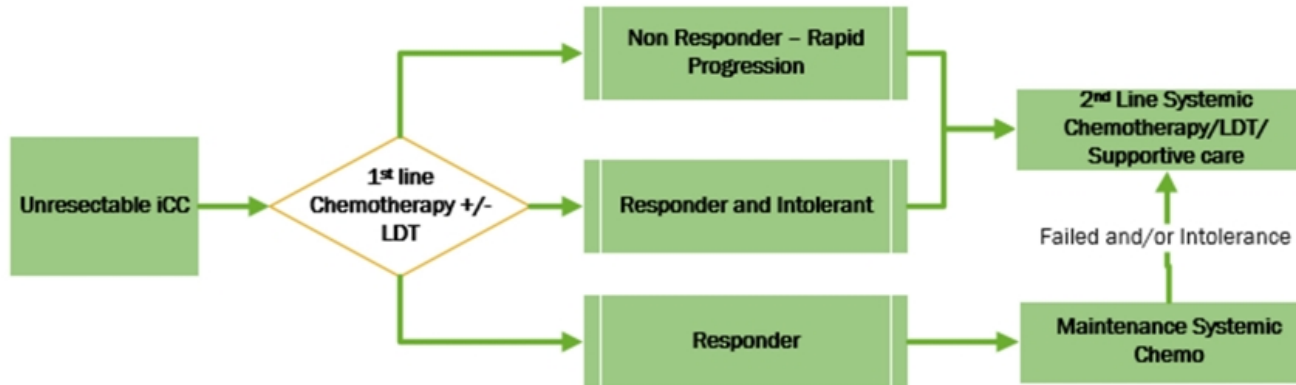
## Hepzato + Best Physician Choice vs Best Physician Choice





# Standard of Care & Epidemiology for iCC

## NCCN Standard of Care



### Population Base

| US Incidence = 3.5k new iCC Cases                | TAM      |
|--|----------|
| 90-95% Unresectable or resection with recurrence | 3.2-3.3K |

### Current Treatment Options

- Therapy Goal = Disease control
- 80% respond to 1st line therapy
- 75% will have disease progression by 1 year

Gupta et al HepatoBiliary Surg Nutr 2017;6(2):101-104

Delcath

## Advanced ICC – 2<sup>nd</sup> Line

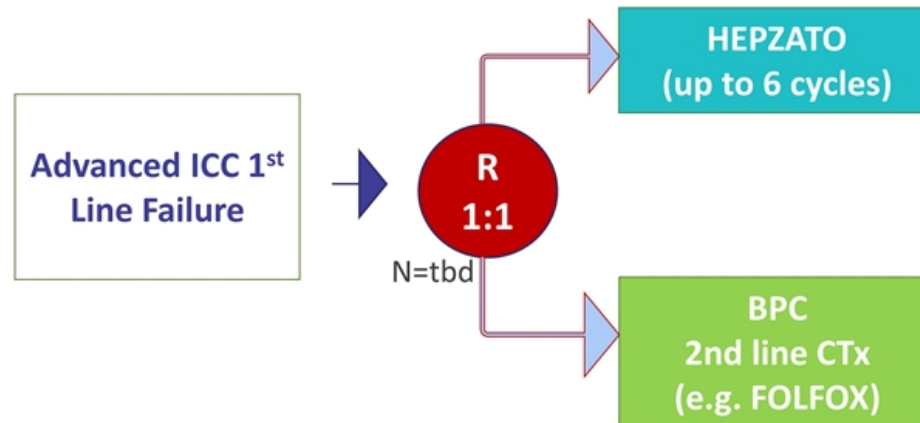
### CHEMOSAT in ICC - European Experience<sup>31,32,32</sup>

| N  | ORR | DCR | CR |
|----|-----|-----|----|
| 20 | 30% | 75% | 3  |

### Hepzato 2<sup>nd</sup> Line vs Best Physician's Choice

#### Inclusion criteria

- Liver dominant disease
- 1 prior line of CTx (e.g. gemcitabine or 5FU-containing based regimen)
- Adequate liver function
- ECOG 0-1



#### 1<sup>o</sup> Endpoint:

PFS@ 6 mo and 12 mo

#### 2<sup>o</sup> Endpoints:

- OS
- ORR
- QoL
- Safety

## Critical IITs

### Hepatic ctDNA Validation

- The liver clears 70% - 80% of ctDNA
- Systemic ctDNA levels should be higher than hepatic vein levels unless there is residual disease in the liver
- The study will collect samples from CRC patients with confirmed liver and non liver mets
- Validation will enable a study targeting stage II/III CRC patients with hepatic MRD - metachronous liver metastases occur 50% in patients post primary resection<sup>34</sup>,
- Hepatic MRD in CRC up to 40K patient TAM

### Treating ctDNA+ OM

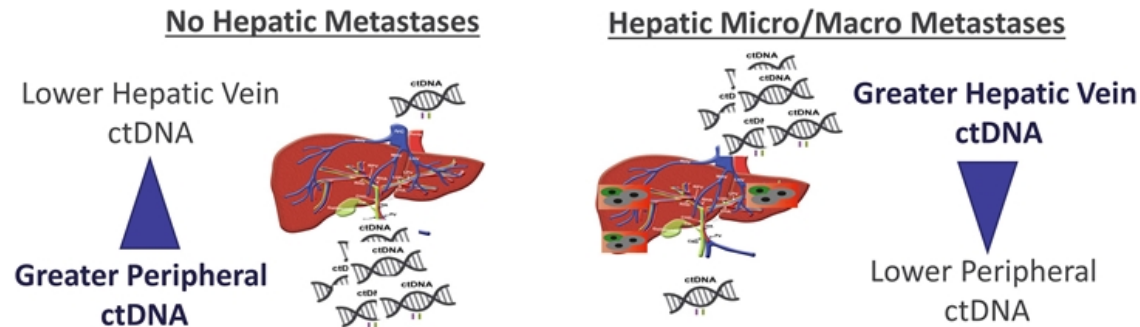
- ~90% of mOM patients present with liver mets
- ctDNA has high specificity for disease recurrence
- ctDNA is likely detectable well prior to radiological evidence enabling earlier treatment

### I/O Combination

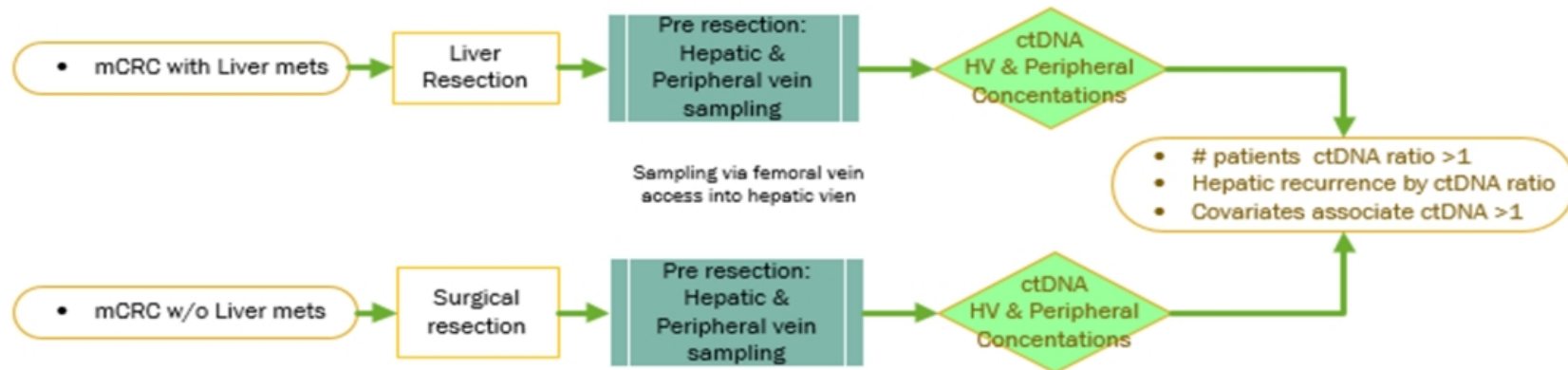
- I/O agents lose efficacy when liver mets are present due to the immunomodulating role of the liver
- The study will be a basket trial for any patients on I/O therapy with liver mets
- Goal will be to make HEPZATO SOC for any patient with liver mets on I/O therapy

# Detecting Liver Minimal Residual Disease

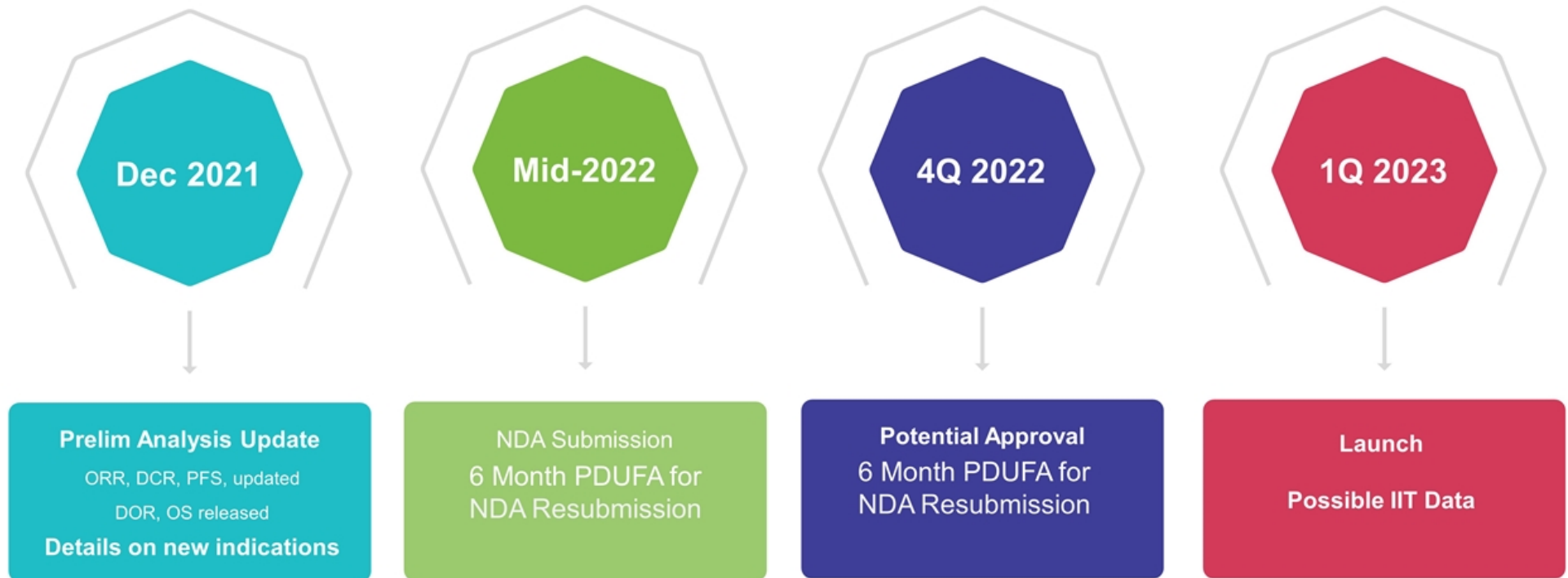
Enabling Technology = ctDNA



~20-30K/year Stage II & III patients recur with liver metastases



## FOCUS Study – Upcoming News Flow



Delcath

## Capital Structure and Share Information - September 30, 2021

| Share Listing - Current                | DCTH (NASDAQ)    |
|--|------------------|
| Shares Outstanding <sup>1</sup>        | 8.81M            |
| Cash and Cash Equivalents <sup>2</sup> | \$29.0M          |
| Warrants Outstanding <sup>3</sup>      | 3.61M            |
| Stock Options Granted                  | 1.70M            |
| 2020 Cash Burn (YTD) <sup>4</sup>      | \$16.2M          |
| Debt <sup>5</sup>                      | \$17.0M          |
| 52 week Low – High <sup>6</sup>        | \$8.28 - \$25.18 |
| 30d Average Daily Volume <sup>7</sup>  | 27,533           |

<sup>1</sup> As of September 30, 2021; includes 7.3M of Common plus 1.2M, Preferred E & E-1 & 0.3M Pre-funded Warrants as converted

<sup>2</sup> As of September 30, 2021; (10-Q filing on November 9, 2021) Includes \$4.2M of restricted cash

<sup>3</sup> As of September 30, 2021; Warrants at a \$10 exercise price

<sup>4</sup> YTD Net cash used in operating activities through Q3, 2021

<sup>5</sup> Includes \$5.0M of notes convertible at \$11.98 per common share equivalent

<sup>6</sup> Used NASDAQ price information starting on September 30, 2020 - September 30, 2021

<sup>7</sup> 30-day average calculated between August 19, 2021 - September 30, 2021

Delcath



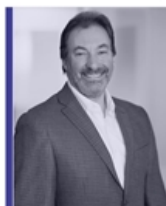
# Multi-Disciplinary, Experienced Leadership Team

**GERARD MICHEL**  
Chief Executive Officer



- » 30+ yrs. pharma/medtech experience
- » C-suite roles at Vericel Corp, Biodel, & NPS
- » M.S. Microbiology, B.S. Biology & Geology from the Univ. of Rochester School of Medicine
- » M.B.A. Simon School of Business & Leadership

**JOHN PURPURA**  
Chief Operating Officer



- » Past VP and Exec Director roles of Reg. Affairs for Bracco Diagnostics
- » Held senior roles Sanofi-Aventis, Bolar Pharma, Luitpold Pharma & Eon Labs
- » M.S. Mgmt. & Policy and B.S. Chemistry and Biology at the State University of NY at Stony Brook

**JOHNNY JOHN, MD**  
SVP Clinical Development & Medical Affairs



- » 15+ yrs. experience in oncology drug development and clinical trials
- » 11 years of personal clinical practice
- » Received M.D. from Mangalore University, India; post-grad training at the University of IL

**KEVIN MUIR**  
VP, Commercial Operations



- » 20+ yrs. of medtech/bioTx sales & marketing experience.
- » Held senior leadership roles at BTG, ClearFlow, Aragon Surgical, Kensey Nash Corporation, and Kyphon.
- » Field Artillery officer in the U.S. Army
- » B.S. in Management Systems Engineering at the U.S. Military Academy at West Point

## BOARD OF DIRECTORS

|                           |          |
|---------------------------|----------|
| Dr. Roger G. Stoll, Ph.D. | Chairman |
| John R. Sylvester         | Director |
| Elizabeth Czerepak        | Director |
| Steven Salamon            | Director |
| Dr. Gil Aharon, Ph.D.     | Director |
| Gerard Michel             | CEO      |

Delcath

# Delcath: A Unique Opportunity



Novel platform in interventional oncology



Multiple near-term catalysts (Final data and NDA filing, new indications)



Safety and efficacy supported by multiple trials and commercial usage



Initial orphan indication allows for targeted marketing effort and rapid uptake



Platform has potential utility in multiple indications

Delcath

THANK YOU

Delcath

# References

1. <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4107710/>
2. Cancer.net Editorial Board (2020) Eye Cancer - Statistics. In: Cancer.Net. <https://www.cancer.net/cancer-types/eye-cancer/statistics>. Accessed 22 Jun 2020
3. Ocular Melanoma Foundation. Treatment of Metastatic Disease. In: OMF – Metastatic Treatment. <http://www.ocularmelanoma.org/metstreatment.htm>. Accessed 22 Jun 2020
4. Patel N, Benipal B. Incidence of Cholangiocarcinoma in the USA from 2001 to 2015: A US Cancer Statistics Analysis of 50 States. *Cureus*. 2019;11(1):e3962. Published 2019 Jan 25.
5. United States Census Bureau. (2019) Monthly Population Estimates for the United States: April 1, 2010 to December 1, 2020 (NA-EST2019-01).
6. Cancer.net Editorial Board. (2020) Neuroendocrine Tumors - Statistics. In: Cancer.Net. <https://www.cancer.net/cancer-types/neuroendocrine-tumors/statistics>. Accessed 22 Jun 2020
7. Saeed A, Buell JF, Kandil E. Surgical treatment of liver metastases in patients with neuroendocrine tumors. *Ann Transl Med*. 2013;1(1):6. doi:10.3978/j.issn.2305-5839.2013.01.08
8. Surveillance, Epidemiology, and End Results (SEER) Program Populations (1969-2018) ([www.seer.cancer.gov/popdata](http://www.seer.cancer.gov/popdata)), National Cancer Institute, DCCPS, Surveillance Research Program, released December 2019.
9. Adam R, Aloia T, Krissat J, Bralet MP, Paule B, Giacchetti S, Delvart V, Azoulay D, Bismuth H, Castaing D. Is liver resection justified for patients with hepatic metastases from breast cancer? *Ann Surg*. 2006 Dec;244(6):897-907; discussion 907-8. doi: 10.1097/01.sla.0000246847.02058.1b. PMID: 17122615; PMCID: PMC1856635.
10. Insa A, Lluch A, Prosper F, Marugan I, Martinez-Agullo A, Garcia-Conde J. Prognostic factors predicting survival from first recurrence in patients with metastatic breast cancer: analysis of 439 patients. *Breast Cancer Res Treat*. 1999 Jul;56(1):67-78. doi: 10.1023/a:1006285726561. PMID: 10517344.
11. Clark GM, Sledge GW Jr, Osborne CK, McGuire WL. Survival from first recurrence: relative importance of prognostic factors in 1,015 breast cancer patients. *J Clin Oncol*. 1987 Jan;5(1):55-61. doi: 10.1200/JCO.1987.5.1.55. PMID: 3806159.
12. Cancer.net Editorial Board. (2020) Colorectal Cancer - Statistics. In: Cancer.Net. <https://www.cancer.net/cancer-types/colorectal-cancer/statistics>. Accessed 22 Jun 2020
13. Ismaili N. Treatment of colorectal liver metastases. *World J Surg Oncol*. 2011;9:154. Published 2011 Nov 24. doi:10.1186/1477-7819-9-154
14. Lane AM, Kim IK, Gragoudas ES. Survival Rates in Patients After Treatment for Metastasis From Uveal Melanoma. *JAMA Ophthalmol*. 2018 Sep 1;136(9):981- 986
15. Karydis I, Gangi A, Wheeler MJ, et al. Percutaneous hepatic perfusion with Melphalan in uveal melanoma: A safe and effective treatment modality in an orphan disease. *J Surg Oncol*. 2018;117(6):1170-1178. doi:10.1002/jso.24956



# References

16. Oweira H, Petrausch U, Helbling D, Schmidt J, Mannhart M, Mehrabi A, Schöb O, Giryas A, Decker M, Abdel-Rahman O. Prognostic value of site-specific metastases in pancreatic adenocarcinoma: A Surveillance Epidemiology and End Results database analysis. *World J Gastroenterol*. 2017 Mar 14;23(10):1872-1880. doi: 10.3748/wjg.v23.i10.1872. PMID: 28348494; PMCID: PMC5352929.
17. Xu L, T, Funchain P, F, Bena J, F, Li M, Tarhini A, Berber E, Singh A, D: Uveal Melanoma Metastatic to the Liver: Treatment Trends and Outcomes. *Ocul Oncol Pathol* 2019;5:323-332. doi: 10.1159/000495113
18. Data on file: presented at American Society of Clinical Oncology – June 4, 2021.
19. Preliminary analysis of FOCUS trial released 3/31/21
20. Hughes S., et al. *Ann Surg Oncol*. 2016 Apr;23(4):1309-19. doi: 10.1245/s10434-015-4968-3.8-3
21. Tulokas S, Mäenpää H, et al. Selective internal radiation therapy (SIRT) as treatment for hepatic metastases of uveal melanoma: a Finnish nation-wide retrospective
22. Sacco J, et al. *Annals of Oncology* (Dec 2020) 31 (suppl\_7): S1441-S1451. 10.1016/annonc/annonc392
23. Shibayama Y, Namikawa K, Sone M, et al. Efficacy and toxicity of transarterial chemoembolization therapy using cisplatin and gelatin sponge in patients with liver metastases from uveal melanoma in an Asian population. *Int J Clin Oncol*. 2017 Jun;22(3):577-584. doi: 10.1007/s10147-017-1095-0. Epub 2017 Jan 31. PMID: 28144882.
24. Meta-analysis: Data on file
25. Piperno - Neumann, et. al. AACR Annual Meeting 2021
26. Ferrucci, P. et al. A New Option for the Treatment of Intrahepatic Cholangiocarcinoma: Percutaneous Hepatic Perfusion with CHEMOSAT Delivery System. *Cells* 2021, 10, 70. <https://doi.org/10.3390/cells10010070>
27. Vogel, A., Gupta, S., Zeile, M. et al. Chemosaturation Percutaneous Hepatic Perfusion: A Systematic Review. *Adv Ther* 33, 2122–2138 (2016). <https://doi.org/10.1007/s12325-016-0424-4>
28. Reddy SK, Kesmodel SB, Alexander HR. Isolated hepatic perfusion for patients with liver metastases. *Therapeutic Advances in Medical Oncology*. July 2014:180-194. doi:10.1177/1758834014529175
29. <https://www.cancer.org/cancer/liver-cancer/about/what-is-key-statistics.html>
30. Bethlehem M., et al. *Cancers* 2021, 13(18), 4726; <https://doi.org/10.3390/cancers13184726>
31. Marquardt, S., Kirstein, M.M., Brüning, R. et al. Percutaneous hepatic perfusion (chemosaturation) with melphalan in patients with intrahepatic cholangiocarcinoma: European multicentre study on safety, short-term effects and survival. *Eur Radiol* 29, 1882–1892 (2019). <https://doi.org/10.1007/s00330-018-5729->
32. Vogl TJ, Zangos S, Scholtz JE et al. Chemosaturation with Percutaneous Hepatic Perfusion of Melphalan for Hepatic Metastases: Experience from Two European Centers. *Fortschr Röntgenstr* 2014; 186: 937–944
33. Personal communication, data on file, [Pier Francesco Ferrucci](#), Tumor Biotherapy Unit, Department of Experimental Oncology, European Institute of Oncology, IRCCS, 20141 Milan, Italy
34. Zeng, X.; Ward, S.E.; Zhou, J.; Cheng, A.S.L. Liver Immune Microenvironment and Metastasis from Colorectal Cancer-Pathogenesis and Therapeutic Perspectives. *Cancers* 2021, 13, 2418. <https://doi.org/10.3390/cancers13102418>