



# ASCELIA PHARMA

Share ticker: ACE  
Nasdaq Stockholm (small cap)

## ASCELIA PHARMA

COMPANY PRESENTATION

### ADVANCING ORPHAN ONCOLOGY

MAY 2020



# FORWARD LOOKING STATEMENTS

This presentation, which includes all information and data on the following slides, any oral statements made when presenting these slides, and any other material distributed or statements made at, or in connection with, such presentation (the “Presentation”), relates to Ascelia Pharma AB (publ) (hereinafter, together with its subsidiaries, the “Company”) is furnished to you solely for your information and may not be reproduced or redistributed, in whole or in part, to any other person without the prior written consent of the Company. You should not rely upon it or use it to form the definitive basis for any decision, contract, commitment or action whatsoever, with respect to any transaction or otherwise.

The information included in this Presentation may contain certain forward-looking statements relating to the business, financial performance and results of the Company and/or the industry in which it operates. Forward-looking statements concern future circumstances and results and other statements that are not historical facts, sometimes identified by the words “believes”, “expects”, “predicts”, “intends”, “projects”, “plans”, “estimates”, “aims”, “foresees”, “anticipates”, “targets”, and similar expressions. The forward-looking statements contained in this Presentation, including assumptions, opinions and views of the Company or cited from third party sources are solely opinions and forecasts which are subject to risks, uncertainties and other factors that may cause actual events to differ materially from any anticipated development. Neither the Company nor any of its affiliates, directors, employees or advisors provides any assurance that the assumptions underlying such forward-looking statements are free from errors nor do any of them accept any responsibility for the future accuracy of the opinions expressed in this Presentation or the actual occurrence of the forecasted developments. This Presentation speaks as of the applicable reporting date, and there may have been changes in matters which affect the Company subsequent to the date of this Presentation. Neither the issue nor delivery of this Presentation shall under any circumstance create any implication that the information contained herein is correct as of any time subsequent to the date hereof or that the affairs of the Company have not since changed, and the Company does not intend, and does not assume any obligation, to update or correct any information included in this Presentation.

Each person should make their own independent assessment of the merits of the Company and should consult their own professional advisors. By receiving this Presentation, you acknowledge that you will be solely responsible for your own assessment of the market and the market position of the Company and that you will conduct your own analysis and be solely responsible for forming your own opinion of the potential future performance of the Company’s business.

# ASCELIA PHARMA: ADVANCING ORPHAN ONCOLOGY

## A global health burden

---

44 million people live with cancer; 18 million are diagnosed each year<sup>1</sup>

USD 150 bn spent yearly on cancer therapies alone<sup>2</sup>

**Orphan drugs represent 12 of 15 new active substances in oncology launched in the US in 2018<sup>2</sup>**

## Dedicated to unmet needs in orphan oncology

---

### Drugs with a clear development and market pathway

- Advancing liver imaging with orphan MRI contrast agent with no competition (in ongoing Phase 3)
- Advancing chemotherapy with novel tablet for gastric cancer (Phase 2 ready)

### Capabilities to bring new compounds to market

- World class cross-functional team
- Headquartered in Malmö, Sweden
- Listed on NASDAQ STOCKHOLM in 2019 (ticker: ACE)
- Solid financial position with SEK 169 million in liquid assets

#### Sources

- 1) <https://canceratlas.cancer.org/the-burden/the-burden-of-cancer/> (2018 figures)
- 2) Global Oncology Trends 2019, IQVIA (2018 figures)

# CLINICAL STAGE PORTFOLIO ADDRESSING CLEAR UNMET NEEDS

Drug candidate	Indication	Phase 1	Phase 2	Phase 3	Filing	Launch
<b>Mangoral</b> <ul style="list-style-type: none"> <li>• Only <u>non</u>-gadolinium imaging drug</li> <li>• No competing products</li> <li>• \$350-500M market with upside potential</li> <li>• De-risked Phase 3 clinical program</li> <li>• Orphan Drug Designation</li> </ul>	<b>Visualisation of focal liver lesions</b> <ul style="list-style-type: none"> <li>• Liver metastases</li> <li>• Primary liver cancer</li> <li>• Benign lesions</li> </ul>	✓	✓	2020 - H2 2021	H1 2022	Q4 2022 - H1 2023
<b>Oncoral</b> <ul style="list-style-type: none"> <li>• Novel tablet chemotherapy formulation</li> <li>• Phase 1 completed with promising results</li> <li>• Gastric cancer is an Orphan indication</li> </ul>	<b>Treatment of gastric cancer</b>	✓	2021 - 2023	Strong case for development and commercialisation partnering after phase 2		

# STRONG AND EXPERIENCED MANAGEMENT AND BOARD

## EXECUTIVE MANAGEMENT



**Magnus Corfitzen**  
Chief Executive Officer



**Kristian Borbos**  
Chief Financial Officer



**Carl Bjartmar, MD, Ph.D**  
Chief Medical Officer



**Julie Waras Brogren**  
Chief Commercial Officer



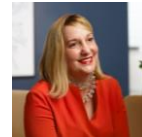
**Mikael Widell**  
Head of IR and Communications



## BOARD OF DIRECTORS



**Peter Benson**  
Chairman



**Lauren Barnes**  
Director



**Bo Jesper Hansen, MD, Ph.D**  
Director



**Hans Maier, Ph.D, Prof.**  
Director



**Helena Wennerström**  
Director



**Niels Mengel**  
Director



**René Spogård**  
Director





An elderly couple is walking through a lush, green park. The woman, on the left, is wearing a red top and light-colored pants, and is smiling. The man, on the right, is wearing a blue and white checkered shirt and blue jeans, and is also smiling. They are holding hands. In the foreground, there is a picnic basket filled with food and drinks. The background is filled with large, leafy trees under a bright sky.

# MANGORAL

LIVER MRI CONTRAST AGENT IN  
PHASE 3 CLINICAL STUDIES



# LIVER METASTASES – A MAJOR CHALLENGE IN ONCOLOGY

## LIVER METASTASES COMMON IN MANY CANCER TYPES

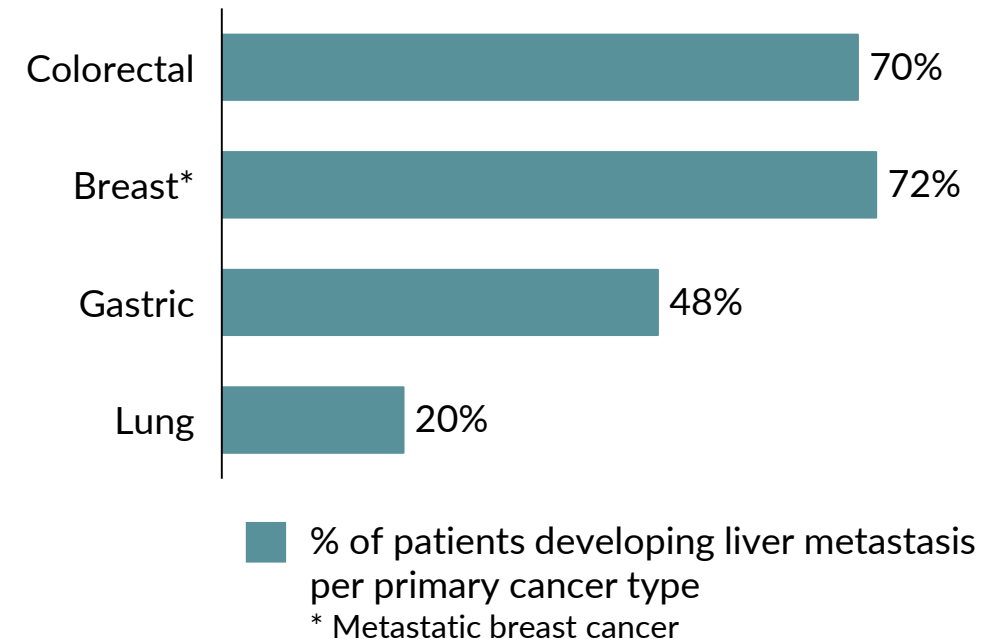
The liver is the **most frequent** organ for metastases after lymph node<sup>3</sup> and often the first site of metastasis

- 70% of patients with colon cancer will develop liver metastases<sup>1</sup>
- Liver metastases are also common in other cancer types such as lung cancer, gastric cancer, metastatic breast cancer<sup>2,3</sup> etc.

Liver metastases often the **cause of mortality** (not primary tumour)<sup>4</sup>



### Incidence of liver metastasis in various primary cancers<sup>1-4</sup>



1) Riihimäki, M. et al. Patterns of metastasis in colon and rectal cancer. *Sci. Rep.* 6, 29765; doi: 10.1038/srep29765 (2016); *Journal of Pathology*, 2014, 232:23-31

2) *Oncotarget*, 2016, 7(32):52307; *Lung Cancer*, 2014, 86:78-84 (6):29765

3) Guy diSibio and Samuel W. French (2008) Metastatic Patterns of Cancers: Results From a Large Autopsy Study. *Archives of Pathology & Laboratory Medicine*: June 2008, Vol. 132, No. 6, pp. 931-939

4) Rahbari et al. Metastatic Spread Emerging From Liver Metastases of Colorectal Cancer: Does the Seed Leave the Soil Again? *Annals of Surgery*: February 2016 - Volume 263 - Issue 2 - p 345-352

# LIVER METASTASES: HOW TO FIND AND WHAT TO DO

## DETECT AND LOCALISE

**Liver MRI** is the **most sensitive** method for detection of liver metastases<sup>2)</sup>

**Gadolinium** based imaging drugs are given to maximise accuracy of liver metastasis detection in MRI



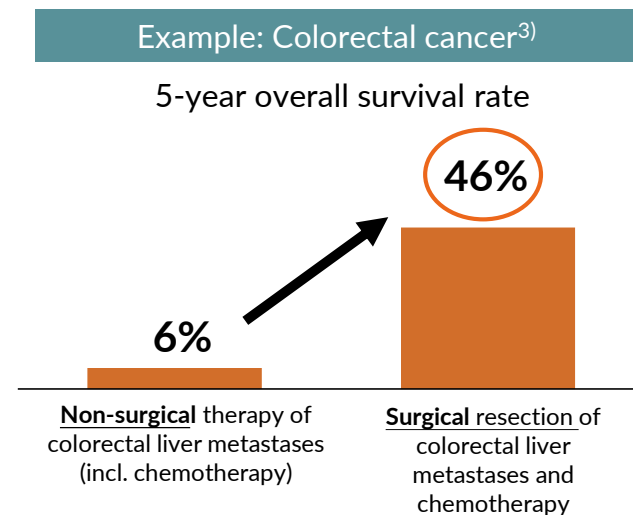
## TREAT

Treatment options for liver metastases are:

- Surgical resection (only if detected early)
- Localised therapies (ablation, embolisation, radiation)
- Drug therapy

## IMPROVE SURVIVAL

**Accurate, early detection** of liver metastases significantly impact treatment decisions and patient survival



1) Guy diSibio and Samuel W. French (2008) Metastatic Patterns of Cancers: Results From a Large Autopsy Study. Archives of Pathology & Laboratory Medicine: June 2008, Vol. 132, No. 6, pp. 931-939

2) Albiin N et al. Manganese chloride tetrahydrate (CMC-001) enhanced liver MRI: evaluation of efficacy and safety in healthy volunteers. MAGMA. 2012 Mar 8

3) Clinical Colorectal Cancer, Vol. 15, No. 4, Dec 2016, e183-192



# GADOLINIUM – THE STANDARD OF CARE CONTRAST AGENT IS NOT SAFE FOR ALL PATIENTS

## SAFETY RISKS ASSOCIATED WITH GADOLINIUM

- Gadolinium (toxic heavy metal) based contrast agents (GBCA) in **renally impaired patients and patients with acute kidney injury** are linked to Nephrogenic Systemic Fibrosis (NSF)
  - An aggressive fibrosing disease in which fibrous connective tissue seeks to replace normal tissue of the skin, muscle and deep inner organs
  - A serious and potentially fatal condition multiorgan disorder with no treatments available
- Since 2006 there have been 3,125 cases of NSF as a result of GBCA use in the US, 745 of which resulted in death
- Gadolinium's link to NSF has been acknowledged by FDA and EMA, with black box warnings and label changes issued



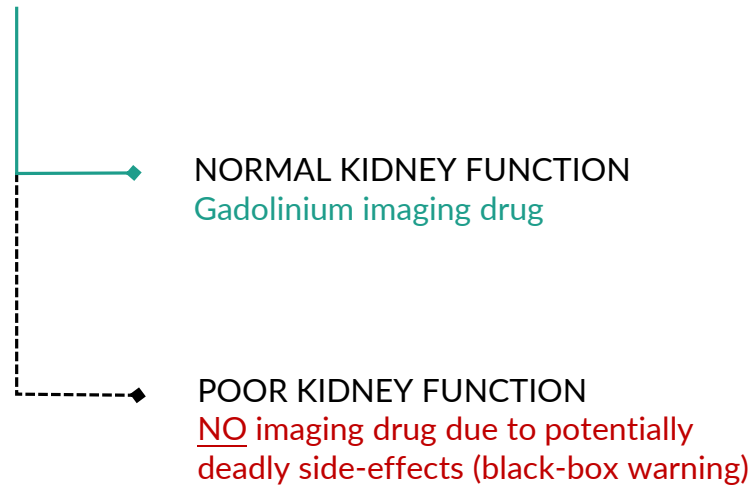
*Clinical presentation of NSF*



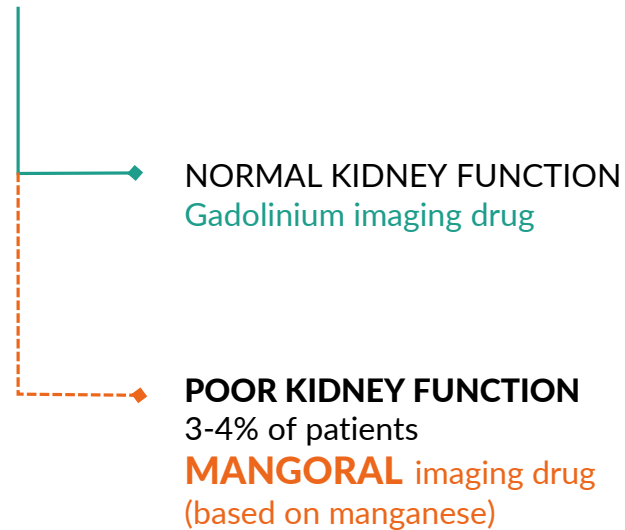
*NSF causes the skin to gradually become fibrotic and adheres to the underlying fascia causing hyperpigmentation, blistering and ulceration*

# MANGORAL – MANGANESE BASED LIVER CONTRAST AGENT

## TODAY



## TOMORROW



**Mangoral** aims to be the only standard of care liver MRI imaging drug for patients with impaired kidney function



280,000

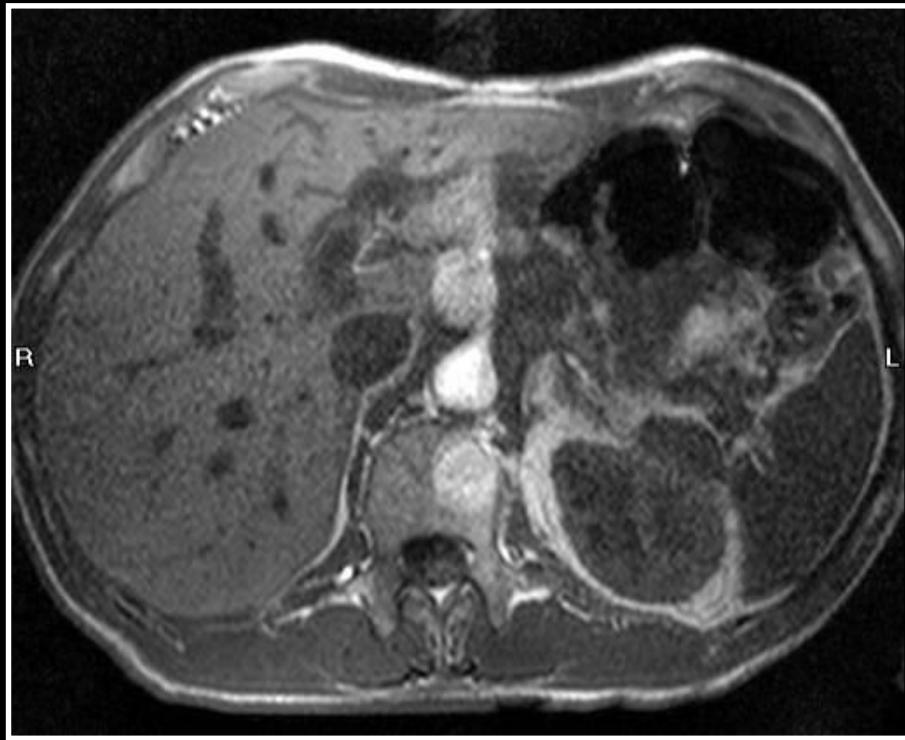
patients with impaired kidney function in major markets

**WARNING: NEPHROGENIC SYSTEMIC FIBROSIS (NSF)**  
*See full prescribing information for complete boxed warning.*  
Gadolinium-based contrast agents (GBCAs) increase the risk for NSF among patients with impaired elimination of the drugs. Avoid use of GBCAs in these patients unless the diagnostic information is essential and not available with non-contrasted MRI or other modalities.

- The risk for NSF appears highest among patients with:
  - Chronic, severe kidney disease (GFR < 30 mL/min/1.73m<sup>2</sup>), or
  - Acute kidney injury.
- Screen patients for acute kidney injury and other conditions that may reduce renal function.
- For patients at risk for chronically reduced renal function (for example, age >60 years, hypertension or diabetes), estimate the glomerular filtration rate (GFR) through laboratory testing (5.1).

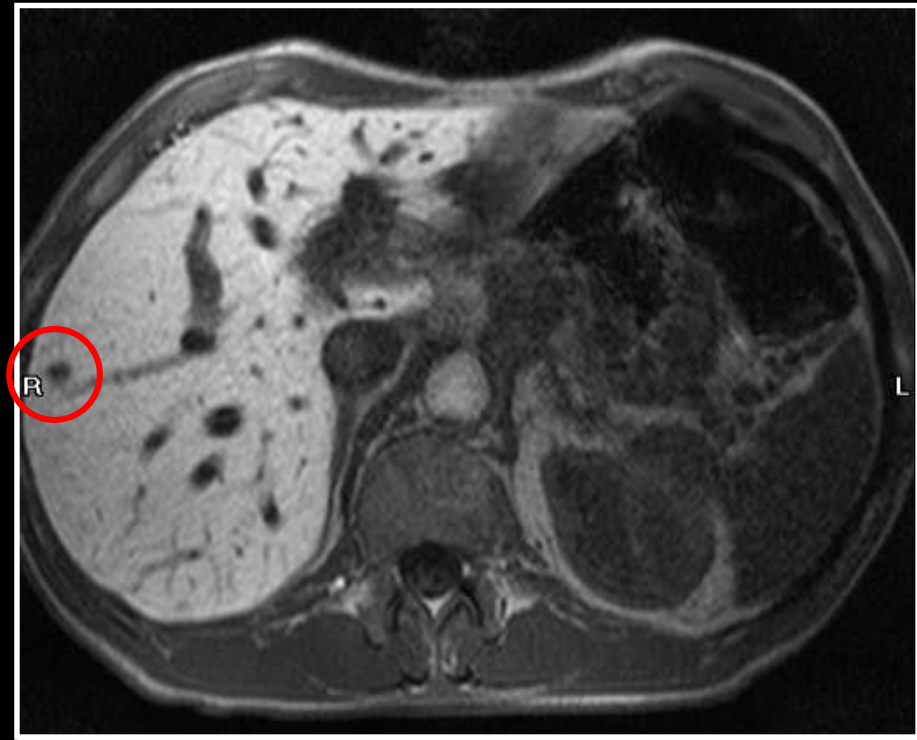
# MANGORAL MAKES A REAL DIFFERENCE

## PATIENT EXAMPLE FROM PHASE II STUDY



**Unenhanced** liver MRI

*(standard of care today in target patient population)*



**Mangoral** enhanced liver MRI

**Liver metastasis appear with Mangoral**



# MANGORAL CLINICAL ACTIVITIES

Study	Objective of the study	Site location and no. of patients	Time schedule
<b>Pivotal Phase 3 study ("SPARKLE")</b>	Assess efficacy and safety of Mangoral in patients with severely reduced kidney function and with known or suspected liver lesions	Global multicentre study in up to 200 patients	<ul style="list-style-type: none"> <li>• Study ongoing</li> <li>• Study results expected in H2-2021</li> </ul>
<b>Hepatic study</b>	Assess the influence of hepatic impairment on the safety, pharmacokinetics and pharmacodynamics of Mangoral	Open-label study on 24 healthy and hepatically impaired participants at the Texas Liver Institute, San Antonio, US	<ul style="list-style-type: none"> <li>• Study ongoing</li> <li>• Study expected to be completed in 2020</li> </ul>
<b>Food effect study</b>	Assess the effect of food intake on Mangoral uptake	Study contract to be awarded	<ul style="list-style-type: none"> <li>• Study preparations ongoing</li> <li>• Short study, expected to be completed in 2020</li> </ul>

These studies, together with the already completed Phase 1 and 2 studies, ensure a comprehensive data package for the regulatory submissions in key markets

# DE-RISKED PHASE 3 STUDY UNDERPINNED BY STRONG DATA FROM COMPLETED STUDIES AND STUDY DESIGN

## Strong data package for Mangoral

Six phase 1 and 2 clinical studies completed

Consistent strong efficacy readout and safety profile

Blind read study of all imaging data presented at major conferences

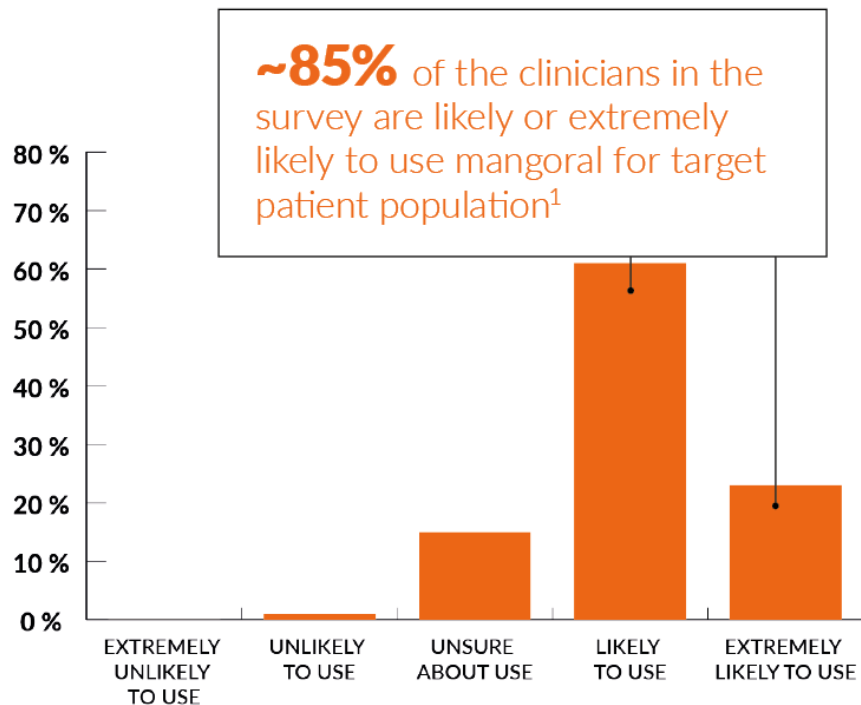
- The study with 178 persons further underlined that Mangoral significantly improves MRI performance
- 33% more lesions were detected after Mangoral enhanced MRI
- **Mangoral significantly improved lesion visualisation**  
Delineation: p-value <0.0001  
Conspicuity: p-value <0.0001

## Phase 3 registration-enabling study (study ongoing)

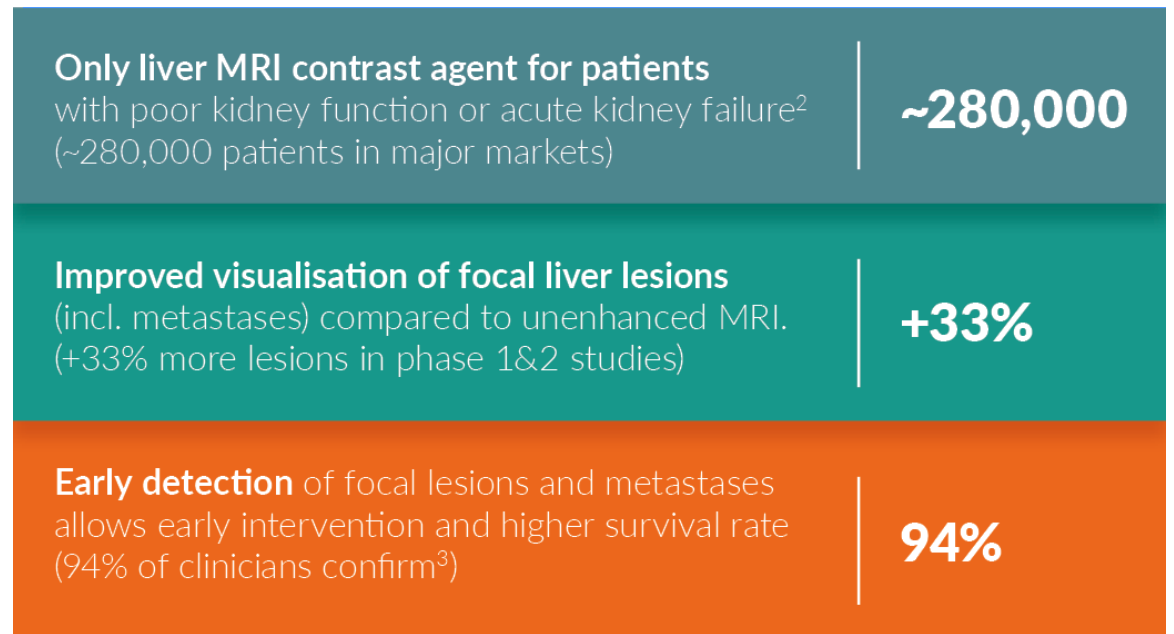
Number of patients	Global study in up to 200 patients
Endpoint	<b>Lesion visualisation</b> <ul style="list-style-type: none"><li>• Lesion border delineation (border sharpness of lesions)</li><li>• Conspicuity (lesion contrast compared to liver background)</li></ul>
Comparator	Unenhanced MRI + Mangoral MRI vs. Unenhanced MRI
Follow-up	72 hours
Randomisation	<b>No</b> - each patient at his/her own control
Validation	Phase 3 program has been discussed with FDA and EMA

# ADDRESSING UNMET NEEDS OF 280,000 PATIENTS IN KEY MARKETS

## Confirmed unmet medical need



## Value to payers, physicians and patients



<sup>1</sup> Market research by Back Bay Life Science Advisors with interview of 84 radiologists across the US regarding clinical practices in liver MRI scanning, the use of gadolinium and mangoral product profile. Notes: 1) Survey answers to question: 'What is your overall opinion of this product for its target population of patients with known or suspected liver metastases and severe renal insufficiency or acute kidney injury?' 2) Based on regulatory drug class warning on use of gadolinium-based contrast agents in patients with renal impairment (an eGFR <30 ml/min/1.73 m<sup>2</sup>) or acute kidney failure. 3) Survey answers to 'Using contrast MRI is important for early intervention, to detect small lesions, which if removed can be curative e.g. colorectal cancer metastases?'



# MANGORAL IS THE ONLY PRODUCT IN A \$350-500M MARKET

## Addressable market

8.5 million people

... in key markets have suspected liver metastases or primary liver cancer (based on relevant primary cancers, colorectal, breast, lung and gastric)

280,000 of these

... equivalent to 3-4% have poor kidney function (stage 5, 4 or 3 worsening or acute kidney injury - with higher prevalence in elderly and cancer populations)

need MRI

... for diagnosis, monitoring and surveillance

at \$1,500 - 3,000 per dose

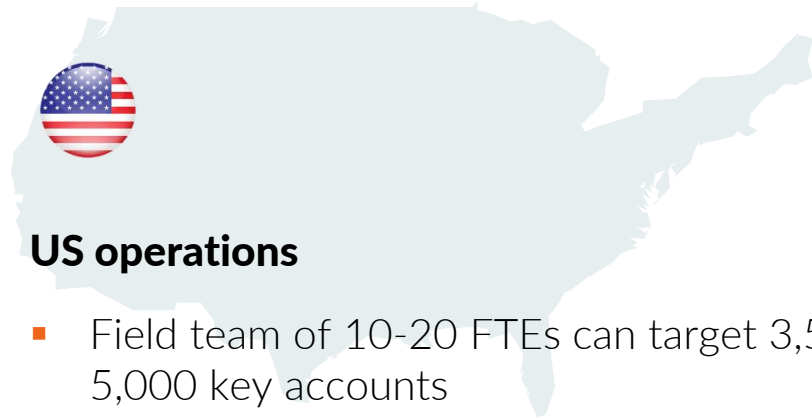
... based on unmet needs in patient population and value of contrast enhanced MRI

**\$350-500 million**

**... addressable market annually with no competing drug**

# OUTLOOK FOR MARKET OPTIMAL LAUNCH STRATEGY

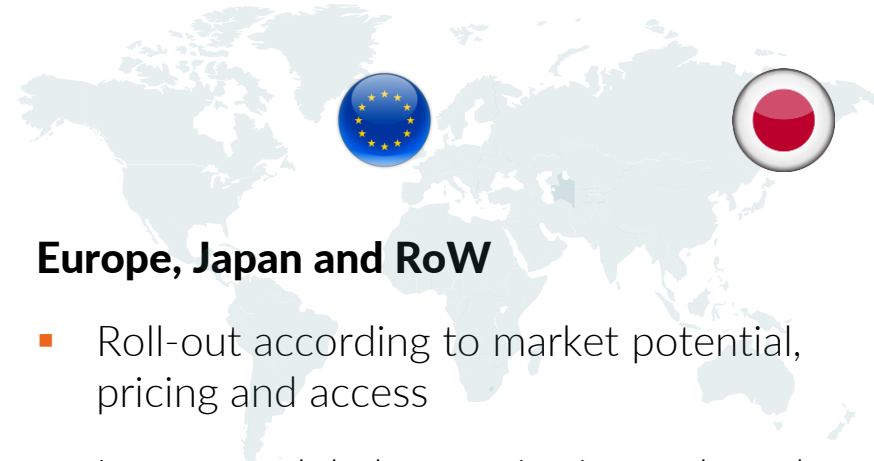
## Strong case for own US commercialisation



### US operations

- Field team of 10-20 FTEs can target 3,500-5,000 key accounts
- Target major hospitals with nephrology departments and independent specialist clinics
- US capability to include commercial and cross-functional support team

## Optimal RoW uptake with partnering



### Europe, Japan and RoW

- Roll-out according to market potential, pricing and access
- Leverage global synergies in pre-launch and launch
- Ascelia vs. partner roles evaluated to maximise value

# PREPARING FOR COMMERCIALISATION

## PREPARE THE PRODUCT

2020

- Phase 3 study SPARKLE (ongoing)
- Detail pricing & access strategy
- Define roll-out priorities & projections
- Develop pre-launch & launch plans
- Build blueprint for US operations and RoW partnering
- Expand key opinion leader network

## PREPARE THE MARKET

2021

- Complete phase 3 study
- Prepare NDA filing
- Advance dialogue with payers & clinical decision makers
- Build US capability & RoW partnering
- Develop supply & logistics partnering

## PREPARE THE LAUNCH

2022

- Reach timely market authorisation
- Secure supply & logistics operations
- Mobilise US operations
- Develop RoW partnership operations
- Execute cross-functional launch
  - Payer adoption
  - Medical advocacy
  - Early adoption and preference



ADVANCING  
ORAL CHEMOTHERAPY

# ONCORAL

CHEMOTHERAPY TABLET FOR GASTRIC CANCER  
READY FOR PHASE 2

# ONCORAL – NOVEL IRINOTECAN TABLET READY FOR PHASE 2

## NOVEL ORAL PATENTED FORMULATION



Formulated as a **tablet** for convenient dosing and health-economic benefits



**Promising safety potential** of oral administration



Potential for **all-tablet chemo-combination**

## PHARMACEUTICAL INGREDIENT HAS PROVEN EFFECT



Irinotecan shown to be effective in **killing cancer cells**



Expected to be efficacious and safe **together** with other well-recognized anti-cancer drugs



**Orphan drug indication** for gastric cancer by the FDA and EMA

With promising Phase 1 results, we are now preparing for Phase 2



# ENCOURAGING ONCORAL PHASE 1 STUDY RESULTS

## Phase 1 single agent study published in Jan 2019

- Results showed that Oncoral was well tolerated; side effects were generally mild to moderate, manageable and similar in type to those observed with intravenous irinotecan
- Hematological toxicities were few and all were mild to moderate
- Pharmacokinetic (PK) data showed consistent daily exposures during treatment at days 1 and 14 with no drug accumulation
- The active metabolite, SN-38, interpatient variability was in the same range as after infusion of irinotecan
- In this heavily pre-treated patient population, Oncoral indicated activity even among patients previously treated with irinotecan infusion
- The study was presented at ESMO congress in October 2018



## Phase 1 combination study published in April 2019

- The combination of Oncoral with another oral chemotherapy, capecitabine, was encouraging which could enable an all-oral chemotherapy combination
- The study data demonstrated reassuring tolerability of Oncoral together with capecitabine
- The combination with capecitabine could become a more convenient and patient friendly treatment option compared to the intravenous formulations of these compounds
- The encouraging tolerability profile justifies further clinical studies to assess the efficacy of this treatment regimen

# TARGETING >\$4 BILLION MARKET IN GASTRIC CANCER

## DISEASE CHARACTERISTICS

- Gastric cancer is the 6<sup>th</sup> most prevalent cancer in the world<sup>1)</sup>
- Gastric cancer is the 3<sup>rd</sup> most frequent cause of cancer death<sup>1)</sup>
- The 5-year survival of gastric cancer is approximately 20%<sup>2)</sup>

## MARKET OPPORTUNITY

- Market for gastric cancer treatment forecast to increase to >\$ 4 billion in 2024<sup>3)</sup>
- Drug treatment typically combination of 2-3 drugs

### Key growth drivers

---

- ① Increase in overall incidence of gastric cancer
- ② Anticipated increase in treatment rates
- ③ Extended treatment duration
- ④ New lines of more expensive therapies
- ⑤ Increased number of patients receiving branded therapy

1) IARC (2012)

2) Clinical Colorectal Cancer 2015; 14(4): 239-50

3) GlobalData - Gastric and Gastroesophageal Junction Adenocarcinoma – Global Drug Forecast and Market Analysis to 2024



A man in a red and white checkered shirt and dark pants is walking through a field of tall, golden grass. He is holding the hand of a young child in a blue jacket. The background shows a line of trees with yellowing leaves, suggesting an autumn setting. A semi-transparent white banner is overlaid across the middle of the image, containing the title text.

# PRIORITIES 2020 AND INVESTMENT HIGHLIGHTS



# Priorities in 2020



First patient in the Phase 3 SPARKLE study



First participant in the hepatic study



Work diligently with study sites during Covid-19 and enrol additional patients



Pre-launch activities and preparations for Mangoral (market launch planned for Q4 2022 - H1 2023)



Prepare Phase 2 study for Oncoral (planned start in 2021)

# INVESTMENT HIGHLIGHTS



## Ascelia Pharma (ticker: ACE) – Advancing orphan oncology

- Drugs targeting unmet medical needs with known mode of action and low development risk
- Solid financial position: SEK 169 million in liquid assets per 31 Mar 2020



## Mangoral – Phase 3 non-gadolinium liver imaging drug

- \$350-500 million annual addressable market
- No competing drugs
- Ongoing Phase 3 program with high likelihood of success – study results expected in H2 2021
- Orphan Drug Designation



## Oncoral – Phase 2 ready oral chemotherapy for gastric cancer

- Novel tablet formulation with significant patient and hospital benefits
- Effective molecule for killing cancer
- Promising Phase 1 results and preparing for Phase 2

# ASCELIA PHARMA

[ascelia.com](https://ascelia.com)

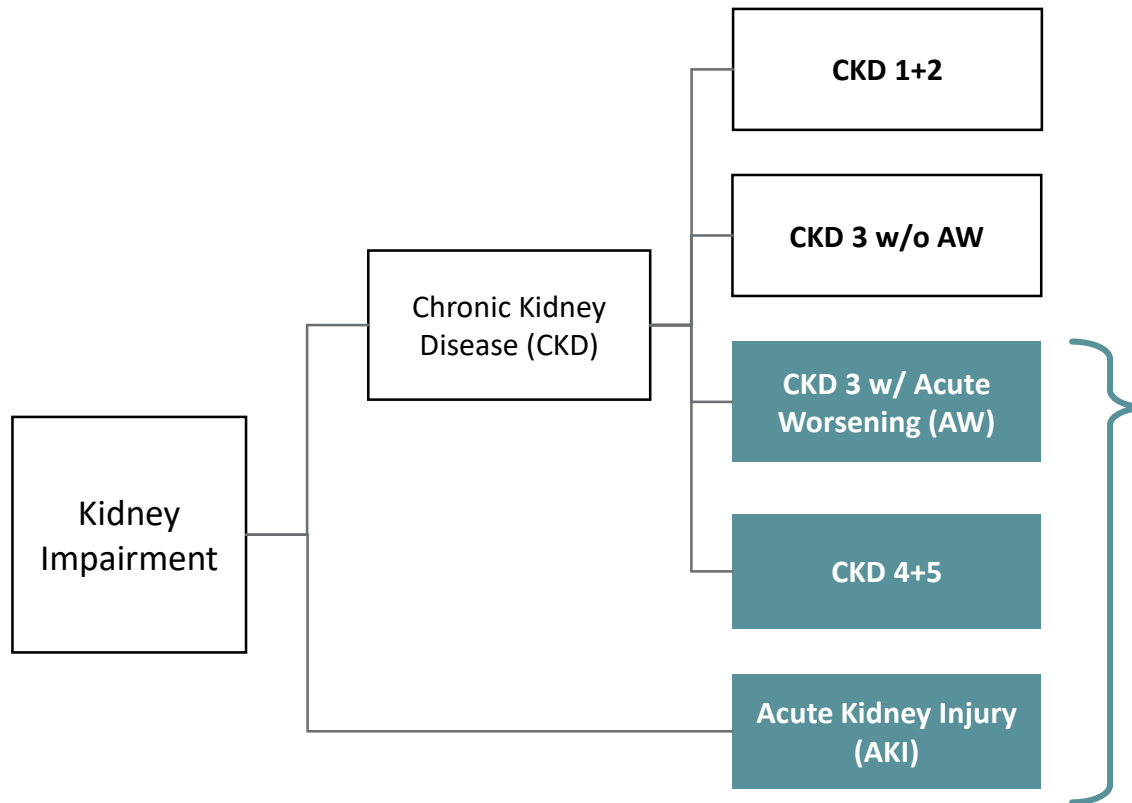




# APPENDIX – MARKET MODEL

# TARGET SEGMENT

PATIENTS WITH KNOWN OR SUSPECTED **FOCAL LIVER LESIONS** AND SEVERELY **IMPAIRED KIDNEY FUNCTION**



## Mangoral Target Segment:

**MRI and CT liver scans** for high-risk patients with severely impaired kidney function:

- eGFR < 30 mL/min/1.73m<sup>2</sup>, or
- Acute Kidney Injury



# MEASURING KIDNEY IMPAIRMENT IN PATIENTS

- According to *American Society of Nephrology*, CKD 4 is defined as  $eGFR < 30 \text{ mL/min/1.73m}^2$  on two occasions separated by  $\geq 90$  days and that is not associated with a transient, reversible condition such as volume depletion
- eGFR is estimated based on serum creatinine measurement, which is considered good for measuring stable patients over time, but is less good in capturing acute changes in kidney function, where the calculated eGFR is higher than the *true GFR*
- Kidney function can be temporarily reduced due to factors such as nephrotoxic drugs (including chemo and antibiotics), surgery, hospitalization, trauma and dehydration



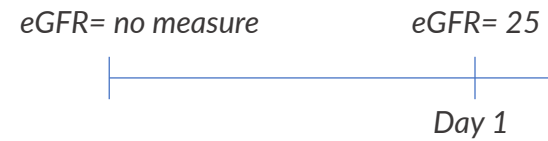
# DEFINITION OF CHRONIC KIDNEY DISEASE 4/5

All of these patients are candidates for Mangoral, but only one is categorized as a CKD 4/5 patient

CKD 4 patient



Not CKD patient



Not CKD patient



CKD 3 patient








# MANGORAL PATIENT POPULATION

- Mangoral can be used for visualization of all focal liver lesions, but is expected to be predominantly used in patients with known or suspected cancer in the liver. The majority of patients will have liver metastases and a small minority will have primary liver cancer
- Today, kidney function is assessed in clinical practice in patients at risk of kidney impairment by measuring serum creatinine and calculating eGFR
- eGFR is reliable in patients in stable condition and with a history of previous eGFR measurements for assessing development in kidney function. It is less reliable in situations where the patient either
  - Has no previous eGFR measurements, or
  - Is in unstable condition that can have acute effects on kidney function
- The estimate of eligible patients for Mangoral therefore includes both CKD 4/5 patients as well as patients with acute changes in kidney function

# \$350-500M ADRESSABLE MARKET WITH NO COMPETITION

## Top-down analysis

Est. cancer patients in 2020 ('000) <sup>1)</sup>		 <sup>2)</sup>	
Colorectal cancer	570	1,441	491
Liver cancer	64	133	120
Other relevant cancers (breast, lung and gastric cancer)	1,792	2,875	992
<b>Total</b>	<b>2,426</b>	<b>4,449</b>	<b>1,603</b>
	<div style="border: 1px solid black; padding: 5px; margin: 5px 0;"> <b>1</b> Age-distributed prevalence of CKD in the chosen cancer indications         </div>		
	<div style="border: 1px solid black; padding: 5px; margin: 5px 0;"> <b>2</b> Adjust for higher cancer risk in patients with CKD         </div>		
<b>Addressable CKD patients ('000)<sup>4)</sup></b>			
CKD 4/5	37	46	30
CKD 3 with acute worsening	55	69	45
<b>Total CKD patients for Mangoral®</b>	<b>92</b>	<b>115</b>	<b>75</b>
	<div style="border: 1px solid black; padding: 5px; margin: 5px 0;"> <b>3</b> Scans per CKD patient – adjusted for the frequency of scans and rate of newly diagnosed patients in the cancer indications         </div>		
<b># Annual Mangoral® scans ('000)</b>	<b>64</b>	<b>95</b>	<b>61</b>
	<div style="border: 1px solid black; padding: 5px; margin: 5px 0;"> <b>4</b> \$1,500 - \$3,000 per dose of Mangoral based on Value-based-pricing         </div>		

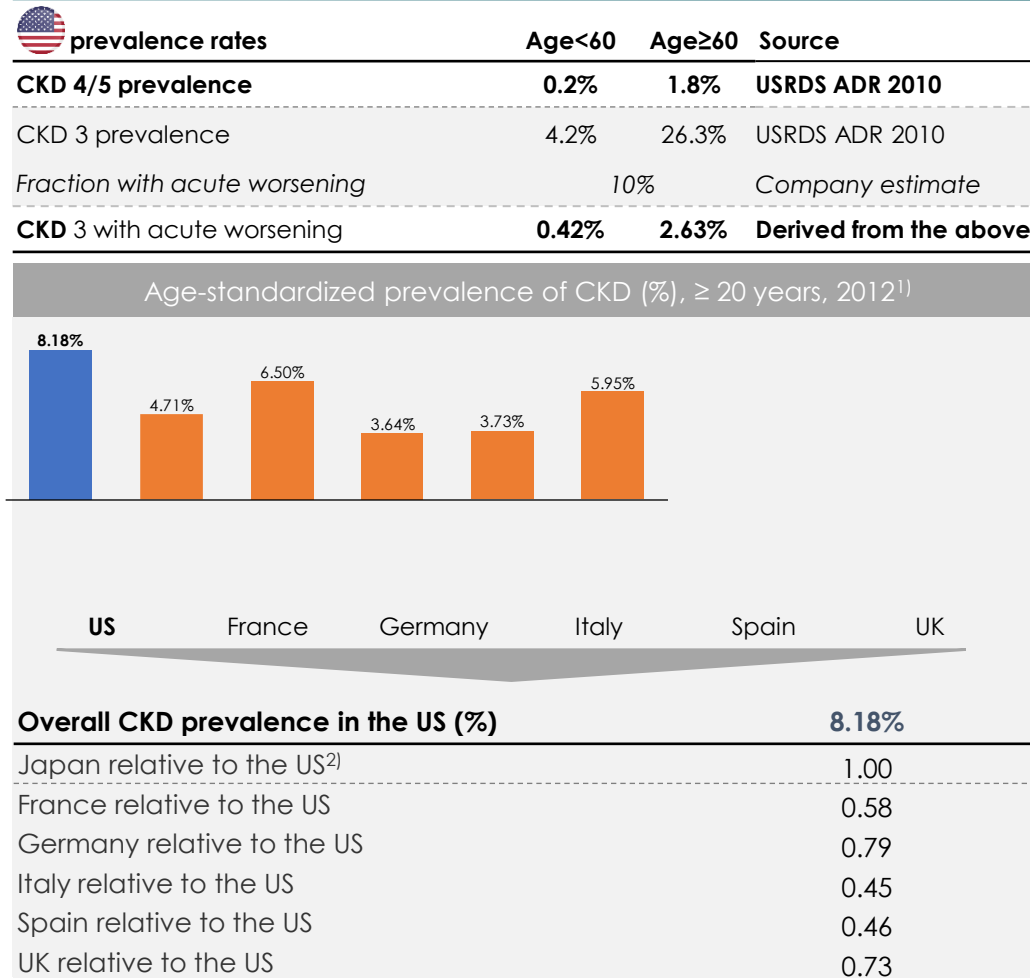
## Explanatory comments

- 1** The prevalence of CKD (Chronic Kidney Disease) varies across the geographies, as well as the age distribution of the population
- 2** The risk of cancer in CKD varies across cancer indications, and is larger in Colorectal than in e.g. Breast and Liver<sup>3)</sup>
- 3** The actual number of expected scans have been adjusted for the different frequency of scanning per cancer indication, as well as the different # of scans given to patients depending on where they are in the treatment cycle
- 4** Expected pricing level has been discussed with >25 payors in the US and Europe











An addressable market of USD 350-500m

# 1 THE PREVALENCE OF CKD VARIES ACROSS GEOGRAPHIES AND AGE

US-specific figures for CKD stage-specific prevalence have been used as a basis



- The detailed prevalence figures obtained in the US for the various stages of CKD have been used as a basis for the CKD prevalence calculations
- Japan's CKD stage-specific prevalence has been found to be relatively identical to the US
- For the EU countries, their stage-specific CKD prevalence have been calculated using the US figures adjusted for these countries' overall CKD prevalence vs. the US

Geography	Prevalence rates	Age<60	Age≥60
 France	CKD 4/5 prevalence	0.12%	1.04%
 France	CKD 3 with acute worsening prevalence	0.24%	1.52%
 Germany	CKD 4/5 prevalence	0.16%	1.43%
 Germany	CKD 3 with acute worsening prevalence	0.33%	2.09%
 Italy	CKD 4/5 prevalence	0.09%	0.80%
 Italy	CKD 3 with acute worsening prevalence	0.19%	1.17%
 Spain	CKD 4/5 prevalence	0.09%	0.82%
 Spain	CKD 3 with acute worsening prevalence	0.19%	1.20%
 UK	CKD 4/5 prevalence	0.15%	1.31%
 UK	CKD 3 with acute worsening prevalence	0.31%	1.91%

1) Global Data 'Chronic Kidney Disease - Epidemiology Forecast to 2022', August 2013  
 2) Clin Exp Nephrol. 2009 Dec;13(6):621-30, Prevalence of chronic kidney disease in the Japanese general population, Imai et al.

## 2 RISK OF COLORECTAL CANCER IS HIGHER IN THE CKD POPULATION

### Overview

Risk factor for cancer	Age<50	Age≥50	Source
Colorectal cancer	3.7	1.64	Ann Surg Oncol (2013) 20:3885–3891, Risk of Colorectal Cancer in Chronic Kidney Disease: A Matched Cohort Study Based on Administrative Data, Wu et al.
Liver cancer	1.0	1.0	No relevant source, i.e. no adjustment to risk factor
Other relevant cancers (breast, lung and gastric cancer)	1.0	1.0	No relevant source, i.e. no adjustment to risk factor

### Effect on calculation

The risk of developing colorectal cancer has been found to be 1.64-3.7x larger for patients with CKD than others

To adjust for this, the CKD prevalence figures (%) have been multiplied by this factor when the addressable CKD cases within the colorectal cancer population have been calculated

*A detailed example follows on the next slide*

### Abstract

#### Background

To investigate the risk of CRC in patients with CKD

#### Methods

- Study cohort included patients aged ≥18 years diagnosed with CKD between 2004 and 2005 (n = 15,975)
- Comparison cohort (n = 79,875) included five randomly selected age- and gender-matched controls for each patient in the study cohort
- All the subjects were followed up from the date of cohort entry until they developed CRC or until the end of 2006.


#### Results

- 460 patients developed CRC during the study period, of whom 116 were from the CKD cohort and 344 were from the comparison cohort
- The age-matched hazard ratio of CRC after excluding dialysis patients was **1.64** (95% CI 1.27-2.11) in patients aged ≥50 years, and **3.7** (95% CI 1.83-7.49) in patients younger <50 years



# EXAMPLE OF HOW THESE ASSUMPTIONS ARE UTILISED IN THE MODEL

Geography: US / Cancer indication: Colorectal

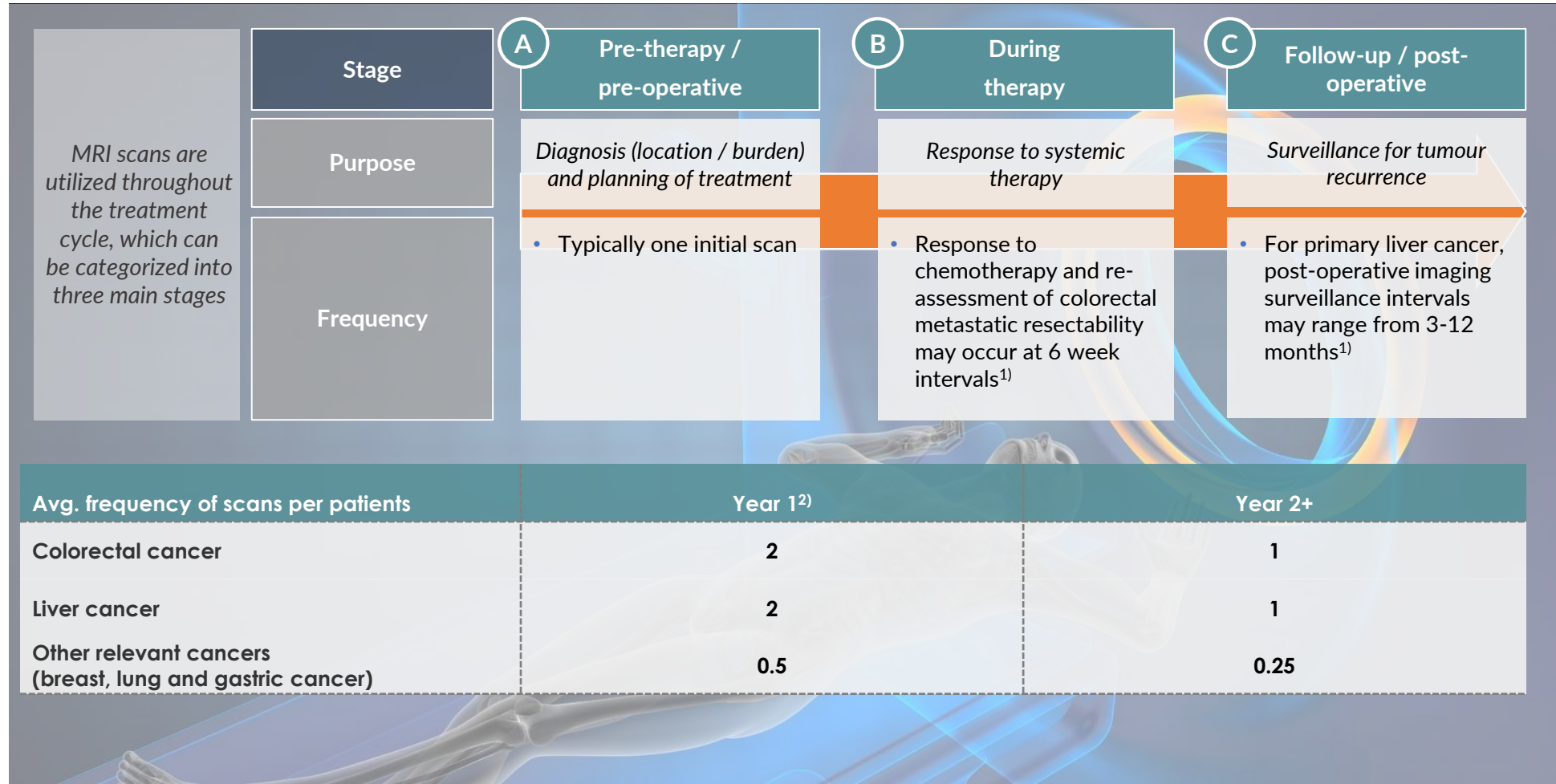
 Patients / age intervals	0-14	15-39	40-44	45-49	50-54	55-59	60-64	65-69	70-74	≥75	Total
Colorectal cancer patients by 2020 ('000) <sup>1)</sup>	0.1	14.8	17.9	32.0	48.3	60.9	71.2	90.2	82.6	234.0	652.1
<b>A</b> CKD 4/5 prevalence	0.2%						1.8%				
CKD 3 prevalence	4.2%						26.3%				
Fraction with acute worsening	10%										
<b>B</b> CKD 3 with acute worsening	0.42%						2.63%				
<b>C</b> Risk factor	3.7					1.64					
Risk-adjusted CKD 4/5 prevalence (%)	0.74%	0.74%	0.74%	0.74%	0.33%	0.33%	2.95%	2.95%	2.95%	2.95%	
Risk-adjusted CKD 3 with acute worsening prevalence (%)	1.55%	1.55%	1.55%	1.55%	0.69%	0.69%	4.31%	4.31%	4.31%	4.31%	
CKD 4/5 patients ('000)	0.001	0.110	0.133	0.237	0.158	0.200	2.102	2.664	2.437	6.908	14.949
CKD 3 with acute worsening patients ('000)	0.001	0.231	0.278	0.498	0.332	0.420	3.072	3.892	3.561	10.093	22.377
<b>Total ('000)</b>	<b>0.002</b>	<b>0.341</b>	<b>0.411</b>	<b>0.735</b>	<b>0.491</b>	<b>0.620</b>	<b>5.174</b>	<b>6.556</b>	<b>5.998</b>	<b>17.001</b>	<b>37.326</b>

Est. total of **37,326** relevant CKD patients in the US colorectal cancer population

1) Based on Globocan 2012.

### 3 IMAGING FREQUENCY DEPENDS ON DISEASE STATE, TREATMENT, PROGNOSIS AND RISK PROFILE

The treatment cycle

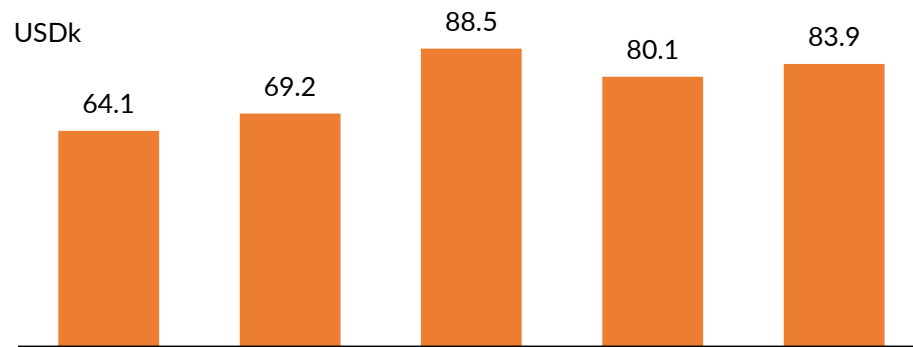


1) AJR Am J Roentgenol, 2014, 203(1):W21-33

2) The newly diagnosed rate for Colorectal, Liver and Other relevant cancers have by Management been assumed to be at 25%, 50% and 25%, respectively. These figures have been derived based on the observed incidence / prevalence across these indications.

# 4 MANGORAL IS AN ORPHAN HIGH VALUE DIAGNOSTIC DRUG WITH EXPECTED STRONG PRICING POWER

Median cost per patient p.a. for orphan drugs, US, 2012-2016<sup>1)</sup>



> 25 payors

- Ascelia Pharma has held discussions with more than 25 payors on Mangoral's market access



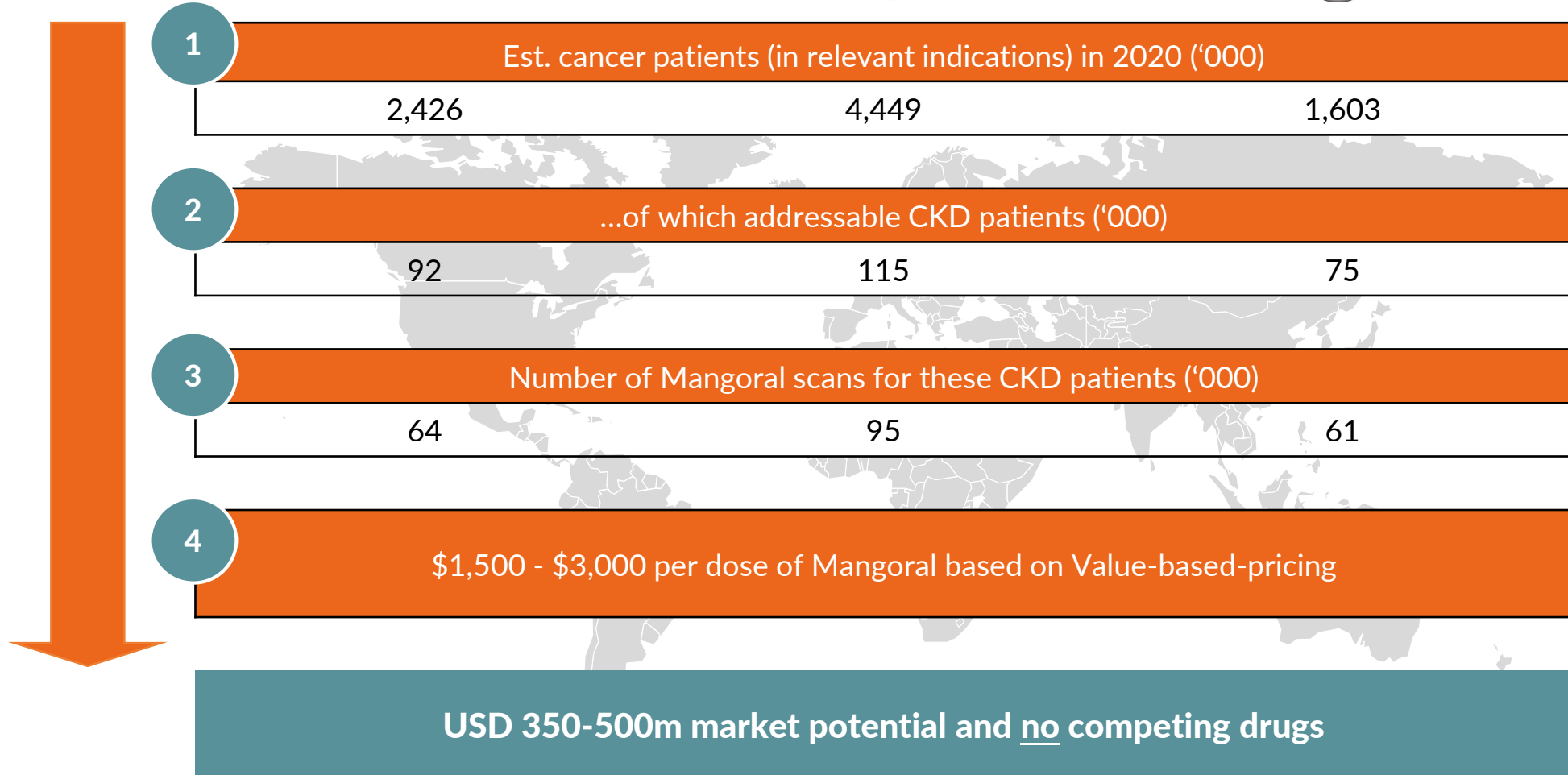
- Feedback from these discussions support the plan for a unique Mangoral reimbursement code and value based pricing

Pricing of other relevant high value diagnostics

Test	Type	Company	Use	Est. price per dose (USDk)
Choline C-11 <sup>2)</sup>	In-vivo (injection)	Zevacor Pharma	PET imaging	5.7
				2.2
Afirma Gene Expression Classifier <sup>3)</sup>	In-vitro	Veracyte	Preoperative microarray test	3.4
Oncotype DX Gene expression microarray <sup>4)</sup>	In-vitro	Genomic Health	Gene expression microarray test	3.7
<b>Axumin (fluciclovine F 18) PET agent<sup>5)</sup></b>	In-vivo (injection)	Blue Earth Diagnostics	PET imaging	<b>3.7</b>

1) Source: EvaluatePharma (Orphan Drug Report 2017)  
 2) SNMI 2016 ([http://snmmi.files.cms-plus.com/docs/hpra/SNMMI%20HOPPS%202016F%20vs%202017P\\_update.pdf](http://snmmi.files.cms-plus.com/docs/hpra/SNMMI%20HOPPS%202016F%20vs%202017P_update.pdf))  
 3) Veracyte 2016 (<http://investor.veracyte.com/releasedetail.cfm?releaseid=975334>)  
 4) Genomic Health 2015 (<http://investor.genomichealth.com/releasedetail.cfm?releaseid=935522>)  
 5) Axumin 2017 (<http://www.axumin.com/pdf/Pass-thru.pdf>)

# IN SUMMARY...





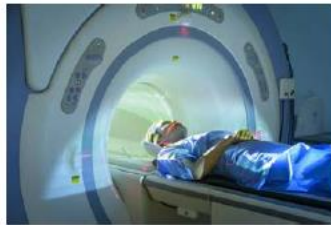
# GADOLINIUM BRAIN ACCUMULATION – UPSIDE POTENTIAL

## Gadolinium under scrutiny for brain accumulation

- Recently, concerns have been raised for gadolinium retention in the brain in all patients regardless of kidney function
- Regulatory agencies have issued warnings and suspended gadolinium-based products
- Concerns about brain accumulation have also sparked media interest (incl. WSJ and Washington Post)

### A Question for Anyone Getting an MRI

Patients need to know if the doctor plans to use contrast, or gadolinium, because it may leave harmful metal deposits; a new FDA warning



THE WALL STREET JOURNAL  
**WSJ**

Millions of magnetic resonance imaging, or MRI, scans are performed annually in the U.S. to look for tumors among other ailments. PHOTO: GETTY IMAGES



By  
*Sumathi Reddy*

Sept. 18, 2017 12:21 p.m. ET

For Ascelia, further limitations to gadolinium products could offer significant upside to the addressable market

## Selected recent regulatory actions



**19 December 2017:**

FDA Drug Safety Communication: FDA warns that gadolinium-based contrast agents (GBCAs) are retained in the body; requires new class warnings



EUROPEAN MEDICINES AGENCY  
SCIENCE. MEDICINES. HEALTH.

**23 November 2017:**

EMA's final opinion confirms suspension and restrictions on use of linear gadolinium agents in body scans



**6 February 2018:**

UK implements EMA decision and Omniscan and intravenous Magnevist are now no longer authorised for use and a product recall of any existing unexpired stock is underway



**8 December 2017:**

Japan's Ministry of Health, Labour and Welfare asks makers of gadolinium-based contrast agents (GBCAs) for MRI scans to revise warning text



**1 April 2018:**

Sweden implements the EMA recommendations. Since gadolinium may ingrain itself in the human brain with unknown long term side effects

# DE-RISKED PHASE 3 STUDY COMPARED TO TYPICAL PHASE 3

Parameter	TYPICAL ONCOLOGY pivotal Phase 3 trial	MANGORAL pivotal Phase 3 trial
Patient sample size	<ul style="list-style-type: none"><li>• 500-1,000+</li></ul>	<ul style="list-style-type: none"><li>✓ Up to 200 patients</li></ul>
Study design	<ul style="list-style-type: none"><li>• Patients randomised into separate study arms thus increasing variation</li></ul>	<ul style="list-style-type: none"><li>✓ Each patient his/her own control thus minimising variation</li></ul>
API	<ul style="list-style-type: none"><li>• Often a novel molecule</li></ul>	<ul style="list-style-type: none"><li>✓ Manganese</li></ul>
Study comparator	<ul style="list-style-type: none"><li>• Add-on to standard-of-care or evaluated head-to-head vs. another active treatment regimen</li></ul>	<ul style="list-style-type: none"><li>✓ Add-on to standard-of-care (unenhanced MRI vs. unenhanced MRI + Mangoral enhanced MRI)</li></ul>
Follow-up time to endpoint	<ul style="list-style-type: none"><li>• Months to years</li></ul>	<ul style="list-style-type: none"><li>✓ Days</li></ul>





# APPENDIX – Q1 2020 HIGHLIGHTS

# SIGNIFICANT MILESTONES REACHED IN 2020

## Key events in Q1-2020



First patient in Mangoral's Phase 3 study SPARKLE



Ascelia Pharma wins the award as Malmö's Best Life Science company

## Key events after the period



First participant in the hepatic study for Mangoral (May 2020)



Patent approval for Oncoral in Japan (Apr 2020)

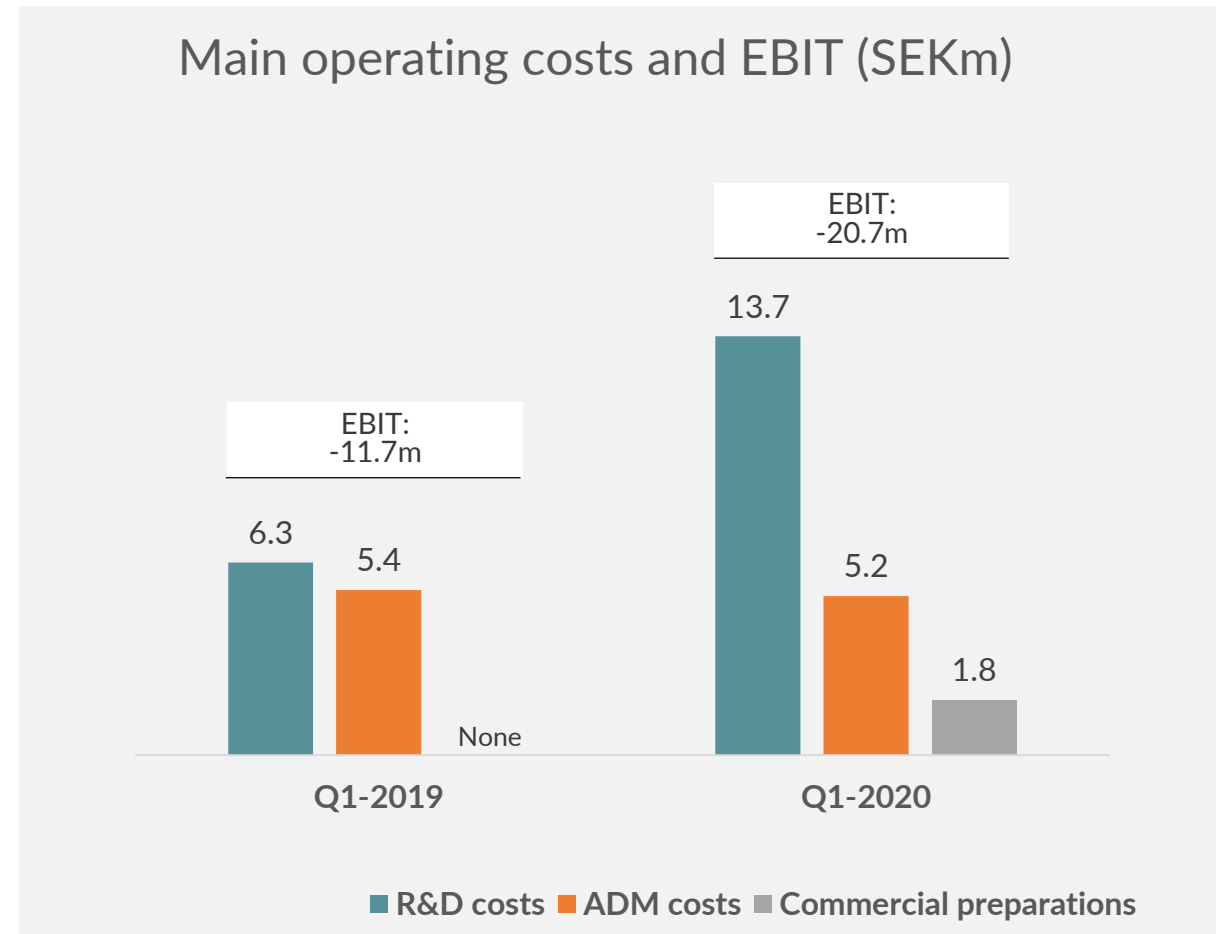


# FINANCIAL HIGHLIGHTS – OPERATING RESULTS

Increased operating loss y/y mainly driven by higher R&D activity for Mangoral's Phase 3 study:

- Preparing and opening of clinical study sites
- Manufacturing preparations
- Regulatory preparations

Also cost for commercial preparations for Mangoral incurred in Q1-2020 (none in Q1-2019)



# FINANCIAL HIGHLIGHTS – LIQUIDITY POSITION

## Continued strong liquidity:

- Liquid assets incl. marketable securities of SEK 169.3 million per 31 Mar 2020
- Liquidity to fund Mangoral clinical development and pre-commercial activities

